G-30 – What is the most optimal intra-operative skin preparation agent for patients undergoing major orthopedic procedures?

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Response/Recommendation: There does not appear to be a superior intra-operative skin preparation agent for patients undergoing major orthopaedic procedures. Rather, the decision to use a particular agent (chlorhexidine gluconate versus iodine-based preparations) should be based on the most likely pathogens of a particular surgical site and should be combined with isopropyl alcohol.

Strength of Recommendation: Moderate

Delegate Vote:

Rationale:

Surgical site infection (SSI) prevention has been extensively investigated, and several steps are recommended for peri-operative skin preparation, including pre-operative washing, appropriate hair removal from incision site, and reducing skin recolonization [1]. Cleansing the surgical site using a chlorhexidine gluconate (CHG) or povidone-iodine (PI) solution prior to incision is considered routine and reduces bacterial load at the surgical site [2-5]. Bacterial cell membranes are disrupted by CHG, whereas PI acts through bacterial protein denaturation [6]. In the most recent Cochrane meta-analysis regarding preoperative skin antisepsis, the authors concluded that the current literature lacked evidence to support the use of a particular solution [7]. However, the consensus was that the skin preparation agent should contain alcohol based on recommendations from the Centers for Disease Control, International Consensus Group Meeting, and prior studies [5, 7-13].

Since this meta-analysis, Charehbili et al. published their cluster-randomized crossover trial consisting of 3,665 patients who underwent breast, vascular, colorectal, gallbladder, or orthopaedic surgery. Patients received either 0.5% CHG or 1% PI in an alcohol-based solution. The authors noted no statistically significant difference between the overall incidence of SSI between groups [14]. The Prep-it Investigators published two randomized controlled trials evaluating the optimal preoperative antiseptic solution for patients who suffered fractures. One study evaluated 1,638 patients who received either 4% CHG or 1% PI and underwent fixation for open fractures. The authors found no statistically significant difference in rates of SSI between groups [15]. The authors' subsequent multicenter randomized controlled trial consisted of 6,785 patients with closed fractures who received eithr 2% CHG or 0.7% PI for skin preparation. The authors noted that skin antisepsis with PI in alcohol resulted in a lower proportion of SSIs than those who received CHG in alcohol [16]. In contrast, Ritter et al. published a prospective randomized trial of 279 consecutive patients who were randomized to receive 2% CHG with alcohol or 1% PI with alcohol for skin preparation intraoperatively. The authors found that skin antisepsis with CHG for surgery for closed trauma of the lower leg and foot led to statistically significant fewer complications of wound healing compared to PI [17]. While a randomized

controlled trial by Shadid et al. demonstrated no statistically significant differences in postoperative wound complications or SSIs in 49 patients undergoing clean foot surgery who received skin preparation with either 0.5% CHG or 1% PI, both with alcohol [18].

Regarding spine surgery, Patrick et al. published their randomized controlled trial of 407 patients who received 2% CHG in alcohol or 1% PI in alcohol for intraoperative skin preparation prior to spine surgery. The authors found that the sequential application of PI and CHG more effectively reduced contamination of surgical wounds than PI alone based on intraoperative skin cultures [10].

Regarding hip and knee arthroplasty, three randomized controlled trials have been published since the prior Cochrane meta-analysis. Cho et al. performed a randomized controlled trial of 150 patients undergoing primary total knee arthroplasty who received intraoperative skin antisepsis with either 10% PI, 2% CHG after 1% PI, or 1% PI after 2% CHG, all with alcohol. Based on rates of bacterial cultures after skin preparation, the authors noted that CHG after PI or PI after CHG were superior to PI alone [19]. Peel et al. published their cluster randomized controlled trial of 780 patients undergoing primary total knee or hip arthroplasty who received either 0.5% CHG in alcohol or 1% PI in alcohol for intraoperative preparation. The authors noted no differences in superficial wound complications between the two groups, but on secondary analysis, the PI group had greater efficacy for preventing SSI [20]. Droll et al. published their randomized controlled trial of 105 patients undergoing primary total hip arthroplasty who received either 2% CHG or 0.7% PI for intraoperative skin preparation. Using intraoperative cultures, the authors noted that PI was more effective than CHG at eliminating skin flora at the hip on initial application, but the two solutions were equally effective by time of closure [21].

