

# Surgical Site Infection in Spine Surgery: Who Is at Risk?

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## Abstract

**Study Design:** Retrospective literature review of spine surgical site infection (SSI).

**Objective:** To perform a review of SSI risk factors and more specifically, categorize them into patient and surgical factors.

**Methods:** A review of published literature on SSI risk factors in adult spine surgery was performed. We included studies that reported risk factors for SSI in adult spinal surgery. Excluded are pediatric patient populations, systematic reviews, and meta-analyses. Overall, we identified 72 cohort studies, 1 controlled-cohort study, 1 matched-cohort study, 1 matched-paired cohort study, 12 case-controlled studies (CCS), 6 case series, and 1 cross-sectional study.

**Results:** Patient-associated risk factors—diabetes mellitus, obesity (body mass index >35 kg/m<sup>2</sup>), subcutaneous fat thickness, multiple medical comorbidities, current smoker, and malnutrition were associated with SSI. Surgical associated factors—pre-operative radiation/postoperative blood transfusion, combined anterior/posterior approach, surgical invasiveness, or levels of instrumentation were associated with increased SSI. There is mixed evidence of age, duration of surgery, surgical team, intraoperative blood loss, dural tear, and urinary tract infection/urinary catheter in association with SSI.

**Conclusion:** SSIs are associated with many risk factors that can be patient or surgically related. Our review was able to identify important modifiable and nonmodifiable risk factors that can be essential in surgical planning and discussion with patients.

## Keywords

infection, cervical, lumbar, thoracic

## Introduction

Surgical site infection (SSI), with its associated morbidity, mortality, hospital length of stay (LOS), and cost, remains a common problem among spine surgery patients. The rate of SSI (superficial and deep) can range from 0.2% to 16.7%, depending on a number of patient-, pathology-, and procedure-related factors.<sup>1,2</sup> The treatment for SSI can be challenging requiring prolonged antibiotics, multiple revision surgeries, prolonged hospital stay, and in some patients, advanced soft tissue reconstructions. Numerous studies have attempted to identify the unique risk factors associated with SSI but are all too often limited to one specific diagnosis or procedure. Among previously identified factors associated with increased risk of SSI are excessive intraoperative blood loss, longer operative time, preoperative smoking, obesity, and higher degree of case complexity (as estimated by the Spine Surgery Invasiveness Index).<sup>3</sup> The purpose of this study is to perform a

review of risk factors for spine SSI and to categorize them into patient- and surgical-related factors.

## Methods

### Study Design and Search Strategy

We conducted a review of all published literature discussing risk factors for SSI in adult spine surgery. The search was

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performed using PubMed from its inception to July 20, 2017. Search terms used were (risk factor) AND (surgical site infection) AND (spine).

### Study Selection

We included studies that reported risk factors for SSI in adult spinal surgery. Exclusion criteria included those which reported on pediatric patient populations, systematic reviews, meta-analyses, those articles published in languages other than English or articles without an abstract.

## Results

### Search Results

The initial PubMed search returned 389 unique titles, of which 138 were included. Of those initially included, 1 was in a language other than English, 4 were meta-analyses, 18 were systematic reviews, 19 reported on pediatric populations, and 2 were excluded as full text could not be obtained. This left 94 unique studies for final and complete review.

### Overview of Included Studies

A total of 72 cohort studies, 1 controlled-cohort study, 1 matched-cohort study, 1 matched-paired cohort study, 12 case-controlled studies (CCS), 6 case series, and 1 cross-sectional study were identified. A summary of these studies can be found in Table 1. Twenty-one studies evaluated only a single potential risk factor, while 73 studies evaluated multiple potential variables as risk factors. Variables identified as associated or not associated with SSI are summarized in Tables 2, 3, and 4, arranged by study.

## Patient-Associated Risk Factors

There were a number of modifiable and nonmodifiable patient-associated risk factors for SSI that were identified, including age, diabetes, nutritional status, smoking, and obesity.

### Age

The relationship between patient age and the risk of SSI is not consistently reported in the literature, with numerous studies that implicating advanced age as a risk factor for SSI, and numerous studies finding no such association. Chaichana et al<sup>15</sup> reviewed 817 consecutive lumbar degenerative cases and found age of >70 years to be an independent risk factor for increased SSI. Manoso et al<sup>53</sup> found that Medicaid patients were at an increased risk for SSI but age alone was not an independent factor. In most studies, it was not possible to parse out the effect of age from other age-related comorbidities. Given the heterogeneity of results, it is not possible to definitively determine the role that age plays in the risk of SSI. The intuitive association between age and SSI is most likely related

to other age-related comorbidities or the accumulation of comorbidities that are globally manifest as patient frailty.

