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## **Abstract**

Surgical-site infection (SSI) after total joint arthroplasty (TJA) continues to pose a challenge and place a substantial burden on patients, surgeons, and the healthcare system. Given the estimated 1.0% to 2.5% annual incidence of SSI after TJA, orthopedists should be cognizant of preventive measures that can help optimize patient outcomes. Advances in surgical technique, sterile protocol, and operative procedures have been instrumental in minimizing SSIs and may account for the recent plateau in rising rates. In this review, we identify and discuss preoperative, intraoperative, and postoperative actions that can be taken to help reduce the incidence of SSIs, and we highlight the economic implications of SSIs that occur after TJA.

urgical-site infection (SSI), a potentially devastating complication of lower extremity total joint arthroplasty (TJA), is estimated to occur in 1% to 2.5% of cases annually.¹ Infection after TJA places a significant burden on patients, surgeons, and the healthcare system. Revision procedures that address infection after total hip arthroplasty (THA) are associated with more hospitalizations, more operations, longer hospital stay, and higher outpatient costs in comparison with primary THAs and revision surgeries for aseptic loosening.² If left untreated, a SSI can go deeper into

the joint and develop into a periprosthetic infection, which can be disastrous and costly. A periprosthetic joint infection study that used 2001 to 2009 Nationwide Inpatient Sample (NIS) data found that the cost of revision procedures increased to \$560 million from \$320 million, and was projected to reach \$1.62 billion by 2020.³ Furthermore, society incurs indirect costs as a result of patient disability and loss of wages and productivity.² Therefore, the issue of infection after TJA is even more crucial in our cost-conscious healthcare environment.

Patient optimization, advances in surgical technique, sterile protocol, and operative procedures have been effective in reducing bacterial counts at incision sites and minimizing SSIs. As a result, infection rates have leveled off after rising for a decade.<sup>4</sup> Although infection prevention modalities have their differences, routine use is fundamental and recommended by the Hospital Infection Control Practices Advisory Committee.<sup>5</sup> Furthermore, both the US Centers for Disease Control and Prevention (CDC) and its Healthcare Infection Control Practices Advisory Committee<sup>6,7</sup> recently updated their SSI prevention guidelines by incorporating evidence-based methodology, an element missing from earlier recommendations.

The etiologies of postoperative SSIs have been discussed ad nauseam, but there are few reports summarizing the literature on infection prevention modalities. In this review, we identify and examine SSI prevention strategies as they relate to lower extremity TJA. Specifically, we discuss the literature on the preoperative, intraoperative, and postoperative actions that can be taken to reduce the incidence of SSIs after TJA. We also highlight the economic implications of SSIs that occur after TJA.

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Table 1. Summary of Studies Reporting on Preoperative Measures to Prevent Surgical-Site Infection

	Study	Patients/ Studies, N	Operation	Interventions	Results
Preoperative skin preparation	Hayek et al <sup>10</sup> (1987)	2015	Various elective inpatient procedures	Preparation with chlorhexidine vs unmedicated soap vs placebo cloth	Decreased infection rates 6 wk after surgery with chlorhexidine (9% vs 11.7% vs 12.8%; P < .05)
	Murray et al <sup>11</sup> (2011)	100	Shoulder surgery	2% chlorhexidine gluconate cloth vs standard shower (soap & water)	Decreased positive surgical-site culture rate with 2% chlorhexidine (66% vs 94%; <i>P</i> = .0008)
	Darouiche et al <sup>12</sup> (2010)	849	Various	Preparation with chlorhexidine-alcohol vs povidone-iodine	Lower rate of SSI with chlorhexidine- alcohol (9.5% vs 16.1%; <i>P</i> = .004; RR, 0.59; 95% CI, 0.41-0.85)
	Zywiel et al <sup>13</sup> (2011)	136	TKA	Advanced cutaneous disinfection protocol vs standard perioperative disinfection	Lower rate of SSI in advanced protocol group (0% vs 3%)
	Kapadia et al <sup>14</sup> (2013)	2545	THA	At-home chlorhexidine cloths plus standard perioperative preparation vs standard perioperative preparation alone	Lower incidence of SSI with at-home chlorhexidine cloths (0.5% vs 1.7%; $P = .04$ )
	Johnson et al <sup>15</sup> (2013)	2293	TKA	At-home chlorhexidine cloths plus standard perioperative preparation vs standard perioperative preparation alone	Lower incidence of SSI with at-home chlorhexidine cloths (0.6% vs 2.2%; $P = .02$ )
	Kapadia et al <sup>16</sup> (2016)	554	THA, TKA	Preadmission 2% chlorhexidine gluconate— impregnated cloths vs preadmission soap & water bath (standard of care)	Lower incidence of infection with chlorhexidine (0.4% vs 2.9%; <i>P</i> = .049; OR, 8.15; 95% CI, 1.01-65.6)

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## **Methods**

For this review, we performed a literature search with PubMed, EBSCOhost, and Scopus. We looked for reports published between the inception of each database and July 2016. Combinations of various search terms were used: surgical site, infection, total joint arthroplasty, knee, hip, preoperative, intraoperative, perioperative, postoperative, preparation, nutrition, ventilation, antibiotic, body exhaust suit, gloves, drain, costs, economic, and payment.

Our search identified 195 abstracts. Drs. Mistry and Chughtai reviewed these to determine which articles were relevant. For any uncertainties, consensus was reached with the help of Dr. Delanois. Of the 195 articles, 103 were potentially relevant, and 54 of the 103 were excluded for being not relevant to preventing SSIs after TJA or for being written in a language other than English. The references in the remaining articles were assessed, and those with potentially relevant titles were selected for abstract review. This step provided another 35 articles. After

## **Take-Home Points**

- SSIs after TJA pose a substantial burden on patients, surgeons, and the healthcare system.
- While different forms of preoperative skin preparation have shown varying outcomes after TJA, the importance of preoperative patient optimization (nutritional status, immune function, etc) cannot be overstated.
- Intraoperative infection prevention measures include cutaneous preparation, gloving, body exhaust suits, surgical drapes, OR staff traffic and ventilation flow, and antibiotic-loaded cement.
- Antibiotic prophylaxis for dental procedures in TJA patients continues to remain a controversial issue with conflicting recommendations.
- SSIs have considerable financial costs and require increased resource utilization. Given the significant economic burden associated with TJA infections, it is imperative for orthopedists to establish practical and cost-effective strategies to prevent these devastating complications.

