

S ADA American Dental Association®

PREVENTION OF ORTHOPAEDIC IMPLANT INFECTION IN PATIENTS UNDERGOING DENTAL **PROCEDURES**

EVIDENCE-BASED GUIDELINE AND EVIDENCE REPORT

Disclaimer

This clinical guideline was developed by a physician and dentist volunteer Work Group and experts in systematic reviews. It is provided as an educational tool based on an assessment of the current scientific and clinical information and accepted approaches to treatment. The recommendations in this guideline are not intended to be a fixed protocol as some patients may require more or less treatment or different means of diagnosis. Patients seen in clinical practice may not be the same as those found in a clinical trial. Patient care and treatment should always be based on a clinician's independent medical judgment given the individual clinical circumstances.

Disclosure Requirement

In accordance with AAOS policy, all individuals whose names appear as authors or contributors to this clinical practice guideline filed a disclosure statement as part of the submission process. All panel members provided full disclosure of potential conflicts of interest prior to beginning work on the recommendations contained within this clinical practice guideline.

Funding Source

No funding from outside commercial sources to support the development of this document.

FDA Clearance

Some drugs or medical devices referenced or described in this clinical practice guideline may not have been cleared by the Food and Drug Administration (FDA) or may have been cleared for a specific use only. The FDA has stated that it is the responsibility of the physician to determine the FDA clearance status of each drug or device he or she wishes to use in clinical practice.

Copyright

All rights reserved. No part of this clinical practice guideline may be reproduced, stored in a retrieval system, or transmitted, in any form, or by any means, electronic, mechanical, photocopying, recording, or otherwise, without prior written permission from the AAOS.

Published 2012 by the American Academy of Orthopaedic Surgeons 6300 North River Road Rosemont, IL 60018 First Edition Copyright 2012 by the American Academy of Orthopaedic Surgeons & American Dental Association

Summary of Recommendations

The following is a summary of the recommendations of the AAOS-ADA clinical practice guideline, Prevention of Orthopaedic Implant Infection in Patients Undergoing Dental Procedures. This summary does not contain rationales that explain how and why these recommendations were developed, nor does it contain the evidence supporting these recommendations. All readers of this summary are strongly urged to consult the full guideline and evidence report for this information. We are confident that those who read the full guideline and evidence report will see that the recommendations were developed using systematic evidence-based processes designed to combat bias, enhance transparency, and promote reproducibility.

This summary of recommendations is not intended to stand alone. Treatment decisions should be made in light of all circumstances presented by the patient. Treatments and procedures applicable to the individual patient rely on mutual communication between patient, physician, dentist and other healthcare practitioners.

1. The practitioner might consider discontinuing the practice of routinely prescribing prophylactic antibiotics for patients with hip and knee prosthetic joint implants undergoing dental procedures.

Grade of Recommendation: Limited

Definition: A **Limited** recommendation means the quality of the supporting evidence that exists is unconvincing, or that well-conducted studies show little clear advantage to one approach versus another.

Evidence from two or more "Low" strength studies with consistent findings, or evidence from a single Moderate quality study recommending for or against the intervention or diagnostic.

Implications: Practitioners should be cautious in deciding whether to follow a recommendation classified as **Limited**, and should exercise judgment and be alert to emerging publications that report evidence. Patient preference should have a substantial influencing role.

2. We are unable to recommend for or against the use of topical oral antimicrobials in patients with prosthetic joint implants or other orthopaedic implants undergoing dental procedures.

Grade of Recommendation: Inconclusive

Definition: An **Inconclusive** recommendation means that there is a lack of compelling evidence resulting in an unclear balance between benefits and potential harm. Evidence from a single low quality study or conflicting findings that do not allow a recommendation for or against the intervention.

Implications: Practitioners should feel little constraint in deciding whether to follow a recommendation labeled as **Inconclusive** and should exercise judgment and be alert to publications that clarify existing evidence for determining balance of benefits versus potential harm. Patient preference should have a substantial influencing role.

3. In the absence of reliable evidence linking poor oral health to prosthetic joint infection, it is the opinion of the work group that patients with prosthetic joint implants or other orthopaedic implants maintain appropriate oral hygiene.

Grade of Recommendation: Consensus

Definition: A **Consensus** recommendation means that expert opinion supports the guideline recommendation even though there is no available empirical evidence that meets the inclusion criteria.

The supporting evidence is lacking and requires the work group to make a recommendation based on expert opinion by considering the known potential harm and benefits associated with the treatment.

Implications: Practitioners should be flexible in deciding whether to follow a recommendation classified as **Consensus**, although they may set boundaries on alternatives. Patient preference should have a substantial influencing role.

Terminology Used in This Guideline

Direct evidence – Evidence that demonstrates a relationship between a dental procedure and orthopaedic implant infection.

Indirect evidence – Evidence that demonstrates a relationship between a dental procedure and a surrogate outcome (i.e. bacteremia).

Incidence – New cases of a disease that occur in an at-risk population during a specified time period (i.e. a new bacteremia after a dental procedure)

Prevalence – Existing cases of a disease in a population during a specified time period (i.e. a bacteremia that exists prior to a dental procedure)

Case-control study – Comparison of a diseased group (cases) to a group without disease (controls)

Surrogate Outcome – An outcome (such as a laboratory measurement) that is used as a substitute for a clinically relevant patient centered outcome

High, Moderate, and Low Strength Studies – Derived from quality and applicability analysis; integrating multiple domains composed of questions related to study design and methods (See Appraising Evidence Quality and Applicability)

Prevention of Orthopaedic Implant Infection in Patients Undergoing Dental Procedures Clinical Practice Guideline Work Group

American Academy of Orthopaedic Surgeons William Watters, III, MD, Co-Chair

Bone and Joint Clinic of Houston 6624 Fannin Street, #2600 Houston, TX 77030

American Dental Association

Michael P. Rethman, DDS, MS, Co-Chair

47-140 Heno Place Kaneohe, HI 96744

American Academy of Orthopaedic Surgeons

Richard Parker Evans, MD

Professor and Margaret Sue Neal Endowed Chair of Orthopaedic Surgery University of Missouri- Kansas City School of Medicine 2301 Holmes Street Kansas City, MO 64108

American Academy of Orthopaedic Surgeons

Calin Moucha, MD

Associate Chief, Joint Replacement Surgery Mount Sinai Medical Center **Assistant Professor** Leni & Peter W. May Department of Orthopaedic Mount Sinai School of Medicine 5 E. 98th Street, Box 1188, 7th Floor New York, NY 10029

American Academy of Orthopaedic Surgeons Richard J. O'Donnell, MD

Chief, UCSF Orthopaedic Oncology Service UCSF Sarcoma Program UCSF Helen Diller Family Comprehensive Cancer Center 1600 Divisadero Street, 4th Floor San Francisco, CA 94115

American Academy of Orthopaedic Surgeons & **Congress of Neurological Surgeons**

Paul A. Anderson, MD

Professor Department of Orthopedics & Rehabilitation University of Wisconsin K4/735 600 Highland Avenue Madison WI 53792

American Dental Association

Elliot Abt, DDS

4709 Golf Road, Suite 1005 Skokie, IL 60076

American Dental Association

Harry C. Futrell, DMD

330 W 23rd Street, Suite J Panama City, FL 32405

American Dental Association

Stephen O. Glenn, DDS

5319 S Lewis Avenue, Suite 222 Tulsa, OK 74105-6543

American Dental Association

John Hellstein, DDS, MS

The University of Iowa, College of Dentistry Department of Oral Pathology, Radiology and Medicine **DSB S356** Iowa City, IA 52242

American Association of Hip and Knee Surgeons

David Kolessar, MD

Geisinger Wyoming Valley Medical Center 1000 East Mountain Boulevard Valley Medical Building Wilkes-Barre, PA 18711

American Association of Neurological Surgeons/Congress of Neurological Surgeons

John E. O'Toole, MD

Assistant Professor of Neurosurgery Rush University Medical Center 1725 W. Harrison Street, Suite 970 Chicago, IL 60612

American Association of Oral and Maxillofacial

Surgeons

Mark J. Steinberg DDS, MD 1240 Meadow Road, Suite 300 Northbrook, IL 60062

College of American Pathologist

Karen C. Carroll MD, FCAP

Johns Hopkins Hospital Department of Pathology-Microbiology Division 600 N Wolfe Street Meyer B1-193 Baltimore, MD 21287

Knee Society

Kevin Garvin, MD

University of Nebraska Medical Center Creighton/Nebraska Health Fund Department of Orthopaedic Surgery 981080 Nebraska Medical Center Omaha, Nebraska 68198-1080

Musculoskeletal Infection Society

Douglas R. Osmon, MD

200 1st Street SW Rochester, MN 55905

Scoliosis Research Society

Anthony Rinella, MD

Illinois Spine & Scoliosis Center 12701 West 143rd Street, Suite 110 Homer Glen, Illinois 60491

Society for Healthcare Epidemiology of America

Angela Hewlett, MD, MS

Assistant Professor, Section of Infectious Diseases University of Nebraska Medical Center 985400 Nebraska Medical Center Omaha, Nebraska 68198

Guidelines Oversight Chair

Michael J. Goldberg, MD

Children's Hospital and Regional Medical Center 1221 1st Avenue, Apt #24E Seattle, WA 98101

AAOS Staff

Deborah S. Cummins, PhD

Director, Research and Scientific Affairs 6300 N. River Road Rosemont, IL 60018

Sharon Song, PhD

Manager, Clinical Practice Guidelines song@aaos.org

Patrick Sluka, MPH

Former AAOS Lead Research Analyst

Kevin Boyer

Former Appropriate Use Criteria Unit Manager Former Interim Clinical Practice Guidelines Manager

Anne Woznica, MLIS

Medical Research Librarian

ADA Staff

Helen Ristic, PhD.

Director, Scientific Information ADA Division of Science 211 E. Chicago Avenue Chicago, IL 60611

Nicholas Buck Hanson, MPH

ADA Lead Research Analyst hansonn@ada.org

Special Recognitions

William Robert Martin, III, MD

Medical Director 317 Massachusetts Avenue NE Washington, D.C. 20002-5769

American Academy of Orthopaedic Surgeons

Additional collaborating organizations involved in this guideline development:

Infectious Disease Society of America (IDSA)

Peer Review Organizations

Participation in the AAOS peer review process does not constitute an endorsement of this guideline by the participating organization.

The following organizations participated in peer review of this clinical practice guideline and gave explicit consent to be listed as peer reviewers:

The Academy of General Dentistry

American Academy of Oral Pathology

American Academy of Orthopaedic Surgeon's Evidence Based Practice Committee

American Academy of Orthopaedic Surgeons' Guidelines Oversight Committee

American Academy of Pediatric Dentistry

American Academy of Periodontology

American Association of Family Physicians

American Association of Hip and Knee Surgeons

American Association of Oral and Maxillofacial Surgeons

American Association of Public Health Dentistry

American College of Prosthodontists

American Dental Association's Council on Scientific Affairs

American Dental Hygienists Association

Canadian Dental Association

Centers for Disease Control and Prevention

College of American Pathologists

Lumbar Spine Research Society

North American Spine Society

Society of Infectious Diseases Pharmacists

The Infectious Diseases Society of America

Participation in the AAOS peer review process does not constitute an endorsement of this guideline by the participating organization.

Table of Contents

| Summary of Recommendations | |
|--------------------------------------------------------|-----|
| Terminology Used in This Guideline | |
| Work Group | |
| Peer Review Organizations | vii |
| Table of Contents | |
| List of Tables | xi |
| List of Figures | xiv |
| Introduction | 1 |
| Overview | |
| Goals and Rationale | 1 |
| Intended Users | |
| Patient Population | |
| Burden of Disease and Etiology | |
| Potential Harms, Benefits, and Contraindications | |
| Preventing Bias in an AAOS Clinical Practice Guideline | 3 |
| Methods | |
| Formulating Preliminary Recommendations | |
| Full Disclosure Information | |
| Study Selection Criteria | |
| Literature Searches | |
| Best Evidence Synthesis | |
| Appraising Evidence Quality and Applicability | |
| Studies of Interventions | |
| Studies of Incidence and Prevalence | |
| Studies of Prognostics | |
| Other Biases In the Published Literature | |
| Grades of Recommendation | |
| Wording of the Final Recommendations | |
| Consensus Recommendations | |
| Voting on the Recommendations | |
| Outcomes Considered | |
| Statistical Methods | |
| Peer Review | |
| Public Commentary | |
| The AAOS Guideline Approval Process | |
| Revision Plans | |
| Guideline Dissemination Plans | |
| Overview of the Evidence | |
| Direct Evidence | |
| Findings | |
| Quality and Applicability | |
| Results | |
| Indirect Evidence: Dental Procedures and Bacteremia | |
| FindingsQuality and Applicability | |
| Quanty and Applicatinity | ,∠c |

| Results | 29 |
|-------------------------------------------------------------------------|-----|
| Indirect Evidence: Risk Factors for Dental Procedure Related Bacteremia | 33 |
| Findings | |
| Quality and Applicability | |
| Results | |
| Indirect Evidence: Prophylaxis for Dental Procedure Related Bacteremia | |
| Findings | |
| Quality and Applicability | |
| Results | |
| Indirect Evidence: Background Microbiology | |
| Findings | |
| Results | |
| Recommendations | |
| Recommendation 1 | 75 |
| Rationale | |
| Findings | |
| Quality and Applicability | |
| Results | |
| Recommendation 2 | |
| Rationale | |
| Findings | |
| Quality and Applicability | |
| Results | |
| Recommendation 3 | |
| Rationale | |
| Findings | |
| Quality and Applicability | |
| Future Research | |
| Appendices | |
| Appendix I | |
| Work Group | |
| Appendix II | |
| Creating Preliminary Recommendations | |
| Appendix III | |
| Study Attrition Diagram | |
| Included Studies Tables | |
| Excluded Studies Tables | |
| Appendix IV | |
| Medical Librarian Search Strategy | |
| Supplemental Search | |
| Appendix V | |
| Evaluating Quality and Applicability | |
| Studies of Interventions | |
| Studies of Incidence and Prevalence | |
| Studies of Prognostics | |
| Annandiy VI | 230 |

| Rules for Opinion Based Consensus Recommendations | 239 |
|------------------------------------------------------------|-----|
| Checklist for Voting on Consensus Recommendations | 240 |
| Appendix VII | 241 |
| Voting with the Nominal Group Technique | |
| Appendix VIII | 242 |
| Structured Peer Review Form | |
| Appendix IX | 246 |
| Peer Review | |
| Public Commentary | 248 |
| Appendix X | |
| AAOS Bodies That Approved This Clinical Practice Guideline | 249 |
| ADA Bodies That Approved This Clinical Practice Guideline | 249 |
| Documentation of Approval | 250 |
| Appendix XI | 251 |
| Supplemental Evidence Tables | 251 |
| Appendix XII | |
| Quality and Applicability Tables for Included Studies | 263 |
| Appendix XIII | 1 |
| Conflict of Interest | |
| References | |

List of Tables

| Table 1 IOM Clinical Practice Guidelines Standards | 7 |
|--------------------------------------------------------------------------------------------------------------|-----------|
| Table 2 IOM Systematic Review Standards | 8 |
| Table 3 Relationship between Quality and Domain Scores for Interventions | 12 |
| Table 4 Relationship between Applicability and Domain Scores for Interventions | 13 |
| Table 5 Relationship between Quality and Domain Scores for Incidence and Prevalence Studies | 13 |
| Table 6 Relationship between Applicability and Domain Scores for Incidence and Prevalence Studies | 14 |
| Table 7 Relationship between Quality and Domain Scores for Prognostic Studies | |
| Table 8 Relationship between Applicability and Domain Scores for Prognostic Studies | |
| Table 9 Strength of Recommendation Descriptions | |
| Table 10 AAOS Guideline Language | |
| Table 10 High and Low Risk Dental Procedures Defined by Berbari, et al. | |
| Table 11 Dental procedures performed and risk of prosthetic hip or knee infection at 6 months and 2 years | |
| Table 12 Antibiotic prophylaxis and risk of prosthetic hip or knee infection at 6 months and 2 years | |
| Table 13 Summary of Risk Factor Significance (Proportion of studies that reported significant results) | |
| Table 14 Risk Factors for Brushing Bacteremia | |
| Table 15 Risk Factors for Chewing Bacteremia | |
| Table 16 Risk Factors for Dental Prophylaxis Bacteremia | |
| Table 17 Risk Factors for Inter-dental Cleaning Bacteremia | |
| Table 18 Risk Factors for Intubation Bacteremia | |
| Table 19 Risk Factors for Oral Surgery Bacteremia | |
| Table 20 Risk Factors for Periodontic Bacteremia | |
| Table 21 Risk Factors for Restorative Bacteremia | 43 |
| Table 22 Risk Factors for Extraction Bacteremia | 44 |
| Table 23 Antibiotic prophylaxis and tooth extraction bacteremia | |
| Table 24 Topical antimicrobials and tooth extraction bacteremia | |
| Table 25 Orthopaedic implant cohort studies | |
| Table 26 Orthopaedic implant case series studies | 58 |
| Table 27 Direct Comparisons of Antibiotic Prophylaxes for the Prevention of Dental-related Bacteremia | 78 |
| Table 28 Indirect (Network) Comparisons of Antibiotic Prophylaxes for the Prevention of Dental-related Ba | cteremia |
| | 80 |
| Table 29 Indirect (Network) Significant Comparisons of Antibiotic Prophylaxes for the Prevention of Dental | -related |
| Bacteremia | 82 |
| Table 30 Conversion of Odds Ratio from Figure 37 to Number Needed to Treat (NNT) | 83 |
| Table 31 Network Meta-Analysis Rankings of Antibiotic Prophylaxes for the Prevention of Dental-related | |
| Bacteremia | |
| Table 32 Direct Comparisons of Topical Antimicrobial Prophylaxes for the Prevention of Dental-related Bac | teremia |
| | 87 |
| Table 33 Indirect (Network) Comparisons of Topical Antimicrobial Prophylaxes for the Prevention of Denta | l-related |
| Bacteremia | |
| Table 34 Indirect (Network) Significant Comparisons of Topical Antimicrobial Prophylaxes for the Prevention | on of |
| Dental-related Bacteremia | |
| Table 35 Conversion of Odds Ratio from Figure 40 to Number Needed to Treat (NNT) | |
| Table 36 Network Meta-Analysis Rankings of Topical Antimicrobial Prophylaxes for the Prevention of Den | |
| related Bacteremia | |
| Table 37 Summary of Oral Health Related Risk Factor (Proportion of studies that reported significant results | |
| Table 38 Oral Health Related Risk Factors for Brushing Bacteremia | |
| Table 39 Oral Health Related Risk Factors for Chewing Bacteremia | |
| Table 40 Oral Health Related Risk Factors for Dental Prophylaxis Bacteremia | |
| Table 41 Oral Health Related Risk Factors for Inter-dental Cleaning Bacteremia | |
| Table 42 Oral Health Related Risk Factors for Intubation Bacteremia | |
| Table 43 Oral Health Related Risk Factors for Oral Surgery Bacteremia | |
| Table 44 Oral Health Related Risk Factors for Periodontic Bacteremia | |
| Table 45 Oral Health Related Risk Factors for Restorative Bacteremia | |
| Table 46 Oral Health Related Risk Factors for Extraction Bacteremia | 103 |

| Table 47 Included Studies for Recommendation 1 | 131 |
|----------------------------------------------------------------------------------------------------|-----------|
| Table 48 Included Studies for Recommendation 2 | 135 |
| Table 49 Included Studies for Recommendation 3 | |
| Table 50 Included Studies for Dental Procedures and Bacteremia | 141 |
| Table 51 Included Studies for Background Microbiology | 149 |
| Table 52 Excluded Studies for Recommendation 1 | 158 |
| Table 53 Excluded Studies for Recommendation 2 | 161 |
| Table 54 Excluded Studies for Recommendation 3 | |
| Table 55 Excluded Studies for Dental Procedures and Bacteremia | 164 |
| Table 56 Excluded Studies for Background Microbiology | 171 |
| Table 57 Excluded Studies Identified During Full Text Review | 199 |
| Table 58 Antibiotic Prophylaxis Network Meta-Analysis Consistency Check | |
| Table 59 Topical Antimicrobial Prophylaxis Network Meta-Analysis Consistency Check | 252 |
| Table 60 Goodness-of-fit Statistics | |
| Table 61 Antibiotic and Topical Antimicrobial Prophylaxis Network Meta-Analysis Consistency Check | 253 |
| Table 62 Bacteremia Incidence Study Details | 255 |
| Table 63 Bacteremia Prevalence Study Details | 257 |
| Table 64 Results of Bacteremia Incidence Random Effects Meta-Analysis | 258 |
| Table 65 Results of Bacteremia Prevalence Random Effects Meta-Analysis | |
| Table 66 Antibiotic Prophylaxis Studies Not Included in Recommendation 1 Network Meta-analysis | |
| Table 67 Topical Antimicrobial Prophylaxis Studies Excluded from Recommendation 2 Network Meta-Ana | lysis 261 |
| Table 68 APPRAISE Table of Prognostic Studies for Recommendation 1, Direct Evidence | |
| Table 69 APPRAISE Table of Treatment Studies for Recommendation 1, Dental Prophylaxis | 263 |
| Table 70 APPRAISE Table of Treatment Studies for Recommendation 1, Intubation | |
| Table 71 APPRAISE Table of Treatment Studies for Recommendation 1, Oral Surgery | 265 |
| Table 72 APPRAISE Table of Treatment Studies for Recommendation 1, Periodontology | |
| Table 73 APPRAISE Table of Treatment Studies for Recommendation 1, Restorative Procedure | |
| Table 74 APPRAISE Table of Treatment Studies for Recommendation 1, Extraction | |
| Table 75 APPRAISE Table of Treatment Studies for Recommendation 2, Brushing | 270 |
| Table 76 APPRAISE Table of Treatment Studies for Recommendation 2, Chewing | 271 |
| Table 77 APPRAISE Table of Treatment Studies for Recommendation 2, Dental Implant | 271 |
| Table 78 APPRAISE Table of Treatment Studies for Recommendation 2, Dental Prophylaxis | 272 |
| Table 79 APPRAISE Table of Treatment Studies for Recommendation 2, Injection | 272 |
| Table 80 APPRAISE Table of Treatment Studies for Recommendation 2, Inter-detal Cleaning | |
| Table 81 APPRAISE Table of Treatment Studies for Recommendation 2, Intubation | |
| Table 82 APPRAISE Table of Treatment Studies for Recommendation 2, Oral Surgery | 274 |
| Table 83 APPRAISE Table of Treatment Studies for Recommendation 2, Orthodontistry | 274 |
| Table 84 APPRAISE Table of Treatment Studies for Recommendation 2, Periodontology | 275 |
| Table 85 APPRAISE Table of Treatment Studies for Recommendation 2, Suture | |
| Table 86 APPRAISE Table of Treatment Studies for Recommendation 2, Tooth Extraction | 276 |
| Table 87 APPRAISE Table of Prognostic Studies for Recommendation 3, Brushing | 279 |
| Table 88 APPRAISE Table of Prognostic Studies for Recommendation 3, Chewing | 280 |
| Table 89 APPRAISE Table of Prognostic Studies for Recommendation 3, Dental Prophylaxis | 281 |
| Table 90 APPRAISE Table of Prognostic Studies for Recommendation 3, Inter-dental Cleaning | 282 |
| Table 91 APPRAISE Table of Prognostic Studies for Recommendation 3, Intubation | 283 |
| Table 92 APPRAISE Table of Prognostic Studies for Recommendation 3, Oral Surgery | 284 |
| Table 93 APPRAISE Table of Prognostic Studies for Recommendation 3, Periodontology | 285 |
| Table 94 APPRAISE Table of Prognostic Studies for Recommendation 3, Restorative Procedure | 286 |
| Table 95 APPRAISE Table of Prognostic Studies for Recommendation 3, Tooth Extraction | |
| Table 96 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Brushing | |
| Table 97 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Brushing | |
| Table 98 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Cleft Palate | 290 |
| Table 99 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Dental Implant | 290 |
| Table 100 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Dental Prophylaxis | 291 |
| Table 101 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Endodontic | |
| Table 102 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline Injections | 293 |

xiii

| Table 103 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Inter-dental Cleaning | 294 |
|--------------------------------------------------------------------------------------------------------------------------------|-----|
| Table 104 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Intubation | 295 |
| Table 105 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Oral Surgery | 296 |
| Table 106 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Orthodontic | |
| Table 107 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Periodontology | 298 |
| Table 108 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Restorative Procedure | 300 |
| Table 109 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Sialography | 301 |
| Table 110 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Suture | 302 |
| Table 111 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Teething | 303 |
| Table 112 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Tooth Extraction | 304 |
| | |
| List of Figures | |
| Figure 1 Overview of the Evidence | 25 |
| Figure 2 Incidence of bacteremia by procedure group | |
| Figure 3 Incidence of bacteremia in single study groups | |
| Figure 4 Prevalence of bacteremia by group | |
| Figure 5 Prevalence of bacteremia in single study groups | |
| Figure 6 Organisms cultured from cohort studies | |
| Figure 7 Organisms cultured from case series studies | |
| Figure 8 Brushing Bacteria (Incidence) | |
| Figure 9 Brushing Bacteria (Prevalence) | |
| Figure 10 Cleft Palate Bacteria (Prevalence) | |
| Figure 11 Dental Implant Bacteria (Incidence) | |
| Figure 12 Dental Prophylaxis Bacteria (Incidence) | |
| Figure 13 Dental Prophylaxis Bacteria (Prevalence) | |
| Figure 14 Endodontic Bacteria (Incidence) | |
| Figure 15 Endodontic Bacteria (Prevalence) | |
| Figure 16 Injection Bacteria (Incidence) | |
| Figure 17 Inter-dental Cleaning Bacteria (Incidence) | |
| Figure 18 Intubation Bacteria (Incidence) | |
| Figure 19 Oral Surgery Bacteria (Incidence) | |
| Figure 20 Oral Surgery Bacteria (Prevalence) | |
| Figure 21 Orthodontic Bacteria (Incidence) | |
| Figure 22 Orthodontic Bacteria (Prevalence) | |
| Figure 23 Periodontic [Scaling & Planing] Bacteria (Incidence) | |
| Figure 24 Periodontic [Gingivectomy] Bacteria (Incidence) | |
| Figure 25 Periodontic [Probing] Bacteria (Incidence) | |
| Figure 26 Periodontic [Scaling & Planing] Bacteria (Prevalence) | |
| Figure 27 Periodontic [Gingivectomy] Bacteria (Prevalence) | |
| Figure 28 Periodontic [Probing] Bacteria (Prevalence) | |
| Figure 29 Sialography Bacteria (Prevalence) | |
| Figure 30 Suture Bacteria (Incidence) | |
| Figure 31 Suture Bacteria (Prevalence) | |
| Figure 32 Teething Bacteria (Prevalence) | |
| Figure 33 Tooth Extraction Bacteria (Incidence) | |
| Figure 34 Tooth Extraction Bacteria (Prevalence) | |
| Figure 35 Network Diagram of Antibiotic Prophylaxis for the Prevention of Dental-related Bacteremia | |
| | |
| Figure 36 Forest Plot of Direct Comparisons of Antibiotics vs. Placebo/No Treatment for the Prevention of I related Bacteremia | |
| Figure 37 Forest Plot of Indirect (Network) Comparisons of Antibiotics vs. Placebo/No Treatment for the Pr | |
| of Dental-related Bacteremia | |
| of Dental-related Bacterenna | |
| Figure 38 Network Diagram of Topical Anumicrobial Prophylaxes for the Prevention of Dental-related Bact | |
| Figure 39 Forest Plot of Direct Comparisons of Topical Antimicrobials vs. No Treatment for the Prevention | |
| Dental-related Bacteremia | |
| DOI:1011 10111100 DUCWICIIII | |

| Figure 40 Forest Plot of Indirect (Network) Comparisons of Topical Antimicrobials vs. No Treatment for the | |
|------------------------------------------------------------------------------------------------------------|----|
| Prevention of Dental-related Bacteremia | 92 |
| | |

INTRODUCTION

OVERVIEW

This clinical practice guideline is based on a systematic review of published studies related to the prevention of orthopaedic implant infection in patients undergoing dental procedures. In addition to providing practice recommendations, this guideline also highlights gaps in the literature and areas that require additional research.

This guideline is intended to be used by all appropriately trained physicians and dentists considering prevention of orthopaedic implant infection in patients undergoing dental procedures.

GOALS AND RATIONALE

The purpose of this clinical practice guideline is to help improve prevention and treatment based on the current best evidence. Current evidence-based practice standards demand that physicians and dentists use the best available evidence in their clinical decision making. To assist them, this clinical practice guideline consists of a systematic review of the available literature related to the prevention of orthopaedic implant infection in patients undergoing dental procedures. The systematic review detailed herein was conducted between October 2010 and July 2011 and demonstrates where there is good evidence, where evidence is lacking, and what topics future research could target to improve the prevention of orthopaedic implant infection in patients undergoing dental procedures. AAOS and ADA staff methodologists and the physician/dentist work group systematically reviewed the available literature and subsequently wrote the following recommendations based on a rigorous, standardized process.

We created this guideline as an educational tool to guide qualified physicians and dentists through a series of treatment decisions in an effort to improve the quality and effectiveness of care. This guideline should not be construed as including all proper methods of care or excluding methods of care reasonably directed to obtaining the same results. The ultimate judgment regarding any specific procedure or treatment must be made in light of all circumstances presented by the patient and the needs and resources particular to the locality or institution.

INTENDED USERS

This guideline is intended to be used by all qualified clinicians considering prevention of orthopaedic implant infection in patients undergoing dental procedures. The guideline is intended to both guide clinical practice and to serve as an information resource for practitioners. An extensive literature base was considered during the development of this guideline. In general, practicing clinicians do not have the resources necessary for such a large project. The AAOS and ADA hope that this guideline will assist practitioners not only in making clinical decisions about their patients, but also in describing, to patients and others, why the chosen treatment represents the best available course of action.

In the interest of collegiality, the ADA elected to follow the rigorous AAOS process for development of this clinical guideline. This guideline is not intended for use as a benefits determination document. Making these determinations involves many factors not considered in the present document, including available resources, business and ethical considerations, and needs.

Evidence for the effectiveness of health-care services is not always present. This is true throughout all areas of medicine and dentistry. Accordingly, all users of this clinical practice guideline are cautioned that an absence of evidence is not evidence of ineffectiveness. An absence means just that; there are no data. It is the AAOS position that rigorously developed clinical practice guidelines should not seek to guide clinical practice when data are absent unless the disease, disorder, or condition in question can result in loss of life or limb. The AAOS incorporates expert opinion into a guideline under these circumstances, and only under these circumstances. Accordingly, when the AAOS states that it cannot recommend for or against a given intervention or service, it is stating that currently available data do not provide clear guidance on which course of action is best, and that it is therefore reluctant to make a recommendation that has potentially national ramifications. The AAOS and ADA believe that when evidence is absent, it is particularly important for treatment decisions to be based on mutual communication between the patient, physician and dentist, with discussion of available treatments and procedures applicable to that patient, and with consideration of the natural history of the disease, costs versus benefits, and current practice patterns. Once the patient has been informed of available therapies and has discussed these options with his/her physician and/or dentist, an informed decision can be made.

PATIENT POPULATION

This document addresses the prevention of orthopaedic implant infection in patients undergoing dental procedures.

BURDEN OF DISEASE AND ETIOLOGY

Approximately 200,000 primary total hip arthroplasties and 400,000 primary total knee arthroplasties were performed in the United States in 2003, with a projected increase to 380,000 hip procedures and over 1,500,000 knee procedures in 2020. Orthopaedic implant infection rates range from 0.3% to 8.3% in the published literature (see Table 26). These infections can be caused by entry of organisms into the wound during surgery, hematogenous spread, recurrence of sepsis in a previously infected joint, or contiguous spread of infection from a local source.

POTENTIAL HARMS, BENEFITS, AND CONTRAINDICATIONS

The goal of prevention of orthopaedic implant infection in patients undergoing dental procedures is avoidance of serious complications resulting from orthopaedic implant infection. Most treatments are associated with some known risks. In addition, contraindications vary widely based on the treatment administered. Therefore, discussion of available treatments applicable to the individual patient rely on mutual communication between the patient, dentist and physician, weighing the potential risks and benefits for that patient.

PREVENTING BIAS IN AN AAOS CLINICAL PRACTICE GUIDELINE

Clinical practice guidelines (CPGs) have come under scrutiny because many of them are not objective. Shaneyfelt and Centor have noted that most current guidelines are not at all like those the Institute of Medicine (IOM) had originally intended, and that they have strayed so far from this original concept that they are mere consensus reports.³ More recently, the IOM has stated that "the quality of CPG development processes and guideline developer adherence to quality standards have remained unsatisfactory and unreliable for decades."⁴ The AAOS understands that only high quality guidelines are credible, and we go to great lengths to ensure the integrity of our guidelines. The purpose of this section is to highlight the processes whereby the AAOS accomplishes this. Additional details about how we combat bias also appear in the Methods section of this guideline.

The AAOS combats bias beginning with the selection of work group members. Applicants for AAOS development work groups who have financial conflicts of interest (COI) related to the guideline topic cannot participate on an AAOS work group if they currently have, or have had a relevant conflict within a year of the start date of guideline development. Applicants also cannot participate if one of their immediate family members has, or has had a relevant conflict of interest.

Financial COI are not the only COI that can influence a guideline. The IOM has noted that income source, long service on government committees or with private insurers, authorship of articles on guideline-related subjects, and biases from personal experience can also cause bias.⁵ This suggests that those with the greatest expertise in any given topic area are also those most likely to introduce bias into guideline development. It also suggests that bias can only be counteracted by processes that are in place throughout the entirety of the development, and not just at the beginning.

One manner whereby the AAOS combats bias throughout guideline development is by having a team that is free of all of the above-mentioned COI conduct the literature searches, evaluate the quality of the literature, and sythesize the data (see Appendix I for a list of the work group members and methodologists who participated in the development of this guideline). Hirsh and Guyatt have suggested that using such conflict-free methodologists is critical to developing an unbiased guideline.⁶

Our use of methodologists changes the traditional role of clinicians in guideline development. The clinicians on an AAOS guideline work group serve as content experts. One of the clinicians' tasks is to frame the scope of the guideline by developing preliminary recommendations (these are the questions that will be addressed by the guideline; see below for further information). Another is to develop the article inclusion criteria. After they have done so, the AAOS medical librarian obtains key words from work group members and uses words, the preliminary recommendations, and inclusion criteria to construct literature search strategies. Clinicians are not permitted to suggest specific articles for inclusion at this time inasmuch as those suggestions are often about articles they have authored or that support a particular point of view.

Methodologists then determine which articles should be recalled and whether a recalled article meets the inclusion criteria. After completing this task, the clinician work group is given a list of the recalled articles that are proposed for inclusion and a list of the recalled studies proposed for exclusion. The work group then reviews these lists and suggests modifications. The purpose of this step is to assure the integrity of the guideline's data set. The methodologists are not obligated to take the work group's suggestions, but they are obligated to explain why they did not. Articles included or excluded as a result of this clinician review are handled as all other included articles or excluded studies. The methodologists also appraise the quality and applicability of each included study (we use "quality" as synonymous with "risk of bias." The latter term is preferred by others but, since quality and risk of bias are measured exactly the same way, the difference between the two seems largely semantic. Similarly, we use the terms "applicability" and "generalizability" as synonyms.)

Quality appraisal is a subject worth special mention because it is a necessary step in performing a systematic review and in developing a clinical practice guideline. One evaluates the quality (or risk of bias) of a study to determine how "believable" its results are, the results of high quality studies are more believable than those of low quality studies. This is why, all other things being equal, a recommendation based on high quality evidence will receive a higher grade than recommendations based on lower quality evidence (see Grades of Recommendation for more information). Biases in quality evaluation can cause overestimates of the confidence one should have in available data, and in a guideline recommendation.

Bias in quality evaluation arises when members of a work group view the papers they authored as being more believable than similar research performed by others, view certain studies as more believable simply because they were conducted by thought leaders in a given medical specialty area, and/or view research results that they are "comfortable" with to be more believable than results with which they are uncomfortable.

The problem of biased quality evaluations is aggravated by the fact that no method for quality/risk of bias assessment has been empirically validated. Ultimately, therefore, all methods of quality/risk of bias assessment, are based on expert opinion (including those based on expert consensus obtained through formal methods like the Delphi method), and they all require judgments that are arbitrary. The method we use is no exception.

Given that all currently available quality evaluation systems are imperfect their susceptibility to bias must be a deciding factor about whether to use them in clinical practice guideline development. The AAOS methodology is guided by the thinking that, if guideline developers have the choice between several methodologically imperfect systems, the least biased system is the best. The burden that falls to readers of clinical practice guidelines is to determine which systems are not. Making this determination requires readers to examine two aspects of quality evaluation; the individual criteria used to evaluate a study, and how those criteria are translated into a final determination of a study's believability.

The criteria used to evaluate a study are often framed as one or more questions about a study's design and/or conduct. At the AAOS, independent methodologists answer these questions. This combats bias by virtually eliminating the intellectual conflicts of interest that can arise when others are providing the answers.

Also preventing bias is the way the quality questions are phrased, and the fact that there are specific criteria (described in almost 300 pages of documentation) for answering each question. The simplest example, the AAOS question "Was there >80% follow-up" illustrates the point. The question is answered "Yes,", "No", or "Unclear." To determine whether a "Yes" or "No" answer is unclear, the methodologist merely looks at the number of patients present at the follow-up time of interest, the number of patients present at the start of the study, and expresses the former as a percentage of the latter. If the article does not report the information required to compute this percentage (or does not directly report the percentage) an "Unclear" answer is supplied. In answering this or any other question in the AAOS quality assessment scheme, the methodologist is merely checking to see if the article provides specific data or makes specific statements. If it does, a "Yes" or "No" answer is supplied. If it does not, an "Unclear" answer is given. This lack of ambiguity in the criteria required to answer each question makes answering each question an almost completely objective exercise.

This stands in sharp contrast to the use of Levels of Evidence systems (also called evidence hierarchies), which are probably the most commonly used way of evaluating study quality in clinical practice guideline development. The vagueness of these systems opens the opportunity for bias. For example, these systems often hold that Level I evidence (i.e., the highest quality evidence) is from a well-designed randomized controlled trial, without ever specifying what "well-designed" means. This lack of specific instructions creates the possibility for bias in grading articles because it allows for an *ad hoc* appraisal of study quality. Furthermore, there are over 50 such systems, individuals do not consistently apply any given system in the same way, many are not sensible to methodologists, and Level I studies, those of the highest level of evidence, do not necessarily report that they used adequate safeguards to prevent bias.

Obviously, simply answering a series of questions about a study does not complete the quality evaluation. All clinical practice guideline developers then use that information to arrive at a final characterization of a study's quality. This can be accomplished in two (and only two) ways, by allowing those who are performing this final characterization to use their judgment, or by not letting them do so. Bias is possible when judgment is allowed. Bias is mitigated in the AAOS system because the final rating is accomplished entirely by a computer that uses a predetermined algorithm.

This aspect of the AAOS system contrasts with the GRADE system, which places the final determination about whether a study has "no", "serious" or "very serious" limitations in the hands of the reviewer. Furthermore, the GRADE system allows the investigator to specify "other sources of bias" (i.e. sources of bias that were not specified *a priori*) and, although this is a theoretically sound way to approach quality evaluation, in practice it too, could allow for *ad hoc* criticisms of a study, and to criticisms that are not evenly applied across all studies. We recognize that we may miss some uncommon study flaws in our evaluation. While this means that our quality evaluation system is not perfectly comprehensive, it does not mean that it is biased. This is yet another example of how the AAOS, faced with a choice among imperfect quality/risk of bias systems, chooses the least biased approach. Given the above mentioned history of guideline development, the AAOS emphasis on elimination of bias seems prudent.

The AAOS system, unlike the GRADE system, also specifically addresses the issue of statistical power (i.e., number of patients enrolled) of a trial. Low statistical power is a common problem in

the medical literature, ¹⁰ and low power studies can lead reviewers to incorrectly conclude that a statistically non-significant result means that a given treatment does not work or, perhaps more serious, to reach positive conclusions about an intervention based on the putative "trends" reported in such studies. We regard low power studies as uninformative, and do not consider them when formulating a final recommendation. (We do, however, include low power studies in meta-analyses, inasmuch as one purpose of a meta-analysis is to overcome the low power of individual studies.)

Like the GRADE system, the AAOS guidelines will include observational studies. However, we do not always do so. Rather, we perform "best evidence" syntheses in AAOS guidelines in which we examine the best available (as opposed to the best possible) evidence. We use the best evidence because it is more "believable" than other evidence. The results of studies that are more believable should not be modified by results that are less believable.

When an AAOS guideline includes uncontrolled studies (e.g., case series) it only includes prospective case series that meet a number of other quality-related criteria. We do not include retrospective case series under any circumstances. Such studies do not establish empirically testable comparisons or relationships a priori, are not based on systematic assignment of patients to treatment groups, and are not designed to fully control measurement bias. There is no specific prohibition against using such studies in the GRADE system. We suggest that all guideline developers who are attempting to produce unbiased guidelines employ similar *a priori* criteria to specify the point at which they consider evidence to be too unreliable to consider.

Also unlike the GRADE system, the AAOS guidelines make provisions for making recommendations based on expert opinion. This recognizes the reality of medicine, wherein certain necessary and routine services (e.g., a history and physical) should be provided even though they are backed by little or no experimental evidence, and wherein certain diseases, disorders, or conditions are so grave that issuing a recommendation in the absence of evidence is more beneficial to patients than not issuing one. To prevent the bias that can result when recommendations based on expert opinion proliferate, we have specific rules for when opinion-based recommendations can be issued (further discussed below) and, perhaps more important, for when they cannot be issued. The AAOS will only issue an opinion-based recommendation when the service in question has virtually no associated harms and is of low cost (e.g., a history and physical) or when the consequences of doing (or not doing) something are so catastrophic that they will result in loss of life or limb.

Clinical practice guidelines have not met quality standards for a long time. In recognition of this, the IOM has developed two checklists, one for systematic reviews¹¹ and another for clinical practice guidelines.⁴ Meeting the items on these checklists should assure readers of a guideline that it is unbiased. Table 1 and Table 2 show the performance of the present AAOS guideline on these standards.

Table 1 IOM Clinical Practice Guidelines Standards

| IOM Guidelines Standard | AAOS Guideline Meets Standard ? |
|------------------------------------------------------------------------------|---------------------------------------------------|
| Establishing transparency | Yes |
| Management of Conflict of Interest | Yes |
| Guideline development group composition | No – AAOS does not involve patient representative |
| Clinical practice guideline – systematic review intersection | Yes |
| Establishing evidence foundations for and rating strength of recommendations | Yes |
| Articulation of recommendations | Yes |
| External review | Yes |
| Updating | Yes |

Table 2 IOM Systematic Review Standards

| IOM Systematic Review Standard | AAOS Systematic Reviews Meet Standard ? |
|------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------|
| Establish a team with appropriate expertise and experience to conduct the systematic review | Yes |
| Manage bias and conflict of interest (COI) of the team conducting the systematic review | Yes |
| Ensure user and stakeholder input as the review is designed and conducted | Yes |
| Manage bias and COI for individuals providing input into the systematic review | Yes |
| Formulate the topic for the systematic review | Yes |
| Develop a systematic review protocol | Yes |
| Submit the protocol for peer review | No – do not have peer review of protocol |
| Make the final protocol publicly available, and add any amendments to the protocol in a timely fashion | Yes |
| Conduct a comprehensive systematic search for evidence | Yes |
| Take action to address potentially biased reporting of research results | No – do not search for unpublished information |
| Screen and select studies | Yes |
| Document the search | Yes |
| Manage data collection | Yes |
| Critically appraise each study | Yes |
| Use a prespecified method to evaluate the body of evidence | Yes |
| Conduct a qualitative synthesis | Yes |
| Decide if, in addition to a qualitative analysis, the systematic review will include a quantitative analysis (meta-analysis) | Yes |
| If conducting a meta-analysis, then do the following: | Yes |
| Prepare final report using a structured format | Partially - no lay public summary |
| Peer review the draft report | Partially - do not use independent third party to manage peer review process |
| Publish the final report in a manner that ensures free public access | Yes |

METHODS

To develop this guideline, the AAOS-ADA work group held an introductory meeting on November 20 and 21, 2010 to establish the scope of the guideline and the systematic reviews. Upon completing the systematic reviews, the work group participated in a two-day recommendation meeting on October 15 and 16, 2011 at which time the final recommendations and rationales were edited, written, and voted on.

FORMULATING PRELIMINARY RECOMMENDATIONS

The work group determined the scope of the guideline by constructing a set of preliminary recommendations. These recommendations specify [what] should be done in [whom], [when], [where], and [how often or how long]. This is similar to the PICO (patients, interventions, comparisons, and outcomes) format used when the scope of a guideline is framed using key questions instead of preliminary recommendations. The preliminary recommendations function as questions for the systematic reviews that underpin each preliminary recommendation, not as final recommendations or conclusions. To avoid "wordsmithing" discussions at the initial work group meeting, the preliminary recommendations are always worded as recommending for something. Appendix II describes the formulation of preliminary recommendations in further detail.

Once established, these preliminary recommendations cannot be modified until the final work group meeting. At this time, they can only be modified in accordance with the available evidence and only in accordance with the AAOS rules for how the wording of a recommendation depends on the grade of recommendation (see below for information about this wording). No modifications of the preliminary recommendations can require new literature searches and, at the final work group meeting, no recommendations can be added that require the use of expert opinion.

FULL DISCLOSURE INFORMATION

All of the work group's preliminary recommendations are represented in this guideline. This ensures full disclosure of the information that the AAOS-ADA work group examined, and assures readers that they are seeing *all* the information, and not just a selected portion of it.

STUDY SELECTION CRITERIA

We developed *a priori* article inclusion criteria for the systematic reviews for each preliminary recommendation. These criteria are our "rules of evidence." Articles that did not meet them are, for the purposes of this guideline, not evidence.

To be included in our systematic reviews (and hence, in this guideline) an article had to be a report of a study that:

- Study must be of patient population of interest (as described by preliminary recommendations)
- Study must report on >50% of the patient population of interest if results are combined
- Article must be a full article report of a clinical study
- Study must appear in a peer-reviewed publication
- Study must be published in English

- Study must be of humans
- Study must not be an in vitro study
- Study must not be a biomechanical study
- Study must not have been performed on cadavers
- Study must be published in or after 1960
- Study results must be quantitatively presented
- Retrospective case series, medical records review, meeting abstracts, historical articles, editorials, letters, and commentaries are excluded
- Registry data is included
- Case series studies that give patients the treatment of interest AND another treatment are excluded
- Case series studies that have non-consecutive enrollment of patients are excluded
- Study should have 10 or more patients per group
- Composite measures or outcomes, even if they are patient-oriented, are excluded

The restriction on English language papers is unlikely to influence the recommendations in the present clinical practice guideline. An umbrella review of systematic reviews on language restriction found that none of the systematic reviews provided empirical evidence that excluding non-English language studies resulted in biased estimates of an intervention's effectiveness. 12

We did not include systematic reviews or meta-analyses conducted by others, or guidelines developed by others. These documents are developed using different inclusion criteria than those specified by the AAOS-ADA work group. Therefore, they may include studies that do not meet our inclusion criteria. We recalled these documents if their abstract suggested that they might address one of our recommendations, and we searched their bibliographies for additional studies.

LITERATURE SEARCHES

We searched for articles published from January 1966 to July 25, 2011. We searched three electronic databases; PubMed, EMBASE, and The Cochrane Central Register of Controlled Trials. Strategies for searching electronic databases were constructed by the AAOS Medical Librarian using previously published search strategies to identify relevant studies. ¹³⁻¹⁸

We supplemented searches of electronic databases with manual screening of the bibliographies of all retrieved publications. We also searched the bibliographies of recent systematic reviews and other review articles for potentially relevant citations. All articles identified were subject to the study selection criteria listed above. As noted above, the guideline work group also examined lists of included and excluded studies for errors and omissions.

We went to these lengths to obtain a complete set of relevant articles. Having a complete set ensures that our guideline is not based on a biased subset of articles. The study attrition diagram in Appendix III provides details about the inclusion and exclusion of the studies considered for this guideline. The search strategies used to identify these studies are provided in Appendix IV.

BEST EVIDENCE SYNTHESIS

We included only the best available evidence for any given outcome addressing a recommendation. Accordingly, we first included the highest quality evidence for any given

outcome if it was available. In the absence of two or more studies that reported an outcome at this quality, we considered studies of the next lowest quality until at least two or more occurrences of an outcome had been acquired. For example, if there were two "Moderate" quality studies that reported an outcome, we did not include "Low" quality studies that also reported this outcome, but if there was only one "Moderate" quality study that reported an outcome, we also included "Low" quality studies.

APPRAISING EVIDENCE QUALITY AND APPLICABILITY STUDIES OF INTERVENTIONS

QUALITY

As noted earlier, we judged quality using questions specified before this guideline topic was selected, and a computer program determined the final quality rating. Accordingly, it is highly unlikely that bias affected our determinations of quality.

We separately evaluated the quality of evidence for each outcome reported by each study. This follows the suggestion of the GRADE working group and others. We evaluated quality using a domain-based approach. Such an approach is used by the Cochrane Collaboration. Unlike the Cochrane Collaboration's scheme, our scheme allows for evaluation of studies of all designs. The domains we used are whether:

- The study was prospective (with prospective studies, it is possible to have an *a priori* hypothesis to test; this is not possible with retrospective studies.)
- The study was of low statistical power
- The assignment of patients to groups was unbiased
- There was blinding to mitigate against a placebo effect
- The patient groups were comparable at the beginning of the study
- The intervention was delivered in such a way that any observed effects could reasonably be attributed to that intervention
- Whether the instruments used to measure outcomes were valid
- Whether there was evidence of investigator bias

Each quality domain is addressed by one or more questions that are answered "Yes," "No," or "Unclear." These questions and the domains that each address are shown in Appendix V.

To arrive at the quality of the evidence for a given outcome, all domains except the "Statistical Power" domain are termed as "flawed" if one or more questions addressing any given domain are answered "No" for a given outcome, or if there are two or more "Unclear" answers to the questions addressing that domain. The "Statistical Power" domain is considered flawed if a given study did not enroll enough patients to detect a standardized difference between means of 0.2.

Domain flaws lead to corresponding reductions in the quality of the evidence. The manner in which we conducted these reductions is shown in Table 3. For example, the evidence reported in a randomized controlled trial (RCT) for any given outcome is rated as "High" quality if zero or one domain is flawed. If two or three domains are flawed for the evidence addressing this outcome, the quality of evidence is reduced to "Moderate," and if four or five domains are

flawed, the quality of evidence is reduced to "Low." The quality of evidence is reduced to "Very Low" if six or more domains are flawed.

Some flaws are so serious that we automatically term the evidence as being of "Very Low" quality, regardless of a study's domain scores. These serious design flaws are:

- Non-consecutive enrollment of patients in a case series
- Case series that gave patients the treatment of interest AND another treatment
- Measuring the outcome of interest one way in some patients and measuring it in another way in other patients
- Low statistical power

Table 3 Relationship between Quality and Domain Scores for Interventions

| Number of Flawed Domains | Quality |
|---------------------------------|----------|
| 0-1 | High |
| 2-3 | Moderate |
| 4-5 | Low |
| >5 | Very Low |

Although we mention levels of evidence in this guideline, we do so only to provide some very general information about study quality to those readers familiar with the levels of evidence system of *The Journal of Bone and Joint Surgery - American*. However, for the reasons noted above, we do not use levels of evidence as when we speak of "quality" in this document, and levels of evidence play no role in our determination of the grade of the final recommendations.

APPLICABILITY

We rated the applicability (also called "generalizability" or "external validity") of the evidence for each outcome reported by each study. As with quality, applicability ratings were determined by a computer program that used predetermined questions about specific applicability domains. We rated applicability as either "High", "Moderate", or "Low" depending on how many domains are flawed. As with quality, a domain is "flawed" if one or more questions addressing that domain is answered "No" or if two or more are answered "Unclear." We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a given outcome, or if there are two or more "Unclear" answers to the questions addressing that domain (see Appendix V for the specific applicability questions we employed and the domains that each question addresses).

Our questions and domains about applicability are those of the PRECIS instrument.²¹ The instrument was originally designed to evaluate the applicability of randomized controlled trials, but it can also be used for studies of other design. The questions in this instrument fall into four domains. These domains and their corresponding questions are shown in Appendix V. As shown in Table 4, the applicability of a study is rated as "High" if it has no flawed domains, as "Low" if all domains are flawed, and as "Moderate" in all other cases.

Table 4 Relationship between Applicability and Domain Scores for Interventions

| Number of Flawed Domains | Applicability |
|--------------------------|---------------|
| 0 | High |
| 1, 2, 3 | Moderate |
| 4 | Low |

STUDIES OF INCIDENCE AND PREVALENCE OUALITY

As with our appraisal of the quality of studies of intervention, our appraisal of studies of incidence and prevalence is a domain-based approach conducted using *a priori* questions (please see Appendix V for the questions we used and the domains to which they apply), and scored by a computer program. The four domains we employed are listed below:

- Outcome (whether the study is measuring the incidence/prevalence of a clinically meaningful event)
- Measurement (whether the study measured the disease/disorder/condition in a way that would lead to accurate estimates of incidence or prevalence)
- Participants (whether those who were studied were representative of the population of interest)
- Investigator Bias (whether author biases could have prejudiced the results)

We characterized a study that has no flaws in any of its domains as being of "High" quality, a study that has one flawed domain as being of "Moderate" quality, a study with two flawed domains as being of "Low" quality, and a study with three or more flawed domains as being of "Very Low" quality (Table 5). We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a given screening/diagnostic/test, or if there are two or more "Unclear" answers to the questions addressing that domain.

We considered some design flaws as so serious that their presence automatically guarantees that a study is characterized as being of "Very Low" quality regardless of its domain scores. These flaws are:

- The outcome of interest could have occurred more than once in a person during the course of the study, and more than the first episode of the outcome was counted in the incidence/prevalence estimate
- The study was a study of the proportion (or number) of people who have a disease, and the study was not a study of point prevalence.

Table 5 Relationship between Quality and Domain Scores for Incidence and Prevalence Studies

| Number of Flawed Domains | Quality |
|---------------------------------|----------|
| 0 | High |
| 1 | Moderate |
| 2 | Low |
| ≥3 | Very Low |

APPLICABILITY

We separately evaluated the applicability of prevalence and incidence studies, and did so using a domain-based approach (please see Appendix V for the relevant questions and the domains they address) that involves predetermined questions and computer scoring. The domains we used for the applicability of prognostics are:

- Participants (i.e. whether the participants in the study were like those seen in the population of interest)
- Analysis (i.e., whether participants were appropriately included and excluded from the analysis)
- Outcome (i.e., whether the incidence/prevalence estimates being made were of a clinically meaningful outcome)

We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a given screening/diagnostic/test, or if there are two or more "Unclear" answers to the questions addressing that domain. We characterized the applicability of a screening/diagnostic test as "High" if none of its domains are flawed, "Low" if all of its domains are flawed, and "Moderate" in all other cases (Table 6).

Table 6 Relationship between Applicability and Domain Scores for Incidence and Prevalence Studies

| Number of Flawed Domains | Applicability |
|--------------------------|---------------|
| 0 | High |
| 1,2 | Moderate |
| 3 | Low |

STUDIES OF PROGNOSTICS *QUALITY*

Our appraisal of studies of prognostics is a domain-based approach conducted using *a priori* questions, and scored by a computer program (please see Appendix V for the questions we used and the domains to which they apply). The six domains we employed are:

- Prospective (A variable is specified as a potential prognostic variable *a priori*. This is not possible with retrospective studies.)
- Power (Whether the study had sufficient statistical power to detect a prognostic variable as statistically significant)
- Analysis (Whether the statistical analyses used to determine that a variable was rigorous to provide sound results)
- Model (Whether the final statistical model used to evaluate a prognostic variable accounted for enough variance to be statistically significant)
- Whether there was evidence of investigator bias

We separately determined a quality score for each prognostic reported by a study. We characterized the evidence relevant to that prognostic variable as being of "High" quality if there are no flaws in any of the relevant domains, as being of "Moderate" quality if one of the relevant

domains is flawed, as "Low" quality if there are two flawed domains, and as "Very Low" quality if three or more relevant domains are flawed (Table 7). We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a given prognostic variable, or if there are two or more "Unclear" answers to the questions addressing that domain.

Table 7 Relationship between Quality and Domain Scores for Prognostic Studies

| Number of Flawed Domains | Quality |
|--------------------------|----------|
| 0 | High |
| 1 | Moderate |
| 2 | Low |
| ≥3 | Very Low |

APPLICABILITY

We separately evaluated the applicability of each prognostic variable reported in a study, and did so using a domain-based approach (please see Appendix V for the relevant questions and the domains they address) that involves predetermined questions and computer scoring. The domains we used for the applicability of prognostics are:

- Patients (i.e. whether the patients in the study and in the analysis were like those seen in clinical practice)
- Analysis (i.e., whether the analysis was not conducted in a way that was likely to describe variation among patients that might be unique to the dataset the authors used)
- Outcome (i.e., whether the prognostic was a predictor of a clinically meaningful outcome)

We characterized the evidence relevant to that prognostic as being of "High" applicability if there are no flaws in any of the relevant domains, as being of "Low" applicability if all three domains are flawed, and as of "Moderate" applicability in all other cases (Table 8). We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a given prognostic variable, or if there are two or more "Unclear" answers to the questions addressing that domain.

Table 8 Relationship between Applicability and Domain Scores for Prognostic Studies

| Applicability |
|---------------|
| High |
| Moderate |
| Low |
| |

OTHER BIASES IN THE PUBLISHED LITERATURE

Despite our efforts to rigorously evaluate the quality of the studies we included, there remains the possibility that some of the articles considered in this guideline are biased. A 2007 umbrella review found that 20 of 23 previous systematic reviews found a positive relationship between pharmaceutical industry support and pro-industry findings, ²² leading the author to conclude that "it is unequivocally the case that sponsorship influences published results." The relationship also seems to exist in orthopaedics, where authors of industry-funded studies of hip and knee

arthroplasty come to positive conclusions more often that authors of studies not funded by industry, ²³ and where the association between trial outcome and funding source exists across subspecialty societies. ²⁴

These apparent biases may not be related to the article's quality²² and, therefore, may not be detected by our evaluations or the quality/risk of bias evaluations performed by others. Accordingly, we follow the suggestion of Montori, et al.²⁵ and do not use the conclusions of the authors of any article. Rather, we use only the information provided in an article's Methods section and in its Results section. Furthermore, we perform our analysis using network meta-analysis, an analytical technique that considers the full range of alternatives rather than just those comparisons selected by industry.²⁶

GRADES OF RECOMMENDATION

A grade of recommendation expresses the degree of confidence one can have in each of the final recommendations. Grades express how likely it is that a recommendation will be overturned by future evidence, and are termed "Strong," "Moderate," or "Limited."

We used the above-discussed quality and applicability ratings in conjunction with consistency, whether the studies reported outcomes that the work group deemed "critical," and the potential for catastrophic harm to determine the final grade of recommendation. More specifically, we began by setting the grade as equal to the quality of the available evidence. In other words, high quality evidence is preliminarily taken as a "Strong" grade, moderate quality as a "Moderate" grade, and low quality as a "Limited" grade. As noted above, very low quality evidence is not included in AAOS guidelines. Accordingly, the final versions of preliminary recommendations that are based on such evidence will either state that the AAOS cannot recommend for or against a given medical service or, assuming that the requirements for a recommendation based on expert opinion are met it will be a consensus-based recommendation. We then adjusted the grade down one step if the evidence is of "Low" applicability, is inconsistent (defined as studies that report qualitatively different effects, a heterogeneous meta-analysis, or a network meta-analysis with statistically significant inconsistency), if there is only one study that addresses a given recommendation, or if a majority of the outcomes deemed "critical" are not reported in the literature. Preliminary grades were adjusted upwards if the evidence is of "High" applicability or if providing the intervention decreases the potential for catastrophic harm (loss of life or limb). Preliminary grades were adjusted downward if the evidence is of "Low" applicability or if the medical service in question is accompanied with catastrophic harm.

For a recommendation of a "Strong" grade, a minimum of two high quality studies are needed. A minimum of two moderate quality studies are required for a "Moderate" grade, and a minimum of two low quality studies are needed for a "Limited" grade. Recommendations addressed by only very low quality studies are consensus-based.

Table 9 Strength of Recommendation Descriptions

| Statement Rating | Description of Evidence Strength | Implication for Practice |
|---------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Strong | Evidence is based on two or more "High" strength studies with consistent findings for recommending for or against the intervention. A Strong recommendation means that the benefits of the recommended approach clearly exceed the potential harm (or that the potential harm clearly exceeds the benefits in the case of a strong negative recommendation), and that the strength of the supporting evidence is high. | Practitioners should follow a Strong recommendation unless a clear and compelling rationale for an alternative approach is present. |
| Moderate | Evidence from two or more "Moderate" strength studies with consistent findings, or evidence from a single "High" quality study for recommending for or against the intervention. | Practitioners should generally follow a Moderate recommendation but remain alert to new information and be sensitive to patient preferences. |
| | A Moderate recommendation means that the benefits exceed the potential harm (or that the potential harm clearly exceeds the benefits in the case of a negative recommendation), but the strength of the supporting evidence is not as strong. | |
| Limited | Evidence from two or more "Low" strength studies with consistent findings, or evidence from a single Moderate quality study recommending for or against the intervention or diagnostic. A Limited recommendation means the quality of the supporting evidence that exists is unconvincing, or that well-conducted studies show little clear advantage to one approach versus another. | Practitioners should be cautious in deciding whether to follow a recommendation classified as Limited , and should exercise judgment and be alert to emerging publications that report evidence. Patient preference should have a substantial influencing role. |

| Inconclusive | Evidence from a single low quality study | Practitioners should feel little |
|------------------------|---------------------------------------------|-----------------------------------------|
| | or conflicting findings that do not allow a | constraint in deciding whether to |
| | recommendation for or against the | follow a recommendation labeled as |
| | intervention. | Inconclusive and should exercise |
| | | judgment and be alert to future |
| | An Inconclusive recommendation means | publications that clarify existing |
| | that there is a lack of compelling | evidence for determining balance of |
| | evidence resulting in an unclear balance | benefits versus potential harm. |
| | between benefits and potential harm. | Patient preference should have a |
| | _ | substantial influencing role. |
| Consensus ¹ | The supporting evidence is lacking and | Practitioners should be flexible in |
| | requires the work group to make a | deciding whether to follow a |
| | recommendation based on expert opinion | recommendation classified as |
| | by considering the known potential harm | Consensus, although they may set |
| | and benefits associated with the | boundaries on alternatives. Patient |
| | treatment. | preference should have a substantial |
| | | influencing role. |
| | A Consensus recommendation means | |
| | that expert opinion supports the guideline | |
| | recommendation even though there is no | |
| | available empirical evidence that meets | |
| | the inclusion criteria. | |
| | | |

The AAOS will issue a consensus-based recommendation only when the service in question has virtually no associated harm and is of low cost (e.g. a history and physical) or when not establishing a recommendation could have catastrophic consequences.

WORDING OF THE FINAL RECOMMENDATIONS

To prevent biased nuances in the way recommendations are worded, the AAOS uses predetermined, specific language for its recommendations. The exact wording is governed by the final grade of the recommendation. This wording, and the corresponding grade, is shown in Table 10.

Table 10 AAOS Guideline Language

| Guideline Language | Grade of Recommendation |
|--------------------------------------------------------------------------------|-------------------------|
| We recommend | Strong |
| We suggest | Moderate |
| The Practitioner <i>might</i> | Limited |
| We are unable to recommend for or against | Inconclusive |
| In the absence of reliable evidence, the opinion of this work group is* | Consensus* |

¹ The AAOS will issue a consensus-based recommendation only when the service in question has virtually no associated harm and is of low cost (e.g. a history and physical) or when not establishing a recommendation could have catastrophic consequences.

*Consensus based recommendations are made only if specific criteria are met (see below).

CONSENSUS RECOMMENDATIONS

Consensus recommendations are recommendations based on expert opinion. As noted above, there are times when it is prudent to make such recommendations. However, liberal use of them can allow for bias. Accordingly, we allow consensus-based recommendations using the procedures described by the United States Preventative Services Task Force (USPSTF). In effect, this means that the AAOS will only issue a consensus-based recommendation under two circumstances. The first is for procedures that have virtually no associated harms, are of relatively low cost, and that reflect current, routine clinical practice. The second is when providing (or not providing) a service could result in loss of life or limb. Because they are based on expert opinion, consensus recommendations are the weakest type of recommendation.

In making such recommendations, the AAOS instructs its clinician work group members to address:

- The potential preventable burden of disease (if the burden is low, a consensus-based recommendation cannot be issued)
- Potential harms (if there are serious harms that result from providing a medical service, a consensus-based recommendation cannot be issued)
- Current practice (a consensus-based recommendation cannot be issued if a service is not currently widely used)
- Why, if warranted, a more costly service is being recommended over a less costly one

The AAOS employs additional rules to combat the bias that may affect such recommendations. The rationale for the recommendation cannot contain references to studies that were not included in the systematic reviews that underpin a guideline. Excluded articles are, in effect, not evidence, and they may not be cited. Also, the final recommendation must use the language shown in Table 10. The rationale cannot contain the language "we recommend," "we suggest," or "the practitioner might" inasmuch as this wording could be confused with the evidence-based recommendations in a guideline. In addition, the rationale must address apparent discrepancies in logic with other recommendations in the guideline. For example, if a guideline does not come to a recommendation in some instances but, in the instance in question, the work group has issued a consensus-based recommendation, the rationale must explain the reason for this difference.