Regarding the shoulder, Blonna et al. published their randomized controlled trial of 40 patients with proximal humerus fractures who underwent intraoperative skin preparation with either 1% PI or 4% CHG followed by PI. The authors noted that although both approaches reduced *Staphylococcus aureus* and *Propionibacterium acnes* skin burden, the double skin preparation was more effective against coagulase-negative *Staphylococcus aureus* [22]. Additionally, Dorfel et al. found that 3.2% PI in alcohol showed benefits over 2% CHG in alcohol in aerobic and anaerobic flora culture positivity of the shoulder based on their randomized cross-over study [23]. A randomized controlled trial of 22 patients by Hancock et al. found no statistically significant reduction in growth of Propionibacterium acnes over 14 days with CHG + benzoyl peroxide compared with CHG alone [24].

Given the heterogeneity of data based on surgical site, there does not appear to be an optimal intra-operative skin preparation agent for all patients undergoing major orthopaedic procedures. In accordance with the review by Dockery et al. on preoperative skin preparation agents and surgical sites, the decision to use a particular skin preparation agent should be based on the most likely pathogens of a surgical site and be combined with isopropyl alcohol [25].

References

1. Wilson J, Topley K, Stott D, Neachell J, Gallagher R: **The OneTogether collaborative** approach to reduce the risk of surgical site infection: identifying the challenges to assuring best practice. *J Infect Prev* 2015, **16**(3):118-125.

- 2. Mangram AJ, Horan TC, Pearson ML, Silver LC, Jarvis WR: Guideline for Prevention of Surgical Site Infection, 1999. Centers for Disease Control and Prevention (CDC) Hospital Infection Control Practices Advisory Committee. *Am J Infect Control* 1999, 27(2):97-132; quiz 133-134; discussion 196.
- 3. Durani P, Leaper D: **Povidone-iodine: use in hand disinfection, skin preparation and antiseptic irrigation**. *Int Wound J* 2008, **5**(3):376-387.
- 4. Milstone AM, Passaretti CL, Perl TM: Chlorhexidine: expanding the armamentarium for infection control and prevention. Clin Infect Dis 2008, 46(2):274-281.
- 5. Berrios-Torres SI, Umscheid CA, Bratzler DW, Leas B, Stone EC, Kelz RR, Reinke CE, Morgan S, Solomkin JS, Mazuski JE *et al*: **Centers for Disease Control and Prevention Guideline for the Prevention of Surgical Site Infection, 2017**. *JAMA Surg* 2017, **152**(8):784-791.
- 6. Bednarek RS, Nassereddin A, Ramsey ML: **Skin Antiseptics**. In: *StatPearls*. edn. Treasure Island (FL); 2025.
- 7. Dumville JC, McFarlane E, Edwards P, Lipp A, Holmes A, Liu Z: **Preoperative skin antiseptics for preventing surgical wound infections after clean surgery**. *Cochrane Database Syst Rev* 2015, **2015**(4):CD003949.
- 8. Saltzman MD, Nuber GW, Gryzlo SM, Marecek GS, Koh JL: **Efficacy of surgical preparation solutions in shoulder surgery**. *J Bone Joint Surg Am* 2009, **91**(8):1949-1953.
- 9. Xu PZ, Fowler JR, Goitz RJ: **Prospective Randomized Trial Comparing the Efficacy of Surgical Preparation Solutions in Hand Surgery**. *Hand* (NY) 2017, **12**(3):258-264.
- 10. Patrick S, McDowell A, Lee A, Frau A, Martin U, Gardner E, McLorinan G, Eames N: Antisepsis of the skin before spinal surgery with povidone iodine-alcohol followed by chlorhexidine gluconate-alcohol versus povidone iodine-alcohol applied twice for the prevention of contamination of the wound by bacteria: a randomised controlled trial. *Bone Joint J* 2017, **99-B**(10):1354-1365.
- 11. Cheng K, Robertson H, St Mart JP, Leanord A, McLeod I: **Quantitative analysis of bacteria in forefoot surgery: a comparison of skin preparation techniques**. *Foot Ankle Int* 2009, **30**(10):992-997.
- 12. Bibbo C, Patel DV, Gehrmann RM, Lin SS: Chlorhexidine provides superior skin decontamination in foot and ankle surgery: a prospective randomized study. Clin Orthop Relat Res 2005, 438:204-208.
- 13. Atkins GJ, Alberdi MT, Beswick A, Blaha JD, Bingham J, Cashman J, Chen AF, Cooper AM, Cotacio GL, Fraguas T *et al*: **General Assembly, Prevention, Surgical Site Preparation: Proceedings of International Consensus on Orthopedic Infections**. *J Arthroplasty* 2019, **34**(2S):S85-S92.
- 14. Charehbili A, Koek MBG, de Mol van Otterloo JCA, Bronkhorst M, van der Zwaal P, Thomassen B, Waasdorp EJ, Govaert JA, Bosman A, van den Bremer J *et al*: Cluster-randomized crossover trial of chlorhexidine-alcohol versus iodine-alcohol for prevention of surgical-site infection (SKINFECT trial). *BJS Open* 2019, **3**(5):617-622.
- 15. Investigators P-I: Aqueous skin antisepsis before surgical fixation of open fractures (Aqueous-PREP): a multiple-period, cluster-randomised, crossover trial. *Lancet* 2022, **400**(10360):1334-1344.
- 16. Investigators P-I, Sprague S, Slobogean G, Wells JL, O'Hara NN, Thabane L, Mullins CD, Harris AD, Wood A, Viskontas D *et al*: **Skin Antisepsis before Surgical Fixation of Extremity Fractures**. *N Engl J Med* 2024, **390**(5):409-420.