Table 1. Summary of Studies Reporting on Preoperative Measures to Prevent Surgical-Site Infection (continued)

	Study	Patients/ Studies, N	Operation	Interventions	Results
Patient optimization	Greene et al <sup>18</sup> (1991)	217	THA, TKA	N/A	Up to 7-fold higher rate of infection after TJA in patients with preoperative malnutrition (lymphocyte count of <1500 cells/mm³ or albumin level of <3.5 g/dL)
	Alfargieny et al <sup>20</sup> (2015)	135	THA, TKA	N/A	Preoperative serum albumin levels were only nutritional biomarker predictive of SSI (P = .011)
	Sørensen <sup>23</sup> (2012)	Meta- analysis of 140 cohort studies	Various	N/A	Smokers had increased risk of wound complications (OR, 2.27; 95% CI, 1.82-2.84), delayed wound healing and dehiscence (OR, 2.07; 95% CI, 1.53-2.81), and infection (OR, 1.79; 95% CI, 1.57-2.04)
					Decreased incidence of SSI with smoking cessation (OR, 0.43; 95% CI, 0.21-0.85)
	Wong et al <sup>24</sup> (2012)	Meta- analysis of 25 studies	Various s	N/A	Decreased incidence of SSI with smoking cessation for at least 4 wk before surgery (OR, 0.69; 95% CI, 0.56-0.84)
	Pugely et al <sup>25</sup> (2015)	25,235	Primary & revision TJA	N/A	Increased risk of SSI with BMI >40 kg/m² (OR, 1.9; 95% CI, 1.3-2.9), electrolyte disturbance (OR, 2.4; 95% CI, 1.0-6.0), and hypertension diagnosis (OR, 1.5; 95% CI, 1.1-2.0)
	Han & Kang <sup>26</sup> (2013)	115	TKA	N/A	Patients with hemoglobin A <sub>1c</sub> levels >8% more likely to have wound complication (OR, 6.07; 95% CI, 1.12-33.0)
	Hwang et al <sup>27</sup> (2015)	462	TKA	N/A	Patients with hemoglobin $A_{1c}$ levels >8% more likely to have SSI (OR, 6.1; 95% CI, 1.6-23.4; $P = .008$ )
					Patients with fasting blood glucose level of $\geq$ 200 mg/dL more likely to have SSI (OR, 9.2; 95% CI, 2.2-38.2; $P = .038$ )
	Moroski et al <sup>29</sup> (2015)	289	Primary & revision TJA	Preoperative 5-day course of intranasal mupirocin decolonization	Reduction in MRSA ( $P = .0073$ ) and MSSA ( $P = .0341$ ) colonization on day of surgery
	Rao et al <sup>30</sup> (2011)	2284	Primary & revision TJA	Preoperative 5-day course of chlorhexidine bath and intranasal mupirocin	Reduction in SSI with implementation of decolonization protocol ( $P = .009$ )

Abbreviations: BMI, body mass index; CI, confidence interval; MRSA, methicillin-resistant Staphylococcus aureus; MSSA, methicillin-sensitive Staphylococcus aureus; N/A, not applicable; OR, odds ratio; RR, relative risk; SSI, surgical-site infection; THA, total hip arthroplasty; TJA, total joint arthroplasty; TKA, total knee arthroplasty.

all exclusions, 48 articles remained. We discuss these in the context of preoperative, intraoperative, and postoperative measures and economic impact.

## **Results**

## **Preoperative Measures**

**Skin Preparation.** Preoperative skin preparation methods include standard washing and rinsing,

antiseptic soaps, and iodine-based or chlorhexidine gluconate-based antiseptic showers or skin cloths. Iodine-based antiseptics are effective against a wide range of Gram-positive and Gram-negative bacteria, fungi, and viruses. These agents penetrate the cell wall, oxidize the microbial contents, and replace those contents with free iodine molecules. Iodophors are free iodine molecules associ-

ated with a polymer (eg, polyvinylpyrrolidone); the iodophor povidone-iodine is bactericidal. Chlorhexidine gluconate-based solutions are effective against many types of yeast, Grampositive and Gram-negative bacteria, and a wide variety of viruses. Both solutions are useful. Patients with an allergy to iodine can use chlorhexidine. Table 1 summarizes the studies on preoperative measures for preventing SSIs.

There is no shortage of evidence of the efficacy of these antiseptics in minimizing the incidence of SSIs. Hayek and colleagues<sup>10</sup> prospectively analyzed use of different preoperative skin preparation methods in 2015 patients. Six weeks after surgery, the infection rate was significantly lower with use of chlorhexidine than with use of an unmedicated bar of soap or placebo cloth (9% vs 11.7% and 12.8%, respectively; P < .05). In a study of 100 patients, Murray and colleagues<sup>11</sup> found the overall bacterial culture rate was significantly lower for those who used a 2% chlorhexidine gluconate cloth before shoulder surgery than for those who took a standard shower with soap (66% vs 94%; P = .0008). Darouiche and colleagues<sup>12</sup> found the overall SSI rate was significantly lower for 409 surgical patients prepared with chlorhexidine-alcohol than for 440 prepared with povidone-iodine (9.5% vs 16.1%; P = .004; relative risk [RR], 0.59; 95% confidence interval [CI], 0.41-0.85).

