

## CADTH RAPID RESPONSE REPORT: SUMMARY WITH CRITICAL APPRAISAL

# Preoperative Interventions for the Prevention of Surgical Site Infections: A Review of Guidelines

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### **Abbreviations**

AGREE II Appraisal of Guidelines for Research & Evaluation 2

CDC Centers for Disease Control and Prevention

CHG Chlorhexidine gluconate

GRADE Grading of Recommendations, Assessment, Development, and

Evaluation

MRSA methicillin-resistant *Staphylococcus aureus*NICE National Institute for Health and Care Excellence

RCT Randomized controlled trial S. aureus Staphylococcus aureus SR Systematic Review SSI Surgical site infection

### **Context and Policy Issues**

Defined as postoperative infections of an incision, organ, or space, <sup>1</sup> surgical site infections (SSIs) are the most common health care-related infections. <sup>2</sup> Occurring in up to 5% of all surgeries, SSIs affect approximately 26,000 to 65,000 Canadian patients annually. <sup>2</sup> Due to increased hospital stays and readmission rates, SSIs cost the health care system between \$350,000 to \$1 million each year. <sup>2</sup> To help reduce morbidity, extended hospitalization, and death, infection control measures have been implemented in surgical settings. <sup>2</sup>

Surgical infection control measures include staff precautions such as practicing hand hygiene and using barrier devices, and patient-specific perioperative infection control interventions that may include nasal decolonization for *Staphylococcus aureus* (*S. aureus*), preoperative washing, skin antisepsis, hair removal, glucose control, bowel preparation, and antibiotic prophylaxis.<sup>3</sup> It has been shown that almost half of SSIs may be prevented by applying evidence-based strategies.<sup>4</sup> SSI prevention measures can be bundled to promote staff and patient adherence, but there is a lack of consensus regarding the appropriate components of an infection control bundle.<sup>3</sup>

This report is an upgrade from a previous CADTH Reference List report published in 2020, and includes one of the research questions from that report.<sup>5</sup> The aim of the current report is to summarize and critically appraise the relevant evidence-based guidelines identified in the previous report<sup>5</sup> regarding preoperative interventions for the prevention of SSIs.

### **Research Question**

What are the evidence-based guidelines regarding preoperative interventions for the prevention of surgical site infections?

### **Key Findings**

Six evidence-based guidelines regarding the use of preoperative interventions for the prevention of surgical site infections were included in this report. Four included guidelines were of high quality, while two guidelines were of moderate quality due to unclear reporting of methodological details.

Of these guidelines, for the purpose of infection prevention, four recommend nasal decolonization with mupirocin, body washing with chlorhexidine gluconate, and bathing with antimicrobial or non-antimicrobial soap prior to surgery. Furthermore, four guidelines recommend the use of alcohol-based solutions for skin antiseptic preparation but



recommend against hair removal unless absolutely required. Three guidelines recommend mechanical bowel preparation with oral antibiotics for elective colorectal surgery, while one guideline recommends against its use with no mention of specific indications or concurrent antibiotic use. Two guidelines made recommendations on perioperative blood glucose control with different target levels. Four guidelines made recommendations on the optimal time for administering antibiotic prophylaxis (i.e., at one or two hours before incision, or at the time of anesthesia). Overall, these recommendations ranged from conditional to strong and were based on evidence that ranged in quality from very low to high (when reported).

### Methods

### Literature Search Methods

The literature search that was conducted for a previous CADTH report<sup>5</sup> was used for this report. A limited literature search was conducted by an information specialist on key resources including Medline via Ovid, the Cochrane Library, the University of York Centre for Reviews and Dissemination (CRD) databases, the websites of Canadian and major international health technology agencies, as well as a focused internet search. The search strategy was comprised of both controlled vocabulary, such as the National Library of Medicine's MeSH (Medical Subject Headings), and keywords. The main search concepts in the previous CADTH report<sup>5</sup> were surgical site infections in the preoperative setting. Methodological filters were applied to limit the retrieval to guidelines only. The search was also limited to English language documents published between January 1, 2015 and February 27, 2020.

### Selection Criteria and Methods

One reviewer screened citations and selected studies. In the first level of screening, titles and abstracts were reviewed and potentially relevant articles were retrieved and assessed for inclusion. The final selection of full-text articles was based on the inclusion criteria presented in Table 1.

**Table 1: Selection Criteria** 

Population	Surgical patients, any age
Intervention	Preoperative bundle components for the prevention of surgical site infections (e.g., nasal decolonization interventions, chlorhexidine gluconate washes/wipes/bathing, preoperative washing with other methods or agents, oral antibiotics)
Comparator	Not applicable
Outcomes	Recommendations regarding preoperative interventions for the prevention of surgical site infections
Study Designs	Evidence-based guidelines

### **Exclusion Criteria**

Articles were excluded if they did not meet the selection criteria outlined in Table 1, they were duplicate publications, or were published prior to 2015. Guidelines with unclear methodology were excluded.



### Critical Appraisal of Individual Studies

The included guidelines were critically appraised by one reviewer using the Appraisal of Guidelines for Research and Evaluation (AGREE) II instrument<sup>6</sup> as a guide. Summary scores were not calculated for the included studies; rather, a review of the strengths and limitations of each included guideline were described narratively.

### **Summary of Evidence**

### Quantity of Research Available

A total of 354 citations were identified in the literature search. Following screening of titles and abstracts, 350 citations were excluded and 4 potentially relevant reports from the electronic search were retrieved for full-text review. Eight potentially relevant publications were retrieved from the grey literature search for full-text review. Of these potentially relevant articles, six publications were excluded for various reasons, and six evidence-based guidelines met the inclusion criteria and were included in this report. Appendix 1 presents the PRISMA<sup>7</sup> flowchart of the study selection.

Additional publications of potential interest are provided in Appendix 5.

### Summary of Study Characteristics

Six evidence-based guidelines were identified and included in this report. 1,8-12 Detailed characteristics of the guidelines are available in Appendix 2.

### Study Design

Six evidence-based guidelines were identified regarding preoperative interventions for the prevention of SSIs.<sup>1,8-12</sup> Two of these guidelines were published in 2019 and were developed by the Asia Pacific Society of Infection Control (APSIC)<sup>11</sup> and the National Institute for Health and Care Excellence (NICE).<sup>9</sup> Two guidelines were published in 2017 and were developed by the American College of Surgeons (ACS) & Surgical Infection Society<sup>12</sup> and the US Centers for Disease Control and Prevention (CDC).<sup>1</sup> The guidelines developed by the WHO<sup>8</sup> and the Ministry of Health Malaysia<sup>10</sup> were published in 2016 and 2015, respectively.

The CDC guidelines were informed by a SR which included systematic searches from 1998 through April 2014 for relevant randomized controlled trials (RCTs) and SRs. The WHO guidelines were informed by a SR which included systematic searches between December 2013 and October 2015 for RCTs and non-randomized studies. The NICE guidelines were informed by systematic searches on March 15, 2018 and screened for RCTs and SRs of RCTs. The Ministry of Health Malaysia guidelines were informed by systematic searches from 2003 onwards and screened for SRs, RCTs, and non-randomized studies. The APSIC guidelines were informed by computerized literature searches on PubMed and a review of other published guidelines (e.g., CDC, WHO), while the ACS guidelines were informed by literature searches on PubMed.

The NICE, <sup>9</sup> CDC, <sup>1</sup> and WHO<sup>8</sup> guidelines used the Grading of Recommendations Assessment, Development and Evaluation (GRADE) approach to evaluate the quality of the evidence. The NICE guideline graded the quality of evidence from very low to high, and reflected the strength of recommendations in the wording (i.e., "offer/advise" was used for strong recommendations with clear evidence of benefit, while "consider" was used if the



evidence was less certain). The CDC guideline graded the quality of evidence from C (i.e., required by state/federal regulation) to A (i.e., high to moderate-quality evidence), and classified the strength of recommendation from no recommendation to strong recommendation. The WHO guideline graded the quality of evidence from very low to high, and classified the strength of recommendation as conditional or strong.

The authors of the APSIC guideline assessed the quality of evidence as category III (i.e., evidence from opinions or expert committee reports) to category I (i.e., evidence from  $\geq$  one RCT), and rated the strength of recommendation from E (i.e., good evidence to recommend against use) to A (i.e., good evidence to recommend use). The guideline development group for the Ministry of Health Malaysia guideline used the US/Canadian Preventive Services Task Force guide to classify the quality of evidence from category III (i.e., evidence from opinions or expert committee reports) to category I (i.e., evidence from  $\geq$  one RCT), and rated the strength of recommendation using the Scottish Intercollegiate Guidelines Network system from C (i.e., evidence from opinions or expert committee reports) to A (i.e., evidence from  $\geq$  one meta-analysis, SR, or RCT). The authors of the ACS guideline did not report assessment of the quality of the evidence or grade the strength of the recommendations. The summary of the evidence or grade the strength of the recommendations.