One consequence of these restrictions is that the AAOS does not typically recommend new medical devices, drugs, or procedures. These procedures are usually supported by little research, and the AAOS is reluctant to make recommendations that could have a national impact based on small amounts of data.

When it is not possible to issue a recommendation (i.e. when the recommendation reads that "we are unable to recommend for or against") the explanation for why a recommendation cannot be given cannot contain an implied recommendation. For example, in the case of a new device, drug, or procedure, the work group may not write a recommendation similar to "Although the treatment *appears to be promising*, there is currently insufficient evidence to recommend for or against its use." The italicized phrase implies that the treatment is effective, whereas not being

able to recommend "for or against" something implies that effectiveness is currently indeterminate.

More details of our rules for making opinion based recommendation can be found in Appendix VI

VOTING ON THE RECOMMENDATIONS

The recommendations and their strength were voted on using a structured voting technique known as the nominal group technique. We present details of this technique in Appendix VII. Voting on guideline recommendations is conducted using a secret ballot and work group members are blinded to the responses of other members. If disagreement between work group members is significant, there is further discussion to see whether the disagreement(s) can be resolved. Up to three rounds of voting are held to attempt to resolve disagreements. If disagreements are not resolved following three voting rounds, no recommendation is adopted. Lack of agreement is a reason that the grade of some recommendations can be labeled "Inconclusive."

Formal votes on all recommendations that are evidence-based or that read "we are unable to recommend for or against" are only on the recommendations. The rationales require only approval of the work group chair and the methodologists unless the recommendation is consensus-based. Both the recommendation and the rationale of a consensus-based recommendation are the subject of formal votes.

OUTCOMES CONSIDERED

In considering the outcomes discussed in this guideline, it is important to distinguish between patient-oriented and surrogate outcomes. Patient-oriented outcomes measure how a patient feels, functions, or survives.²⁹ A patient-oriented outcome "tells clinicians, directly and without the need for extrapolation, that a diagnostic, therapeutic or preventive procedure helps patients live longer or live better."³⁰ Patient-oriented outcomes include pain relief, death, and fractures. Surrogate outcomes are laboratory measurements or physical signs used as substitutes for patient-oriented outcomes. Surrogate outcomes include outcomes like blood cholesterol levels, laboratory and imaging results, and bone mineral densities.

Surrogate outcomes are problematic. An intervention that improves a surrogate outcome does not necessarily improve a patient-oriented outcome. The opposite can be true. Using a surrogate outcome as a study endpoint can make a harmful treatment look beneficial. For example, although the surrogate outcome cardiac sinus rhythm improves when quinidine is given after conversion, mortality is tripled. Similarly, sodium fluoride increases bone mineral density, but it also increases the rate of non-vertebral fractures. This leads to an important (and often overlooked) aspect about surrogate outcomes. To be useful, a surrogate outcome must not only correlate with the patient-oriented outcome of interest, but also the surrogate must predict (capture) the effects of an intervention on that outcome. Additionally, many surrogates may correlate with an outcome, but few predict the effects of an intervention. For these reasons, the AAOS rarely uses surrogate outcomes as endpoints in its clinical practice guidelines. We make an exception, in this guideline, for bacteremia associated with a dental procedure because there is little reliable evidence predicting the effects of bacteremia associated with a dental procedure on orthopaedic implant infections.

STATISTICAL METHODS

When possible, we recalculate the results reported in individual studies and compile them to answer the recommendations. The statistical analysis is conducted using STATA 10.0³³. STATA was used to determine the magnitude, direction, and/or 95% confidence intervals of the treatment effect. For data reported as means (and associated measures of dispersion) the mean difference between groups and the 95% confidence interval was calculated and a two-tailed t-test of independent groups was used to determine statistical significance. When published studies report measures of dispersion other than the standard deviation the value was estimated to facilitate calculation of the treatment effect. In studies that report standard errors or confidence intervals the standard deviation was back-calculated. In studies that only report the median, range, and/or size of the trial, we estimated the means and variances according to a published method.³⁴ In some circumstances statistical testing was conducted by the authors and measures of dispersion were not reported. In the absence of measures of dispersion, the results of the statistical analyses conducted by the authors (i.e. the p-value) are considered as evidence. For proportions, we report the ratio of events along with the percentage. P-values < 0.05 were considered statistically significant.

We performed network meta-analyses (also known as a mixed treatment comparisons analyses) to ascertain the comparative effectiveness of strategies for preventing bacteremia among patients undergoing dental extraction. All of the trials entered into our analyses were randomized controlled trials (most, but not all, were of "Moderate" quality; additional details on their quality are presented in the sections of this guideline that present our results of the appraisal of these studies).

Analyses were performed as described by Lu and Ades³⁵ using Winbugs v 1.4.3. This method preserves the randomization of the original trials. The Markov chains in our model were said to have converged if plots of the Gelman-Rubin statistics indicated that widths of pooled runs and individual runs stabilized around the same value and their ratio was approximately one.³⁶ In general, we performed 100,000 iterations, the first 50,000 of which were discarded as "burn in" iterations for each of the network models we describe. We specified vague priors for the trial baselines and the basic parameters (normal distribution with mean 0 and variance 10,000) and for the random effects standard deviation (uniform distribution: U(0,2)). We use p <0.05 to define statistical significance.

To assess the adequacy of our models, we checked their overall fit by comparing the posterior mean deviance to the number of data points in any given model. These two figures are approximately equal for models that fit the data well. We also checked the statistical consistency of the models using a "back-calculation" method for networks with direct evidence from multi-arm trials.³⁷ This method requires point estimates and dispersions of the trial data being entered into the network meta-analysis. When there were two or more trials comparing two of the same treatments, we obtained these latter two quantities from traditional random effects meta-analytic models computed according to the method of DerSimonian and Laird.³⁸ All traditional meta-analyses were performed using STATA.

We performed separate network meta-analyses for antibiotic prophylaxis and for non-antibiotic prophylaxis (e.g., antiseptic rinses) because the analysis combining both types of prophylaxis resulted in a statistically inconsistent model.

PEER REVIEW

A draft of the present guideline was peer reviewed. Peer review was performed using a structured peer review form (see Appendix VIII). This form requires all peer reviewers to declare their conflicts of interest.

To determine who would serve as peer reviewers, the work group nominated external specialty societies before work on the guideline began. By having work groups specify *organizations* for review (as opposed to individuals), we are attempting to prevent overly favorable reviews that could arise should work group members choose reviewers whom they had personal or professional relationships. We also blind peer reviewers to the identities of the work group members when they peer review the draft.

The outside specialty societies were nominated at the beginning of the process and solicited for names of peer reviewers approximately six weeks before the final recommendation meeting for a guideline. The physician members of the AAOS Guidelines Oversight Committee and the Evidence Based Practice Committee review all draft AAOS clinical practice guidelines. In addition, the ADA Council on Scientific Affairs will review the guideline.

On occasion, some specialty societies (both orthopaedic and non-orthopaedic) ask their evidence-based practice (EBP) committee to provide peer review of our guidelines. The specialty society is responsible for compiling this type of review into one document before it is returned to us. We ask that the Chairpersons of these external EBP committees declare their conflicts of interest and manage the conflicts of interest of their committee members. Some specialty societies ask to post the guideline on their website for review by all of their interested members. Again, the AAOS asks that these reviews be collated into a single response by the specialty society, and that the person responsible for submitting this document to the AAOS disclose his or her financial conflicts of interest. We also ask that this posting be to the "members" only portion of the specialty societies' website because our drafted document represents a "work in progress" and is subject to change as a direct result of the review process. In addition, the draft has not been formally approved by the AAOS Board of Directors or the ADA Board of Trustees. This is not an attempt to restrict input on the draft. Nor do we consider it as a method to imply that outside specialty societies who provide review of the document necessarily agree with the stated recommendations. Hence, the reason all peer review comments and our responses are made publicly available.

AAOS and ADA staff drafted initial responses to comments about methodology. These responses were then reviewed by the work group co-chairs, who also respond to questions concerning clinical practice and techniques. All changes to a recommendation as a result of peer review input were voted on and accepted by a majority of the work group members via teleconference. All changes to any guideline recommendation are based on the evidence in the guideline recommendations. Final changes to the guideline are incorporated, detailed in a summary sheet and forwarded with the document through the rest of the review and approval process.

The AAOS and ADA believe that it is important for guideline developers to demonstrate that they are responsive to peer review. Accordingly, after the AAOS Board of Directors approves a guideline, the AAOS posts all peer reviewer comments on its website (see

http://www.aaos.org/research/guidelines/guide.asp to access these documents) with a point-by-point description of how the AAOS responded to each non-editorial comment made by each reviewer. Reviewers who wish to remain anonymous can notify the AAOS, and their names will be redacted; their comments, our responses and their conflicts of interest will however still be posted for review.

Forty-seven outside organizations were solicited to provide peer reviewers for this document. The draft of this guideline was sent to seventeen review organizations who responded to the solicitation and a total of twenty-three peer reviewers received the document not including the AAOS Evidence-based Practice Committee and Guidelines Oversight Committee members. Eighteen of these reviewers returned comments (see Appendix IX). The disposition of all non-editorial peer review comments was documented and accompanied this guideline through the public commentary and the AAOS guideline approval process.

PUBLIC COMMENTARY

After modifying the draft in response to peer review, the guideline was sent for a thirty day period of "Public Commentary." Public Commentators are blinded to the identities of the work group members. Commentators consist of members of the AAOS Board of Directors (BOD), members of the Council on Research and Quality (CORQ), members of the Board of Councilors (BOC), and members of the Board of Specialty Societies (BOS). AAOS guidelines are automatically forwarded to the AAOS BOD and CORQ for commentary. Members of the BOC and BOS are solicited for interest. If they ask to see the document, it is forwarded to them. In addition, the guideline will be forwarded to the ADA Board of Trustees, Council on Dental Practice, Council on Access, Prevention and Interprofessional Relations, Council on Dental Benefit Programs, and Council on Dental Education and Licensure for commentary.

The draft guideline is, if warranted, modified in response to public commentary by the AAOS Clinical Practice Guidelines Unit, the ADA Division of Science, and the work group members. If changes are made as a result of public comment, these changes are summarized, and those who provided commentary are notified that their input resulted in a change in the guideline. Changes as a result of public commentary are based on evidence in the guideline recommendations. All changes are detailed in a summary sheet that accompanies the document through the approval process.

Over one hundred commentators have had the opportunity to provide input into this guideline. Of these, fifty-eight members received the document and five returned comments (see Appendix IX).

THE AAOS GUIDELINE APPROVAL PROCESS

This final guideline draft was approved by the AAOS Evidence Based Practice Committee, the AAOS Guidelines Oversight Committee, the AAOS Council on Research and Quality, the ADA Council on Scientific Affairs, the AAOS Board of Directors, and the ADA Board of Trustees. Descriptions of these bodies are provided in Appendix X. These reviewing bodies do not have the option to modify the draft guideline during the approval process. They can only vote to approve it or reject it. Accordingly, no changes were made to this guideline during the approval process.

REVISION PLANS

This guideline represents a cross-sectional view of current treatment and may become outdated as new evidence becomes available. This guideline will be revised in accordance with new evidence, changing practice, rapidly emerging treatment options, and new technology. Accordingly, this guideline will be updated or withdrawn in five years in accordance with the standards of the National Guideline Clearinghouse.

GUIDELINE DISSEMINATION PLANS

The primary purpose of the present document is to provide interested readers with full documentation about not only our recommendations, but also about how we arrived at those recommendations. This document is also posted on the AAOS website at http://www.aaos.org/research/guidelines/guide.asp.

Guidelines are first announced by a press release and then published on the AAOS's and the ADA's website. Guideline summaries are published in the Journal of the American Academy of Orthopaedic Surgeons, Journal of the American Dental Association, *AAOS Now* and *ADA News*. In addition, guidelines are disseminated at the AAOS Annual Meeting in various venues such as on Academy Row and at Committee Scientific Exhibits.

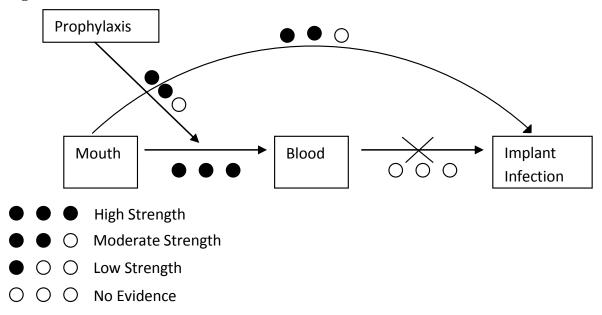
Selected guidelines are disseminated by webinar, an Online Module for the Orthopaedic Knowledge Online website, Radio Media Tours, Media Briefings, and by distributing them at relevant Continuing Medical Education (CME) courses and at the AAOS Resource Center.

Other dissemination efforts outside of the AAOS and ADA include submitting the guideline to the National Guideline Clearinghouse and distributing the guideline at other medical specialty societies' meetings.

OVERVIEW OF THE EVIDENCE

As illustrated in Figure 1, there is varying quality of evidence that explains the proposed association between dental procedures and orthopaedic implant infection. Only one study of direct evidence of moderate strength (represented in Figure 1 below, by the arching arrow) was considered for this guideline. The results of this study show that dental procedures are not risk factors for subsequent implant infection and furthermore that antibiotic prophylaxis does not reduce the risk of subsequent infection. However, multiple high strength studies of indirect evidence link oral procedures to bacteremia, a surrogate measure of risk of orthopaedic implant infection. Furthermore, multiple moderate strength studies of indirect evidence suggest that prophylaxis decreases the incidence of post dental procedure bacteremia. No studies exist that explain the microbiological relationship between bacteremia and orthopaedic implant infection.

Figure 1 Overview of the Evidence



DIRECT EVIDENCE FINDINGS

The results of one study provide direct evidence for the association between dental procedures and antibiotic prophylaxis on prosthetic hip and knee infection. This single-center, case-control study prospectively enrolled patients between 2001and 2006. 339 case patients were diagnosed with a prosthetic hip or knee infection. 339 control patients were hospitalized on an orthopedic service without a prosthetic hip or knee infection. Characteristics of case and control patients were compared, risk factors for prosthetic hip or knee infection were analyzed and multivariate logistic regression was performed to assess the association between variables and odds of infection. The model included covariates of sex, joint age, dental propensity score, body mass index >40, procedure time >4 h, immunocompromised host, American Society of Anesthesiologists score, wound healing complications, prior arthroplasty or surgery on the index joint, use of surgical antibiotic prophylaxis, postoperative urinary tract infection, and distant organ infection. The results from this model show that low and high-risk dental procedures (see Table 11)

performed within 6 months and 2 years of the hospital admission date were not significantly associated with increased risk of prosthetic hip or knee infection compared with no dental procedure (see Table 12 for a summary of the results of the logistic regression model). The model also assessed the association between antibiotic prophylaxis and prosthetic joint infection. Low and high-risk dental procedures with antibiotic prophylaxis were compared with the same procedures without prophylaxis. No significant associations were found (see

Table 13).

Table 11 High and Low Risk Dental Procedures Defined by Berbari, et al.

| High Risk Dental Procedures | Low Risk Dental Procedures | | | | |
|------------------------------------|-----------------------------------|--|--|--|--|
| Dental abscess therapy | Dental fillings | | | | |
| Dental extraction | Endodontic treatment | | | | |
| Dental filing | Fluoride treatment | | | | |
| Dental hygiene | Restorative dentistry | | | | |
| Periodontal treatment | | | | | |
| Mouth surgery | | | | | |
| 1 1 1 1007 | · II . A · | | | | |

based on the 1997 version of the American Heart Association Guideline on Infective Endocarditis

QUALITY AND APPLICABILITY

Only one study of moderate quality and applicability exists that provides direct evidence for an association between dental procedures and prosthetic hip and knee infection. Details of our appraisal of this study are provided in Table 69 of Appendix XII.

RESULTS

Table 12 Dental procedures performed and risk of prosthetic hip or knee infection at 6 months and 2 years

| | Odds Ratio (95% Confidence Interval) | | | | | |
|-----------------------------|--------------------------------------|------|---------------|------|--|--|
| Variable | 6 months | P | 2 years | P | | |
| Low-risk dental procedure | | | | | | |
| Low-risk dental procedure | 1.1 (0.6-2.1) | 0.77 | 0.6 (0.4-1.1) | 0.11 | | |
| without antibiotic | | | | | | |
| prophylaxis | | | | | | |
| Low-risk dental procedure | 0.7 (0.3-1.5) | 0.33 | 0.8 (0.5-1.2) | 0.29 | | |
| with antibiotic prophylaxis | | | | | | |
| High-risk dental procedure | | | | | | |
| High-risk dental procedure | 0.8 (0.4-1.7) | 0.6 | 0.8 (0.4-1.6) | 0.56 | | |
| without antibiotic | | | | | | |
| prophylaxis | | | | | | |
| High-risk dental procedure | 0.5 (0.3-0.9) | 0.01 | 0.7 (0.5-1.1) | 0.14 | | |
| with antibiotic prophylaxis | | | | | | |

Table 13 Antibiotic prophylaxis and risk of prosthetic hip or knee infection at 6 months and 2 years $\frac{1}{2}$

| | Odds Ratio (95% Confidence Interval) | | | |
|------------------------|--------------------------------------|---------------|--|--|
| Variable | 6 Months | 2 Years | | |
| Antibiotic Prophylaxis | | | | |
| Low-risk procedure | 0.7 (0.3-1.5) | 1.2 (0.7-2.2) | | |
| High-risk procedure | 0.7 (0.3-1.4) | 0.9 (0.5-1.6) | | |

INDIRECT EVIDENCE: DENTAL PROCEDURES AND BACTEREMIA FINDINGS

Multiple studies of high quality regarding dental procedures with bacteremia as the outcome are considered for this guideline. Rates of bacteremia after dental procedures varied significantly by and within procedure group. Rates are reported as either incidence or prevalence. We focused primarily on the incidence data because these studies reported new cases of bacteremia as a result of the dental procedure. Studies that reported prevalence did not take the necessary measures to ensure that the study population was free of bacteremia before undergoing their respective dental procedures. Due to the heterogeneity of bacteremia rates within procedure group we were unable to calculate an accurate mean value. Therefore the rates of bacteremia are presented in box plots in Figure 2 & Figure 4. Median incidence rates range from approximately 5% for chewing to upwards of 65% for simple tooth extraction and gingivectomy. Prevalence rates are comparable. Rates of bacteremia were represented by a single study in some cases (see Figure 3 & Figure 5 for details). Individual study details can be found in

Table 63 and Table 64 in Appendix XI.

QUALITY AND APPLICABILITY

Refer to Table 97 to Table 113 in Appendix XII.

RESULTS

Figure 2 Incidence of bacteremia by procedure group

Bacteremia (Incidence)

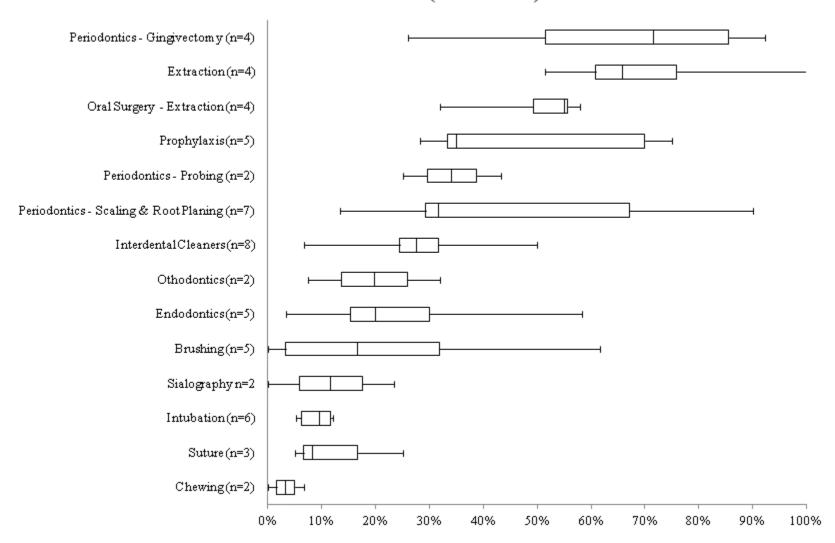


Figure 3 Incidence of bacteremia in single study groups

Bacteremia (Incidence)

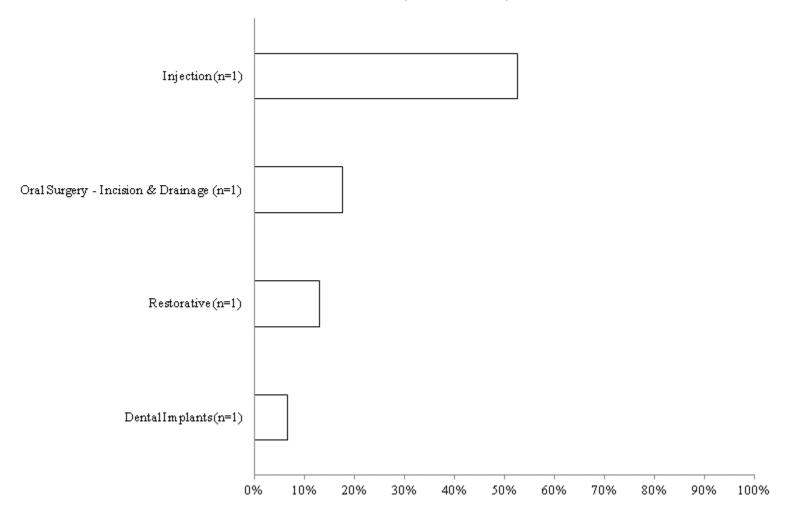


Figure 4 Prevalence of bacteremia by group

Bacteremia (Prevalence)

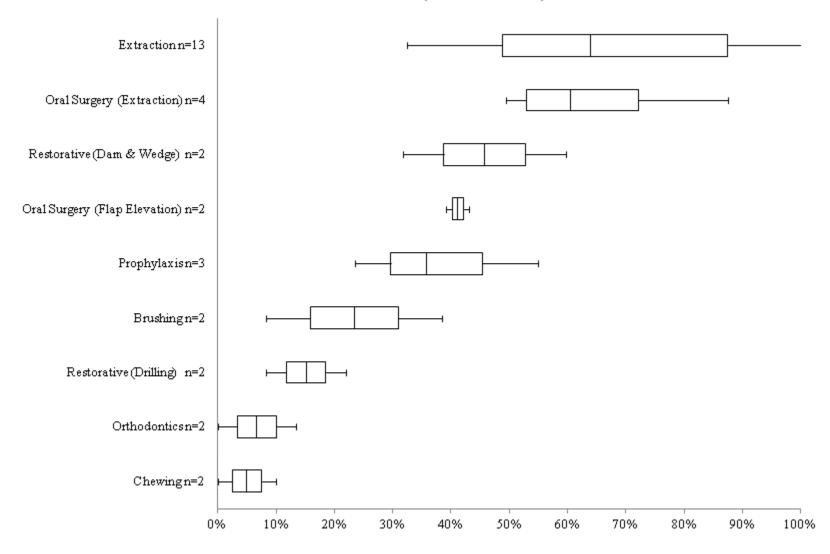
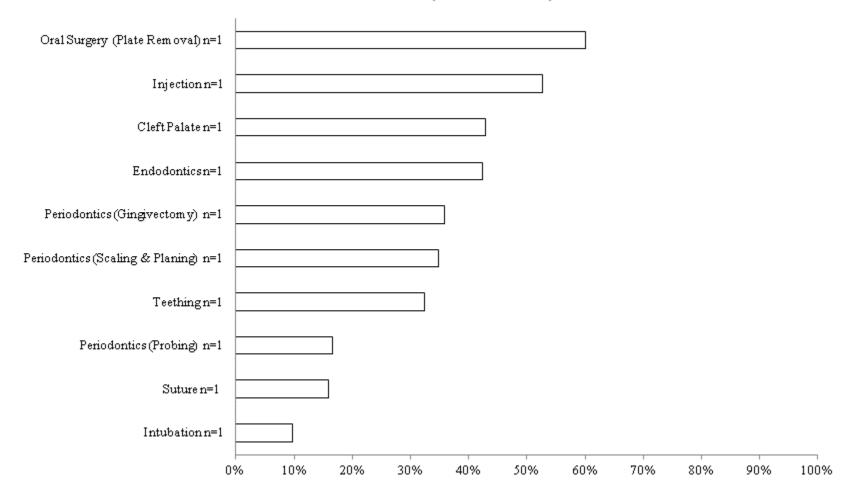


Figure 5 Prevalence of bacteremia in single study groups

Bacteremia (Prevalence)



INDIRECT EVIDENCE: RISK FACTORS FOR DENTAL PROCEDURE RELATED BACTEREMIA FINDINGS

While the quality of the evidence is low, several prognostic studies have addressed a multitude of patient characteristics as potential risk factors for developing bacteremia from dental procedures. These low strength studies report on oral health indicators and general patient characteristics such as age, gender, etc. The results vary across and within procedure groups. Evidence is often contradictory. See Table 14 for a summary of significant findings and Table 15 - Table 23 for details.

QUALITY AND APPLICABILITY

Refer to Table 88 - Table 96 in Appendix XII.

RESULTS
Table 14 Summary of Risk Factor Significance (Proportion of studies that reported significant results)

| Risk Factor | Brushing | Chewing | Dental | Inter- | Intubation | Oral | Periodontic | Restorative | Tooth |
|--------------------------|-----------|---------|-------------|--------------------|----------------|--------------|-------------|-------------|------------|
| | | | Prophylaxis | dental Cleaning | | Surgery | | | Extraction |
| Patient | | | | Resul | ts (% Signific | ant, n/N) | l | | |
| Characteristics | | | | | , 8 | , , | | | |
| Age | 50%, 1/2 | | 33%, 1/3 | 50%, 1/2 | 0%, 0/1 | 0%, 0/2 | 0%, 0/1 | 0%, 0/1 | 33%, 1/3 |
| BMI | 0%, 0/1 | | | | | | | | |
| Cirrhosis | 100%, 1/1 | | | | | | | | |
| Diabetes | | | 0%, 0/1 | | | | | | |
| Gender | 0%, 0/2 | | 0%, 0/3 | 0%, 0/2 | 0%, 0/1 | 0%, 0/2 | | 0%, 0/1 | 0%, 0/3 |
| Inflammatory Disease | | | | | | 100%, 1/1 | | | 100%, 2/2 |
| Mixed | | | | | | | | 0%, 0/1 | |
| Dentition | | | | | | | | | |
| Race | | | | | | | | 0%, 0/1 | |
| Smoking | | | 0%, 0/2 | 0%, 0/1 | | | 0%, 0/1 | | |
| Status | | | | | | | | | |
| Procedure | | | | | | | | | |
| # Teeth | | | | | | 0%, 0/1 | | | 100%, 4/4 |
| Extracted Anaesthesia | | | | | | 0%, 0/1 | | | |
| Anaesthetic Modality | | | | | | 0 /0 , 0/ 1 | | | 100%, 1/1 |
| Anaesthetic Technique | | | | | | | | | 0%, 0/1 |
| Bleeding | 0%, 0/1 | | 50%, 1/2 | 0%, 0/1 | | 100%, 1/1 | 50%, 1/2 | | 50%, 1/2 |
| Bleeding Type | 100%, 1/1 | | | | | | | | 0%, 0/1 |
| Blood Loss | | | | | | 0%, 0/1 | | | 100%, 1/1 |

| Risk Factor | Brushing | Chewing | Dental Prophylaxis | Inter- dental Cleaning | Intubation | Oral Surgery | Periodontic | Restorative | Tooth Extraction |
|--------------------------|-----------|---------|-----------------------|------------------------------|------------|-----------------|-------------|-------------|---------------------|
| Procedure Time | | | 0%, 0/1 | 0%, 0/1 | | 100%, 1/1 | | | |
| Oral Health | | | | | | | | | |
| # Teeth | | | | | | 0%, 0/1 | 0%, 0/1 | | |
| Present | | | | | | | | | |
| Abscess | | | | | | 0%, 0/1 | | | 0%, 0/2 |
| Apical Lucency | 0%, 0/1 | | | | | | | | 0%, 0/1 |
| Calculus Index/Score | 100%, 1/1 | | | | | | | | 0%, 1/1 |
| Caries | 0%, 0/1 | | | | | | | 0%, 0/1 | 0%, 0/1 |
| Caries Depth | 0%, 0/1 | | | | | | | 0%, 0/1 | 0%, 0/1 |
| Clinical Attachment Loss | | | | 0%, 0/1 | | | | | |
| Gingival Index/Score | 25%, 1/4 | | 100%, 1/1 | 0%, 0/1 | | 50%, 1/2 | | 100%, 1/1 | 67%, 2/3 |
| Gingival Size | | | | | | | | 0%, 0/1 | |
| Gingivitis | 0%, 0/1 | 0%, 0/1 | 0%, 0/1 | | | | | | |
| Infected Tooth | | | | | | 100%, 1/1 | | | |
| Odontogenic Disease | | | | | | | | | 0%, 0/1 |
| Oral Health Status | | | | | 0%, 0/1 | 0%, 0/1 | | | 50%, 1/2 |
| Periodontal Diagnosis | | | 0%, 0/1 | | | | | | 0%, 0/1 |
| Periodontitis | 0%, 0/1 | 0%, 0/1 | 100%, 1/1 | 0%, 0/1 | | | 50%, 1/2 | 0%, 0/1 | |
| Plaque Index/Score | 67%, 2/3 | | 50%, 1/2 | 0%, 0/1 | | 0%, 0/1 | | | 0%, 0/3 |

| Risk Factor | Brushing | Chewing | Dental Prophylaxis | Inter- dental Cleaning | Intubation | Oral Surgery | Periodontic | Restorative | Tooth Extraction |
|-------------------|----------|---------|-----------------------|------------------------------|------------|-----------------|-------------|-------------|---------------------|
| Probing | | | 0%, 0/2 | 0%, 0/1 | | | 0%, 0/1 | | 33%, 1/3 |
| Depth | | | | | | | | | |
| Probing | 0%, 0/1 | | | | | | 100%, 1/1 | | 0%, 0/1 |
| Depth Mean | | | | | | | | | |
| Radiolucency | | | | | | | | 0%, 0/1 | |
| Recession | | | 0%, 0/1 | | | | | | |
| Suppuration | | | | | | | | 0%, 0/1 | |
| Swelling | | | | | | | | 0%, 0/1 | |
| Tooth Mobility | 0%, 0/1 | | | | | | | | 0%, 0/1 |

Table 15 Risk Factors for Brushing Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|------------------|----------|----|------------------------------------|------------------------------------------------------|----------------------------|-----------------------------------|
| Ashare 2009 | Low | 48 | ANOVA followed by Bonferroni | Bacteremia (Bacterial Load @ 30s, 5m, 15m) | Cirrhosis | p<0.01 for all time points |
| Ashare 2009 | Low | 48 | unknown | Bacteremia (Bacterial Load @ 30s, 5m, 15m) | Age | NS for all time points |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Age | OR 1.06 p=.017 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | ВМІ | OR 0.99 p=.749 |
| Ashare 2009 | Low | 48 | unknown | Bacteremia (Bacterial Load @ 30s, 5m, 15m) | Gender | NS for all time points |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Sex (risk level=female) | OR 1.09 p=.866 |
| Ashare 2009 | Low | 48 | Linear regression | Bacteremia (Bacterial Load @ 30s, 5m, 15m) | Plaque Index | p<0.01 @ 30s & 5m, NS @ 15m |
| Bhanji 2002 | Low | 50 | logistic regression | Bacteremia | Plaque Score | OR 1.05, p=0.44 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Mean plaque score | OR 2.53 p=.010 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Plaque score ≥ 2 | OR 3.78 p=.008 |
| Ashare 2009 | Low | 48 | Linear regression | Bacteremia (Bacterial Load @ 30s, 5m, 15m) | Gingival Index | NS for all time points |
| Bhanji 2002 | Low | 50 | chi square | Bacteremia | Gingival Score | p=0.96 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Mean gingival score | OR 1.62 p=.203 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Gingival score ≥ 2 | OR 1.61 p=.335 |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|------------------|----------|----|------------------------|---------------------------------------------------------------|----------------------------------|-------------------|
| Silver 1977 | Low | 96 | Critical ratio test | Bacteremia | Gingival Index | p<.01 |
| Forner 2006 | Low | 20 | Fishers exact test | Bacteremia | Gingivitis | NS |
| Forner 2006 | Low | 20 | Fishers exact test | Bacteremia | Periodontitis | NS |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Mean calculus score | OR 1.77 p=.048 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Calculus score ≥ 2 | OR 4.43 p=.004 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Bleeding with toothbrushing | OR 0.89 p=.810 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Bleeding type with toothbrushing | OR 7.96 p=.015 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Mean probing depth | OR 1.02 p=.918 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Tooth mobility score | OR 1.93 p=.200 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Dental caries | OR 4.40 p=.165 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Depth of dental caries | OR 0.43 p=.155 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Apical lucency | OR 2.37 p=.086 |
| Lockhart 2009 | Low | 98 | logistic regression | Bacteremia (Infective Endocarditis related bacteria) | Apical lucency size (mm) | OR 0.87 p=.647 |

Table 16 Risk Factors for Chewing Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|----|-------------------------|------------|---------------|---------|
| Forner | Very Low | 20 | Fisher's exact test | Bacteremia | Periodontitis | NS |
| 2006 | | | | | | |
| Forner | Very Low | 20 | Fisher's exact test | Bacteremia | Gingivitis | NS |
| 2006 | | | | | | |

Table 17 Risk Factors for Dental Prophylaxis Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|----------------|----------|----|-------------------------------------|------------------------|--------------------------------------------------|--------------|
| Cherry 2007 | Low | 60 | Logistic regression | Bacteremia | Age | OR 1.4 p=.05 |
| De Leo 1974 | Low | 39 | Chi square | Bacteremia | Age | 6.31, NS |
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Age | NS |
| Cherry 2007 | Low | 60 | Logistic regression | Bacteremia | Gender | NS |
| De Leo 1974 | Low | 39 | Chi square | Bacteremia | Sex | NS |
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Gender | NS |
| Cherry 2007 | Low | 60 | Logistic regression | Bacteremia | Smoking status | NS |
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Smoking | NS |
| Cherry 2007 | Low | 60 | Logistic regression | Bacteremia | Plaque Index | NS |
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Plaque Index | 0.41 p=.0117 |
| Cherry 2007 | Low | 60 | Logistic regression | Bacteremia | Modified papilla, margin, attached gingiva index | NS |
| Cherry 2007 | Low | 60 | Logistic regression | Bacteremia | Probing depth | NS |
| Cherry 2007 | Low | 60 | Logistic regression | Bacteremia | Recession | NS |
| Cherry 2007 | Low | 60 | Logistic regression | Bacteremia | Bleeding on scaling | NS |
| Forner 2006 | Low | 20 | Fishers exact test | Bacteremia | Periodontitis | p<.001 |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|-----------------|----------|----|-------------------------------------|------------------------|-------------------------|--------------|
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Periodontal diagnosis | NS |
| Forner 2006 | Low | 20 | Fishers exact test | Bacteremia | Gingivitis | NS |
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Gingival Index | 0.53 p<.0001 |
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Bleeding on probing | 0.45 p=.0089 |
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Probing pocket depth >5 | NS |
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Pocket sum score | NS |
| Forner 2006 | Low | 20 | Spearman's correlation coefficients | Bacteremia (magnitude) | Scaling time* | NS |
| Trivedi 1984 | Low | 40 | Chi square | Bacteremia | Diabetes | 4.5 p>0.5 |

^{*}Procedure related risk factor

Table 18 Risk Factors for Inter-dental Cleaning Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|-------------------|----------|----|-------------------------------------|------------|----------------------|------------|
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Periodontitis | 0.17 p=.2 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Age | 0.18 p=.2 |
| Linberger 1973 | Low | 21 | Chi square | Bacteremia | Age | 0.81 p<.04 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Gender | -0.08 p=.5 |
| Linberger 1973 | Low | 21 | Exact method of binomial dist. | Bacteremia | Sex | 1.97, NS |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Smoking status | -0.04 p=.7 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Time spent flossing* | -0.04 p=.8 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Gingival Index | 0.22 p=.09 |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|----------------|----------|----|-------------------------------------|------------|------------------------------------|------------|
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Plaque Index | 0.07 p=.6 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | % of sites bleeding on flossing | 0.17 p=.2 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | # sites bleeding on flossing | 0.17 p=.2 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | % of sites bleeding on probing | 0.16 p=.2 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Pocket depth | 0.09 p=.5 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Clinical attachment loss | 0.06 p=.6 |
| Crasta 2009 | Low | 60 | Spearman's correlation coefficients | Bacteremia | Self-reported daily flossing | −0.12 p=.4 |

^{*}Procedure related risk factor

Table 19 Risk Factors for Intubation Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|-----|------------------|------------|-------------|---------|
| Valdes | Low | 110 | Logistic | Bacteremia | Age | NS |
| 2008 | | | regression | | | |
| Valdes | Low | 110 | Logistic | Bacteremia | Sex | NS |
| 2008 | | | regression | | | |
| Valdes | Low | 110 | Logistic | Bacteremia | Oral health | NS |
| 2008 | | | regression | | status | |

Table 20 Risk Factors for Oral Surgery Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|-----------|----------|-----|------------------|------------|--------------|-------------|
| Enabulele | Low | 170 | chi-square | Bacteremia | Inflammatory | 0.004 p=.05 |
| 2008 | | | | | disease | |
| Enabulele | Low | 170 | chi-square | Bacteremia | Sex | NS |
| 2008 | | | | | | |
| Tomas | Low | 100 | not reported | Bacteremia | Gender | NS |
| 2008 | | | | | | |
| Roberts | Low | 154 | chi-square | Bacteremia | Abscess | 1.878 |
| 1998 | | | | | | p=.1706 |
| Roberts | Low | 154 | Pearson | Bacteremia | Age | 0.29 |
| 1998 | | | correlation | | | |
| | | | coefficient | | | |
| Tomas | Low | 100 | not reported | Bacteremia | Age | NS |
| 2008 | | | | | | |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|-----------------|----------|-----|-------------------------------------|------------|---------------------------------------------------------------------------------------|---------|
| Roberts 1998 | Low | 154 | Scheffe's multiple comparison | Bacteremia | Plaque Index | p=.47 |
| Roberts 1998 | Low | 154 | Scheffe's multiple comparison | Bacteremia | Gingival Index | p<.03 |
| Takai 2005 | Low | 237 | chi-square | Bacteremia | Gingival Index | NS |
| Roberts 1998 | Low | 154 | Scheffe's multiple comparison | Bacteremia | Bleeding Index | p<.04 |
| Takai 2005 | Low | 237 | chi-square | Bacteremia | Oral hygiene index simplified | NS |
| Takai 2005 | Low | 237 | chi-square | Bacteremia | # teeth present | NS |
| Takai 2005 | Low | 237 | chi-square | Bacteremia | Blood loss | NS |
| Takai 2005 | Low | 237 | chi-square | Bacteremia | Duration of procedure* | p<.05 |
| Takai 2005 | Low | 237 | chi-square | Bacteremia | # teeth extracted* | NS |
| Takai 2005 | Low | 237 | chi-square | Bacteremia | Method of procedure* | NS |
| Takai 2005 | Low | 237 | chi-square | Bacteremia | Infection in extracted tooth (periodontitis, periapical infection, and pericoronitis) | p<.01 |
| Takai 2005 | Low | 237 | chi-square | Bacteremia | Anaesthesia for procedure* | NS |

^{*}Procedure related risk factor

Table 21 Risk Factors for Periodontic Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------------|----------|----|------------------------|------------|------------------------|---------------------------------|
| Daly 1997 | Low | 30 | chi-square | Bacteremia | Periodontitis severity | p=.9 |
| Daly 2001 | Low | 40 | logistic regression | Bacteremia | Periodontitis | OR 5.993 CI=1.081- 33.215 |
| Daly 1997 | Low | 30 | t-test | Bacteremia | Bleeding on probing | p=.3 |
| Daly 2001 | Low | 40 | logistic regression | Bacteremia | Bleeding on probing | OR 1.025 CI=1.004- 1.047 |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|----|------------------|------------|---------------|------------|
| Daly | Low | 40 | logistic | Bacteremia | Age | OR 1.008 |
| 2001 | | | regression | | | CI=.960- |
| | | | | | | 1.058 |
| Daly | Low | 40 | logistic | Bacteremia | Sex | NS |
| 2001 | | | regression | | | |
| Daly | Low | 40 | logistic | Bacteremia | Smoking | NS |
| 2001 | | | regression | | status | |
| Daly | Low | 40 | logistic | Bacteremia | # of teeth | OR 1.0 |
| 2001 | | | regression | | | CI=.845- |
| | | | | | | 1.185 |
| Daly | Low | 40 | logistic | Bacteremia | Total probing | OR 1.006 |
| 2001 | | | regression | | depth | CI=.999- |
| | | | | | | 1.013 |
| Daly | Low | 40 | logistic | Bacteremia | Plaque index | OR 3.154 |
| 2001 | | | regression | | | CI=.603- |
| | | | | | | 16.514 |
| Daly | Low | 40 | logistic | Bacteremia | Mean probing | OR 1.444 |
| 2001 | | | regression | | depth per | CI=.1.055- |
| | | | | | tooth | 1.977 |

Table 22 Risk Factors for Restorative Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|-----------------|----------|----|---------------------------------|------------|------------------------------------------------|---------|
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Age | p=.06 |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Sex | NS |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Race | NS |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Gingival Score (0-3) | p=.01 |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Gingival Size (0-3) | NS |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Periodontal disease with probing >3mm | NS |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Mixed Dentition | p=.08 |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Caries Present | NS |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Depth of caries (0-3) | NS |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Periapical radiolucency | NS |
| Brennan 2007 | Very Low | 51 | chi-square or fisher's exact | Bacteremia | Size radiolucency (mm) | NS |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|---------|----------|----|------------------|------------|-------------|---------|
| Brennan | Very Low | 51 | chi-square or | Bacteremia | Swelling | NS |
| 2007 | | | fisher's exact | | | |
| Brennan | Very Low | 51 | chi-square or | Bacteremia | Suppuration | NS |
| 2007 | | | fisher's exact | | | |

Table 23 Risk Factors for Extraction Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|----------|----------|-----|------------------|----------------|------------------|------------|
| Barbosa | Low | 210 | logistic | Bacteremia 30s | Oral health | OR 3.704 |
| 2010 | | | regression | | status | (1.929- |
| | | | (univariate) | | | 7.109) |
| Barbosa | Low | 210 | logistic | Bacteremia 15m | Oral health | OR 2.047 |
| 2010 | | | regression | | status | (1.138- |
| | | | (univariate) | | | 3.683) |
| Wahlmann | Low | 59 | logistic | Bacteremia | Oral Hygiene | NS |
| 1999 | | | regression | | | |
| Wahlmann | Low | 59 | logistic | Bacteremia | Periodontal | NS |
| 1999 | | | regression | | status | |
| Barbosa | Low | 210 | logistic | Bacteremia 30s | Local anesthetic | OR 0.143 |
| 2010 | | | regression | | technique* | (0.063- |
| | | | (univariate) | | | 0.323), OR |
| | | | | | | 0.119 |
| | | | | | | (0.046- |
| | | | | | | 0.309) |
| Barbosa | Low | 210 | logistic | Bacteremia 15m | Local anesthetic | OR 0.179 |
| 2010 | | | regression | | technique* | (0.090- |
| | | | (univariate) | | • | 0.356), OR |
| | | | | | | 0.186 |
| | | | | | | (0.076- |
| | | | | | | 0.455) |
| Barbosa | Low | 210 | logistic | Bacteremia 60m | Local anesthetic | OR 0.118 |
| 2010 | | | regression | | technique* | (0.027- |
| | | | (univariate) | | | 0.520), OR |
| | | | | | | 0.251 |
| | | | | | | (0.055- |
| | | | | | | 1.135) |
| Barbosa | Low | 210 | logistic | Bacteremia 30s | Anesthetic | OR 7.431 |
| 2010 | | | regression | | modality* | (3.453- |
| | | | (univariate) | | | 15.990) |
| Barbosa | Low | 210 | logistic | Bacteremia 15m | Anesthetic | OR 5.518 |
| 2010 | | | regression | | modality* | (3.004- |
| | | | (univariate) | | | 10.133) |
| Barbosa | Low | 210 | logistic | Bacteremia 60m | Anesthetic | OR 6.247 |
| 2010 | | | regression | | modality* | (2.058- |
| | | | (univariate) | | | 18.961) |
| Barbosa | Low | 210 | logistic | Bacteremia 30s | Anesthetic | OR 5.040 |
| 2010 | | | regression | | modality* | (2.068- |
| | | | (multivariate) | | | 12.283) |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|------------|----------|-----|------------------|----------------|-----------------------|--------------|
| Barbosa | Low | 210 | logistic | Bacteremia 15m | Anesthetic | OR 5.368 |
| 2010 | | | regression | | modality* | (2.361- |
| | | | (multivariate) | | | 12.211) |
| Barbosa | Low | 210 | logistic | Bacteremia 60m | Anesthetic | OR 6.464 |
| 2010 | | | regression | | modality* | (1.333- |
| | | | (multivariate) | | | 31.346) |
| Barbosa | Low | 210 | logistic | Bacteremia 15m | # of | OR 1.126 |
| 2010 | | | regression | | extractions* | (1.046- |
| | | | (univariate) | | | 1.212) |
| Barbosa | Low | 210 | logistic | Bacteremia 60m | # of | OR 1.128 |
| 2010 | | | regression | | extractions* | (1.042- |
| | | | (univariate) | | | 1.222) |
| Coulter | Low | 58 | Spearman's | Bacteremia | # of teeth | r=0.08 |
| 1990 | | | correlation | | extracted* | |
| | | | coefficient | | | |
| Okabe 1995 | Low | 183 | Mann-Whitney | Bacteremia | # of | 4367.5 |
| | | | | | extractions* | p<.0001 |
| Wahlmann | Low | 59 | logistic | Bacteremia | # of | Significant |
| 1999 | | | regression | | extractions* | for Control |
| | | | | | | grp |
| Coulter | Low | 58 | chi-square | Bacteremia | Plaque Index | NS |
| 1990 | | | 1 | | | |
| Lockhart | Low | 96 | logistic | Bacteremia | Mean plaque | OR 0.74 |
| 2009 | | | regression | | score | p=.236 |
| Lockhart | Low | 96 | logistic | Bacteremia | Plaque score ≥ 2 | OR 0.90 |
| 2009 | | | regression | | 1 | p=.811 |
| Roberts | Low | 154 | Scheffe's | Bacteremia | Plaque Index | p=.47 |
| 1998 | | | multiple | | 1 | r |
| | | | comparison | | | |
| Coulter | Low | 58 | chi-square | Bacteremia | Gingival Index | NS |
| 1990 | | | 1 | | | |
| Lockhart | Low | 96 | logistic | Bacteremia | Mean gingival | OR 0.71 |
| 2009 | | | regression | | score | p=.217 |
| Lockhart | Low | 96 | logistic | Bacteremia | Gingival score | OR 0.76 |
| 2009 | | | regression | | ≥ 2 | p=.518 |
| Roberts | Low | 154 | Scheffe's | Bacteremia | Gingival Index | p<.03 |
| 1998 | | | multiple | | | r |
| 1,,,, | | | comparison | | | |
| Coulter | Low | 58 | Fisher's | Bacteremia | Abscess | p=0.2088 |
| 1990 | | | | | | r 3.2333 |
| Roberts | Low | 154 | chi-square | Bacteremia | Abscess | 1.878 |
| 1998 | | | 1 | | | p=.1706 |
| Enabulele | Low | 170 | chi-square | Bacteremia | Inflammatory | 0.004 p=.05 |
| 2008 | | 1,0 | Jin Square | Zuctoronnu | disease | 0.00 i p=.05 |
| Okabe 1995 | Low | 183 | Fisher's | Bacteremia | Inflammatory | p<.0005 |
| Chaoc 1773 | 2011 | | | 2 uctoronnu | disease | P |
| Enabulele | Low | 170 | chi-square | Bacteremia | Sex | NS |
| 2008 | LOW | 1/0 | om square | Ductoronna | JOA | 110 |
| 2000 | | I | 1 | | | 1 |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|------------|----------|-----|------------------|------------|------------------|--------------|
| Lockhart | Low | 96 | logistic | Bacteremia | Sex (risk | OR 1.64 |
| 2009 | | | regression | | level=female) | p=.241 |
| Okabe 1995 | Low | 183 | Fisher's | Bacteremia | Sex | 0.624, NS |
| Lockhart | Low | 70 | chi-square or | Bacteremia | Surgery Time < | p=.04 |
| 1996 | | | Fisher's exact | | 3m* | |
| Lockhart | Low | 70 | chi-square or | Bacteremia | Surgery Time > | p=.04 |
| 1996 | | | Fisher's exact | | 6m* | 1 |
| Okabe 1995 | Low | 183 | Mann-Whitney | Bacteremia | Duration of | 4050 p<.05 |
| | | | | | procedure* | • |
| Wahlmann | Low | 59 | logistic | Bacteremia | Duration of | NS |
| 1999 | | | regression | | procedure* | |
| Lockhart | Low | 70 | chi-square or | Bacteremia | Odontogenic | NS |
| 1996 | | | Fisher's exact | | disease severity | |
| Lockhart | Low | 96 | logistic | Bacteremia | Age | OR 1.03 |
| 2009 | | | regression | | | p=.211 |
| Okabe 1995 | Low | 183 | Mann-Whitney | Bacteremia | Age | 4517.5 |
| | | | | | | p<.0005 |
| Roberts | Low | 154 | Pearson | Bacteremia | Age | 0.29 |
| 1998 | | | correlation | | | |
| | | | coefficient | | | |
| Lockhart | Low | 96 | logistic | Bacteremia | BMI | OR 0.99 |
| 2009 | | | regression | | | p=.630 |
| Lockhart | Low | 96 | logistic | Bacteremia | Mean calculus | OR 0.93 |
| 2009 | | | regression | | score | p=.724 |
| Lockhart | Low | 96 | logistic | Bacteremia | Calculus score | OR 0.82 |
| 2009 | | | regression | | ≥ 2 | p=.715 |
| Lockhart | Low | 96 | logistic | Bacteremia | Bleeding with | NA |
| 2009 | | | regression | | toothbrushing | |
| Lockhart | Low | 96 | logistic | Bacteremia | Bleeding type | NA |
| 2009 | | | regression | | with | |
| | | | | | toothbrushing | |
| Okabe 1995 | Low | 183 | Mann-Whitney | Bacteremia | Blood loss (ml) | 3997.5 p<.05 |
| Roberts | Low | 154 | Scheffe's | Bacteremia | Bleeding Index | p<.04 |
| 1998 | | | multiple | | | |
| | | | comparison | | | |
| Lockhart | Low | 96 | logistic | Bacteremia | Mean probing | OR 0.95 |
| 2009 | | | regression | | depth | p=.735 |
| Lockhart | Low | 96 | logistic | Bacteremia | Tooth mobility | OR 1.01 |
| 2009 | | | regression | | score | p=.978 |
| Lockhart | Low | 96 | logistic | Bacteremia | Dental caries | OR 1.66 |
| 2009 | | | regression | | | p=.452 |
| Lockhart | Low | 96 | logistic | Bacteremia | Depth of dental | OR 0.21 |
| 2009 | | | regression | | caries | p=.156 |
| Lockhart | Low | 96 | logistic | Bacteremia | Apical lucency | OR 0.86 |
| 2009 | | | regression | | | p=.724 |
| Lockhart | Low | 96 | logistic | Bacteremia | Apical lucency | OR 1.00 |
| 2009 | | | regression | | size (mm) | p=.995 |

^{*}Procedure related risk factor

INDIRECT EVIDENCE: PROPHYLAXIS FOR DENTAL PROCEDURE RELATED BACTEREMIA FINDINGS

We recognize the diversity of opinion concerning the clinical importance of bacteremia as a surrogate outcome for orthopaedic implant infection and understand the clinician's concern and rationale for wanting to prevent bacteremia. Multiple studies of moderate quality regarding prophylaxis for the prevention of bacteremia post dental procedure suggest that antibiotic and topical antimicrobial prophylaxis are effective in reducing the rate of bacteremia after simple tooth extraction. There was insufficient data to investigate the effects of prophylaxis in regard to other dental procedure groups via a meta-analysis. However, simple tooth extraction resulted in the second highest median incidence of bacteremia and the highest median prevalence of bacteremia for all procedure groups (see Figure 2 & Figure 4). Table 24 describes the included studies related to antibiotic prophylaxis for the prevention of bacteremia upon tooth extraction. Table 25 describes the included studies related to topical antimicrobials for the prevention of bacteremia upon tooth extraction.

We performed network meta-analyses in order to determine which prophylactic treatments are most effective. An initial attempt was made to combine both antibiotics and topical antimicrobials into a single network meta-analysis. The exact cause of the inconsistency could not be determined and therefore all results of antibiotic and topical antimicrobial prophylaxis are presented independent of one another. The implication of this inconsistency is that formal (as well as casual) indirect comparisons of treatment effects can be misleading and are thus avoided in this clinical practice guideline. See Table 62 in the Appendix XI for results of the consistency check. Further details on the results of these independent network meta-analyses are presented in Recommendation 1 and Recommendation 2.

QUALITY AND APPLICABILITY

Refer to Table 70 - Table 87 in Appendix XII.

RESULTS
Table 24 Antibiotic prophylaxis and tooth extraction bacteremia

| Study | N | Strength | Outcome (specific type) | Active Antibiotic (%, n/N) | Control (%, n/N) | Route of Administration | Time of Administration | Results |
|------------------|-----|----------|----------------------------|----------------------------------------------------------------------------------|--------------------------------------|-------------------------------------------------------------------------------------------------------|---------------------------------|-------------------------------------------------------------------|
| Lockhart 2008 | 179 | High | Bacteremia | Amoxicillin (56%, 50/90) | Placebo (80%, 71/89) | Oral | 1 hour before procedure | Favors Amoxicillin |
| Lockhart 2004 | 100 | High | Bacteremia | Amoxicillin (33%, 16/49) | Placebo (84%, 43/51) | Oral elixir | 1 hour before intubation | Favors Amoxicillin |
| Aitken 1995 | 40 | Moderate | Bacteremia | Erythromycin (60%, 12/20) Clindamycin (40%, 8/20) | N/A | Oral | 1-1.5 hours before procedure | Favors Clindamycin over Erythromycin |
| Cannell 1991 | 60 | Moderate | Bacteremia | Erythromycin (60%, 13/20) Josamycin (70%, 14/20) | Placebo (65%, 13/20) | Oral | 1-1.5 hours before procedure | Erythromycin and Josamycin marginally more effective than placebo |
| Casolari 1989 | 106 | Moderate | Bacteremia | Penicillin (48%, 12/25) Antiseptic rinse (44%, 11/25) | No Treatment (67.9%, 38/56) | Oral | 1 hour before procedure | Favors Penicillin and Antiseptic over control |
| Coulter 1990 | 58 | Moderate | Bacteremia | Penicillin or Amoxicillin or Amoxicillin or Erythromycin (35%, 9/26) | No Treatment (63%, 20/32) | Intramuscular (Penicillin), Oral (Amoxicillin), Intravenous (Amoxicillin), Intravenous (Erythromycin) | Unclear | Favors Antibiotics over control |

| Study | N | Strength | Outcome (specific type) | Active Antibiotic (%, n/N) | Control (%, n/N) | Route of Administration | Time of Administration | Results |
|-----------|-----|----------|------------------------------------------|--------------------------------------------------------------------------------------------------|--------------------------------------|----------------------------|----------------------------|---------------------------------------------------------------------------------------------------|
| Diz 2006 | 221 | Moderate | Bacteremia | Amoxicillin (46.4%, 26/56) Clindamycin (85.1%, 46/54) Moxifloxacin (56.9%, 33/58) | No Treatment (96.2%, 51/53) | Oral | 1-2 hours before procedure | Favors Amoxicillin over control and Clindamycin, Favors Moxifloxacin over control and Clindamycin |
| Hall 1996 | 39 | Moderate | Bacteremia | Cefaclor (79%, 16/20) | Placebo (85%, 16/19) | Oral | 1 hour before procedure | No difference |
| | | | Bacteremia (viridans streptococci) | Cefaclor (79%, 16/20) | Placebo (50%, 10/19) | Oral | 1 hour before procedure | No difference |
| | | | Bacteremia (anaerobic) | Cefaclor (74%, 15/20) | Placebo (75%, 14/19) | Oral | 1 hour before procedure | No difference |
| Hall 1996 | 38 | Moderate | Bacteremia | Erythromycin (79%, 15/19) Clindamycin (84%, 16/19) | N/A | Oral | 1.5 hours before procedure | No difference |
| | 38 | | Bacteremia (viridans streptococci) | Erythromycin (79%, 15/19) Clindamycin (74%, 14/19) | N/A | Oral | 1.5 hours before procedure | No difference |
| Hall 1993 | 60 | Moderate | Bacteremia | Penicillin (90%, 18/20) Amoxicillin (85%, 17/20) | Placebo (95%, 19/20) | Oral | 1 hour before procedure | No difference |
| | 60 | | Bacteremia (viridans streptococci) | Penicillin (70% 14/20) Amoxicillin (55%, 11/20) | Placebo (70%, 14/20) | Oral | 1 hour before procedure | No difference |

| Study | N | Strength | Outcome (specific type) | Active Antibiotic (%, n/N) | Control (%, n/N) | Route of Administration | Time of Administration | Results |
|-----------------|-----|----------|----------------------------|-------------------------------------------------------------------------------------------------------------------------|-------------------------------------|----------------------------------------------------------|-------------------------------------------------------------------------------------------------------|-----------------------------------------------------------|
| | 60 | | Bacteremia (anaerobic) | Penicillin (85%, 17/20) Amoxicillin (75%, 15/20) | Placebo (85%, 17/20) | Oral | 1 hour before procedure | No difference |
| Head 1984 | 65 | Moderate | Bacteremia (anaerobic) | Penicillin V (20%) Metronidazole (52%) | Placebo (84%) | Oral | 1 hour before procedure | Favors Penicillin over Metronidazole |
| Jokinen 1970 | 152 | Moderate | Bacteremia | Penicillin (40%, 15/38) Penicillin with local prophylaxis (5%, 2/38) | No Treatment (87%, 66/76) | Oral (Penicillin), topical (local prophylaxis) | 45-90 minutes before procedure PLUS daily doses prior to operation day (penicillin) | Favors prophylaxis |
| Khairat 1966 | 242 | Moderate | Bacteremia | Erythromycin 250mg (37.5%, 6.16) Erythromycin 500mg (41%, 7/17) Erythromycin 1000mg (33%, 3/9) Tetracycline (3%, 3/100) | No Treatment (64%, 64/100) | Oral (Erythromycin), Intravenous (Tetracycline) | 1.5-4 hours before procedure (Erythromycin), 3 minutes before procedure (Tetracycline) | Favors prophylaxis |
| Maskell 1986 | 30 | Moderate | Bacteremia | Teicoplanin (60%, 6/10) Amoxicillin (40%, 4/10) | No Treatment (100%, 10/10) | Intramuscular (Teicoplanin), Oral (Amoxicillin) | 1 hour before procedure | Favors Amoxicillin over Teicoplanin over control |
| Roberts 1987 | 94 | Moderate | Bacteremia | Amoxicillin (2%, 1/47) | No Treatment (38%, 18/47) | Oral | 2 hours before procedure | Favors Amoxicillin |

| Study | N | Strength | Outcome (specific type) | Active Antibiotic (%, n/N) | Control (%, n/N) | Route of Administration | Time of Administration | Results |
|-------------------|-----|----------|------------------------------------------|--------------------------------------------------------------------------|--------------------------------------|--------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------|--------------------------------------------------------|
| Shanson 1987 | 120 | Moderate | Bacteremia (viridans streptococci) | Amoxicillin (25%, 10/40) Teicoplanin (2.5%, 1/40) | No Treatment (32.5%, 13/40) | Intramuscular (Amoxicillin), Intravenous bolus (Teicoplanin) | 25-40 minutes before procedure (Amoxicillin), 5-10 minutes before procedure (Teicoplanin) | Favors Teicoplanin over Amoxicillin over control |
| Shanson 1985 | 82 | Moderate | Bacteremia (streptococcal) | Erythromycin (15%, 6/40) | Placebo (43%, 18/42) | Oral | 1 hour before procedure | Favors Erythromycin |
| Shanson 1978 | 120 | Moderate | Bacteremia (streptococcal) | Penicillin V (12%, 5/40) Amoxicillin (5%, 2/40) | No Treatment (40%, 16/40) | Oral | 1 hour before procedure | Favors Penicillin and Amoxicillin over control |
| | 60 | | Bacteremia (anaerobic) | Penicillin V (20%, 4/20) Amoxicillin (15%, 3/20) | No Treatment (50%, 10/20) | Oral | 1 hour before procedure | Favors Penicillin and Amoxicillin over control |
| | 60 | | Bacteremia | Penicillin (20%, 4/20) Amoxicillin (25%, 5/20) | No Treatment (70%, 14/20) | Oral | 1 hour before procedure | Favors Penicillin and Amoxicillin over control |
| Vergis 29 2001 | | Moderate | Bacteremia | Oral Amoxicillin (10%, 1/10) Topical Amoxicillin (60%, 6/10) | No Treatment (89%, 8/9) | Oral, Rinse | 1 hour before procedure (Oral), 1 and 2 hours before procedure (Rinse) | Favors oral Amoxicillin over topical and control |
| | 36 | | Bacteremia (intent-to- treat) | Oral Amoxicillin (9%, 1/11) Topical Amoxicillin (53%, 8/15) | No Treatment (90%, 9/10) | Oral, Rinse | 1 hour before procedure (Oral), 1 and 2 hours before procedure (Rinse) | Favors oral Amoxicillin over topical and control |

| Study | N | Strength | Outcome (specific type) | Active Antibiotic (%, n/N) | Control (%, n/N) | Route of Administration | Time of Administration | Results |
|------------------|-----|----------|----------------------------|-------------------------------------------------------------------------------------|-------------------------------------|-------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------|
| DeVries 1972 | 200 | Low | Bacteremia | Lincomycin (8%, 2/25) Clindamycin Prolonged (0%, 0/25) Clindamycin Short (8%, 4/50) | No Treatment (49%, 49/100) | Oral | Daily for 3 days AND 1 hour before procedure (lincomycin), Daily for 3 days AND 2 hours before procedure (prolonged Clindamycin), 2 hours before procedure (short Clindamycin) | Favors Lincomycin and Clindamycin over control |
| Wahlmann 1999 | 60 | Low | Bacteremia | Cefuroxime (33%, 10/30) | Placebo (86%, 25/30) | Intravenous | 10 minutes before procedure | Favors Cefuroxime |

$Table\ 25\ Topical\ antimicrobials\ and\ tooth\ extraction\ bacteremia$

| Study | N | Strength | Outcome | Active Treatment (n/N, %) | Control (n/N, %) | Application | Results |
|------------------|-----|----------|------------|----------------------------------------------------------------------------------|------------------------------------|-------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Lockhart 1996 | 70 | High | Bacteremia | Chlorhexidine (31/37, 84%) | Placebo (31/33, 94%) | Mouth rinse | No difference |
| Casolari 1989 | 106 | Moderate | Bacteremia | Chlorhexidine OR Povidone-iodine (11/25, 44%) Penicillin antibiotic (12/25, 48%) | No treatment (38/56, 68%) | Irrigation of gingival crevice and retention of solution in mouth for a few minutes | Favors Chlorhexidine and Povidone-Iodine over control |

| Study | N | Strength | Outcome | Active Treatment (n/N, %) | Control (n/N, %) | Application | Results |
|--------------------|-----|----------|------------|---------------------------------------------------------------------------------------------------------------------------------|------------------------------------|-------------------------------------------------------------------------------------|----------------------------------------------------------------------------|
| Jokinen 1978 | 152 | Moderate | Bacteremia | Organic Iodine (21/38, 55%) Operative Field Isolation (13/38, 34%) Isolation+Iodine (12/38, 32%) Isolation+Chlorhex (5/38, 13%) | N/A | Mouth rinse | No difference |
| Macfarlane 1984 | 60 | Moderate | Bacteremia | Chlorhexidine (5/20, 25%) Povidone-Iodine (8/20, 40%) | Saline (16/20, 80%) | Irrigation of gingival crevice and retention of solution in mouth for a few minutes | Favors Povidone-Iodine over saline, Favors Chlorhexidine over saline |
| Rahn 1995 | 120 | Moderate | Bacteremia | Chlorhexidine (18/40, 45%) Povidone-Iodine (11/40, 27.5%) | Water (21/40, 52.5%) | Irrigation of gingival crevice and retention of solution in mouth for a few minutes | Favors Povidone-Iodine |
| Scopp 1971 | 64 | Moderate | Bacteremia | Povidone-Iodine (9/32, 28%) | Placebo (18/32, 56%) | Mouth rinse and irrigation of gingiva | Favors Povidone-Iodine rinse over placebo |
| Sweet 1978 | 100 | Moderate | Bacteremia | Chloramine-T rinse (12/25, 48%) Chloramine-T brush (12/25, 48%) Lugol's solution (20/25, 80%) | No Treatment (21/25, 84%) | Chloramine-T rinse and brushing, Irrigation with Lugol's solution | Favors Chloramine-T (brush or rinse) over control and Lugol's solution |
| Tomas 2007 | 106 | Moderate | Bacteremia | Chlorhexidine (42/53, 79%) | No Treatment (51/53, 96%) | Mouth filled | Favors Chlorhexidine |
| Cutcher 1971 | 100 | Low | Bacteremia | Phenolated (27/50, 54%) | No Treatment (39/50, 78%) | Mouth rinse | Favors Phenolated rinse |

| Study | N | Strength | Outcome | Active Treatment (n/N, %) | Control (n/N, %) | Application | Results |
|-----------------------|-----|----------|------------|--------------------------------------------------------------------------------------|---------------------------------------------------------------|-----------------------------------------------|---------------------------------------------------------------------------|
| Francis 1973 | 175 | Low | Bacteremia | Sodium perborate- ascorbic acid (9/50, 18%) | No Treatment (51/100, 51%) Saline (15/25, 60%) | Mouth rinse and irrigation of gingival sulcus | Favors Sodium perborate- ascorbic acid over saline and no treatment |
| Jones 1970 | 201 | Low | Bacteremia | Phenolated (12/67, 18%) | No Treatment (44/67, 66%) Saline (31/67, 46%) | Mouth rinse and irrigation of gingival sulcus | Favors Phenolated rinse over control, Favors saline rinse over control |
| Nasif 1977 | 120 | Low | Bacteremia | Hydrogen Peroxide (13/60, 22%) | No Treatment (26/60, 43%) | Irrigation of gingival sulcus | Favors Hydrogen Peroxide |
| Yamalik 1992 (941) | 80 | Low | Bacteremia | Povidone-Iodine (7/20, 35%) Hydrogen peroxide (10/20, 50%) Chlorhexidine (8/20, 40%) | No Treatment (14/20, 70%) | Irrigation of gingival sulcus | Favors Povidone-Iodine over control |

INDIRECT EVIDENCE: BACKGROUND MICROBIOLOGY FINDINGS

There was no direct evidence to explain the proposed association between bacteremia and orthopaedic implant infection, therefore we summarized the microbiological information pertaining to cases and rates of bacteremia and implant infection when available. Thirteen orthopaedic implant cohort studies were included that followed up on almost 13,000 implants, twelve of which provided detailed information on any infections that resulted over the course of the study. Approximately 53% of organisms responsible for the infections were *Staphylococcus* species. Overall rate of infection was approximately 1.5%. Of the studies that distinguished early from late infections we were able to calculate rates of 0.4% and 0.9% respectively. The definition of late infection varied greatly. In some cases it was not defined and in others it ranged from >3months to >18months. See Table 26 and 26 for details.