Chlorhexidine gluconate-impregnated cloths have also had promising results, which may be attributed to general ease of use and potentially improved patient adherence. Zywiel and colleagues<sup>13</sup> reported no SSIs in 136 patients who used these cloths at home before total knee arthroplasty (TKA) and 21 SSIs (3.0%) in 711 patients who did not use the cloths. In a study of 2545 THA patients, Kapadia and colleagues<sup>14</sup> noted a significantly lower incidence of SSIs with at-home preoperative use of chlorhexidine cloths than with only in-hospital perioperative skin preparation (0.5% vs 1.7%; P = .04). In 2293 TKAs, Johnson and colleagues<sup>15</sup> similarly found a lower incidence of SSIs with at-home preoperative use of chlorhexidine cloths (0.6% vs 2.2%; P = .02). In another prospective, randomized trial, Kapadia and colleagues<sup>16</sup> compared 275 patients who used chlorhexidine cloths the night before and the morning of lower extremity TJA surgery with 279 patients who underwent standardof-care preparation (preadmission bathing with antibacterial soap and water). The chlorhexidine cohort had a lower overall incidence of infection (0.4% vs 2.9%; P = .049), and the standard-of-care

cohort had a stronger association with infection (odds ratio [OR], 8.15; 95% CI, 1.01-65.6).

Patient Optimization. Poor nutritional status may compromise immune function, potentially resulting in delayed healing, increased risk of infection, and, ultimately, negative postoperative outcomes. Malnutrition can be diagnosed on the basis of a prealbumin level of <15 mg/dL (normal, 15-30 mg/ dL), a serum albumin level of <3.4 g/dL (normal, 3.4-5.4 g/dL), or a total lymphocyte count under 1200 cells/µL (normal, 3900-10,000 cells/µL). 17-19 Greene and colleagues<sup>18</sup> found that patients with preoperative malnutrition had up to a 7-fold higher rate of infection after TJA. In a study of 135 THAs and TKAs, Alfargieny and colleagues<sup>20</sup> found preoperative serum albumin was the only nutritional biomarker predictive of SSI (P = .011). Furthermore, patients who take immunomodulating medications (eg, for inflammatory arthropathies) should temporarily discontinue them before surgery in order to lower their risk of infection.21

Smoking is well established as a major risk factor for poor outcomes after surgery. It is postulated that the vasoconstrictive effects of nicotine and the hypoxic effects of carbon monoxide contribute to poor wound healing.22 In a meta-analysis of 4 studies, Sørensen<sup>23</sup> found smokers were at increased risk for wound complications (OR, 2.27; 95% CI, 1.82-2.84), delayed wound healing and dehiscence (OR, 2.07; 95% CI, 1.53-2.81), and infection (OR, 1.79; 95% CI, 1.57-2.04). Moreover, smoking cessation decreased the incidence of SSIs (OR, 0.43; 95% CI, 0.21-0.85). A metaanalysis by Wong and colleagues<sup>24</sup> revealed an inflection point for improved outcomes in patients who abstained from smoking for at least 4 weeks before surgery. Risk of infection was lower for these patients than for current smokers (OR, 0.69; 95% CI, 0.56-0.84).

Other comorbidities contribute to SSIs as well. In their analysis of American College of Surgeons National Surgical Quality Improvement Program registry data on 25,235 patients who underwent primary and revision lower extremity TJA, Pugely and colleagues<sup>25</sup> found that, in the primary TJA cohort, body mass index (BMI) of >40 kg/m² (OR, 1.9; 95% CI, 1.3-2.9), electrolyte disturbance (OR, 2.4; 95% CI, 1.0-6.0), and hypertension diagnosis (OR, 1.5; 95% CI, 1.1-2.0) increased the risk of SSI within 30 days. Furthermore, diabetes mellitus delays collagen synthesis, impairs lymphocyte function, and impairs wound healing, which may lead to poor recovery and higher risk of infection.<sup>26</sup> In a

study of 167 TKAs performed in 115 patients with type 2 diabetes mellitus, Han and Kang<sup>26</sup> found that wound complications were 6 times more likely in those with hemoglobin  $A_{1c}$  (HbA $_{1c}$ ) levels higher than 8% than in those with lower HbA $_{1c}$  levels (OR, 6.07; 95% CI, 1.12-33.0). In a similar study of 462 patients with diabetes, Hwang and colleagues<sup>27</sup> found a higher likelihood of superficial SSIs in patients with HbA $_{1c}$  levels  $\geq$ 8% (OR, 6.1; 95% CI, 1.6-23.4; P = .008). This association was also found in patients with a fasting blood glucose level of  $\geq$ 200 mg/dL (OR, 9.2; 95% CI, 2.2-38.2; P = .038).

Methicillin-resistant Staphylococcus aureus (MRSA) is thought to account for 10% to 25% of all periprosthetic infections in the United States.<sup>28</sup> Nasal colonization by this pathogen increases the risk for SSIs; however, decolonization protocols have proved useful in decreasing the rates of colonization. Moroski and colleagues<sup>29</sup> assessed the efficacy of a preoperative 5-day course of intranasal mupirocin in 289 primary or revision TJA patients. Before surgery, 12 patients had positive MRSA cultures, and 44 had positive methicillin-sensitive S aureus (MSSA) cultures. On day of surgery, a significant reduction in MRSA (P = .0073) and MSSA (P = .0341) colonization was noted. Rao and colleagues<sup>30</sup> found that the infection rate decreased from 2.7% to 1.2% in 2284 TJA patients treated with a decolonization protocol (P = .009).