The rating systems for quality of evidence and strength of recommendations, if available, are reported in Appendix 4. Decisions on the recommendations were reached through consensus in five guidelines, <sup>1,8-10,12</sup> while the methodology for formulating the recommendations was not reported in the APSIC guideline. <sup>11</sup>

### Country of Origin

The WHO guideline<sup>8</sup> is meant to apply globally, while the ACS<sup>12</sup> and CDC<sup>1</sup> guidelines are meant to apply to the United States. The other three guidelines are meant to apply to the Asia Pacific region,<sup>11</sup> the United Kingdom,<sup>9</sup> and Malaysia.<sup>10</sup>

### Patient Population

The target population covered by the six guidelines was patients undergoing surgical procedures. 1,8-12 The Ministry of Health Malaysia guideline specifically focused on patients undergoing oral and maxillofacial surgical procedures. 10 The intended users of four guidelines are surgical staff who provide care for surgical patients. 1,8-12 In two guidelines, the intended users were not explicitly stated, but they appear to be intended for surgical staff. 11,12 The CDC, 1 WHO, 8 and Ministry of Health Malaysia 10 guidelines are also intended for professional societies and organizations, anesthesiologists and pharmacists, and dental practitioners and educators, respectively.

### Interventions and Comparators

The six guidelines considered a variety of preoperative interventions that can be used to help prevent SSIs.<sup>1,8-12</sup> These included preoperative body washing, mechanical bowel preparation, antibiotic prophylaxis, hair removal, skin antiseptic, glucose control, and nasal decolonization.

### **Outcomes**

All six guidelines considered the incidence of SSIs.<sup>1,8-12</sup> Three guidelines also considered SSI-related deaths,<sup>1,8,9</sup> and two guidelines<sup>1,9</sup> considered duration of hospital stay, hospital readmission, antimicrobial resistance, and adverse events. Additionally, the NICE guideline



considered postoperative antibiotic use, infection complications, and other nosocomial infections as outcomes.<sup>9</sup>

### Summary of Critical Appraisal

Additional details regarding the AGREE II evaluation of the included guidelines are provided in Appendix 3.

The guideline development groups for all six guidelines were comprised of experts from multidisciplinary areas. All guidelines provided a clear description of their objectives, specified the target populations, and provided unambiguous and easily identifiable recommendations. 1,8-12 The views of the funding sources did not appear to have influenced the guidelines' contents. 1,8-12 However, the views and preferences of the target population were not sought and the target users of the guidelines were not explicitly defined in two guidelines. 11,12

With respect to rigour of guideline development, the APSIC<sup>11</sup> and the ACS<sup>12</sup> guideline provided brief details of their methodology without search timeframes. Additionally, the authors of these two guidelines conducted literature searches in one database (i.e., PubMed), which may have resulted in omission of relevant information. The methodology for formulating the recommendations was not reported in the APSIC guideline.<sup>11</sup> The authors of the ACS guideline did not assess the quality of the evidence or grade the strength of the recommendations.<sup>12</sup> In the NICE guideline, the supporting evidence used to inform the recommendations on preoperative bathing, hair removal, mechanical bowel preparation, and antibiotic prophylaxis was not updated since the original 2008 guideline publication.<sup>13</sup> Furthermore, the quality of the aforementioned supporting evidence was not available online. The recommendations on preoperative washing, nasal decolonization, and skin antisepsis were updated in the current NICE guideline.<sup>9</sup> All six guidelines were externally reviewed prior to publication.<sup>1,8-12</sup> However, three guidelines did not explicitly describe procedures for guideline updates.<sup>10-12</sup>

In terms of guideline applicability, all six guidelines presented monitoring criteria for their recommendations. The NICE<sup>9</sup> CDC,<sup>1</sup> and WHO<sup>8</sup> guideline described facilitators or barriers to their application and provided tools for putting recommendations into practice. Four guidelines did not consider the potential resource implications of applying the recommendations.<sup>1,10-12</sup> Finally, the six guidelines were developed for use in the Asia Pacific region,<sup>11</sup> Malaysia,<sup>10</sup> United Kingdom,<sup>9</sup> United States,<sup>1,12</sup> or globally;<sup>8</sup> therefore, the generalizability of the recommendations to the Canadian context is unclear.

### Summary of Findings

The recommendations regarding the use of preoperative interventions for the prevention of SSIs from the six included evidence-based guidelines are highlighted below. 1.8-12 Detailed summaries of the recommendations and the evidence on which the recommendations were based are presented in Appendix 4.

### Guidelines

### Recommendations Regarding S. aureus and/or Antimicrobial Resistance Screening

Two identified evidence-based guidelines provided recommendations regarding preoperative *S. aureus* and/or antimicrobial resistance screening. The APSIC and ACS guidelines recommend that hospitals evaluate their baseline *S. aureus*, *methicillin-resistant* 



S. aureus (MRSA), and SSI rates to determine if the implementation of *S. aureus* screening and decolonization procedures is appropriate. <sup>11,12</sup> Furthermore, the APSIC guideline recommends that surveillance on mupirocin resistance rates should be taken into consideration when implementing decolonization measures. <sup>11</sup> The APSIC recommendations <sup>11</sup> were rated B in strength and were based on level II evidence, while the ACS guideline <sup>12</sup> did not report the recommendation strength or evidence level.

### Recommendations Regarding Nasal Decolonization and Preoperative Body Washing

Four identified evidence-based guidelines provided recommendations regarding nasal decolonization, 8,9,11,12 and five provided recommendations on preoperative body washing. 1,8,9,11,12

Regarding nasal decolonization, the NICE guideline recommends considering using nasal decolonization with mupirocin along with preoperative body washing with chlorhexidine gluconate (CHG) if *S. aureus* is a likely cause of SSIs (recommendation strength: consider; evidence level: very low to high).<sup>9,14</sup> The APSIC and WHO guidelines recommend nasal decolonization with mupirocin 2% ointment, with or without CHG body washing, for cardiothoracic and orthopedic surgical patients with known *S. aureus* nasal carriage (recommendation strength: A or strong; evidence level: I or moderate).<sup>8,11</sup> The WHO guideline states that this recommendation can also be considered for other surgical procedures (recommendation strength: conditional; evidence level: moderate).<sup>8</sup> The ACS guideline recommends clinicians consider using nasal mupirocin, with or without CHG body washing (strength of recommendation and level of evidence not reported).<sup>12</sup> For SSI bundles to be effective, interventions should be adhered to and performed close to surgery date (strength of recommendation and level of evidence not reported).<sup>12</sup> The CDC did not make recommendations regarding nasal decolonization.<sup>1</sup>

Regarding preoperative body washing, four guidelines (APSIC, NICE, CDC, and WHO) recommend surgical patients bathe with soap before their procedure. 1,8,9,11 These recommendations ranged from conditional to strong and were based on evidence that ranged in quality from very low to moderate where reported (evidence level not reported in the NICE guideline). 1,8,9,11 Three guidelines (APSIC, CDC, and WHO) state that patients can use antimicrobial or non-antimicrobial soap for their preoperative bath (recommendation strength ranged from conditional to strong; evidence level ranged from very low to moderate). 1,8,11 Due to uncertainty in the available evidence, the CDC and WHO were not able to make recommendations regarding the use of CHG washcloths. 1,8 Furthermore, the ACS guideline highlighted that body washing with CHG, when not part of an SSI bundle, reduces skin pathogen levels but not SSI rates (strength of recommendation and level of evidence not reported). 12

### **Recommendations Regarding Skin Antiseptic Preparation**

Four identified evidence-based guidelines provided recommendations regarding preoperative skin antisepsis. 8,9,11,12 Unless contraindicated, alcohol-based skin antiseptic solutions should be used to prepare the surgical site prior to incision as per APSIC, NICE, WHO, and ACS guidelines (recommendation strength was A or strong; evidence level ranged from very low to high where reported; recommendation strength and evidence level not reported in the ACS guideline). 8,9,11,12 Contraindications to the use of alcohol-based solutions may include surgical sites close to or involving mucous membrane, 9,12 cornea, 12 or ear. 12 Three guidelines made recommendations on the specific types of skin antiseptic solutions to use. 8,9,12 As the first choice for skin antisepsis, the NICE9 and WHO8 guidelines recommend using alcohol-based CHG solutions, while the ACS12 guideline recommends



alcohol-based CHG or iodine solutions (recommendation strength was strong; evidence level ranged from very low to high where reported; recommendation strength not reported in the ACS and NICE guidelines and evidence level not reported in the ACS guideline). If alcohol-based CHG solutions are contraindicated, the NICE guideline recommends using alcohol-based povidone-iodine, aqueous CHG, or aqueous povidone-iodine solutions (strength of recommendation: not reported; evidence levels: very low to high). Similarly, if alcohol-based solutions are contraindicated, the ACS guideline also recommends aqueous CHG over the use of aqueous iodine solutions (strength of recommendation and level of evidence not reported).