### General Comorbidities

Koutsoumbelis et al<sup>42</sup> reviewed 3128 patients undergoing lumbar fusion at a single institution. The authors found several comorbidities that are associated with increased SSI, including diabetes mellitus (DM), chronic obstructive pulmonary disease (COPD), coronary artery disease (CAD), and osteoporosis. The hypothesis of osteoporosis and the association with SSI is thought to be related to loss of collagen in skin as well as bone, leading to aberrant wound healing.<sup>42</sup> Klemencsics et al<sup>41</sup> concluded that patients with DM, CAD, arrhythmia, chronic liver disease, and autoimmune disease were at a higher risk of SSI. Furthermore, patients with multiple comorbidities are at an increased risk for SSI. Kurtz et al<sup>46</sup> found that patients with Charleston comorbidity index (CCI) of 5 versus 0 had an adjusted hazard ratio of 2.48 in developing a postoperative SSI.

### Diabetes Mellitus

It has been clearly established in the literature DM is an independent risk factor for SSI. There are several presumed pathophysiolgies for this. Microvascular disease associated with DM can impair nutrition and oxygen delivery to the peripheral tissues and reduce the systemic ability to resist infection. Hyperglycemia can impair leukocyte functions such as adherence, chemotaxis, and phagocytosis. Furthermore, DM can lead to impaired collagen synthesis and fibroblast proliferation that delays wound healing. Browne et al<sup>14</sup> reviewed the Nationwide Inpatient Sample (NIS) database of 11 000 patients who underwent lumbar fusion. The reported that DM was associated with increased SSI, blood transfusion, increased LOS and nonroutine discharge. Chen et al<sup>17</sup> found that patients with DM had an adjusted relative risk of 4.1 of developing an SSI. Golinvaux et al<sup>27</sup> further delineated the risk factors by reporting that insulin dependent DM portends a higher SSI risk than non-insulin-dependent diabetes. In patients with the diagnosis of DM, preoperative glycemic control is essential in minimizing the risk of SSI. Since HbA1c reflects the average blood glucose over a period of 6 to 12 weeks, it is an important indicator of how well diabetes is being managed. Hikata et al<sup>33</sup> found that patients with DM had a higher rate of SSI than nondiabetics (16.7% vs 3.2%). Furthermore, while immediate perioperative glycemic control did not differ between those DM patients that did or did not develop an SSI, the immediate preoperative HbA1C was significantly higher in those who developed SSI (7.6%) than in those who did not (6.9%). In the same study, SSI developed in none of the patients with HbA1C <7.0% and in 35.5% of patients with HbA1C >7.0%. Thus, pre- and perioperative glycemic control are significant modifiable risk factors for SSI and should be part of a systematic infection prevention strategy.