## Intraoperative Measures

Cutaneous Preparation. The solutions used in perioperative skin preparation are similar to those used preoperatively: povidone-iodine, alcohol, and chlorhexidine. The efficacy of these preparations varies. Table 2 summarizes the studies on intraoperative measures for preventing SSIs. In a prospective study, Saltzman and colleagues<sup>31</sup> randomly assigned 150 shoulder arthroplasty patients to one of 3 preparations: 0.75% iodine scrub with 1% iodine paint (Povidone-Iodine; Tyco Healthcare Group), 0.7% iodophor with 74% iodine povacrylex (DuraPrep; 3M Health Care), or chlorhexidine gluconate with 70% isopropyl alcohol (ChloraPrep; Enturia). All patients had their skin area prepared and swabbed for culture before incision. Although no one in any group developed a SSI, patients in the chlorhexidine group had the lowest overall incidence of positive skin cultures. That incidence (7%) and the incidence of patients in the iodophor group (19%) were significantly lower than that of patients in the iodine group (31%) (P < .001 for

both). Conversely, another study<sup>32</sup> found a higher likelihood of SSI with chlorhexidine than with povidone-iodine (OR, 4.75; 95% CI, 1.42-15.92; *P* = .012). This finding is controversial, but the body of evidence led the CDC to recommend use of an alcohol-based solution for preoperative skin preparation.<sup>6</sup>

The literature also highlights the importance of technique in incision-site preparation. In a prospective study, Morrison and colleagues<sup>33</sup> randomly assigned 600 primary TJA patients to either (1) use of alcohol and povidone-iodine before draping, with additional preparation with iodine povacrylex (DuraPrep) and isopropyl alcohol before application of the final drape (300-patient intervention group) or (2) only use of alcohol and povidone-iodine before draping (300-patient control group). At the final follow-up, the incidence of SSI was significantly lower in the intervention group than in the control group (1.8% vs 6.5%; P = .015). In another study that assessed perioperative skin preparation methods, Brown and colleagues<sup>34</sup> found that airborne bacteria levels in operating rooms were >4 times higher with patients whose legs were prepared by a scrubbed, gowned leg-holder than with patients whose legs were prepared by an unscrubbed, ungowned leg-holder (P = .0001).

Hair Removal. Although removing hair from surgical sites is common practice, the literature advocating it varies. A large comprehensive review<sup>35</sup> revealed no increased risk of SSI with removing vs not removing hair (RR, 1.65; 95% Cl, 0.85-3.19). On the other hand, some hair removal methods may affect the incidence of infection. For example, use of electric hair clippers is presumed to reduce the risk of SSIs, whereas traditional razors may compromise the epidermal barriers and create a pathway for bacterial colonization. 5,36,37 In the aforementioned review,35 SSIs were more than twice as likely to occur with hair removed by shaving than with hair removed by electric clippers (RR, 2.02; 95% CI, 1.21-3.36). Cruse and Foord<sup>38</sup> found a higher rate of SSIs with hair removed by shaving than with hair removed by clipping (2.3% vs 1.7%). Most surgeons agree that, if given the choice, they would remove hair with electric clippers rather than razors.

**Gloves.** Almost all orthopedists double their gloves for TJA cases. Over several studies, the incidence of glove perforation during orthopedic procedures has ranged from 3.6% to 26%, <sup>39-41</sup> depending on the operating room personnel and glove layering studied. Orthopedists must know this startling finding, as surgical glove perforation



Table 2. Summary of Studies Reporting on Intraoperative Measures to Prevent Surgical-Site Infection

	Study	Patients/ Studies, N	Operation	Interventions/Endpoints	Results
Perioperative skin preparation	Saltzman et al <sup>31</sup> (2009)	150	TSA	Preparation with iodine vs iodophor vs chlorhexidine gluconate	Lower incidence of positive skin cultures with chlorhexidine gluconate or iodophor preparation vs iodine preparation ( $P < .001$ for both)
	Carroll et al <sup>32</sup> (2014)	964	THA, TKA	0.5% chlorhexidine vs 1% iodine	Increased likelihood of SSI with chlorhexidine preparation (OR, 4.75; 95% CI, 1.42-15.92; <i>P</i> = .012)
	Morrison et al <sup>33</sup> (2016)	600	Primary TJA	Skin preparation before draping and additional preparation before application of final drape vs skin preparation only before draping	Reduced incidence of SSI with repeat skin preparation (1.8% vs 6.5%; $P = .015$ )
	Brown et al <sup>34</sup> (1996)	29	THA, TKA	Unscrubbed, ungowned leg- holder vs scrubbed, gowned leg-holder during preparation	Airborne bacteria levels >4 times higher with unscrubbed, ungowned leg-holder $(P = .0001)$
Hair removal	Tanner et al <sup>35</sup> (2006)	Review of 11 studies	Various	Removal of hair from surgical site	No increased risk of SSI with hair removal vs no hair removal (RR, 1.65; 95% CI, 0.85-3.19)
					Increased risk of SSI with shaving vs use of electrical clippers (RR, 2.02; 95% CI, 1.21-3.36)
	Cruse & Foord <sup>38</sup> (1973)	23,649 surgical wounds	Various	Hair removal by shaving vs hair removal with electric clippers	Higher rate of SSI with shaving vs use of electric clippers (2.3% vs 1.7%)
Gloves and body exhaust suits	Carter et al <sup>42</sup> (2012)	3863 gloves	Primary and revision TJA	Incidence of glove perforation	Higher incidence of glove perforation with revision vs primary TJA (8.9% vs $3.7\%$ ; $P = .04$ )
	Hooper et al <sup>47</sup> (2011)	88,311	THA, TKA	Body exhaust suits	Increase in early revision THA for deep infection with use of body exhaust suits (0.186% vs 0.064%; P < .0001)
					Increase in early revision TKA for deep infection with use of body exhaust suits (0.243% vs 0.098%; P < .0001)
	Miner et al <sup>48</sup> (2007)	8288	TKA	Body exhaust suits	No difference in risk of SSI with use of body exhaust suits (RR, 0.75; 95% CI, 0.34-1.62)

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is associated with an increase in the rate of SSIs, from 1.7% to 5.7%. <sup>38</sup> Carter and colleagues <sup>42</sup> found the highest risk of glove perforation occurs when double-gloved attending surgeons, adult reconstruction fellows, and registered nurses initially assist during primary and revision TJA. In their study, outer and inner glove layers were perforated 2.5% of the time. All outer-layer perforations were noticed, but inner-layer perforations went unnoticed 81% of the time, which poses a potential hazard for both patients and healthcare personnel.