### **Recommendations Regarding Hair Removal**

Four identified evidence-based guidelines provided recommendations regarding preoperative hair removal.<sup>8,9,11,12</sup> The APSIC, NICE, ACS, and WHO guidelines recommend that hair should not be removed unless absolutely needed due to interference with the surgical procedure (recommendation strength was B or strong; evidence level ranged from III to moderate where reported; recommendation strength not reported in the ACS guideline and evidence level not reported in the ACS and NICE guidelines).<sup>8,9,11,12</sup> Specifically, clippers should be used while razors should be avoided as per APSIC, NICE, ACS, and WHO guidelines (recommendation strength was A or strong; evidence level was I or moderate where reported; recommendation strength not reported in the ACS guideline and evidence level not reported in the ACS and NICE guidelines).<sup>8,9,11,12</sup> As for the timing of hair removal, the NICE guideline<sup>9</sup> recommends that it be performed on the day of surgery (recommendation strength: strong; evidence level: not reported), while the APSIC guideline <sup>11</sup> states that no recommendation can be made (recommendation strength: C; evidence level: III).

### **Recommendations Regarding Mechanical Bowel Preparation**

Four identified evidence-based guidelines provided recommendations regarding preoperative mechanical bowel preparation. <sup>8,9,11,12</sup> The APSIC, ACS, and WHO guidelines recommend the use of mechanical bowel preparation combined with oral antibiotics for elective colorectal surgical patients (recommendation strength A or conditional; evidence level was I or moderate where reported; recommendation strength and evidence level not reported in the ACS guideline), <sup>8,11,12</sup> and the WHO guideline recommends against using mechanical bowel preparation alone without oral antibiotics for this population (recommendation strength: strong; evidence level: moderate). However, the NICE guideline recommends against the routine use of mechanical bowel preparation to help prevent SSIs, but did not specify the specific indication or whether this was irrespective of oral antibiotic use (recommendation strength: strong; evidence level: not reported). <sup>9</sup>

### **Recommendations Regarding Glucose Control**

Two identified evidence-based guidelines provided recommendations regarding preoperative glucose control immediately prior to surgery. The ACS guideline recommends maintaining perioperative blood glucose levels between 110 to 150 mg/dL (6.1 to 8.3 mmol/L) for all patients living with or without diabetes and less than 180 mg/dL (10.0 mmol/L) for patients undergoing cardiac surgery (strength of recommendation and level of evidence not reported). Furthermore, the ACS guideline states that blood glucose targets of less than 110 mg/dL (6.1 mmol/L) increase the risk of hypoglycemia without reducing the rates of SSIs (strength of recommendation and level of evidence not reported). The CDC guideline recommends maintaining blood glucose levels under 200 mg/dL (11.1 mmol/L) without regard for diabetes status (recommendation strength: strong:



evidence level: A).<sup>1</sup> Due to the lack of evidence from RCTs, the CDC was not able to make recommendations regarding lower glucose targets, or regarding the optimal timing, duration, or delivery method of glucose control interventions.<sup>1</sup>

### **Recommendations Regarding Antibiotic Prophylaxis**

Six identified evidence-based guidelines provided recommendations regarding preoperative antibiotic prophylaxis. 1,8-12

The APSIC and ACS guidelines recommend that, when indicated, antibiotic prophylaxis should be administered within one hour prior to incision, except for vancomycin and fluoroquinolones that need to be given within two hours before incision (recommendation strength: A; evidence level: I where reported; recommendation strength and evidence level not reported in the ACS guideline). The WHO guideline recommends antibiotic prophylaxis be administered within two hours before incision with consideration of the half-life of the antibiotic (recommendation strength: strong; evidence level: moderate). The NICE guideline recommends considering the administration of a dose of antibiotic prophylaxis intravenously when anesthesia is started (recommendation strength: consider; evidence level: not reported). The CDC guideline recommends that the administration of parenteral antibiotic prophylaxis should be timed to ensure bactericidal concentration is reached when the incision is made (recommendation strength: strong; evidence level: low to very low-quality); however, no recommendation could be made on specific timing.

The CDC guideline recommends that the indication to use parenteral antibiotic prophylaxis should be based on published guidelines (recommendation strength: strong; evidence level: low to very low-quality). Specifically, the CDC recommends that parenteral antibiotic prophylaxis should be given for all cesarean sections (recommendation strength: strong; evidence level: high). The NICE guideline recommends the administration of antibiotic prophylaxis before contaminated procedures, clean-contaminated procedures, or clean procedures that involve prosthesis or implant placement (recommendation strength: strong; evidence level: not reported). However, the NICE guideline recommends against the routine use of antibiotic prophylaxis for clean non-prosthetic uncomplicated procedures (recommendation strength: strong; evidence level: not reported).

The NICE<sup>9</sup> guideline recommends that the choice of antibiotic should take into consideration local antibiotic formularies and possible side effects (recommendation strength: strong; evidence level: not reported), while the ACS guideline<sup>12</sup> recommends taking into account the type of surgical procedure and most likely etiological pathogens (strength of recommendation and level of evidence were not reported). Although the CDC guideline<sup>1</sup> was not able to make a recommendation on the effect of weight-adjusted parenteral antibiotic prophylaxis dosing due to the lack of RCTs, the ACS guideline recommends that antibiotic prophylaxis dosing should be weight-adjusted (strength of recommendation and level of evidence was not reported).<sup>12</sup>

The sixth guideline published by the Ministry of Health Malaysia focused on the use of antibiotic prophylaxis for the prevention of oral surgery-related SSIs. <sup>10</sup> The guideline makes grade A recommendations stating that antibiotic prophylaxis is indicated for dental implants (evidence level: I), bone grafts (evidence level: I), clean-contaminated procedures (evidence levels: I, II-2, and III), and head and neck cancers (evidence levels: II-1, II-3, and III). <sup>10</sup> The guideline makes grade B recommendations stating that antibiotic prophylaxis is indicated for medically compromised patients (evidence levels: I, II-2, and III), extended duration procedures (evidence levels: II-2 and III), and facial bone fracture procedures



(evidence levels: II-2, II-3, and III). <sup>10</sup> The guideline makes grade B recommendations stating that amoxicillin, penicillin G, or clindamycin (evidence levels: I, II-1, II-3, and III) should be given within one hour before incision (evidence levels: II-2, II-3, and III) and at the full therapeutic dose (evidence levels: II-2 and II-3). <sup>10</sup> Alternatively, cloxacillin, cefazolin or clindamycin should be administered if the procedure involves the skin (recommendation strength: C; evidence level: III). <sup>10</sup>

### Limitations

One evidence-based guideline was created by oral surgeons and dental health specialists and was focused on the prevention of oral surgery-related SSIs; therefore, these recommendations may not be generalizable to other surgical procedures. Apart from one guideline that is intended for global use, the other five guidelines were developed for use in the Asia Pacific region, Malaysia, United Kingdom, or United States. United States. United States if these guidelines developed outside of Canada are generalizable to the Canadian context. Thus, considering the limitations mentioned, the recommendations summarized in this report need to be interpreted with caution.