**Table I.** Characteristics of Studies Included in Review.

Author	Study Design	Level of Evidence	Group Demographics (Overall)			Group Demographics (Control)-If Applicable			Non-significant Variables	Significant Variables	Spinal levels	Approach	Instrumentation?	Indication for Surgery	Surgical Procedure	
			Number of Patients	Mean Age (y)	Special Characteristics	Number of Patients	Mean Age (y)	Number of Patients								
Abdul-Jabbar et al <sup>4</sup> 2012	Retro, Cohort	Uni-/multivariate logistic regression	III	66.28	56.5	Administrative claims database	193		6435	>7), bone or connective tissue cancer, approach (A/P combined)	CAD, DM, surgical region, smoking obesity, IA, diagnosis, transfusion, procedure type	C/T/L	A/P/Comb	Some	Degenerative, deformity, tumor	Decompression, fusion, deformity
Aoude et al <sup>5</sup> 2016	Retro, Cohort	Multivariate logistic regression	III	13.695	NS	NSQIP database, focused on blood transfusion				Transfusion (lumbar fusion only, not thoracic)	—	T/L	A/P/comb	Y	NS (excluded trauma)	Fusion
Asonguha et al <sup>6</sup> 2016	Retro, Controlled Cohort	Multivariate logistic regression	III	238						Epidural steroid paste, renal disease, immunosuppression	Procedure type, preoperative admission to hospital, surgical duration, EBL, diuretic, CHF, age, BMI, HTN, CAD, smoking, asthma, COPD	L	P	N	Degenerative	Decompression
Atkinson et al <sup>7</sup> 2016	Retro, Cohort	Uni-/bivariate logistic regression	III	152	60.6	Spinal metastases				Number of levels operated, surgical region (thoracic)	Age, gender, emergency surgery, Waterlow score, BMI, ETOH, smoking ASA, preoperative albumin, preoperative protein, preoperative WBC, preoperative CRP, incision length, interval between admission and surgery, number of staff in operating room	C/T/L	A/P	NS	Metastases	NS
Babu et al <sup>8</sup> 2012	Retro, Cohort	Uni-/multivariate logistic regression	IV	20	47	Tracheostomy/ACDF for SCI				—	Early tracheostomy	C	A	Y	Trauma, degenerative	Decompression, fusion
Barnes et al <sup>9</sup> 2012	Retro, Cohort	Multivariate logistic regression	III	90	44.8		15		75	Philadelphia collar, trauma	—	C	P	Y	Trauma, degenerative	Decompression, fusion
Bernrey et al <sup>10</sup> 2008	Retro, Cohort	ANOVA	III	71	40.28 ± 19.22	Tracheostomy in SCI (quadriplegia)				—	Early tracheostomy	C	A/P/comb	Y	Trauma	Decompression, fusion
Bian et al <sup>11</sup> 2003	Retro, Cohort	Uni-/multivariate logistic regression	III	256	43	Trauma	24	55	232	37	Delay to surgery (>60 hours), postoperative ICU stay (> day), number of surgical teams (orthopedic only vs combined orthopedic/neurosurgery)	C/T/L	A/P/comb	Y	Trauma	Decompression, fusion
Bohl et al <sup>12</sup> 2016	Retro, CCS	Poisson regression	III	10.825	NS	NSQIP database, focused on malnutrition				Albumin (<3.5 g/dL)	—	L	P	NS	Degenerative, deformity	Fusion
Boston et al <sup>13</sup> 2009	Retro, CCS	Multivariate logistic regression	III				55	44	179	45	Workers' compensation, method of hair removal, smoking, incontinence	NS	NS	NS	NS	Fusion, laminectomy, other
Browne et al <sup>14</sup> 2007	Retro, Cohort	Multivariate logistic regression	III	197.46	48.95 ± 18.16	Focused on DM				DM	—	L	?	?	Degenerative, deformity	Fusion
Chaidara et al <sup>15</sup> 2014	Retro, Cohort	Multivariate logistic regression	III	817	56 ± 14		37		780	Age (>70), DM, obesity, prior spine surgery, LOS (> days)	L	P	Y	Degenerative	Decompression, fusion	
Chen et al <sup>16</sup> 2009	Retro, Cohort	Multivariate logistic regression	II	244	NS					DM, EBL	L	P	Y	Degenerative, deformity	Fusion	
Chen et al <sup>17</sup> 2011	Retro, Cohort	Uni-/multivariate logistic regression	III	45	49.6	Sacral chordoma	16			Albumin, prior surgery, surgical duration (>6 hours)	Gender, obesity, smoking alcohol, DM, tumor size, radiation, instrumentation	S	P	Some	Tumor (sacral chordoma)	Tumor resection

(continued)