In addition, there was a significant increase in the incidence of glove perforations for attending surgeons during revision TJA vs primary TJA (8.9% vs 3.7%; P = .04). This finding may be expected given the complexity of revision procedures, the presence of sharp bony and metal edges, and the longer operative times. Giving more attention to glove perforations during arthroplasties may mitigate the risk of SSI. As soon as a perforation is noticed, the glove should be removed and replaced.

Body Exhaust Suits. Early TJAs had infection

Table 2. Summary of Studies Reporting on Intraoperative Measures to Prevent Surgical-Site Infection (continued)

	Study	Patients/ Studies, N	Operation	Interventions/Endpoints	Results
Surgical drapes	Blom et al <sup>50</sup> (2000)	N/A	N/A	7 different surgical drapes	Increased bacterial penetration with woven cloth drapes vs nonwoven disposable drapes
	Blom et al <sup>51</sup> (2002)	N/A	N/A	Effect of different wetting agents on bacterial penetration and growth with reusable polyester/cotton surgical drapes	Enhanced bacterial penetration rate and heavy growth (>100,000 colony-forming units) after 30 minutes with drapes wet with blood or normal saline
	Fairclough et al <sup>52</sup> (1986)	235	Hip surgery requiring implant	Povidone-iodine preparation vs iodophor-impregnated adhesive drape	Surgical-site bacterial contamination reduced from 15% to 1.6% with iodophor-impregnated drape
	Webster & Alghamdi <sup>53</sup> (2007)	4195	Various	Adhesive drapes with/ without antimicrobial properties vs no drapes	No reduced risk of SSI with iodine- impregnated drapes (RR, 1.03; 95% CI, 0.06-1.66; $P = .89$ )
					Increased risk of infection with overall use of adhesive draping (RR, 1.23; 95% CI, 1.02-1.48; $P = .03$ )
Ventilation flow	Evans <sup>54</sup> (2011)	Review of 48 studies	Various TJAs	Laminar vs nonlaminar airflow	Laminar airflow can decrease bacterial count in air and wound
	Lidwell et al <sup>46</sup> (1982)	8055	TKA	Laminar vs nonlaminar airflow	Lower incidence of SSI with laminar- airflow operating rooms (0.6% vs 2.3%; P < .001)
	Miner et al <sup>48</sup> (2007)	8288	TKA	Laminar vs nonlaminar airflow	No difference in risk of SSI with laminar- airflow operating rooms (RR, 1.57; 95% CI, 0.75-3.31)
	Hooper et al <sup>47</sup> (2011)	88,311	THA, TKA	Laminar vs nonlaminar airflow	Increased incidence of early infections with laminar airflow for both TKA (0.193% vs 0.100%; <i>P</i> = .019) and THA (0.148% vs 0.061%; <i>P</i> < .001)
Staff traffic volume	Lynch et al <sup>55</sup> (2009)	28	Various	Frequency of opening operating room door	Door may remain open for up to 20 minutes per case
	Young & O'Regan <sup>56</sup> (2010)	46	Cardiac	Frequency of opening operating room door	Door is open for almost 10.7% of each hour
	Pryor & Messmer <sup>57</sup> (1998)	2864	Various	Personnel in operating room	SSI rate of 6.27% with >17 people entering vs 1.52% with <9 people entering
Antibiotic prophylaxis	Gorenoi et al <sup>60</sup> (2010)	Review of 10 studies	TKA	Antibiotic prophylaxis vs no prophylaxis	Antibiotic prophylaxis is highly effective, but a particular antibiotic could not be recommended
	AlBuhairan et al <sup>61</sup> (2008)	Review of 26 studies	TJA	Antibiotic prophylaxis vs no prophylaxis	8% reduction in absolute risk of wound infection and 81% reduction in RR with antibiotic prophylaxis ( <i>P</i> < .00001)
Antibiotic-loaded bone cement	Parvizi et al <sup>64</sup> (2008)	Meta- analysis of 19 studies	THA	ALBC vs regular bone cement	Decreased infection rate with ALBC (1.2% vs 2.3%)
	Namba et al <sup>65</sup> (2009)	22,889	TKA	ALBC vs regular bone cement	Higher incidence of deep infection with ALBC (1.4% vs 0.7%; $P = .002$ )
	Zhou et al <sup>66</sup> (2015)	Meta- analysis of 5 studies	TKA	ALBC vs regular bone cement	No difference in incidence of deep SSI (1.32% vs 1.89%; RR, 0.75; 95% CI, 0.43-1.33; <i>P</i> = .33)

Abbreviations: ALBC, antibiotic-laden bone cement; CI, confidence interval; N/A, not applicable; OR, odds ratio; RR, relative risk; SSI, surgical-site infection; THA, total hip arthroplasty; TJA, total joint arthroplasty; TKA, total knee arthroplasty; TSA, total shoulder arthroplasty.

rates approaching 10%.43 Bacterial-laden particles shed from surgical staff were postulated to be the cause, 44,45 and this idea prompted the development of new technology, such as body exhaust suits, which have demonstrated up to a 20-fold reduction in airborne bacterial contamination and decreased incidence of deep infection, from 1% to 0.1%, as compared with conventional surgical attire.46 However, the efficacy of these suits was recently challenged. Hooper and colleagues<sup>47</sup> assessed >88,000 TJA cases in the New Zealand Joint Registry and found a significant increase in early revision THA for deep infection with vs without use of body exhaust suits (0.186% vs 0.064%; P < .0001). The incidence of revision TKAs for deep infections with use of these suits was similar (0.243% vs 0.098%; P < .001). Many of the surgeons surveyed indicated their peripheral vision was limited by the suits, which may contribute to sterile field contamination. By contrast, Miner and colleagues<sup>48</sup> were unable to determine an increased risk of SSI with use of body exhaust suits (RR, 0.75; 95% CI, 0.34-1.62), though there was a trend toward more infections without suits. Moreover, these suits are effective in reducing mean air bacterial counts (P =.014), but it is not known if this method correlates with mean wound bacterial counts (r = -.011) and therefore increases the risk of SSI.49