Eighteen studies addressing only infected orthopaedic implants were included and totaled approximately 1090 cases of implant infections. All eighteen studies provided detailed information on the infection. Approximately 64% of the infections were *Staphylococcus* species. Of the studies that distinguished early from late infections, 36.7% were early and 63.3% were late. The definition of late infection varied greatly. It ranged from >4 weeks to >1 year. See Table 27 and Figure 7 for details.

Incidence and prevalence of bacteremia varied greatly by procedure and study, as did the organism responsible for the bacteremia. Data is presented by procedure group. For studies that provided the necessary information, data were pooled and represent the proportion of bacteremic study participants that were found positive for the respective infecting organism. Microbiology data that was available from patients who received a form of prophylaxis was not included. No clear association between the organisms found in the prosthetic implant infections and bacteremia exists. However, the majority of the organisms found in implant infections are *Staphylococcus* and the majority of the organisms found as the cause of bacteremias are *Streptococcus*. See Figure 8 - Figure 34 for microbiological details on bacteremia.

RESULTS
Table 26 Orthopaedic implant cohort studies

| Author | Year | Implant | Study N | Infected N | % Population Infected | Early Infection | Late Infection | % Late infection | Late Infection Criteria |
|-----------|------|---------------|---------|---------------|-----------------------|--------------------|-------------------|------------------|-------------------------------|
| Ainscow | 1984 | Hip & Knee | 1112 | 22 | 2.0% | 11 | 11 | 1.0% | ≥3 months |
| Choong | 2007 | Hip | 819 | 14 | 1.7% | NA | NA | NA | NA |
| Goodman | 2006 | Hip | 17 | 1 | 5.9% | NA | NA | NA | NA |
| Hamilton | 2008 | Hip & Knee | 1993 | 29 | 1.5% | 11 | 18 | 0.9% | ≥3 months |
| Klenerman | 1991 | Hip & Knee | 174 | 2 | 1.1% | 0 | 2 | 1.1% | ≥3 months |
| Mont | 1999 | Hip | 109 | 1 | 0.9% | NA | NA | NA | NA |
| Petrie | 1998 | Knee | 1837 | 40 | 2.2% | NA | NA | NA | NA |
| Sancheti | 2009 | Knee | 297 | 1 | 0.3% | 0 | 1 | 0.3% | ≥7 months |
| Smith | 1997 | Hip | 66 | 2 | 3.0% | 0 | 2 | 3.0% | ≥18 months |
| Soultanis | 2003 | Spine | 60 | 5 | 8.3% | NA | 5 | 8.3% | ≥1 year |
| Uckay | 2009 | Hip & Knee | 6101 | 71 | 1.2% | 21 | 50 | 0.8% | ≥3 months |
| Wagner | 2000 | Hip | 78 | 1 | 1.3% | 0 | 1 | 1.3% | NA |
| Wimmer | 1998 | Spine | 110 | 1 | 0.9% | 0 | 1 | 0.9% | ≥17 months |

Figure 6 Organisms cultured from cohort studies

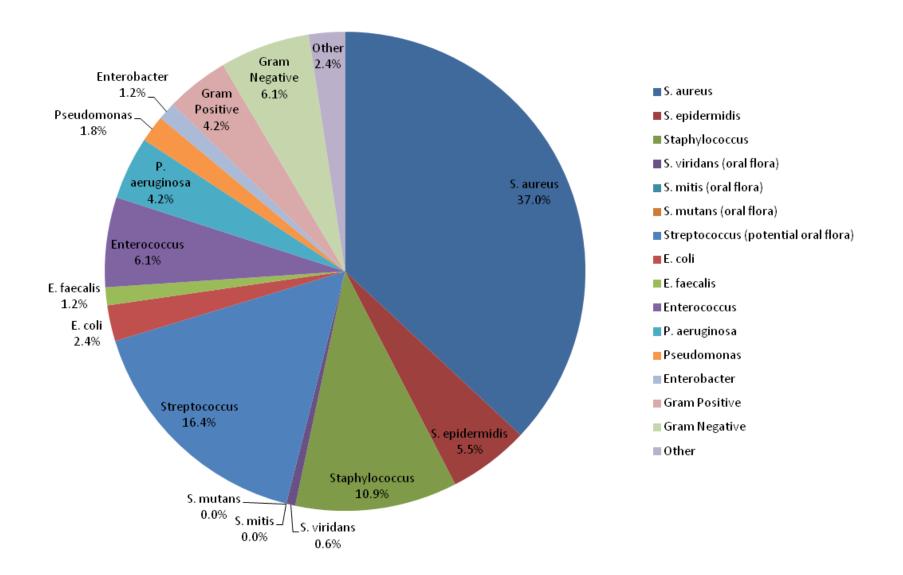
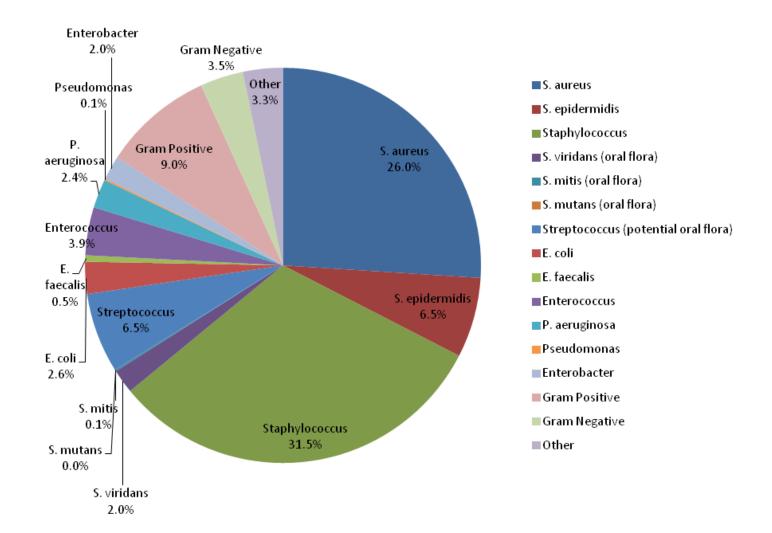
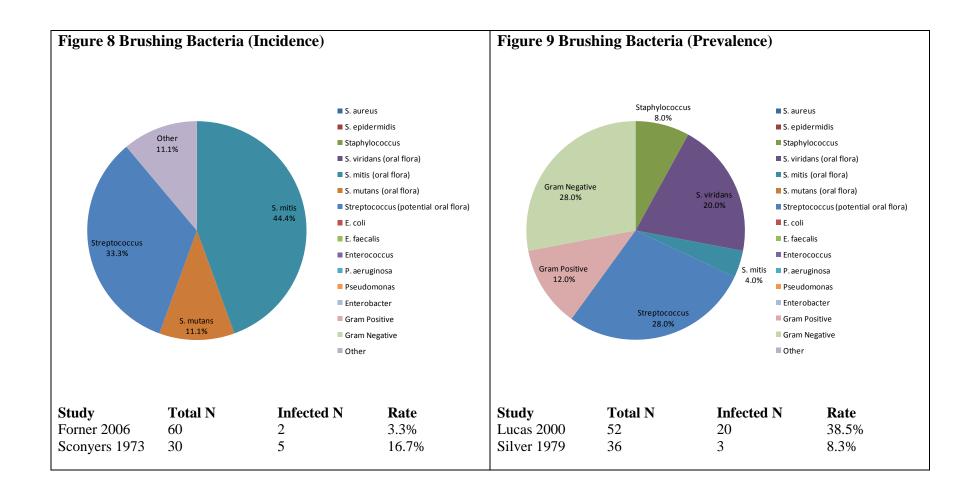


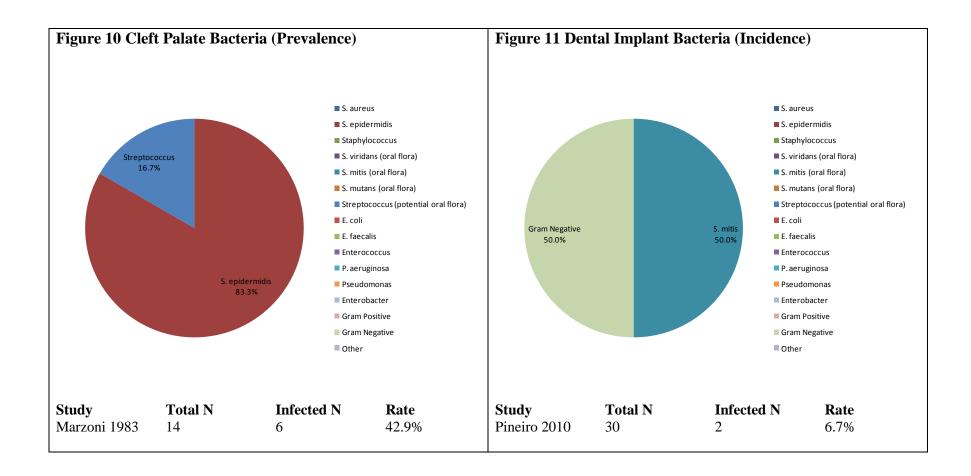
Table 27 Orthopaedic implant case series studies

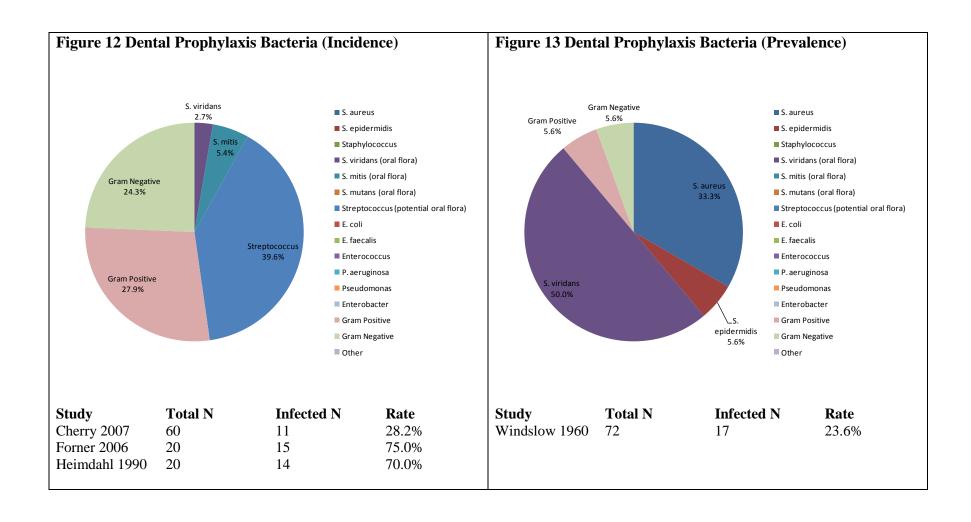
| Author | Year | Site | Study N | Infected N | Early Infection | Late Infection | % Late Infection | Late Infection Criteria |
|---------------------|------|----------------------------------|---------|------------|--------------------|-------------------|------------------|-------------------------------|
| Berbari | 2009 | Hip & Knee | 339 | 339 | 151 | 188 | 55.5% | ≥12 months |
| Chiu | 2007 | Knee | 40 | 40 | 10 | 30 | 75.0% | ≥4 weeks |
| Cordero- Ampuero | 2009 | Hip | 36 | 36 | 0 | 36 | 100.0% | ≥3 months |
| Cordero- Ampuero | 2007 | Hip & Knee | 40 | 40 | 0 | 40 | 100.0% | ≥3 months |
| Crockarell | 1998 | Hip | 42 | 42 | 19 | 23 | 54.8% | ≥1 month |
| Fink | 2008 | Knee | 40 | 40 | 0 | 40 | 100.0% | ≥2 months |
| Hoad-Reddick | 2005 | Knee | 59 | 59 | NA | NA | NA | NA |
| Insall | 1983 | Knee | 11 | 11 | 3 | 8 | 72.7% | ≥3 months |
| Jerosch | 2003 | Shoulder | 12 | 12 | 2 | 10 | 83.3% | 4 weeks |
| Mont | 1997 | Knee | 24 | 24 | 10 | 14 | 58.3% | ≥29 days |
| Munoz- Mahamud | 2011 | Hip, Knee, Other | 79 | 79 | 69 | 10 | 12.7% | ≥3 months |
| Rao | 2003 | Hip, Knee, Elbow | 36 | 36 | 13 | 23 | 63.9% | ≥1 year |
| Rodriguez | 2009 | Hip , Knee, Shoulder | 50 | 50 | 0 | 50 | 100.0% | ≥5 years |
| Soriano | 2007 | Knee, Hip, Shoulder, other | 85 | 85 | NA | NA | NA | NA |
| Soriano | 2006 | Hip & Knee | 47 | 47 | NA | NA | NA | NA |
| Waldman | 2000 | Knee | 16 | 16 | 4 | 12 | 75.0% | ≥6 months |
| Windsor | 1990 | Knee | 29 | 29 | NA | NA | NA | NA |
| Wroblewski | 1986 | Hip | 102 | 102 | NA | NA | NA | NA |

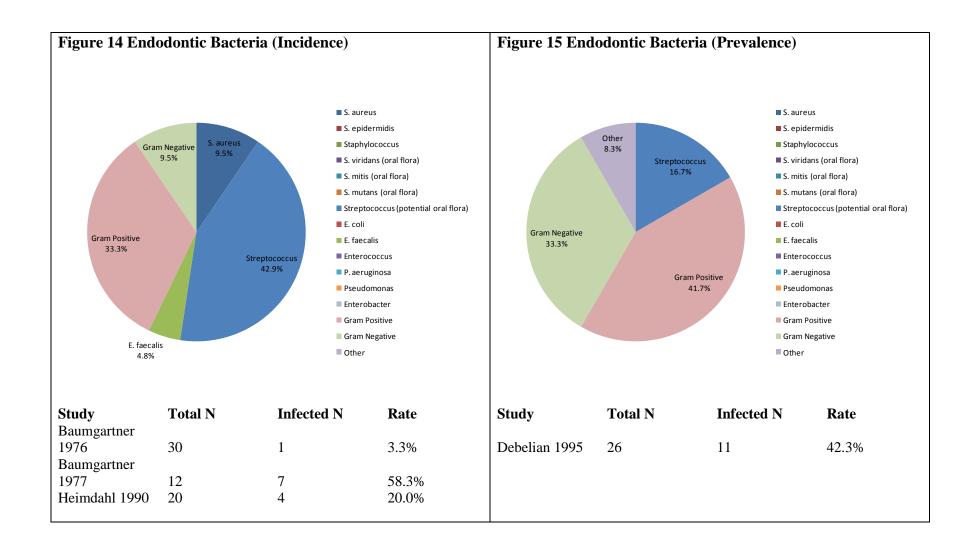
Figure 7 Organisms cultured from case series studies

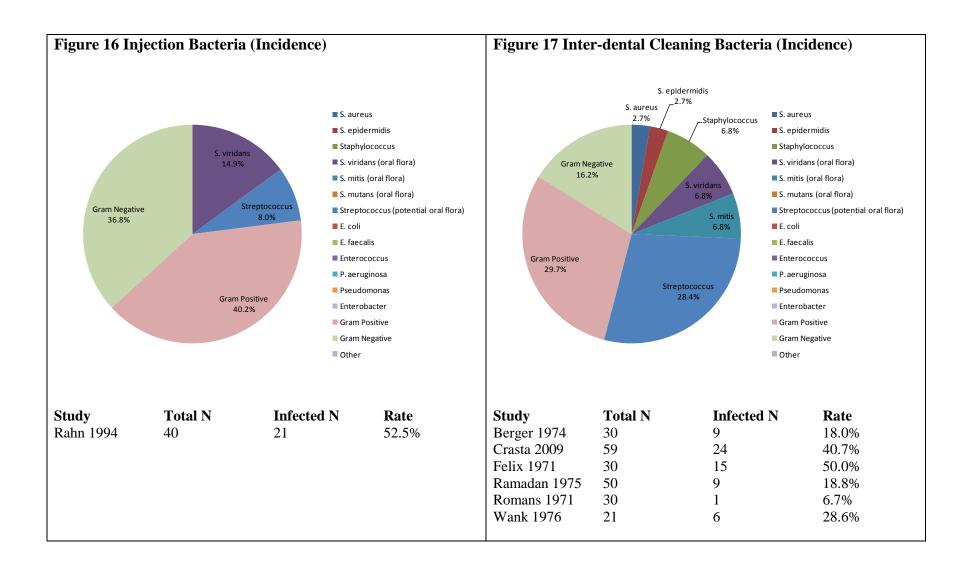


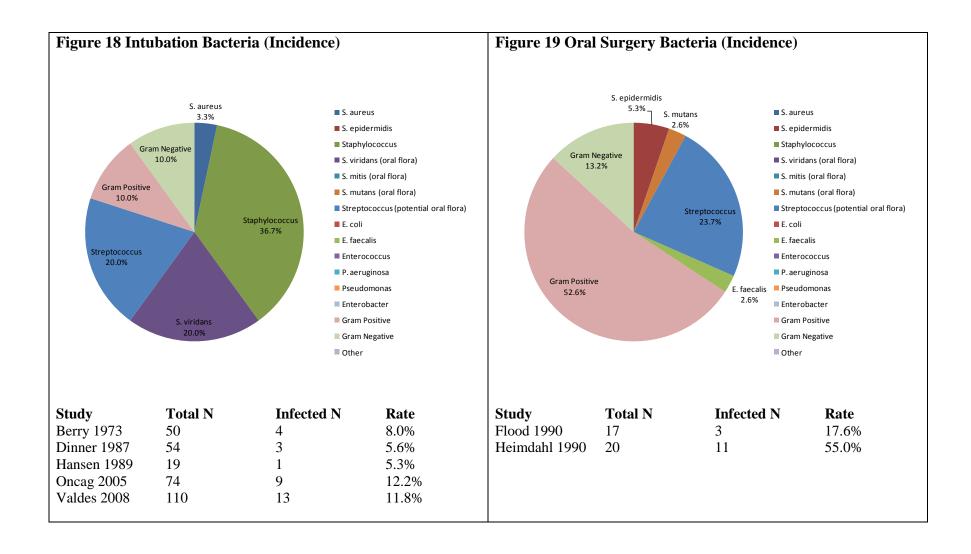


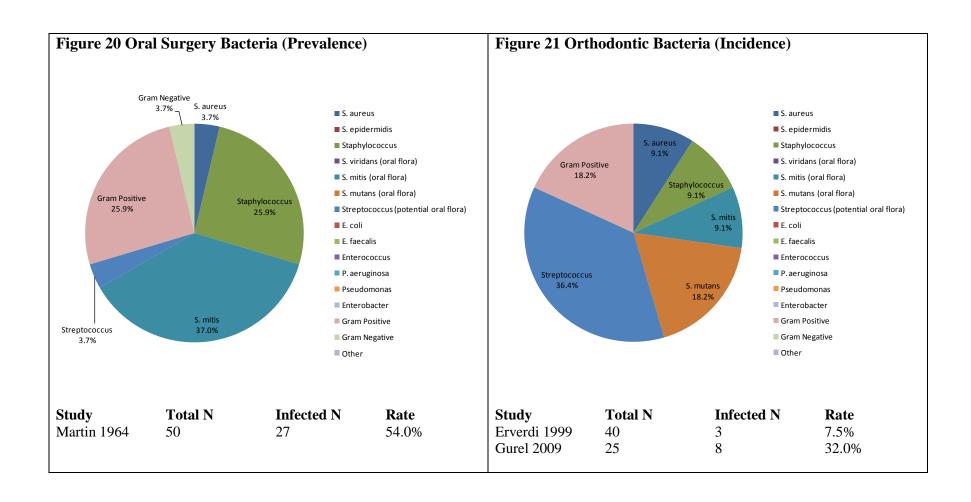


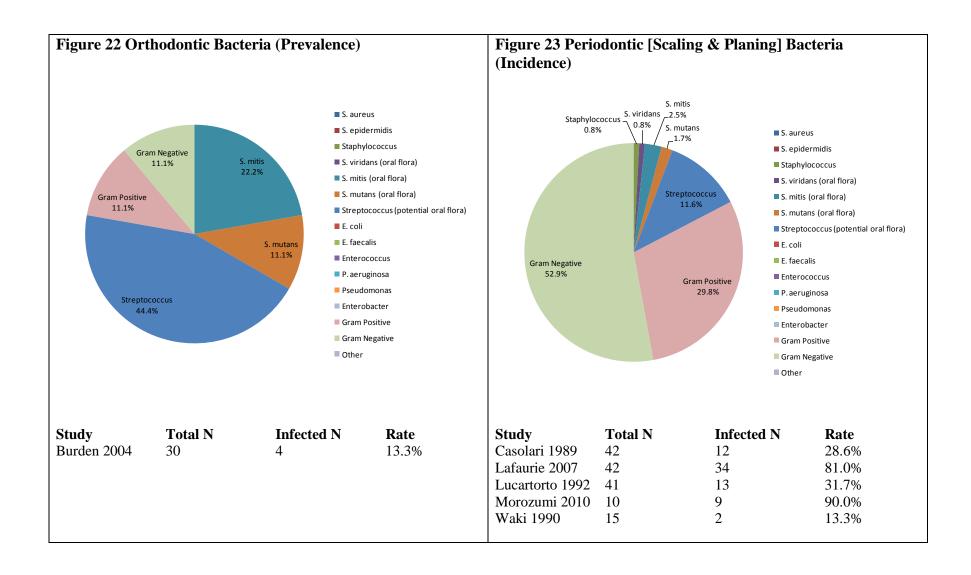


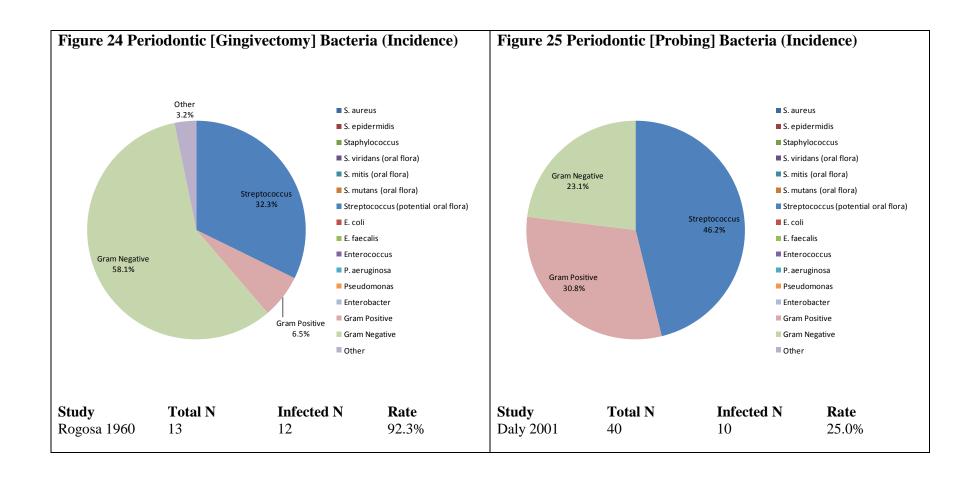


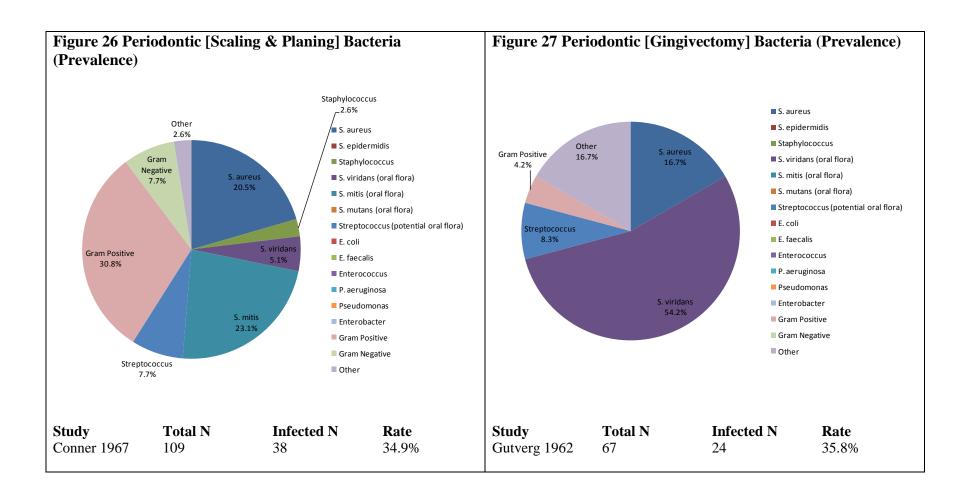


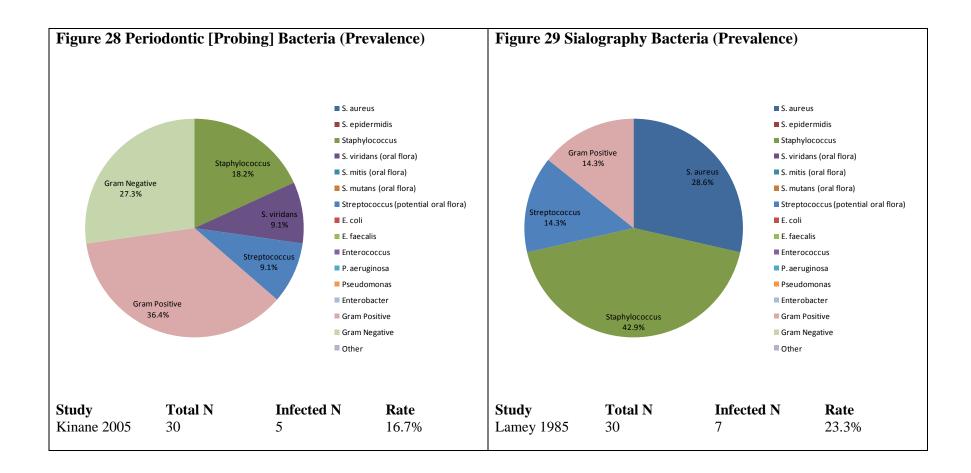


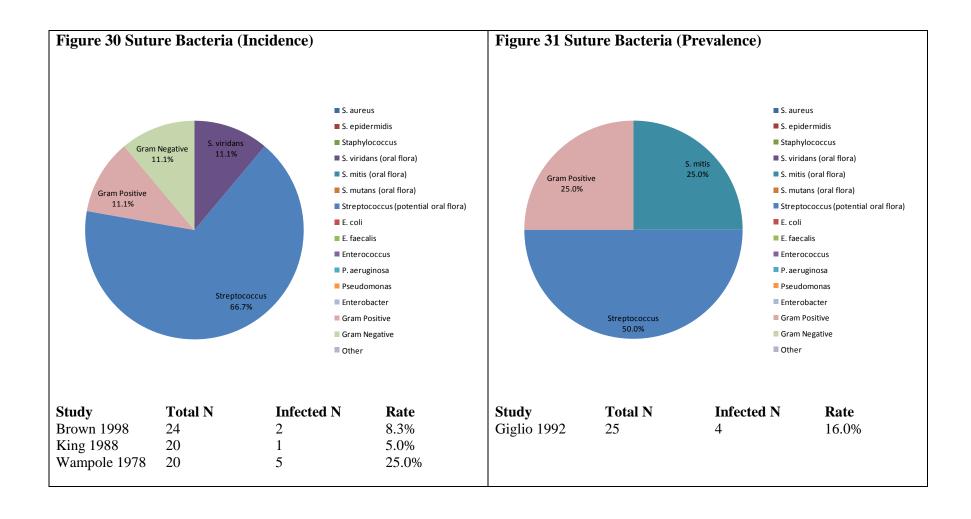


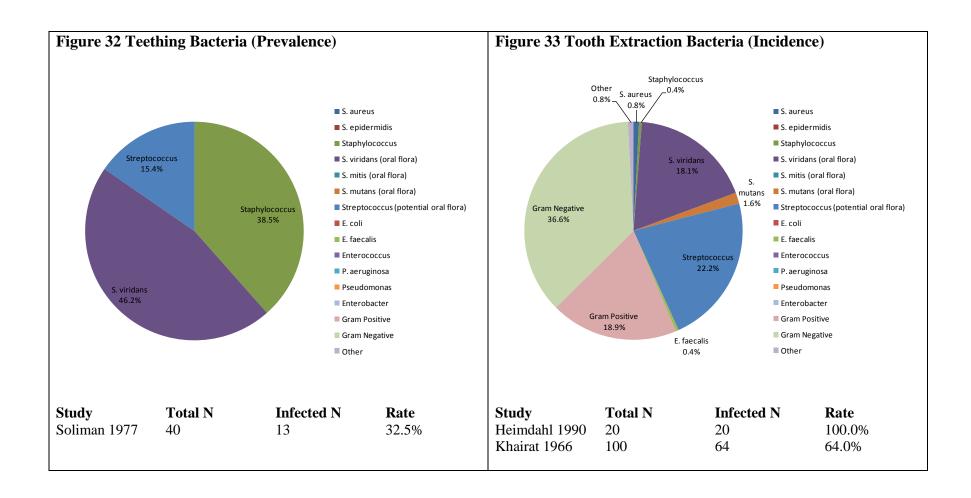


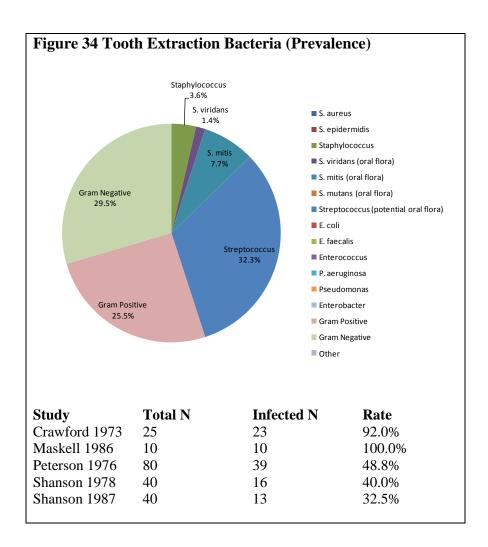












RECOMMENDATIONS

The following recommendations are not intended to stand alone. Treatment decisions should be made in light of all circumstances presented by the patient. Treatments and procedures applicable to the individual patient rely on mutual communication between patient, physician, dentist and other healthcare practitioners.

RECOMMENDATION 1

The practitioner might consider discontinuing the practice of routinely prescribing prophylactic antibiotics for patients with hip and knee prosthetic joint implants undergoing dental procedures.

Grade of Recommendation: Limited

RATIONALE

Moderate strength evidence finds that dental procedures are unrelated to implant infection and that antibiotic prophylaxis prior to dental procedures does not reduce the risk of subsequent implant infection. There is no direct evidence to support otherwise. High strength evidence suggests that antibiotic prophylaxis reduces the incidence of post-dental procedure related bacteremia, but there is no evidence that these bacteremias are related to prosthetic joint infections.

A single well-conducted case-control study provides direct evidence for this recommendation.³⁹ Case-control studies are appropriate to answer questions regarding risk factors or etiology. Study enrollment consisted of 339 patients with prosthetic hip or knee infections (cases) and 339 patients with hip or knee arthroplasties without infection (controls) hospitalized on an orthopaedic service during the same time period. The comparison between these groups was for differences in dental visits (exposure) in terms of high and low-risk dental procedures, with and without antibiotic prophylaxis. Results reported as odds ratios with 95% confidence interval, demonstrate no statistically significant differences between groups. Neither dental procedures nor antibiotic prophylaxis prior to dental procedures were associated with risk of prosthetic hip or knee infections. The authors performed a sample size calculation and withdrawals were low, minimizing attrition bias. The prospective nature of this study minimized recall bias. Additionally, blinding of the treatment group to those assessing outcomes limits detection bias.

Although this one study of direct evidence was of moderate quality, it did have limitations. The authors conducted covariate analysis on some subgroups of higher risk patients. The number of patients in these subgroups, however, was relatively small, and there is insufficient data to suggest that these patients are at higher risk of experiencing hematogenous infections.

There is high quality evidence that demonstrates the occurrence of bacteremia with dental procedures. Historically, there has been a suggestion that bacteremias can cause hematogenous seeding of total joint implants, both in the early postoperative period and for many years following implantation. It was felt that the most critical period was up to two years after joint placement. In addition, bacteremias may occur during normal daily activities such as chewing and tooth brushing. It is likely that these daily activities induce many more bacteremias than dental procedure associated bacteremias. While evidence supports a strong association between certain dental procedures and bacteremia, there is no evidence to demonstrate a direct link between dental procedure associated bacteremia and infection of prosthetic joints or other orthopaedic implants. Multiple studies of moderate and high quality evidence suggest that antibiotic prophylaxis decreases the risk of dental procedure associated bacteremias. However, dental procedure associated bacteremia is a surrogate outcome for prosthetic joint infection. Surrogate outcomes may or may not relate to a clinically relevant patient outcome. Of additional concern is a positive surrogate outcome (e.g. reduced bacteremias) that could mask a negative patient-centered outcome (e.g. implant infection).

This recommendation is limited to patients with hip and knee prostheses because the single study of direct evidence included only patients with these types of orthopaedic implants. There is no direct evidence that met our inclusion criteria for patients with other types of orthopaedic implants.

FINDINGS

As illustrated in Figure 1 there is varying quality of evidence that explains the purported association between dental procedures and orthopaedic implant infection. Only one moderate quality study of direct evidence was considered for this recommendation. The results of this study conclude that dental procedures are not risk factors for subsequent orthopaedic implant infection and furthermore that antibiotic prophylaxis prior to dental procedures does not reduce the risk of implant infection. However, multiple high quality studies of indirect evidence link oral procedures to bacteremia (see Figure 2 - Figure 5). Furthermore, multiple moderate quality studies of indirect evidence suggest that antibiotic prophylaxis prevents post-dental procedure bacteremia. Details of our analysis on antibiotic prophylaxis are presented in the results section below.

QUALITY AND APPLICABILITY

NETWORK META-ANALYSIS

Of the 21 studies included for this recommendation, 2 were of high quality and moderate applicability, 17 were of moderate quality and moderate applicability, and 2 were of low quality and moderate applicability. For details see Table 69 and Table 75 of Appendix XII.

RESULTS

NETWORK META-ANALYSIS

Twenty one studies that investigated the efficacy of antibiotic prophylaxis for prevention of dental procedure related bacteremia were included that compared antibiotics to controls or other antibiotics. Direct and indirect comparisons were drawn from network meta-analysis as diagramed in Figure 35. The network meta-analysis allowed us to compare treatments that were not in the same study. More detailed information on this method can be found in the "Statistical Methods" section of this guideline. Table 28, Table 29, and Table 30 summarize the results of these comparisons. Figure 36 and Figure 37 graphically depict the direct and indirect antibiotic comparisons vs. placebo/no treatment. Odds ratios were converted to number needed to treat (NNT) for a more clinically meaningful interpretation (see Table 31). Rankings of the antibiotics are presented in Table 32. These rankings do not indicate statistical significance.

The overall network model was consistent. See Table 59 in Appendix XI. Goodness-of-fit statistics are also presented in Appendix XI (see

Table 61). These results suggest that our model fits the available data. Individual study results can be found in Table 24. Individual study results that could not be meta-analyzed can be found in Table 67 in Appendix XI.

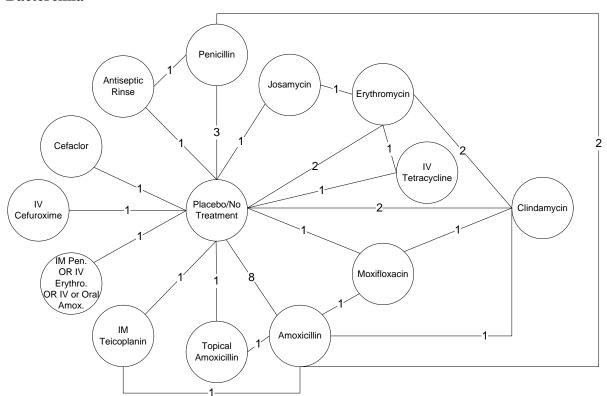


Figure 35 Network Diagram of Antibiotic Prophylaxis for the Prevention of Dental-related Bacteremia

Circles denote the treatments studied. Lines between circles denote treatment comparisons that are addressed by direct evidence. The numbers on these lines show the number of trials that compared the two treatments denoted in the circles.

Table 28 Direct Comparisons of Antibiotic Prophylaxes for the Prevention of Dentalrelated Bacteremia

| Comparison | Studies | Odds Ratio (95% CI) |
|------------------------------------------------|----------------|-------------------------|
| Amoxicillin vs. Placebo/No Treatment | 8 | 0.093* (0.041, 0.212) |
| Penicillin vs. Placebo/No Treatment | 3 | 0.282 (0.109, 0.731) |
| Erythromycin vs. Placebo/No Treatment | 2 | 0.512 (0.188, 1.396) |
| Clindamycin vs. Placebo/No Treatment | 2 | 0.121 (0.049, 0.299) |
| Josamycin vs. Placebo/No Treatment | 1 | 1.256 (0.334, 4.733) |
| Moxifloxacin vs. Placebo/No Treatment | 1 | 0.052 (0.011, 0.233) |
| Cefaclor vs. Placebo/No Treatment | 1 | 0.75 (0.144, 3.903) |
| IV Tetracycline vs. Placebo/No Treatment | 1 | 0.017 (0.005, 0.059) |
| IV Cefuroxime vs. Placebo/No Treatment | 1 | 0.1 (0.029, 0.34) |
| IM Teicoplanin vs. Placebo/No Treatment | 1 | 0.069 (0.003, 1.498) |
| Topical Amoxicillin vs. Placebo/No Treatment | 1 | 0.127 (0.013, 1.269) |
| Antiseptic Rinse vs. Placebo/No Treatment | 1 | 0.372 (0.141, 0.98) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. | | |
| Placebo/No Treatment | 1 | 0.318 (0.108, 0.935) |
| Penicillin vs. Amoxicillin | 2 | 0.997 (0.308, 3.229) |
| Clindamycin vs. Amoxicillin | 1 | 6.635 (2.654, 16.586) |
| Moxifloxacin vs. Amoxicillin | 1 | 1.523 (0.728, 3.189) |
| IM Teicoplanin vs. Amoxicillin | 1 | 2.25 (0.376, 13.465) |
| Topical Amoxicillin vs. Amoxicillin | 1 | 11.429 (1.155, 113.115) |
| Antiseptic Rinse vs. Penicillin | 1 | 0.851 (0.28, 2.591) |
| Clindamycin vs. Erythromycin | 2 | 0.7 (0.23, 2.129) |
| Josamycin vs. Erythromycin | 1 | 1.256 (0.334, 4.733) |
| IV Tetracycline vs. Erythromycin | 1 | 0.05 (0.014, 0.186) |
| Moxifloxacin vs. Clindamycin | 1 | 0.23 (0.092, 0.572) |

^{*}Heterogeneity (I²>50%)

Figure 36 Forest Plot of Direct Comparisons of Antibiotics vs. Placebo/No Treatment for the Prevention of Dental-related Bacteremia

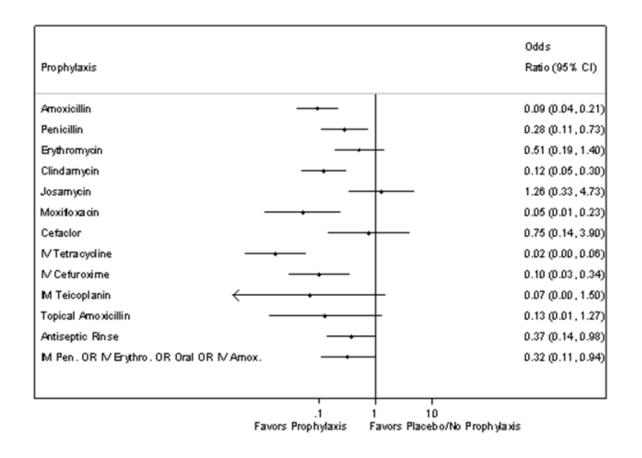


Table 29 Indirect (Network) Comparisons of Antibiotic Prophylaxes for the Prevention of Dental-related Bacteremia

| Comparison | Odds Ratio |
|-------------------------------------------------------------|----------------------|
| Amoxicillin vs. Placebo/No Treatment | 0.071 (0.026, 0.167) |
| Penicillin vs. Placebo/No Treatment | 0.176 (0.042, 0.685) |
| Erythromycin vs. Placebo/No Treatment | 0.426 (0.111, 1.593) |
| Clindamycin vs. Placebo/No Treatment | 0.235 (0.063, 0.842) |
| Josamycin vs. Placebo/No Treatment | 0.838 (0.093, 7.636) |
| Moxifloxacin vs. Placebo/No Treatment | 0.068 (0.009, 0.447) |
| Cefaclor vs. Placebo/No Treatment | 0.719 (0.047, 10.31) |
| IV Tetracycline vs. Placebo/No Treatment | 0.016 (0.001, 0.146) |
| IV Cefuroxime vs. Placebo/No Treatment | 0.089 (0.007, 0.952) |
| IM Teicoplanin vs. Placebo/No Treatment | 0.099 (0.006, 1.266) |
| Topical Amoxicillin vs. Placebo/No Treatment | 0.326 (0.028, 3.479) |
| Antiseptic Rinse vs. Placebo/No Treatment | 0.239 (0.028, 1.899) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Placebo/No | |
| Treatment | 0.301 (0.029, 3.064) |
| Penicillin vs. Amoxicillin | 2.478 (0.594, 11.06) |
| Erythromycin vs. Amoxicillin | 5.983 (1.345, 30.20) |
| Clindamycin vs. Amoxicillin | 3.303 (0.816, 14.87) |
| Josamycin vs. Amoxicillin | 11.77 (1.161, 134.5) |
| Moxifloxacin vs. Amoxicillin | 0.968 (0.143, 6.753) |
| Cefaclor vs. Amoxicillin | 10.10 (0.608, 180.9) |
| IV Tetracycline vs. Amoxicillin | 0.226 (0.019, 2.615) |
| IV Cefuroxime vs. Amoxicillin | 1.254 (0.099, 17.20) |
| IM Teicoplanin vs. Amoxicillin | 1.393 (0.101, 17.27) |
| Topical Amoxicillin vs. Amoxicillin | 4.585 (0.422, 52.35) |
| Antiseptic Rinse vs. Amoxicillin | 3.363 (0.375, 32.49) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Amoxicillin | 4.229 (0.372, 54.81) |
| Erythromycin vs. Penicillin | 2.413 (0.370, 16.34) |
| Clindamycin vs. Penicillin | 1.333 (0.214, 8.542) |
| Josamycin vs. Penicillin | 4.749 (0.368, 64.07) |
| Moxifloxacin vs. Penicillin | 0.390 (0.038, 3.811) |
| Cefaclor vs. Penicillin | 4.075 (0.198, 83.42) |
| IV Tetracycline vs. Penicillin | 0.091 (0.006, 1.290) |
| IV Cefuroxime vs. Penicillin | 0.506 (0.031, 8.068) |
| IM Teicoplanin vs. Penicillin | 0.562 (0.028, 9.679) |
| Topical Amoxicillin vs. Penicillin | 1.850 (0.119, 28.41) |
| Antiseptic Rinse vs. Penicillin | 1.357 (0.163, 11.47) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Penicillin | 1.706 (0.118, 26.84) |
| Clindamycin vs. Erythromycin | 0.552 (0.137, 2.181) |
| Josamycin vs. Erythromycin | 1.968 (0.222, 17.81) |
| Moxifloxacin vs. Erythromycin | 0.161 (0.016, 1.389) |
| Cefaclor vs. Erythromycin | 1.688 (0.082, 33.61) |
| IV Tetracycline vs. Erythromycin | 0.037 (0.003, 0.353) |
| IV Cefuroxime vs. Erythromycin | 0.209 (0.013, 3.158) |
| IM Teicoplanin vs. Erythromycin | 0.232 (0.010, 4.034) |
| Topical Amoxicillin vs. Erythromycin | 0.766 (0.048, 11.33) |
| Antiseptic Rinse vs. Erythromycin | 0.562 (0.045, 6.494) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Erythromycin | 0.707 (0.050, 10.30) |

| Comparison | Odds Ratio |
|--------------------------------------------------------------------|----------------------|
| Josamycin vs. Clindamycin | 3.564 (0.329, 39.80) |
| Moxifloxacin vs. Clindamycin | 0.293 (0.037, 2.090) |
| Cefaclor vs. Clindamycin | 3.055 (0.150, 60.82) |
| IV Tetracycline vs. Clindamycin | 0.068 (0.005, 0.779) |
| IV Cefuroxime vs. Clindamycin | 0.379 (0.024, 5.691) |
| IM Teicoplanin vs. Clindamycin | 0.421 (0.021, 6.972) |
| Topical Amoxicillin vs. Clindamycin | 1.388 (0.091, 19.96) |
| Antiseptic Rinse vs. Clindamycin | 1.018 (0.085, 11.63) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Clindamycin | 1.280 (0.091, 18.65) |
| Moxifloxacin vs. Josamycin | 0.082 (0.004, 1.368) |
| Cefaclor vs. Josamycin | 0.857 (0.025, 27.41) |
| IV Tetracycline vs. Josamycin | 0.019 (0.000, 0.373) |
| IV Cefuroxime vs. Josamycin | 0.106 (0.004, 2.789) |
| IM Teicoplanin vs. Josamycin | 0.118 (0.003, 3.333) |
| Topical Amoxicillin vs. Josamycin | 0.389 (0.014, 9.954) |
| Antiseptic Rinse vs. Josamycin | 0.285 (0.013, 5.870) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Josamycin | 0.359 (0.014, 8.688) |
| Cefaclor vs. Moxifloxacin | 10.43 (0.386, 301.8) |
| IV Tetracycline vs. Moxifloxacin | 0.233 (0.012, 4.379) |
| IV Cefuroxime vs. Moxifloxacin | 1.295 (0.061, 28.38) |
| IM Teicoplanin vs. Moxifloxacin | 1.438 (0.056, 31.72) |
| Topical Amoxicillin vs. Moxifloxacin | 4.735 (0.237, 100.0) |
| Antiseptic Rinse vs. Moxifloxacin | 3.472 (0.211, 60.64) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Moxifloxacin | 4.366 (0.230, 95.01) |
| IV Tetracycline vs. Cefaclor | 0.022 (0.000, 0.756) |
| IV Cefuroxime vs. Cefaclor | 0.124 (0.003, 4.517) |
| IM Teicoplanin vs. Cefaclor | 0.137 (0.002, 5.562) |
| Topical Amoxicillin vs. Cefaclor | 0.454 (0.012, 16.46) |
| Antiseptic Rinse vs. Cefaclor | 0.333 (0.011, 10.07) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Cefaclor | 0.418 (0.012, 14.52) |
| IV Cefuroxime vs. IV Tetracycline | 5.551 (0.208, 150.8) |
| IM Teicoplanin vs. IV Tetracycline | 6.165 (0.174, 189.2) |
| Topical Amoxicillin vs. IV Tetracycline | 20.28 (0.735, 561.7) |
| Antiseptic Rinse vs. IV Tetracycline | 14.87 (0.686, 330.9) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. IV Tetracycline | 18.70 (0.769, 484.9) |
| IM Teicoplanin vs. IV Cefuroxime | 1.110 (0.028, 35.26) |
| Topical Amoxicillin vs. IV Cefuroxime | 3.658 (0.116, 107.7) |
| Antiseptic Rinse vs. IV Cefuroxime | 2.682 (0.109, 62.67) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. IV Cefuroxime | 3.370 (0.120, 96.44) |
| Topical Amoxicillin vs. IM Teicoplanin | 3.293 (0.108, 117.0) |
| Antiseptic Rinse vs. IM Teicoplanin | 2.415 (0.093, 75.71) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. IM Teicoplanin | 3.034 (0.098, 116.1) |
| Antiseptic Rinse vs. Topical Amoxicillin | 0.733 (0.031, 17.77) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Topical Amoxicillin | 0.922 (0.034, 28.33) |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. vs. Antiseptic Rinse | 1.257 (0.056, 29.51) |

Table 30 Indirect (Network) Significant Comparisons of Antibiotic Prophylaxes for the Prevention of Dental-related Bacteremia

Comparison Amoxicillin favored over Placebo/No Treatment Penicillin favored over Placebo/No Treatment Clindamycin favored over Placebo/No Treatment Moxifloxacin favored over Placebo/No Treatment IV Tetracycline favored over Placebo/No Treatment IV Cefuroxime favored over Placebo/No Treatment Amoxicillin favored over Erythromycin Amoxicillin favored over Josamycin IV Tetracycline favored over Erythromycin

IV Tetracycline favored over Clindamycin IV Tetracycline favored over Josamycin IV Tetracycline favored over Cefaclor

Figure 37 Forest Plot of Indirect (Network) Comparisons of Antibiotics vs. Placebo/No Treatment for the Prevention of Dental-related Bacteremia

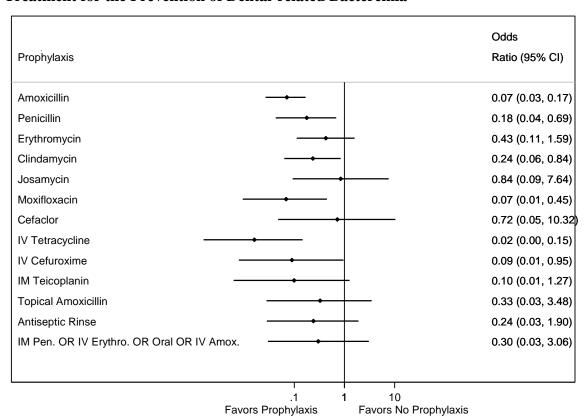


Table 31 Conversion of Odds Ratio from Figure 37 to Number Needed to Treat (NNT)

| Treatment | NNT |
|--------------------------------------------|------|
| Amoxicillin | 1.8 |
| Penicillin | 2.5 |
| Erythromycin | 5.0 |
| Clindamycin | 3.0 |
| Josamycin | 14.0 |
| Moxifloxacin | 1.9 |
| Cefaclor | 9.3 |
| IV Tetracycline | 1.5 |
| IV Cefuroxime | 2.1 |
| IM Teicoplanin | 2.2 |
| Topical Amoxicillin | 4.0 |
| Antiseptic Rinse | 3.2 |
| IM Pen. OR IV Erythro. OR Oral OR IV Amox. | 3.7 |

Table 32 Network Meta-Analysis Rankings of Antibiotic Prophylaxes for the Prevention of Dental-related Bacteremia

| | Rank 1 | | | | | | | | | | | | | 14 |
|--------------------|-----------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|---------|
| Prophylaxis | (Best) | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 | 12 | 13 | (Worst) |
| Placebo/No | | | | | | | | | | | | | | |
| Treatment | 0.0% | 0.0% | 0.0% | 0.0% | 0.0% | 0.0% | 0.0% | 0.1% | 0.5% | 2.8% | 10.8% | 27.3% | 38.4% | 20.2% |
| Amoxicillin | 1.7% | 14.2% | 28.9% | 28.5% | 16.4% | 6.8% | 2.5% | 0.7% | 0.2% | 0.0% | 0.0% | 0.0% | 0.0% | 0.0% |
| Penicillin | 0.3% | 1.9% | 4.2% | 8.1% | 13.4% | 17.3% | 16.8% | 14.1% | 10.4% | 6.8% | 4.0% | 1.7% | 0.6% | 0.2% |
| Erythromycin | 0.0% | 0.1% | 0.3% | 0.6% | 1.5% | 3.3% | 6.3% | 10.3% | 15.6% | 20.3% | 20.4% | 13.2% | 6.1% | 2.0% |
| Clindamycin | 0.0% | 0.4% | 1.5% | 3.7% | 8.1% | 12.9% | 16.7% | 18.3% | 16.7% | 11.6% | 6.3% | 2.6% | 0.9% | 0.3% |
| Josamycin | 0.1% | 0.3% | 0.6% | 1.0% | 1.5% | 2.3% | 3.2% | 4.4% | 6.0% | 8.6% | 11.5% | 14.5% | 17.4% | 28.5% |
| Moxifloxacin | 8.2% | 22.1% | 18.9% | 15.4% | 11.8% | 8.5% | 5.8% | 3.7% | 2.3% | 1.5% | 0.9% | 0.5% | 0.2% | 0.1% |
| Cefaclor | 0.5% | 1.4% | 1.7% | 2.1% | 2.8% | 3.7% | 4.5% | 5.5% | 6.7% | 8.1% | 9.9% | 11.1% | 13.8% | 28.0% |
| IV Tetracycline | 67.7% | 16.2% | 6.5% | 3.6% | 2.3% | 1.4% | 0.9% | 0.6% | 0.3% | 0.2% | 0.1% | 0.1% | 0.0% | 0.0% |
| IV Cefuroxime | 8.8% | 18.3% | 13.0% | 11.2% | 10.9% | 9.0% | 7.3% | 6.0% | 4.7% | 3.9% | 2.9% | 1.9% | 1.3% | 0.9% |
| IM Teicoplanin | 9.3% | 15.5% | 11.6% | 10.3% | 10.0% | 8.6% | 7.4% | 6.6% | 5.7% | 4.7% | 3.9% | 2.8% | 2.0% | 1.6% |
| Topical | | | | | | | | | | | | | | |
| Amoxicillin | 1.1% | 3.0% | 3.8% | 4.7% | 6.3% | 7.6% | 8.2% | 8.9% | 9.6% | 10.4% | 10.3% | 9.3% | 8.1% | 8.6% |
| Antiseptic Rinse | 1.1% | 3.4% | 4.6% | 5.8% | 8.2% | 10.3% | 11.3% | 11.1% | 11.0% | 10.4% | 8.8% | 6.3% | 4.3% | 3.2% |
| IM Pen. OR IV | | | | | | | | | | | | | | |
| Erythro. OR Oral | | | | | | | | | | | | | | |
| OR IV Amox. | 1.1% | 3.1% | 4.2% | 4.9% | 6.8% | 8.2% | 9.1% | 9.6% | 10.2% | 10.7% | 10.4% | 8.7% | 6.8% | 6.5% |

RECOMMENDATION 2

We are unable to recommend for or against the use of topical oral antimicrobials in patients with prosthetic joint implants or other orthopaedic implants undergoing dental procedures.

Grade of Recommendation: Inconclusive

RATIONALE

There is high quality evidence that demonstrates the occurrence of bacteremias with dental procedures. However, there is no evidence to demonstrate a direct link between dental procedure associated bacteremia and infection of prosthetic joints or other orthopaedic implants.

There is conflicting evidence regarding the effect of antimicrobial mouth rinse on the incidence of bacteremia associated dental procedures. One high quality study reports no difference in the incidence of bacteremia following antimicrobial mouth rinsing in patients undergoing dental extractions. Conversely, numerous studies suggest that topical antimicrobial prophylaxis decreases the incidence of dental procedure associated bacteremia. However, there is no evidence that application of antimicrobial mouth rinses before dental procedures prevents infection of prosthetic joints or other orthopaedic implants.

FINDINGS

As illustrated in Figure 1 there is varying quality of evidence that explains the relationship between dental procedures and orthopaedic implant infection. Only one moderate quality study of direct evidence was considered for this recommendation. The results of this study conclude that dental procedures are not risk factors for subsequent orthopaedic implant infection. However, multiple high quality studies of indirect evidence link oral procedures to bacteremia (see Figure 2 - Figure 5). Furthermore, multiple studies of indirect evidence of moderate strength suggest that topical antimicrobial prophylaxis prevents post-dental procedure bacteremia. Details of our analysis on topical antimicrobial prophylaxis are presented in the results section below.

QUALITY AND APPLICABILITY

NETWORK META-ANALYSIS

Of the 12 studies included for this recommendation, 1 was of high quality and moderate applicability, 7 were of moderate quality and moderate applicability, and 4 were of low quality and moderate applicability. For details see Table 69 and Table 87 of Appendix XII.

RESULTS

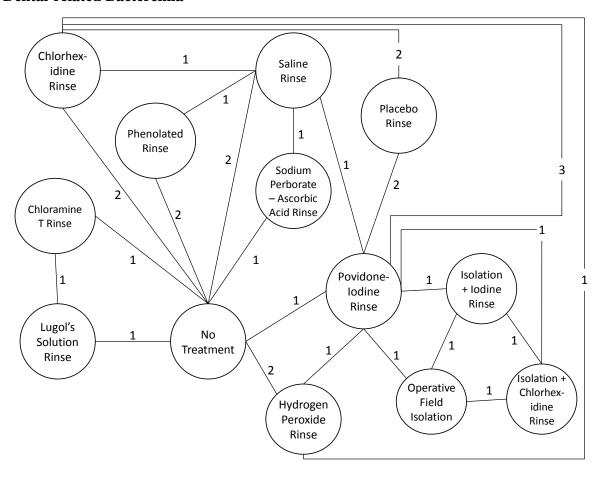
NETWORK META-ANALYSIS

Twelve studies were included that investigated the efficacy of topical antimicrobials for prevention of dental procedure related bacteremia. Direct and indirect comparisons were drawn from network meta-analysis as diagramed in Figure 38. Table 33, Table 34, and Table 35 summarize the results of these comparisons. Figure 39 and Figure 40 graphically depict the direct and indirect topical antimicrobial comparisons vs. no treatment. Odds ratios were converted to number needed to treat (NNT) for a more clinically meaningful interpretation (see Table 36). Rankings of the topicals are presented in Table 37. These rankings do not indicate statistical significance.

The overall network model was consistent. See Table 60 in Appendix XI. Goodness-of-fit statistics are also presented in Appendix XI (see

Table 61). These results suggest that our model fits the available data. Individual study results can be found in Table 25. Individual study results that could not be meta-analyzed can be found in Table 68 in Appendix XI.

Figure 38 Network Diagram of Topical Antimicrobial Prophylaxes for the Prevention of Dental-related Bacteremia



Circles denote the treatments studied. Lines between circles denote treatment comparisons that are addressed by direct evidence. The numbers on these lines show the number of trials that compared the two treatments denoted in the circles.