### **Conclusions and Implications for Decision or Policy Making**

This report provides a summary and critical appraisal of guidelines identified in a previous CADTH Reference List report;<sup>5</sup> a companion CADTH report regarding the clinical effectiveness and cost-effectiveness of pre-operative nasal decolonization (with or without chlorhexidine gluconate washes or wipes) for the prevention of surgical site infections is published elsewhere. This report included six evidence-based guidelines regarding the use of preoperative interventions for the prevention of SSIs. The Prevention published by the University of Toronto's Best Practice in Surgery was excluded from this report due to unclear methodology (see Appendix 5). Informed partly by the WHO8 and NICE9 guidelines, this guideline included similar recommendations on nasal decolonization, skin antisepsis, hair removal, and antibiotic prophylaxis tailored for University of Toronto affiliated hospitals. The prevention of SUS and SU

Of the included guidelines, four recommend nasal decolonization with mupirocin and body washing with CHG.<sup>8,9,11,12</sup> Additionally, four guidelines recommend bathing with antimicrobial or non-antimicrobial soap before surgery.<sup>1,8,9,11</sup> Two guidelines recommend taking into account the rates of *S. aureus*, MRSA, SSIs, and/or mupirocin resistance in the implementation of screening and decolonization measures.<sup>11,12</sup> Four guidelines recommend alcohol-based solutions for skin antisepsis and recommend against hair removal.<sup>8,9,11,12</sup> Three guidelines<sup>8,11,12</sup> recommend mechanical bowel preparation with oral antibiotics for elective colorectal surgery, while one guideline<sup>9</sup> recommends against its routine use but did not specify for which indications and did not mention oral antibiotic use. Two guidelines recommend maintaining perioperative blood glucose levels at 110 to 150 mg/dL (6.1 to 8.3 mmol/L)<sup>12</sup> or below 200 mg/dL (11.1 mmol/L)<sup>1</sup> without regard for diabetic status. Four guidelines recommend administering antibiotics at one<sup>11,12</sup> or two<sup>8</sup> hours prior to incision, or at the time of anesthesia.<sup>9</sup> Focusing on antibiotic prophylaxis for oral surgical procedures, one guideline recommends the use of amoxicillin, penicillin G, or clindamycin, or cloxacillin, cefazolin, or clindamycin for procedures involving the skin.<sup>10</sup>

Overall, these recommendations ranged from conditional to strong and were based on evidence that ranged in quality from very low to high where reported (recommendation strength and/or evidence level was not reported for all ACS recommendations and some



NICE recommendations).<sup>1,8-12</sup> In general, there was agreement across guidelines for the use of preoperative interventions for the prevention of SSIs; however, the variation in the strengths of recommendations and heterogeneity in the quality of evidence should be considered when interpreting the findings of this report. Guidelines developed with rigorous methodology that are specific to the Canadian context would provide additional guidance in preventing SSIs in a more local context.



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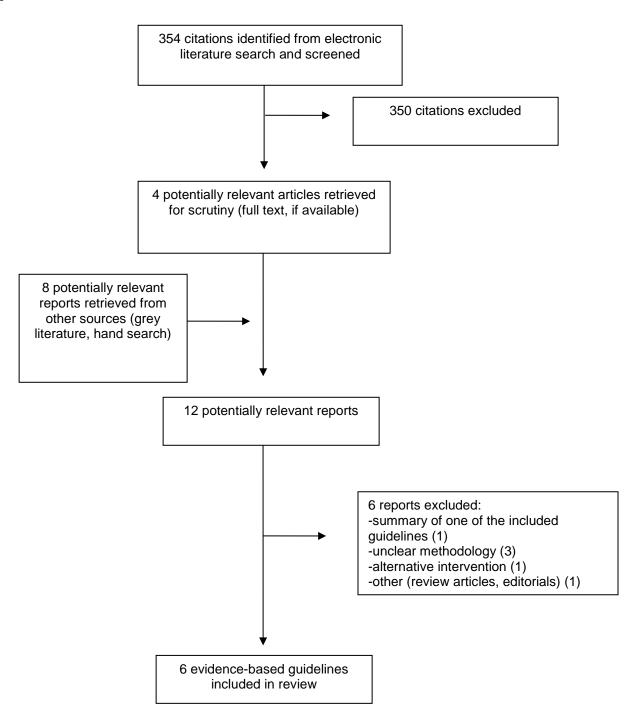
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# **Appendix 1: Selection of Included Studies**





# **Appendix 2: Characteristics of Included Publications**

**Table 2: Characteristics of Included Guideline** 

Intended Users, Target Population, Country, Funding Source	Intervention and Practice Considered	Major Outcomes Considered	Evidence Collection, Selection, and Synthesis	Evidence Quality Assessment	Recommendations Development and Evaluation	Guideline Validation
		Asia Pacific So	ciety of Infection Co	ontrol (APSIC) Guideline, 201	911	
Intended users: Surgical staff who provide care for surgical patients  Target population: Patients undergoing surgical procedures  Asia Pacific region  Funding source: Educational grant from 3 M Asia Pacific	The guideline provided recommendation s regarding preoperative washing, mechanical bowel preparation, antibiotic prophylaxis, hair removal, skin antiseptic, and nasal decolonization.	Incidence of SSIs	The APSIC working group conducted literature searches in PubMed and screened published guidelines (e.g., WHO, CDC, Cochrane). SRs, RCTs and non- randomized studies were eligible for inclusion.	Categories for quality of evidence: I: Evidence derived from ≥ one RCT II: Evidence derived from ≥ one well-designed non-randomized study, cohort or case-controlled study, or critical results from uncontrolled studies III: Opinions of respected authorities based on clinical experience, descriptive studies, or expert committee reports	The working group engaged in discussions in person and via email to complete the guideline.  Recommendation grading system: A: Good evidence to support recommendation for use B: Moderate evidence to support recommendation for use C: Insufficient evidence to recommend for or against use D: Moderate evidence to recommend against use E: Good evidence to recommend against use	The guideline was validated by two external reviewers, APSIC Executive Committee, and nationa Infection Control societies in Asia Pacific.
	Nat	tional Institute fo	or Health and Care I	Excellence (NICE) Guideline,	2019 <sup>9</sup>	
Intended users: Health care professionals, commissioners and providers, and those undergoing surgery  Target population: Adults, young people, and children undergoing surgical procedures  United Kingdom	The guideline provided recommendation s regarding preoperative washing, mechanical bowel preparation, antibiotic prophylaxis, hair removal, antiseptic skin preparation, and	Incidence of SSIs, mortality, duration of hospital stay, postoperative antibiotic use, hospital readmission, infection complications, antimicrobial resistance, adverse events, and	Literature searches were conducted on March 15, 2018 in various databases (e.g., Medline, Embase, Cochrane Database of Systematic Reviews). Retrieved articles were screened for RCTs and SRs of RCTs.	Evidence quality was assessed using the GRADE approach and presented in GRADE tables with quality of evidence ranked from very low to high.	The GDG develops recommendations based on scientific evidence and other evidence such as expert testimony and stakeholder views. The guideline development group reaches an agreement on the strength of recommendations through an informal consensus process.  The strength of recommendations is reflected in the wording:	Draft NICE guidelines are posted online for review by registered stakeholders. Albeit not mentioned in this guideline, NICE occasionally solicits external

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Intended Users, Target Population, Country, Funding Source	Intervention and Practice Considered	Major Outcomes Considered	Evidence Collection, Selection, and Synthesis	Evidence Quality Assessment	Recommendations Development and Evaluation	Guideline Validation
Funding source: The United Kingdom government	nasal decolonization.	other nosocomial infections			Offer/Advise: Strong recommendation (i.e., clear evidence of benefit) Consider: Evidence of benefit is less certain	experts for review.
	America	n College of Su	rgeons (ACS) & Su	rgical Infection Society Guide	line, 2017 <sup>12</sup>	•
Intended users: Surgical staff who provide care for surgical patients  Target population: Patients undergoing surgical procedures  United States  Funding source: ACS & Surgical Infection Society	The guideline provided recommendation s regarding preoperative washing, bowel preparation, antibiotic prophylaxis, hair removal, skin preparation, nasal decolonization, glucose control, and MRSA screening.	Incidence of SSIs	Literature searches were conducted in PubMed on specific topics to fill knowledge gaps in previous guidelines. This is an update to a previous version of the guideline. Retrieved articles were screened for SRs, RCTs and non-randomized studies.	Quality assessment of the included evidence was not mentioned.	Consensus agreement was reached by internal and external experts for this guideline update.  There was no mention of using a grading system for strength of recommendations.	The guideline was validated by an internal expert panel and external expert reviewers.
		Centers for Dise	ease Control and Pro	evention (CDC) Guideline, 20	)17 <sup>1</sup>	
Intended users: Health care professionals, professional societies/organization s  Target population: Patients undergoing surgical procedures  United States  Funding source: CDC	The guideline provided recommendation s regarding preoperative washing, antibiotic prophylaxis, and glucose control.	Incidence of SSIs, mortality, duration of hospital stay, hospital readmission, adverse events, and antimicrobial resistance	A systematic review was conducted from 1998 through April 2014 in various databases (e.g., Medline, Embase, CINAHL, Cochrane Library). Retrieved articles were screened for RCTs and SRs.	Evidence quality was assessed using a modified GRADE approach.  Categories for quality of evidence:  A: High to moderate-quality evidence that suggest net benefit or harm  B: Low-quality evidence that suggest net benefit or harm, or an accepted practice supported by low to very low-quality evidence  C: Required by state/federal regulation	The guideline development group develops recommendations based on the literature review and a consensus process.  Recommendation grading system:  I: Strong recommendation II: Weak recommendation: supported by any quality evidence that suggest a trade-off between risk and benefit No recommendation: Low to very low-quality evidence with an uncertain trade-off of risk and benefit, or no	The guideline was reviewed by a panel of experts, HICPAC members, and members of the public.