Surgical Drapes. Surgical draping, including cloths, iodine-impregnated materials, and woven or unwoven materials, is the standard of care worldwide. The particular draping technique usually varies by surgeon. Plastic drapes are better barriers than cloth drapes, as found in a study by Blom and colleagues<sup>50</sup>: Bacterial growth rates were almost 10 times higher with use of wet woven cloth drapes than with plastic surgical drapes. These findings were supported in another, similar study by Blom and colleagues<sup>51</sup>: Wetting drapes with blood or normal saline enhanced bacterial penetration. In addition, wetting drapes with chlorhexidine or iodine reduced but did not eliminate bacterial penetration. Fairclough and colleagues<sup>52</sup> emphasized that iodine-impregnated drapes reduced surgical-site bacterial contamination from 15% to 1.6%. However, a Cochrane review<sup>53</sup> found these drapes had no effect on the SSI rate (RR, 1.03; 95% CI, 0.06-1.66; P = .89), though the risk of infection was slightly higher with adhesive draping than with no drape (RR, 1.23; 95% CI, 1.02-1.48; P = .03).

**Ventilation Flow.** Laminar-airflow systems are widely used to prevent SSIs after TJA. Horizontal-flow and vertical-flow ventilation provides

and maintains ultra-clean air in the operating room. Evans<sup>54</sup> found the bacterial counts in the air and the wound were lower with laminar airflow than without this airflow. The amount of airborne bacterial colony-forming units and dust large enough to carry bacteria was reduced to 1 or 2 particles more than 2  $\mu$ m/m³ with use of a typical laminar-airflow system. In comparing 3922 TKA patients in laminar-airflow operating rooms with 4133 patients in conventional rooms, Lidwell and colleagues<sup>46</sup> found a significantly lower incidence of SSIs in patients in laminar-airflow operating rooms (0.6% vs 2.3%; P < .001).

Conversely, Miner and colleagues<sup>48</sup> did not find a lower risk of SSI with laminar-airflow systems (RR, 1.57; 95% CI, 0.75-3.31). In addition, in their analysis of >88,000 cases from the New Zealand Joint Registry, Hooper and colleagues<sup>47</sup> found that the incidence of early infections was higher with laminar-airflow systems than with standard airflow systems for both TKA (0.193% vs 0.100%; P = .019) and THA (0.148% vs 0.061%; P < .001). They postulated that vertically oriented airflow may have transmitted contaminated particles into the surgical sites. Additional evidence may be needed to resolve these conflicting findings and determine whether clean-air practices provide significant clinical benefit in the operating room.

Staff Traffic Volume. When staff enters or exits the operating room or makes extra movements during a procedure, airflow near the wound is disturbed and no longer able to remove sufficient airborne pathogens from the sterile field. The laminarairflow pattern may be disrupted each time the operating room doors open and close, potentially allowing airborne pathogens to be introduced near the patient. Lynch and colleagues<sup>55</sup> found the operating room door opened almost 50 times per hour, and it took about 20 seconds to close each time. As a result, the door may remain open for up to 20 minutes per case, causing substantial airflow disruption and potentially ineffective removal of airborne bacterial particles. Similarly, Young and O'Regan<sup>56</sup> found the operating room door opened about 19 times per hour and took 20 seconds to close each time. The theater door was open an estimated 10.7% of each hour of sterile procedure. Presence of more staff also increases airborne bacterial counts. Pryor and Messmer<sup>57</sup> evaluated a cohort of 2864 patients to determine the effect of number of personnel in the operating theater on the incidence of SSIs. Infection rates were 6.27% with >17 different people entering the room and

Table 3. Summary of Studies Reporting on Postoperative Measures to Prevent Surgical-Site Infection

	Study	Patients, N	Operation	Interventions	Results
Antibiotic prophylaxis and drain management	Berbari et al <sup>71</sup> (2010)	678	THA, TKA	Antibiotic prophylaxis for dental procedures	No decreased risk of subsequent THA infection (OR, 0.9; 95% CI, 0.5-1.6)
					No decreased risk of subsequent TKA infection (OR, 1.2; 95% CI, 0.7-2.2)
	Jaberi et al <sup>74</sup> (2008)	10,325	THA, TKA	Treatments for draining wound to prevent subsequent SSI	Of 300 patients (2.9%) with persistent drainage, 83 (28%) required surgical débridement

Abbreviations: CI, confidence interval; OR, odds ratio; SSI, surgical-site infection; THA, total hip arthroplasty; TKA, total knee arthroplasty.

1.52% with <9 different people entering the room. Restricting the number of people in the room may be one of the easiest and most efficient ways to prevent SSI.

Systemic Antibiotic Prophylaxis. Perioperative antibiotic use is vital in minimizing the risk of infection after TJA. The Surgical Care Improvement Project recommended beginning the first antimicrobial dose either within 60 minutes before surgical incision (for cephalosporin) or within 2 hours before incision (for vancomycin) and discontinuing the prophylactic antimicrobial agents within 24 hours after surgery ends. 58,59 However, Gorenoi and colleagues<sup>60</sup> were unable to recommend a way to select particular antibiotics, as they found no difference in the effectiveness of various antibiotic agents used in TKA. A systematic review by AlBuhairan and colleagues<sup>61</sup> revealed that antibiotic prophylaxis (vs no prophylaxis) reduced the absolute risk of a SSI by 8% and the relative risk by 81% (P < 0.0001). These findings are supported by evidence of the efficacy of perioperative antibiotics in reducing the incidence of SSI.62,63 Antibiotic regimens should be based on susceptibility and availability, depending on hospital prevalence of infections. Even more, patients should receive prophylaxis in a timely manner. Finally, bacteriostatic antibiotics (vancomycin) should not be used on their own for preoperative prophylaxis.