Table 33 Direct Comparisons of Topical Antimicrobial Prophylaxes for the Prevention of Dental-related Bacteremia

| Comparison | Studies | Odds Ratio (95% CI) |
|---------------------------------------------------------------|----------------|-----------------------|
| Saline Rinse vs. No Treatment | 2 | 0.778* (0.249, 2.429) |
| Chlorhexidine Rinse vs. No Treatment | 2 | 0.219 (0.08, 0.597) |
| Povidone-Iodine Rinse vs. No Treatment | 1 | 0.231 (0.061, 0.869) |
| Chloramine T Rinse/Brush vs. No Treatment | 1 | 0.176 (0.053, 0.586) |
| Lugol's Solution Rinse vs. No Treatment | 1 | 0.762 (0.179, 3.249) |
| Hydrogen Peroxide Rinse vs. No Treatment | 2 | 0.379 (0.192, 0.748) |
| Sodium Perborate-Ascorbic Acid Rinse vs. No Treatment | 1 | 0.211 (0.093, 0.479) |
| Phenolated Rinse vs. No Treatment | 2 | 0.192* (0.067, 0.545) |
| Chlorhexidine Rinse vs. Saline Rinse | 1 | 0.083 (0.019, 0.37) |
| Povidone-Iodine Rinse vs. Saline Rinse | 1 | 0.167 (0.041, 0.686) |
| Sodium Perborate-Ascorbic Acid Rinse vs. Saline Rinse | 1 | 0.146 (0.05, 0.43) |
| Phenolated Rinse vs. Saline Rinse | 1 | 0.253 (0.115, 0.557) |
| Povidone-Iodine Rinse vs. Chlorhexidine Rinse | 3 | 0.812 (0.352, 1.872) |
| Hydrogen Peroxide Rinse vs. Chlorhexidine Rinse | 1 | 1.5 (0.429, 5.248) |
| Chlorhexidine Rinse vs. Placebo Rinse | 2 | 0.623 (0.286, 1.356) |
| Hydrogen Peroxide Rinse vs. Povidone-Iodine Rinse | 1 | 1.857 (0.522, 6.612) |
| Povidone-Iodine Rinse vs. Placebo Rinse | 2 | 0.325 (0.162, 0.651) |
| Lugol's Solution Rinse vs. Chloramine T Rinse/Brush | 1 | 4.333 (1.405, 13.36) |
| Operative Field Isolation vs. Organic Iodine Rinse | 1 | 0.420 (0.166, 1.062) |
| Isolation + Iodine Rinse vs. Organic Iodine Rinse | 1 | 0.373 (0.146, 0.953) |
| Isolation + Chlorhexidine Rinse vs. Organic Iodine Rinse | 1 | 0.122 (0.039, 0.382) |
| Isolation + Iodine Rinse vs. Operative Field Isolation | 1 | 0.887 (0.340, 2.314) |
| Isolation + Chlorhexidine Rinse vs. Operative Field Isolation | 1 | 0.291 (0.091, 0.925) |
| Isolation + Chlorhexidine Rinse vs. Isolation + Iodine Rinse | 1 | 0.328 (0.102, 1.050) |

^{*}Heterogeneity (I² >50%)

Figure 39 Forest Plot of Direct Comparisons of Topical Antimicrobials vs. No Treatment for the Prevention of Dental-related Bacteremia

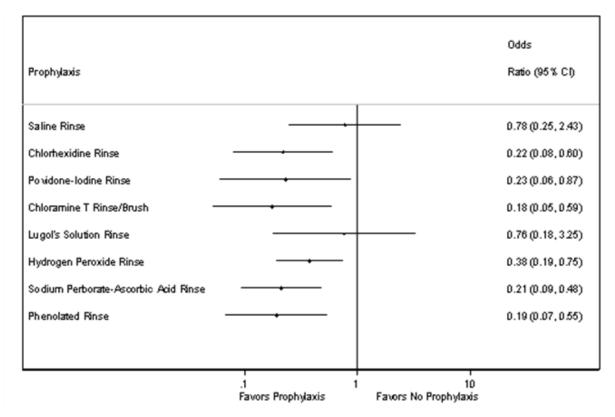


Table 34 Indirect (Network) Comparisons of Topical Antimicrobial Prophylaxes for the Prevention of Dental-related Bacteremia

| Comparison | Odds Ratio (95% CI) | | | |
|----------------------------------------------------------------|----------------------|--|--|--|
| Saline Rinse vs. No Treatment | 0.960 (0.402, 2.447) | | | |
| Chlorhexidine Rinse vs. No Treatment | 0.170 (0.059, 0.459) | | | |
| Povidone-Iodine Rinse vs. No Treatment | 0.143 (0.049, 0.412) | | | |
| Chloramine T Rinse/Brush vs. No Treatment | 0.158 (0.025, 0.891) | | | |
| Lugol's Solution Rinse vs. No Treatment | 0.740 (0.100, 5.269) | | | |
| Hydrogen Peroxide Rinse vs. No Treatment | 0.331 (0.108, 0.989) | | | |
| Sodium Perborate-Ascorbic Acid Rinse vs. No Treatment | 0.174 (0.041, 0.710) | | | |
| Phenolated Rinse vs. No Treatment | 0.216 (0.078, 0.612) | | | |
| Placebo Rinse vs. No Treatment | 0.400 (0.107, 1.485) | | | |
| Operative Field Isolation vs. No Treatment | 0.058 (0.008, 0.377) | | | |
| Isolation + Iodine Rinse vs. No Treatment | 0.051 (0.007, 0.331) | | | |
| Isolation + Chlorhexidine Rinse vs. No Treatment | 0.015 (0.002, 0.113) | | | |
| Chlorhexidine Rinse vs. Saline Rinse | 0.177 (0.051, 0.548) | | | |
| Povidone-Iodine Rinse vs. Saline Rinse | 0.149 (0.043, 0.475) | | | |
| Chloramine T Rinse/Brush vs. Saline Rinse | 0.165 (0.020, 1.147) | | | |
| Lugol's Solution Rinse vs. Saline Rinse | 0.771 (0.085, 6.586) | | | |
| Hydrogen Peroxide Rinse vs. Saline Rinse | 0.345 (0.084, 1.290) | | | |
| Sodium Perborate-Ascorbic Acid Rinse vs. Saline Rinse | 0.181 (0.040, 0.768) | | | |
| Phenolated Rinse vs. Saline Rinse | 0.225 (0.067, 0.718) | | | |
| Placebo Rinse vs. Saline Rinse | 0.417 (0.095, 1.714) | | | |
| Operative Field Isolation vs. Saline Rinse | 0.061 (0.008, 0.414) | | | |
| Isolation + Iodine Rinse vs. Saline Rinse | 0.054 (0.007, 0.364) | | | |
| Isolation + Chlorhexidine Rinse vs. Saline Rinse | 0.016 (0.001, 0.125) | | | |
| Povidone-Iodine Rinse vs. Chlorhexidine Rinse | 0.842 (0.349, 2.097) | | | |
| Chloramine T Rinse/Brush vs. Chlorhexidine Rinse | 0.932 (0.117, 7.113) | | | |
| Lugol's Solution Rinse vs. Chlorhexidine Rinse | 4.340 (0.474, 40.44) | | | |
| Hydrogen Peroxide Rinse vs. Chlorhexidine Rinse | 1.946 (0.553, 7.127) | | | |
| Sodium Perborate-Ascorbic Acid Rinse vs. Chlorhexidine Rinse | 1.023 (0.190, 5.708) | | | |
| Phenolated Rinse vs. Chlorhexidine Rinse | 1.267 (0.324, 5.328) | | | |
| Placebo Rinse vs. Chlorhexidine Rinse | 2.348 (0.854, 6.862) | | | |
| Operative Field Isolation vs. Chlorhexidine Rinse | 0.344 (0.057, 2.095) | | | |
| Isolation + Iodine Rinse vs. Chlorhexidine Rinse | 0.304 (0.050, 1.870) | | | |
| Isolation + Chlorhexidine Rinse vs. Chlorhexidine Rinse | 0.093 (0.013, 0.635) | | | |
| Chloramine T Rinse/Brush vs. Povidone-Iodine Rinse | 1.106 (0.136, 8.524) | | | |
| Lugol's Solution Rinse vs. Povidone-Iodine Rinse | 5.150 (0.552, 48.47) | | | |
| Hydrogen Peroxide Rinse vs. Povidone-Iodine Rinse | 2.308 (0.625, 8.364) | | | |
| Sodium Perborate-Ascorbic Acid Rinse vs. Povidone-Iodine Rinse | 1.214 (0.219, 6.739) | | | |
| Phenolated Rinse vs. Povidone-Iodine Rinse | 1.504 (0.370, 6.328) | | | |
| Placebo Rinse vs. Povidone-Iodine Rinse | 2.787 (1.037, 7.675) | | | |

| Comparison | Odds Ratio (95% CI) |
|-----------------------------------------------------------------------------|----------------------|
| Operative Field Isolation vs. Povidone-Iodine Rinse | 0.409 (0.084, 1.962) |
| Isolation + Iodine Rinse vs. Povidone-Iodine Rinse | 0.361 (0.074, 1.709) |
| Isolation + Chlorhexidine Rinse vs. Povidone-Iodine Rinse | 0.110 (0.019, 0.592) |
| Lugol's Solution Rinse vs. Chloramine T Rinse/Brush | 4.655 (0.872, 26.95) |
| Hydrogen Peroxide Rinse vs. Chloramine T Rinse/Brush | 2.087 (0.261, 17.56) |
| Sodium Perborate-Ascorbic Acid Rinse vs. Chloramine T Rinse/Brush | 1.098 (0.114, 11.16) |
| Phenolated Rinse vs. Chloramine T Rinse/Brush | 1.360 (0.182, 11.11) |
| Placebo Rinse vs. Chloramine T Rinse/Brush | 2.519 (0.287, 23.78) |
| Operative Field Isolation vs. Chloramine T Rinse/Brush | 0.369 (0.027, 4.997) |
| Isolation + Iodine Rinse vs. Chloramine T Rinse/Brush | 0.326 (0.024, 4.499) |
| Isolation + Chlorhexidine Rinse vs. Chloramine T Rinse/Brush | 0.100 (0.006, 1.473) |
| Hydrogen Peroxide Rinse vs. Lugol's Solution Rinse | 0.448 (0.047, 4.263) |
| Sodium Perborate-Ascorbic Acid Rinse vs. Lugol's Solution Rinse | 0.235 (0.020, 2.740) |
| Phenolated Rinse vs. Lugol's Solution Rinse | 0.292 (0.031, 2.784) |
| Placebo Rinse vs. Lugol's Solution Rinse | 0.540 (0.051, 5.766) |
| Operative Field Isolation vs. Lugol's Solution Rinse | 0.079 (0.005, 1.206) |
| Isolation + Iodine Rinse vs. Lugol's Solution Rinse | 0.070 (0.004, 1.070) |
| Isolation + Chlorhexidine Rinse vs. Lugol's Solution Rinse | 0.021 (0.001, 0.358) |
| Sodium Perborate-Ascorbic Acid Rinse vs. Hydrogen Peroxide Rinse | 0.526 (0.090, 3.089) |
| Phenolated Rinse vs. Hydrogen Peroxide Rinse | 0.651 (0.150, 2.980) |
| Placebo Rinse vs. Hydrogen Peroxide Rinse | 1.206 (0.267, 5.595) |
| Operative Field Isolation vs. Hydrogen Peroxide Rinse | 0.177 (0.023, 1.340) |
| Isolation + Iodine Rinse vs. Hydrogen Peroxide Rinse | 0.156 (0.020, 1.188) |
| Isolation + Chlorhexidine Rinse vs. Hydrogen Peroxide Rinse | 0.048 (0.005, 0.405) |
| Phenolated Rinse vs. Sodium Perborate-Ascorbic Acid Rinse | 1.238 (0.231, 6.868) |
| Placebo Rinse vs. Sodium Perborate-Ascorbic Acid Rinse | 2.293 (0.346, 15.19) |
| Operative Field Isolation vs. Sodium Perborate-Ascorbic Acid Rinse | 0.336 (0.032, 3.340) |
| Isolation + Iodine Rinse vs. Sodium Perborate-Ascorbic Acid Rinse | 0.297 (0.029, 2.956) |
| Isolation + Chlorhexidine Rinse vs. Sodium Perborate-Ascorbic Acid Rinse | 0.091 (0.007, 0.987) |
| Placebo Rinse vs. Phenolated Rinse | 1.852 (0.359, 9.290) |
| Operative Field Isolation vs. Phenolated Rinse | 0.271 (0.032, 2.172) |
| Isolation + Iodine Rinse vs. Phenolated Rinse | 0.240 (0.028, 1.919) |
| Isolation + Chlorhexidine Rinse vs. Phenolated Rinse | 0.073 (0.007, 0.648) |
| Operative Field Isolation vs. Placebo Rinse | 0.146 (0.022, 0.921) |
| Isolation + Iodine Rinse vs. Placebo Rinse | 0.129 (0.019, 0.819) |
| Isolation + Chlorhexidine Rinse vs. Placebo Rinse | 0.039 (0.005, 0.277) |
| Isolation + Iodine Rinse vs. Operative Field Isolation | 0.883 (0.181, 4.271) |
| Isolation + Chlorhexidine Rinse vs. Operative Field Isolation | 0.271 (0.046, 1.478) |
| Isolation + Chlorhexidine Rinse vs. Isolation + Iodine Rinse | 0.306 (0.052, 1.677) |

Table 35 Indirect (Network) Significant Comparisons of Topical Antimicrobial Prophylaxes for the Prevention of Dental-related Bacteremia

Comparison

Chlorhexidine Rinse vs. No Treatment Povidone-Iodine Rinse vs. No Treatment Chloramine T Rinse/Brush vs. No Treatment Hydrogen Peroxide Rinse vs. No Treatment Sodium Perborate-Ascorbic Acid Rinse vs. No Treatment Phenolated Rinse vs. No Treatment Operative Field Isolation vs. No Treatment Isolation + Iodine Rinse vs. No Treatment Isolation + Chlorhexidine Rinse vs. No Treatment Chlorhexidine Rinse vs. Saline Rinse Povidone-Iodine Rinse vs. Saline Rinse Sodium Perborate-Ascorbic Acid Rinse vs. Saline Rinse Phenolated Rinse vs. Saline Rinse Operative Field Isolation vs. Saline Rinse Isolation + Iodine Rinse vs. Saline Rinse Isolation + Chlorhexidine Rinse vs. Saline Rinse Isolation + Chlorhexidine Rinse vs. Chlorhexidine Rinse Povidone-Iodine Rinse vs. Placebo Rinse Isolation + Chlorhexidine Rinse vs. Povidone-Iodine Rinse Isolation + Chlorhexidine Rinse vs. Lugol's Solution Rinse Isolation + Chlorhexidine Rinse vs. Hydrogen Peroxide Rinse Isolation + Chlorhexidine Rinse vs. Sodium Perborate-Ascorbic Acid Rinse Isolation + Chlorhexidine Rinse vs. Phenolated Rinse Operative Field Isolation vs. Placebo Rinse Isolation + Iodine Rinse vs. Placebo Rinse Isolation + Chlorhexidine Rinse vs. Placebo Rinse

Figure 40 Forest Plot of Indirect (Network) Comparisons of Topical Antimicrobials vs. No Treatment for the Prevention of Dental-related Bacteremia

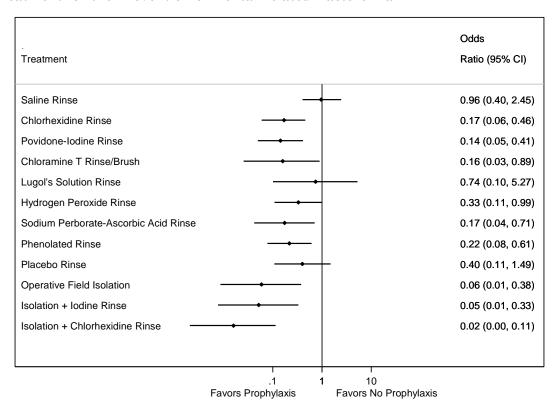


Table 36 Conversion of Odds Ratio from Figure 40 to Number Needed to Treat (NNT)

| Treatment | NNT |
|--------------------------------------|------|
| Saline Rinse | 70.0 |
| Chlorhexidine Rinse | 2.5 |
| Povidone-Iodine Rinse | 2.3 |
| Chloramine T Rinse/Brush | 2.5 |
| Lugol's Solution Rinse | 11.7 |
| Hydrogen Peroxide Rinse | 3.9 |
| Sodium Perborate-Ascorbic Acid Rinse | 2.5 |
| Phenolated Rinse | 2.8 |
| Placebo Rinse | n/a |
| Operative Field Isolation | 1.8 |
| Isolation + Iodine Rinse | 1.8 |
| Isolation + Chlorhexidine Rinse | 1.5 |

Table 37 Network Meta-Analysis Rankings of Topical Antimicrobial Prophylaxes for the Prevention of Dental-related Bacteremia

| | Rank 1 | | | | | | | | | | | | 13 |
|------------------------------------------------------|-----------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|---------|
| Prophylaxis | (Best) | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 | 12 | (Worst) |
| None | 0% | 0% | 0% | 0% | 0% | 0% | 0% | 0% | 1% | 3% | 20% | 45% | 32% |
| Saline Rinse Chlorhexidine | 0% | 0% | 0% | 0% | 0% | 0% | 0% | 1% | 2% | 7% | 25% | 34% | 31% |
| Rinse Povidone-Iodine | 0% | 1% | 4% | 11% | 20% | 23% | 19% | 13% | 6% | 2% | 0% | 0% | 0% |
| Rinse Chloramine T | 0% | 1% | 4% | 25% | 27% | 20% | 13% | 7% | 3% | 1% | 0% | 0% | 0% |
| Rinse/Brush Lugol's Solution | 3% | 8% | 9% | 16% | 11% | 10% | 10% | 11% | 10% | 8% | 2% | 1% | 0% |
| Rinse Hydrogen | 0% | 0% | 1% | 1% | 2% | 2% | 3% | 5% | 8% | 13% | 21% | 11% | 32% |
| Peroxide Rinse Sodium Perborate- Ascorbic Acid | 0% | 0% | 1% | 2% | 3% | 6% | 10% | 16% | 25% | 23% | 10% | 2% | 1% |
| Rinse | 1% | 5% | 7% | 14% | 13% | 13% | 13% | 12% | 10% | 6% | 3% | 1% | 0% |
| Phenolated Rinse | 0% | 2% | 3% | 8% | 11% | 15% | 18% | 18% | 14% | 8% | 3% | 0% | 0% |
| Placebo Rinse Operative Field | 0% | 0% | 0% | 1% | 1% | 4% | 8% | 13% | 21% | 28% | 16% | 5% | 3% |
| Isolation Isolation + Iodine | 4% | 31% | 37% | 12% | 6% | 4% | 2% | 2% | 1% | 1% | 0% | 0% | 0% |
| Rinse Isolation + Chlorhexidine | 6% | 41% | 31% | 10% | 4% | 3% | 2% | 1% | 1% | 0% | 0% | 0% | 0% |
| Rinse | 84% | 11% | 3% | 1% | 0% | 0% | 0% | 0% | 0% | 0% | 0% | 0% | 0% |

RECOMMENDATION 3

In the absence of reliable evidence linking poor oral health to prosthetic joint infection, it is the opinion of the work group that patients with prosthetic joint implants or other orthopaedic implants maintain appropriate oral hygiene.

Grade of Recommendation: Consensus

RATIONALE

The lack of evidence relating oral bacteremias to prosthetic joint or other orthopaedic implant infections is the basis for the consensus rationale for this recommendation.

Oral hygiene measures are low cost, provide potential benefit, are consistent with current practice, and are in accordance with good oral health.

There is evidence of the relationship of oral microflora to bacteremia. This bacteremia may be associated with poor oral hygiene. This implies that improvement of oral hygiene (or maintenance of good oral hygiene) may be beneficial in reducing bacteremias.

FINDINGS

No direct evidence was found in support of Recommendation 3. However, several prognostic studies of indirect evidence are included that explore whether or not oral health status can predict development of bacteremia after dental procedures. These low strength studies address oral health indicators as potential risk factors for developing bacteremia as a result of undergoing a dental procedure. The results of these studies are inconsistent and summarized in the results section below. See Table 38 for a summary of study results and Table 39 - Table 47 for more detail. By optimizing oral health, one could eliminate these potential risk factors and therefore reduce their risk of developing a dental procedure related bacteremia.

QUALITY AND APPLICABILITY

Refer to Table 88 - Table 96 in Appendix XII.

Table 38 Summary of Oral Health Related Risk Factor (Proportion of studies that reported significant results)

| | Brushing | Chewing | Dental Prophylaxis | Inter- dental Cleaning | Intubation | Oral Surgery | Periodontic | Restorative | Tooth Extraction |
|--------------------------|-----------|---------|-----------------------|------------------------------|----------------|-----------------|-------------|-------------|---------------------|
| Risk Factor | | | 1 | | ts (%Significa | nt, n/N) | | | |
| # Teeth Present | | | | | | 0%, 0/1 | 0%, 0/1 | | |
| Abscess | | | | | | 0%, 0/1 | | | 0%, 0/2 |
| Apical Lucency | 0%, 0/1 | | | | | | | | 0%, 0/1 |
| Calculus Index/Score | 100%, 1/1 | | | | | | | | 0%, 1/1 |
| Caries | 0%, 0/1 | | | | | | | 0%, 0/1 | 0%, 0/1 |
| Caries Depth | 0%, 0/1 | | | | | | | 0%, 0/1 | 0%, 0/1 |
| Clinical Attachment Loss | | | | 0%, 0/1 | | | | | |
| Gingival Index/Score | 25%, 1/4 | | 100%, 1/1 | 0%, 0/1 | | 50%, 1/2 | | 100%, 1/1 | 67%, 2/3 |
| Gingival Size | | | | | | | | 0%, 0/1 | |
| Gingivitis | 0%, 0/1 | 0%, 0/1 | 0%, 0/1 | | | | | | |
| Infected Tooth | | | | | | 100%, 1/1 | | | |
| Odontogenic Disease | | | | | | | | | 0%, 0/1 |
| Oral Health Status | | | | | 0%, 0/1 | 0%, 0/1 | | | 50%, 1/2 |
| Periodontal Diagnosis | | | 0%, 0/1 | | | | | | 0%, 0/1 |
| Periodontitis | 0%, 0/1 | 0%, 0/1 | 100%, 1/1 | 0%, 0/1 | | | 50%, 1/2 | 0%, 0/1 | |
| Plaque Index/Score | 67%, 2/3 | | 50%, 1/2 | 0%, 0/1 | | 0%, 0/1 | | | 0%, 0/3 |
| Probing Depth | | | 0%, 0/2 | 0%, 0/1 | | | 0%, 0/1 | | 33%, 1/3 |
| Probing Depth Mean | 0%, 0/1 | | | , | | | 100%, 1/1 | | 0%, 0/1 |
| Radiolucency | | | | 1 | | | | 0%, 0/1 | |
| Recession | | | 0%, 0/1 | | | | | , | |

| | Brushing | Chewing | Dental Prophylaxis | Inter- dental Cleaning | Intubation | Oral Surgery | Periodontic | Restorative | Tooth Extraction |
|----------------|----------|---------|-----------------------|------------------------------|------------|-----------------|-------------|-------------|---------------------|
| Suppuration | | | | | | | | 0%, 0/1 | |
| Swelling | | | | | | | | 0%, 0/1 | |
| Tooth Mobility | 0%, 0/1 | | | | | | | | 0%, 0/1 |

Table 39 Oral Health Related Risk Factors for Brushing Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|-------------|----------|-----|----------------------|-------------------------|---------------|--------------|
| | | | | Bacteremia | | p<0.01 @ 30s |
| Ashare | | | Linear | (Bacterial Load @ | | & 5m, NS @ |
| 2009 | Low | 48 | regression | 30s, 5m, 15m) | Plaque Index | 15m |
| Bhanji | | | logistic | | | OR 1.05, |
| 2002 | Low | 50 | regression | Bacteremia | Plaque Score | p=0.44 |
| | | | | Bacteremia | | |
| | | | | (Infective | | |
| Lockhart | | | logistic | Endocarditis | Mean plaque | OR 2.53 |
| 2009 | Low | 98 | regression | related bacteria) | score | p=.010 |
| | | | | Bacteremia | | |
| | | | | (Infective | | |
| Lockhart | | | logistic | Endocarditis | Plaque score | OR 3.78 |
| 2009 | Low | 98 | regression | related bacteria) | ≥ 2 | p=.008 |
| | | | | Bacteremia | | |
| Ashare | | | Linear | (Bacterial Load @ | Gingival | NS for all |
| 2009 | Low | 48 | regression | 30s, 5m, 15m) | Index | time points |
| Bhanji | | | | | Gingival | |
| 2002 | Low | 50 | chi square | Bacteremia | Score | p=0.96 |
| | | | | Bacteremia | | |
| | | | | (Infective | | |
| Lockhart | | 0.0 | logistic | Endocarditis | Mean gingival | OR 1.62 |
| 2009 | Low | 98 | regression | related bacteria) | score | p=.203 |
| | | | | Bacteremia | | |
| v 11 . | | | | (Infective | G: 1 | OD 1 61 |
| Lockhart | _ | 00 | logistic | Endocarditis | Gingival | OR 1.61 |
| 2009 | Low | 98 | regression | related bacteria) | $score \ge 2$ | p=.335 |
| Silver | T | 06 | Cuiti and matin that | D (| Gingival | |
| 1977 | Low | 96 | Critical ratio test | Bacteremia | Index | p<.01 |
| Forner 2006 | T | 20 | Fishers exact test | Bacteremia | Cim minuisia | NC |
| Forner | Low | 20 | Fishers exact test | Dacterenna | Gingivitis | NS |
| 2006 | Low | 20 | Eighorg areat tost | Bacteremia | Periodontitis | NS |
| 2000 | Low | 20 | Fishers exact test | Bacteremia | remodolitius | NS |
| | | | | | | |
| Lockhart | | | logistic | (Infective Endocarditis | Mean calculus | OR 1.77 |
| 2009 | Low | 98 | regression | related bacteria) | score | p=.048 |
| 2007 | LOW | 70 | 10810331011 | Bacteremia | SCOIC | p0+0 |
| | | | | (Infective | | |
| Lockhart | | | logistic | Endocarditis | Calculus | OR 4.43 |
| 2009 | Low | 98 | regression | related bacteria) | $score \ge 2$ | p=.004 |
| 2007 | LOW | 70 | 10510331011 | Bacteremia | 50010 = 2 | p=.00+ |
| | | | | (Infective | | |
| Lockhart | | | logistic | Endocarditis | Bleeding with | OR 0.89 |
| 2009 | Low | 98 | • | related bacteria) | toothbrushing | p=.810 |
| 2009 | Low | 98 | regression | related bacteria) | toothbrushing | p=.810 |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|----------|----------|----|------------------|-------------------|----------------|---------|
| | | | | Bacteremia | | |
| | | | | (Infective | Bleeding type | |
| Lockhart | | | logistic | Endocarditis | with | OR 7.96 |
| 2009 | Low | 98 | regression | related bacteria) | toothbrushing | p=.015 |
| | | | | Bacteremia | | |
| | | | | (Infective | | |
| Lockhart | | | logistic | Endocarditis | Mean probing | OR 1.02 |
| 2009 | Low | 98 | regression | related bacteria) | depth | p=.918 |
| | | | | Bacteremia | | |
| | | | | (Infective | | |
| Lockhart | | | logistic | Endocarditis | Tooth | OR 1.93 |
| 2009 | Low | 98 | regression | related bacteria) | mobility score | p=.200 |
| | | | | Bacteremia | | |
| | | | | (Infective | | |
| Lockhart | | | logistic | Endocarditis | | OR 4.40 |
| 2009 | Low | 98 | regression | related bacteria) | Dental caries | p=.165 |
| | | | | Bacteremia | | |
| | | | | (Infective | | |
| Lockhart | | | logistic | Endocarditis | Depth of | OR 0.43 |
| 2009 | Low | 98 | regression | related bacteria) | dental caries | p=.155 |
| | | | | Bacteremia | | |
| | | | | (Infective | | |
| Lockhart | | | logistic | Endocarditis | Apical | OR 2.37 |
| 2009 | Low | 98 | regression | related bacteria) | lucency | p=.086 |
| | | | | Bacteremia | | |
| | | | | (Infective | Apical | |
| Lockhart | | | logistic | Endocarditis | lucency size | OR 0.87 |
| 2009 | Low | 98 | regression | related bacteria) | (mm) | p=.647 |

Table 40 Oral Health Related Risk Factors for Chewing Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|----|---------------------|------------|---------------|---------|
| Forner | | | | | | |
| 2006 | Very Low | 20 | Fisher's exact test | Bacteremia | Periodontitis | NS |
| Forner | | | | | | |
| 2006 | Very Low | 20 | Fisher's exact test | Bacteremia | Gingivitis | NS |

Table 41 Oral Health Related Risk Factors for Dental Prophylaxis Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|----|------------------|-------------|---------------|--------------|
| Cherry | | | Logistic | | | |
| 2007 | Low | 60 | regression | Bacteremia | Plaque Index | NS |
| | | | Spearman's | | | |
| Forner | | | correlation | Bacteremia | | |
| 2006 | Low | 20 | coefficients | (magnitude) | Plaque Index | 0.41 p=.0117 |
| | | | | | Modified | |
| | | | | | papilla, | |
| | | | | | margin, | |
| Cherry | | | Logistic | | attached | |
| 2007 | Low | 60 | regression | Bacteremia | gingiva index | NS |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|----|--------------------|-------------|---------------|--------------|
| Cherry | | | Logistic | | | |
| 2007 | Low | 60 | regression | Bacteremia | Probing depth | NS |
| Cherry | | | Logistic | | | |
| 2007 | Low | 60 | regression | Bacteremia | Recession | NS |
| Cherry | | | Logistic | | Bleeding on | |
| 2007 | Low | 60 | regression | Bacteremia | scaling | NS |
| Forner | | | | | | |
| 2006 | Low | 20 | Fishers exact test | Bacteremia | Periodontitis | p<.001 |
| | | | Spearman's | | | |
| Forner | | | correlation | Bacteremia | Periodontal | |
| 2006 | Low | 20 | coefficients | (magnitude) | diagnosis | NS |
| Forner | | | | | | |
| 2006 | Low | 20 | Fishers exact test | Bacteremia | Gingivitis | NS |
| | | | Spearman's | | | |
| Forner | | | correlation | Bacteremia | Gingival | |
| 2006 | Low | 20 | coefficients | (magnitude) | Index | 0.53 p<.0001 |
| | | | Spearman's | | | |
| Forner | | | correlation | Bacteremia | Bleeding on | |
| 2006 | Low | 20 | coefficients | (magnitude) | probing | 0.45 p=.0089 |
| | | | Spearman's | | Probing | |
| Forner | | | correlation | Bacteremia | pocket depth | |
| 2006 | Low | 20 | coefficients | (magnitude) | >5 | NS |
| | | | Spearman's | | | |
| Forner | | | correlation | Bacteremia | Pocket sum | |
| 2006 | Low | 20 | coefficients | (magnitude) | score | NS |

Table 42 Oral Health Related Risk Factors for Inter-dental Cleaning Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|----|------------------|------------|---------------|------------|
| | | | Spearman's | | | |
| Crasta | | | correlation | | | |
| 2009 | Low | 60 | coefficients | Bacteremia | Periodontitis | 0.17 p=.2 |
| | | | Spearman's | | | |
| Crasta | | | correlation | | Gingival | |
| 2009 | Low | 60 | coefficients | Bacteremia | Index | 0.22 p=.09 |
| | | | Spearman's | | | |
| Crasta | | | correlation | | | |
| 2009 | Low | 60 | coefficients | Bacteremia | Plaque Index | 0.07 p=.6 |
| | | | Spearman's | | % of sites | |
| Crasta | | | correlation | | bleeding on | |
| 2009 | Low | 60 | coefficients | Bacteremia | flossing | 0.17 p=.2 |
| | | | Spearman's | | # sites | |
| Crasta | | | correlation | | bleeding on | |
| 2009 | Low | 60 | coefficients | Bacteremia | flossing | 0.17 p=.2 |
| | | | Spearman's | | % of sites | |
| Crasta | | | correlation | | bleeding on | |
| 2009 | Low | 60 | coefficients | Bacteremia | probing | 0.16 p=.2 |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|----|------------------|------------|----------------|------------|
| | | | Spearman's | | | |
| Crasta | | | correlation | | | |
| 2009 | Low | 60 | coefficients | Bacteremia | Pocket depth | 0.09 p=.5 |
| | | | Spearman's | | Clinical | |
| Crasta | | | correlation | | attachment | |
| 2009 | Low | 60 | coefficients | Bacteremia | loss | 0.06 p=.6 |
| | | | Spearman's | | | |
| Crasta | | | correlation | | Self-reported | |
| 2009 | Low | 60 | coefficients | Bacteremia | daily flossing | -0.12 p=.4 |

Table 43 Oral Health Related Risk Factors for Intubation Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|-----|------------------|------------|-------------|---------|
| Valdes | | | Logistic | | Oral health | |
| 2008 | Low | 110 | regression | Bacteremia | status | NS |

Table 44 Oral Health Related Risk Factors for Oral Surgery Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|---------------|----------|-----|------------------|----------------|-------------------------|---------|
| Roberts | | | | | | 1.878 |
| 1998 | Low | 154 | chi-square | Bacteremia | Abscess | p=.1706 |
| | | | Pearson | | | |
| Roberts | | | correlation | | | |
| 1998 | Low | 154 | coefficient | Bacteremia | Age | 0.29 |
| Tomas | | | | | | |
| 2008 | Low | 100 | not reported | Bacteremia | Age | NS |
| | | | Scheffe's | | | |
| Roberts | | | multiple | | | |
| 1998 | Low | 154 | comparison | Bacteremia | Plaque Index | p=.47 |
| | | | Scheffe's | | | |
| Roberts | | | multiple | | Gingival | |
| 1998 | Low | 154 | comparison | Bacteremia | Index | p<.03 |
| Takai | | | | | Gingival | |
| 2005 | Low | 237 | chi-square | Bacteremia | Index | NS |
| | | | Scheffe's | | | |
| Roberts | | 1 | multiple | | Bleeding | |
| 1998 | Low | 154 | comparison | Bacteremia | Index | p<.04 |
| | | | | | Oral hygiene | |
| Takai | | 225 | | | index | |
| 2005 | Low | 237 | chi-square | Bacteremia | simplified | NS |
| Takai | T | 227 | -1-1 | Da eta na nais | # 4 41 | NIC |
| 2005 Takai | Low | 237 | chi-square | Bacteremia | # teeth present | NS |
| | T and | 237 | ala: a aura na | Dagtamamia | Dia ad ia aa | NC |
| 2005 | Low | 237 | chi-square | Bacteremia | Blood loss Infection in | NS |
| | | | | | extracted tooth | |
| | | | | | (periodontitis, | |
| | | | | | periapical | |
| Takai | | | | | infection, and | |
| 2005 | Low | 237 | chi-square | Bacteremia | pericoronitis) | p<.01 |
| 2003 | LOW | 231 | ciii-square | Dactelellla | pericoronius) | h<.01 |

Table 45 Oral Health Related Risk Factors for Periodontic Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|--------|----------|----|------------------|------------|---------------|------------|
| Daly | | | | | Periodontitis | |
| 1997 | Low | 30 | chi-square | Bacteremia | severity | p=.9 |
| | | | | | | OR 5.993 |
| Daly | | | logistic | | | CI=1.081- |
| 2001 | Low | 40 | regression | Bacteremia | Periodontitis | 33.215 |
| Daly | | | | | Bleeding on | |
| 1997 | Low | 30 | t-test | Bacteremia | probing | p=.3 |
| | | | | | | OR 1.025 |
| Daly | | | logistic | | Bleeding on | CI=1.004- |
| 2001 | Low | 40 | regression | Bacteremia | probing | 1.047 |
| | | | | | | OR 1.0 |
| Daly | | | logistic | | | CI=.845- |
| 2001 | Low | 40 | regression | Bacteremia | # of teeth | 1.185 |
| | | | | | | OR 1.006 |
| Daly | | | logistic | | Total probing | CI=.999- |
| 2001 | Low | 40 | regression | Bacteremia | depth | 1.013 |
| | | | | | | OR 3.154 |
| Daly | | | logistic | | | CI=.603- |
| 2001 | Low | 40 | regression | Bacteremia | Plaque index | 16.514 |
| | | | | | Mean probing | OR 1.444 |
| Daly | | | logistic | | depth per | CI=.1.055- |
| 2001 | Low | 40 | regression | Bacteremia | tooth | 1.977 |

Table 46 Oral Health Related Risk Factors for Restorative Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|---------|----------|----|------------------|------------|----------------|---------|
| Brennan | | | chi-square or | | Gingival | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | Score (0-3) | p=.01 |
| Brennan | | | chi-square or | | Gingival Size | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | (0-3) | NS |
| | | | | | Periodontal | |
| | | | | | disease with | |
| Brennan | | | chi-square or | | probing | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | >3mm | NS |
| Brennan | | | chi-square or | | Mixed | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | Dentition | p=.08 |
| Brennan | | | chi-square or | | | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | Caries Present | NS |
| Brennan | | | chi-square or | | Depth of | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | caries (0-3) | NS |
| Brennan | | | chi-square or | | Periapical | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | radiolucency | NS |
| | | | | | Size | |
| Brennan | | | chi-square or | | radiolucency | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | (mm) | NS |
| Brennan | | | chi-square or | | | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | Swelling | NS |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|---------|----------|----|------------------|------------|-------------|---------|
| Brennan | | | chi-square or | | | |
| 2007 | Very Low | 51 | fisher's exact | Bacteremia | Suppuration | NS |

Table 47 Oral Health Related Risk Factors for Extraction Bacteremia

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|----------|----------|-----|-------------------------|----------------|-----------------------|----------|
| | | | logistic | | | OR 3.704 |
| Barbosa | | | regression | | Oral health | (1.929- |
| 2010 | Low | 210 | (univariate) | Bacteremia 30s | status | 7.109) |
| | | | logistic | | | OR 2.047 |
| Barbosa | | | regression | | Oral health | (1.138- |
| 2010 | Low | 210 | (univariate) | Bacteremia 15m | status | 3.683) |
| Wahlmann | | | logistic | | | |
| 1999 | Low | 59 | regression | Bacteremia | Oral Hygiene | NS |
| Wahlmann | | | logistic | | Periodontal | |
| 1999 | Low | 59 | regression | Bacteremia | status | NS |
| Coulter | | | | | | |
| 1990 | Low | 58 | chi-square | Bacteremia | Plaque Index | NS |
| Lockhart | | | logistic | | Mean plaque | OR 0.74 |
| 2009 | Low | 96 | regression | Bacteremia | score | p=.236 |
| Lockhart | | | logistic | | | OR 0.90 |
| 2009 | Low | 96 | regression | Bacteremia | Plaque score ≥ 2 | p=.811 |
| | | | Scheffe's | | | 1 |
| Roberts | | | multiple | | | |
| 1998 | Low | 154 | comparison | Bacteremia | Plaque Index | p=.47 |
| Coulter | | | • | | 1 | |
| 1990 | Low | 58 | chi-square | Bacteremia | Gingival Index | NS |
| Lockhart | | | logistic | | Mean gingival | OR 0.71 |
| 2009 | Low | 96 | regression | Bacteremia | score | p=.217 |
| Lockhart | | | logistic | | Gingival score | OR 0.76 |
| 2009 | Low | 96 | regression | Bacteremia | ≥ 2 | p=.518 |
| | | | Scheffe's | | | 1 |
| Roberts | | | multiple | | | |
| 1998 | Low | 154 | comparison | Bacteremia | Gingival Index | p<.03 |
| Coulter | | | • | | <u> </u> | 1 |
| 1990 | Low | 58 | Fisher's | Bacteremia | Abscess | p=0.2088 |
| Roberts | | | | | | 1.878 |
| 1998 | Low | 154 | chi-square | Bacteremia | Abscess | p=.1706 |
| Lockhart | | | chi-square or | | Odontogenic | • |
| 1996 | Low | 70 | Fisher's exact | Bacteremia | disease severity | NS |
| Lockhart | | | logistic | | Mean calculus | OR 0.93 |
| 2009 | Low | 96 | regression | Bacteremia | score | p=.724 |
| Lockhart | | | logistic | | Calculus score | OR 0.82 |
| 2009 | Low | 96 | regression | Bacteremia | ≥ 2 | p=.715 |
| Lockhart | | | logistic | | Bleeding with | |
| 2009 | Low | 96 | regression | Bacteremia | toothbrushing | NA |
| | | | | | Bleeding type | |
| Lockhart | | | logistic | | with | |
| 2009 | Low | 96 | regression | Bacteremia | toothbrushing | NA |

| Author | Strength | N | Statistical Test | Outcome | Risk Factor | Results |
|------------|----------|-----|------------------|------------|-----------------|--------------|
| Okabe 1995 | Low | 183 | Mann-Whitney | Bacteremia | Blood loss (ml) | 3997.5 p<.05 |
| | | | Scheffe's | | | |
| Roberts | | | multiple | | | |
| 1998 | Low | 154 | comparison | Bacteremia | Bleeding Index | p<.04 |
| Lockhart | | | logistic | | Mean probing | OR 0.95 |
| 2009 | Low | 96 | regression | Bacteremia | depth | p=.735 |
| Lockhart | | | logistic | | Tooth mobility | OR 1.01 |
| 2009 | Low | 96 | regression | Bacteremia | score | p=.978 |
| Lockhart | | | logistic | | | OR 1.66 |
| 2009 | Low | 96 | regression | Bacteremia | Dental caries | p=.452 |
| Lockhart | | | logistic | | Depth of dental | OR 0.21 |
| 2009 | Low | 96 | regression | Bacteremia | caries | p=.156 |
| Lockhart | | | logistic | | | OR 0.86 |
| 2009 | Low | 96 | regression | Bacteremia | Apical lucency | p=.724 |
| Lockhart | | | logistic | | Apical lucency | OR 1.00 |
| 2009 | Low | 96 | regression | Bacteremia | size (mm) | p=.995 |

FUTURE RESEARCH

The grades of recommendation in this clinical practice guideline are "limited" at best due to the lack of evidence in some cases and conflicting evidence in others. Only one study that met the inclusion criteria attempted to define the relationship, or lack thereof, between dental procedures and subsequent orthopaedic implant infections and preventive effect of antibiotic prophylaxis. Relying on evidence that does not directly address this relationship to inform clinical practice assumes that bacteremia is an appropriate surrogate outcome for prosthetic joint or other orthopaedic implant associated infection. Additional research is necessary to assess the pros and cons of providing antimicrobial prophylaxis for this study population and definitively determine if there is an association between dental procedures and orthopaedic implant infections.

Specifically:

- Prospective, controlled (ideally randomized), adequately powered trials investigating the effect of prophylactic interventions with the primary outcome of implant infection.
- Research investigating the relationship between bacteremias and orthopaedic implant infection.
- Research determining if bacteremia is an appropriate surrogate outcome for orthopaedic implant infection
- Research investigating the relationship between oral health and orthopaedic implant infection
- Cost-benefit analysis of antimicrobial prophylaxis for patients with orthopaedic implants undergoing dental procedures

APPENDICES

APPENDIX I WORK GROUP

Prevention of Orthopaedic Implant Infection in Patients Undergoing Dental Procedures Clinical Practice Guideline Work Group

<u>American Academy of Orthopaedic Surgeons</u> William Watters, III, MD, Co-Chair

Bone and Joint Clinic of Houston 6624 Fannin Street, #2600 Houston, TX 77030

American Dental Association

Michael P. Rethman, DDS, MS, Co-Chair

47-140 Heno Place Kaneohe, HI 96744

American Academy of Orthopaedic Surgeons

Richard Parker Evans, MD

Professor and Margaret Sue Neal Endowed Chair of Orthopaedic Surgery University of Missouri- Kansas City School of Medicine 2301 Holmes Street Kansas City, MO 64108

American Academy of Orthopaedic Surgeons

Calin Moucha, MD

Associate Chief, Joint Replacement Surgery Mount Sinai Medical Center Assistant Professor Leni & Peter W. May Department of Orthopaedic Surgery Mount Sinai School of Medicine 5 E. 98th Street, Box 1188, 7th Floor New York, NY 10029

American Academy of Orthopaedic Surgeons Richard J. O'Donnell, MD

Chief, UCSF Orthopaedic Oncology Service UCSF Sarcoma Program UCSF Helen Diller Family Comprehensive Cancer Center 1600 Divisadero Street, 4th Floor San Francisco, CA 94115

American Academy of Orthopaedic Surgeons & Congress of Neurological Surgeons

Paul A. Anderson, MD

Professor Department of Orthopedics & Rehabilitation University of Wisconsin K4/735 600 Highland Avenue Madison WI 53792

American Dental Association

Elliot Abt, DDS

4709 Golf Road, Suite 1005 Skokie, IL 60076

American Dental Association

Harry C. Futrell, DMD

330 W 23rd Street, Suite J Panama City, FL 32405

American Dental Association

Stephen O. Glenn, DDS

5319 S Lewis Avenue, Suite 222 Tulsa, OK 74105-6543

American Dental Association

John Hellstein, DDS, MS

The University of Iowa, College of Dentistry Department of Oral Pathology, Radiology and Medicine DSB S356 Iowa City, IA 52242

American Association of Hip and Knee Surgeons

David Kolessar, MD

Geisinger Wyoming Valley Medical Center 1000 East Mountain Boulevard Valley Medical Building Wilkes-Barre, PA 18711

American Association of Neurological Surgeons/Congress of Neurological Surgeons

John E. O'Toole, MD

Assistant Professor of Neurosurgery Rush University Medical Center 1725 W. Harrison Street, Suite 970 Chicago, IL 60612

<u>American Association of Oral and Maxillofacial</u> <u>Surgeons</u>

Mark J. Steinberg DDS, MD

1240 Meadow Road, Suite 300 Northbrook, IL 60062

College of American Pathologist

Karen C. Carroll MD, FCAP

Johns Hopkins Hospital

Department of Pathology-Microbiology Division

600 N Wolfe Street Meyer B1-193 Baltimore, MD 21287

Knee Society

Kevin Garvin, MD

University of Nebraska Medical Center Creighton/Nebraska Health Fund Department of Orthopaedic Surgery 981080 Nebraska Medical Center Omaha, Nebraska 68198-1080

Musculoskeletal Infection Society

Douglas R. Osmon, MD

200 1st Street SW Rochester, MN 55905

Scoliosis Research Society

Anthony Rinella, MD

Illinois Spine & Scoliosis Center 12701 West 143rd Street, Suite 110 Homer Glen, Illinois 60491

Society for Healthcare Epidemiology of America

Angela Hewlett, MD, MS

Assistant Professor, Section of Infectious Diseases University of Nebraska Medical Center 985400 Nebraska Medical Center Omaha, Nebraska 68198

Guidelines Oversight Chair

Michael J. Goldberg, MD

Children's Hospital and Regional Medical Center 1221 1st Avenue, Apt #24E Seattle, WA 98101

AAOS Staff

Deborah S. Cummins, PhD

Director, Research and Scientific Affairs 6300 N. River Road Rosemont, IL 60018

Sharon Song, PhD

Manager, Clinical Practice Guidelines song@aaos.org

Patrick Sluka, MPH

Former AAOS Lead Research Analyst

Kevin Bover

Former Appropriate Use Criteria Unit Manager Former Interim Clinical Practice Guidelines Manager

Anne Woznica, MLIS

Medical Research Librarian

ADA Staff

Helen Ristic, PhD.

Director, Scientific Information ADA Division of Science 211 E. Chicago Avenue Chicago, IL 60611

Nicholas Buck Hanson, MPH

ADA Lead Research Analyst hansonn@ada.org

Special Recognitions

William Robert Martin, III, MD

American Academy of Orthopaedic Surgeons Medical Director 317 Massachusetts Avenue NE Washington, D.C. 20002-5769

APPENDIX II

CREATING PRELIMINARY RECOMMENDATIONS

In an effort to ensure the broadest literature search possible and to evaluate the many aspects related to preventing orthopaedic implant infection in patients undergoing dental procedures, the work group constructed a causal pathway for orthopaedic implant infection consisting of the following factors:

- Patients
- Patient Characteristics Increasing Risk of Infection
- Prophylactic Interventions
- Effect of Intervention on:
 - o Bacteria/Fungi in the Mouth
 - o Bacteremia/Fungemia in the Blood
 - o Implant Infection

The factors and their components were then combined to create a series of questions from which our literature searches were derived. The components of each factor listed above are illustrated in the figure below. The questions for which we derived our literature searches are listed below.

Preliminary recommendations were then created based on the interventions selected for the causal pathway. Remaining questions not directly related to an intervention (e.g. questions about no intervention, the relationship between bacteremia and implant infection) were assessed in order to further inform the discussion among work group members when they met at the final recommendation meeting.

Causal Pathway

PATIENTS ...

... at risk for oral bacteremias

(i.e. everybody)

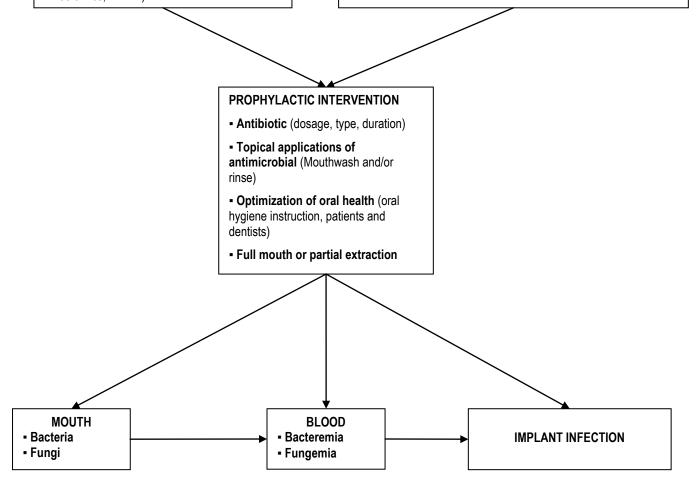
Oral/Dental Procedure (e.g. gingival manipulation or mucosal incision) Daily activities (naturally occurring) (Epidemiology studies for bacteremia)

... with Bone and Joint Implant

Prosthetic Joint Implant
(including silastic implants)
Massive structural allografts
Spinal instrumentation (e.g. rods)
Trauma device (e.g. plates, screws)
Bone void fillers (e.g. allografts, bone glass, ceramics, PMMA)

FACTORS INCREASING RISK OF INFECTION

- Immunocompromised (diabetes mellitus Type I and II, autoimmune disease, organ, transplant, chemotherapy, bone marrow transplant, HIV, steroid, obesity, hemophilia, malnutrition, tobacco exposure, alcohol, elderly, leukemia, radiation therapy, cancer, immunomodulated therapy,)
- Oral health status (gingivitis, periodontitis, caries, nonodontogenic infection, odontogenic infection)
- Edentulous
- History of previous implant infection
- Time from implant
- Multiple implants
- At risk prosthesis (revision prosthesis, prosthesis mechanically failed, megaprosthesis, endoprosthetic reconstruction)
- Bisphosphonate therapy



Questions Derived from Causal Pathway

Relationships Between Mouth, Blood, and Implant Infection

- 1. What is the relationship between bacteria in the mouth and implant infection?
- 2. What is the relationship between fungi in the mouth and implant infection?
- 3. What is the relationship between bacteria in the mouth (after an oral/dental procedure) and bacteremia?
- 4. What is the relationship between fungi in the mouth (after an oral/dental procedure) and fungemia?
- 5. What is the relationship between bacteremia from an oral source after an oral/dental procedure and implant infection?
- 6. What is the relationship between fungemia from an oral source after an oral/dental procedure and implant infection?

Patients Without Bone and Joint Implants

- 7. In patients without an implant having an oral/dental procedure or undertaking daily activities who have immunocompromising factors, what are the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 8. In patients without an implant having an oral/dental procedure or undertaking daily activities who have immunocompromising factors, what are the incidence, nature, duration, and magnitude of fungi in the mouth?
- 9. In patients without an implant having an oral/dental procedure or undertaking daily activities who have immunocompromising factors, what are the incidence, nature, duration, and magnitude of bacteremia in the blood?
- 10. In patients without an implant having an oral/dental procedure or undertaking daily activities who have immunocompromising factors, what are the incidence, nature, duration, and magnitude of fungemia in the blood?
- 11. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what are the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 12. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what are the incidence, nature, duration, and magnitude of fungi in the mouth?
- 13. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what are the incidence, nature, duration, and magnitude of bacteremia in the blood?

- 14. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what are the incidence, nature, duration, and magnitude of fungemia in the blood?
- 15. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what are the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 16. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what are the incidence, nature, duration, and magnitude of fungi in the mouth?
- 17. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what are the incidence, nature, duration, and magnitude of bacteremia in the blood?
- 18. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what are the incidence, nature, duration, and magnitude of fungemia in the blood?
- 19. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what are the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 20. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what are the incidence, nature, duration, and magnitude of fungi in the mouth?
- 21. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what are the incidence, nature, duration, and magnitude of bacteremia in the blood?
- 22. In patients without an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what are the incidence, nature, duration, and magnitude of fungemia in the blood?

Patients With Bone and Joint Implants

- 23. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 24. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungi in the mouth?

- 25. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteremia?
- 26. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungemia?
- 27. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of no intervention on the incidence, nature, duration, and magnitude of implant infection?
- 28. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 29. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 30. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteremia?
- 31. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungemia?
- 32. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of implant infection?
- 33. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 34. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 35. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteremia?
- 36. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungemia?

- 37. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of implant infection?
- 38. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 39. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 40. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteremia?
- 41. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungemia?
- 42. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of implant infection?
- 43. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 44. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 45. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteremia?
- 46. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungemia?
- 47. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have immunocompromising factors, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of implant infection?
- 48. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteria in the mouth?

- 49. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 50. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteremia?
- 51. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungemia?
- 52. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of no intervention on the incidence, nature, duration, and magnitude of implant infection?
- 53. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 54. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 55. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteremia?
- 56. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungemia?
- 57. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of implant infection?
- 58. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 59. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 60. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteremia?

- 61. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungemia?
- 62. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of implant infection?
- 63. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 64. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 65. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteremia?
- 66. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungemia?
- 67. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of implant infection?
- 68. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 69. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 70. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteremia?
- 71. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungemia?
- 72. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have Poor/Good oral health status, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of implant infection?

- 73. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have a history of previous implant infection, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 74. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have a history of previous implant infection, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 75. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have a history of previous implant infection, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteremia?
- 76. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have a history of previous implant infection, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungemia?
- 77. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have a history of previous implant infection, what is the effect of no intervention on the incidence, nature, duration, and magnitude of implant infection?
- 78. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 79. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 80. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteremia?
- 81. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungemia?
- 82. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of implant infection?
- 83. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 84. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungi in the mouth?

- 85. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteremia?
- 86. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungemia?
- 87. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of implant infection?
- 88. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 89. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 90. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteremia?
- 91. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungemia?
- 92. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of implant infection?
- 93. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 94. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 95. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteremia?
- 96. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungemia?

- 97. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have history of previous implant infection, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of implant infection?
- 98. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 99. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 100. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteremia?
- 101. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungemia?
- 102. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of no intervention on the incidence, nature, duration, and magnitude of implant infection?
- 103. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 104. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 105. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteremia?
- 106. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungemia?
- 107. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of implant infection?
- 108. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteria in the mouth?

- 109. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 110. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteremia?
- 111. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungemia?
- 112. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of implant infection?
- 113. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 114. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 115. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteremia?
- 116. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungemia?
- 117. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of implant infection?
- 118. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 119. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 120. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteremia?

- 121. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungemia?
- 122. In patients with an implant having an oral/dental procedure or undertaking daily activities and [insert duration] since implant surgery, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of implant infection?
- 123. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 124. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 125. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteremia?
- 126. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungemia?
- 127. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of no intervention on the incidence, nature, duration, and magnitude of implant infection?
- 128. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 129. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 130. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteremia?
- 131. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungemia?
- 132. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of implant infection?

- 133. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 134. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 135. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteremia?
- 136. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungemia?
- 137. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of implant infection?
- 138. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 139. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 140. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteremia?
- 141. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungemia?
- 142. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of implant infection?
- 143. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 144. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungi in the mouth?

- 145. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteremia?
- 146. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungemia?
- 147. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have multiple implants, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of implant infection?
- 148. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 149. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 150. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteremia?
- 151. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungemia?
- 152. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of no intervention on the incidence, nature, duration, and magnitude of implant infection?
- 153. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 154. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 155. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteremia?
- 156. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungemia?

- 157. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of implant infection?
- 158. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 159. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 160. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteremia?
- 161. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungemia?
- 162. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of implant infection?
- 163. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 164. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 165. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteremia?
- 166. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungemia?
- 167. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of implant infection?
- 168. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteria in the mouth?

- 169. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 170. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteremia?
- 171. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungemia?
- 172. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have "at-risk" prosthesis, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of implant infection?
- 173. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 174. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 175. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteremia?
- 176. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungemia?
- 177. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of no intervention on the incidence, nature, duration, and magnitude of implant infection?
- 178. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 179. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 180. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteremia?

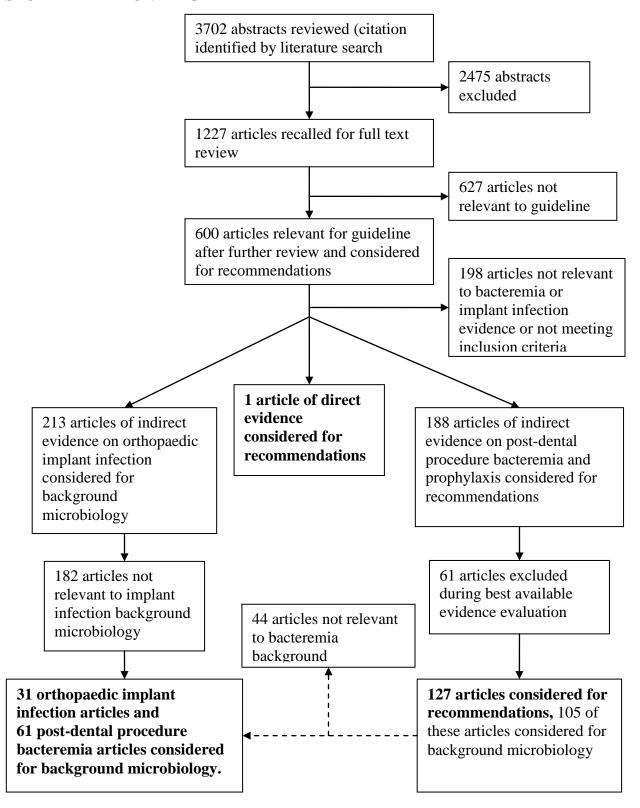
- 181. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungemia?
- 182. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of implant infection?
- 183. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 184. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 185. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteremia?
- 186. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungemia?
- 187. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of implant infection?
- 188. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 189. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 190. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteremia?
- 191. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungemia?
- 192. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of implant infection.

- 193. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 194. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 195. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteremia?
- 196. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungemia?
- 197. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have no teeth (edentulous), what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of implant infection?
- 198. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 199. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 200. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of no intervention on the incidence, nature, duration, and magnitude of bacteremia?
- 201. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of no intervention on the incidence, nature, duration, and magnitude of fungemia?
- 202. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of no intervention on the incidence, nature, duration, and magnitude of implant infection?
- 203. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 204. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungi in the mouth?

- 205. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of bacteremia?
- 206. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of fungemia?
- 207. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of antibiotic prophylaxis on the incidence, nature, duration, and magnitude of implant infection?
- 208. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 209. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 210. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of bacteremia?
- 211. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of fungemia?
- 212. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of topical application of antimicrobials on the incidence, nature, duration, and magnitude of implant infection?
- 213. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 214. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 215. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of bacteremia?
- 216. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of fungemia?

- 217. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of optimization of oral health instructions on the incidence, nature, duration, and magnitude of implant infection?
- 218. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteria in the mouth?
- 219. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungi in the mouth?
- 220. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of bacteremia?
- 221. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of fungemia?
- 222. In patients with an implant having an oral/dental procedure or undertaking daily activities and who have bisphosphonate therapy, what is the effect of extraction (full or partial mouth) on the incidence, nature, duration, and magnitude of implant infection?