					NDTLI	
Intended Users, Target Population, Country, Funding Source	Intervention and Practice Considered	Major Outcomes Considered	Evidence Collection, Selection, and Synthesis	Evidence Quality Assessment	Recommendations Development and Evaluation	Guideline Validation
					published evidence deemed critical	
			WHO Guidel	ine, 2016 <sup>8</sup>		
Intended users: Surgeons, nurses, support staff, anesthesiologists, pharmacists, and other professionals providing surgical care  Target population: Patients undergoing surgical procedures  Global  Funding sources: WHO and Fleming Fund of the United Kingdom Government	The guideline provided recommendation s regarding preoperative washing, bowel preparation, antibiotic prophylaxis, hair removal, skin preparation, and nasal decolonization.	Incidence of SSIs and SSI- related deaths	A SREG made up of researchers and professionals conducted multiple systematic reviews between December 2013 and October 2015 to provide the evidence for this guideline. Literature searches were conducted in various databases (e.g., Medline, Excerpta Medica Database, CINAHL, Cochrane Central Register of Controlled Trials, WHO regional databases) for studies published after January 1, 1990. Retrieved articles were screened for RCTs and nonrandomized studies.	Evidence quality was assessed using GRADE.  Categories for quality of evidence: High: very confident that the true effect lies close to the effect estimate Moderate: moderately confident that the true effect lies close to the effect estimate Low: the true effect may differ considerably from the effect estimate Very low: the true effect likely differs considerably from the effect estimate	The GDG develops recommendations based on the literature review and a consensus process.  Recommendation grading system: Strong recommendation: GDG was confident that the benefits outweigh the risks Conditional recommendation (may use the terminology "suggests considering"): GDG considered that the benefits probably outweighed the risks	The guideline was reviewed by an External Peer Review Group with five technical experts with extensive knowledge in surgery and infection prevention and control measures.
	<u> </u>	Minis	stry of Health Malay	rsia Guideline, 2015 <sup>10</sup>		
Intended users: Oral and maxillofacial surgeons, dental practitioners, and	The guideline provided recommendation s regarding	Incidence of oral SSIs	Literature searches were conducted in various databases	Evidence quality was assessed using the US/Canadian Preventive Services Task Force	The guideline development group and review committee develop recommendations based on the literature	The guideline was reviewed

Intended Users, Target Population, Country, Funding Source	Intervention and Practice Considered	Major Outcomes Considered	Evidence Collection, Selection, and Synthesis	Evidence Quality Assessment	Recommendations Development and Evaluation	Guideline Validation
academics involved in dentist training  Target population: Patients undergoing oral and maxillofacial surgical procedures  Malaysia  Funding source: Malaysia Ministry of Health	antibiotic prophylaxis.		(e.g., PubMed/Medline, Cochrane Database of Systematic Reviews, ISI Web of Knowledge, OVID) for articles published from 2003 onwards. Retrieved articles were screened for SRs, RCTs and non-randomized studies.	guide.  Categories for quality of evidence: I: Evidence derived from ≥ one RCT II-1: Evidence derived from ≥ one well-designed non-randomized study II-2: Evidence derived from ≥ one well-designed cohort or case-controlled study, preferably from ≥ one research group II-3: Evidence derived from multiple time series with or without the intervention III: Opinions of respected authorities based on clinical experience, descriptive studies, or expert committee reports	review and a consensus process.  Recommendation grading system (SIGN): A: Evidence from ≥ one meta-analysis, systematic review, or RCT that is pertinent to the target population B: Evidence from well-conducted clinical trials that are pertinent to the target population and demonstrate consistency of results; or evidence extrapolated from meta-analyses, systematic reviews or RCTs C: Evidence from opinions of respected authorities or expert committee reports	externally by a clinician and academic.

ACS = American College of Surgeons; APSIC = Asia Pacific Society of Infection Control; ASHP = American Society of Health-System Pharmacists; CDC = Centers for Disease Control and Prevention; CINAHL = Cumulative Index to Nursing and Allied Health Literature; CPSI = Canadian Patient Safety Institute; GDG = Guidelines Development Group; GRADE = Grading of Recommendations, Assessment, Development, and Evaluation; HICPAC = Healthcare Infection Control Practices Advisory Committee; MRSA = methicillin-resistant Staphylococcus aureus; NICE = National Institute for Health and Care Excellence; NR = not reported; RCT = randomized controlled trial; SIGN = Scottish Intercollegiate Guidelines Network; SR = systematic review; SREG = Systematic Reviews Expert Group; SSI = surgical site infection.



# **Appendix 3: Critical Appraisal of Included Publications**

Table 3: Strengths and Limitations of Guidelines using AGREE II<sup>6</sup>

	l		Guid	eline		
Item	APSIC, 2019 <sup>11</sup>	NICE, 2019 <sup>9</sup>	ACS, 2017 <sup>12</sup>	CDC, 2017 <sup>1</sup>	WHO, 2016 <sup>8</sup>	Ministry of Health Malaysia, 2015 <sup>10</sup>
Domain 1: Scope and Pur	pose					
The overall objective(s) of the guideline is (are) specifically described.	Yes	Yes	Yes	Yes	Yes	Yes
2. The health question(s) covered by the guideline is (are) specifically described.	Yes	Yes	Yes	Yes	Yes	Yes
3. The population (patients, public, etc.) to whom the guideline is meant to apply is specifically described.	Yes	Yes	Yes	Yes	Yes	Yes
Domain 2: Stakeholder In	volvement					
4. The guideline development group includes individuals from all relevant professional groups.	Yes	Yes	Yes	Yes	Yes	Yes
5. The views and preferences of the target population (patients, public, etc.) have been sought.	No	Yes	No	Yes	Yes	No
6. The target users of the guideline are clearly defined.	Apparent but not explicitly described	Yes	Apparent but not explicitly described	Yes	Yes	Yes
Domain 3: Rigour of Deve	lopment					
7. Systematic methods were used to search for evidence.	Unclear	Yes	Unclear	Yes	Yes	Yes
8. The criteria for selecting the evidence are clearly described.	No	Yes	No	Yes	Yes	Yes
9. The strengths and limitations of the body of evidence are clearly described.	Yes	Yes	No	Yes	Yes	Yes
10. The methods for formulating the recommendations are clearly described.	Unclear	Yes	Yes	Yes	Yes	Yes



			Guid	eline		
ltem	APSIC, 2019 <sup>11</sup>	NICE, 2019 <sup>9</sup>	ACS, 2017 <sup>12</sup>	CDC, 2017 <sup>1</sup>	WHO, 2016 <sup>8</sup>	Ministry of Health Malaysia, 2015 <sup>10</sup>
11. The health benefits, side effects, and risks have been considered in formulating the recommendations.	Yes	Yes	Yes	Yes	Yes	Yes
12. There is an explicit link between the recommendations and the supporting evidence.	Yes	Yes	Yes	Yes	Yes	Yes
13. The guideline has been externally reviewed by experts prior to its publication.	Yes	Yes	Yes	Yes	Yes	Yes
14. A procedure for updating the guideline is provided.	No	Yes	No	Yes	Yes	No
Domain 4: Clarity of Prese	entation	'			,	
15. The recommendations are specific and unambiguous.	Yes	Yes	Yes	Yes	Yes	Yes
16. The different options for management of the condition or health issue are clearly presented.	Yes	Yes	Yes	Yes	Yes	Yes
17. Key recommendations are easily identifiable.	Yes	Yes	Yes	Yes	Yes	Yes
Domain 5: Applicability		,				
18. The guideline describes facilitators and barriers to its application.	No	Yes	No	Yes	Yes	No
19. The guideline provides advice and/or tools on how the recommendations can be put into practice.	No	Yes	No	Yes	Yes	No
20. The potential resource implications of applying the recommendations have been considered.	No	Yes	No	No	Yes	No
21. The guideline presents monitoring and/or auditing criteria.	Yes	Yes	Yes	Yes	Yes	Yes
Domain 6: Editorial Indepe	endence					



	Guideline						
Item	APSIC, 2019 <sup>11</sup>	NICE, 2019 <sup>9</sup>	ACS, 2017 <sup>12</sup>	CDC, 2017 <sup>1</sup>	WHO, 2016 <sup>8</sup>	Ministry of Health Malaysia, 2015 <sup>10</sup>	
22. The views of the funding body have not influenced the content of the guideline.	Yes	Yes	Yes	Yes	Yes	Yes	
23. Competing interests of guideline development group members have been recorded and addressed.	Yes	Yes	Yes	Yes	Yes	Yes	

ACS = American College of Surgeons; AGREE II = Appraisal of Guidelines for Research and Evaluation II; APSIC = Asia Pacific Society of Infection Control; CDC = Centers for Disease Control and Prevention; NICE = National Institute for Health and Care Excellence.