Antibiotic Cement. Antibiotic-loaded bone cement (ALBC), which locally releases antimicrobials in high concentration, is often used in revision joint arthroplasty, but use in primary joint arthroplasty remains controversial. In a study of THA patients, Parvizi and colleagues<sup>64</sup> found infection rates of 1.2% with 2.3% with and without use of ALBC, respectively. Other studies have had opposing results. Namba and colleagues<sup>65</sup> evaluated 22,889 primary TKAs, 2030 (8.9%) of which used ALBC. The incidence of deep infection was significantly

higher with ALBC than with regular bone cement (1.4% vs 0.7%; P = .002). In addition, a meta-analysis of >6500 primary TKA patients, by Zhou and colleagues, <sup>66</sup> revealed no significant difference in the incidence of deep SSIs with use of ALBC vs regular cement (1.32% vs 1.89%; RR, 0.75; 95% CI, 0.43-1.33; P = .33). More evidence is needed to determine the efficacy of ALBC in primary TJA. International Consensus Meeting on Periprosthetic Joint Infection participants recommended use of ALBC in high-risk patients, including patients who are obese or immunosuppressed or have diabetes or a prior history of infection. <sup>67</sup>

## **Postoperative Measures**

Antibiotic Prophylaxis. The American Academy of Orthopaedic Surgeons (AAOS) and the American Dental Association (ADA) have suggestions for antibiotic prophylaxis for patients at increased risk for infection. As of 2015, the ADA no longer recommends antibiotic prophylaxis for patients with prosthetic joint implants, 68 whereas the AAOS considers all patients with TJA to be at risk. 69 For TJA patients, the AAOS recommends administering antibiotic prophylaxis at least 1 hour before a dental procedure and discontinuing it within 24 hours after the procedure ends. 69 Single preoperative doses are acceptable for outpatient procedures. 70 Table 3 summarizes the studies that reported on postoperative measures for preventing SSI.

Although recommendations exist, the actual risk of infection resulting from dental procedures and the role of antibiotic prophylaxis are not well defined. Berbari and colleagues<sup>71</sup> found that antibiotic prophylaxis in high- or low-risk dental procedures did not decrease the risk of subsequent THA infection (OR, 0.9; 95% CI, 0.5-1.6) or TKA infection (OR, 1.2; 95% CI, 0.7-2.2). Moreover, the risk of infection was no higher for patients who had a prosthetic hip or knee and underwent a high- or

low-risk dental procedure without antibiotic prophylaxis (OR, 0.8; 95% CI, 0.4-1.6) than for similar patients who did not undergo a dental procedure (OR, 0.6; 95% CI, 0.4-1.1). Some studies highlight the low level of evidence supporting antibiotic prophylaxis during dental procedures. However, there is no evidence of adverse effects of antibiotic prophylaxis. Given the potential high risk of infection after such procedures, a more robust body of evidence is needed to reach consensus.

Evacuation Drain Management. Prolonged use of surgical evacuation drains may be a risk factor for SSI. Therefore, early drain removal is paramount. Higher infection rates with prolonged drain use have been found in patients with persistent wound drainage, including malnourished, obese, and over-anticoagulated patients. Patients with wounds persistently draining for >1 week should undergo superficial wound irrigation and débridement. Jaberi and colleagues<sup>74</sup> assessed 10,325 TJA patients and found that the majority of persistent drainage ceased within 1 week with use of less invasive measures, including oral antibiotics and local wound care. Furthermore, only 28% of patients with persistent drainage underwent surgical débridement. It is unclear if this practice alone is appropriate. Infection should always be suspected and treated aggressively, and cultures should be obtained from synovial fluid before antibiotics are started, unless there is an obvious superficial infection that does not require further work-up.67

## **Economic Impact**

SSIs remain a significant healthcare issue, and the social and financial costs are staggering. Without appropriate measures in place, these complications will place a larger burden on the healthcare system primarily as a result of longer hospital stays, multiple procedures, and increased resource utilization.<sup>75</sup> Given the risk of progression to prosthetic joint infection, early preventive interventions must be explored.

Several studies have addressed the economic implications of SSIs after TJA as well as the impact of preventive interventions (**Table 4**). Using the NIS database, Kurtz and colleagues<sup>4</sup> found that not only were hospital stays significantly longer for infected (vs noninfected) knee arthroplasties (7.6 vs 3.9 days; P < .0001), but hospital charges were 1.52 times higher (P < .0001), and results were similar for infected (vs noninfected) hips (9.7 vs 4.3 days; 1.76 times higher charges; P < .0001 for both). Kapadia and colleagues<sup>76</sup> matched 21

TKA patients with periprosthetic infections with 21 noninfected TKA patients at a single institution and found the infected patients had more readmissions (3.6 vs 0.1; P < .0001), longer hospitalizations (5.3)vs 3.0 days; P = .0002), more days in the hospital within 1 year of arthroplasty (23.7 vs 3.4 days; P < .0001), and more clinic visits (6.5 vs 1.3; P < .0001) .0001). Furthermore, the infected patients had a significantly higher mean annual cost of treatment (\$116,383 vs \$28,249; P < .0001). Performing a Markov analysis, Slover and colleagues<sup>77</sup> found that the decreased incidence of infection and the potential cost savings associated with preoperative S aureus screening and a decolonization protocol were able to offset the costs acquired by the screening and decolonization protocol. Similarly, Cummins and colleagues<sup>78</sup> evaluated the effects of ALBC on overall healthcare costs; if revision surgery was the primary outcome of all infections, use of ALBC (vs cement without antibiotics) resulted in a cost-effectiveness ratio of \$37,355 per quality-adjusted life year. Kapadia and colleagues<sup>79</sup> evaluated the economic impact of adding 2% chlorhexidine gluconate-impregnated cloths to an existing preoperative skin preparation protocol for TKA. One percent of non-chlorhexidine patients and 0.6% of chlorhexidine patients developed an infection. The reduction in incidence of infection amounted to projected net savings of almost \$2.1 million per 1000 TKA patients. Nationally, annual healthcare savings were expected to range from \$0.78 billion to \$3.18 billion with implementation of this protocol.