APPENDIX III STUDY ATTRITION DIAGRAM



INCLUDED STUDIES TABLES

RECOMMENDATION 1

Table 48 Included Studies for Recommendation 1

| Author(s) | Year | Title |
|------------------------------------------------------------------------------------------------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------------------------|
| Berbari EF;Osmon DR;Carr A;Hanssen AD;Baddour LM;Greene D;Kupp LI;Baughan LW;Harmsen WS;Mandrekar JN;Therneau TM;Steckelberg JM;Virk A;Wilson WR; | 2010 | Dental procedures as risk factors for prosthetic hip or knee infection: a hospital-based prospective case-control study |
| Morozumi T;Kubota T;Abe D;Shimizu T;Komatsu Y;Yoshie H; | 2010 | Effects of irrigation with an antiseptic and oral administration of azithromycin on bacteremia caused by scaling and root planing |
| Lockhart PB;Brennan MT;Sasser HC;Fox PC;Paster BJ;Bahrani-Mougeot FK; | 2008 | Bacteremia associated with toothbrushing and dental extraction |
| Brennan MT;Kent ML;Fox PC;Norton HJ;Lockhart PB; | 2007 | The impact of oral disease and nonsurgical treatment on bacteremia in children |
| Diz DP;Tomas C;Limeres PJ;Medina HJ;Fernandez FJ;Alvarez FM; | 2006 | Comparative efficacies of amoxicillin, clindamycin, and moxifloxacin in prevention of bacteremia following dental extractions |
| Lockhart PB;Brennan MT;Kent ML;Norton HJ;Weinrib DA; | 2004 | Impact of amoxicillin prophylaxis on the incidence, nature, and duration of bacteremia in children after intubation and dental procedures |
| Vergis EN;Demas PN;Vaccarello SJ;Yu VL; | 2001 | Topical antibiotic prophylaxis for bacteremia after dental extractions |
| Wahlmann U;Al-Nawas B;Jutte M;Wagner W; | 1999 | Clinical and microbiological efficacy of single dose cefuroxime prophylaxis for dental surgical procedures |

Table 48 Included Studies for Recommendation 1

| Author(s) | Year | Title |
|-----------------------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------------------------------|
| Hall G;Heimdahl A;Nord CE; | 1996 | Effects of prophylactic administration of cefaclor on transient bacteremia after dental extraction |
| Hall G;Nord CE;Heimdahl A; | 1996 | Elimination of bacteraemia after dental extraction: comparison of erythromycin and clindamycin for prophylaxis of infective endocarditis |
| Aitken C;Cannell H;Sefton AM;Kerawala C;Seymour A;Murphy M;Whiley RA;Williams JD; | 1995 | Comparative efficacy of oral doses of clindamycin and erythromycin in the prevention of bacteraemia |
| Hall G;Hedstrom SA;Heimdahl A;Nord CE; | 1993 | Prophylactic administration of penicillins for endocarditis does not reduce the incidence of postextraction bacteremia |
| Goker K;Guvener O; | 1992 | Antibacterial effects of ofloxacin, clindamycin and sultamicillin on surgical removal of impacted third molars |
| Katoh H; | 1992 | Incidence of transient bacteremia following dental surgery-prophylactic use of cefuroxime, ceftriaxone or clindamycin |
| Cannell H;Kerawala C;Sefton AM;Maskell JP;Seymour A;Sun ZM;Williams JD; | 1991 | Failure of two macrolide antibiotics to prevent post-extraction bacteraemia |
| Coulter WA;Coffey A;Saunders ID;Emmerson AM; | 1990 | Bacteremia in children following dental extraction |
| Casolari C;Neglia R;Forabosco A;Galetti R;Fabio U; | 1989 | Incidence of oral bacteremia and antimicrobial prophylaxis |
| Roberts GJ;Radford P;Holt R; | 1987 | Prophylaxis of dental bacteraemia with oral amoxycillin in children |

Table 48 Included Studies for Recommendation 1

| Author(s) | Year | Title |
|--------------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------------------------------------------|
| Shanson DC;Shehata A;Tadayon M;Harris M; | 1987 | Comparison of intravenous teicoplanin with intramuscular amoxycillin for the prophylaxis of streptococcal bacteraemia in dental patients |
| Maskell JP;Carter JL;Boyd RB;Williams RJ; | 1986 | Teicoplanin as a prophylactic antibiotic for dental bacteraemia |
| Josefsson K;Heimdahl A;von KL;Nord CE; | 1985 | Effect of phenoxymethylpenicillin and erythromycin prophylaxis on anaerobic bacteraemia after oral surgery |
| Shanson DC;Akash S;Harris M;Tadayon M; | 1985 | Erythromycin stearate, 1.5 g, for the oral prophylaxis of streptococcal bacteraemia in patients undergoing dental extraction: efficacy and tolerance |
| Head TW;Bentley KC;Millar EP;deVries JA; | 1984 | A comparative study of the effectiveness of metronidazole and penicillin V in eliminating anaerobes from postextraction bacteremias |
| Appleman MD;Sutter VL;Sims TN; | 1982 | Value of antibiotic prophylaxis in periodontal surgery |
| Baltch AL;Schaffer C;Hammer MC;Sutphen NT;Smith RP;Conroy J;Shayegani M; | 1982 | Bacteremia following dental cleaning in patients with and without penicillin prophylaxis |
| Shanson DC;Cannon P;Wilks M; | 1978 | Amoxycillin compared with penicillin V for the prophylaxis of dental bacteraemia |
| DeVries J;Francis LE;Lang D; | 1972 | Control of post-extraction bacteraemias in the penicillin- hypersensitive patient |
| Jokinen MA; | 1970 | Bacteremia following dental extraction and its prophylaxis |

Table 48 Included Studies for Recommendation 1

| Author(s) | Year | Title |
|-----------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------------|
| Khairat O; | 1966 | An effective antibiotic cover for the prevention of endocarditis following dental and other post-operative bacteraemias |
| Martin WJ;Schirger A; | 1964 | Prevention of bacteremia after oral surgery |
| Gutverg M; | 1962 | Studies on bacteremia following oral surgery: Some prophylactic approaches to bacteremia and the results of tissue examination of excised gingival |

Table 49 Included Studies for Recommendation 2

| Author(s) | Year | Title |
|-----------------------------------------------------------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Fine DH;Furgang D;McKiernan M;Tereski-Bischio D;Ricci-Nittel D;Zhang P;Araujo MW; | 2010 | An investigation of the effect of an essential oil mouthrinse on induced bacteraemia: a pilot study |
| Morozumi T;Kubota T;Abe D;Shimizu T;Komatsu Y;Yoshie H; | 2010 | Effects of irrigation with an antiseptic and oral administration of azithromycin on bacteremia caused by scaling and root planing |
| Pineiro A;Tomas I;Blanco J;Alvarez M;Seoane J;Diz P; | 2010 | Bacteraemia following dental implants' placement |
| Cherry M;Daly CG;Mitchell D;Highfield J; | 2007 | Effect of rinsing with povidone-iodine on bacteraemia due to scaling: a randomized-controlled trial |
| Tomas I;Alvarez M;Limeres J;Tomas M;Medina J;Otero JL;Diz P; | 2007 | Effect of a chlorhexidine mouthwash on the risk of postextraction bacteremia |
| Fourrier F;Dubois D;Pronnier P;Herbecq P;Leroy O;Desmettre T;Pottier-Cau E;Boutigny H;Di PC;Durocher A;Roussel-Delvallez M; | 2005 | Effect of gingival and dental plaque antiseptic decontamination on nosocomial infections acquired in the intensive care unit: a double-blind placebo-controlled multicenter study |
| Erverdi N;Acar A;Isguden B;Kadir T; | 2001 | Investigation of bacteremia after orthodontic banding and debanding following chlorhexidine mouth wash application |
| Brown AR;Papasian CJ;Shultz P;Theisen FC;Shultz RE; | 1998 | Bacteremia and intraoral suture removal: can an antimicrobial rinse help? |
| Fine DH;Korik I;Furgang D;Myers R;Olshan A;Barnett ML;Vincent J; | 1996 | Assessing pre-procedural subgingival irrigation and rinsing with an antiseptic mouthrinse to reduce bacteremia |
| Lockhart PB; | 1996 | An analysis of bacteremias during dental extractions. A double-blind, placebo-controlled study of chlorhexidine |

Table 49 Included Studies for Recommendation 2

| Author(s) | Year | Title |
|------------------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------------|
| Rahn R;Schneider S;Diehl O;Schafer V;Shah PM; | 1995 | Preventing post-treatment bacteremia: comparing topical povidone-iodine and chlorhexidine |
| Yamalik MK;Yucetas S;Abbasoglu U; | 1992 | Effects of various antiseptics on bacteremia following tooth extraction |
| Lofthus JE;Waki MY;Jolkovsky DL;Otomo- Corgel J;Newman MG;Flemmig T;Nachnani S; | 1991 | Bacteremia following subgingival irrigation and scaling and root planing |
| Waki MY;Jolkovsky DL;Otomo-Corgel J;Lofthus JE;Nachnani S;Newman MG;Flemmig TF; | 1990 | Effects of subgingival irrigation on bacteremia following scaling and root planing |
| Casolari C;Neglia R;Forabosco A;Galetti R;Fabio U; | 1989 | Incidence of oral bacteremia and antimicrobial prophylaxis |
| MacFarlane TW;Ferguson MM;Mulgrew CJ; | 1984 | Post-extraction bacteraemia: role of antiseptics and antibiotics |
| Jokinen MA; | 1978 | Prevention of postextraction bacteremia by local prophylaxis |
| Sweet JB;Gill VJ;Chusid MJ;Elin RJ; | 1978 | Nitroblue tetrazolium and Limulus assays for bacteremia after dental extraction: effect of topical antiseptics |
| Nasif AS; | 1977 | The incidence of post-extraction bacteremia after irrigation of the gingival sulcus with hydrogen peroxide solution |
| Brenman HS;Randall E; | 1974 | Local degerming with povidone-iodine, II. Prior to gingivectomy |
| Huffman GG;Wood WH;Hausler WJ;Jensen J; | 1974 | The effects of preoperative rinsing with cetylpyridinium chloride on bacteremia associated with the surgical removal of impacted third molars |

Table 49 Included Studies for Recommendation 2

| Author(s) | Year | Title |
|-------------------------------------------|------|------------------------------------------------------------------------------------------------------------------|
| Madsen KL; | 1974 | Effect of chlorhexidine mouthrinse and periodontal treatment upon bacteremia produced by oral hygiene procedures |
| Francis LE;DeVries J;Lang D; | 1973 | An oral antiseptic for the control of post-extraction bacteraemia |
| Cutcher JL;Goldberg JR;Lilly GE;Jones JC; | 1971 | Control of bacteremia associated with extraction of teeth. II |
| Scopp IW;Orvieto LD; | 1971 | Gingival degerming by povidone-iodine irrigation: bacteremia reduction in extraction procedures |
| Jones JC;Cutcher JL;Goldberg JR;Lilly GE; | 1970 | Control of bacteremia associated with extraction of teeth |

Table 50 Included Studies for Recommendation 3

| Author(s) | Year | Title |
|---------------------------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------|
| Barbosa M;Carmona IT;Amaral B;Limeres J;Alvarez M;Cerqueira C;Diz P; | 2010 | General anesthesia increases the risk of bacteremia following dental extractions |
| Ashare A;Stanford C;Hancock P;Stark D;Lilli K;Birrer E;Nymon A;Doerschug KC;Hunninghake GW; | 2009 | Chronic liver disease impairs bacterial clearance in a human model of induced bacteremia |
| Crasta K;Daly CG;Mitchell D;Curtis B;Stewart D;Heitz-Mayfield LJ; | 2009 | Bacteraemia due to dental flossing |
| Lockhart PB;Brennan MT;Thornhill M;Michalowicz BS;Noll J;Bahrani-Mougeot FK;Sasser HC; | 2009 | Poor oral hygiene as a risk factor for infective endocarditis- related bacteremia |
| Enabulele OI;Aluyi HSA;Omokao O; | 2008 | Incidence of bacteraemia following teeth extraction at the dental clinic of the University of Benin Teaching Hospital, Benin city, Nigeria |
| Tomas I;Pereira F;Llucian R;Poveda R;Diz P;Bagan JV; | 2008 | Prevalence of bacteraemia following third molar surgery |
| Valdes C;Tomas I;Alvarez M;Limeres J;Medina J;Diz P; | 2008 | The incidence of bacteraemia associated with tracheal intubation |
| Brennan MT;Kent ML;Fox PC;Norton HJ;Lockhart PB; | 2007 | The impact of oral disease and nonsurgical treatment on bacteremia in children |
| Cherry M;Daly CG;Mitchell D;Highfield J; | 2007 | Effect of rinsing with povidone-iodine on bacteraemia due to scaling: a randomized-controlled trial |

Table 50 Included Studies for Recommendation 3

| Author(s) | Year | Title |
|----------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------|
| Forner L;Larsen T;Kilian M;Holmstrup P; | 2006 | Incidence of bacteremia after chewing, tooth brushing and scaling in individuals with periodontal inflammation |
| Takai S;Kuriyama T;Yanagisawa M;Nakagawa K;Karasawa T; | 2005 | Incidence and bacteriology of bacteremia associated with various oral and maxillofacial surgical procedures |
| Bhanji S;Williams B;Sheller B;Elwood T;Mancl L; | 2002 | Transient bacteremia induced by toothbrushing a comparison of the Sonicare toothbrush with a conventional toothbrush |
| Daly CG;Mitchell DH;Highfield JE;Grossberg DE;Stewart D; | 2001 | Bacteremia due to periodontal probing: a clinical and microbiological investigation |
| Wahlmann U;Al-Nawas B;Jutte M;Wagner W; | 1999 | Clinical and microbiological efficacy of single dose cefuroxime prophylaxis for dental surgical procedures |
| Roberts GJ;Watts R;Longhurst P;Gardner P; | 1998 | Bacteremia of dental origin and antimicrobial sensitivity following oral surgical procedures in children |
| Daly C;Mitchell D;Grossberg D;Highfield J;Stewart D; | 1997 | Bacteraemia caused by periodontal probing |
| Lockhart PB; | 1996 | An analysis of bacteremias during dental extractions. A double-blind, placebo-controlled study of chlorhexidine |
| Okabe K;Nakagawa K;Yamamoto E; | 1995 | Factors affecting the occurrence of bacteremia associated with tooth extraction |
| Coulter WA;Coffey A;Saunders ID;Emmerson AM; | 1990 | Bacteremia in children following dental extraction |
| Trivedi DN; | 1984 | Bacteraemia due to operative procedure |

Table 50 Included Studies for Recommendation 3

| Author(s) | Year | Title |
|----------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------|
| Silver JG;Martin AW;McBride BC; | 1977 | Experimental transient bacteraemias in human subjects with varying degrees of plaque accumulation and gingival inflammation |
| De Leo AA;Schoenknecht FD;Anderson MW;Peterson JC; | 1974 | The incidence of bacteremia following oral prophylaxis on pediatric patients |
| Lineberger LT;De Marco TJ; | 1973 | Evaluation of transient bacteremia following routine periodontal procedures |

DENTAL PROCEDURES AND BACTEREMIA

Table 51 Included Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title |
|-----------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------|
| Barbosa M;Carmona IT;Amaral B;Limeres J;Alvarez M;Cerqueira C;Diz P; | 2010 | General anesthesia increases the risk of bacteremia following dental extractions |
| Morozumi T;Kubota T;Abe D;Shimizu T;Komatsu Y;Yoshie H; | 2010 | Effects of irrigation with an antiseptic and oral administration of azithromycin on bacteremia caused by scaling and root planing |
| Pineiro A;Tomas I;Blanco J;Alvarez M;Seoane J;Diz P; | 2010 | Bacteraemia following dental implants' placement |
| Crasta K;Daly CG;Mitchell D;Curtis B;Stewart D;Heitz-Mayfield LJ; | 2009 | Bacteraemia due to dental flossing |
| Gurel HG;Basciftci FA;Arslan U; | 2009 | Transient bacteremia after removal of a bonded maxillary expansion appliance |
| Nixon PP;Littler P;Davies K;Krishnam MS; | 2009 | Does sialography require antibiotic prophylaxis? |
| Sonbol H;Spratt D;Roberts GJ;Lucas VS; | 2009 | Prevalence, intensity and identity of bacteraemia following conservative dental procedures in children |
| Enabulele OI;Aluyi HSA;Omokao O; | 2008 | Incidence of bacteraemia following teeth extraction at the dental clinic of the University of Benin Teaching Hospital, Benin city, Nigeria |
| Lockhart PB;Brennan MT;Sasser HC;Fox PC;Paster BJ;Bahrani-Mougeot FK; | 2008 | Bacteremia associated with toothbrushing and dental extraction |
| Tomas I;Pereira F;Llucian R;Poveda R;Diz P;Bagan JV; | 2008 | Prevalence of bacteraemia following third molar surgery |

Table 51 Included Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title |
|------------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------|
| Valdes C;Tomas I;Alvarez M;Limeres J;Medina J;Diz P; | 2008 | The incidence of bacteraemia associated with tracheal intubation |
| Cherry M;Daly CG;Mitchell D;Highfield J; | 2007 | Effect of rinsing with povidone-iodine on bacteraemia due to scaling: a randomized-controlled trial |
| Lafaurie GI;Mayorga-Fayad I;Torres MF;Castillo DM;Aya MR;Baron A;Hurtado PA; | 2007 | Periodontopathic microorganisms in peripheric blood after scaling and root planing |
| Tomas I;Alvarez M;Limeres J;Potel C;Medina J;Diz P; | 2007 | Prevalence, duration and aetiology of bacteraemia following dental extractions |
| Tomas I;Alvarez M;Limeres J;Tomas M;Medina J;Otero JL;Diz P; | 2007 | Effect of a chlorhexidine mouthwash on the risk of postextraction bacteremia |
| Forner L;Larsen T;Kilian M;Holmstrup P; | 2006 | Incidence of bacteremia after chewing, tooth brushing and scaling in individuals with periodontal inflammation |
| Forner L;Nielsen CH;Bendtzen K;Larsen T;Holmstrup P; | 2006 | Increased plasma levels of IL-6 in bacteremic periodontis patients after scaling |
| Murphy AM;Daly CG;Mitchell DH;Stewart D;Curtis BH; | 2006 | Chewing fails to induce oral bacteraemia in patients with periodontal disease |
| Oncag O;Aydemir S;Ersin N;Koca H; | 2006 | Bacteremia incidence in pediatric patients under dental general anesthesia |
| Kinane DF;Riggio MP;Walker KF;MacKenzie D;Shearer B; | 2005 | Bacteraemia following periodontal procedures |
| Oncag O;Cokmez B;Aydemir S;Balcioglu T; | 2005 | Investigation of bacteremia following nasotracheal intubation |

Table 51 Included Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title |
|------------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------|
| Savarrio L;MacKenzie D;Riggio M;Saunders WP;Bagg J; | 2005 | Detection of bacteraemias during non-surgicalroot canal treatment |
| Takai S;Kuriyama T;Yanagisawa M;Nakagawa K;Karasawa T; | 2005 | Incidence and bacteriology of bacteremia associated with various oral and maxillofacial surgical procedures |
| Burden DJ;Coulter WA;Johnston CD;Mullally B;Stevenson M; | 2004 | The prevalence of bacteraemia on removal of fixed orthodontic appliances |
| Rajasuo A;Nyfors S;Kanervo A;Jousimies-Somer H;Lindqvist C;Suuronen R; | 2004 | Bacteremia after plate removal and tooth extraction |
| Rajasuo A;Perkki K;Nyfors S;Jousimies-Somer H;Meurman JH; | 2004 | Bacteremia following surgical dental extraction with an emphasis on anaerobic strains |
| Bhanji S;Williams B;Sheller B;Elwood T;Mancl L; | 2002 | Transient bacteremia induced by toothbrushing a comparison of the Sonicare toothbrush with a conventional toothbrush |
| Daly CG;Mitchell DH;Highfield JE;Grossberg DE;Stewart D; | 2001 | Bacteremia due to periodontal probing: a clinical and microbiological investigation |
| Lucas V;Roberts GJ; | 2000 | Odontogenic bacteremia following tooth cleaning procedures in children |
| Roberts GJ;Gardner P;Longhurst P;Black AE;Lucas VS; | 2000 | Intensity of bacteraemia associated with conservative dental procedures in children |
| Erverdi N;Kadir T;Ozkan H;Acar A; | 1999 | Investigation of bacteremia after orthodontic banding |
| Brown AR;Papasian CJ;Shultz P;Theisen FC;Shultz RE; | 1998 | Bacteremia and intraoral suture removal: can an antimicrobial rinse help? |

Table 51 Included Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title |
|------------------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------|
| Roberts GJ;Simmons NB;Longhurst P;Hewitt PB; | 1998 | Bacteraemia following local anaesthetic injections in children |
| Roberts GJ;Watts R;Longhurst P;Gardner P; | 1998 | Bacteremia of dental origin and antimicrobial sensitivity following oral surgical procedures in children |
| Daly C;Mitchell D;Grossberg D;Highfield J;Stewart D; | 1997 | Bacteraemia caused by periodontal probing |
| Roberts GJ;Holzel HS;Sury MR;Simmons NA;Gardner P;Longhurst P; | 1997 | Dental bacteremia in children |
| Debelian GJ;Olsen I;Tronstad L; | 1995 | Bacteremia in conjunction with endodontic therapy |
| Rahn R;Schneider S;Diehl O;Schafer V;Shah PM; | 1995 | Preventing post-treatment bacteremia: comparing topical povidone-iodine and chlorhexidine |
| Ali MT;Tremewen DR;Hay AJ;Wilkinson DJ; | 1992 | The occurrence of bacteraemia associated with the use of oral and nasopharyngeal airways |
| Giglio JA;Rowland RW;Dalton HP;Laskin DM; | 1992 | Suture removal-induced bacteremia: a possible endocarditis risk |
| Lucartorto FM;Franker CK;Maza J; | 1992 | Postscaling bacteremia in HIV-associated gingivitis and periodontitis |
| Roberts GJ;Gardner P;Simmons NA; | 1992 | Optimum sampling time for detection of dental bacteraemia in children |
| Lofthus JE;Waki MY;Jolkovsky DL;Otomo- Corgel J;Newman MG;Flemmig T;Nachnani S; | 1991 | Bacteremia following subgingival irrigation and scaling and root planing |

Table 51 Included Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title |
|---------------------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------------------------------|
| Coulter WA;Coffey A;Saunders ID;Emmerson AM; | 1990 | Bacteremia in children following dental extraction |
| Flood TR;Samaranayake LP;MacFarlane TW;McLennan A;MacKenzie D;Carmichael F; | 1990 | Bacteraemia following incision and drainage of dento-alveolar abscesses |
| Heimdahl A;Hall G;Hedberg M;Sandberg H;Soder PO;Tuner K;Nord CE; | 1990 | Detection and quantitation by lysis-filtration of bacteremia after different oral surgical procedures |
| Waki MY;Jolkovsky DL;Otomo-Corgel J;Lofthus JE;Nachnani S;Newman MG;Flemmig TF; | 1990 | Effects of subgingival irrigation on bacteremia following scaling and root planing |
| Casolari C;Neglia R;Forabosco A;Galetti R;Fabio U; | 1989 | Incidence of oral bacteremia and antimicrobial prophylaxis |
| Hansen CP;Westh H;Brok KE;Jensen R;Bertelsen S; | 1989 | Bacteraemia following orotracheal intubation and oesophageal balloon dilatation |
| King RC;Crawford JJ;Small EW; | 1988 | Bacteremia following intraoral suture removal |
| Dinner M;Tjeuw M;Artusio JF; | 1987 | Bacteremia as a complication of nasotracheal intubation |
| Shanson DC;Shehata A;Tadayon M;Harris M; | 1987 | Comparison of intravenous teicoplanin with intramuscular amoxycillin for the prophylaxis of streptococcal bacteraemia in dental patients |
| Maskell JP;Carter JL;Boyd RB;Williams RJ; | 1986 | Teicoplanin as a prophylactic antibiotic for dental bacteraemia |
| Josefsson K;Heimdahl A;von KL;Nord CE; | 1985 | Effect of phenoxymethylpenicillin and erythromycin prophylaxis on anaerobic bacteraemia after oral surgery |

Table 51 Included Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title |
|---------------------------------------------------------------|------|---------------------------------------------------------------------------------------------------------------------------------------------------|
| Lamey PJ;MacFarlane TW;Patton DW;Samaranayake LP;Ferguson MM; | 1985 | Bacteraemia consequential to sialography |
| Trivedi DN; | 1984 | Bacteraemia due to operative procedure |
| Marzoni FA;Kelly DR; | 1983 | Bacteremia following cleft palate repaira prospective study |
| Sconyers JR;Albers DD;Kelly R; | 1979 | Relationship of bacteremia to toothbrushing in clinically healthy patients |
| Silver JG;Martin AW;McBride BC; | 1979 | Experimental transient bacteraemias in human subjects with clinically healthy gingivae |
| Shanson DC;Cannon P;Wilks M; | 1978 | Amoxycillin compared with penicillin V for the prophylaxis of dental bacteraemia |
| Wampole HS;Allen AL;Gross A; | 1978 | The incidence of transient bacteremia during periodontal dressing change |
| Baumgartner JC;Heggers JP;Harrison JW; | 1977 | Incidence of bacteremias related to endodontic procedures. II. Surgical endodontics |
| Soliman NA;el-Batawy YA;Abdallah AK; | 1977 | Studies on bacteremia following oral surgery: Some prophylactic approaches to bacteremia and the results of tissue examination of excised gingiva |
| Baumgartner JC;Heggers JP;Harrison JW; | 1976 | The incidence of bacteremias related to endodontic procedures. I. Nonsurgical endodontics |
| Peterson LJ;Peacock R; | 1976 | The incidence of bacteremia in pediatric patients following tooth extraction |

Table 51 Included Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title |
|------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------------|
| Wank HA;Levison ME;Rose LF;Cohen DW; | 1976 | A quantitative measurement of bacteremia and its relationship to plaque control |
| Ramadan AE;Zaki SA;Nour ZM; | 1975 | A study of transient bacteremia following the use of dental floss silk and interdental stimulators |
| Berger SA;Weitzman S;Edberg SC;Casey JI; | 1974 | Bacteremia after the use of an oral irrigation device. A controlled study in subjects with normal-appearing gingiva: comparison with use of toothbrush |
| Crawford JJ;Sconyers JR;Moriarty JD;King RC;West JF; | 1974 | Bacteremia after tooth extractions studied with the aid of prereduced anaerobically sterilized culture media |
| De Leo AA;Schoenknecht FD;Anderson MW;Peterson JC; | 1974 | The incidence of bacteremia following oral prophylaxis on pediatric patients |
| Berry FA;Blankenbaker WL;Ball CG; | 1973 | Comparison of bacteremia occurring with nasotracheal and orotracheal intubation |
| Francis LE;DeVries J;Lang D; | 1973 | An oral antiseptic for the control of post-extraction bacteraemia |
| Lineberger LT;De Marco TJ; | 1973 | Evaluation of transient bacteremia following routine periodontal procedures |
| Sconyers JR;Crawford JJ;Moriarty JD; | 1973 | Relationship of bacteremia to toothbrushing in patients with periodontitis |
| Degling TE; | 1972 | Orthodontics, bacteremia, and the heart damaged patient |
| DeVries J;Francis LE;Lang D; | 1972 | Control of post-extraction bacteraemias in the penicillin- hypersensitive patient |

Table 51 Included Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title |
|--------------------------------------------------------------|------|---------------------------------------------------------------------------------------------------------------------------------------------------|
| The American Academy of Periodontology | 1972 | Oral irrigation and bacteremia |
| Felix JE;Rosen S;App GR; | 1971 | Detection of bacteremia after the use of an oral irrigation device in subjects with periodontitis |
| Romans AR;App GR; | 1971 | Bacteremia, a result from oral irrigation in subjects with gingivitis |
| Wada K;Tomizawa M;Sasaki I; | 1968 | Study on bacteriemia in patients with pyorrhea alveolaris caused by surgical operations |
| Conner HD;Haberman S;Collings CK;Winford TE; | 1967 | Bacteremias following periodontal scaling in patients with healthy appearing gingiva |
| Khairat O; | 1966 | The non-aerobes of post-extraction bacteremia |
| Martin WJ;Schirger A; | 1964 | PREVENTION OF BACTEREMIA AFTER ORAL SURGERY |
| Bender IB;SELTZER S;TASHMAN S;MELOFF G; | 1963 | Dental procedures in patients with rheumatic heart disease |
| Gutverg M; | 1962 | Studies on bacteremia following oral surgery: Some prophylactic approaches to bacteremia and the results of tissue examination of excised gingiva |
| ROGOSA M;HAMPP EG;NEVIN TA;WAGNER HN;DRISCOLL EJ;Baer PN; | 1960 | Blood sampling and cultural studies in the detection of postoperative bacteremias |
| Winslow MB;KOBERNICK SD; | 1960 | Bacteremia after prophylaxis |

BACKGROUND MICROBIOLOGY

Table 52 Included Studies for Background Microbiology

| Author(s) | Year | Title |
|------------------------------------------------------------------------------------------------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------------|
| Munoz-Mahamud E;Garcia S;Bori G;Martinez- Pastor JC;Zumbado JA;Riba J;Mensa J;Soriano A; | 2011 | Comparison of a low-pressure and a high-pressure pulsatile lavage during debridement for orthopaedic implant infection |
| Berbari EF;Osmon DR;Carr A;Hanssen AD;Baddour LM;Greene D;Kupp LI;Baughan LW;Harmsen WS;Mandrekar JN;Therneau TM;Steckelberg JM;Virk A;Wilson WR; | 2010 | Dental procedures as risk factors for prosthetic hip or knee infection: a hospital-based prospective case-control study |
| Morozumi T;Kubota T;Abe D;Shimizu T;Komatsu Y;Yoshie H; | 2010 | Effect of irrigation with antiseptic and oral administration of azithromycin on bacteremia caused by scaling and root planing |
| Pineiro A;Tomas I;Blanco J;Alvarez M;Seoane J;Diz P; | 2010 | Bacteraemia following dental implants' placement |
| Cordero-Ampuero J;Esteban J;Garcia-Cimbrelo E; | 2009 | Oral antibiotics are effective for highly resistant hip arthroplasty infections |
| Crasta K;Daly CG;Mitchell D;Curtis B;Stewart D;Heitz-Mayfield LJ; | 2009 | Bacteraemia due to dental flossing |
| Gurel HG;Basciftci FA;Arslan U; | 2009 | Transient bacteremia after removal of a bonded maxillary expansion appliance |
| Rodriguez D;Pigrau C;Euba G;Cobo J;Garcia- Lechuz J;Palomino J;Riera M;Del Toro MD;Granados A;Ariza X; | 2009 | Acute Hematogenous Prosthetic Joint Infection: Prospective Evaluation of Medical and Surgical Management |
| Sancheti KH;Laud NS;Bhende H;Reddy G;Pramod N;Mani JN; | 2009 | The INDUS knee prosthesis - Prospective multicentric trial of a posteriorly stabilized high-flex design: 2 years follow-up |

Table 52 Included Studies for Background Microbiology

| Author(s) | Year | Title |
|-----------------------------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------------------|
| Uckay I;Lubbeke A;Emonet S;Tovmirzaeva L;Stern R;Ferry T;Assal M;Bernard L;Lew D;Hoffmeyer P; | 2009 | Low incidence of haematogenous seeding to total hip and knee prostheses in patients with remote infections |
| Enabulele OI;Aluyi HSA;Omokao O; | 2008 | Incidence of bacteraemia following teeth extraction |
| Fink B;Makowiak C;Fuerst M;Berger I;Schafer P;Frommelt L; | 2008 | The value of synovial biopsy, joint aspiration and C-reactive protein in the diagnosis of late peri-prosthetic infection of total knee replacements |
| Hamilton H;Jamieson J; | 2008 | Deep infection in total hip arthroplasty |
| Valdes C;Tomas I;Alvarez M;Limeres J;Medina J;Diz P; | 2008 | The incidence of bacteraemia associated with tracheal intubation |
| Cherry M;Daly CG;Mitchell D;Highfield J; | 2007 | Effect of rinsing with povidone-iodine on bacteraemia due to scaling: a randomized-controlled trial |
| Chiu FY;Chen CM; | 2007 | Surgical debridement and parenteral antibiotics in infected revision total knee arthroplasty |
| Choong PF;Dowsey MM;Carr D;Daffy J;Stanley P; | 2007 | Risk factors associated with acute hip prosthetic joint infections and outcome of treatment with a rifampinbased regimen |
| Cordero-Ampuero J;Esteban J;Garcia-Cimbrelo E;Munuera L;Escobar R; | 2007 | Low relapse with oral antibiotics and two-stage exchange for late arthroplasty infections in 40 patients after 2-9 years |
| Lafaurie GI;Mayorga-Fayad I;Torres MF;Castillo DM;Aya MR;Baron A;Hurtado PA; | 2007 | Periodontopathic microorganisms in peripheric blood after scaling and root planning |

Table 52 Included Studies for Background Microbiology

| Author(s) | Year | Title |
|--------------------------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------|
| Soriano A;Gomez J;Gomez L;Azanza JR;Perez R;Romero F;Pons M;Bella F;Velasco M;Mensa J; | 2007 | Efficacy and tolerability of prolonged linezolid therapy in the treatment of orthopedic implant infections |
| Forner L;Larsen T;Kilian M;Holmstrup P; | 2006 | Incidence of bacteremia after chewing, tooth brushing and scaling in individuals with periodontal inflammation |
| Forner L;Nielsen CH;Bendtzen K;Larsen T;Holmstrup P; | 2006 | Increased plasma levels of IL-6 in bacteremic periodontis patients after scaling |
| Goodman SB;Oh KJ;Imrie S;Hwang K;Shegog M; | 2006 | Revision total hip arthroplasty in juvenile chronic arthritis: 17 revisions in 11 patients followed for 4-12 years |
| Oncag O;Aydemir S;Ersin N;Koca H; | 2006 | Bacteremia incidence in pediatric patients under dental general anesthesia |
| Soriano A;Garcia S;Bori G;Almela M;Gallart X;Macule F;Sierra J;Martinez JA;Suso S;Mensa J; | 2006 | Treatment of acute post-surgical infection of joint arthroplasty |
| Hoad-Reddick DA;Evans CR;Norman P;Stockley I; | 2005 | Is there a role for extended antibiotic therapy in a two-stage revision of the infected knee arthroplasty? |
| Kinane DF;Riggio MP;Walker KF;MacKenzie D;Shearer B; | 2005 | Bacteremia folloing periodontal procedures |
| Oncag O;Cokmez B;Aydemir S;Balcioglu T; | 2005 | Bacteremia incidence in pediatric patients under dental general anesthesia |
| Burden DJ;Coulter WA;Johnston CD;Mullally B;Stevenson M; | 2004 | The prevalence of bacteraemia on removal of fixed orthodontic appliances |

Table 52 Included Studies for Background Microbiology

| Author(s) | Year | Title |
|------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------|
| Rajasuo A;Nyfors S;Kanervo A;Jousimies-Somer H;Lindqvist C;Suuronen R; | 2004 | Bacteremia after plate removal and tooth extraction |
| Jerosch J;Schneppenheim M; | 2003 | Management of infected shoulder replacement |
| Rao N;Crossett LS;Sinha RK;Le Frock JL; | 2003 | Long-term suppression of infection in total joint arthroplasty |
| Soultanis K;Mantelos G;Pagiatakis A;Soucacos PN; | 2003 | Late infection in patients with scoliosis treated with spinal instrumentation |
| Daly CG;Mitchell DH;Highfield JE;Grossberg DE;Stewart D; | 2001 | Bacteremia due to periodontal probing: a clinical and micobiological investigation |
| Lucas V;Roberts GJ; | 2000 | Odontogenic bacteremia following tooth cleaning procedures in children |
| Wagner M;Wagner H; | 2000 | Medium-term results of a modern metal-on-metal system in total hip replacement |
| Waldman BJ;Hostin E;Mont MA;Hungerford DS; | 2000 | Infected total knee arthroplasty treated by arthroscopic irrigation and debridement |
| Erverdi N;Kadir T;Ozkan H;Acar A; | 1999 | Investigation of bacteremia after orthodontic banding |
| Mont MA;Yoon TR;Krackow KA;Hungerford DS; | 1999 | Clinical experience with a proximally porous-coated second- generation cementless total hip prosthesis: minimum 5-year follow-up |
| Crockarell JR;Hanssen AD;Osmon DR;Morrey BF; | 1998 | Treatment of infection with debridement and retention of the components following hip arthroplasty |

Table 52 Included Studies for Background Microbiology

| Author(s) | Year | Title |
|---------------------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------|
| Petrie RS;Hanssen AD;Osmon DR;Ilstrup D; | 1998 | Metal-backed patellar component failure in total knee arthroplasty: a possible risk for late infection |
| Smith JA;Dunn HK;Manaster BJ; | 1998 | Cementless femoral revision arthroplasty. 2- to 5-year results with a modular titanium alloy stem |
| Wimmer C;Nogler M;Frischhut B; | 1998 | Influence of antibiotics on infection in spinal surgery: a prospective study of 110 patients |
| Daly C;Mitchell D;Grossberg D;Highfield J;Stewart D; | 1997 | Bacteremia caused by periodontal probing |
| Mont MA; Waldman B; Banerjee C; Pacheco IH; Hungerford DS; | 1997 | Multiple irrigation, debridement, and retention of components in infected total knee arthroplasty |
| Debelian GJ;Olsen I;Tronstad L; | 1995 | Bacteremia in conjunction with endodontic therapy |
| Goker K;Guvener O; | 1992 | Antibacterial Effects of Ofloxacin, Clindamycin and Sultamicillin on Surgical Removal of Impacted Third Molars |
| Klenerman L;Seal D;Sullens K; | 1991 | Combined prophylactic effect of ultraclean air and cefuroxime for reducing infection in prosthetic surgery |
| Flood TR;Samaranayake LP;MacFarlane TW;McLennan A;MacKenzie D;Carmichael F; | 1990 | Bacteraemia following incision and drainage of dento-alveolar abscesses |
| Heimdahl A;Hall G;Hedberg M;Sandberg H;Soder PO;Tuner K;Nord CE; | 1990 | Detection and Quantitation by Lysis-Filtration of Bacteremia after Different Oral Surgical Procedures |
| Waki MY;Jolkovsky DL;Otomo-Corgel J;Lofthus JE;Nachnani S;Newman MG;Flemmig TF; | 1990 | Effects of subgingival irrigation on bacteremia following scaling and rootplaning |

Table 52 Included Studies for Background Microbiology

| Author(s) | Year | Title |
|---------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------------------|
| Windsor RE;Insall JN;Urs WK;Miller DV;Brause BD; | 1990 | Two-stage reimplantation for the salvage of total knee arthroplasty complicated by infection. Further follow-up and refinement of indications |
| Casolari C;Neglia R;Forabosco A;Galetti R;Fabio U; | 1989 | Incidence of oral bacteremia and antimicrobial prophylaxis |
| Hansen CP;Westh H;Brok KE;Jensen R;Bertelsen S; | 1989 | Bacteraemia following orotracheal intubation and oesophageal balloon dilatation |
| Dinner M;Tjeuw M;Artusio JF; | 1987 | Bacteremia as a Complication of Nasotrachael intubation |
| Shanson DC;Shehata A;Tadayon M;Harris M; | 1987 | Comparison of intravenous teicoplanin with intramuscular amoxycillin for the prophylaxis of streptococcal bacteraemia in dental patients |
| Maskell JP;Carter JL;Boyd RB;Williams RJ; | 1986 | Teicoplanin as a prophylactic antibiotic for dental bacteraemia |
| Wroblewski BM; | 1986 | One-stage revision of infected cemented total hip arthroplasty |
| Lamey PJ;MacFarlane TW;Patton DW;Samaranayake LP;Ferguson MM; | 1985 | Bacteraemia consequential to sialography |
| Ainscow DA;Denham RA; | 1984 | The risk of haematogenous infection in total joint replacements |
| Insall JN;Thompson FM;Brause BD; | 1983 | Two-stage reimplantation for the salvage of infected total knee arthroplasty |
| Marzoni FA;Kelly DR; | 1983 | Bacteremia following cleft palate repaira prospective study |

Table 52 Included Studies for Background Microbiology

| Author(s) | Year | Title |
|------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------------|
| Silver JG;Martin AW;McBride BC; | 1979 | Experimental transient bacteraemias in human subjects with clinically healthy gingivae |
| Shanson DC;Cannon P;Wilks M; | 1978 | Amoxycillin compared with penicillin V for the prophylaxis of dental bacteraemia |
| Baumgartner JC;Heggers JP;Harrison JW; | 1977 | Incidence of bacteremias related to endodontic procedures. II. Surgical endodontics |
| Soliman NA;el-Batawy YA;Abdallah AK; | 1977 | Bacteriologic study of the systemic disturbances accompanying primary teething |
| Baumgartner JC;Heggers JP;Harrison JW; | 1976 | The incidence of bacteremias related to endodontic procedures. I. Nonsurgical endodontics |
| Peterson LJ;Peacock R; | 1976 | The incidence of bacteremia in pediatric patients following tooth extraction |
| Wank HA;Levison ME;Rose LF;Cohen DW; | 1976 | A quantitative measurement of bacteremia and its relationship to plaque control |
| Ramadan AE;Zaki SA;Nour ZM; | 1975 | A study of transient bacteremia following the use of dental floss silk and interdental stimulators |
| Berger SA;Weitzman S;Edberg SC;Casey JI; | 1974 | Bacteremia after the use of an oral irrigation device. A controlled study in subjects with normal-appearing gingiva: comparison with use of toothbrush |
| Brenman HS;Randall E; | 1974 | Local degerming with providone-iodine II. Prior ro gingivectomy |

Table 52 Included Studies for Background Microbiology

| Author(s) | Year | Title |
|------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------|
| De Leo AA;Schoenknecht FD;Anderson MW;Peterson JC; | 1974 | The incidence of bacteremia following oral prophylaxis on pediatric patients |
| Berry FA;Blankenbaker WL;Ball CG; | 1973 | A Comparison of Bacteremia Occurring With Nasotracheal and Orotracheal Intubation |
| Crawford JJ;Sconyers JR;Moriarty JD;King RC;West JF; | 1973 | Bacteremia after tooth extractions studied with the aid of prereduced anaerobically sterilized culture media |
| Lineberger LT;De Marco TJ; | 1973 | Evaluation of transient bacteremia following routine periodontal procedures |
| Sconyers JR;Crawford JJ;Moriarty JD; | 1973 | Relationship of bacteremia to toothbrushing in patients with periodontitis |
| Felix JE;Rosen S;App GR; | 1971 | Detection of bacteremia after the use of an oral irrigation device in subjects with periodontitis |
| Romans AR;App GR; | 1971 | Bacteremia, a result from oral irrigation in subjects with gingivitis |
| Conner HD;Haberman S;Collings CK;Winford TE; | 1967 | Bacteremias following periodontal scaling in patients with healthy appearing gingiva |
| Khairat O; | 1966 | The non-aerobes of post-extraction bacteremia |
| Martin WJ;Schirger A; | 1964 | Prevention of bacteremia after oral surgery |
| Gutverg M; | 1962 | Studies on bacteremia following oral surgery: some prophylactic approaches to bacteremia and the result of tissue examination of excised gingiva |

Table 52 Included Studies for Background Microbiology

| Author(s) | Year | Title |
|--------------------------------------------------------------|------|-----------------------------------------------------------------------------------|
| ROGOSA M;HAMPP EG;NEVIN TA;WAGNER HN;DRISCOLL EJ;Baer PN; | 1960 | Blood sampling and cultural studies in the detection of postoperative bacteremias |
| Winslow MB;KOBERNICK SD; | 1960 | Bacteremia after prophylaxis |

EXCLUDED STUDIES TABLES

RECOMMENDATION 1

Table 53 Excluded Studies for Recommendation 1

| Author(s) | Year | Title | Reason for Exclusion |
|-----------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------|
| Bahrani-Mougeot FK;Paster BJ;Coleman S;Ashar J;Barbuto S;Lockhart PB; | 2008 | Diverse and novel oral bacterial species in blood following dental procedures | Relevant data previously published |
| Jeon HS;Hong SP;Cho BO;Mulyukin A;Choi JY;Kim SG; | 2005 | Hematogenous infection of the human temporomandibular joint | Not best available evidence |
| Roberts GJ;Holzel HS;Sury MR;Simmons NA;Gardner P;Longhurst P; | 1997 | Dental bacteremia in children | Split mouth design |
| Aoki T;Kobayashi I; | 1996 | Blood culture positive rate of 3 media (Bactec(registered trademark), FAN(registered trademark), and VITAL ANA(registered trademark)) after tooth extraction using imipenem | n<10 |
| Kaneko A;Sasaki J;Yamazaki J;Kobayashi I; | 1995 | Intravenous administration of vancomycin is ineffective against bacteremia following tooth extraction | No control group |
| Nohara T;Kobayashi I; | 1995 | Transient bacteremia after tooth extraction with intravenous cefuroxime prophylaxis | No control group |
| Shirai T;Kobayashi I; | 1995 | Transient bacteremia after tooth extraction using ceftriaxone intravenously | No control group |

Table 53 Excluded Studies for Recommendation 1

| Author(s) | Year | Title | Reason for Exclusion |
|------------------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------|
| Sasaki J;Otsuka T;Ozawa H;Takakura J;Kobayashi I; | 1994 | Transient bacteremia after tooth extraction using ampicillin intravenously | No control group |
| Sefton AM;Maskell JP;Kerawala C;Cannell H;Seymour A;Sun ZM;Williams JD; | 1990 | Comparative efficacy and tolerance of erythromycin and josamycin in the prevention of bacteraemia following dental extraction | Duplicate publication |
| Gismondo MR;Nicoletti G; | 1989 | Prophylaxis of dental bacteremia | Insufficient data for analysis |
| Baltch AL;Pressman HL;Schaffer C;Smith RP;Hammer MC;Shayegani M;Michelsen P; | 1988 | Bacteremia in patients undergoing oral procedures. Study following parenteral antimicrobial prophylaxis as recommended by the American Heart Association, 1977 | Insufficient data for analysis |
| Hess J;Holloway Y;Dankert J; | 1983 | Incidence of postextraction bacteremia under penicillin cover in children with cardiac disease | No control group |
| Baltch AL;Pressman HL;Hammer MC;Sutphen NC;Smith RP;Shayegani M; | 1982 | Bacteremia following dental extractions in patients with and without penicillin prophylaxis | Insufficient data for analysis |
| Tolman DE;Schirger A;Martin WJ;Washington JA; | 1972 | Ampicillin administered prophylactically in oral surgery | No control group |
| Martin WJ;Waite DE;Miller JJ;Schirger A; | 1971 | Oral surgery. Cloxacillin for prophylaxis | No control group |

Table 53 Excluded Studies for Recommendation 1

| Author(s) | Year | Title | Reason for Exclusion |
|-------------------------------------------|------|---------------------------------------------------------------------------------------|-----------------------------|
| Benson DD;Waite DE;Hall WH;Carroll GW; | 1970 | Omnipen (ampicillin) for prophylaxis. Prior to oral surgery | No control group |
| Elliott RH;Dunbar JM; | 1968 | Streptococcal bacteraemia in children following dental extractions | Not best available evidence |
| Schirger A; Waite DE; Martin WJ; | 1968 | Erythromycin for prophylaxis prior to oral surgery in patients allergic to panicillin | No control group |
| Waite DE;Schirger A;Martin WJ; | 1967 | Cloxacillin for prophylaxis in oral surgery | No control group |
| Schirger A;Martin WJ;ROYER RO;NEEDHAM GM; | 1960 | Bacterial invasion of blood after oral surgical procedures | Duplicate publication |

Table 54 Excluded Studies for Recommendation 2

| Author(s) | Year | Title | Reason for Exclusion |
|-----------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------|
| Assaf M;Yilmaz S;Kuru B;Ipci SD;Noyun U;Kadir T; | 2007 | Effect of the diode laser on bacteremia associated with dental ultrasonic scaling: a clinical and microbiological study | Split mouth design |
| Aguada E;Olona IL;Salazar MB; | 1997 | Gingival degerming by povidone-iodine irrigation: bacteremia reduction in extraction procedures | Blood drawn from sulcus |
| Rahn R;Diehl O;Schafer V;Shah PM;Fleischer W;Reimer K; | 1994 | The effect of topical Povidone-Iodine and Chlorhexidine on the incidence of bacteremia following dental treatment procedures | Duplicate publication |
| Allison C;Simor AE;Mock D;Tenenbaum HC; | 1993 | Prosol-chlorhexidine irrigation reduces the incidence of bacteremia during ultrasonic scaling with the Cavi-Med: a pilot investigation | Split mouth design |
| Reinhardt RA;Bolton RW;Hlava G; | 1982 | Effect of nonsterile versus sterile water irrigation with ultrasonic scaling on postoperative bacteremias | Split mouth design |
| Witzenberger T;O'Leary TJ;Gillette WB; | 1982 | Effect of a local germicide on the occurrence of bacteremia during subgingival scaling | Split mouth design |
| Madsen KL; | 1975 | Effect of chlorhexidine mouthrinse and periodontal treatment upon bacteremia produced by oral hygiene procedures | Duplicate publication |

Table 54 Excluded Studies for Recommendation 2

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------|-----------------------------|
| Tamini HA;Norwood RS;August AA;Dunkin RT;Eversole LR;Moser EH; | 1975 | Use of antiseptics before injection to minimize incidence of bacteremia | Split mouth design |
| Bartlett RC;Howell RM; | 1973 | Topical vancomycin as a deterrent to bacteremias following dental procedures | Split mouth design |
| Eldirini AH; | 1968 | Effectiveness of epinephrine in local anesthetic solutions on the bacteremia following dental extraction | Not topical antimicrobial |
| Winslow MB;Millstone SH; | 1965 | Bacteremia after prophylaxis | No control group |
| Louis JD; | 1960 | The influence of epinephrine on the incidence of bacteremia | Not topical antimicrobial |

Table 55 Excluded Studies for Recommendation 3

| Author(s) | Year | Title | Reason for Exclusion |
|------------------------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------|--------------------------------------------|
| Lafaurie GI;Mayorga-Fayad I;Torres MF;Castillo DM;Aya MR;Baron A;Hurtado PA; | 2007 | Periodontopathic microorganisms in peripheric blood after scaling and root planing | No statistical test for prognostic factors |
| Tomas I;Alvarez M;Limeres J;Potel C;Medina J;Diz P; | 2007 | Prevalence, duration and aetiology of bacteraemia following dental extractions | Not best available evidence |
| Murphy AM;Daly CG;Mitchell DH;Stewart D;Curtis BH; | 2006 | Chewing fails to induce oral bacteraemia in patients with periodontal disease | No statistical test for prognostic factors |
| Roberts GJ;Gardner P;Longhurst P;Black AE;Lucas VS; | 2000 | Intensity of bacteraemia associated with conservative dental procedures in children | No statistical test for prognostic factors |
| Witzenberger T;O'Leary TJ;Gillette WB; | 1982 | Effect of a local germicide on the occurrence of bacteremia during subgingival scaling | Split mouth design |
| Wank HA;Levison ME;Rose LF;Cohen DW; | 1976 | A quantitative measurement of bacteremia and its relationship to plaque control | Not best available evidence |
| Madsen KL; | 1974 | Effect of chlorhexidine mouthrinse and periodontal treatment upon bacteremia produced by oral hygiene procedures | Not best available evidence |

DENTAL PROCEDURES AND BACTEREMIA

Table 56 Excluded Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title | Reason for Exclusion |
|---------------------------------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------------|-----------------------------|
| Fine DH;Furgang D;McKiernan M;Tereski-Bischio D;Ricci-Nittel D;Zhang P;Araujo MW; | 2010 | An investigation of the effect of an essential oil mouthrinse on induced bacteraemia: a pilot study | Not best available evidence |
| Jones DJ;Munro CL;Grap MJ;Kitten T;Edmond M; | 2010 | Oral care and bacteremia risk in mechanically ventilated adults | Not best available evidence |
| Ashare A;Stanford C;Hancock P;Stark D;Lilli K;Birrer E;Nymon A;Doerschug KC;Hunninghake GW; | 2009 | Chronic liver disease impairs bacterial clearance in a human model of induced bacteremia | Not best available evidence |
| Bahrani-Mougeot FK;Paster BJ;Coleman S;Ashar J;Barbuto S;Lockhart PB; | 2008 | Diverse and novel oral bacterial species in blood following dental procedures | Duplicate publication |
| Lucas VS;Gafan G;Dewhurst S;Roberts GJ; | 2008 | Prevalence, intensity and nature of bacteraemia after toothbrushing | Not best available evidence |
| Assaf M;Yilmaz S;Kuru B;Ipci SD;Noyun U;Kadir T; | 2007 | Effect of the diode laser on bacteremia associated with dental ultrasonic scaling: a clinical and microbiological study | Split mouth design |
| Lucas VS;Kyriazidou A;Gelbier M;Roberts GJ; | 2007 | Bacteraemia following debanding and gold chain adjustment | Not best available evidence |
| Diz DP;Tomas C;Limeres PJ;Medina HJ;Fernandez FJ;Alvarez FM; | 2006 | Comparative efficacies of amoxicillin, clindamycin, and moxifloxacin in prevention of bacteremia following dental extractions | Duplicate publication |

Table 56 Excluded Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title | Reason for Exclusion |
|-------------------------------------------------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------|---------------------------------------|
| Roberts GJ;Jaffray EC;Spratt DA;Petrie A;Greville C;Wilson M;Lucas VS; | 2006 | Duration, prevalence and intensity of bacteraemia after dental extractions in children | Insufficient data for analysis |
| Hartzell JD;Torres D;Kim P;Wortmann G; | 2005 | Incidence of bacteremia after routine tooth brushing | Not best available evidence |
| Rosa EA;Rached RN;Tanaka O;Fronza F;Fronza F;Araujo AR; | 2005 | Preliminary investigation of bacteremia incidence after removal of the Haas palatal expander | n<10 |
| Lucas VS;Omar J;Vieira A;Roberts GJ; | 2002 | The relationship between odontogenic bacteraemia and orthodontic treatment procedures | Not best available evidence |
| Erverdi N;Acar A;Isguden B;Kadir T; | 2001 | Investigation of bacteremia after orthodontic banding and debanding following chlorhexidine mouth wash application | Not best available evidence |
| Vergis EN;Demas PN;Vaccarello SJ;Yu VL; | 2001 | Topical antibiotic prophylaxis for bacteremia after dental extractions | n<10 |
| Erverdi N;Biren S;Kadir T;Acar A; | 2000 | Investigation of bacteremia following orthodontic debanding | Not best available evidence |
| Messini M;Skourti I;Markopulos E;Koutsia-Carouzou C;Kyriakopoulou E;Kostaki S;Lambraki D;Georgopoulos A; | 1999 | Bacteremia after dental treatment in mentally handicapped people | Cannot determine bacteremia incidence |

Table 56 Excluded Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------|
| Roberts GJ;Holzel HS;Sury MR;Simmons NA;Gardner P;Longhurst P; | 1997 | Dental bacteremia in children | Split mouth design |
| McLaughlin JO;Coulter WA;Coffey A;Burden DJ; | 1996 | The incidence of bacteremia after orthodontic banding | Not best available evidence |
| Okabe K;Nakagawa K;Yamamoto E; | 1995 | Factors affecting the occurrence of bacteremia associated with tooth extraction | Not best available evidence |
| Morishima T;Sasaki J; | 1994 | Transient bacteremia after tooth extraction | Cannot determine bacteremia incidence |
| Rahn R;Diehl O;Schafer V;Shah PM;Fleischer W;Reimer K; | 1994 | The effect of topical Povidone-Iodine and Chlorhexidine on the incidence of bacteremia following dental treatment procedures | Duplicate publication |
| Allison C;Simor AE;Mock D;Tenenbaum HC; | 1993 | Prosol-chlorhexidine irrigation reduces the incidence of bacteremia during ultrasonic scaling with the Cavi-Med: a pilot investigation | Split mouth design |
| Yamalik MK;Yucetas S;Abbasoglu U; | 1992 | Effects of various antiseptics on bacteremia following tooth extraction | Not best available evidence |
| Schlein RA;Kudlick EM;Reindorf CA;Gregory J;Royal GC; | 1991 | Toothbrushing and transient bacteremia in patients undergoing orthodontic treatment | Not best available evidence |

Table 56 Excluded Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------|
| Hunter KM;Holborow DW;Kardos TB;Lee-Knight CT;Ferguson MM; | 1989 | Bacteraemia and tissue damage resulting from air polishing | Not best available evidence |
| Baltch AL;Pressman HL;Schaffer C;Smith RP;Hammer MC;Shayegani M;Michelsen P; | 1988 | Bacteremia in patients undergoing oral procedures. Study following parenteral antimicrobial prophylaxis as recommended by the American Heart Association, 1977 | Insufficient data for analysis |
| Lewis HJ;Culligan GA;Pochee E;de Wet FA;Crewe-Brown HH; | 1987 | A microbiological investigation of post- extraction bacteraemia in black subjects | Not best available evidence |
| Roberts GJ;Radford P;Holt R; | 1987 | Prophylaxis of dental bacteraemia with oral amoxycillin in children | Not best available evidence |
| Chung A;Kudlick EM;Gregory JE;Royal GC;Reindorf CA; | 1986 | Toothbrushing and transient bacteremia in patients undergoing orthodontic treatment | Not best available evidence |
| Appleman MD;Sutter VL;Sims TN; | 1982 | Value of antibiotic prophylaxis in periodontal surgery | Not best available evidence |
| Baltch AL;Schaffer C;Hammer MC;Sutphen NT;Smith RP;Conroy J;Shayegani M; | 1982 | Bacteremia following dental cleaning in patients with and without penicillin prophylaxis | Not best available evidence |
| Reinhardt RA;Bolton RW;Hlava G; | 1982 | Effect of nonsterile versus sterile water irrigation with ultrasonic scaling on postoperative bacteremias | Split mouth design |

Table 56 Excluded Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title | Reason for Exclusion |
|---------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------|---------------------------------------|
| Witzenberger T;O'Leary TJ;Gillette WB; | 1982 | Effect of a local germicide on the occurrence of bacteremia during subgingival scaling | Split mouth design |
| Carroll GC;Sebor RJ; | 1980 | Dental flossing and its relationship to transient bacteremia | n<10 |
| Sweet JB;Gill VJ;Chusid MJ;Elin RJ; | 1978 | Nitroblue tetrazolium and Limulus assays for bacteremia after dental extraction: effect of topical antiseptics | Not best available evidence |
| Hockett RN;Loesche WJ;Sodeman TM; | 1977 | Bacteraemia in asymptomatic human subjects | Insufficient data for analysis |
| Nasif AS; | 1977 | The incidence of post-extraction bacteremia after irrigation of the gingival sulcus with hydrogen peroxide solution | Not best available evidence |
| Silver JG;Martin AW;McBride BC; | 1977 | Experimental transient bacteraemias in human subjects with varying degrees of plaque accumulation and gingival inflammation | Not best available evidence |
| Speck WT;Spear SS;Krongrad E;Mandel L;Gersony WM; | 1976 | Transient bacteremia in pediatric patients after dental extraction | Not best available evidence |
| Faigel HC;Gaskill WF; | 1975 | Bacteremia in pediatric patients following dental manipulations | Cannot determine bacteremia incidence |

Table 56 Excluded Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------|---------------------------------------|
| Madsen KL; | 1975 | Effect of chlorhexidine mouthrinse and periodontal treatment upon bacteremia produced by oral hygiene procedures | Duplicate publication |
| Symington JM; | 1975 | Streptococci isolated from post- extraction bacteraemias | Insufficient data for analysis |
| Tamini HA;Norwood RS;August AA;Dunkin RT;Eversole LR;Moser EH; | 1975 | Use of antiseptics before injection to minimize incidence of bacteremia | Split mouth design |
| Madsen KL; | 1974 | Effect of chlorhexidine mouthrinse and periodontal treatment upon bacteremia produced by oral hygiene procedures | Cannot determine bacteremia incidence |
| Bartlett RC;Howell RM; | 1973 | Topical vancomycin as a deterrent to bacteremias following dental procedures | Split mouth design |
| Berry FA;Yarbrough S;Yarbrough N;Russell CM;Carpenter MA;Hendley JO; | 1973 | Transient bacteremia during dental manipulation in children | Cannot determine bacteremia incidence |
| Farrington FH; | 1973 | The incidence of transient bacteremia following pulpotomies on primary teeth | Not best available evidence |
| Cutcher JL;Goldberg JR;Lilly GE;Jones JC; | 1971 | Control of bacteremia associated with extraction of teeth. II | Not best available evidence |
| Hurwitz GA;Speck WT;Keller GB; | 1971 | Absence of bacteremia in children after prophylaxis | Not best available evidence |

Table 56 Excluded Studies for Dental Procedures and Bacteremia

| Author(s) | Year | Title | Reason for Exclusion |
|------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------|--------------------------------|
| Speck WT;Hurwitz GA;Keller GB; | 1971 | Transient bacteremia in pediatric patients following dental manipulatin | Not best available evidence |
| Tamimi HA;Thomassen PR;Moser EH; | 1969 | Bacteremia study using a water irrigation device | Not best available evidence |
| de Vries JA;Francis LE;Platonow M; | 1968 | Adjunctive use of antibiotics in traumatic dental procedures | Insufficient data for analysis |
| Eldirini AH; | 1968 | Effectiveness of epinephrine in local anesthetic solutions on the bacteremia following dental extraction | Not best available evidence |
| Khairat O; | 1966 | An effective antibiotic cover for the prevention of endocarditis following dental and other post-operative bacteraemias | Not best available evidence |