# **Appendix 4: Main Study Findings and Authors' Conclusions**

### **Table 4: Summary of Recommendations in Included Guidelines**

Recommendations and supporting evidence	Quality of evidence and strength of recommendation
Asia Pacific Society of Infection Control (APSIC) Guide	line, 2019 <sup>11</sup>
Evidence-based guideline regarding preoperative measures for the prevention of SSIs.	
<ul> <li>1. Preoperative washing: "It is necessary for patients who will undergo surgery to have at least 1 preoperative bath with soap (antimicrobial or non-antimicrobial)."<sup>11</sup> (p2)</li> <li>This recommendation was informed by one article<sup>17</sup></li> </ul>	Quality of evidence: II     Recommendation strength: B
<ul> <li>2. Mechanical bowel preparation: "Combination mechanical bowel preparation and oral antibiotic preparation are recommended for all elective colorectal surgery in adults." (p2)</li> <li>This recommendation was informed by one article 18</li> </ul>	2. Quality of evidence: I Recommendation strength: A
<ul> <li>3. Hair removal:</li> <li>a. "Hair removal should be avoided unless hair interferes with the operative procedure."</li> <li>b. "If hair removal is necessary, a razor should be avoided and an electric clipper should be used."</li> <li>c. "No recommendation regarding the timing of hair removal by clipper is made." (p3)</li> <li>These recommendations were informed by two articles 19,20</li> </ul>	3. a. Quality of evidence: III Recommendation strength: B b. Quality of evidence: I Recommendation strength: A c. Quality of evidence: III Recommendation strength: C
4. <b>Nasal decolonization:</b> a. "Hospitals should evaluate their SSI, <i>Staphylococcus aureus (S. aureus)</i> and MRSA rates, and mupirocin resistant rate, if available, to determine whether implementation of a screening program is appropriate." b. "Patients undergoing cardiothoracic and orthopedic surgery with known nasal carriage of <i>S. aureus</i> should receive perioperative intranasal application of mupirocin 2% ointment with or without a combination of CHG body wash." ( <i>p3</i> )  • These recommendations were informed by three articles <sup>21-23</sup>	4. a. Quality of evidence: II Recommendation strength: B  b. Quality of evidence: I Recommendation strength: A
<ul> <li>5. Skin antiseptic preparation: "Alcohol based skin antiseptic preparations should be used, unless contraindicated." (p4)</li> <li>This recommendation was informed by three articles<sup>24-26</sup></li> </ul>	5. Quality of evidence: I Recommendation strength: A
6. <b>Antibiotic prophylaxis:</b> "Administration of prophylaxis antimicrobials should only be performed when indicated. Prophylactic antimicrobials should be administered within 1 h before incision for all antimicrobials except vancomycin and fluoroquinolones where it should be administered within 2 h. Re-dosing should be considered to maintain adequate tissue levels based on agent half-life. A single dose of antimicrobial prophylactic is adequate for most surgical procedures." 11 (p4)  • These recommendations were informed by two articles 12,26	6. Quality of evidence: I Recommendation strength: A
National Institute for Health and Care Excellence (NICE) G	uideline, 2019 <sup>9</sup>
Evidence-based guideline regarding preoperative measures for the prevention and treatment of SSIs. These recommendations were informed by a review of the published SRs and RCTs.  1. Preoperative washing and nasal decolonization:	The wording of recommendations reflects the recommendation strength Offer/Advise: Strong recommendation (i.e., clear evidence of benefit) Consider: Evidence of benefit is less

certain



Recommendations and supporting evidence	Quality of evidence and
	strength of recommendation
<ul> <li>a. "Advise patients to shower or have a bath (or help patients to shower, bath or bed bath) using soap, either the day before, or on the day of, surgery." (p5)</li> <li>This recommendation was informed by a SR referenced in the original 2008 guideline 13</li> </ul>	a. The quality of evidence was assessed, but was not reported online
<ul> <li>b. "Consider nasal mupirocin in combination with a chlorhexidine body wash before procedures in which <i>Staphylococcus aureus</i> is a likely cause of a surgical site infection. This should be locally determined and take into account the type of procedure, individual patient risk factors, the increased risk of side effects in preterm infants, and the potential impact of infection. Maintain surveillance on antimicrobial resistance associated with the use of mupirocin."9 (<i>p6</i>)</li> <li>These recommendations were informed by an evidence review conducted by NICE<sup>14</sup></li> </ul>	b. The quality of the supporting evidence ranged from very low to high
2. <b>Hair removal:</b> "Do not use hair removal routinely to reduce the risk of surgical site infection. If hair has to be removed, use electric clippers with a single-use head on the day of surgery. Do not use razors for hair removal, because they increase the risk of surgical site infection." 9 (p6)	2. The quality of evidence was assessed, but was not reported online
<ul> <li>These recommendations were informed by a SR and an RCT referenced in the original 2008 guideline<sup>13</sup></li> </ul>	
<ul> <li>3. Mechanical bowel preparation: "Do not use mechanical bowel preparation routinely to reduce the risk of surgical site infection." (p7)</li> <li>These recommendations were informed by 12 RCTs referenced in the original 2008 guideline<sup>13</sup></li> </ul>	3. The quality of evidence was assessed, but was not reported online
4. Antibiotic prophylaxis:  a. "Give antibiotic prophylaxis to patients before: clean surgery involving the placement of a prosthesis or implant, clean-contaminated surgery, and contaminated surgery."  b. "Do not use antibiotic prophylaxis routinely for clean non-prosthetic uncomplicated	4. The quality of evidence was assessed, but was not reported online
surgery."  c. "Use the local antibiotic formulary and always take into account the potential adverse effects when choosing specific antibiotics for prophylaxis."  d. "Consider giving a single dose of antibiotic prophylaxis intravenously on starting anaesthesia. However, give prophylaxis earlier for operations in which a tourniquet is	
used."  e. "Before giving antibiotic prophylaxis, take into account the timing and pharmacokinetics (for example, the serum half-life) and necessary infusion time of the antibiotic. Give a repeat dose of antibiotic prophylaxis when the operation is longer than the half-life of the antibiotic given."  f. "Give antibiotic treatment (in addition to prophylaxis) to patients having surgery on a	
dirty or infected wound." g. "Inform patients before the operation, whenever possible, if they will need antibiotic prophylaxis, and afterwards if they have been given antibiotics during their operation." (p. 7)	
<ul> <li>These recommendations were informed by 18 SRs and 22 RCTs referenced in the original 2008 guideline<sup>13</sup></li> </ul>	
<ul> <li>5. Antiseptic skin preparation:</li> <li>a. "Prepare the skin at the surgical site immediately before incision using an antiseptic preparation."</li> <li>b. "Be aware of the risks of using skin antiseptics in babies, in particular the risk of severe chemical injuries with the use of chlorhexidine (both alcohol-based and aqueous solutions) in preterm babies."</li> </ul>	5. The quality of the supporting evidence ranged from very low to high



Recommendations ar	Recommendations and supporting evidence					
c. "When deciding which antiseptic skin pre table [below]." (p9)  These recommendations were info NICE <sup>14</sup>						
Indication	Antiseptic Option					
First option	Alcohol-based solution of chlorhexidine					
If chlorhexidine is contraindicated	Alcohol-based solution of povidone-iodine					
If the area is close to a mucous membrane	Aqueous solution of chlorhexidine					
If an alcohol-based solution and chlorhexidine are both unsuitable	Aqueous solution of povidone-iodine					
American College of S	urgeons (ACS) & Surgical Infection Socie	ty Guideline, 2017 <sup>12</sup>				
Evidence-based guideline regarding preoper treatment of SSIs. These recommendations literature.		The quality of the evidence and the strength of the recommendations were not reported.				
Preoperative washing: "Routine preoper part of a decolonization protocol or preoper pathogen concentrations, but has not been     This recommendation was informed."	shown to reduce SSI."12 (p61)					
2. MRSA screening: a. "Decision about whether or not to implen and decolonization protocols should depen b. "Clinical practice guidelines from the Am recommend screening and nasal mupirocin patients before total joint replacement and c. "MRSA bundles (screening, decolonization highly effective if adhered to, otherwise the d. "No standard decolonization protocol sumupirocin alone vs nasal mupirocin plus che. "Decolonization protocols should be comeffective." f. "Vancomycin should not be administered						
patients."12 (p61)  • These recommendations were info	ormed by four articles <sup>23,28-30</sup>					
Bowel preparations: "Combination med recommended for all elective colectomies."     This recommendation was informed.						
4. Hair removal: "Hair removal should be a hair removal is necessary, clippers should learn This recommendation was informed."						
5. <b>Skin preparation:</b> "Alcohol-containing p contraindication exists (eg fire hazard, surfaclear superior agent (chlorhexidine vs iodin cannot be included in the preparation, chlorunless contraindications exist." (p62)  • This recommendation was informed.						