Improved patient selection may be an important factor in reducing SSIs. In an analysis of 8494 joint arthroplasties, Malinzak and colleagues<sup>80</sup> noted that patients with a BMI of  $\geq$ 50 kg/m² had an increased OR of infection of 21.3 compared to those with BMI <50 kg/m². Wagner and colleagues<sup>81</sup> analyzed 21,361 THAs and found that, for every BMI unit over 25 kg/m², there was an 8% increased risk of joint infection (P < .001). Although it is unknown if there is an association between reduction in preoperative BMI and reduction in postoperative complication risk, it may still be worthwhile and cost-effective to modify this and similar risk factors before elective procedures.

Market forces are becoming a larger consideration in healthcare and are being driven by provider competition. Because Treatment outcomes, quality of care, and healthcare prices have gained attention as a means of estimating potential costs. Because In 2011, the Centers for Medicare & Medicaid Services

Table 4. Summary of Studies Reporting on Economic Impact of Surgical-Site Infection and/or Effect of Preventive Measures

Study	Patients, N	Operation	Endpoints	Results	
Kurtz et al <sup>4</sup> (2008)	9190	THA, TKA	Incidence and costs of SSI	Greater LOS (7.6 vs 3.9 days; $P$ < .0001) and higher hospital charges ( $P$ < .0001) for infected TKAs than for noninfected TKAs	
				Greater LOS (9.7 vs 4.3 days; $P$ < .0001) and higher hospital charges ( $P$ < .0001) for infected THAs than for noninfected THAs	
Kapadia et al <sup>76</sup> (2014)	42	TKA	Readmissions, infections, days in hospital, number of clinic visits, and annual costs for infected vs noninfected TKAs	Infected patients had more readmissions (3.6 vs 0.1; $P < .0001$ ), longer hospitalization (5.3 vs 3.0 days; $P = .0002$ ), more days in hospital within 1 year of arthroplasty (23.7 vs 3.4; $P < .0001$ ), and more clinic visits (6.5 vs 1.3; $P < .0001$ )	
				Higher annual costs for infected patients (\$116,383 vs \$28,249; $P < .0001$ )	
Slover et al <sup>77</sup> (2011)	365	THA, TKA	Potential costs of decolonization protocol	Decreased incidence of infection and potential cost savings of preoperative <i>Staphylococcus aureus</i> screening and decolonization protocol are sufficient to offset costs acquired by protocol	
Cummins et al <sup>78</sup> (2009)	N/A	THA	Cost-effectiveness of use of ALBC if revision THA is considered primary outcome of all infections	Cost-effectiveness ratio of \$37,355 per quality-adjusted life year with use of ALBC vs cement without antibiotics	
Kapadia et al <sup>79</sup> (2013)	2213	TKA	Cost vs benefit of including chlorhexidine protocol in preoperative skin preparation protocol	Incidence of SSI reduced from 1% to 0.6%, with projected net saving of \$2.1 million per 1000 TKA patients	
Malinzak et al <sup>80</sup> (2009)	8494	THA, TKA	Selection of patients with comorbidities	Increased odds ratio of infection (18.3; $P$ < .0001) for patients with BMI of >50 kg/m² than for patients with BMI of <50 kg/m²	
				Patients with diabetes 3 times more likely to develop infection than patients without diabetes ( $P = .0027$ )	
Wagner et al <sup>81</sup> (2016)	21,361	THA	Association of BMI and postoperative complications	8% increased risk of THA infection for every unit of BMI >25 kg/m $^2$ ( $P < .001$ )	

Abbreviations: ALBC, antibiotic-laden bone cement; BMI, body mass index; LOS, length of stay; N/A, not applicable; SSI, surgical-site infection; THA, total hip arthroplasty; TKA, total knee arthroplasty.

(CMS) advanced the Bundled Payments for Care Improvement (BPCI) initiative, which aimed to provide better coordinated care of higher quality and lower cost. B4 This led to development of the Comprehensive Care for Joint Replacement (CJR) program, which gives beneficiaries flexibility in choosing services and ensures that providers adhere to required standards. During its 5-year test period beginning in 2016, the CJR program is projected to save CMS \$153 million. 4 Under this program, the institution where TJA is performed is responsible for all the costs of related care from time of surgery through 90 days after hospital dis-

charge—which is known as an "episode of care." If the cost incurred during an episode exceeds an established target cost (as determined by CMS), the hospital must repay Medicare the difference. Conversely, if the cost of an episode is less than the established target cost, the hospital is rewarded with the difference. Bundling payments for a single episode of care in this manner is thought to incentivize providers and hospitals to give patients more comprehensive and coordinated care. Given the substantial economic burden associated with joint arthroplasty infections, it is imperative for orthopedists to establish practical and

cost-effective strategies that can prevent these disastrous complications.

## **Conclusion**

SSIs are a devastating burden to patients, surgeons, and other healthcare providers. In recent years, new discoveries and innovations have helped mitigate the incidence of these complications of THA and TKA. However, the incidence of SSIs may rise with the increasing use of TJAs and with the development of new drug-resistant pathogens. In addition, the increasing number of TJAs performed on overweight and high-risk patients means the costs of postoperative infections will be substantial. With new reimbursement models in place, hospitals and providers are being held more accountable for the care they deliver during and after TJA. Consequently, more emphasis should be placed on techniques that are proved to minimize the incidence of SSIs.

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This paper will be judged for the Resident Writer's Award.