- 17. Ritter B, Herlyn PKE, Mittlmeier T, Herlyn A: **Preoperative skin antisepsis using chlorhexidine may reduce surgical wound infections in lower limb trauma surgery when compared to povidone-iodine a prospective randomized trial**. *Am J Infect Control* 2020, **48**(2):167-172.
- 18. Shadid MB, Speth M, Voorn GP, Wolterbeek N: Chlorhexidine 0.5%/70% Alcohol and Iodine 1%/70% Alcohol Both Reduce Bacterial Load in Clean Foot Surgery: A Randomized, Controlled Trial. *J Foot Ankle Surg* 2019, 58(2):278-281.
- 19. Cho MR, Choi WK, Che SH, Song SK: **Efficacy of skin preparation solutions in patients with total knee replacement: A randomized controlled trial**. *J Orthop Surg* (*Hong Kong*) 2023, **31**(1):10225536231165358.
- 20. Peel TN, Dowsey MM, Buising KL, Cheng AC, Choong PFM: Chlorhexidine-alcohol versus iodine-alcohol for surgical site skin preparation in an elective arthroplasty (ACAISA) study: a cluster randomized controlled trial. Clin Microbiol Infect 2019, 25(10):1239-1245.
- 21. Droll KP, Abouassaly M, Cullinan C, Puskas D, Dubois S: **Efficacy of surgical skin preparation solutions in hip arthroplasty: a prospective randomized trial**. *Can J Surg* 2022, **65**(6):E756-E762.
- 22. Blonna D, Allizond V, Bellato E, Banche G, Cuffini AM, Castoldi F, Rossi R: Single versus Double Skin Preparation for Infection Prevention in Proximal Humeral Fracture Surgery. *Biomed Res Int* 2018, **2018**:8509527.
- 23. Dorfel D, Maiwald M, Daeschlein G, Muller G, Hudek R, Assadian O, Kampf G, Kohlmann T, Harnoss JC, Kramer A: Comparison of the antimicrobial efficacy of povidone-iodine-alcohol versus chlorhexidine-alcohol for surgical skin preparation on the aerobic and anaerobic skin flora of the shoulder region. *Antimicrob Resist Infect Control* 2021, **10**(1):17.
- 24. Hancock DS, Rupasinghe SL, Elkinson I, Bloomfield MG, Larsen PD: **Benzoyl peroxide** + **chlorhexidine versus chlorhexidine alone skin preparation to reduce Propionibacterium acnes: a randomized controlled trial**. *ANZ J Surg* 2018, **88**(11):1182-1186.
- 25. Dockery DM, Allu S, Vishwanath N, Li T, Berns E, Glasser J, Spake CSL, Antoci V, Born CT, Garcia DR: Review of Pre-Operative Skin Preparation Options Based on Surgical Site in Orthopedic Surgery. Surg Infect (Larchmt) 2021, 22(10):1004-1013.