BACKGROUND MICROBIOLOGY

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Le D;Smith K;Tanzer D;Tanzer M; | 2011 | Modular femoral sleeve and stem implant provides long-term total hip survivorship | Insufficient data for analysis |
| Aslam S;Reitman C;Darouiche RO; | 2010 | Risk factors for subsequent diagnosis of prosthetic joint infection | Retrospective study |
| Barbosa M;Carmona IT;Amaral B;Limeres J;Alvarez M;Cerqueira C;Diz P; | 2010 | General anesthesia increases the risk of bacteremia following dental extractions | Insufficient data on bacteremia for background microbiology |
| Burnett RS;Aggarwal A;Givens SA;McClure JT;Morgan PM;Barrack RL; | 2010 | Prophylactic antibiotics do not affect cultures in the treatment of an infected TKA: a prospective trial | Insufficient data for analysis |
| Cordero-Ampuero J;Esteban J;Garcia-Rey E; | 2010 | Results after late polymicrobial, gram- negative, and methicillin-resistant infections in knee arthroplasty | Insufficient data for analysis |
| Erhart J;Jaklitsch K;Schurz M;Vecsei V;Ehall R; | 2010 | Cementless two-staged total hip arthroplasty with a short term interval period for chronic deep periprosthetic infection. Technique and long-term results | Review |
| Estes CS;Beauchamp CP;Clarke HD;Spangehl MJ; | 2010 | A two-stage retention debridement protocol for acute periprosthetic joint infections | Retrospective study |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|-------------------------------------------------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------|--------------------------------|
| Goddard NJ;Mann HA;Lee CA; | 2010 | Total knee replacement in patients with end-stage haemophilic arthropathy: 25-year results | Retrospective study |
| McCleery MA;Leach WJ;Norwood T; | 2010 | Rates of infection and revision in patients with renal disease undergoing total knee replacement in Scotland | Insufficient data for analysis |
| Ocguder A;Firat A;Tecimel O;Solak S;Bozkurt M; | 2010 | Two-stage total infected knee arthroplasty treatment with articulating cement spacer | Insufficient data for analysis |
| Ritter MA;Farris A; | 2010 | Outcome of Infected Total Joint Replacement | Retrospective study |
| Rodriguez D;Pigrau C;Euba G;Cobo J;Garcia-Lechuz J;Palomino J;Riera M;Del Toro MD;Granados A;Ariza X; | 2010 | Acute haematogenous prosthetic joint infection: prospective evaluation of medical and surgical management | Duplicate Publication |
| Sousa R;Pereira A;Massada M;da Silva MV;Lemos R;Costa e Castro; | 2010 | Empirical antibiotic therapy in prosthetic joint infections | Retrospective study |
| Zywiel MG;Johnson AJ;Stroh DA;Martin J;Marker DR;Mont MA; | 2010 | Prophylactic oral antibiotics reduce reinfection rates following two-stage revision total knee arthroplasty | Retrospective study |
| Bin D;Noble PC; | 2009 | Aseptic loosening of cemented stem following cemented hip arthroplasty: Analysis of 36 revised specimens | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|---------------------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------|
| Byren I;Bejon P;Atkins BL;Angus B;Masters S;McLardy-Smith P;Gundle R;Berendt A; | 2009 | One hundred and twelve infected arthroplasties treated with 'DAIR' (debridement, antibiotics and implant retention): antibiotic duration and outcome | Retrospective study |
| Carrington NC;Sierra RJ;Gie GA;Hubble MJ;Timperley AJ;Howell JR; | 2009 | The Exeter Universal cemented femoral component at 15 to 17 years: an update on the first 325 hips | Insufficient data for analysis |
| Cavusoglu AT;Er MS;Inal S;Ozsoy MH;Dincel VE;Sakaogullari A; | 2009 | Pin site care during circular external fixation using two different protocols | Insufficient data for analysis |
| Chen WS;Fu TH;Wang JW; | 2009 | Two-stage reimplantation of infected hip arthroplasties | Insufficient data for analysis |
| Dale H;Hallan G;Hallan G;Espehaug B;Havelin LI;Engesaeter LB; | 2009 | Increasing risk of revision due to deep infection after hip arthroplasty | Retrospective study |
| Dauchy FA;Dupon M;Dutronc H;de BB;Lawson-Ayayi S;Dubuisson V;Souillac V; | 2009 | Association between psoas abscess and prosthetic hip infection: a case-control study | Insufficient data for analysis |
| Goebel D;Schultz W; | 2009 | The Mayo cementless femoral component in active patients with osteoarthritis | Insufficient data for analysis |
| Hooper GJ;Rothwell AG;Stringer M;Frampton C; | 2009 | Revision following cemented and uncemented primary total hip replacement: a seven-year analysis from the New Zealand Joint Registry | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|-----------------------------------------------------------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Martinez-Pastor JC;Munoz-Mahamud E;Vilchez F;Garcia-Ramiro S;Bori G;Sierra J;Martinez JA;Font L;Mensa J;Soriano A; | 2009 | Outcome of acute prosthetic joint infections due to gram-negative bacilli treated with open debridement and retention of the prosthesis | Retrospective study |
| Nixon PP;Littler P;Davies K;Krishnam MS; | 2009 | Does sialography require antibiotic prophylaxis? | Insufficient data on bacteremia for background microbiology |
| Ong KL;Kurtz SM;Lau E;Bozic KJ;Berry DJ;Parvizi J; | 2009 | Prosthetic joint infection risk after total hip arthroplasty in the Medicare population | Insufficient data for analysis |
| Ren W;Blasier R;Peng X;Shi T;Wooley PH;Markel D; | 2009 | Effect of oral erythromycin therapy in patients with aseptic loosening of joint prostheses | Insufficient data for analysis |
| Sonbol H;Spratt D;Roberts GJ;Lucas VS; | 2009 | Prevalence, intensity and identity of bacteraemia following conservative dental procedures in children | Insufficient data on bacteremia for background microbiology |
| Stefansdottir A;Johansson D;Knutson K;Lidgren L;Robertsson O; | 2009 | Microbiology of the infected knee arthroplasty: report from the Swedish Knee Arthroplasty Register on 426 surgically revised cases | Retrospective study |
| Tintle SM;Forsberg JA;Potter BK;Islinger RB;Andersen RC; | 2009 | Prosthesis retention, serial debridement, and antibiotic bead use for the treatment of infection following total joint arthroplasty | Review |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Yoo JJ;Kwon YS;Koo KH;Yoon KS;Kim YM;Kim HJ; | 2009 | One-stage cementless revision arthroplasty for infected hip replacements | Review |
| Zeller V;Lavigne M;Leclerc P;Lhotellier L;Graff W;Ziza JM;Desplaces N;Mamoudy P; | 2009 | Group B streptococcal prosthetic joint infections: a retrospective study of 30 cases | Retrospective study |
| Brook I; | 2008 | Microbiology and management of joint and bone infections due to anaerobic bacteria | Review |
| Gosheger G;Goetze C;Hardes J;Joosten U;Winkelmann W;von EC; | 2008 | The influence of the alloy of megaprostheses on infection rate | Retrospective study |
| Lau TW;Leung F;Chan CF;Chow SP; | 2008 | Wound complication of minimally invasive plate osteosynthesis in distal tibia fractures | Retrospective study |
| Leclercq S;Benoit JY;de Rosa JP;Euvrard P;Leteurtre C;Girardin P; | 2008 | Results of the Evora dual-mobility socket after a minimum follow-up of five years | Insufficient data for analysis |
| Lockhart PB;Brennan MT;Sasser HC;Fox PC;Paster BJ;Bahrani-Mougeot FK; | 2008 | Bacteremia associated with toothbrushing and dental extraction | Insufficient data on bacteremia for background microbiology |
| Oussedik SI;Haddad FS; | 2008 | The use of linezolid in the treatment of infected total joint arthroplasty | Retrospective study |
| Parvizi J;Ghanem E;Azzam K;Davis E;Jaberi F;Hozack W; | 2008 | Periprosthetic infection: are current treatment strategies adequate? | Retrospective study |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|-------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Poeschl PW;Ploder O;Seemann R;Poeschl E; | 2008 | Maxillomandibular fixation using intraoral cortical bone screws and specially designed metal hooks (Ottenhaken) in the conservative treatment of mandibular fractures | Insufficient data for analysis |
| Ritter MA;Meneghini RM; | 2008 | Twenty-year survivorship of cementless anatomic graduated component (AGC) total knee replacement | Insufficient data for analysis |
| Schafer P;Fink B;Sandow D;Margull A;Berger I;Frommelt L; | 2008 | Prolonged bacterial culture to identify late periprosthetic joint infection: a promising strategy | Insufficient data for analysis |
| Tomas I;Pereira F;Llucian R;Poveda R;Diz P;Bagan JV; | 2008 | Prevalence of bacteraemia following third molar surgery | Insufficient data on bacteremia for background microbiology |
| Aboltins CA;Page MA;Buising KL;Jenney AW;Daffy JR;Choong PF;Stanley PA; | 2007 | Treatment of staphylococcal prosthetic joint infections with debridement, prosthesis retention and oral rifampicin and fusidic acid | Retrospective study |
| Byrne AM;Morris S;McCarthy T;Quinlan W;O'byrne JM; | 2007 | Outcome following deep wound contamination in cemented arthroplasty | Study on perioperative contamination |
| Cook JL;Scott RD;Long WJ; | 2007 | Late hematogenous infections after total knee arthroplasty: experience with 3013 consecutive total knees | Retrospective study |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Frances A;Moro E;Cebrian JL;Marco F;Garcia-Lopez A;Serfaty D;Lopez-Duran L; | 2007 | Reconstruction of bone defects with impacted allograft in femoral stem revision surgery | Insufficient data for analysis |
| Kowalski TJ;Berbari EF;Huddleston PM;Steckelberg JM;Mandrekar JN;Osmon DR; | 2007 | The management and outcome of spinal implant infections: contemporary retrospective cohort study | Retrospective study |
| Rao N;Hamilton CW; | 2007 | Efficacy and safety of linezolid for Gram-positive orthopedic infections: a prospective case series | Insufficient data for analysis |
| Renvert S;Roos-Jansaker AM;Lindahl C;Renvert H;Rutger PG; | 2007 | Infection at titanium implants with or without a clinical diagnosis of inflammation | Insufficient data for analysis |
| Sundararaj GD;Babu N;Amritanand R;Venkatesh K;Nithyananth M;Cherian VM;Lee VN; | 2007 | Treatment of haematogenous pyogenic vertebral osteomyelitis by single-stage anterior debridement, grafting of the defect and posterior instrumentation | Insufficient data for analysis |
| Tomas I;Alvarez M;Limeres J;Potel C;Medina J;Diz P; | 2007 | Prevalence, duration and aetiology of bacteraemia following dental extractions | Insufficient data on bacteremia for background microbiology |
| Tomas I;Alvarez M;Limeres J;Tomas M;Medina J;Otero JL;Diz P; | 2007 | Effect of a chlorhexidine mouthwash on the risk of postextraction bacteremia | Insufficient data on bacteremia for background microbiology |
| You JH;Lee GC;So RK;Cheung KW;Hui M; | 2007 | Linezolid versus vancomycin for prosthetic joint infections: a cost analysis | Simulation model |
| | | | |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|-----------------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Barberan J; | 2006 | Management of infections of osteoarticular prosthesis | Review |
| Barberan J;Aguilar L;Carroquino G;Gimenez MJ;Sanchez B;Martinez D;Prieto J; | 2006 | Conservative treatment of staphylococcal prosthetic joint infections in elderly patients | Retrospective study |
| Comba F;Buttaro M;Pusso R;Piccaluga F; | 2006 | Acetabular reconstruction with impacted bone allografts and cemented acetabular components: a 2- to 13-year follow-up study of 142 aseptic revisions | Insufficient data for analysis |
| Diz DP;Tomas C;Limeres PJ;Medina HJ;Fernandez FJ;Alvarez FM; | 2006 | Comparative efficacies of amoxicillin, clindamycin, and moxifloxacin in prevention of bacteremia following dental extractions | Insufficient data on bacteremia for background microbiology |
| Engesaeter LB;Espehaug B;Lie SA;Furnes O;Havelin LI; | 2006 | Does cement increase the risk of infection in primary total hip arthroplasty? Revision rates in 56,275 cemented and uncemented primary THAs followed for 0-16 years in the Norwegian Arthroplasty Register | Insufficient data for analysis |
| Fulkerson E;Valle CJ;Wise B;Walsh M;Preston C;Di Cesare PE; | 2006 | Antibiotic susceptibility of bacteria infecting total joint arthroplasty sites | Review |
| Laffer RR;Graber P;Ochsner PE;Zimmerli W; | 2006 | Outcome of prosthetic knee-associated infection: evaluation of 40 consecutive episodes at a single centre | Retrospective study |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|---------------------------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Lin C;Hsu H;Huang C;Chen S; | 2006 | Late-onset infection of total knee arthroplasty caused by the Klebsiella pneumoniae bacteremia | Insufficient data for analysis |
| Lindeboom JA;Frenken JW;Tuk JG;Kroon FH; | 2006 | A randomized prospective controlled trial of antibiotic prophylaxis in intraoral bone-grafting procedures: preoperative single-dose penicillin versus preoperative single-dose clindamycin | Insufficient data for analysis |
| Lotke PA;Carolan GF;Puri N; | 2006 | Impaction grafting for bone defects in revision total knee arthroplasty | Insufficient data for analysis |
| Murphy AM;Daly CG;Mitchell DH;Stewart D;Curtis BH; | 2006 | Chewing fails to induce oral bacteraemia in patients with periodontal disease | Insufficient data on bacteremia for background microbiology |
| Rallis G;Mourouzis C;Papakosta V;Papanastasiou G;Zachariades N; | 2006 | Reasons for miniplate removal following maxillofacial trauma: a 4-year study | Insufficient data for analysis |
| Theodossy T;Jackson O;Petrie A;Lloyd T; | 2006 | Risk factors contributing to symptomatic plate removal following sagittal split osteotomy | Insufficient data for analysis |
| Bassetti M;Vitale F;Melica G;Righi E;Di BA;Molfetta L;Pipino F;Cruciani M;Bassetti D; | 2005 | Linezolid in the treatment of Gram- positive prosthetic joint infections | Retrospective study |
| Belthur MV;Bradish CF;Gibbons PJ; | 2005 | Late orthopaedic sequelae following meningococcal septicaemia. A multicentre study | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------------------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Chu VH;Crosslin DR;Friedman JY;Reed SD;Cabell CH;Griffiths RI;Masselink LE;Kaye KS;Corey GR;Reller LB;Stryjewski ME;Schulman KA;Fowler VG; | 2005 | Staphylococcus aureus bacteremia in patients with prosthetic devices: costs and outcomes | Insufficient data for analysis |
| Khatri M;Stirrat AN; | 2005 | Souter-Strathclyde total elbow arthroplasty in rheumatoid arthritis: medium-term results | Retrospective study |
| Marculescu CE;Berbari EF;Hanssen AD;Steckelberg JM;Osmon DR; | 2005 | Prosthetic joint infection diagnosed postoperatively by intraoperative culture | Retrospective study |
| Silva M;Luck JV; | 2005 | Long-term results of primary total knee replacement in patients with hemophilia | Review |
| Takai S;Kuriyama T;Yanagisawa M;Nakagawa K;Karasawa T; | 2005 | Incidence and bacteriology of bacteremia associated with various oral and maxillofacial surgical procedures | Insufficient data on bacteremia for background microbiology |
| Durbhakula SM;Czajka J;Fuchs MD;Uhl RL; | 2004 | Spacer endoprosthesis for the treatment of infected total hip arthroplasty | Retrospective study |
| Forster H;Marotta JS;Heseltine K;Milner R;Jani S; | 2004 | Bactericidal activity of antimicrobial coated polyurethane sleeves for external fixation pins | Insufficient data for analysis |
| Ikavalko M;Belt EA;Kautiainen H;Lehto MU; | 2004 | Souter arthroplasty for elbows with severe destruction | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|-------------------------------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Muschik M;Luck W;Schlenzka D; | 2004 | Implant removal for late-developing infection after instrumented posterior spinal fusion for scoliosis: reinstrumentation reduces loss of correction. A retrospective analysis of 45 cases | Retrospective study |
| Pavoni GL;Giannella M;Falcone M;Scorzolini L;Liberatore M;Carlesimo B;Serra P;Venditti M; | 2004 | Conservative medical therapy of prosthetic joint infections: retrospective analysis of an 8-year experience | Retrospective study |
| Rajasuo A;Perkki K;Nyfors S;Jousimies- Somer H;Meurman JH; | 2004 | Bacteremia Following Surgical Dental Extraction with an Emphasis on Anaerobic Stra | Insufficient data on bacteremia for background microbiology |
| Rao N;Ziran BH;Hall RA;Santa ER; | 2004 | Successful treatment of chronic bone and joint infections with oral linezolid | Insufficient data for analysis |
| Savarrio L;MacKenzie D;Riggio M;Saunders WP;Bagg J; | 2004 | Detection of bacteraemias during non- surgicalroot canal treatment | Insufficient data on bacteremia for background microbiology |
| Stavrev VP;Stavrev PV; | 2004 | Complications in total hip replacement | Insufficient data for analysis |
| Bago J;Ramirez M;Pellise F;Villanueva C; | 2003 | Survivorship analysis of Cotrel- Dubousset instrumentation in idiopathic scoliosis | Retrospective study |
| Davis III CM;Berry DJ;Harmsen WS; | 2003 | Cemented revision of failed uncemented femoral components of total hip arthroplasty | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Engesaeter LB;Lie SA;Espehaug B;Furnes O;Vollset SE;Havelin LI; | 2003 | Antibiotic prophylaxis in total hip arthroplasty: Effects of antibiotic prophylaxis systemically and in bone cement on the revision rate of 22,170 primary hip replacements followed 0-14 years in the Norwegian Arthroplasty Register | Insufficient data for analysis |
| Gallo J;Kolar M;Novotny R;Rihakova P;Ticha V; | 2003 | Pathogenesis of prosthesis-related infection | Review |
| Ross JJ;Saltzman CL;Carling P;Shapiro DS; | 2003 | Pneumococcal septic arthritis: review of 190 cases | Retrospective study |
| Bhanji S;Williams B;Sheller B;Elwood T;Mancl L; | 2002 | Transient bacteremia induced by toothbrushing a comparison of the Sonicare toothbrush with a conventional toothbrush | Insufficient data on bacteremia for background microbiology |
| Husted H;Toftgaard JT; | 2002 | Clinical outcome after treatment of infected primary total knee arthroplasty | Retrospective study |
| Norian JM;Ries MD;Karp S;Hambleton J; | 2002 | Total knee arthroplasty in hemophilic arthropathy | Retrospective study |
| Perkins TR;Gunckle W; | 2002 | Unicompartmental knee arthroplasty: 3- to 10-year results in a community hospital setting | Insufficient data for analysis |
| van Koeveringe AJ;Ochsner PE; | 2002 | Revision cup arthroplasty using Burch- Schneider anti-protrusio cage | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|---------------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Acklin YP;Berli BJ;Frick W;Elke R;Morscher EW; | 2001 | Nine-year results of Muller cemented titanium Straight Stems in total hip replacement | Insufficient data for analysis |
| Chiu KY;Ng TP;Tang WM;Poon KC;Ho WY;Yip D; | 2001 | Charnley total hip arthroplasty in Chinese patients less than 40 years old | Insufficient data for analysis |
| Fowler VG;Fey PD;Reller LB;Chamis AL;Corey GR;Rupp ME; | 2001 | The intercellular adhesin locus ica is present in clinical isolates of Staphylococcus aureus from bacteremic patients with infected and uninfected prosthetic joints | Insufficient data for analysis |
| Ikavalko M;Lehto MU; | 2001 | Fractured rheumatoid elbow: treatment with Souter elbow arthroplastya clinical and radiologic midterm follow-up study | Insufficient data for analysis |
| Murdoch DR;Roberts SA;Fowler Jr VGJ;Shah MA;Taylor SL;Morris AJ;Corey GR; | 2001 | Infection of orthopedic prostheses after Staphylococcus aureus bacteremia | Insufficient data for analysis |
| Richards BR;Emara KM; | 2001 | Delayed infections after posterior TSRH spinal instrumentation for idiopathic scoliosis: revisited | Retrospective study |
| Vergis EN;Demas PN;Vaccarello SJ;Yu VL; | 2001 | Topical antibiotic prophylaxis for bacteremia after dental extractions | Insufficient data on bacteremia for background microbiology |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Abumi K;Saita M;Iida T;Kaneda K; | 2000 | Reduction and fixation of sacroiliac joint dislocation by the combined use of S1 pedicle screws and the galveston technique | Insufficient data for analysis |
| De LF;Viola R;Pellizzer G;Lazzarini L;Tramarin A;Fabris P; | 2000 | Regional prophylaxis with teicoplanin in monolateral or bilateral total knee replacement: an open study | Insufficient data for analysis |
| Gordon JE;Kelly-Hahn J;Carpenter CJ;Schoenecker PL; | 2000 | Pin site care during external fixation in children: results of a nihilistic approach | Insufficient data for analysis |
| Houshian S;Zawadski AS;Riegels- Nielsen P; | 2000 | Duration of postoperative antibiotic therapy following revision for infected knee and hip arthroplasties | Retrospective study |
| Mohler DG;Kessler JI;Earp BE; | 2000 | Augmented amputations of the lower extremity | Retrospective study |
| Roberts GJ;Gardner P;Longhurst P;Black AE;Lucas VS; | 2000 | Intensity of bacteraemia associated with conservative dental procedures in children | Insufficient data on bacteremia for background microbiology |
| Aydinli U;Karaeminogullari O;Tiskaya K; | 1999 | Postoperative deep wound infection in instrumented spinal surgery | Retrospective study |
| Brown EC;Lachiewicz PF; | 1999 | Precoated femoral component in total hip arthroplasty. Results of 5- to 9-year followup | Retrospective study |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|-----------------------------------------------------------------------|------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Dearborn JT;Harris WH; | 1999 | High placement of an acetabular component inserted without cement in a revision total hip arthroplasty. Results after a mean of ten years | Retrospective study |
| Fehring TK;Calton TF;Griffin WL; | 1999 | Cementless fixation in 2-stage reimplantation for periprosthetic sepsis | Retrospective study |
| Hyman JL;Salvati EA;Laurencin CT;Rogers DE;Maynard M;Brause DB; | 1999 | The arthroscopic drainage, irrigation, and debridement of late, acute total hip arthroplasty infections: average 6-year follow-up | Retrospective study |
| Isiklar ZU;Demirors H;Akpinar S;Tandogan RN;Alparslan M; | 1999 | Two-stage treatment of chronic staphylococcal orthopaedic implant-related infections using vancomycin impregnated PMMA spacer and rifampin containing antibiotic protocol | Insufficient data for analysis |
| Leopold SS;Berger RA;Rosenberg AG;Jacobs JJ;Quigley LR;Galante JO; | 1999 | Impaction allografting with cement for revision of the femoral component. A minimum four-year follow-up study with use of a precoated femoral stem | Insufficient data for analysis |
| Lucas V;Roberts GJ; | 1999 | Odontogenic bacteremia following tooth cleaning procedures in children | Insufficient data on bacteremia for background microbiology |
| Mont MA;Yoon TR;Krackow KA;Hungerford DS; | 1999 | Eliminating patellofemoral complications in total knee arthroplasty: clinical and radiographic results of 121 consecutive cases using the Duracon system | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Segawa H;Tsukayama DT;Kyle RF;Becker DA;Gustilo RB; | 1999 | Infection after total knee arthroplasty. A retrospective study of the treatment of eighty-one infections | Retrospective study |
| Vena VE;Hsu J;Rosier RN;O'Keefe RJ; | 1999 | Pelvic reconstruction for severe periacetabular metastatic disease | Retrospective study |
| Bohm P;Bosche R; | 1998 | Survival analysis of the Harris-Galante I acetabular cup | Insufficient data for analysis |
| Hartofilakidis G;Stamos K;Karachalios T; | 1998 | Treatment of high dislocation of the hip in adults with total hip arthroplasty. Operative technique and long-term clinical results | Insufficient data for analysis |
| Kofoed H;Sorensen TS; | 1998 | Ankle arthroplasty for rheumatoid arthritis and osteoarthritis: prospective long-term study of cemented replacements | Insufficient data for analysis |
| Lo NN;Tan JS;Tan SK;Vathsala A; | 1998 | Results of total hip replacement in renal transplant recipients | Insufficient data for analysis |
| Roberts GJ;Simmons NB;Longhurst P;Hewitt PB; | 1998 | Bacteraemia following local anaesthetic injections in children | Insufficient data on bacteremia for background microbiology |
| Roberts GJ;Watts R;Longhurst P;Gardner P; | 1998 | Bacteremia of dental origin and antimicrobial sensitivity following oral surgical procedures in children | Insufficient data on bacteremia for background microbiology |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------|
| Stein A;Bataille JF;Drancourt M;Curvale G;Argenson JN;Groulier P;Raoult D; | 1998 | Ambulatory treatment of multidrug- resistant Staphylococcus-infected orthopedic implants with high-dose oral co-trimoxazole (trimethoprim- sulfamethoxazole) | Insufficient data for analysis |
| Diduch DR;Insall JN;Scott WN;Scuderi GR;Font-Rodriguez D; | 1997 | Total knee replacement in young, active patients. Long-term follow-up and functional outcome | Insufficient data for analysis |
| Drancourt M;Stein A;Argenson JN;Roiron R;Groulier P;Raoult D; | 1997 | Oral treatment of Staphylococcus spp. infected orthopaedic implants with fusidic acid or ofloxacin in combination with rifampicin | Insufficient data for analysis |
| Grunig R;Morscher E;Ochsner PE; | 1997 | Three-to 7-year results with the uncemented SL femoral revision prosthesis | Insufficient data for analysis |
| Kaandorp CJ;Dinant HJ;van de Laar MA;Moens HJ;Prins AP;Dijkmans BA; | 1997 | Incidence and sources of native and prosthetic joint infection: a community based prospective survey | Insufficient data for analysis |
| Madey SM;Callaghan JJ;Olejniczak JP;Goetz DD;Johnston RC; | 1997 | Charnley total hip arthroplasty with use of improved techniques of cementing. The results after a minimum of fifteen years of follow-up | Insufficient data for analysis |
| McLaughlin JR;Lee KR; | 1997 | Total hip arthroplasty with an uncemented femoral component. Excellent results at ten-year follow-up | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Nijhof MW;Oyen WJ;van KA;Claessens RA;van der Meer JW;Corstens FH; | 1997 | Hip and knee arthroplasty infection. In- 111-IgG scintigraphy in 102 cases | Retrospective study |
| Ozaki T;Hillmann A;Bettin D;Wuisman P;Winkelmann W; | 1997 | Intramedullary, antibiotic-loaded cemented, massive allografts for skeletal reconstruction. 26 cases compared with 19 uncemented allografts | Insufficient data for analysis |
| Roberts GJ;Holzel HS;Sury MR;Simmons NA;Gardner P;Longhurst P; | 1997 | Dental bacteremia in children | Insufficient data on bacteremia for background microbiology |
| Hauser R;Berchtold W;Schreiber A; | 1996 | Incidence of deep sepsis in uncemented total hip arthroplasty using clean air facility as a function of antibiotic prophylaxis | Retrospective study |
| Lai KA;Shen WJ;Yang CY;Lin RM;Lin CJ;Jou IM; | 1996 | Two-stage cementless revision THR after infection. 5 recurrences in 40 cases followed 2.5-7 years | Retrospective study |
| Lu H;Mehdi G;Zhou D;Lin J; | 1996 | Simultaneous bilateral total knee arthroplasty for rheumatoid arthritis | Insufficient data for analysis |
| Silverton C;Rosenberg AO;Barden RM;Sheinkop MB;Galante JO; | 1996 | The prosthesis-bone interface adjacent to tibial components inserted without cement. Clinical and radiographic follow-up at nine to twelve years | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|------------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------|--------------------------------|
| Tsukayama DT;Estrada R;Gustilo RB; | 1996 | Infection after total hip arthroplasty. A study of the treatment of one hundred and six infections | Retrospective study |
| Wimmer C;Gluch H; | 1996 | Management of postoperative wound infection in posterior spinal fusion with instrumentation | Retrospective study |
| Aglietti P;Buzzi R;Segoni F;Zaccherotti G; | 1995 | Insall-Burstein posterior-stabilized knee prosthesis in rheumatoid arthritis | Insufficient data for analysis |
| Hanssen AD;Trousdale RT;Osmon DR; | 1995 | Patient outcome with reinfection following reimplantation for the infected total knee arthroplasty | Retrospective study |
| Bell RS;Davis A;Allan DG;Langer F;Czitrom AA;Gross AE; | 1994 | Fresh osteochondral allografts for advanced giant cell tumors at the knee | Insufficient data for analysis |
| Ivarsson I;Wahlstrom O;Djerf K;Jacobsson SA; | 1994 | Revision of infected hip replacement. Two-stage procedure with a temporary gentamicin spacer | Retrospective study |
| Mauriello JA;Hargrave S;Yee S;Mostafavi R;Kapila R; | 1994 | Infection after insertion of alloplastic orbital floor implants | Retrospective study |
| Nasser S; | 1994 | The incidence of sepsis after total hip replacement arthroplasty | Insufficient data for analysis |
| Petrou G;Gavras M;Diamantopoulos A;Kapetsis T;Kremmydas N;Kouzoupis A; | 1994 | Uncemented total hip replacements and thigh pain | Retrospective study |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|---------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Rahn R;Schneider S;Diehl O;Schafer V;Shah PM; | 1994 | Preventing post-treatment bacteremia: comparing topical povidone-iodine and chlorhexidine | Insufficient data on bacteremia for background microbiology |
| Stromberg CN;Herberts P; | 1994 | A multicenter 10-year study of cemented revision total hip arthroplasty in patients younger than 55 years old. A follow-up report | Insufficient data for analysis |
| Drancourt M;Stein A;Argenson JN;Zannier A;Curvale G;Raoult D; | 1993 | Oral rifampin plus ofloxacin for treatment of Staphylococcus-infected orthopedic implants | Insufficient data for analysis |
| Laus M;Pignatti G;Tigani D;Alfonso C;Giunti A; | 1993 | Anterior decompression and plate fixation in fracture dislocations of the lower cervical spine | Insufficient data for analysis |
| Moeckel B;Huo MH;Salvati EA;Pellicci PM; | 1993 | Total hip arthroplasty in patients with diabetes mellitus | Insufficient data for analysis |
| Putz PA; | 1993 | A pilot study of oral fleroxacin given once daily in patients with bone and joint infections | Insufficient data for analysis |
| Riska EB; | 1993 | Ceramic endoprosthesis in total hip arthroplasty | Insufficient data for analysis |
| Ali MT;Tremewen DR;Hay AJ;Wilkinson DJ; | 1992 | The occurrence of bacteremia associated with the use of oral and nasopharyngeal airways | Insufficient data on bacteremia for background microbiology |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Davey PG;Rowley DR;Phillips GA; | 1992 | Teicoplaninhome therapy for prosthetic joint infections | Insufficient data for analysis |
| Lucartorto FM;Franker CK;Maza J; | 1992 | Postscaling becteremia in HIV-associated gingivitis and periodontitis | Insufficient data on bacteremia for background microbiology |
| Mason JC;Dollery CT;So A;Cohen J;Bloom SR;Bulpitt C;Russell-Jones R;Oakley CM; | 1992 | An infected prosthetic hip. Is there a role for prophylactic antibiotics? | Retrospective study |
| Mombelli A;Lang NP; | 1992 | Antimicrobial treatment of peri-implant infections | Insufficient data for analysis |
| Roberts GJ;Gardner P;Simmons NA; | 1992 | Optimum sampling time for detection of dental bacteraemia in children | Insufficient data on bacteremia for background microbiology |
| Schmalzried TP;Amstutz HC;Au MK;Dorey FJ; | 1992 | Etiology of deep sepsis in total hip arthroplasty. The significance of hematogenous and recurrent infections | Retrospective study |
| Armstrong RA; Whiteside LA; | 1991 | Results of cementless total knee arthroplasty in an older rheumatoid arthritis population | Insufficient data for analysis |
| Lofthus JE;Waki MY;Jolkovsky DL;Otomo-Corgel J;Newman MG;Flemmig T;Nachnani S; | 1991 | Bacteremia following subgingival irrigation and scaling and root planing | Insufficient data on bacteremia for background microbiology |
| Mathiesen EB;Lindgren JU;Blomgren GG;Reinholt FP; | 1991 | Corrosion of modular hip prostheses | Retrospective study |

Table 57 Excluded Studies for Background Microbiology

| _Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Rasul AT;Tsukayama D;Gustilo RB; | 1991 | Effect of time of onset and depth of infection on the outcome of total knee arthroplasty infections | Retrospective study |
| Sanderson PJ; | 1991 | Infection in orthopaedic implants | Review |
| Swanson AB;de Groot SG;Masada K;Makino M;Pires PR;Gannon DM;Sattel AB; | 1991 | Constrained total elbow arthroplasty | Insufficient data for analysis |
| Coulter WA;Coffey A;Saunders ID;Emmerson AM; | 1990 | Bacteremia in children following dental extraction | Insufficient data on bacteremia for background microbiology |
| Kelly PJ;Fitzgerald RH;Cabanela ME;Wood MB;Cooney WP;Arnold PG;Irons GB; | 1990 | Results of treatment of tibial and femoral osteomyelitis in adults | Insufficient data for analysis |
| Mnaymneh W;Emerson RH;Borja F;Head WC;Malinin TI; | 1990 | Massive allografts in salvage revisions of failed total knee arthroplasties | Retrospective study |
| Stern SH;Insall JN; | 1990 | Total knee arthroplasty in obese patients | Insufficient data for analysis |
| Wilson MG; Kelley K; Thornhill TS; | 1990 | Infection as a complication of total knee- replacement arthroplasty. Risk factors and treatment in sixty-seven cases | Retrospective study |
| Wymenga AB;Van Dijke BJ;Van Horn JR;Slooff TJ; | 1990 | Prosthesis-related infection. Etiology, prophylaxis and diagnosis (a review) | Review |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------|
| Emery SE;Chan DP;Woodward HR; | 1989 | Treatment of hematogenous pyogenic vertebral osteomyelitis with anterior debridement and primary bone grafting | Insufficient data for analysis |
| Lian G;Cracchiolo A;Lesavoy M; | 1989 | Treatment of major wound necrosis following total knee arthroplasty | Retrospective study |
| Eskola A;Santavirta S;Konttinen YT;Tallroth K;Hoikka V;Lindholm ST; | 1988 | Cementless total replacement for old tuberculosis of the hip | Insufficient data for analysis |
| Goulet JA;Pellicci PM;Brause BD;Salvati EM; | 1988 | Prolonged suppression of infection in total hip arthroplasty | Retrospective study |
| Gustilo RB;Pasternak HS; | 1988 | Revision total hip arthroplasty with titanium ingrowth prosthesis and bone grafting for failed cemented femoral component loosening | Insufficient data for analysis |
| Kester MA;Cook SD;Harding AF;Rodriguez RP;Pipkin CS; | 1988 | An evaluation of the mechanical failure modalities of a rotating hinge knee prosthesis | Insufficient data for analysis |
| Larsson SE;Larsson S;Lundkvist S; | 1988 | Unicompartmental knee arthroplasty. A prospective consecutive series followed for six to 11 years | Insufficient data for analysis |
| Maderazo EG;Judson S;Pasternak H; | 1988 | Late infections of total joint prostheses. A review and recommendations for prevention | Review |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|-----------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------|
| Madoff S;Hooper DC; | 1988 | Nongenitourinary infections caused by Mycoplasma hominis in adults | Retrospective study |
| Schutzer SF;Harris WH; | 1988 | Deep-wound infection after total hip replacement under contemporary aseptic conditions | Retrospective study |
| Bengtson S;Blomgren G;Knutson K;Wigren A;Lidgren L; | 1987 | Hematogenous infection after knee arthroplasty | Retrospective study |
| Catto BA;Jacobs MR;Shlaes DM; | 1987 | Streptococcus mitis. A cause of serious infection in adults | Insufficient data for analysis |
| Sherrer Y;Bloch D;Strober S;Fries J; | 1987 | Comparative toxicity of total lymphoid irradiation and immunosuppressive drug treated patients with intractable rheumatoid arthritis | Insufficient data for analysis |
| Stuyck J;Verbist L;Mulier JC; | 1987 | Treatment of chronic osteomyelitis with ciprofloxacin | Retrospective study |
| Unger AS;Inglis AE;Ranawat CS;Johanson NA; | 1987 | Total hip arthroplasty in rheumatoid arthritis. A long-term follow-up study | Insufficient data for analysis |
| Grogan TJ;Dorey F;Rollins J;Amstutz HC; | 1986 | Deep sepsis following total knee arthroplasty. Ten-year experience at the University of California at Los Angeles Medical Center | Retrospective study |
| Terayama K; | 1986 | Experience with Charnley low-friction arthroplasty in Japan | Insufficient data for analysis |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Fitzgerald RH;Jones DR; | 1985 | Hip implant infection. Treatment with resection arthroplasty and late total hip arthroplasty | Retrospective study |
| Josefsson K;Heimdahl A;von KL;Nord CE; | 1985 | Effect of phenoxymethylpenicillin and erythromycin prophylaxis on anaerobic bacteraemia after oral surgery | Insufficient data on bacteremia for background microbiology |
| Lachiewicz PF;Inglis AE;Insall JN;Sculco TP;Hilgartner MW;Bussel JB; | 1985 | Total knee arthroplasty in hemophilia | Retrospective study |
| Amstutz HC;Thomas BJ;Jinnah R;Kim W;Grogan T;Yale C; | 1984 | Treatment of primary osteoarthritis of the hip. A comparison of total joint and surface replacement arthroplasty | Retrospective study |
| Cluzel RA;Lopitaux R;Sirot J;Rampon S; | 1984 | Rifampicin in the treatment of osteoarticular infections due to staphylococci | Insufficient data for analysis |
| Inman RD;Gallegos KV;Brause BD;Redecha PB;Christian CL; | 1984 | Clinical and microbial features of prosthetic joint infection | Retrospective study |
| Poss R;Thornhill TS;Ewald FC;Thomas WH;Batte NJ;Sledge CB; | 1984 | Factors influencing the incidence and outcome of infection following total joint arthroplasty | Retrospective study |
| Trivedi DN; | 1984 | Bacteraemia due to operative procedure | Insufficient data on bacteremia for background microbiology |
| Glynn MK;Sheehan JM; | 1983 | An analysis of the causes of deep infection after hip and knee arthroplasties | Retrospective study |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|------------------------------------------------------|------|---------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Ritter MA;Sieber JM; | 1983 | A non-surgical approach to preventing hematogenous infections in total joint replacements | Retrospective study |
| Thomas BJ;Moreland JR;Amstutz HC; | 1983 | Infection after total joint arthroplasty from distal extremity sepsis | Retrospective study |
| Miley GB;Scheller AD;Turner RH; | 1982 | Medical and surgical treatment of the septic hip with one-stage revision arthroplasty | Retrospective study |
| Soreide O;Lillestol J;Alho A;Hvidsten K; | 1982 | Migration of the femoral stem in hip arthroplasties. Analysis of associations with structural, radiological and follow-up variables | Insufficient data for analysis |
| Stinchfield FE;Bigliani LU;Neu HC;Goss TP;Foster CR; | 1980 | Late hematogenous infection of total joint replacement | Retrospective study |
| Hughes PW;Salvati EA;Wilson PD;Blumenfeld EL; | 1979 | Treatment of subacute sepsis of the hip by antibiotics and joint replacement. Criteria for diagnosis with evaluation of twenty-six cases | Retrospective study |
| Sconyers JR; Albers DD; Kelly R; | 1979 | Relationship of bacteremia to toothbrushing in clinically healthy patients | Insufficient data on bacteremia for background microbiology |
| Mallory TH; | 1978 | Excision arthroplasty with delayed wound closure for the infected total hip replacement | Retrospective study |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------|------|-----------------------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Marmor L;Berkus D; | 1978 | Hematogenous infection of total knee implants | Retrospective study |
| Wampole HS;Allen AL;Gross A; | 1978 | The incidence of transient bacteremia during periodontal dressing change | Insufficient data on bacteremia for background microbiology |
| Visuri T;Antila P;Laurent LE; | 1976 | A comparison of dicloxacillin and ampicillin in the antibiotic prophylaxis of total hip replacement | Insufficient data for analysis |
| Millender LH;Nalebuff EA;Hawkins RB;Ennis R; | 1975 | Infection after silicone prosthetic arthroplasty in the hand | Retrospective study |
| Wilson PD;Aglietti P;Salvati EA; | 1974 | Subacute sepsis of the hip treated by antibiotics and cemented prosthesis | Retrospective study |
| Francis LE;DeVries J;Lang D; | 1973 | An oral antiseptic for the control of post- extraction bacteraemia | Insufficient data on bacteremia for background microbiology |
| Roberts GJ;Simmons NB;Longhurst P;Hewitt PB; | 1973 | Evaluation of transient bacteremia following routine periodontal procedures | Insufficient data on bacteremia for background microbiology |
| America Academy of Periodontology | 1972 | Oral irrigation and bacteremia | Insufficient data on bacteremia for background microbiology |
| Degling TE; | 1972 | Orthodontics, bacteremia, and the heart damaged patient | Insufficient data on bacteremia for background microbiology |
| DeVries J;Francis LE;Lang D; | 1972 | Control of post-extraction bacteraemias in the penicillin-hypersensitive patient | Insufficient data on bacteremia for background microbiology |

Table 57 Excluded Studies for Background Microbiology

| Author(s) | Year | Title | Reason for Exclusion |
|-----------------------------------------|------|---------------------------------------------------------------------------------------|-------------------------------------------------------------|
| Wada K;Tomizawa M;Sasaki I; | 1968 | Study on bacteriemia in patients of pyorrhea Alveolaris caused by surgical operations | Insufficient data on bacteremia for background microbiology |
| Bender IB;SELTZER S;TASHMAN S;MELOFF G; | 1963 | Dental procedures in patients with rheumatic heart disease | Insufficient data on bacteremia for background microbiology |

PUBLICATIONS EXCLUDED DURING FULL TEXT REVIEW

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|-------------------------------------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------|
| Ashrafi SS;Nakib N; | 2011 | Need for antibiotic premedication for patients having periodontal dental procedures | Review |
| Castillo DM;Sanchez-Beltran MC;Castellanos JE;Sanz I;Mayorga- Fayad I;Sanz M;Lafaurie GI; | 2011 | Detection of specific periodontal microorganisms from bacteraemia samples after periodontal therapy using molecular-based diagnostics | Comparison of testing methods |
| Esteban J;Cordero-Ampuero J; | 2011 | Treatment of prosthetic osteoarticular infections | Review |
| Garg A;Guez G; | 2011 | Debate rages over antibiotic prophylaxis in patients with total joint replacements | Commentary |
| Mercuri LG;Psutka D; | 2011 | Perioperative, Postoperative, and Prophylactic Use of Antibiotics in Alloplastic Total Temporomandibular Joint Replacement Surgery: A Survey and Preliminary Guidelines | Survey |
| Swan J;Dowsey M;Babazadeh S;Mandaleson A;Choong PF; | 2011 | Significance of sentinel infective events in haematogenous prosthetic knee infections | Retrospective study |
| Zywiel MG;Johnson AJ;Stroh DA;Martin J;Marker DR;Mont MA; | 2011 | Prophylactic oral antibiotics reduce reinfection rates following two-stage revision total knee arthroplasty | Retrospective sudy |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|-------------------------------------------------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Akiyama T;Miyamoto H;Fukuda K;Sano N;Katagiri N;Shobuike T;Kukita A;Yamashita Y;Taniguchi H;Goto M; | 2010 | Development of a novel PCR method to comprehensively analyze salivary bacterial flora and its application to patients with odontogenic infections | Not relevant to bacteremia or implant infection evidence |
| Akutsu Y;Matsubara H;Shuto K;Shiratori T;Uesato M;Miyazawa Y;Hoshino I;Murakami K;Usui A;Kano M;Miyauchi H; | 2010 | Pre-operative dental brushing can reduce the risk of postoperative pneumonia in esophageal cancer patients | Not relevant to bacteremia or implant infection evidence |
| de Oliveira CE;Gasparoto TH;Dionisio TJ;Porto VC;Vieira NA;Santos CF;Lara VS; | 2010 | Candida albicans and denture stomatitis: evaluation of its presence in the lesion, prosthesis, and blood | Not relevant to bacteremia or implant infection evidence |
| Ebersole JL;Stevens J;Steffen MJ;Dawson ID;Novak MJ; | 2010 | Systemic endotoxin levels in chronic indolent periodontal infections | Not relevant to bacteremia or implant infection evidence |
| Bebek B;Bago I;Skaljac G;Plecko V;Miletic I;Anic I; | 2009 | Antimicrobial effect of 0.2% chlorhexidine in infected root canals | Not relevant to bacteremia or implant infection evidence |
| Brook I; | 2009 | The bacteriology of salivary gland infections | Review |
| Herzke CA;Chen LF;Anderson DJ;Choi Y;Sexton DJ;Kaye KS; | 2009 | Empirical antimicrobial therapy for bloodstream infection due to methicillin-resistant Staphylococcus aureus: no better than a coin toss | Not relevant to bacteremia or implant infection evidence |
| Huddleston PM;Clyburn TA;Evans RP;Moucha CS;Prokuski LJ;Joseph J;Sale K; | 2009 | Surgical site infection prevention and control: An emerging paradigm | Review |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|-----------------------------------------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Kuong EE;Ng FY;Yan CH;Fang CX;Chiu PK; | 2009 | Antibiotic prophylaxis after total joint replacements | Review |
| Myburgh HP;Butow KW; | 2009 | Cleft soft palate reconstruction: prospective study on infection and antibiotics | Not relevant to bacteremia or implant infection evidence |
| Nakano K;Ooshima T; | 2009 | Serotype classification of Streptococcus mutans and its detection outside the oral cavity | Not applicable |
| Parahitiyawa NB;Jin LJ;Leung WK;Yam WC;Samaranayake LP; | 2009 | Microbiology of odontogenic bacteremia: beyond endocarditis | Review |
| Anirudhan D;Bakhshi S;Xess I;Broor S;Arya LS; | 2008 | Etiology and outcome of oral mucosal lesions in children on chemotherapy for acute lymphoblastic leukemia | Not relevant to bacteremia or implant infection evidence |
| Bahrani-Mougeot FK;Paster BJ;Coleman S;Ashar J;Knost S;Sautter RL;Lockhart PB; | 2008 | Identification of oral bacteria in blood cultures by conventional versus molecular methods | Not relevant to bacteremia or implant infection evidence |
| Bahrani-Mougeot FK;Thornhill M;Sasser H;Marriott I;Brennan MT;Papagerakis S;Coleman S;Fox PC;Lockhart PB; | 2008 | Systemic host immuno-inflammatory response to dental extractions and periodontitis | Not relevant to bacteremia or implant infection evidence |
| Cogulu D;Uzel A;Oncag O;Eronat C; | 2008 | PCR-based identification of selected pathogens associated with endodontic infections in deciduous and permanent teeth | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------------------------------------------------------------------|------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Jones DJ;Munro CL; | 2008 | Oral care and the risk of bloodstream infections in mechanically ventilated adults: A review | Review |
| Lalani T;Chu VH;Grussemeyer CA;Reed SD;Bolognesi MP;Friedman JY;Griffiths RI;Crosslin DR;Kanafani ZA;Kaye KS;Ralph CG;Fowler VG; | 2008 | Clinical outcomes and costs among patients with Staphylococcus aureus bacteremia and orthopedic device infections | Cost analysis |
| Lee MK;Ide M;Coward PY;Wilson RF; | 2008 | Effect of ultrasonic debridement using a chlorhexidine irrigant on circulating levels of lipopolysaccharides and interleukin-6 | Not relevant to bacteremia or implant infection evidence |
| Montefusco V;Gay F;Spina F;Miceli R;Maniezzo M;Teresa AM;Farina L;Piva S;Palumbo A;Boccadoro M;Corradini P; | 2008 | Antibiotic prophylaxis before dental procedures may reduce the incidence of osteonecrosis of the jaw in patients with multiple myeloma treated with bisphosphonates | Retrospective study |
| Sassone LM;Fidel RA;Faveri M;Guerra R;Figueiredo L;Fidel SR;Feres M; | 2008 | A microbiological profile of symptomatic teeth with primary endodontic infections | Not relevant to bacteremia or implant infection evidence |
| Tosello A;Chevaux JM;Montal S;Foti B; | 2008 | Assessment of oral status and oro- pharyngeal candidosis in elderly in short- term hospital care | Not relevant to bacteremia or implant infection evidence |
| Uckay I;Pittet D;Bernard L;Lew D;Perrier A;Peter R; | 2008 | Antibiotic prophylaxis before invasive dental procedures in patients with arthroplasties of the hip and knee | Review |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|-------------------------------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Wright TI;Baddour LM;Berbari EF;Roenigk RK;Phillips PK;Jacobs MA;Otley CC; | 2008 | Antibiotic prophylaxis in dermatologic surgery: advisory statement 2008 | Advisory Statement |
| Yilmaz S;Oren H;Demircioglu F;Irken G; | 2008 | Assessment of febrile neutropenia episodes in children with acute leukemia treated with BFM protocols | Retrospective study |
| Kuriyama T;Williams DW;Yanagisawa M;Iwahara K;Shimizu C;Nakagawa K;Yamamoto E;Karasawa T; | 2007 | Antimicrobial susceptibility of 800 anaerobic isolates from patients with dentoalveolar infection to 13 oral antibiotics | Not relevant to bacteremia or implant infection evidence |
| Faveri M;Feres M;Shibli JA;Hayacibara RF;Hayacibara MM;de Figueiredo LC; | 2006 | Microbiota of the dorsum of the tongue after plaque accumulation: An experimental study in humans | Not relevant to bacteremia or implant infection evidence |
| Flynn TR;Shanti RM;Hayes C; | 2006 | Severe odontogenic infections, part 2: prospective outcomes study | Not relevant to bacteremia or implant infection evidence |
| Flynn TR;Shanti RM;Levi MH;Adamo AK;Kraut RA;Trieger N; | 2006 | Severe odontogenic infections, part 1: prospective report | Not relevant to bacteremia or implant infection evidence |
| Khemaleelakul S;Baumgartner JC;Pruksakom S; | 2006 | Autoaggregation and coaggregation of bacteria associated with acute endodontic infections | Not relevant to bacteremia or implant infection evidence |
| Marculescu CE;Berbari EF;Hanssen AD;Steckelberg JM;Harmsen SW;Mandrekar JN;Osmon DR; | 2006 | Outcome of prosthetic joint infections treated with debridement and retention of components | Retrospective study |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|-----------------------------------------------------------------------------------------------------------------------------------------------------------|------|---------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Rocas IN;Baumgartner JC;Xia T;Siqueira JF; | 2006 | Prevalence of selected bacterial named species and uncultivated phylotypes in endodontic abscesses from two geographic locations | Not relevant to bacteremia or implant infection evidence |
| Saito D;Leonardo RT;Rodrigues JL;Tsai SM;Hofling JF;Goncalves RB; | 2006 | Identification of bacteria in endodontic infections by sequence analysis of 16S rDNA clone libraries | Not relevant to bacteremia or implant infection evidence |
| Sakamoto M;Rocas IN;Siqueira JF;Benno Y; | 2006 | Molecular analysis of bacteria in asymptomatic and symptomatic endodontic infections | Not relevant to bacteremia or implant infection evidence |
| Sakr MR;El-Aiady AA;Ragab SH;Gomaa HE;El Din HG; | 2006 | Fungal and bacterial infection in malnourished children and its relation to severity of the disease | Not relevant to bacteremia or implant infection evidence |
| Sixou JL;Aubry-Leuliette A;De Medeiros-Battista O;Lejeune S;Jolivet- Gougeon A;Solhi-Pinsard H;Gandemer V;Barbosa-Rogier M;Bonnaure-Mallet M; | 2006 | Capnocytophaga in the dental plaque of immunocompromised children with cancer | Not relevant to bacteremia or implant infection evidence |
| Not available | 2005 | Antibacterial prophylaxis for dental, GI, and GU procedures | Review |
| Chakraborty P;Chattopadhyay UK; | 2005 | A study on the polymicrobial etiology of root canal infections in anterior non-vital teeth in a government hospital in Kolkata, India | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Chu FC;Tsang CS;Chow TW;Samaranayake LP; | 2005 | Identification of cultivable microorganisms from primary endodontic infections with exposed and unexposed pulp space | Not relevant to bacteremia or implant infection evidence |
| Ferrari PH;Cai S;Bombana AC; | 2005 | Effect of endodontic procedures on enterococci, enteric bacteria and yeasts in primary endodontic infections | Not relevant to bacteremia or implant infection evidence |
| Huang ST;Lee HC;Lee NY;Liu KH;Ko WC; | 2005 | Clinical characteristics of invasive Haemophilus aphrophilus infections | Retrospective study |
| Iwai T;Inoue Y;Umeda M;Huang Y;Kurihara N;Koike M;Ishikawa I; | 2005 | Oral bacteria in the occluded arteries of patients with Buerger disease | Not relevant to bacteremia or implant infection evidence |
| Nowak E;Niepsuj K;Nolewajka-Lasak I;Rheinbaben FV; | 2005 | The effectiveness of preoperative rinsing with skinsept oral on reducing the bacterial flora and eradicating Helicobacter pylori in the oral cavity | Not relevant to bacteremia or implant infection evidence |
| Shariff G;Brennan MT;Louise KM;Fox PC;Weinrib D;Burgess P;Lockhart PB; | 2004 | Relationship between oral bacteria and hemodialysis access infection | Not relevant to bacteremia or implant infection evidence |
| Apisarnthanarak A;Mayfield JL;Garison T;McLendon PM;DiPersio JF;Fraser VJ;Polish LB; | 2003 | Risk factors for Stenotrophomonas maltophilia bacteremia in oncology patients: a case-control study | Not relevant to bacteremia or implant infection evidence |
| Candoni A;Fili C;Trevisan R;Silvestri F;Fanin R; | 2003 | Fusobacterium nucleatum: a rare cause of bacteremia in neutropenic patients with leukemia and lymphoma | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Krcmery V;Gogova M;Ondrusova A;Buckova E;Doczeova A;Mrazova M;Hricak V;Fischer V;Marks P;Kovacik J;Schramekova E;Vitekova D;Sedlak T;Duris I;Samudovsky J;Semanova M;Kovac M;Duris T;Herman O;Cernoskova M;Sefara J;Kojsova M;Baranikova D;Ayazi M;Neuschlova D | 2003 | Etiology and Risk Factors of 339 Cases of Infective Endocarditis: Report from a 10-year National Prospective Survey in the Slovak Republic | Not applicable |
| Listgarten MA;Loomer PM; | 2003 | Microbial identification in the management of periodontal diseases. A systematic review | Review |
| Seymour RA; Whitworth JM; Martin M; | 2003 | Antibiotic prophylaxis for patients with joint prostheses: still a dilemma for dental practitioners (Brief record) | Retrospective study |
| Brook I; | 2002 | Aerobic and anaerobic microbiology of suppurative sialadenitis | Not relevant to bacteremia or implant infection evidence |
| Fouad AF;Barry J;Caimano M;Clawson M;Zhu Q;Carver R;Hazlett K;Radolf JD; | 2002 | PCR-based identification of bacteria associated with endodontic infections | Not relevant to bacteremia or implant infection evidence |
| Geerts SO;Nys M;De MP;Charpentier J;Albert A;Legrand V;Rompen EH; | 2002 | Systemic release of endotoxins induced by gentle mastication: association with periodontitis severity | Not relevant to bacteremia or implant infection evidence |
| Kucukkaya M;Kabukcuoglu Y;Tezer M;Kuzgun U; | 2002 | Management of childhood chronic tibial osteomyelitis with the Ilizarov method | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Munson MA;Pitt-Ford T;Chong B;Weightman A;Wade WG; | 2002 | Molecular and cultural analysis of the microflora associated with endodontic infections | Not relevant to bacteremia or implant infection evidence |
| Peters LB; Wesselink PR; van Winkelhoff AJ; | 2002 | Combinations of bacterial species in endodontic infections | Not relevant to bacteremia or implant infection evidence |
| Reebye UN;Ollerhead TR;Hughes CV;Cottrell DA; | 2002 | The microbial composition of mandibular third molar pericoronal infections | Not relevant to bacteremia or implant infection evidence |
| Roberts G;Holzel H; | 2002 | Intravenous antibiotic regimens and prophylaxis of odontogenic bacteraemia | Retrospective study |
| Sunde PT;Olsen I;Debelian GJ;Tronstad L; | 2002 | Microbiota of periapical lesions refractory to endodontic therapy | Not applicable |
| Tada A;Watanabe T;Yokoe H;Hanada N;Tanzawa H; | 2002 | Oral bacteria influenced by the functional status of the elderly people and the type and quality of facilities for the bedridden | Not relevant to bacteremia or implant infection evidence |
| van SD;Kaandorp C;Krijnen P; | 2002 | Cost-effectiveness of antibiotic prophylaxis for bacterial arthritis | Cost analysis |
| Fernandes-Naglik L;Downes J;Shirlaw P;Wilson R;Challacombe SJ;Kemp GK;Wade WG; | 2001 | The clinical and microbiological effects of a novel acidified sodium chlorite mouthrinse on oral bacterial mucosal infections | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|------------------------------------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Fujiwara T;Nakano K;Kawaguchi M;Ooshima T;Sobue S;Kawabata S;Nakagawa I;Hamada S; | 2001 | Biochemical and genetic characterization of serologically untypable Streptococcus mutans strains isolated from patients with bacteremia | Not relevant to bacteremia or implant infection evidence |
| Glass RT;Bullard JW;Hadley CS;Mix EW;Conrad RS; | 2001 | Partial spectrum of microorganisms found in dentures and possible disease implications | Not relevant to bacteremia or implant infection evidence |
| Krijnen P;Kaandorp CJ;Steyerberg EW;van SD;Moens HJ;Habbema JD; | 2001 | Antibiotic prophylaxis for haematogenous bacterial arthritis in patients with joint disease: a cost effectiveness analysis | Cost analysis |
| Lana MA;Ribeiro-Sobrinho AP;Stehling R;Garcia GD;Silva BK;Hamdan JS;Nicoli JR;Carvalho MA;Farias LM; | 2001 | Microorganisms isolated from root canals presenting necrotic pulp and their drug susceptibility in vitro | Not relevant to bacteremia or implant infection evidence |
| Peciuliene V;Reynaud AH;Balciuniene I;Haapasalo M; | 2001 | Isolation of yeasts and enteric bacteria in root-filled teeth with chronic apical periodontitis | Not relevant to bacteremia or implant infection evidence |
| Pitten FA;Kramer A; | 2001 | Efficacy of cetylpyridinium chloride used as oropharyngeal antiseptic | Not relevant to bacteremia or implant infection evidence |
| Wrobel CJ;Chappell ET;Taylor W; | 2001 | Clinical presentation, radiological findings, and treatment results of coccidioidomycosis involving the spine: report on 23 cases | Retrospective study |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Kuriyama T;Karasawa T;Nakagawa K;Saiki Y;Yamamoto E;Nakamura S; | 2000 | Bacteriologic features and antimicrobial susceptibility in isolates from orofacial odontogenic infections | Not relevant to bacteremia or implant infection evidence |
| Labarca JA;Leber AL;Kern VL;Territo MC;Brankovic LE;Bruckner DA;Pegues DA; | 2000 | Outbreak of Stenotrophomonas maltophilia bacteremia in allogenic bone marrow transplant patients: role of severe neutropenia and mucositis | n<10 |
| Lucht U; | 2000 | The Danish Hip Arthroplasty Register | Review |
| Monsenego P; | 2000 | Presence of microorganisms on the fitting denture complete surface: study 'in vivo' | Not relevant to bacteremia or implant infection evidence |
| Mullally BH;Dace B;Shelburne CE;Wolff LF;Coulter WA; | 2000 | Prevalence of periodontal pathogens in localized and generalized forms of early-onset periodontitis | Not relevant to bacteremia or implant infection evidence |
| Osaki T;Yoneda K;Yamamoto T;Ueta E;Kimura T; | 2000 | Candidiasis may induce glossodynia without objective manifestation | Not relevant to bacteremia or implant infection evidence |
| Peltroche-Llacsahuanga H;Reichhart E;Schmitt W;Lutticken R;Haase G; | 2000 | Investigation of infectious organisms causing pericoronitis of the mandibular third molar | Not relevant to bacteremia or implant infection evidence |
| Rupf S;Kannengiesser S;Merte K;Pfister W;Sigusch B;Eschrich K; | 2000 | Comparison of profiles of key periodontal pathogens in periodontium and endodontium | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|------------------------------------------------------------------------------------------------------|------|-----------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Siqueira JF;Rocas IN;Souto R;de UM;Colombo AP; | 2000 | Checkerboard DNA-DNA hybridization analysis of endodontic infections | Not relevant to bacteremia or implant infection evidence |
| Sunde PT;Olsen I;Lind PO;Tronstad L; | 2000 | Extraradicular infection: a methodological study | Not relevant to bacteremia or implant infection evidence |
| Sunde PT;Tronstad L;Eribe ER;Lind PO;Olsen I; | 2000 | Assessment of periradicular microbiota by DNA-DNA hybridization | Not relevant to bacteremia or implant infection evidence |
| Bentley KC;Head TW;Aiello GA; | 1999 | Antibiotic prophylaxis in orthognathic surgery: a 1-day versus 5-day regimen | Not relevant to bacteremia or implant infection evidence |
| Hall G;Heimdahl A;Nord CE; | 1999 | Bacteremia after oral surgery and antibiotic prophylaxis for endocarditis | Review |
| LaPorte DM;Waldman BJ;Mont MA;Hungerford DS; | 1999 | Infections associated with dental procedures in total hip arthroplasty | Retrospective study |
| Lockhart PB;Durack DT; | 1999 | Oral microflora as a cause of endocarditis and other distant site infections | Review |
| Nicolatou-Galitis O;Bakiri M;Belegrati M;Nikolatos G;Spyropoulos C;Fisfis M;Kalmantis T;Velegraki A; | 1999 | Oropharyngeal candidiasis in patients with hematological immunosuppression. A pilot study | Not relevant to bacteremia or implant infection evidence |
| Reit C;Molander A;Dahlen G; | 1999 | The diagnostic accuracy of microbiologic root canal sampling and the influence of antimicrobial dressings | Not relevant to bacteremia or implant infection evidence |
| Amir J;Yagupsky P; | 1998 | Invasive Kingella kingae infection associated with stomatitis in children | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Pinero J; | 1998 | Nd:YAG-assisted periodontal curettage to prevent bacteria before cardiovascular surgery | Not relevant to bacteremia or implant infection evidence |
| Segreti J;Nelson JA;Trenholme GM; | 1998 | Prolonged suppressive antibiotic therapy for infected orthopedic prostheses | Retrospective study |
| Chaudhry R;Kalra N;Talwar V;Thakur R; | 1997 | Anaerobic flora in endodontic infections | Not relevant to bacteremia or implant infection evidence |
| Drucker DB;Gomes BP;Lilley JD; | 1997 | Role of anaerobic species in endodontic infection | Not relevant to bacteremia or implant infection evidence |
| Jacobson J;Patel B;Asher G;Woolliscroft JO;Schaberg D; | 1997 | Oral staphylococcus in older subjects with rheumatoid arthritis | Not relevant to bacteremia or implant infection evidence |
| Kulak Y;Arikan A;Kazazoglu E; | 1997 | Existence of Candida albicans and microorganisms in denture stomatitis patients | Not relevant to bacteremia or implant infection evidence |
| Moritz A;Gutknecht N;Schoop U;Goharkhay K;Doertbudak O;Sperr W; | 1997 | Irradiation of infected root canals with a diode laser in vivo: results of microbiological examinations | Not relevant to bacteremia or implant infection evidence |
| Waldman BJ;Mont MA;Hungerford DS; | 1997 | Total knee arthroplasty infections associated with dental procedures | Retrospective study |
| Bollen CM;Vandekerckhove BN;Papaioannou W;Van EJ;Quirynen M; | 1996 | Full- versus partial-mouth disinfection in the treatment of periodontal infections. A pilot study: long-term microbiological observations | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|-------------------------------------------------------------------------------------|------|---------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Deacon JM;Pagliaro AJ;Zelicof SB;Horowitz HW; | 1996 | Prophylactic use of antibiotics for procedures after total joint replacement | Review |
| Debelian GJ;Olsen I;Tronstad L; | 1996 | Electrophoresis of whole-cell soluble proteins of microorganisms isolated from bacteremias in endodontic therapy | Not relevant to bacteremia or implant infection evidence |
| Rajasuo A;Jousimies-Somer H;Savolainen S;Leppanen J;Murtomaa H;Meurman JH; | 1996 | Bacteriologic findings in tonsillitis and pericoronitis | Not relevant to bacteremia or implant infection evidence |
| Yoneyama T;Hashimoto K;Fukuda H;Ishida M;Arai H;Sekizawa K;Yamaya M;Sasaki H; | 1996 | Oral hygiene reduces respiratory infections in elderly bed-bound nursing home patients | Not relevant to bacteremia or implant infection evidence |
| Quirynen M;Bollen CM;Vandekerckhove BN;Dekeyser C;Papaioannou W;Eyssen H; | 1995 | Full- vs. partial-mouth disinfection in the treatment of periodontal infections: short-term clinical and microbiological observations | Not relevant to bacteremia or implant infection evidence |
| Richard P;Amador D;Moreau P;Milpied N;Felice MP;Daeschler T;Harousseau JL;Richet H; | 1995 | Viridans streptococcal bacteraemia in patients with neutropenia | Not relevant to bacteremia or implant infection evidence |
| Bartzokas CA;Johnson R;Jane M;Martin MV;Pearce PK;Saw Y; | 1994 | Relation between mouth and haematogenous infection in total joint replacements | n<10 |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Sjostrom K;Ou J;Whitney C;Johnson B;Darveau R;Engel D;Page RC; | 1994 | Effect of treatment on titer, function, and antigen recognition of serum antibodies to Actinobacillus actinomycetemcomitans in patients with rapidly progressive periodontitis | Not relevant to bacteremia or implant infection evidence |
| Brook I;Frazier EH; | 1993 | Anaerobic osteomyelitis and arthritis in a military hospital: a 10-year experience | Not relevant to bacteremia or implant infection evidence |
| Donnelly JP;Muus P;Horrevorts AM;Sauerwein RW;de Pauw BE; | 1993 | Failure of clindamycin to influence the course of severe oromucositis associated with streptococcal bacteraemia in allogeneic bone marrow transplant recipients | Not relevant to bacteremia or implant infection evidence |
| Helovuo H;Hakkarainen K;Paunio K; | 1993 | Changes in the prevalence of subgingival enteric rods, staphylococci and yeasts after treatment with penicillin and erythromycin | Not relevant to bacteremia or implant infection evidence |
| Holan G;Kadari A;Engelhard D;Chosack A; | 1993 | Temperature elevation in children following dental treatment under general anesthesia with or without prophylactic antibiotics | Not relevant to bacteremia or implant infection evidence |
| Lo Bue AM;Sammartino R;Chisari G;Gismondo MR;Nicoletti G; | 1993 | Efficacy of azithromycin compared with spiramycin in the treatment of odontogenic infections | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| O'Sullivan EA;Duggal MS;Bailey CC;Curzon ME;Hart P; | 1993 | Changes in the oral microflora during cytotoxic chemotherapy in children being treated for acute leukemia | Not relevant to bacteremia or implant infection evidence |
| Ueta E;Osaki T;Yoneda K;Yamamoto T; | 1993 | Prevalence of diabetes mellitus in odontogenic infections and oral candidiasis: an analysis of neutrophil suppression | Not relevant to bacteremia or implant infection evidence |
| Bergmann OJ;Ellegaard B;Dahl M;Ellegaard J; | 1992 | Gingival status during chemical plaque control with or without prior mechanical plaque removal in patients with acute myeloid leukaemia | Not relevant to bacteremia or implant infection evidence |
| Epstein JB;Vickars L;Spinelli J;Reece D; | 1992 | Efficacy of chlorhexidine and nystatin rinses in prevention of oral complications in leukemia and bone marrow transplantation | Not relevant to bacteremia or implant infection evidence |
| Hashioka K;Yamasaki M;Nakane A;Horiba N;Nakamura H; | 1992 | The relationship between clinical symptoms and anaerobic bacteria from infected root canals | Not relevant to bacteremia or implant infection evidence |
| Norden C;Nelson JD;Mader JT;Calandra GB; | 1992 | Evaluation of new anti-infective drugs for the treatment of infections of prosthetic hip joints. Infectious Diseases Society of America and the Food and Drug Administration | Review |
| Ufomata D;Akerele JO; | 1992 | Bacteriological investigation of infected root canals in Benin City, Nigeria | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------------------------------|------|------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Baumgartner JC;Falkler WA; | 1991 | Bacteria in the apical 5 mm of infected root canals | Not relevant to bacteremia or implant infection evidence |
| Bergmann OJ; | 1991 | Alterations in oral microflora and pathogenesis of acute oral infections during remission-induction therapy in patients with acute myeloid leukaemia | Not relevant to bacteremia or implant infection evidence |
| Hirai K;Tagami A;Okuda K; | 1991 | Isolation and classification of anaerobic bacteria from pulp cavities of nonvital teeth in man | Not relevant to bacteremia or implant infection evidence |
| Jacobson JJ;Schweitzer SO;Kowalski CJ; | 1991 | Chemoprophylaxis of prosthetic joint patients during dental treatment: a decision-utility analysis | Decision utiltiy analysis |
| Thyne GM;Ferguson JW; | 1991 | Antibiotic prophylaxis during dental treatment in patients with prosthetic joints | Review |
| Bell SM;Gatus BJ;Shepherd BD; | 1990 | Antibiotic prophylaxis for the prevention of late infections of prosthetic joints | Retrospective study |
| Jacobson JJ;Schweitzer S;DePorter DJ;Lee JJ; | 1990 | Antibiotic prophylaxis for dental patients with joint prostheses? A decision analysis | Decision utiltiy analysis |
| Peterson DE;Minah GE;Reynolds MA;Weikel DS;Overholser CD;DePaola LG;Wade JC;Suzuki JB; | 1990 | Effect of granulocytopenia on oral microbial relationships in patients with acute leukemia | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|---------------------------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Wilkins J;Patzakis MJ; | 1990 | Peripheral teflon catheters. Potential source for bacterial contamination of orthopedic implants? | Irrelevant study population |
| Bergmann OJ; | 1989 | Oral infections and fever in immunocompromised patients with haematologic malignancies | Not relevant to bacteremia or implant infection evidence |
| Brown AT;Sims RE;Raybould TP;Lillich TT;Henslee PJ;Ferretti GA; | 1989 | Oral gram-negative bacilli in bone marrow transplant patients given chlorhexidine rinses | Not relevant to bacteremia or implant infection evidence |
| Daoud A;Saighi-Bouaouina A; | 1989 | Treatment of sequestra, pseudarthroses, and defects in the long bones of children who have chronic hematogenous osteomyelitis | Retrospective study |
| Etemadzadeh H;Meurmann JH;Murtomaa H;Torkko H;Lappi L;Roos M; | 1989 | Effect on plaque growth and salivary micro-organisms of amine fluoride- stannous fluoride and chlorhexidine- containing mouthrinses | Not relevant to bacteremia or implant infection evidence |
| Gerlach KL;Schaal KP;Walz C;Pape HD; | 1989 | Treatment of severe odontogenic infections with amoxicillin/clavulanic acid | Not relevant to bacteremia or implant infection evidence |
| Heimdahl A;Mattsson T;Dahllof G;Lonnquist B;Ringden O; | 1989 | The oral cavity as a port of entry for early infections in patients treated with bone marrow transplantation | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------|------|-------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Lindqvist C;Soderholm AL;Slatis P; | 1989 | Dental X-ray status of patients admitted for total hip replacement | Not relevant to bacteremia or implant infection evidence |
| Lo Bue AM;Chisari G;Fiorenza G;Ferlito S;Gismondo MR; | 1989 | The activity of ofloxacin compared to spiramycin in oral surgery | Not relevant to bacteremia or implant infection evidence |
| Steele MT;Sainsbury CR;Robinson WA;Salomone JA;Elenbaas RM; | 1989 | Prophylactic penicillin for intraoral wounds | Not relevant to bacteremia or implant infection evidence |
| Tsevat J;Durand-Zaleski I;Pauker SG; | 1989 | Cost-effectiveness of antibiotic prophylaxis for dental procedures in patients with artificial joints | Cost analysis |
| Weikel DS;Peterson DE;Rubinstein LE;Metzger-Samuels C;Overholser CD; | 1989 | Incidence of fever following invasive oral interventions in the myelosuppressed cancer patient | Not relevant to bacteremia or implant infection evidence |
| Appelbaum PC;Spangler SK;Strauss M; | 1988 | Reduction of oral flora with ciprofloxacin in healthy volunteers | Not relevant to bacteremia or implant infection evidence |
| Bergmann OJ; | 1988 | Oral infections and septicemia in immunocompromised patients with hematologic malignancies | Not relevant to bacteremia or implant infection evidence |
| Cioffi GA;Terezhalmy GT;Taybos GM; | 1988 | Total joint replacement: a consideration for antimicrobial prophylaxis | Review |
| De LM; | 1988 | Clinical and microbiological effects of in vivo miocamycin therapy on oral infections and in surgical prophylaxis | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------------------------------------------------------|------|---------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Ferretti GA;Ash RC;Brown AT;Parr MD;Romond EH;Lillich TT; | 1988 | Control of oral mucositis and candidiasis in marrow transplantation: a prospective, double-blind trial of chlorhexidine digluconate oral rinse | Not relevant to bacteremia or implant infection evidence |
| Jacobson JJ;Schweitzer S;DePorter DJ;Lee JJ; | 1988 | Chemoprophylaxis of dental patients with prosthetic joints: a simulation model | Simulation model |
| Ranta H;Haapasalo M;Ranta K;Kontiainen S;Kerosuo E;Valtonen V;Suuronen R;Hovi T; | 1988 | Bacteriology of odontogenic apical periodontitis and effect of penicillin treatment | Not relevant to bacteremia or implant infection evidence |
| Rosen S;Ogg-Bell K;Heller A;Weisenstein P;Beck FM; | 1988 | Use of an organic iodine compound to decrease oral microflora in the implant patient | Not relevant to bacteremia or implant infection evidence |
| Dumbach J;Spitzer W; | 1987 | Short-term antibiotic prophylaxis in elective oral and maxillofacial surgery with mezlocillin and oxacillin | Not relevant to bacteremia or implant infection evidence |
| Ebersole JL;Taubman MA;Smith DJ;Frey DE;Haffajee AD;Socransky SS; | 1987 | Human serum antibody responses to oral microorganisms. IV. Correlation with homologous infection | Not relevant to bacteremia or implant infection evidence |
| Foster RJ;Collins FJ;Bach AW; | 1987 | Concurrent oral surgery and orthopaedic treatment in the multiply injured patient: is there an increased incidence of orthopaedic sepsis? | Irrelevant study population |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Jacobson JJ;Matthews LS; | 1987 | Bacteria isolated from late prosthetic joint infections: dental treatment and chemoprophylaxis | Retrospective study |
| Kerver AJH;Rommes JH;Mevissen- Verhage EAE; | 1987 Intensive care patients - A prospective study Characterization by pyocine typing and serotyping of oral and sputum strains of Pseudomonas aeruginosa isolated from cystic fibrosis patients | | Not relevant to bacteremia or implant infection evidence |
| Komiyama K;Habbick BF;Martin T;Tumber SK; | | | Not relevant to bacteremia or implant infection evidence |
| Maniloff G;Greenwald R;Laskin R;Singer C; | | | n<10 |
| Quayle AA;Russell C;Hearn B; | sensitivities to cefotetan and seven other | odontogenic soft tissue infections: Their sensitivities to cefotetan and seven other antibiotics, and implications for therapy | Not relevant to bacteremia or implant infection evidence |
| Fong IW;Ledbetter WH;Vandenbroucke AC;Simbul M;Rahm V; | 1986 | Ciprofloxacin concentrations in bone and muscle after oral dosing | Not applicable |
| Jacobson JJ;Millard HD;Plezia R;Blankenship JR; | 1986 | Dental treatment and late prosthetic joint infections | Retrospective study |
| Santosh S;Saini OP;Manjit C;Uma S; | 1986 | Bacteriological status of closed root canals of non-vital teeth | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|----------------------------------------|------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Wittmann DH;Kotthaus E; | 1986 | Further methodological improvement in antibiotic bone concentration measurements: penetration of ofloxacin into bone and cartilage | Not relevant to bacteremia or implant infection evidence |
| Heimdahl A;von KL;Satoh T;Nord CE; | 1985 | Clinical appearance of orofacial infections of odontogenic origin in relation to microbiological findings | Not relevant to bacteremia or implant infection evidence |
| Kovatch AL;Wald ER;Albo VC; | 1985 | Oral trimethoprim sulfamethoxazole for prevention of bacterial infection during the induction phase of cancer chemotherapy in children | Not relevant to bacteremia or implant infection evidence |
| Mangini P;Cicchetti M;Bottaro L; | 1985 | A multicenter, randomized parallel double-blind study comparing three antibiotics, cephemic-cofosfolactamine, fosfomycin and cephalexin, in the treatment of systemic infections | Not relevant to bacteremia or implant infection evidence |
| McGowan DA;Hendrey ML; | 1985 | Is antibiotic prophylaxis required for dental patients with joint replacements? | Retrospective study |
| Woodman AJ;Vidic J;Newman HN;Marsh PD; | 1985 | Effect of repeated high dose prophylaxis with amoxycillin on the resident oral flora of adult volunteers | Not relevant to bacteremia or implant infection evidence |
| Cannon PD;Black HJ;Kitson K;Ward CS; | 1984 | Serum amoxycillin levels following oral loading dose prior to outpatient general anaesthesia for dental extractions | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------|------|----------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Newman MG; | 1984 | Anaerobic oral and dental infection | Review |
| Terezhalmy GT;Hall EH; | 1984 | The asplenic patient: a consideration for antimicrobial prophylaxis | Review |
| Crawford I;Russell C; | 1983 | Streptococci isolated from the bloodstream and gingival crevice of man | Not relevant to bacteremia or implant infection evidence |
| Hunt DE;Meyer RA; | 1983 | Continued evolution of the microbiology of oral infections | Not relevant to bacteremia or implant infection evidence |
| Southall PJ;Mahy NJ;Davies RM;Speller DC; | 1983 | Resistance in oral streptococci after repeated two-dose amoxycillin prophylaxis | Not relevant to bacteremia or implant infection evidence |
| von KL;Nord CE; | 1983 | Ornidazole compared to phenoxymethylpenicillin in the treatment of orofacial infections | Not relevant to bacteremia or implant infection evidence |
| Erasmus M;Lichter D;Rock R;Rumbak A;Rumbak J; | 1982 | An investigation to determine the frequency of resistance of plaque bacteria to certain antimicrobial drugs | Not relevant to bacteremia or implant infection evidence |
| Greenberg MS;Cohen SG;McKitrick JC;Cassileth PA; | 1982 | The oral flor as a source of septicemia in patients with acute leukemia | Not relevant to bacteremia or implant infection evidence |
| Stobberinoh EE;Eggink CO; | 1982 | The value of the bacteriological culture in endodontics. II. The bacteriological flora of endodontic specimens | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion | |
|--------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|--|
| Bystrom A;Sundqvist G; | 1981 | Bacteriologic evaluation of the efficacy of mechanical root canal instrumentation in endodontic therapy | Not relevant to bacteremia or implant infection evidence | |
| Newman KA;Schimpff SC;Young VM;Wiernik PH; | 1981 | Lessons learned from surveillance cultures in patients with acute nonlymphocytic leukemia. Usefulness for epidemiologic, preventive and therapeutic research | Not relevant to bacteremia or implant infection evidence | |
| von KL;Nord CE;Nordenram A; | 1981 | Anaerobic bacteria in dentoalveolar infections | Not relevant to bacteremia or implant infection evidence | |
| Jacobsen PL;Murray W; | Prophylactic coverage of dental patients with artificial joints: a retrospective analysis of thirty-three infections in hip prostheses | | Retrospective study | |
| Kannangara DW;Thadepalli H;McQuirter JL; | 1980 | Bacteriology and treatment of dental infections | Not relevant to bacteremia or implant infection evidence | |
| Krekmanov L;Hallander HO; | 1980 | Relationship between bacterial contamination and alveolitis after third molar surgery | Not relevant to bacteremia or implant infection evidence | |
| Carlsson AK;Lidgren L;Lindberg L; | 1977 | Prophylactic antibiotics against early and late deep infections after total hip replacements | Retrospective study | |
| Sabiston CB;Grigsby WR; | 1977 | The microbiology of dentalpyogenic infections | Not relevant to bacteremia or implant infection evidence | |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|---------------------------------------|------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Altonen M;SAXEN L;Kosunen T;Ainamo J; | 1976 | Effect of two antimicrobial rinses and oral prophylaxis on preoperative degerming of saliva | Not relevant to bacteremia or implant infection evidence |
| Billick SB;Borchardt KA;Poenisch P; | 1976 | Asymptomatic orophyarngeal flora in patients admitted to hospital | Not relevant to bacteremia or implant infection evidence |
| Williams BL;Pantalone RM;Sherris JC; | 1976 | Subgingival microflora and periodontitis | Not relevant to bacteremia or implant infection evidence |
| Gabrielson ML;Stroh E; | 1975 | Antibiotic efficacy in odontogenic infections | Not relevant to bacteremia or implant infection evidence |
| Mejare B; | 1975 | Streptococcus faecalis and Streptococcus faecium in infected dental root canals at filling and their susceptibility to azidocillin and some comparable antibiotics | Not relevant to bacteremia or implant infection evidence |
| Murray PR; Washington JA; | 1975 | Microscopic and baceriologic analysis of expectorated sputum | Not relevant to bacteremia or implant infection evidence |
| Sabiston CB;Gold WA; | 1974 | Anaerobic bacteria in oral infections | Not relevant to bacteremia or implant infection evidence |
| Sims W; | 1974 | The clinical bacteriology of purulent oral infections | Not relevant to bacteremia or implant infection evidence |
| Turner JE;Mincer HH; | 1974 | Prevalence and antibiotic susceptibility of microorganisms isolated from oral infectious disease | Not relevant to bacteremia or implant infection evidence |