Recommendations and supporting evidence	Quality of evidence and strength of recommendation
<ul> <li>6. Antibiotic prophylaxis:</li> <li>a. "Administer prophylactic antibiotics only when indicated."</li> <li>b. "Choice of prophylactic antibiotic should be dictated by the procedure and pathogens most likely to cause SSI."</li> <li>c. "Prophylactic antibiotic should be administered within 1 hour before incision or within 2 hours for vancomycin or fluoroquinolones."</li> <li>d. "Prophylactic antibiotic dosing should be weight-adjusted.</li> <li>e. "Re-dose antibiotics to maintain adequate tissue levels based on agent half-life or for every 1,500 mL blood loss."</li> <li>f. "There is no evidence that prophylactic antibiotic administration after incision closure decreases SSI risk; prophylactic antibiotics should be discontinued at time of incision closure (exceptions include implant-based breast reconstruction, joint arthroplasty, and cardiac procedures where optimal duration of antibiotic therapy remains unknown)."12 (p62)</li> <li>• These recommendations were informed by three articles<sup>26,34,35</sup></li> <li>7. Glucose control: "Hyperglycemia in the immediate preoperative period is associated with an increased risk of SSI. Target perioperative blood glucose should be between</li> </ul>	
with an increased risk of SSI. Target perioperative blood glucose should be between 110 to 150 mg/dL in all patients, regardless of diabetic status, except in cardiac surgery patients where the target perioperative blood glucose is <180 mg/dL. Target blood glucose rates <110 mg/dL have been tied to adverse outcomes and increased episodes of hypoglycemia and do not decrease SSI risk." (p62)  This recommendation was informed by 13 articles 26,36-47	
Centers for Disease Control and Prevention (CDC) Guid	eline, 2017 <sup>1</sup>
Evidence-based guideline regarding preoperative measures for the prevention of SSIs. The systematic review used to inform specific recommendations was presented in the online supplemental section. <sup>48</sup>	
1. Antibiotic prophylaxis (parenteral): a. "Administer preoperative antimicrobial agents only when indicated based on published clinical practice guidelines and timed such that a bactericidal concentration of the agents is established in the serum and tissues when the incision is made." b. "No further refinement of timing can be made for preoperative antimicrobial agents based on clinical outcomes."	Quality of evidence: B (accepted practice) Recommendation strength: I
<ul> <li>No recommendation/unresolved issue</li> <li>c. "Administer the appropriate parenteral prophylactic antimicrobial agents before skin incision in all cesarean section procedures."</li> <li>d. "The literature search did not identify randomized controlled trials that evaluated the benefits and harms of weight-adjusted parenteral antimicrobial prophylaxis dosing and its effect on the risk of SSI." (p786)</li> <li>No recommendation/unresolved issue</li> </ul>	c. Quality of evidence: A (high-quality evidence) Recommendation strength: I
<ul> <li>2. Glucose control:</li> <li>a. "Implement perioperative glycemic control and use blood glucose target levels less than 200 mg/dL in patients with and without diabetes."</li> <li>b. "The search did not identify randomized controlled trials that evaluated lower (&lt;200mg/dL) or narrower blood glucose target levels than recommended in this guideline nor the optimal timing, duration, or delivery method of perioperative glycemic control for the prevention of SSI." (p787)</li> <li>No recommendation/unresolved issue</li> </ul>	2. a. Quality of evidence: A (high to moderate-quality evidence) Recommendation strength: I



Recommendations and supporting evidence	Quality of evidence and strength of recommendation
<ul> <li>3. Preoperative washing:</li> <li>a. "Advise patients to shower or bathe (full body) with soap (antimicrobial or nonantimicrobial) or an antiseptic agent on at least the night before the operative day."</li> <li>b. "Randomized controlled trial evidence suggested uncertain trade-offs between the benefits and harms regarding the optimal timing of the preoperative shower or bath, the total number of soap or antiseptic agent applications, or the use of chlorhexidine gluconate washcloths for the prevention of SSI." (p787)</li> <li>No recommendation/unresolved issue</li> </ul>	3. a. Quality of evidence: B (accepted practice) Recommendation strength: I
WHO Guideline, 2016 <sup>8</sup>	
Evidence-based guideline regarding preoperative measures for the prevention of SSIs and SSI-related deaths.	
<ul> <li>1. Preoperative washing:</li> <li>a. "It is good clinical practice for patients to bathe or shower prior to surgery. The panel suggests that either a plain or antimicrobial soap may be used for this purpose."</li> <li>This recommendation was informed by a SR<sup>49</sup></li> <li>b. "The panel decided not to formulate a recommendation on the use of chlorhexidine gluconate (CHG)-impregnated cloths for the purpose of reducing SSI due to the limited and very low quality evidence." (p58)</li> </ul>	Quality of evidence: Moderate quality     Recommendation strength:     Conditional
<ul> <li>2. Nasal decolonization:</li> <li>a. "The panel recommends that patients undergoing cardiothoracic and orthopaedic surgery with known nasal carriage of <i>S. aureus</i> should receive perioperative intranasal applications of mupirocin 2% ointment with or without a combination of CHG body wash."</li> <li>b. "The panel suggests considering to treat also patients with known nasal carriage of <i>S. aureus</i> undergoing other types of surgery with perioperative intranasal applications of mupirocin 2% ointment with or without a combination of CHG body wash." (p63)</li> <li>These recommendations were informed by a SR<sup>50</sup></li> </ul>	2. a. Quality of evidence: Moderate quality Recommendation strength: Strong b. Quality of evidence: Moderate quality Recommendation strength: Conditional
<ul> <li>3. Optimal time for antibiotic prophylaxis:</li> <li>a. "The panel recommends the administration of SAP prior to the surgical incision when indicated (depending on the type of operation)."</li> <li>b. "The panel recommends the administration of SAP within 120 minutes before incision, while considering the half-life of the antibiotic." (p71)</li> <li>These recommendations were informed by a SR<sup>51</sup></li> </ul>	3. a. Low quality Recommendation strength: Strong b. Moderate quality Recommendation strength: Strong
<ul> <li>4. Mechanical bowel preparation and oral antibiotics:</li> <li>a. "The panel suggests that preoperative oral antibiotics combined with mechanical bowel preparation (MBP) should be used to reduce the risk of SSI in adult patients undergoing elective colorectal surgery."</li> <li>b. "The panel recommends that MBP alone (without administration of oral antibiotics) should not be used for the purpose of reducing SSI in adult patients undergoing elective colorectal surgery."<sup>8</sup> (p76)</li> <li>These recommendations were informed by a SR<sup>52</sup></li> <li>5. Hair removal: "The panel recommends that in patients undergoing any surgical</li> </ul>	4. a. Quality of evidence: Moderate quality Recommendation strength: Conditional b. Moderate quality Recommendation strength: Strong  5. Quality of evidence: Moderate
procedure, hair should either not be removed or, if absolutely necessary, it should be removed only with a clipper. Shaving is strongly discouraged at all times, whether preoperatively or in the operating room (OR)."8 (p82)  • These recommendations were informed by a SR <sup>53</sup>	quality Recommendation strength: Strong



Recommendations and supporting evidence	Quality of evidence and strength of recommendation	
6. <b>Skin antiseptic:</b> "The panel recommends alcohol-based antiseptic solutions based on CHG for surgical site skin preparation in patients undergoing surgical procedures." <sup>8</sup>	Quality of evidence: Low to moderate quality     Recommendation strength: Strong	
<ul> <li>(p87)</li> <li>These recommendations were informed by a SR<sup>54</sup></li> </ul>		
Ministry of Health Malaysia, 2015 <sup>10</sup>		
Evidence-based guideline regarding antibiotic prophylaxis for the prevention of oral SSIs.		
<ul> <li>1. "Antibiotic prophylaxis is indicated for all surgical procedures carried out on medically compromised patients especially those with ASA score of 3 or more." (p3)</li> <li>This recommendation was informed by two SRs, one guideline, and one primary study</li> </ul>	Quality of evidence: Level I, II-2, and III     Recommendation strength: Grade B	
<ul> <li>2. "Antibiotic prophylaxis is not indicated for clean surgery in healthy patients." (p4)</li> <li>This recommendation was informed by five primary studies</li> </ul>	2. Quality of evidence: Level II-2, II-3, and III	
<ul> <li>3. "Antibiotic prophylaxis is not indicated for lower third molar surgery."<sup>10</sup> (<i>p5</i>)</li> <li>This recommendation was informed by one SR and seven primary studies</li> </ul>	Recommendation strength: Grade B 3. Quality of evidence: Level I, II-2, and III	
<ul> <li>4. "Antibiotic prophylaxis is not recommended for routine periodontal surgery." (p5)</li> <li>This recommendation was informed by two primary studies</li> <li>5. "Antibiotic prophylaxis may be indicated for minor surgery with a high degree of difficulty in which the duration of the surgery is predicted to be long." (p6)</li> <li>This recommendation was informed by two primary studies</li> </ul>	Recommendation strength: Grade A 4. Quality of evidence: Level III Recommendation strength: Grade B 5. Quality of evidence: Level II-2 and III Recommendation strength: Grade B	
<ul> <li>6. "Antibiotic prophylaxis is indicated for surgery to place dental implants." (p6)</li> <li>This recommendation was informed by two SRs</li> </ul>	Quality of evidence: Level I     Recommendation strength: Grade A	
<ul> <li>7. "Antibiotic prophylaxis is indicated for minor oral surgical procedures in which a bone graft is inserted." (p7)</li> <li>This recommendation was informed by one primary study</li> </ul>	7. Quality of evidence: Level I Recommendation strength: Grade A	
8. "Antibiotic prophylaxis is indicated for major clean contaminated maxillofacial surgery." 10 (p7)	Quality of evidence: Level I, II-2,     and III	
<ul> <li>This recommendation was informed by two SRs and three primary studies</li> </ul>	Recommendation strength: Grade A	
9. "Antibiotic prophylaxis is indicated in all forms of head and neck cancer surgery." ( <i>p8</i> )	9. Quality of evidence: Level II-1, II-3,	
This recommendation was informed by four primary studies	and III Recommendation strength: Grade A	
<ul> <li>10. "Antibiotic prophylaxis is indicated for open reduction and internal fixation of facial bone fractures. Antibiotics should not be continued postoperatively." (p9)</li> <li>This recommendation was informed by four primary studies</li> </ul>	10. Quality of evidence: Level II-2, II-3, and III Recommendation strength: Grade B	
<ul> <li>11.</li> <li>a. "Amoxicillin, Penicillin G and Clindamycin are appropriate choices of antibiotics for oral surgical prophylaxis." (p13)</li> <li>This recommendation was informed by six primary studies</li> <li>b. "Cloxacillin, cefazolin or clindamycin should be considered if the surgery extends onto the skin." (p13)</li> <li>This recommendation was informed by one primary study</li> </ul>	11. Quality of evidence: Level I, II-1, II-3, and III a. Recommendation strength: Grade B b. Recommendation strength: Grade C	



Recommendations and supporting evidence	Quality of evidence and strength of recommendation
<ul> <li>12. "The dose of antibiotic to be administered for surgical prophylaxis should be at the full therapeutic dose of the antibiotic." (p13) <ul> <li>This recommendation was informed by two primary studies</li> </ul> </li> <li>13. "The first dose of the antibiotic should be administered within 60 minutes prior to the surgical incision." (p15) <ul> <li>This recommendation was informed by seven primary studies</li> </ul> </li> <li>14. <ul> <li>a. "Additional doses of prophylactic antibiotics should be administered if the length of surgery exceeds either two half-lives or half the therapeutic interval of the drug." (p15)</li> <li>b. "The additional dose strength should be the same as the initial prophylactic dose of the antibiotic." (p16)</li> <li>These recommendations were informed by three primary studies</li> </ul> </li> </ul>	12. Quality of evidence: Level II-2 and II-3 Recommendation strength: Grade B 13. Quality of evidence: Level II-2, II-3, and III Recommendation strength: Grade B 14. Quality of evidence: Level II-1, II-2, and II-3 a. Recommendation strength: Grade B b. Recommendation strength: Grade C

ACS = American College of Surgeons; APBI = accelerated partial breast radiation; APSIC = Asia Pacific Society of Infection Control; ASA = American Society of Anesthesiologists; CDC = Centers for Disease Control and Prevention; CHG = chlorhexidine gluconate; MBP = mechanical bowel preparation; MRSA = methicillin-resistant Staphylococcus aureus; NICE = National Institute for Health and Care Excellence; NR = not reported; NRS = non-randomized study; OP = operating room; PPA = perioperative prophylactic antibiotic; RCT = randomized controlled trial; SAP = surgical antibiotic prophylaxis; SR = systematic review; SSI = surgical site infection.



# **Appendix 5: Additional References of Potential Interest**

### **Previous CADTH Reports**

- Li KX, Grobelna A. Decolonization for the treatment of methicillin resistant staphylococcus aureus: clinical effectiveness and guidelines [CADTH rapid response report: reference list]. Ottawa (ON): CADTH; 2019 Jan: <a href="https://cadth.ca/sites/default/files/pdf/htis/2019/RA1005%20ARO%20Decolonization%20Final.pdf">https://cadth.ca/sites/default/files/pdf/htis/2019/RA1005%20ARO%20Decolonization%20Final.pdf</a> Accessed 2020 Mar 3.
- Banerjee S, Argáez C. Topical antibiotics for infection prevention: a review of the clinical effectiveness and guidelines [CADTH rapid response report: summary with critical appraisal]. Ottawa (ON): CADTH; 2017 Mar: <a href="https://www.ncbi.nlm.nih.gov/books/NBK487430/">https://www.ncbi.nlm.nih.gov/books/NBK487430/</a> Accessed 2020 Mar 3.
- Chlorhexidine gluconate wipes for infection prevention in acute and critical care: a review of clinical effectiveness and cost-effectiveness [CADTH rapid response report: summary with critical appraisal]. Ottawa (ON): CADTH; 2016 Apr: <a href="https://cadth.ca/sites/default/files/pdf/htis/apr-2016/RC0769%20Chlorhexidine%20Wipes%20Final.pdf">https://cadth.ca/sites/default/files/pdf/htis/apr-2016/RC0769%20Chlorhexidine%20Wipes%20Final.pdf</a> Accessed 2020 Mar 3.

### Overview of Systematic Reviews

 Gillespie BM, Walker RM, McInnes E, et al. Preoperative and postoperative recommendations to surgical wound care interventions: a systematic meta-review of Cochrane reviews. *Int J Nurs Stud.* 2020 Feb;102:103486. <u>PubMed: PM31810020</u>

### Guidelines

### Unclear Methodology

- Consensus guideline on preoperative antibiotics and surgical site infection in breast surgery. Columbia (MD): The American Society of Breast Surgeons; 2018: <a href="https://www.breastsurgeons.org/docs/statements/Consensus-Guideline-on-Preoperative-Antibiotics-and-Surgical-Site-Infection-in-Breast-Surgery.pdf">https://www.breastsurgeons.org/docs/statements/Consensus-Guideline-on-Preoperative-Antibiotics-and-Surgical-Site-Infection-in-Breast-Surgery.pdf</a> Accessed 2020 Mar 3.
- Bonnar P, Dhar P, Rotstein O, et al. Surgical site infection prevention. Toronto (ON): University of Toronto Best Practice in Surgery; 2017 Sep: <a href="http://bestpracticeinsurgery.ca/wp-content/uploads/2017/11/SSI-BPS-CPG-Nov20.pdf">http://bestpracticeinsurgery.ca/wp-content/uploads/2017/11/SSI-BPS-CPG-Nov20.pdf</a> Accessed 2020 Mar 3.
- Centre for Health Protection. Recommendations on prevention of surgical site infection. 2<sup>nd</sup> ed. Kowloon, Hong Kong: Scientific Committee on Infection Control, and Infection Control Branch, Centre for Health Protection, Department of Health; 2017 Sep: <a href="https://www.chp.gov.hk/files/pdf/recommendations">https://www.chp.gov.hk/files/pdf/recommendations</a> on prevention of surgical site infection 2nd edition.pdf Accessed 2020 Mar 3.



### Summary of WHO 2016 Guideline

 Allegranzi B, Bischoff P, de Jonge S, et al. New WHO recommendations on preoperative measures for surgical site infection prevention: an evidence-based global perspective. *Lancet Infect Dis.* 2016 Dec;16(12):e276-e287.
 PubMed: PM27816413

### Alternative Intervention

Don't order peri-operative antibiotics beyond a 24-hour post-operative period for noncomplicated instrumented cases in patients who are not at high risk for infection or
wound contamination. Administration of a single preoperative dose for spine cases
without instrumentation is adequate. Toronto (ON): Choosing Wisely Canada; 2019 Jul:
<a href="https://choosingwiselycanada.org/spine/">https://choosingwiselycanada.org/spine/</a> Accessed 2020 Mar 3.