Table 58 Excluded Studies Identified During Full Text Review

| Author(s) | Year | Title | Reason for Exclusion |
|--------------------------------------------------|--------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------|
| Stone HH;Geheber CE;Kolb LD;Kitchens WR; | 1973 Alimentary tract colonization by Candida albicans | | Not relevant to bacteremia or implant infection evidence |
| Khairat O; | 1967 | Bacteroides corrodens isolated from bacteriaemias | Duplicate publication |
| Diener J;Schwartz SM;Shelanski M;Steinberg G; | 1964 | Bacteremia and oral sepsis with particular reference to the possible reduction of systemic disease originating from the oral cavity | Not relevant to bacteremia or implant infection evidence |
| Garrod LP;WATERWORTH PM; | 1962 | The risks of dental extraction during penicillin treatment | Not relevant to bacteremia or implant infection evidence |

APPENDIX IV MEDICAL LIBRARIAN SEARCH STRATEGY PUBMED/MEDLINE STRATEGY:

#1

Dentistry[mh] OR Mouth[mh] OR "Dental Care"[mh] OR "Mouth Diseases/therapy"[mh] OR "Mouth Neoplasms/therapy"[mh] OR "Dental implants"[mh] OR "Dental Prosthesis"[mh] OR "Nonodontogenic Cysts"[mh] OR "Odontogenic Cysts"[mh] OR "Dental Health Surveys"[mh] OR "oral bacteria" OR "dental caries" OR ((oral[titl] OR dental[titl]) NOT medline[sb]) OR "Teeth Extraction"[ot] OR Tooth[ot] OR Dentistry[ot] OR Endodontics[ot] OR jsubsetd

#2

flossing[tiab] OR toothbrush*[tiab] OR brushing[tiab] OR dental[tiab] OR oral[tiab] OR periodont*[tiab] OR endodont*[tiab] OR gingiv*[tiab] OR mouth[tiab] OR hematogenous[tiab]

#3

"Bacterial Infections"[mh:noexp] OR Bacteremia[mh] OR Fungemia[mh] OR bacteremia[tiab] OR bacteraemia[tiab] OR fungemia[tiab] OR fungaemia[tiab] OR (Septicemia[mh:noexp] AND 1966[mhda]:1991[mhda]) OR Bacteremia[ot] OR "Streptococcal Infections"[ot] OR Septicemia[ot]

#4

"Anti-bacterial agents"[pa] OR "Anti-bacterial agents"[mh] OR "Antifungal Agents"[mh] OR "Anti-Infective Agents, Local"[mh] OR "Anti-Infective Agents"[mh:noexp] OR (Premedication[mh] AND 1973[mhda]:1995[mhda]) OR "Antibiotic Prophylaxis"[mh] OR ("Postoperative Complications"[mh] AND " Anti-bacterial agents/therapeutic use"[mh] AND 1968[mhda]:1975[mhda]) OR (antibiotic*[tiab] AND prophyla*[tiab]) OR "Prosthesis-Related Infections"[mh] OR Infection Control[mh] OR (Infection[mh:noexp] AND 1966[mhda]:1991[mhda])

#5

"Prostheses and Implants" [mh:noexp] OR "Bone Nails" [mh] OR "Bone Plates" [mh] OR "Bone Screws" [mh] OR "Internal Fixators" [mh] OR "Joint Prosthesis" [mh] OR Arthroplasty [mh] OR arthroplasty [tiab] OR ((joint [tiab] OR knee [tiab] OR hip [tiab]) AND (artificial [tiab] OR replacement [tiab] OR prosthe* [tiab])) OR (("Tissue Scaffolds" [mh] OR instrumentation [tiab] OR rod [tiab] OR rods [tiab] OR allograft* [tiab] OR "bone glass" OR (bone [tiab] AND void [tiab] AND filler* [tiab])) AND "Orthopedic Procedures" [mh]) OR "Bone Transplantation" [mh] OR ("Prosthesis Implantation" [mh] OR (silastic [tiab] AND (implant* [tiab] OR prosthes* [tiab])) AND ("Musculoskeletal System" [mh] OR Extremities [mh]))

#6

#1 AND #3

#7

#5 AND (#4 OR #3) AND (#2 OR #1)

#8

#6 OR #7

#9

English[lang]

#10

(animal[mh] NOT human[mh]) OR cadaver[mh] OR cadaver*[titl] OR ((comment[pt] OR editorial[pt] OR letter[pt] OR "historical article"[pt]) NOT "clinical trial"[pt]) OR addresses[pt] OR news[pt] OR "newspaper article"[pt] OR pmcbook

#11

#8 AND #9 NOT #10

Results sorted by study type

#12

Medline[tw] OR systematic review[tiab] OR Meta-analysis[pt]

#13

"Clinical Trial"[pt] OR (clinical[tiab] AND trial[tiab]) OR random*[tw] OR "Therapeutic use"[sh]

#14

#11 AND #12

#15

#11 AND #13 NOT #12

#16

#11 NOT (#13 OR #12)

EMBASE SEARCH STRATEGY

#1

Dentistry/exp OR Mouth/exp OR 'Dental Care'/exp OR 'mouth disease'/dm_dt,dm_su,dm_th,dm_rh,dm_dm OR 'mouth tumor'/dm_dt,dm_su,dm_th,dm_rh,dm_dm OR 'odontogenic cyst'/de OR 'odontogenic tumor'/de

#2

flossing:ti,ab OR toothbrush*:ti,ab OR dental:ti,ab OR peridont*:ti,ab OR endodont*:ti,ab OR gingiv*:ti,ab OR mouth:ti,ab OR hematogenous:ti,ab OR 'oral bacteria'

#3

'Bacterial Infection'/de OR Bacteremia/exp OR Fungemia/exp OR bacteremia:ti,ab OR bacteraemia:ti,ab OR fungemia:ti,ab OR fungaemia:ti,ab

#4

'Antiinfective Agent'/exp OR 'Antibiotic Prophylaxis'/de OR 'antibiotic prophylaxis' OR 'Prosthesis Infection'/de OR Infection/de

#5

'Joint Prosthesis'/exp OR 'Bone Nail'/de OR 'Bone Plate'/de OR 'Bone Screw'/de OR 'Internal Fixator'/de OR 'Pedicle Screw'/de OR 'Bone Graft'/exp OR 'tissue scaffold'/de OR 'bone void filler' OR ('silicone prosthesis'/de AND 'musculoskeletal system'/exp)

#6

#1 AND #3

#7

#5 AND (#4 OR #3) AND (#2 OR #1)

#8

#6 OR #7

#9

English:la AND [humans]/lim AND [embase]/lim

#10

cadaver/de OR 'in vitro study'/exp OR 'abstract report'/de OR book/de OR editorial/de OR note/de OR (letter/de NOT 'types of study'/exp)

#11

#8 AND #9 NOT #10

Results sorted by study type

#12

'meta analysis':ti,ab,de OR 'systematic review':ti,ab,de OR medline:ti,ab,de

#13

random*:ti,ab,de OR 'clinical trial':ti,ab,de OR 'health care quality'/exp

#14

#11 AND #12

#15

(#11 AND #13) NOT #12

#16

#11 NOT (#12 OR #13)

COCHRANE LIBRARY STRATEGY

(dental OR periodont* OR gingiv* OR mouth) AND (bacteremia OR bacteraemia OR fungemia OR fungaemia)

SUPPLEMENTAL SEARCH PUBMED/MEDLINE

#1

"Prostheses and Implants" [mh:noexp] OR "Bone Nails" [mh] OR "Bone Plates" [mh] OR "Bone Screws" [mh] OR "Internal Fixators" [mh] OR "Joint Prosthesis" [mh] OR Arthroplasty [mh] OR arthroplasty [tiab] OR ((joint [tiab] OR knee [tiab] OR hip [tiab]) AND (artificial [tiab] OR replacement [tiab] OR prosthe* [tiab])) OR (("Tissue Scaffolds" [mh] OR instrumentation [tiab] OR rod [tiab] OR rods [tiab] OR allograft* [tiab] OR "bone glass" OR (bone [tiab] AND void [tiab] AND filler* [tiab])) AND "Orthopedic Procedures" [mh]) OR "Bone Transplantation" [mh] OR ("Prosthesis Implantation" [mh] OR (silastic [tiab] AND (implant* [tiab] OR prosthes* [tiab])) AND ("Musculoskeletal System" [mh] OR Extremities [mh]))

#2

Bacteremia[mh] OR Fungemia[mh] OR bacteremia[tiab] OR bacteraemia[tiab] OR fungemia[tiab] OR fungaemia[tiab] OR hematogenous[tiab] OR haematogenous[tiab] OR "late infection" OR (late[titl] AND infection[titl])

#3

#1 AND #2

#4

"1960"[PDat]: "2011"[PDat] AND English[lang]

#5

(animal[mh] NOT human[mh]) OR cadaver[mh] OR cadaver*[titl] OR ((comment[pt] OR editorial[pt] OR letter[pt] OR "historical article"[pt]) NOT "clinical trial"[pt]) OR case reports[pt] OR addresses[pt] OR news[pt] OR "newspaper article"[pt] OR pmcbook

#6

#3 AND #4 NOT #5

#7

Medline[tw] OR systematic review[tiab] OR Meta-analysis[pt]

#8 (removed "therapeutic use"[sh] from published filter search string)
"Clinical Trial"[pt] OR (clinical[tiab] AND trial[tiab]) OR random*[tw]

#9 (added keywords for joint registries)

cohort studies[mh] OR cohort* OR (epidemiologic methods[mh:noexp] AND 1966[pdat]:1989[pdat]) OR case-control studies[mh] OR ((case OR cases) AND (control OR

controls OR controlled)) OR ((case OR cases) AND series*) OR registr* OR register*

#10

Microbiological Techniques[mh]

#11

#6 AND #7

#12

#6 AND #8 NOT #7

#13

#6 AND #9 NOT (#7 OR #8)

#14

#6 AND #10 NOT (#7 OR #8 OR #9)

#15

#6 NOT (#7 OR #8 OR #9 OR #10)

EMBASE

#1

Arthroplasty/exp OR 'Joint Prosthesis'/exp OR 'Bone Nail'/de OR 'Bone Plate'/de OR 'Bone Screw'/de OR 'Internal Fixator'/de OR 'Pedicle Screw'/de OR 'Bone Graft'/exp OR 'tissue scaffold'/de OR 'bone void filler' OR ('silicone prosthesis'/de AND 'musculoskeletal system'/exp)

#2

Bacteremia/exp OR Fungemia/exp OR bacteremia:ti,ab OR bacteraemia:ti,ab OR fungemia:ti,ab OR fungaemia:ti,ab

#3

#1 AND #2

#4

English:la AND [humans]/lim AND [embase]/lim

#5

cadaver/de OR 'in vitro study'/exp OR 'abstract report'/de OR book/de OR editorial/de OR note/de OR (letter/de NOT 'types of study'/exp)

#6

#3 AND #4 NOT #5

Results sorted by study type

```
#7
```

'meta analysis':ti,ab,de OR 'systematic review':ti,ab,de OR medline:ti,ab,de

#8

random*:ti,ab,de OR 'clinical trial':ti,ab,de OR 'health care quality'/exp

#9

'cohort analysis'/exp OR 'longitudinal study'/exp OR 'prospective study'/exp OR 'follow up'/exp OR cohort* OR 'case control study'/exp OR (case* AND control*)

#10

'microbiological examination'/exp

#11

#6 AND #7

#12

#6 AND #8 NOT #7

#13

#6 AND #9 NOT (#7 OR #8)

#14

#6 AND #10 NOT (#7 OR #8 OR #9)

#15

#6 NOT (#7 OR #8 OR #9 OR #10)

APPENDIX V EVALUATING QUALITY AND APPLICABILITY STUDIES OF INTERVENTIONS OUALITY

We judged quality using questions specified before this topic was selected and a computer program determined the final quality rating. We separately evaluated the quality of evidence for each outcome reported by each study. This follows the suggestion of the GRADE working group. We evaluated quality using a domain-based approach using a scheme to allow for evaluation of intervention studies of all designs. The domains we used are whether:

- The study was prospective (with prospective studies, it is possible to have an *a priori* hypothesis to test; this is not possible with retrospective studies.)
- The study was of low statistical power
- The assignment of patients to groups was unbiased
- There was blinding to mitigate against a placebo effect
- The patient groups were comparable at the beginning of the study
- The intervention was delivered in such a way that any observed effects could reasonably be attributed to that intervention
- Whether the instruments used to measure outcomes were valid
- Whether there was evidence of investigator bias

Each quality domain is addressed by one or more questions that are answered "Yes," "No," or "Unclear." These questions and the domains that each address are shown below.

To arrive at the quality of the evidence for a given outcome, all domains except the "Statistical Power" domain are termed as "flawed" if one or more questions addressing any given domain are answered "No" for a given outcome, or if there are two or more "Unclear" answers to the questions addressing that domain. The "Statistical Power" domain is considered flawed if a given study did not enroll enough patients to detect a standardized difference between means of 0.2.

Domain flaws lead to corresponding reductions in the quality of the evidence. The manner in which we conducted these reductions is shown in the table below. For example, the evidence reported in a randomized controlled trial (RCT) for any given outcome is rated as "High" quality if zero or one domain is flawed. If two or three domains are flawed for the evidence addressing this outcome, the quality of evidence is reduced to "Moderate," and if four or five domains are flawed, the quality of evidence is reduced to "Low." The quality of evidence is reduced to "Very Low" if six or more domains are flawed.

Some flaws are so serious that we automatically term the evidence as being of "Very Low" quality, regardless of a study's domain scores. These serious design flaws are:

- Non-consecutive enrollment of patients in a case series
- Case series that gave patients the treatment of interest AND another treatment
- Measuring the outcome of interest one way in some patients and measuring it in another way in other patients
- Low statistical power

Quality Questions and Domains for Four Designs of Studies of Interventions Parallel.

| | | O and to a second | 0 | 111-411 | 0 |
|--------------------------------|-----------------------------------|-------------------|---------------------|------------|--------|
| Damain | Overtions | Contemporary | Crossover | Historical | Case |
| Domain | Question: | Controls | Trials | Controls | Series |
| Group Assignment | Stochastic | Yes | Yes | No | No |
| Group Assignment | Quasi-random Assignment | No | No | No | na* |
| Group Assignment | Matched Groups | No | No | Yes | No |
| Group Assignment | Consecutive Enrollment | na | na | na | Yes |
| Prospective | Prospective | Yes | Yes | Yes | Yes |
| Blinding | Blinded Patients | Yes | Yes | No | No |
| Blinding | Blinded Assessors | Yes | Yes | No | No |
| Blinding | Blinding Verified | Yes | Yes | No | No |
| Group Comparability | Allocation Concealment | Yes | Yes | No | No |
| Group Comparability | >80% Follow-up | Yes | Yes | No | Yes |
| Group Comparability | <20% Completion Difference | Yes | Yes | No | No |
| Group Comparability | Similar Baseline Outcome Values | Yes | na | Yes | No |
| Group Comparability | Comparable Pt. Characteristics | Yes | na | Yes | No |
| Group Comparability | Same Control Group Results | na | Yes | na | na |
| Group Comparability | Same Experimental Group Results | na | Yes | na | na |
| Treatment Integrity | Same Centers | Yes | Yes | Yes | No |
| To a store a set but a suite : | Same Treatment Duration in and | | | | |
| Treatment Integrity | across All Groups | Yes | Yes | Yes | No |
| | Same Concomitant Treatment to All | | | | |
| Treatment Integrity | Groups (controlled studies only) | Yes | Yes | Yes | na |
| | No Confounding Treatment (case | 103 | 103 | 103 | Πα |
| Treatment Integrity | series only) | na | na | na | Yes |
| Measurement | Same Instruments | Yes | Yes | Yes | Yes |
| Measurement | Valid Instrument | Yes | Yes | Yes | Yes |
| Bias | Article & Abstract Agree | Yes | Yes | Yes | Yes |
| Bias | All Outcomes Reported | Yes | Yes | Yes | Yes |
| Bias | • | Yes | Yes | Yes | Yes |
| | A Priori Analysis | | | | |
| Statistical Power | Statistically Significant | High | High | High | High |
| Statistical Power | Number of patients in analysis | See per | ow for further info | rmation | |
| *"na" means | not asked | | | | |

Relationship between Quality and Domain Scores for Studies of Interventions

| Number of Flawed Domains | Strength of Evidence |
|--------------------------|----------------------|
| 0-1 | High |
| 2-3 | Moderate |
| 4-5 | Low |
| >5 | Very Low |

APPLICABILITY

We rated the applicability (also called "generalizability" or "external validity") of the evidence for each outcome reported by each study. As with quality, a computer program that used predetermined questions about specific applicability domains determined applicability ratings. We rated applicability as either "High", "Moderate", or "Low" depending on how many domains are flawed. As with quality, a domain is "flawed" if one or more questions addressing that domain is answered "No: or if two or more are answered "Unclear." We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a

given outcome, or if there are two or more "Unclear" answers to the questions addressing that domain

Our questions and domains about applicability are those of the PRECIS instrument. The instrument was originally designed to evaluate the applicability of randomized controlled trials, but it can also be used for studies of other design. The questions in this instrument fall into four domains. These domains and their corresponding questions are shown below. The applicability of a study is rated as "High" if it has no flawed domains, as "Low" if all domains are flawed, and as "Moderate" in all other cases as shown in the table below.

Applicability Questions and Domains for Studies of Interventions

| Question | Domain |
|----------------------------------------|-----------------------------|
| All Types of Patients Enrolled | Participants |
| Flexible Instructions to Practitioners | Interventions and Expertise |
| Full Range of Expt'l Practitioners | Interventions and Expertise |
| Usual Practice Control | Interventions and Expertise |
| Full Range of Control Practitioners | Interventions and Expertise |
| No Formal Follow-up | Interventions and Expertise |
| Usual and Meaningful Outcome | Interventions and Expertise |
| Compliance Not Measured | Compliance and Adherence |
| No Measure of Practitioner Adherence | Compliance and Adherence |
| All Patients in Analysis | Analysis |

Relationship between Applicability and Domain Scores for Interventions

| Number of Flawed Domains | Applicability |
|---------------------------------|---------------|
| 0 | High |
| 1, 2, 3 | Moderate |
| 4 | Low |

STUDIES OF INCIDENCE AND PREVALENCE QUALITY

Our appraisal of studies of incidence and prevalence is a domain-based approach conducted using *a priori* questions and scored by a computer program. The four domains we employed are:

- Outcome (whether the study is measuring the incidence/prevalence of a clinically meaningful event)
- Measurement (whether the study measured the disease/disorder/condition in a way that would lead to accurate estimates of incidence or prevalence)
- Participants (whether those who were studied were representative of the population of interest)
- Investigator Bias (whether author biases could have prejudiced the results)

Quality Questions and Domains for Studies of Incidence and Prevalence

| Question | Domain | Incidence | Prevalence |
|-------------------------------------------------|--------------|-----------|------------|
| Outcome Could Occur >1 Time in a Participant | None* | Yes | Yes |
| Study of Proportions or Number of Episodes | None | Yes | Yes |
| Only First Episode Counted | Measurement | Yes | Yes |
| Standard Methods for Collecting Outcomes Data | Outcome | Yes | Yes |
| Consistent Outcome Definitions | Outcome | Yes | Yes |
| Data Obtained from People or Records | None | Yes | Yes |
| Free from Response Bias | Measurement | Yes | Yes |
| Free from Information Bias | Measurement | Yes | Yes |
| Valid Instrument | Measurement | Yes | Yes |
| Valid Database Entries | Measurement | Yes | Yes |
| Study of In-Hospital Events | None | Yes | Yes |
| Use of Medical Records/Administrative Databases | Measurement | Yes | Yes |
| Appropriately Timed Outcome | Measurement | Yes | No |
| Chronic or Acute Disease | None | No | Yes |
| Study of Point Prevalence | None | No | Yes |
| Can Estimate Be Affected by Disease Severity | None | Yes | Yes |
| Correction for Disease Severity | Measurement | Yes | Yes |
| Population or Sample Data | None | Yes | Yes |
| Random Selection of Participants | Participants | Yes | Yes |
| >80% of Patients in Analysis | Participants | Yes | Yes |
| Free of Financial Conflicts of Interest | Bias | Yes | Yes |
| A Priori Analysis | Bias | Yes | Yes |
| Consistent Abstract, Results, Discussion | Bias | Yes | Yes |

^{*}An entry of "None" means that the question is not used in determining quality but, rather, is used for other purposes. A "Yes" entry in the above table means that a question is asked, a "No" entry means that it is not asked.

We characterized a study that has no flaws in any of its domains as being of "High" quality, a study that has one flawed domain as being of "Moderate" quality, a study with two flawed domains as being of "Low" quality, and a study with three or more flawed domains as being of "Very Low" quality. We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a given screening/diagnostic/test, or if there are two or more "Unclear" answers to the questions addressing that domain.

We considered some design flaws as so serious that their presence automatically guarantees that a study is characterized as being of "Very Low" quality regardless of its domain scores. These flaws are:

- The outcome of interest could have occurred more than once in a person during the course of the study, and more than the first episode of the outcome was counted in the incidence/prevalence estimate
- The study was a study of the proportion (or number) of people who have a disease, and the study was not a study of point prevalence.

Relationship between Quality and Domain Scores for Studies of Incidence and Prevalence

| Number of Flawed Domains | Quality |
|--------------------------|----------|
| 0 | High |
| 1 | Moderate |
| 2 | Low |
| ≥3 | Very Low |

APPLICABILITY

We separately evaluated the applicability of prevalence and incidence studies, and did so using a domain-based approach that involves predetermined questions and computer scoring. The domains we used for the applicability of prognostics are:

- Participants (i.e. whether the participants in the study were like those seen in the population of interest)
- Analysis (i.e., whether participants were appropriately included and excluded from the analysis)
- Outcome (i.e., whether the incidence/prevalence estimates being made were of a clinically meaningful outcome)

Applicability Questions and Domains for Studies of Incidence and Prevalence

| Question | Domain |
|-------------------------------|----------|
| Full Spectrum of Patients | Patients |
| All Patients in Analysis | Patients |
| No Stepwise Analysis | Analysis |
| Unambiguous Coding Scheme | Analysis |
| Model Validated | Analysis |
| Clinically Meaningful Outcome | Outcome |

We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a given screening/diagnostic/test, or if there are two or more "Unclear" answers to the questions addressing that domain. We characterized the applicability of a screening/diagnostic test as "High" if none of its domains are flawed, "Low" if all of its domains are flawed, and "Moderate" in all other cases.

Relationship between Applicability and Domain Scores for Studies of Incidence and Prevalence

| Number of Flawed Domains | Applicability |
|--------------------------|---------------|
| 0 | High |
| 1,2 | Moderate |
| 3 | Low |

STUDIES OF PROGNOSTICS

QUALITY

Our appraisal of studies of prognostics is a domain-based approach conducted using *a priori* questions, and scored by a computer program. The five domains we employed are:

- Prospective (A variable is specified as a potential prognostic variable *a priori*. This is not possible with retrospective studies.)
- Power (Whether the study had sufficient statistical power to detect a prognostic variable as statistically significant)
- Analysis (Whether the statistical analyses used to determine that a variable was rigorous to provide sound results)
- Model (Whether the final statistical model used to evaluate a prognostic variable accounted for enough variance to be statistically significant)
- Whether there was evidence of investigator bias

Quality Questions and Domains for Studies of Prognostics

| Question | Domain |
|-------------------------------------------------------|-------------------|
| Prospective | Prospective |
| At Least 10 Patients per Important Variable | Power |
| At Least 10 Events* | Power |
| All Important Variables Screened for Entry Into Model | Analysis |
| Interactions Tested | Analysis |
| Collinearity Absent | Analysis |
| Primary Analysis (not subgroup or post hoc) | Analysis |
| Statistically Significant Fit | Model |
| Article and Abstract Agree | Investigator Bias |
| Results Reported for All Variables Studies | Investigator Bias |
| Blinded Data Analysts** | Investigator Bias |

^{*}Asked only if the variable predicted by the prognostic is dichotomous.

We separately determined a quality score for each prognostic reported by a study. We characterized the evidence relevant to that prognostic variable as being of "High" quality if there are no flaws in any of the relevant domains, as being of "Moderate" quality if one of the relevant domains is flawed, as "Low" quality if there are two flawed domains, and as "Very Low" quality if three or more relevant domains are flawed. We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a given prognostic variable, or if there are two or more "Unclear" answers to the questions addressing that domain.

Relationship between Quality and Domain Scores for Studies of Prognostics

| Number of Flawed Domains | Quality |
|--------------------------|----------|
| 0 | High |
| 1 | Moderate |
| 2 | Low |
| ≥3 | Very Low |

^{**}Asked only if the prognostic variable is derived from a study that attempts to predict which patients respond best to a treatment.

APPLICABILITY

We separately evaluated the applicability of each prognostic variable reported in a study, and did so using a domain-based approach that involves predetermined questions and computer scoring. The domains we used for the applicability of prognostics are:

- Patients (i.e. whether the patients in the study and in the analysis were like those seen in actual clinical practice)
- Analysis (i.e., whether the analysis was not conducted in a way that was likely to describe variation among patients that might be unique to the dataset the authors used)
- Outcome (i.e., whether the prognostic was a predictor of a clinically meaningful outcome)

Applicability Questions and Domains for Studies of Prognostics

| Question | Domain |
|-------------------------------|----------|
| Full Spectrum of Patients | Patients |
| All Patients in Analysis | Patients |
| No Stepwise Analysis | Analysis |
| Unambiguous Coding Scheme | Analysis |
| Model Validated | Analysis |
| Clinically Meaningful Outcome | Outcome |

We characterized the evidence relevant to that prognostic as being of "High" applicability if there are no flaws in any of the relevant domains, as being of "Low" applicability if all three domains are flawed, and as of "Moderate" applicability in all other cases. We characterized a domain as "flawed" if one or more questions addressing any given domain are answered "No" for a given prognostic variable, or if there are two or more "Unclear" answers to the questions addressing that domain.

Relationship between Domain Scores and Applicability for Studies of Prognostics

| Number of Flawed Domains | Applicability |
|---------------------------------|---------------|
| 0 | High |
| 1,2 | Moderate |
| 3 | Low |

APPENDIX VI

RULES FOR OPINION BASED CONSENSUS RECOMMENDATIONS

A guideline can contain recommendations that are backed by little or no data. Under such circumstances, work groups often issue opinion-based recommendations. Although doing so is sometimes acceptable in an evidence-based guideline (expert opinion is a form of evidence), it is also important to avoid constructing a guideline that liberally uses expert opinion; research shows that expert opinion is often incorrect.

Opinion-based recommendations are developed only if they address a vitally important aspect of patient care. For example, constructing an opinion-based recommendation in favor of taking a history and physical is warranted. Constructing an opinion-based recommendation in favor of a specific modification of a surgical technique is seldom warranted. To ensure that an opinion-based recommendation is absolutely necessary, the AAOS has adopted rules to guide the content of the rationales that underpin such recommendations. These rules are based on those outlined by the US Preventive Services Task Force (USPSTF).²⁷ Specifically, rationales based on expert opinion must:

- Not contain references to or citations from articles not included in the systematic review that underpins the recommendation.
- Not contain the AAOS guideline language "We Recommend", "We suggest" or "The practitioner might".
- Contain an explanation of the potential preventable burden of disease. This involves considering both the incidence and/or prevalence of the disease, disorder, or condition and considering the associated burden of suffering. To paraphrase the USPSTF, when evidence is insufficient, provision of a treatment (or diagnostic) for a serious condition might be viewed more favorably than provision of a treatment (or diagnostic) for a condition that does not cause as much suffering. The AAOS (like the USPSTF) understand that evaluating the "burden of suffering" is subjective and involves judgment. This evaluation should be informed by patient values and concerns. The considerations outlined in this bullet make it difficult to recommend new technologies. It is not appropriate for a guideline to recommend widespread use of a technology backed by little data and for which there is limited experience.
- Address potential harms. In general, "When the evidence is insufficient, an intervention with a large potential for harm (such as major surgery) might be viewed less favorably than an intervention with a small potential for harm (such as advice to watch less television)."²⁷
- Address apparent discrepancies in the logic of different recommendations. Accordingly, if there are no relevant data for several recommendations and the work group chooses to issue an opinion-based recommendation in some cases but chooses not to make a recommendation in other cases, the rationales for the opinion-based recommendations must explain why this difference exists. Information garnered from the previous bullet points will be helpful in this regard.

- Consider current practice. The USPSTF specifically states that clinicians justifiably fear that not doing something that is done on a widespread basis will lead to litigation.²⁷ The consequences of not providing a service that is neither widely available nor widely used are less serious than the consequences of not providing a treatment accepted by the medical profession and thus expected by patients. Discussions of available treatments and procedures rely on mutual communication between the patient's guardian and physician, and on weighing the potential risks and benefits for a given patient. The patient's "expectation of treatment" must be tempered by the treating physician's guidance about the reasonable outcomes that the patient can expect.
- Justify, why a more costly device, drug, or procedure is being recommended over a less costly one whenever such an opinion-based recommendation is made.

Work group members write the rationales for opinion based recommendations on the first day of the final work group meeting. When the work group re-convenes on the second day of its meeting, it will vote on the rationales. The typical voting rules will apply. If the work group cannot adopt a rationale after three votes, the rationale and the opinion-based recommendation will be withdrawn, and a "recommendation" stating that the group can neither recommend for or against the recommendation in question will appear in the guideline.

Discussions of opinion-based rationales may cause some members to change their minds about whether to issue an opinion-based recommendation. Accordingly, at any time during the discussion of the rationale for an opinion-based recommendation, any member of the work group can make a motion to withdraw that recommendation and have the guideline state that the work group can neither recommend for or against the recommendation in question.

CHECKLIST FOR VOTING ON CONSENSUS RECOMMENDATIONS

- 1. When voting on the rationale, please consider the following:
- 2. Does the recommendation affect a substantial number of patients or address treatment (or diagnosis) of a condition that causes death and/or considerable suffering?
- 3. Does the recommendation address the potential harms that will be incurred if it is implemented and, if these harms are serious, does the recommendation justify;
 - a. why the potential benefits outweigh the potential harms and/or
 - b. why an alternative course of treatment (or diagnostic workup) that involves less serious or fewer harms is not being recommended?
- 4. Does the rationale explain why the work group chose to make a recommendation in the face of minimal evidence while, in other instances, it chose to make no recommendation in the face of a similar amount of evidence?
- 5. Does the rationale explain that the recommendation is consistent with current practice?
- 6. If relevant, does the rationale justify why a more costly device, drug, or procedure is being recommended over a less costly one?

APPENDIX VII

VOTING WITH THE NOMINAL GROUP TECHNIQUE

Voting on guideline recommendations will be conducted using a modification of the nominal group technique (NGT), a method previously used in guideline development.²⁸ Briefly each member of the guideline Work Group ranks his or her agreement with a guideline recommendation on a scale ranging from 1 to 9 (where 1 is "extremely inappropriate" and 9 is "extremely appropriate"). Consensus is obtained if the number of individuals who do not rate a measure as 7, 8, or 9 is statistically non-significant (as determined using the binomial distribution). Because the number of Work Group members who are allowed to dissent with the recommendation depends on statistical significance, the number of permissible dissenters varies with the size of the work group. The number of permissible dissenters for several work group sizes is given in the table below:

| Group Size | Number of Permissible Dissenters |
|------------|-------------------------------------|
| < 4 | group size not allowed |
| 4-5 | 0 |
| 6-8 | 1 |
| 9-11 | 1 |
| 12-14 | 2 |
| 15-16 | 3 |
| 17-19 | 4 |
| 20-22 | 5 |
| 23-24 | 6 |
| 25-27 | 7 |
| 28-29 | 8 |
| 30-32 | 9 |
| 33-34 | 10 |
| 35-36 | 11 |

The NGT is conducted by first having members vote on a given recommendation without discussion. If the number of dissenters is "permissible", the recommendation is adopted without further discussion. If the number of dissenters not permissible, there is further discussion to see whether the disagreement(s) can be resolved. Three rounds of voting are held to attempt to resolve disagreements. If disagreements are not resolved after three voting rounds, no recommendation is adopted.

APPENDIX VIII STRUCTURED PEER REVIEW FORM

Review of any AAOS confidential draft allows us to improve the overall guideline but <u>does not imply endorsement</u> by any given individual or any specialty society who participates in our review processes. The AAOS review process may result in changes to the documents; therefore, endorsement cannot occur until the AAOS Board of Directors officially approves the final guideline. The ADA will also employ a formal approval process.

Please note that if you return a review:

- Your review comments will be published on the AAOS website, and may be published on the ADA website, with our explanation of why we did or did not change the draft document in response to your comments.
- Your conflicts of interest disclosures will be published on the AAOS website, and may be published on the ADA website, with your review comments.

| Reviewer Information: Name of Reviewer: | |
|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------|
| Address: | |
| City: State: Zip Code: | |
| Phone: Fax: E-mail: | |
| Specialty Area/Discipline: | |
| Work setting: Credentials: | |
| May we list you as a Peer Reviewer in the final Guidelines? PLEASE READ: If you do not wish to be listed, your name will be removed for iden. However, your review comments, our responses and your COI will still be available you public review on our website with the posted Guideline if you complete this review. | v 1 1 |
| Are you reviewing this guideline as a representative of a professional society? | ☐ Yes ☐ No |
| If yes, may we list your society as a reviewer of this guideline? | Yes No |
| Society Name: (Listing the specialty society as a reviewing society does not imply or otherwise indica | te endorsement of this guideline.) |
| Conflicts of Interest (COI): All Reviewers must declare their conflicts of interest. If the boxes below are not checked and/or the reviewer does not attach his/her conflict be addressed by the AAOS nor will the reviewer's name or society be listed as a review guideline, only the chairperson or lead of the review must declare their relevant COI. | s of interest, the reviewer's comments will not |
| ☐ I have declared my conflicts of interest on page 2 of this form. | |
| ☐ I have declared my conflicts of interest in the AAOS database; my customer | # is |
| ☐ I understand that the AAOS will post my declared conflicts of interest with n this guideline on the AAOS website. | ny comments concerning review of |

REVIEWER CONFLICT OF INTEREST - The Orthopaedic Disclosure Program

Each item below requires an answer. Please report information for the last 12-months. ☐ Yes ☐ No Do you or a member of your immediate family receive royalties for any pharmaceutical, biomaterial or orthopaedic product or device? If YES, please identify product or device: ☐ Yes ☐ No Within the past twelve months, have you or a member of your immediate family served on the speakers bureau or have you been paid an honorarium to present by any pharmaceutical, biomaterial or orthopaedic product or device company? If YES, please identify company: ☐ Yes ☐ No Are you or a member of your immediate family a PAID EMPLOYEE for any pharmaceutical, biomaterial or orthopaedic device or equipment company, or supplier? If YES, please identify company or supplier: ☐ Yes ☐ No Are you or a member of your immediate family a PAID CONSULTANT for any pharmaceutical, biomaterial or orthopaedic device or equipment company, or supplier? If YES, please identify company or supplier: Are you or a member of your immediate family an UNPAID CONSULTANT for any pharmaceutical, Yes No biomaterial or orthopaedic device or equipment company, or supplier? If YES, please identify company or supplier: ☐ Yes ☐ No Do you or a member of your immediate family own stock or stock options in any pharmaceutical, biomaterial or orthopaedic device or equipment company, or supplier (excluding mutual funds) If YES, please identify company or supplier: Do you or a member of your immediate family receive research or institutional support as a principal Yes No investigator from any pharmaceutical, biomaterial or orthopaedic device or equipment company, or supplier? If YES, please identify company or supplier: ☐ Yes ☐ No Do you or a member of your immediate family receive any other financial or material support from any pharmaceutical, biomaterial or orthopaedic device and equipment company or supplier? If YES, please identify company or supplier: ☐ Yes ☐ No Do you or a member of your immediate family receive any royalties, financial or material support from any medical and/or orthopaedic publishers? If YES, please identify publisher: Yes No Do you or a member of your immediate family serve on the editorial or governing board of any medical and/or orthopaedic publication? If YES, please identify: ☐ Yes ☐ No Do you or a member of your immediate family serve on the Board of Directors or a committee of any medical and/or orthopaedic professional society? If YES, please identify:

Structured Peer Review Form Instructions

Please read and review this Draft Clinical Practice Guideline with particular focus on your area of expertise. Your responses will be used to assess the validity, clarity and accuracy of the interpretation of the evidence. If applicable, **please specify the draft page and line numbers in your comments**. Please feel free to also comment on the overall structure and content of the document. If you need more space than is provided, please attach additional pages.

| | Disagree | Somewhat Disagree | Somewh Agree | at Agree |
|---------------------------------------------------------------------------------------------------------------------------------------|----------|----------------------|-----------------|-------------|
| 1. The recommendations are clearly stated | | | | |
| 2. There is an explicit link between the recommendations and the supporting evidence | | | | |
| 3. Given the nature of the topic and the data, all clinically important outcomes are considered | | | | |
| 4. The guideline's target audience is clearly described | | | | |
| 5. The patients to whom this guideline is meant to apply are specifically described | | | | |
| 6. The criteria used to select articles for inclusion are appropriate | | | | |
| 7. The reasons why some studies were excluded are clearly described | | | | |
| 8. All important studies that met the article inclusion criteria are included | | | | |
| 9. The validity of the studies is appropriately appraised | | | | |
| 10. The methods are described in such a way as to be reproducible. | | | | |
| 11. The statistical methods are appropriate to the material and the objectives of this guideline | | | | |
| 12. Important parameters (e.g., setting, study population, study design) that could affect study results are systematically addressed | | | | |
| 13. Health benefits, side effects, and risks are adequately addressed | | | | |
| 14. The writing style is appropriate for health care professionals. | | | | |
| 15. The grades assigned to each recommendation are appropriate | | | | |

COMMENTS

PLEASE RETURN ALL COMMENTS IN WORD FORMAT

Please provide a brief explanation of both your positive and negative answers in the preceding section. If applicable, please specify the draft page and line numbers in your comments. Please feel free to also comment on the overall structure and content of the Guideline

| OVERALL ASSESSMENT |
|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Would you recommend these guidelines for use in clinical practice? (Check one) |
| ☐ Strongly recommend |
| Recommend (with provisions or alterations) |
| ☐ Would not recommend |
| Unsure |
| Note: Your answer to this question does not constitute an endorsement of this guideline. We ask this question as a means of monitoring the clinical relevance of our guideline. |

APPENDIX IX PEER REVIEW

Participation in the AAOS-ADA peer review process does not constitute an endorsement of this guideline by the participating organization.

Peer review of the draft guideline is completed by external organizations with an interest in the guideline. Outside peer reviewers are solicited for each AAOS guideline and consist of experts in the guideline's topic area. These experts represent professional societies other than AAOS and are nominated by the guideline work group prior to beginning work on the guideline. For this guideline, twenty-one outside peer review organizations were invited to review the draft guideline and all supporting documentation. Nine societies participated in the review of this guideline draft and seven explicitly consented to be listed as a peer review organization in this appendix. Two organizations did not give explicit consent that the organization name could be listed in this publication.

The organizations that reviewed the document and consented to be listed as a peer review organization are listed below:

American Academy of Family Physicians (AAFP)

American Association of Hip and Knee Surgeons (AAHKS)

American Association of Oral and Maxillofacial Surgeons (AOMS)

American Academy of Oral Pathology (AAOP)

American Academy of Pediatric Dentistry (AAPD)

American Association of Public Health Dentistry (AAPHD)

American Dental Association (ADA)

American Dental Hygienists' Association (ADHA)

Canadian Dental Association (CDA)

Infectious Disease Society of America (IDSA)

Lumbar Spine Research Society (LSRS)

North American Spine Society (NASS)

College of American Pathologists (CAP)

American Academy of Periodontology (AAP)

American College of Prosthodontists (ACP)

Society of Infectious Disease Pharmacists (SIDP)

Individuals who participated in the peer review of this document and gave their consent to be listed as reviewers of this document are:

Robert Rich, Jr. MD
Brian J. McGrory, MD
Louis G. Mercuri, DDS, MS
Sook-Bin Woo, DMD
A. Charles Post, DDS
Paul A. Moore, DMD, PhD, MPH
Paul S. Farsai, DMD, MPH
Denise Bowers, RDH, PhD
Charles Shuler, DMD, PhD

Susan Sutherland, DDS, MSc
John S. Kirkpatrick, MD, MS
Charles A. Reitman, MD
James M. Horton, MD
Tushar Patel, MD
Jamie Baisden, MD, FACS
John Steele, MD, PhD
Frank Scannaapieco, DMD, PhD
Mijin Choi, DDS, MS, FACP
Erika J. Ernst PharmD, BCPS

Participation in the AAOS-ADA guideline peer review process does not constitute an endorsement of the guideline by the participating organizations or the individuals listed above nor does it in any way imply the reviewer supports this document.

PUBLIC COMMENTARY

A period of public commentary follows the peer review of the draft guideline. If significant non-editorial changes are made to the document as a result of public commentary, these changes are also documented and forwarded to the AAOS and ADA bodies that approve the final guideline.

Public commentators who gave explicit consent to be listed in this document include the following:

Arlen D. Hanssen, MD Thomas K. Fehring, MD Laura MB Gehrig, MD Marc M. DeHart, MD

Participation in the AAOS-ADA guideline public commentary review process does not constitute an endorsement of the guideline by the participating organizations or the individual listed nor does it in any way imply the reviewer supports this document.

APPENDIX X

AAOS BODIES THAT APPROVED THIS CLINICAL PRACTICE GUIDELINE

Guidelines Oversight Committee

The AAOS Guidelines Oversight Committee (GOC) consists of sixteen AAOS members. The overall purpose of this Committee is to oversee the development of the clinical practice guidelines, performance measures, health technology assessments and utilization guidelines.

Evidence Based Practice Committee

The AAOS Evidence Based Practice Committee (EBPC) consists of ten AAOS members. This Committee provides review, planning, and oversight for all activities related to quality improvement in orthopaedic practice, including, but not limited to evidence-based guidelines, performance measures, and outcomes..

Council on Research and Quality

To enhance the mission of the AAOS, the Council on Research and Quality promotes the most ethically and scientifically sound basic, clinical, and translational research possible to ensure the future care for patients with musculoskeletal disorders. The Council also serves as the primary resource to educate its members, the public, and public policy makers regarding evidenced-based medical practice, orthopaedic devices and biologics regulatory pathways and standards development, patient safety, occupational health, technology assessment, and other related areas of importance.

Board of Directors

The 16 member AAOS Board of Directors manages the affairs of the AAOS, sets policy, and determines and continually reassesses the Strategic Plan.

ADA BODIES THAT APPROVED THIS CLINICAL PRACTICE GUIDELINE

Council on Scientific Affairs

The Council on Scientific Affairs (CSA) consists of seventeen ADA members. The CSA serves the public, the dental profession and other health professions as the primary source of timely, relevant and emerging information on the science of dentistry and promotion of oral health.

The CSA provides recommendations to the ADA's policymaking bodies on scientific issues. The Council also promotes, reviews, evaluates, and conducts studies on scientific matters.

DOCUMENTATION OF APPROVAL

AAOS-ADA Work Group Draft Completed: February 12, 2012 Peer Review Completed: March 15, 2012 August 27, 2012 Public Commentary Completed: AAOS Guidelines Oversight Committee: September 22, 2012 September 22, 2012 AAOS Evidence Based Practice Committee: AAOS Council on Research and Quality: October 26, 2012 December 7, 2012 AAOS Board of Directors: November 13, 2012 ADA Council on Scientific Affairs

APPENDIX XI SUPPLEMENTAL EVIDENCE TABLES

Table 59 Antibiotic Prophylaxis Network Meta-Analysis Consistency Check

| | | | | Direct | | | | |
|------------------|--------|-------|--------|--------|---------|-------|-------|------|
| | MC | | Direct | SD (Ln | | SD | | |
| Comparison | Mean | MC SD | Ln OR | OR) | Omega | Omega | Z | p |
| Placebo vs: | | | | | | | | |
| Amoxicillin | -2.638 | 0.465 | -2.375 | 0.420 | -1.174 | 0 | 0.000 | 1.00 |
| Penicillin | -1.738 | 0.695 | -1.266 | 0.486 | -0.451 | 0 | 0.000 | 1.00 |
| Erythromycin | -0.852 | 0.664 | -0.669 | 0.512 | -0.267 | 0 | 0.000 | 1.00 |
| Clindamycin | -1.453 | 0.650 | -2.112 | 0.462 | 0.672 | 0 | 0.000 | 1.00 |
| Josamycin | -0.187 | 1.114 | 0.228 | 0.677 | -0.243 | 0 | 0.000 | 1.00 |
| Moxifloxacin | -2.676 | 0.961 | -2.957 | 0.765 | 0.485 | 0 | 0.000 | 1.00 |
| IV Tetracycline | -4.123 | 1.144 | -4.075 | 0.635 | -0.022 | 0 | 0.000 | 1.00 |
| IM Teicoplanin | -2.312 | 1.343 | -2.674 | 1.570 | -1.347 | 3.030 | 0.444 | 0.66 |
| Topical | | | | | | | | |
| Amoxicillin | -1.118 | 1.217 | -2.064 | 1.174 | 12.797 | 0 | 0.000 | 1.00 |
| Antiseptic Rinse | -1.424 | 1.048 | -0.989 | 0.494 | -0.124 | 0 | 0.000 | 1.00 |
| Amoxicillin vs: | | | | | | | | |
| Penicillin | 0.900 | 0.731 | -0.003 | 0.600 | 1.862 | 0 | 0.000 | 1.00 |
| Clindamycin | 1.185 | 0.725 | 1.892 | 0.467 | -0.503 | 0 | 0.000 | 1.00 |
| Moxifloxacin | -0.038 | 0.959 | 0.421 | 0.377 | -0.084 | 0 | 0.000 | 1.00 |
| IM Teicoplanin | 0.327 | 1.299 | 0.811 | 0.913 | -0.472 | 0 | 0.000 | 1.00 |
| Topical | | | | | | | | |
| Amoxicillin | 1.521 | 1.215 | 2.436 | 1.170 | -11.541 | 0 | 0.000 | 1.00 |
| Penicillin vs: | | | | | | | | |
| Antiseptic Rinse | 0.314 | 1.06 | -0.161 | 0.568 | 0.191 | 0 | 0.000 | 1.00 |
| Erythromycin vs: | | | | | | | | |
| Clindamycin | -0.601 | 0.696 | -0.357 | 0.568 | -0.484 | 0 | 0.000 | 1.00 |
| Josamycin | 0.665 | 1.113 | 0.228 | 0.677 | 0.257 | 0 | 0.000 | 1.00 |
| IV Tetracycline | -3.271 | 1.148 | -2.996 | 0.670 | -0.142 | 0 | 0.000 | 1.00 |
| Clindamycin vs: | | | | | | | | |
| Moxifloxacin | -1.223 | 0.994 | -1.470 | 0.465 | 0.069 | 0 | 0.000 | 1.00 |

Table 60 Topical Antimicrobial Prophylaxis Network Meta-Analysis Consistency Check

| | | | | Direct | | | | |
|-----------------------------------------------------------|------------|----------|----------|------------|-------|-------------|------|------|
| | 3.50 | 3.40 | Direct | SD | | CIP. | | |
| | MC Mean | MC SD | Ln OR | (Ln OR) | Omega | SD Omega | Z | n |
| No Treatment vs: | Mean | SD | OK | OK) | Omega | Omega | L | р |
| Saline Rinse | -0.04 | 0.45 | -0.25 | 0.58 | -0.54 | 0.93 | 0.58 | 0.56 |
| Chlorhexidine Rinse | -0.04 | 0.43 | -1.52 | 0.50 | -0.54 | 0.93 | 0.00 | 1.00 |
| Povidone-Iodine Rinse | -1.77 | 0.52 | -1.47 | 0.68 | 1.24 | 1.09 | 1.14 | 0.26 |
| Chloramine T Rinse/Brush | -1.94 | 0.90 | -1.74 | 0.61 | -0.09 | 0.00 | 0.00 | 1.00 |
| | -0.30 | 1.00 | -0.27 | 0.74 | -0.03 | 0.00 | 0.00 | 1.00 |
| Lugol's Solution Rinse | | | | | | | | |
| Hydrogen Peroxide Rinse Sodium Perborate-Ascorbic Acid | -1.10 | 0.56 | -0.97 | 0.35 | -0.08 | 0.00 | 0.00 | 1.00 |
| Rinse | -1.75 | 0.71 | -1.56 | 0.42 | -0.10 | 0.00 | 0.00 | 1.00 |
| Phenolated Rinse | -1.53 | 0.51 | -1.65 | 0.53 | -1.76 | 2.05 | 0.86 | 0.39 |
| | | | -100 | | | | | |
| Saline Rinse vs: | | | | | | | | |
| Chlorhexidine Rinse | -1.73 | 0.60 | -2.49 | 0.76 | -1.96 | 1.22 | 1.60 | 0.11 |
| Povidone-Iodine Rinse | -1.90 | 0.60 | -1.79 | 0.72 | 0.36 | 1.31 | 0.28 | 0.78 |
| Sodium Perborate-Ascorbic Acid | | | | | | | | |
| Rinse | -1.71 | 0.74 | -1.92 | 0.55 | 0.27 | 0.00 | 0.00 | 1.00 |
| Phenolated Rinse | -1.49 | 0.59 | -1.37 | 0.40 | -0.10 | 0.00 | 0.00 | 1.00 |
| | | | | | | | | |
| Chlorhexidine Rinse vs: | | | | | | | | |
| Povidone-Iodine Rinse | -0.17 | 0.45 | -0.21 | 0.43 | 0.30 | 0.00 | 0.00 | 1.00 |
| Hydrogen Peroxide Rinse | 0.67 | 0.64 | 0.41 | 0.64 | 22.85 | 0.00 | 0.00 | 1.00 |
| Placebo Rinse | 0.85 | 0.53 | 0.47 | 0.40 | 0.51 | 0.00 | 0.00 | 1.00 |
| | | | | | | | | |
| Povidone-Iodine Rinse vs: | | | | | | | | |
| Hydrogen Peroxide Rinse | 0.84 | 0.65 | 0.62 | 0.65 | 17.68 | 0.00 | 0.00 | 1.00 |
| Placebo Rinse | 1.03 | 0.51 | 1.12 | 0.35 | -0.09 | 0.00 | 0.00 | 1.00 |
| Operative Field Isolation | -0.89 | 0.79 | -0.87 | 0.47 | -0.02 | 0.00 | 0.00 | 1.00 |
| Isolation + Iodine Rinse | -1.02 | 0.79 | -0.98 | 0.48 | -0.02 | 0.00 | 0.00 | 1.00 |
| Isolation + Chlorhexidine Rinse | -2.20 | 0.87 | -2.10 | 0.58 | -0.08 | 0.00 | 0.00 | 1.00 |
| | | | | | | | | |
| Chloramine T Rinse/Brush vs: | | | | | | | | |
| Lugol's Solution Rinse | 1.54 | 0.87 | 1.47 | 0.57 | 0.06 | 0.00 | 0.00 | 1.00 |
| | | | | | | | | |
| Operative Field Isolation vs: | | | | | | | | |
| Isolation + Iodine Rinse | -0.12 | 0.79 | -0.12 | 0.49 | 0.00 | 0.00 | 0.00 | 1.00 |
| Isolation + Chlorhexidine Rinse | -1.31 | 0.87 | -1.23 | 0.59 | -0.06 | 0.00 | 0.00 | 1.00 |
| | | | | | | | | |
| Isolation + Iodine Rinse vs : | | | | | | | | |
| Isolation + Chlorhexidine Rinse | -1.18 | 0.87 | -1.11 | 0.59 | -0.06 | 0.00 | 0.00 | 1.00 |

Table 61 Goodness-of-fit Statistics

| | Data Points | Residual Deviance | |
|-------------------------------------------|-------------|-------------------|--|
| Antibiotic Prophylaxis Network | 43 | 43.03 | |
| Topical Antimicrobial Prophylaxis Network | 33 | 31.31 | |

Table 62 Antibiotic and Topical Antimicrobial Prophylaxis Network Meta-Analysis Consistency Check

| | | | | Direct | | | | | |
|----------------------------|-------|------|--------|--------|--------|--------|-------|------|------|
| | | | Direct | SD | | | | | |
| | MC | MC | Ln | (Ln | Direct | | SD | | |
| | Mean | SD | OR | OR) | Var | Omega | Omega | Z | p |
| Placebo Pill/No Treatment | | | | | | | | | |
| vs: | | | | | | | | | |
| Amoxicillin | -2.56 | 0.37 | -2.38 | 0.42 | 0.18 | 0.81 | 0.89 | 0.91 | 0.36 |
| Penicillin | -1.68 | 0.57 | -1.27 | 0.49 | 0.24 | -1.09 | 0.00 | 0.00 | 1.00 |
| Erythromycin | -0.85 | 0.53 | -0.67 | 0.51 | 0.26 | -3.21 | 0.00 | 0.00 | 1.00 |
| Clindamycin | -1.44 | 0.51 | -2.11 | 0.46 | 0.21 | 2.82 | 0.00 | 0.00 | 1.00 |
| Josamycin | -0.18 | 0.91 | 0.23 | 0.68 | 0.46 | -0.50 | 0.00 | 0.00 | 1.00 |
| Moxifloxacin | -2.61 | 0.74 | -2.96 | 0.77 | 0.59 | -5.85 | 3.13 | 1.87 | 0.06 |
| IV Tetracycline | -4.13 | 0.95 | -4.07 | 0.63 | 0.40 | -0.04 | 0.00 | 0.00 | 1.00 |
| IM Teicoplanin | -2.16 | 1.16 | -2.67 | 1.57 | 2.47 | -1.12 | 2.32 | 0.48 | 0.63 |
| Topical Amoxicillin | -1.07 | 1.02 | -2.06 | 1.17 | 1.38 | -4.14 | 2.40 | 1.72 | 0.08 |
| Chlorhexidine or Povidone- | | | | | | | | | |
| Iodine Rinse | -1.38 | 0.83 | -0.99 | 0.49 | 0.24 | -0.22 | 0.00 | 0.00 | 1.00 |
| Saline Rinse | -0.01 | 0.50 | -0.80 | 0.36 | 0.13 | 0.80 | 0.00 | 0.00 | 1.00 |
| Chlorhexidine Rinse | -1.78 | 0.57 | -1.52 | 0.51 | 0.26 | -1.16 | 0.00 | 0.00 | 1.00 |
| Povidone-Iodine Rinse | -1.93 | 0.59 | -1.47 | 0.68 | 0.46 | 1.83 | 1.35 | 1.36 | 0.17 |
| Chloramine T Rinse/Brush | -1.84 | 0.96 | -1.74 | 0.61 | 0.38 | -0.07 | 0.00 | 0.00 | 1.00 |
| Lugol's Solution Rinse | -0.29 | 1.06 | -0.27 | 0.74 | 0.55 | -0.01 | 0.00 | 0.00 | 1.00 |
| Hydrogen Peroxide Rinse | -1.11 | 0.62 | -0.97 | 0.35 | 0.12 | -0.06 | 0.00 | 0.00 | 1.00 |
| Phenolated Rinse | -1.52 | 0.57 | -1.65 | 0.53 | 0.28 | 0.85 | 0.00 | 0.00 | 1.00 |
| Sodium Perborate-Ascorbic | | | | | | | | | |
| Acid Rinse | -1.76 | 0.79 | -1.56 | 0.42 | 0.17 | -0.08 | 0.00 | 0.00 | 1.00 |
| Amoxicillin vs: | | | | | | | | | |
| Penicillin | 0.88 | 0.60 | 0.00 | 0.60 | 0.36 | 507.16 | 0.00 | 0.00 | 1.00 |
| Clindamycin | 1.12 | 0.57 | 1.89 | 0.47 | 0.22 | -1.55 | 0.00 | 0.00 | 1.00 |
| Moxifloxacin | -0.05 | 0.73 | 0.42 | 0.38 | 0.14 | -0.17 | 0.00 | 0.00 | 1.00 |
| IM Teicoplanin | 0.39 | 1.12 | 0.42 | 0.91 | 0.14 | -0.17 | 0.00 | 0.00 | 1.00 |
| - | | 1.12 | 2.44 | 1.17 | 1.37 | 4.03 | 2.41 | 1.68 | 0.09 |
| Topical Amoxicillin | 1.48 | 1.02 | 2.44 | 1.1/ | 1.37 | 4.03 | ∠.41 | 1.08 | 0.09 |

| | | | D : (| Direct | | | | | |
|--------------------------------------------|---------------|------------|--------------------|------------------|---------------|-------|-------------|---------|------|
| | MC Mean | MC SD | Direct Ln OR | SD (Ln OR) | Direct Var | Omega | SD Omega | ${f Z}$ | р |
| | IVICUII | D D | <u> </u> | O1 () | , ui | omegu | Omega | | |
| Penicillin vs: | | | | | | | | | |
| Chlorhexidine or Povidone- Iodine Rinse | 0.30 | 0.85 | -0.16 | 0.57 | 0.32 | 0.37 | 0.00 | 0.00 | 1.00 |
| Erythromycin vs: | | | | | | | | | |
| Clindamycin | -0.59 | 0.56 | -0.36 | 0.57 | 0.32 | 11.20 | 3.94 | 2.84 | 0.00 |
| • | -0.39 0.67 | 0.50 | 0.23 | 0.57 | 0.32 | 0.55 | 0.00 | 0.00 | 1.00 |
| Josamycin | | | | | | | | | |
| IV Tetracycline | -3.28 | 0.96 | -3.00 | 0.67 | 0.45 | -0.27 | 0.00 | 0.00 | 1.00 |
| Clindamycin vs: | | | | | | | | | |
| Moxifloxacin | -1.17 | 0.77 | -1.47 | 0.46 | 0.22 | 0.17 | 0.00 | 0.00 | 1.00 |
| Saline Rinse vs: | | | | | | | | | |
| Chlorhexidine Rinse | -1.77 | 0.65 | -2.49 | 0.76 | 0.58 | -2.66 | 1.47 | 1.81 | 0.07 |
| Povidone-Iodine Rinse | -1.92 | 0.66 | -1.79 | 0.72 | 0.52 | 0.79 | 1.80 | 0.44 | 0.66 |
| Phenolated Rinse | -1.51 | 0.66 | -1.37 | 0.40 | 0.16 | -0.08 | 0.00 | 0.00 | 1.00 |
| Sodium Perborate-Ascorbic | | | | | | | | | |
| Acid Rinse | -1.75 | 0.82 | -1.92 | 0.55 | 0.30 | 0.14 | 0.00 | 0.00 | 1.00 |
| Chlorhexidine Rinse vs: | | | | | | | | | |
| Povidone-Iodine Rinse | -0.15 | 0.49 | -0.21 | 0.43 | 0.18 | 0.19 | 0.00 | 0.00 | 1.00 |
| Hydrogen Peroxide Rinse | 0.67 | 0.70 | 0.41 | 0.64 | 0.41 | 1.25 | 0.00 | 0.00 | 1.00 |
| Placebo Rinse | 0.88 | 0.58 | 0.47 | 0.40 | 0.16 | 0.37 | 0.00 | 0.00 | 1.00 |
| Povidone-Iodine Rinse vs: | | | | | | | | | |
| Hydrogen Peroxide Rinse | 0.82 | 0.71 | 0.62 | 0.65 | 0.42 | 0.91 | 0.00 | 0.00 | 1.00 |
| Placebo Rinse | 1.03 | 0.56 | 1.12 | 0.35 | 0.13 | -0.07 | 0.00 | 0.00 | 1.00 |
| Operative Field Isolation | -0.90 | 0.87 | -0.87 | 0.47 | 0.22 | -0.01 | 0.00 | 0.00 | 1.00 |
| Isolation + Iodine Rinse | -1.02 | 0.87 | -0.98 | 0.48 | 0.23 | -0.01 | 0.00 | 0.00 | 1.00 |
| Isolation + Chlorhexidine | | | 017 0 | | | | | | -100 |
| Rinse | -2.20 | 0.94 | -2.10 | 0.58 | 0.34 | -0.06 | 0.00 | 0.00 | 1.00 |
| Chloramine T Rinse/Brush | | | | | | | | | |
| vs: | | | | | | | | | |
| Lugol's Solution Rinse | 1.55 | 0.94 | 1.47 | 0.57 | 0.33 | 0.05 | 0.00 | 0.00 | 1.00 |
| Operative Field Isolation vs: | | | | | | | | | |
| Isolation + Iodine Rinse | -0.12 | 0.88 | -0.12 | 0.49 | 0.24 | 0.00 | 0.00 | 0.00 | 1.00 |
| Isolation + Chlorhexidine | | _ | | | | | | | |
| Rinse | -1.31 | 0.95 | -1.23 | 0.59 | 0.35 | -0.05 | 0.00 | 0.00 | 1.00 |

Direct Direct SD MCMC(Ln **Direct** SD Ln OR) Omega Omega Z SD Mean \mathbf{OR} Var p **Isolation + Iodine Rinse vs:** Isolation + Chlorhexidine Rinse -1.18 0.95 -1.11 0.59 0.35 -0.04 0.000.00 1.00

Table 63 Bacteremia Incidence Study Details

| Study | Procedure | N | n | Rate | LowerCI | UpperCI | SD |
|----------------|-------------|-----|----|----------|----------|----------|----------|
| Bhanji 2002 | brushing | 47 | 29 | 0.617021 | 0.474266 | 0.742093 | 0.068325 |
| Forner 2006 | brushing | 60 | 2 | 0.033333 | 0.009189 | 0.113638 | 0.026646 |
| Lockhart 2008 | brushing | 88 | 28 | 0.318182 | 0.230226 | 0.421348 | 0.048756 |
| Sconyers 1973 | brushing | 30 | 5 | 0.166667 | 0.073365 | 0.335644 | 0.066909 |
| Sconyers 1979 | brushing | 50 | 0 | 0 | 0 | 0.071348 | 0.018202 |
| Forner 2006 | chewing | 60 | 4 | 0.066667 | 0.026229 | 0.159254 | 0.033936 |
| Murphy 2006 | chewing | 21 | 0 | 0 | 0 | 0.154639 | 0.03945 |
| | dental | | | | | | |
| Pineiro 2010 | implants | 30 | 2 | 0.066667 | 0.018477 | 0.213235 | 0.049684 |
| Cherry 2007 | prophylaxis | 60 | 11 | 0.333333 | 0.218739 | 0.544864 | 0.05922 |
| De Leo 1974 | prophylaxis | 39 | 11 | 0.282051 | 0.165435 | 0.437753 | 0.06947 |
| Forner 2006 | prophylaxis | 19 | 21 | 0.35 | 0.241678 | 0.476374 | 0.100626 |
| Forner 2006 | prophylaxis | 20 | 15 | 0.75 | 0.531299 | 0.888138 | 0.091032 |
| Heimdahl | | | | | | | |
| 1990 | prophylaxis | 20 | 14 | 0.7 | 0.481027 | 0.854523 | 0.095281 |
| Baumgartner | | | | | | | |
| 1976 | endodontics | 30 | 1 | 0.033333 | 0.005909 | 0.166704 | 0.04102 |
| Baumgartner | | | | | | | |
| 1977 | endodontics | 12 | 7 | 0.583333 | 0.319511 | 0.80674 | 0.124295 |
| Bender 1963 | endodontics | 98 | 15 | 0.153061 | 0.095007 | 0.237289 | 0.036297 |
| Heimdahl | | | | | | | |
| 1990 | endodontics | 20 | 4 | 0.2 | 0.080658 | 0.416017 | 0.085552 |
| Savarrio 2004 | endodontics | 30 | 9 | 0.3 | 0.166647 | 0.478758 | 0.079621 |
| Bender 1963 | extraction | 33 | 17 | 0.515152 | 0.352184 | 0.67496 | 0.082342 |
| Casolari 1989 | extraction | 56 | 38 | 0.678571 | 0.548226 | 0.78599 | 0.060655 |
| Heimdahl | | | | | | | |
| 1990 | extraction | 20 | 20 | 1 | 0.838875 | 1 | 0.041105 |
| Khairat 1966 | extraction | 100 | 64 | 0.64 | 0.542354 | 0.727288 | 0.047178 |
| Rahn 1994 | injection | 40 | 21 | 0.525 | 0.374974 | 0.670645 | 0.075428 |
| American | | | | | | | |
| Academy of | | | | | | | |
| Periodontology | interdental | | | | | | |
| 1972 | cleaner | 60 | 17 | 0.283333 | 0.185068 | 0.407673 | 0.056788 |
| | interdental | | | | | | |
| Berger 1974 | cleaner | 30 | 8 | 0.266667 | 0.141827 | 0.44448 | 0.077209 |
| | interdental | | | | | | |
| Crasta 2009 | cleaner | 59 | 24 | 0.40678 | 0.29089 | 0.534066 | 0.062036 |

| Study | Procedure | N | n | Rate | LowerCI | UpperCI | SD |
|------------------------|------------------------|------------|----|----------|----------|----------|-----------|
| | interdental | | | | | | |
| Felix 1971 | cleaner | 30 | 15 | 0.5 | 0.331541 | 0.668459 | 0.08595 |
| Lineberger | interdental | | | | | | |
| 1973 | cleaner | 30 | 8 | 0.266667 | 0.141827 | 0.44448 | 0.077209 |
| D 1 1055 | interdental | ~ 0 | | 0.40 | 0.00==00 | 0.0000 | 0.050.00 |
| Ramadan 1975 | cleaner | 50 | 9 | 0.18 | 0.097702 | 0.307961 | 0.053638 |
| D 10 - 1 | interdental | 20 | | 0.0 | | | 0.040.504 |
| Romans 1971 | cleaner interdental | 30 | 2 | 0.066667 | | | 0.049684 |
| Wank 1976 | cleaner | 21 | 6 | 0.285714 | 0.138139 | 0.499564 | 0.092202 |
| Ali 1992 | intubation | 36 | 0 | 0.111111 | 0.044066 | 0.253148 | 0.053338 |
| Berry 1973 | intubation | 50 | 4 | 0.08 | 0.03155 | 0.188382 | 0.040009 |
| Dinner 1987 | intubation | 54 | 3 | 0.055556 | 0.019073 | 0.151072 | 0.033674 |
| Hansen 1989 | intubation | 19 | 1 | 0.052632 | 0.009352 | 0.246387 | 0.060469 |
| Oncag 2005 | intubation | 74 | 9 | 0.121622 | 0.065323 | 0.215266 | 0.038251 |
| Valdes 2008 | intubation | 110 | 13 | 0.118182 | 0.070381 | 0.19175 | 0.030962 |
| Enabulele | | | | | | | |
| 2008 | oral surgery | 50 | 16 | 0.32 | 0.207582 | 0.458103 | 0.063909 |
| Heimdahl | | | | | | | |
| 1990 | oral surgery | 20 | 11 | 0.55 | 0.342085 | 0.741802 | 0.10197 |
| Josefsson 1985 | oral surgery | 20 | 11 | 0.55 | 0.342085 | 0.741802 | 0.10197 |
| Takai 2005 | oral surgery | 57 | 33 | 0.578947 | 0.449801 | 0.698124 | 0.063349 |
| Erverdi 1999 | orthodontics | 40 | 3 | 0.075 | 0.025836 | 0.198642 | 0.044084 |
| Gürel 2009 | orthodontics | 25 | 8 | 0.32 | 0.172052 | 0.515897 | 0.087717 |
| | periodontics | | | | | | |
| | scaling root | | | | | | |
| Bender 1963 | planing | 15 | 8 | 0.533333 | 0.30117 | 0.751905 | 0.114985 |
| | periodontics | | | | | | |
| | scaling root | | | | | | |
| Casolari 1989 | planing | 42 | 12 | 0.285714 | 0.17167 | 0.435672 | 0.067349 |
| | periodontics | | | | | | |
| | scaling root | | | | | | |
| Lafaurie 2007 | planing | 42 | 34 | 0.809524 | 0.666992 | 0.90018 | 0.059488 |
| | periodontics | | | | | | |
| | scaling root | | _ | | | | |
| Lofthus 1991 | planing | 10 | 3 | 0.3 | 0.107791 | 0.603222 | 0.126387 |
| | periodontics | | | | | | |
| Lucartorto | scaling root | | | | | | |
| 1992 | planing | 41 | 13 | 0.317073 | 0.195646 | 0.469842 | 0.069949 |
| | periodontics | | | | | | |
| Morozumi | scaling root | | | | | | |
| 2010 | planing | 10 | 9 | 0.9 | 0.59585 | 0.982124 | 0.098541 |
| | periodontics | | | | | | |
| | scaling root | | | | | | |
| Waki 1990 | planing | 15 | 2 | 0.133333 | 0.037361 | 0.37882 | 0.087108 |
| | periodontics | | | | | | |
| Bender 1963 | gingivectomy | 12 | 10 | 0.833333 | 0.551969 | 0.953035 | 0.102314 |
| Lineberger | periodontics | | | | | 0.0 | |
| 1973 | gingivectomy | 10 | 6 | 0.6 | 0.312674 | 0.83182 | 0.132437 |

| Study | Procedure | N n Rate | | LowerCI | UpperCI | SD | |
|--------------|--------------|----------|----|----------|----------|----------|----------|
| | periodontics | | | | | | |
| Rogosa 1960 | gingivectomy | 13 | 12 | 0.923077 | 0.66686 | 0.98629 | 0.081489 |
| | periodontics | | | | | | |
| Wada 1968 | gingivectomy | 77 | 20 | 0.25974 | 0.174892 | 0.367422 | 0.049116 |
| | periodontics | | | | | | |
| Daly 1997 | probing | 30 | 13 | 0.433333 | 0.273775 | 0.608027 | 0.08527 |
| | periodontics | | | | | | |
| Daly 2001 | probing | 40 | 10 | 0.25 | 0.141871 | 0.40194 | 0.066345 |
| Oncag 2006 | restorative | 23 | 3 | 0.130435 | 0.045377 | 0.321275 | 0.070383 |
| Brown 1998 | suture | 24 | 2 | 0.083333 | 0.023159 | 0.258488 | 0.060034 |
| King 1988 | suture | 20 | 1 | 0.05 | 0.008881 | 0.236131 | 0.057973 |
| Wampole 1978 | suture | 20 | 5 | 0.25 | 0.111862 | 0.468701 | 0.091032 |

Table 64 Bacteremia Prevalence Study Details

| Study | Procedure | N | n | Rate | LowerCI | LowerCI UpperCI | |
|----------------|--------------|-----|-----|----------|----------|-----------------|----------|
| Lucas 2000 | brushing | 52 | 20 | 0.384615 | 0.264705 | 0.520401 | 0.06523 |
| Silver 1979 | brushing | 36 | 3 | 0.083333 | 0.028749 | 0.218267 | 0.048347 |
| Degling 1972 | chewing | 40 | 0 | 0 | 0 | 0.087622 | 0.022353 |
| Trivedi 1984 | chewing | 20 | 2 | 0.1 | 0.027866 | 0.301034 | 0.069687 |
| Marzoni 1983 | cleft palate | 14 | 6 | 0.428571 | 0.213808 | 0.674094 | 0.117421 |
| Lucas 1999 | prophylaxis | 103 | 33 | 0.320388 | 0.238131 | 0.415562 | 0.045264 |
| Trivedi 1984 | prophylaxis | 40 | 22 | 0.55 | 0.398291 | 0.692947 | 0.075169 |
| Winslow 1960 | prophylaxis | 72 | 17 | 0.236111 | 0.152967 | 0.345988 | 0.049241 |
| Debelian 1995 | endodontics | 26 | 11 | 0.423077 | 0.255444 | 0.610514 | 0.09058 |
| Barbosa 2010 | extraction | 210 | 149 | 0.709524 | 0.644796 | 0.766723 | 0.031104 |
| Coulter 1990 | extraction | 32 | 20 | 0.625 | 0.452544 | 0.770661 | 0.081154 |
| Crawford | | | | | | | |
| 1973 | extraction | 25 | 23 | 0.92 | 0.750339 | 0.97778 | 0.058022 |
| DeVries 1972 | extraction | 100 | 49 | 0.49 | 0.39422 | 0.58652 | 0.049057 |
| Khairat 1966 | extraction | 100 | 64 | 0.64 | 0.542354 | 0.727288 | 0.047178 |
| Maskell 1986 | extraction | 10 | 10 | 1 | 0.722467 | 1 | 0.070801 |
| Peterson 1976 | extraction | 80 | 39 | 0.4875 | 0.381079 | 0.595067 | 0.05459 |
| Roberts 1992 | extraction | 229 | 84 | 0.366812 | 0.307068 | 0.430951 | 0.031603 |
| Roberts 1987 (| extraction | 47 | 18 | 0.382979 | 0.257907 | 0.525734 | 0.068325 |
| Shanson 1987 | extraction | 40 | 13 | 0.325 | 0.200845 | 0.479823 | 0.071169 |
| Shanson 1978 | extraction | 20 | 14 | 0.7 | 0.481027 | 0.854523 | 0.095281 |
| Tomas 2007 | extraction | 53 | 51 | 0.962264 | 0.872457 | 0.98959 | 0.029881 |
| Trivedi 1984 | extraction | 40 | 35 | 0.875 | 0.738879 | 0.945405 | 0.052686 |
| Roberts 1998 | injection | 93 | 49 | 0.526882 | 0.42637 | 0.625261 | 0.050738 |
| Roberts 1997 | intubation | 31 | 3 | 0.096774 | 0.033465 | 0.248999 | 0.054984 |
| | oral surgery | | | | | | |
| Martin 1964 | extraction | 50 | 27 | 0.54 | 0.403989 | 0.670303 | 0.067939 |
| | oral surgery | | | | | | |
| Rajasuo 2004 | extraction | 16 | 14 | 0.875 | 0.639772 | 0.965023 | 0.082974 |
| | oral surgery | | | | | | |
| Roberts 1998 | extraction | 103 | 51 | 0.495146 | 0.400516 | 0.590125 | 0.04837 |
| | oral surgery | | | | | | |
| Tomas 2008 | extraction | 100 | 67 | 0.67 | 0.573053 | 0.754369 | 0.046255 |

| Study | Procedure | N n Rate | | LowerCI | UpperCI | SD | |
|--------------|-----------------------------|----------|----|----------|----------|----------|----------|
| | oral surgery | | | | | | |
| Roberts 1997 | flap elevation oral surgery | 51 | 20 | 0.392157 | 0.270273 | 0.529148 | 0.066041 |
| Roberts 1998 | flap elevation | 51 | 22 | 0.431373 | 0.305012 | 0.567347 | 0.066923 |
| Roberts 1996 | oral surgery | 31 | 22 | 0.431373 | 0.303012 | 0.307347 | 0.000923 |
| Rajasuo 2004 | plate removal | 10 | 6 | 0.6 | 0.312674 | 0.83182 | 0.132437 |
| Burden 2004 | orthodontics | 30 | 4 | 0.133333 | 0.053097 | 0.296813 | 0.062174 |
| Degling 1972 | orthodontics | 10 | 0 | 0 | 0 | 0.277533 | 0.070801 |
| | periodontics | | | | | | |
| Kinane 2005 | probing | 30 | 5 | 0.166667 | 0.073365 | 0.335644 | 0.066909 |

Table 65 Results of Bacteremia Incidence Random Effects Meta-Analysis

| | | Pooled | | |
|-------------------------------------|--------------|--------------------------|----------------------------|----------------|
| Dwooduwo Cwoun | n studies | Incidence of Bacteremia* | 95% Confidence Interval | \mathbf{I}^2 |
| Procedure Group | | | | |
| Brushing | 5 | 21.8% | 5.2 - 38.4% | 96.3% |
| Chewing | 2 | 3.6% | 0 – 10.1% | 39.1% |
| Prophylaxis | 5 | 47.7% | 29.0 - 66.4% | 85.8% |
| Endodontics | 5 | 22.1% | 8.8 – 35.5% | 83.4% |
| Extraction | 4 | 71.4% | 49.4 – 93.4% | 94.1% |
| Interdental Cleaners | 8 | 27.5% | 17.8% - 37.1% | 77.2% |
| Intubation | 6 | 9.3% | 6.1% - 12.5% | 0.0% |
| Oral Surgery - Extraction | 4 | 49.2% | 35.2 – 63.3% | 68.9% |
| Orthodontics | 2 | 18.6% | 0 - 42.5% | 83.9% |
| Periodontics – Scaling/Root Planing | 7 | 46.9% | 24.4 – 69.4% | 92.6% |
| Periodontics – Gingivectomy | 4 | 65.1% | 27.6 – 100% | 95.2% |
| Periodontics – Probing | 2 | 33.4% | 15.5 - 51.3% | 65.3% |
| Sialography | 2 | 10.6% | 0 – 33.4% | 88.5% |
| Suture | 3 | 10.8% | 0.7 - 21% | 43.4% |

 Table 66 Results of Bacteremia Prevalence Random Effects Meta-Analysis

 Procedure Group
 n
 Pooled
 95%
 I²

| | studies | Incidence of Bacteremia* | Confidence Interval | |
|-----------------------------------------------------|---------|--------------------------|------------------------|-------|
| Brushing | 2 | 23.1% | 0 – 52.6% | 92.7% |
| Chewing | 2 | 2.8% | 0 – 11.6% | 46.4% |
| Prophylaxis | 3 | 35.9% | 20.5 – 51.3% | 83.6% |
| Extraction | 13 | 65.3% | 51.8 – 78.8% | 96.1% |
| Oral Surgery – Extraction | 4 | 63.7% | 49.3 – 78.0% | 83.9% |
| Oral Surgery – Flap Elevation | 2 | 41.2% | 31.9 – 50.4% | 0.0% |
| Orthodontics | 2 | 7.1% | 0 - 20.1% | 50.1% |
| Restorative – Drilling | 2 | 14.7% | 1.4 - 28.0% | 84.5% |
| Restorative – Rubber Dam and Matrix Band & Wedge | 2 | 45.6% | 18.2 – 73.0% | 93.9% |

Table 67 Antibiotic Prophylaxis Studies Not Included in Recommendation 1 Network Meta-analysis

| | | | | | | Control (%, | |
|----------------|-------------|-----|----------|-------------------------|--------------------------------|----------------|---------------|
| Procedure | Study | N | Strength | Outcome (specific type) | Active Antibiotic (%, n/N) | n/N) | Results |
| Dental | | | | | | No Treatment | Favors |
| Prophylaxis | Baltch 1982 | 56 | Low | Bacteremia | Penicillin G (10.7%, 3/28) | (60.7%, 17/28) | Penicillin G |
| | Lockhart | | | | | Placebo (18%, | Favors |
| Intubation | 2004 | 100 | High | Bacteremia | Amoxicillin (4%, 2/49) | 9/51) | Amoxicillin |
| | | | - | | Ofloxacin (40%,10/25) | | |
| | | | | | Clindamycin (40%, 10/25) | Placebo (44%, | |
| Oral Surgery | Goker 1992 | 100 | Moderate | Bacteremia | Sultamicillin (36%, 9/25) | 11/25) | No difference |
| | Josefsson | | | | Penicillin (50%, 10/20) | No treatment | |
| Oral Surgery | 1985 | 60 | Moderate | Bacteremia | Erythromycin (55%, 11/20) | (55%, 11/20) | No difference |
| | | | | | IV Cefuroxime (4%, 1/24) IV | | |
| | | | | | Ceftriaxone (0%, 0/21) IV | | |
| Oral Surgery | Katoh 1992 | 62 | Moderate | Bacteremia | Clindamycin (6%, 1/17) | N/A | No difference |
| | | | | | 600mg penicillin (16%, 8/50) | No treatment | Favors |
| Oral Surgery | Martin 1964 | 127 | Moderate | Bacteremia | 300mg penicillin (19%, 5/27) | (54%, 27/50) | Penicillin |
| | Appleman | | | | | Placebo (44%, | |
| Periodontology | 1982 | 31 | Moderate | Bacteremia | Cephalexin (36%, 10/28) | 11/25) | No difference |
| | | | | | Mysteclin plus dental | | |
| | Gutverg | | | | prophylaxis (10%, 5/52) | No Treatment | Favors |
| Periodontology | 1962 | 163 | Moderate | Bacteremia | Mysteclin (5%, 3/57) | (36%, 24/67) | Mysteclin |
| | | | | | Azithromycin (20%, 2/10) | | |
| | Morozumi | | | | Essential Oil Antiseptic (70%, | No Treatment | Favors |
| Periodontology | 2010 | 30 | High | Bacteremia | 7/10) | (90%, 9/10) | Azithromycin |
| | Brennan | | | | | Placebo (20%, | Favors |
| Restorative | 2007 | 100 | Moderate | Bacteremia | Amoxicillin (6%, 3/49) | 10/51) | Amoxicillin |

Table 68 Topical Antimicrobial Prophylaxis Studies Excluded from Recommendation 2 Network Meta-Analysis

| Procedure | Study | N | Strength | Outcome (specific type) | Active Treatment (%, n/N) or (mean, SD) | Control (%, n/N) or (mean, SD) | Results |
|----------------|--------------|-----------|-----------|----------------------------|-----------------------------------------|-----------------------------------|---------------|
| | | | _ | | | No Treatment (34%, 10/29) | |
| Brushing | Madsen 1974 | 29 | Low | Bacteremia Bacteremia | , , , | | No difference |
| | | | | (Aerobic | | Placebo (35.1, | |
| Chewing | Fine 2010 | 22 | Moderate | CFU/ml) | Essential Oil Rinse (8.0, 11.12) | 36.29) | Favors Rinse |
| C | | | | Bacteremia | | , | |
| | | | | (Anaerobic | | Placebo (30.3, | |
| Chewing | Fine 2010 | 22 | Moderate | CFU/ml) | Essential Oil Rinse (6.0, 7.92) | 34.74) | Favors Rinse |
| | | | | | | No Treatment (7%, | |
| Dental Implant | Pineiro 2010 | 50 | Moderate | Bacteremia | Chlorhexidine (0%, 0/20) | 2/30) | No difference |
| D 4 1 | | | | | | | Favors |
| Dental | C1 2007 | 60 | 3.6.1 | D | D :1 I !: (100/ 2/20) | g 1: (200/ 0/20) | Povidone- |
| Prophylaxis | Cherry 2007 | 60 | Moderate | Bacteremia | Povidone-Iodine (10%, 3/30) | Saline (30%, 9/30) | Iodine |
| Dental | | | | Bacteremia (Aerobic | | Dlagaba (29.72 | |
| Prophylaxis | Fine 1996 | 18 | Moderate | CFU/ml) | Essential Oil Rinse (4.67, 2.14) | Placebo (38.72, 17.82) | Favors Rinse |
| Tiophylaxis | 1411C 1990 | 10 | Moderate | Bacteremia | Essential Off Killse (4.07, 2.14) | 17.02) | ravois Kinse |
| Dental | | | | (Anaerobic | | Placebo (14.89, | |
| Prophylaxis | Fine 1997 | 18 | Moderate | CFU/ml) | Essential Oil Rinse (1.61, 1.54) | 7.86) | Favors Rinse |
| Topilyiums | 1 1110 1777 | 10 | Wisaciate | Cr C/III) | Essential off Tampe (1.01, 1.01) | 7.00) | Favors |
| | | | | | Chlorhexidine (45%, 18/40) | | Povidone- |
| Injection | Rahn 1995 | 120 | Moderate | Bacteremia | Povidone-Iodine (28%, 11/40) | Water (53%, 21/40) | Iodine |
| Inter-dental | | | | | , | No Treatment | |
| Cleaning | Madsen 1974 | 29 | Low | Bacteremia | Chlorhexidine (24%, 7/29) | (34%, 10/29) | No difference |
| _ | Fourrier | | | | | Placebo (18%, | |
| Intubation | 2005 | 228 | High | Bacteremia | Antiseptic Rinse (18%, 20/114) | 21/114) | No difference |
| | Huffman | | | | Cetylpyridinium Chloride (83%, | | |
| Oral surgery | 1974 | 25 | Low | Bacteremia | 10/12) | Saline (70%, 9/13) | No difference |
| | | | | | | No Treatment (7%, | |
| Orthodontistry | Erverdi 2001 | 150 | Low | Bacteremia | Chlorhexidine (3%, 2/80) | 5/70) | No difference |
| | _ | | | | | | Favors |
| | Brenman | | | | D. 1.1 V. 11 (200) 575 5 | Placebo (58%, | Povidone- |
| Periodontology | 1974 | 52 | Moderate | Bacteremia | Povidone-Iodine (23%, 6/26) | 15/26) | Iodine |

| Procedure | Study | N | Strength | Outcome (specific type) | Active Treatment (%, n/N) or (mean, SD) | Control (%, n/N) or (mean, SD) | Results |
|----------------|--------------|----|-------------|-------------------------|-----------------------------------------------|-----------------------------------|---------------|
| | State | | ~ v. v. gv. | (specific type) | Chlorhexidine (20%, 2/10) | No Treatment | |
| Periodontology | Lofthus 1991 | 30 | Moderate | Bacteremia | Water (40%, 4/10) Azithromycin (20%, 2/10) | (30%, 3/10) | No difference |
| | Morozumi | | | | Essential Oil Antiseptic (70%, | No Treatment | Favors |
| Periodontology | 2010 | 30 | High | Bacteremia | 7/10) | (90%, 9/10) | Azithromycin |
| | | | - | | Chlorhexidine (27%, 4/15) | No Treatment | |
| Periodontology | Waki 1990 | 54 | Moderate | Bacteremia | Water (15%, 2/13) | (13%, 2/15) | No difference |
| • | | | | | | No Treatment (9%, | |
| Suture | Brown 1998 | 55 | Moderate | Bacteremia | Chlorhexidine (15%, 4/27) | 2/22) | No difference |

APPENDIX XII

QUALITY AND APPLICABILITY TABLES FOR INCLUDED STUDIES

Table 69 APPRAISE Table of Prognostic Studies for Recommendation 1, Direct Evidence

- •: Domain free of flaws
- o: Domain flaws present

| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
|--------------|-------------|-------|----------|-------------------|-------|----------|----------|----------|----------|---------------|
| Berbari 2010 | • | • | 0 | • | • | Moderate | • | 0 | • | Moderate |

Table 70 APPRAISE Table of Treatment Studies for Recommendation 1, Dental Prophylaxis

•: Domain free of flaws Compliance and Adherence Intervention and Expertise o: Domain flaws present Group Comparability Treatment Integrity **Group Assignment** Investigator Bias Measurement **Participants** Hypothesis Blinding Analysis Study **Outcome** Quality **Applicability** Baltch 1982 Moderate Bacteremia 0 0 0 0 Low 0

Table 71 APPRAISE Table of Treatment Studies for Recommendation 1, Intubation

| o: Domain flaws | present | | | | | | | | | | rtise | ence. | | |
|------------------|------------|------------|----------------------|----------|---------------|-----------|-------------|--------------|---------|--------------|--------------|------------|---------|---------------|
| | | | nt | | bility | rity | | | | | Expertise | Adherenc | | |
| | | | ssignment | | omparability | Integrity | ınt | r Bias | | S | n and | e and | | |
| | | Hypothesis | \blacktriangleleft | ing | \mathcal{C} | Treatment | Measurement | Investigator | | Participants | Intervention | Compliance | /sis | |
| | | [ypo | Group | Blinding | Group | reat | Ieası | nvest | | artic | nter | omp | nalysis | |
| Study | Outcome | | <u> </u> | <u> </u> | <u> </u> | | | 7 | Quality | <u> </u> | T | | | Applicability |
| Lockhart 2004 | Bacteremia | • | • | • | • | • | • | • | High | • | Ο | • | • | Moderate |

Table 72 APPRAISE Table of Treatment Studies for Recommendation 1, Oral Surgery

| o: Domain flaws | present Outcome | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | Quality | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | Applicability |
|-------------------|-----------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| Goker 1992 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |
| Josefsson 1985 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |
| Katoh 1992 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |
| Martin 1964 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |

Table 73 APPRAISE Table of Treatment Studies for Recommendation 1, Periodontology

| o: Domain flaws p | Outcome | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | Quality | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | Applicability |
|-------------------|------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| Appleman 1981 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |
| Gutverg 1962 | Bacteremia | • | 0 | 0 | 0 | • | • | • | Moderate | • | 0 | • | • | Moderate |
| Morozumi 2010 | Bacteremia | • | • | 0 | • | • | • | • | High | • | 0 | • | • | Moderate |

Table 74 APPRAISE Table of Treatment Studies for Recommendation 1, Restorative Procedure

•: Domain free of flaws

| o: Domain flaws | present | | | | | | | | | | rtise | dherence | | |
|-----------------|------------|------------|-----------|----------|--------------|--------------|-------------|--------------|----------|--------------|--------------|------------|---------|---------------|
| | | | ent | | ability | grity | | S. | | | d Expertise | A | | |
| | | .s. | ssignment | | omparability | ıt Integrity | nent | tor Bias | | nts | ion and | ice and | | |
| | | Hypothesis | roup As | Blinding | Group Co | Treatment | Measurement | Investigator | | Participants | Intervention | Compliance | nalysis | |
| Study | Outcome | Hy | Gr | Bli | Gr | Tr | Me | In | Quality | Pa | Int | ပိ | An | Applicability |
| Brennan 2007 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |

Table 75 APPRAISE Table of Treatment Studies for Recommendation 1, Extraction

| o: Domain flaws pr | Pesent | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | Quality | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | Applicability |
|--------------------|------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| Study | Outcome | | | | | | | | Quality | | | | | Аррисавииу |
| Aitken 1995 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |
| Cannell 1991 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |

| o: Domain flaws p | present | sis | Group Assignment | | Group Comparability | Treatment Integrity | ement | Investigator Bias | | ants | Intervention and Expertise | Compliance and Adherence | 70 | |
|-------------------|------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| Study | Outcome | Hypothesis | Group 4 | Blinding | Group (| Treatmo | Measurement | Investig | Quality | Participants | Interver | Complia | Analysis | Applicability |
| Casolari 1989 | Bacteremia | • | 0 | 0 | 0 | • | • | • | Moderate | • | 0 | 0 | 0 | Moderate |
| Coulter 1990 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |
| DeVries 1972 | Bacteremia | • | 0 | • | 0 | 0 | • | 0 | Low | • | 0 | • | 0 | Moderate |
| Dios 2006 | Bacteremia | • | 0 | • | 0 | • | • | • | Moderate | • | 0 | • | • | Moderate |
| Hall 1993 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |
| Hall 1996 | Bacteremia | • | 0 | • | 0 | • | • | • | Moderate | • | 0 | • | 0 | Moderate |
| Hall 1996 | Bacteremia | • | 0 | • | 0 | • | • | • | Moderate | • | 0 | • | 0 | Moderate |
| Head 1984 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |
| Jokinen 1970 | Bacteremia | • | 0 | 0 | 0 | • | • | • | Moderate | • | 0 | 0 | 0 | Moderate |
| Khairat 1966 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |

| o: Domain flaws p | present | sis | Group Assignment | | Group Comparability | Treatment Integrity | ment | Investigator Bias | | ants | Intervention and Expertise | Compliance and Adherence | | |
|-------------------|------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| Study | Outcome | Hypothesis | Group A | Blinding | Group (| Treatme | Measurement | Investig | Quality | Participants | Interven | Complia | Analysis | Applicability |
| Lockhart 2004 | Bacteremia | • | • | • | • | • | • | • | High | • | • | • | • | Moderate |
| Lockhart 2008 | Bacteremia | • | • | • | • | • | • | • | High | • | • | • | • | Moderate |
| Maskell 1986 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |
| Roberts 1987 | Bacteremia | • | 0 | 0 | • | • | • | 0 | Moderate | • | 0 | 0 | • | Moderate |
| Shanson 1978 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |
| Shanson 1985 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |
| Shanson 1987 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |
| Vergis 2001 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | 0 | Moderate |
| Wahlmann 1999 | Bacteremia | • | 0 | 0 | 0 | • | • | 0 | Low | • | 0 | 0 | 0 | Moderate |

Table 76 APPRAISE Table of Treatment Studies for Recommendation 2, Brushing

| o: Domain flaws | Present Outcome | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | Quality | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | Applicability |
|-----------------|-----------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|---------|--------------|----------------------------|--------------------------|----------|---------------|
| Madsen 1974 | Bacteremia | • | 0 | 0 | 0 | • | • | 0 | low | • | 0 | • | • | Moderate |

Table 77 APPRAISE Table of Treatment Studies for Recommendation 2, Chewing

| •: Domain free o | of flaws | | | | | | | | | | ė | ce | | |
|------------------|------------|------------|-----------|----------|-----------|----------------------|-------------|--------------|----------|--------------|--------------|------------|---------|---------------|
| o: Domain flaws | present | | | | | | | | | | Expertise | dherence | | |
| | | | . | | ility | ţ | | | | | (xp) | dhe | | |
| | | | nen | | parabilit | Integri | | Bias | | | and I | y pı | | |
| | | | ssignment | | npa | Inte | ent | | | Ñ | | e and | | |
| | | esis | Assi | 5,0 | Com | ent | em. | gato | | bant | ntio | anc | S | |
| | | Hypothesis | dno | Blinding | dno | Treatment | Measurement | Investigator | | Participants | Intervention | Compliance | nalysis | |
| Study | Outcome | Hyl | Gre | Blir | Group | Tre | Me | Inv | Quality | Par | Inte | C_{0} | Ans | Applicability |
| | | | | | | | | | ~ • | | | | | |
| Fine 2010 | Bacteremia | • | 0 | • | • | • | | 0 | Moderate | • | 0 | | | Moderate |

Table 78 APPRAISE Table of Treatment Studies for Recommendation 2, Dental Implant

| •: Domain free of | f flaws | | | | | | | | | | ė | ce | | |
|-------------------|------------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| o: Domain flaws | present Outcome | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | Quality | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | Applicability |
| Pineiro 2010 | Bacteremia | • | 0 | 0 | 0 | • | • | • | Moderate | • | 0 | • | • | Moderate |

Table 79 APPRAISE Table of Treatment Studies for Recommendation 2, Dental Prophylaxis

•: Domain free of flaws

| o: Domain flaws p | resent | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | |
|-------------------|------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| Study | Outcome | H | 5 | B | g | | Σ | II | Quality | P | Ir | Ö | V | Applicability |
| Cherry 2007 | Bacteremia | • | 0 | 0 | 0 | • | • | • | Moderate | • | 0 | • | • | Moderate |
| Fine 1996 | Bacteremia | • | • | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |

Table 80 APPRAISE Table of Treatment Studies for Recommendation 2, Injection

| o: Domain flaws p | Outcome | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | Quality | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | Applicability |
|-------------------|------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| Rahn 1995 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |

Table 81 APPRAISE Table of Treatment Studies for Recommendation 2, Inter-detal Cleaning

•: Domain free of flaws

| o: Domain flaws Study | present Outcome | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | Quality | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | Applicability |
|-----------------------|------------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|---------|--------------|----------------------------|--------------------------|----------|---------------|
| Madsen 1974 | Bacteremia | • | 0 | 0 | 0 | • | • | 0 | low | • | 0 | • | • | Moderate |

Table 82 APPRAISE Table of Treatment Studies for Recommendation 2, Intubation

| o: Domain flaws pr | resent Outcome | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | Quality | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | Applicability |
|--------------------|-----------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|---------|--------------|----------------------------|--------------------------|----------|---------------|
| Fourrier 2005 | Bacteremia | • | • | • | • | • | • | • | High | • | 0 | • | • | Moderate |

Table 83 APPRAISE Table of Treatment Studies for Recommendation 2, Oral Surgery

| •: Domain free of | flaws | | | | | | | | | | | | | |
|-------------------|------------|------------|-----------|----------|------------|-----------|-------------|--------------|---------|----------|----------|--------|---------|---------------|
| o: Domain flaws | present | | | | | | | | | | rtise | nce | | |
| | | | | | > | | | | | | <u> </u> | ere | | |
| | | | ıţ | | parability | ity | | | | | Exp | Adh | | |
| | | | ssignment | | ırak | Integrity | | Bias | | | and | pu | | |
| | | 70 | ign | | ompa | | ent | | | t | _ | e an | | |
| | | esis | Ass | Þi | رة ر | ent | em. | gate | | pan | ention | anc | S | |
| | | Hypothesis | dn | Blinding | dn | Treatment | Measurement | Investigator | | ticij | × | Compli | nalysis | |
| | | Hyp | Group | 3lin | Group | [ře | Mea | nve | | Part | Inter | Çon | √na | |
| Study | Outcome | | | <u> </u> | | | | Ι | Quality | | | | 7 | Applicability |
| Huffman | Dastanania | | \sim | | \sim | | | \sim | T | | \circ | | | Madausta |

Low

Moderate

Table 84 APPRAISE Table of Treatment Studies for Recommendation 2, Orthodontistry

0

Bacteremia

1974

•: Domain free of flaws Compliance and Adherence o: Domain flaws present Group Comparability Treatment Integrity **Group Assignment** Investigator Bias Measurement **Participants** Hypothesis Blinding Analysis Study **Applicability Outcome** Quality Erverdi 2001 Bacteremia Moderate 0 Low 0

Table 85 APPRAISE Table of Treatment Studies for Recommendation 2, Periodontology

| o: Domain flaws | present | Hypothesis | up Assignment | Blinding | up Comparability | Treatment Integrity | Measurement | Investigator Bias | | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | |
|------------------|------------|------------|---------------|----------|------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| Study | Outcome | Hyl | Group | Blir | Group | Tre | Me | Inv | Quality | Par | Inte | Cor | Ans | Applicability |
| Brenman 1974 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |
| Lofthus 1991 | Bacteremia | • | 0 | 0 | 0 | • | • | • | Moderate | • | 0 | • | • | Moderate |
| Morozumi 2010 | Bacteremia | • | • | 0 | • | • | • | • | High | • | 0 | • | • | Moderate |
| Waki 1990 | Bacteremia | • | 0 | • | 0 | • | • | • | Moderate | • | 0 | • | • | Moderate |

Table 86 APPRAISE Table of Treatment Studies for Recommendation 2, Suture

•: Domain free of flaws

| o: Domain flaws J | present | | | | | | | | | | rtise | dherence | | |
|-------------------|------------|------------|-----------|----------|--------------|--------------|-------------|--------------|----------|--------------|--------------|------------|---------|---------------|
| | | | ent | | ability | grity | | S | | | d Expertise | Ā | | |
| | | is | ssignment | | omparability | nt Integrity | nent | tor Bias | | nts | ion and | ıce and | | |
| | | Hypothesis | roup As | Blinding | Group C | Treatment | Measurement | Investigator | | Participants | Intervention | Compliance | nalysis | |
| Study | Outcome | Hy | Gr | Bli | Gr | Tr | M | Im | Quality | Pa | Int | ప | An | Applicability |
| Brown 1998 | Bacteremia | • | 0 | • | 0 | • | • | • | Moderate | • | 0 | • | • | Moderate |

Table 87 APPRAISE Table of Treatment Studies for Recommendation 2, Tooth Extraction

| o: Domain flaws | present | | ment | | Comparability | Integrity | | Bias | | | and Expertise | and Adherence | | |
|------------------|------------|------------|------------------|----------|---------------|---------------|-------------|----------------|----------|--------------|----------------|---------------|----------|---------------|
| Study | Outcome | Hypothesis | Group Assignment | Blinding | Group Compa | Treatment Int | Measurement | Investigator B | Quality | Participants | Intervention a | Compliance an | Analysis | Applicability |
| Casolari 1989 | Bacteremia | • | 0 | 0 | 0 | • | • | • | Moderate | • | 0 | • | • | Moderate |
| Cutcher 1971 | Bacteremia | • | 0 | 0 | 0 | • | • | 0 | Low | • | 0 | • | • | Moderate |

| o: Domain flaws p | present | | | | ĸ | | | | | | pertise | nerence | | |
|--------------------|------------|------------|------------------|----------|---------------------|---------------------|-------------|-------------------|----------|--------------|----------------------------|--------------------------|----------|---------------|
| Study | Outcome | Hypothesis | Group Assignment | Blinding | Group Comparability | Treatment Integrity | Measurement | Investigator Bias | Quality | Participants | Intervention and Expertise | Compliance and Adherence | Analysis | Applicability |
| Francis 1973 | Bacteremia | • | 0 | 0 | 0 | • | • | 0 | Low | • | 0 | • | • | Moderate |
| Jokinen 1970 | Bacteremia | • | 0 | 0 | 0 | • | • | • | Moderate | • | 0 | • | • | Moderate |
| Jones 1970 | Bacteremia | • | 0 | 0 | 0 | • | • | 0 | Low | • | 0 | • | • | Moderate |
| Lockhart 1996 | Bacteremia | • | • | • | • | • | • | 0 | High | • | 0 | • | • | Moderate |
| MacFarlane 1984 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |
| Nasif 1977 | Bacteremia | • | 0 | 0 | 0 | • | • | 0 | Low | • | 0 | • | • | Moderate |
| Rahn 1995 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |
| Scopp 1971 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |
| Sweet 1978 | Bacteremia | • | 0 | • | 0 | • | • | 0 | Moderate | • | 0 | • | • | Moderate |
| Tomas 2007 | Bacteremia | • | 0 | 0 | • | • | • | • | Moderate | • | 0 | • | • | Moderate |

| o: Domain flaws | present | | | | | | | | | | rtise | rence | | |
|-----------------|------------|------------|-----------------|----------|---------------------|-----------|-------------|-----------|---------|--------------|--------------|------------|---------|---------------|
| | | | ent | | billity | rity | | ø | | | l Expertise | Adher | | |
| | | 70 | ssignme | | omparabili | Integri | ent | or Bias | | ts | on and | e and | | |
| | | Hypothesis | \triangleleft | Blinding | $\ddot{\mathbf{C}}$ | Treatment | Measurement | estigator | | Participants | Intervention | Compliance | nalysis | |
| - | | lype | Group | Slinc | Group | lrea | Jea | Inve | | arti | nter | Jom | \nal | |
| Study | Outcome | H | | | | | | | Quality | | | | 4 | Applicability |
| Yamalik 1992 | Bacteremia | • | 0 | 0 | 0 | • | • | 0 | Low | • | 0 | • | • | Moderate |

Table 88 APPRAISE Table of Prognostic Studies for Recommendation 3, Brushing

•: Domain free of flaws

| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
|------------------|-------------|-------|----------|-------------------|-------|---------|----------|----------|----------|---------------|
| Ashare 2009 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Bhanji 2002 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Forner 2006 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Lockhart 2009 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Silver 1977 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |

Table 89 APPRAISE Table of Prognostic Studies for Recommendation 3, Chewing

- •: Domain free of flaws
- o: Domain flaws present

| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
|-------------|-------------|-------|----------|-------------------|-------|----------|----------|----------|----------|---------------|
| Forner 2006 | • | 0 | 0 | 0 | • | Very Low | • | 0 | 0 | Moderate |

Table 90 APPRAISE Table of Prognostic Studies for Recommendation 3, Dental Prophylaxis

- •: Domain free of flaws
- o: Domain flaws present

| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
|--------------|-------------|-------|----------|-------------------|-------|---------|----------|----------|----------|---------------|
| Cherry 2007 | • | 0 | 0 | • | • | Low | • | 0 | 0 | Moderate |
| De Leo 1974 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Forner 2006 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Trivedi 1984 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |

Table 91 APPRAISE Table of Prognostic Studies for Recommendation 3, Inter-dental Cleaning

- •: Domain free of flaws
- o: Domain flaws present

| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
|--------------------|-------------|-------|----------|-------------------|-------|---------|----------|----------|----------|---------------|
| Crasta 2009 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Lineberger 1973 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |

Table 92 APPRAISE Table of Prognostic Studies for Recommendation 3, Intubation

- •: Domain free of flaws
- o: Domain flaws present

| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
|-------------|-------------|-------|----------|-------------------|-------|---------|----------|----------|----------|---------------|
| Valdes 2008 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |

Table 93 APPRAISE Table of Prognostic Studies for Recommendation 3, Oral Surgery

- •: Domain free of flaws
- o: Domain flaws present

| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
|-------------------|-------------|-------|----------|-------------------|-------|---------|----------|----------|----------|---------------|
| Enabulele 2008 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Roberts 1998 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Takai 2005 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Tomas 2008 | • | • | 0 | 0 | • | Low | • | Ο | Ο | Moderate |

Table 94 APPRAISE Table of Prognostic Studies for Recommendation 3, Periodontology

- •: Domain free of flaws
- o: Domain flaws present

| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
|-----------|-------------|-------|----------|-------------------|-------|---------|----------|----------|----------|---------------|
| Daly 1997 | • | 0 | 0 | 0 | • | Low | • | 0 | 0 | Moderate |
| Daly 2001 | • | • | 0 | 0 | • | Low | • | 0 | 0 | Moderate |

Table 95 APPRAISE Table of Prognostic Studies for Recommendation 3, Restorative Procedure

- •: Domain free of flaws
- o: Domain flaws present

| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
|-----------------|-------------|-------|----------|-------------------|-------|----------|----------|----------|----------|---------------|
| Brennan 2007 | • | • | 0 | 0 | 0 | Very Low | • | 0 | 0 | Moderate |

Table 96 APPRAISE Table of Prognostic Studies for Recommendation 3, Tooth Extraction

- •: Domain free of flaws
- o: Domain flaws present

| | 9 | | | or Bias | | | | | | |
|-------------------|-------------|-------|----------|-------------------|-------|----------|----------|----------|----------|---------------|
| Study | Prospective | Power | Analysis | Investigator Bias | Model | Quality | Patients | Analysis | Outcomes | Applicability |
| Barbosa 2010 | • | • | 0 | 0 | • | Moderate | • | 0 | 0 | Moderate |
| Coulter 1990 | • | • | 0 | 0 | • | Moderate | • | 0 | 0 | Moderate |
| Enabulele 2008 | • | • | 0 | 0 | • | Moderate | • | 0 | 0 | Moderate |
| Lockhart 1996 | • | • | 0 | 0 | • | Moderate | • | 0 | 0 | Moderate |
| Lockhart 2009 | • | • | 0 | 0 | • | Moderate | • | 0 | 0 | Moderate |
| Okabe 1995 | • | • | 0 | 0 | • | Moderate | • | 0 | 0 | Moderate |
| Roberts 1998 | • | • | 0 | 0 | • | Moderate | • | 0 | 0 | Moderate |
| Wahlmann 1999 | • | • | 0 | 0 | • | Moderate | • | 0 | 0 | Moderate |

Table 97 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Brushing

| V 2 0 222W112 22W | vo present | | | | | 3ias | | | | | |
|-------------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
| Bhanji 2002 | Bacteremia | Ι | • | • | • | • | High | • | 0 | 0 | Moderate |
| Forner 2006 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Lockhart 2008 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Lucas 2000 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Sconyers 1979 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Sconyers 1973 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Silver 1979 | Bacteremia | P | • | • | • | • | High | • | 0 | 0 | Moderate |

Table 98 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Brushing

•: Domain free of flaws

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|-----------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Degling 1972 | Bacteremia | P | • | • | • | • | High | • | 0 | 0 | Moderate |
| Forner 2006 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Murphy 2006 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Trivedi 1984 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

Table 99 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Cleft Palate

•: Domain free of flaws

o: Domain flaws present

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|-----------------|------------|--------------------------|---------|-------------|--------------|-------------------|---------|----------|----------|--------------|---------------|
| Marzoni 1983 | Bacteremia | P | • | • | 0 | 0 | Low | • | 0 | 0 | Moderate |

Table 100 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Dental Implant

•: Domain free of flaws

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|--------------|------------|--------------------------|---------|-------------|--------------|-------------------|---------|----------|----------|---------------------|---------------|
| Pineiro 2010 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |

Table 101 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Dental Prophylaxis

| | • | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | | Analysis | Outcomes | Participants | |
|------------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Study | Outcome | In P | Ō | M | Pa | In | Quality | Aı | Ō | Pa | Applicability |
| Cherry 2007 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| De Leo 1974 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Forner 2006 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Forner 2006 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Heimdahl 1990 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Lucas 1999 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Trivedi 1984 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Winslow 1960 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

Table 102 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Endodontic

| | vs present | | | | | as | | | | | |
|---------------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
| Baumgartner 1977 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Baumgartner 1976 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Bender 1963 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Debelian 1995 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Heimdahl 1990 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Savarrio 2005 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |

Table 103 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Injections

•: Domain free of flaws

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|-----------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Roberts 1998 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Rahn 1995 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

Table 104 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Inter-dental Cleaning

| o. Domain maws | present | ą | | ent | ıts | or Bias | | | | ıts | |
|------------------------------------------------------|------------|--------------------------|---------|-------------|--------------|-------------------|-----------|----------|----------|--------------|---------------|
| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
| Berger 1974 | Bacteremia | I | | | | 0 | Moderate | , | 0 | 0 | Moderate |
| Deiger 1974 | Bacterenna | 1 | | | | O | Wioderate | | O | O | Wioderate |
| Crasta 2009 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Felix 1971 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Lineberger 1973 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Ramadan 1975 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Romans 1971 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| The American Academy of Periodontology 1972 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Wank 1976 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

Table 105 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Intubation

•: Domain free of flaws

| | o present | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | | Analysis | Outcomes | Participants | |
|-----------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Study | Outcome | Inci /Pre | Out | Mea | Part | Inve | Quality | Ana | Out | Part | Applicability |
| Ali 1992 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Berry 1973 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Dinner 1987 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Hansen 1989 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Oncag 2005 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Roberts 1997 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Valdes 2008 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |

Table 106 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Oral Surgery

•: Domain free of flaws

| | • | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | | Analysis | Outcomes | Participants | |
|-------------------|------------|--------------------------|----------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Study | Outcome | Ir A | <u> </u> | 2 | <u>P</u> | I I | Quality | ▼ | <u> </u> | P | Applicability |
| Enabulele 2008 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Flood 1990 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Heimdahl 1990 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Josefsson 1985 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Martin 1964 | Bacteremia | P | • | • | • | 0 | Moderate | 0 | 0 | 0 | Low |
| Rajasuo 2004 | Bacteremia | P | • | 0 | • | • | Moderate | • | 0 | 0 | Moderate |
| Rajasuo 2004 | Bacteremia | P | • | 0 | • | • | Moderate | • | 0 | 0 | Moderate |
| Roberts 1997 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Roberts 1998 | Bacteremia | P | • | 0 | • | 0 | Low | • | 0 | 0 | Moderate |
| Takai 2005 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Tomas 2008 | Bacteremia | P | • | 0 | • | 0 | Low | • | 0 | 0 | Moderate |

Table 107 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Orthodontic

•: Domain free of flaws

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|-----------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Burden 2004 | Bacteremia | P | • | • | • | • | High | • | 0 | 0 | Moderate |
| Degling 1972 | Bacteremia | P | • | • | • | • | High | • | 0 | 0 | Moderate |
| Erverdi 1999 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Gürel 2009 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

Table 108 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Periodontology

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|--------------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Bender 1963 | Bacteremia | I | _ | | | | High | | 0 | 0 | Moderate |
| Casolari 1989 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Conner 1967 | Bacteremia | P | • | • | • | • | High | • | 0 | 0 | Moderate |
| Daly 1997 | Bacteremia | Ι | • | • | • | • | High | • | 0 | 0 | Moderate |
| Daly 2001 | Bacteremia | I | • | • | 0 | • | Moderate | • | 0 | 0 | Moderate |
| Gutverg 1962 | Bacteremia | P | • | • | • | • | High | • | 0 | 0 | Moderate |
| Kinane 2005 | Bacteremia | P | • | 0 | • | 0 | Moderate | 0 | 0 | 0 | Low |
| Lafaurie 2007 | Bacteremia | I | • | • | • | 0 | High | 0 | 0 | 0 | Moderate |
| Lineberger 1973 | Bacteremia | I | • | • | 0 | 0 | Moderate | 0 | 0 | 0 | Moderate |
| Lofthus 1991 | Bacteremia | I | • | • | • | 0 | High | 0 | 0 | 0 | Moderate |
| Lucartorto 1992 | Bacteremia | I | • | • | • | 0 | High | 0 | 0 | 0 | Moderate |

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|------------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| | Outcome | | | | | | Quanty | | | | Applicability |
| Morozumi 2010 | Bacteremia | I | • | • | • | 0 | High | 0 | 0 | 0 | Moderate |
| Rogosa 1960 | Bacteremia | I | • | • | 0 | 0 | Moderate | 0 | 0 | 0 | Moderate |
| Wada 1968 | Bacteremia | I | • | • | 0 | 0 | Moderate | 0 | 0 | 0 | Moderate |
| Waki 1990 | Bacteremia | I | • | • | • | 0 | High | 0 | 0 | 0 | Moderate |

Table 109 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Restorative Procedure

•: Domain free of flaws

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|-----------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Oncag 2006 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Roberts 2000 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | • | Moderate |
| Sonbol 2009 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

Table 110 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Sialography

•: Domain free of flaws

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Lamey 1985 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Nixon 2009 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

Table 111 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Suture

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|-----------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Brown 1998 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Giglio 1992 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| King 1988 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Wampole 1978 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

Table 112 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Teething

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|-----------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Soliman 1977 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

Table 113 APPRAISE Table of Incidence/Prevalence Studies for Control/Baseline, Tooth Extraction

| G. I | 0 | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | 0 P | Analysis | Outcomes | Participants | A 71 1 171 |
|------------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Study | Outcome | | | | | | Quality | 7 | | | Applicability |
| Barbosa 2010 | Bacteremia | P | • | 0 | • | • | Moderate | • | 0 | 0 | Moderate |
| Bender 1963 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Casolari 1989 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Coulter 1990 | Bacteremia | P | • | • | • | • | High | • | 0 | 0 | Moderate |
| Crawford 1974 | Bacteremia | P | • | 0 | • | • | Moderate | • | 0 | 0 | Moderate |
| DeVries 1972 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Francis 1973 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Heimdahl 1990 | Bacteremia | I | • | • | • | • | High | • | 0 | 0 | Moderate |
| Khairat 1966 | Bacteremia | I | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Maskell 1896 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

| Study | Outcome | Incidence /Prevalence | Outcome | Measurement | Participants | Investigator Bias | Quality | Analysis | Outcomes | Participants | Applicability |
|------------------|------------|--------------------------|---------|-------------|--------------|-------------------|----------|----------|----------|--------------|---------------|
| Peterson 1976 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Roberts 1992 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Shanson 1978 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Shanson 1987 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |
| Tomas 2007 | Bacteremia | P | • | 0 | • | • | Moderate | • | 0 | 0 | Moderate |
| Tomas 2007 | Bacteremia | P | • | 0 | • | • | Moderate | • | 0 | 0 | Moderate |
| Trivedi 1984 | Bacteremia | P | • | • | • | 0 | Moderate | • | 0 | 0 | Moderate |

APPENDIX XIII CONFLICT OF INTEREST

All members of the AAOS work group disclosed any conflicts of interest prior to the development of the recommendations for this guideline. Conflicts of interest are disclosed with the American Academy of Orthopaedic Surgeons via a private on-line reporting database and also verbally at the recommendation approval meeting.

Disclosure Items: (n) = Respondent answered 'No' to all items indicating no conflicts. 1= Royalties from a company or supplier; 2= Speakers bureau/paid presentations for a company or supplier; 3A= Paid employee for a company or supplier; 3B= Paid consultant for a company or supplier; 3C= Unpaid consultant for a company or supplier; 4= Stock or stock options in a company or supplier; 5= Research support from a company or supplier as a PI; 6= Other financial or material support from a company or supplier; 7= Royalties, financial or material support from publishers; 8= Medical/Orthopaedic publications editorial/governing board; 9= Board member/committee appointments for a society.

William C. Watters, III, MD, Work Group Co-Chair: 1 (Stryker); 3B (Palladian; Stryker); 4 (Intrinsic Orthopedics); 8 (Official Disability Guidelines; Spine; The Spine Journal); 9 (American Board of Spine Surgery; North American Spine Society); Submitted on: 08/11/2011.

Michael P. Rethman, DDS, MS, Work Group Co-Chair: 3B (Colgate-Palmolive); 4 (Colgate-Palmolive; Pfizer); 9 (American Dental Association Foundation); Submitted on: 02/05/2013.

Elliott Abt, DDS: (n) Submitted on: 10/19/2011.

Harry C. Futrell, DMD: (n) Submitted on: 10/04/2011.

Stephen O. Glenn, DDS: 8 (Key/ Alliance of the American Dental Association); 9 (American Dental Association); Submitted on: 10/19/2011.

John Hellstein, DDS, MS: 9 (American Academy of Oral and Maxillofacial Pathology; American Board of Oral and Maxillofacial Pathology; American Dental Association Council on Scientific Affairs; Basal Cell Carcinoma Nevus Syndrome Life Support Network); Submitted on: 10/04/2011.

Mark J. Steinberg, DDS, MD: 9 (American Association of Oral and Maxillofacial Surgeons); Submitted on: 04/19/2011.

Richard Parker Evans, MD: 2 (Johnson & Johnson; Smith & Nephew)

Michael J. Goldberg, MD: 8 (Journal Children's Orthopaedics; Journal of Pediatric Orthopedics); 9 (AAOS); Submitted on: 04/27/2011.

Calin Stefan Moucha, MD: 2 (3M) 4 (Auxillium); 9 (AAOS); Submitted on: 10/02/2011.

Richard J. O'Donnell, MD: 9 (National Comprehensive Cancer Network; Northern California Chapter, Western Orthopaedic Association; Orthopaedic Surgical Osseointegration Society; Sarcoma Alliance); Submitted on: 10/04/2011.

Paul A. Anderson, MD: 1 (Pioneer; Stryker); 3B (Aesculap/B.Braun); 3C (Expanding Orthopedics; SI Bone; Spatatec; Titan Surgical); 4 (Pioneer Surgical; SI Bone; Spartec; Titan Surgical); 8 (Clinical Orthopaedics and Related Research; Journal of Bone and Joint Surgery - American; Journal of Orthopaedics and Traumatology; Journal of Spinal Disorders; Neurosurgery; Spine; Spine Arthropalsty Journal); 9 (American Academy of Orthopaedic Surgeons, American Society for Testing and Materials; North American Spine Society; Spine Arthroplasty Society; Spine Section of American Association of Neurological Surgeons/Congress of Neurological Surgeons); Submitted on: 04/07/2011.

John E. O'Toole, MD: 1 (Globus Medical); 3B (Globus Medical; Pioneer Surgical); 3C (Medtronic); Submitted on: 10/19/2011.

David J. Kolessar, MD: 4 (Zimmer); Submitted on: 04/07/2011.

Karen C. Carroll, MD, FCAP: 7 (ASM Press; McGraw-Hill); 8 (Infectious Diseases in Clinical Practice; Journal Clinical Microbiology/ASM Press); Submitted on: 10/05/2011.

Kevin Garvin, MD: 1 (Biomet); 8 (Wolters Kluwer Health - Lippincott Williams & Wilkins); 9 (AAOS; AAOS; American Orthopaedic Association; American Orthopaedic Association); Submitted on: 09/21/2011.

Douglas R. Osmon, MD: 9 (Musculoskeletal infection society); Submitted on: 10/05/2011.

Anthony Rinella, MD: (n); Submitted on: 10/05/2011.

Angela Hewlett, MD, MS: 9 (Society for Healthcare Epidemiology of America); Submitted on: 10/04/2011.

William Robert Martin, III, MD: 9 (National Board of Medical Examiners); Submitted on: 03/12/2010.

Deborah S. Cummins, PhD: (n); Submitted on 11/15/2012.

Sharon Song, PhD: (n); Submitted on 1/28/2013.

Patrick Sluka, MPH: (n); Submitted on 10/19/2011.

Kevin Boyer, MPH: (n); Submitted on 03/05/2012.

Anne Woznica, MLIS: (n); Submitted on 10/03/2012.

Helen Ristic, PhD: (n); Submitted on 01/15/2013.

Nicholas Buck Hanson, MPH: (n); Submitted on 01/14/2013.

REFERENCES

- (1) Kurtz S, Ong K, Lau E, Mowat F, Halpern M. Projections of primary and revision hip and knee arthroplasty in the United States from 2005 to 2030. *J Bone Joint Surg Am* 2007;89(4):780-785.
- (2) Della Valle CJ, Zuckerman JD, Di Cesare PE. Periprosthetic Sepsis. *Clin Orthop* 2004;(420):26-31.
- (3) Shaneyfelt TM, Centor RM. Reassessment of clinical practice guidelines: go gently into that good night. *JAMA* 2009;301(8):868-869.
- (4) Institute of Medicine. Clinical Practice Guidelines We Can Trust. Washington, D.C.: National Academies Press; 2011.
- (5) Institute of Medicine. Conflict of Interest in Medical Research, Education, and Practice. Washington, D.C.: National Academies Press; 2009.
- (6) Hirsh J, Guyatt G. Clinical experts or methodologists to write clinical guidelines? *Lancet* 2009;374(9686):273-275.
- (7) Atkins D, Eccles M, Flottorp S et al. Systems for grading the quality of evidence and the strength of recommendations I: critical appraisal of existing approaches The GRADE Working Group. *BMC Health Serv Res* 2004;4(1):38.
- (8) Poolman RW, Struijs PA, Krips R, Sierevelt IN, Lutz KH, Bhandari M. Does a "Level I Evidence" rating imply high quality of reporting in orthopaedic randomised controlled trials? *BMC Med Res Methodol* 2006;6:44.
- (9) GRADE handbook for grading quality of evidence and strength of recommendation. The GRADE Working Group; 2009.
- (10) Moher D, Dulberg CS, Wells GA. Statistical power, sample size, and their reporting in randomized controlled trials. *JAMA* 1994;272(2):122-124.
- (11) Institute of Medicine. Finding What Works in Health Care: Standards for Systematic Reviews. Washington, D.C.: National Academies Press; 2011.
- (12) Morrison A, Moulton K, Clark M et al. English-Language Restriction When Conducting Systematic Review-based Meta-analyses: Systemataic Review of Published Studies. Ottawa: Canadian Agency for Drugs and Technologies in Health; 2009.
- (13) Kastner M, Wilczynski NL, Walker-Dilks C, McKibbon KA, Haynes B. Agespecific search strategies for Medline. *J Med Internet Res* 2006;8(4):e25.

- (14) Haynes RB, McKibbon KA, Wilczynski NL, Walter SD, Werre SR. Optimal search strategies for retrieving scientifically strong studies of treatment from Medline: analytical survey. *BMJ* 2005;330(7501):1179.
- (15) Montori VM, Wilczynski NL, Morgan D, Haynes RB. Optimal search strategies for retrieving systematic reviews from Medline: analytical survey. *BMJ* 2005;330(7482):68.
- (16) Wong SS, Wilczynski NL, Haynes RB. Comparison of top-performing search strategies for detecting clinically sound treatment studies and systematic reviews in MEDLINE and EMBASE. *J Med Libr Assoc* 2006;94(4):451-455.
- (17) Kastner M, Wilczynski NL, Walker-Dilks C, McKibbon KA, Haynes B. Agespecific search strategies for Medline. *J Med Internet Res* 2006;8(4):e25.
- (18) Wong SS, Wilczynski NL, Haynes RB. Optimal CINAHL search strategies for identifying therapy studies and review articles. *J Nurs Scholarsh* 2006;38(2):194-199.
- (19) Treadwell JR, Tregear SJ, Reston JT, Turkelson CM. A system for rating the stability and strength of medical evidence. *BMC Med Res Methodol* 2006;6:52.
- (20) Higgins J, Altman D. Assessing risk of bias in included studies. In: Higgins J, Green S, editors. *Cochrane Handbook for SYstematic Reviews of Interventions*. John Wiley & Sons; 2008. 187-241.
- (21) Thorpe KE, Zwarenstein M, Oxman AD et al. A pragmatic-explanatory continuum indicator summary (PRECIS): a tool to help trial designers. *J Clin Epidemiol* 2009;62(5):464-475.
- (22) Sismondo S. Pharmaceutical company funding and its consequences: a qualitative systematic review. *Contemp Clin Trials* 2008;29(2):109-113.
- (23) Ezzet KA. The prevalence of corporate funding in adult lower extremity research and its correlation with reported results. *J Arthroplasty* 2003;18(7 Suppl 1):138-145.
- (24) Khan SN, Mermer MJ, Myers E, Sandhu HS. The roles of funding source, clinical trial outcome, and quality of reporting in orthopedic surgery literature. *Am J Orthop (Belle Mead NJ)* 2008;37(12):E205-E212.
- (25) Montori VM, Jaeschke R, Schunemann HJ et al. Users' guide to detecting misleading claims in clinical research reports. *BMJ* 2004;329(7474):1093-1096.
- (26) Salanti G, Higgins JP, Ades AE, Ioannidis JP. Evaluation of networks of randomized trials. *Stat Methods Med Res* 2008;17(3):279-301.

- (27) Petitti DB, Teutsch SM, Barton MB, Sawaya GF, Ockene JK, DeWitt T. Update on the methods of the U.S. Preventive Services Task Force: insufficient evidence. *Ann Intern Med* 2009;150(3):199-205.
- (28) Murphy MK, Black LA, Lamping DL, McKee CM, Sanderson C.F., Askam J. Consensus development methods, and their use in clinical guideline development. *Health Technol Assess* 1998.
- (29) Fleming TR, DeMets DL. Surrogate end points in clinical trials: are we being misled? *Ann Intern Med* 1996;125(7):605-613.
- (30) Shaughnessy AF, Slawson DC. What happened to the valid POEMs? A survey of review articles on the treatment of type 2 diabetes. *BMJ* 2003;327(7409):266.
- (31) Grimes DA, Schulz KF. Surrogate end points in clinical research: hazardous to your health. *Obstet Gynecol* 2005;105(5 Pt 1):1114-1118.
- (32) Prentice RL. Surrogate endpoints in clinical trials: definition and operational criteria. *Stat Med* 1989;8(4):431-440.
- (33) Stata Statistical Software: Release 10 [computer program]. College Station, TX: StatCorp LP; 2007.
- (34) Hozo SP, Djulbegovic B, Hozo I. Estimating the mean and variance from the median, range, and the size of a sample. *BMC Med Res Methodol* 2005;5(1):13.
- (35) Lu G, Ades AE. Combination of direct and indirect evidence in mixed treatment comparisons. *Stat Med* 2004;23(20):3105-3124.
- (36) Brooks S, Gelman A. Alternative methods for monitoring convergence of iterative simulations. *Journal of Computational and Graphical Statistics* 1998;7:434-455.
- (37) Dias S, Welton NJ, Caldwell DM, Ades AE. Checking consistency in mixed treatment comparison meta-analysis. *Stat Med* 2010;29(7-8):932-944.
- (38) DerSimonian R., Laird N. Meta-Analysis in Clinical Trials. *Controlled Clinical Trials* 1986;7:177-188.
- (39) Berbari EF, Osmon DR, Carr A et al. Dental procedures as risk factors for prosthetic hip or knee infection: a hospital-based prospective case-control study. *Clin Infect Dis* 2010;50(1):8-16.