**Table 1.** (continued)

Author	Study Design	Group Demographics (Overall)			Group Demographics (Infected)			Group Demographics (Control)—If Applicable			Non-significant Variables	Indication for Surgery	Surgical Procedure	
		Level of Evidence	Number of Patients	Mean Age (y)	Special Characteristics	Number of Patients	Mean Age (y)	Significant Variables	Mean Age (y)	Approach	Instrumentation?			
Craig et al <sup>3</sup> 2012	Retro, Cohort	Multivariate logistic regression	III	152	49.5	63	53.5	BMI ( $>35 \text{ kg/m}^2$ ), HTN, surgical region (thoracic, lumbosacral), SII ( $>1$ ), renal disease	49.4	A/P comb	Some	Degenerative, tumor, trauma	Decompression, fusion, deformity correction	
De La Garza Ramos et al <sup>8</sup> 2015	Retro, Cohort	Student's t test, chi-square, univariate analysis, log-binomial model	IV	732	NS	Focused on obesity	—	—	—	L	P	Y	Degenerative	Fusion
De La Garza Ramos et al <sup>9</sup> 2016	Retro, Cohort	Multivariate logistic regression	III	36 440	NS	264	61.2 $\pm$ 11.6	36 176	60.5 $\pm$ 11.9	Chronic steroid use, surgical duration, renal disease (lumbar only), hemato-oncological disease (lumbar only), DM (lumbar only), obesity (lumbar only), LOS (lumbar only)	—	—	—	
De La Garza Ramos et al <sup>10</sup> 2017	Retro, CCS	Multivariate logistic regression	III	293	NS	Three-column osteotomy complex spine deformity	15	57 $\pm$ 14	278	61 $\pm$ 13	Obesity (class II), multilevel 3-column osteotomy	C/L	A/P comb	Decompression or fusion
Demura et al <sup>21</sup> 2009	Retro, Cohort	Uni-/multivariate logistic regression	III	113	56	Spinal metastases	8	—	113	DM, preoperative radiation	—	—	Deformity correction and fusion	
Dubroy et al <sup>12</sup> 2015	Pro, Cohort	Uni-/multivariate logistic regression	II	518	47.8 $\pm$ 19.1	Acute spinal trauma injury	25	493	Age, DM, surgical duration	—	—	—	Decompression, fusion, tumour resection (en bloc, debulking)	
Ee et al <sup>23</sup> 2014	Retro, CCS	Multivariate logistic regression	III	—	—	—	27	61.6 $\pm$ 13.7	162	56.8 $\pm$ 14.9	Open surgery (compared with MIS), DM, number of levels operated, BMI	C/T/L	A/P	Decompression or fusion
Fang et al <sup>24</sup> 2005	Retro, CCS	Uni-/multivariate logistic regression	III	—	—	Both adult and pediatric patients; only including adult results	21	—	29	Age ( $>60$ ) y, prior infection, EOH	—	—	Deformity, degenerative, disc disease	
Felling et al <sup>25</sup> 2012	Pro, Cohort	Pearson chi-square, multivariate regression	II	302	57	Cervical spondylotic myelopathy	—	—	—	Approach (posterior)	C	A/P comb	ACDF, other	
Fishman et al <sup>26</sup> 2017	Retro, Cohort	Chi-square	III	56	NS	Major deformity surgery ( $>8$ level fusion) focused on allogeneic transfusion	—	—	Allogenic transfusion	C/T/L	P	Y	Cervical spondylotic myelopathy	
Glassman et al <sup>27</sup> 2016	Retro, Cohort	Binary logistic regression	III	2653	NS	Based on 3 databases (Denmark, Japan, United States)	—	—	Gender, LOS, BMI, number of levels fused	—	A/P lateral/	NS	Degenerative	Fusion

(continued)

Table I. (continued)

Author	Study Design	Analysis	Level of Evidence	Group Demographics (Overall)	Group Demographics (Infected)	Group Demographics (Control)–If Applicable	Non-significant Variables	Significant Variables	Surgical Procedure
				Number of Patients	Mean Age (y)	Number of Patients	Mean Age (y)	Mean Age (y)	Approach
Golinvalx et al <sup>38</sup> 2014	Retro, Cohort	Multivariate logistic regression	III	15480	NS	NSQIP database, focused on DM (insulin vs non-insulin dependent)	NS	NS	A/P lateral/ comb
Grusky et al <sup>39</sup> 2012	Retro, Cohort	Step-down binary logistic regression	III	6666	NS	NS	NS	NS	Fusion
Haddad et al <sup>40</sup> 2016	Retro, Cohort	Multivariate logistic regression	III	187327	NS	NS database	Case order (in lumbar decompression only), approach (posterior, in cervical or lumbar fusion only), revision (cervical only), surgical duration (lumbar decompression or fusion), ASA (lumbar fusion only), age (lumbar fusion only)	Surgical duration (cervical only), age (lumbar decompression or cervical only), ASA (lumbar decompression or cervical only), gender, revision (lumbar decompression or fusion only)	Decompression, fusion
Hayashi et al <sup>41</sup> 2015	Retro, Cohort	Multivariate logistic regression	III	125	53.8	Total en bloc spondectomy for vertebral tumor (primary or metastases)	8	117	Decompression, fusion, stabilization
Hijas-Gomez et al <sup>42</sup> 2017	Retro, Cohort	Uni-/multivariate logistic regression	III	892	55	NS	Instrumentation, approach (A/P combined)	Age, tumor histology, prior surgery, surgical duration	Fusion, en bloc tumor resection
Hilata et al <sup>43</sup> 2014	Retro, Cohort	Chi-square, Mann-Whitney, Fisher's exact test	III	347	NS	NS	NS	NS	En bloc spondectomy
Jahet et al <sup>44</sup> 2016	Retro, Cohort	Multivariate logistic regression	III	3057	60.71	NSQIP database	54	54	Decompression, fusion
Kanian et al <sup>45</sup> 2009	Retro, Cohort	Chi square, T-test	III	997	NS	NS	DM, preoperative HbA1c	Age, gender, BMI, obesity, preoperative LOS, insulin use, steroid use, prior surgery, preoperative muscle weakness, preoperative incontinence, number of levels used, surgical duration, EBL, transfusion	Decompression, fusion
Kearns et al <sup>36</sup> 2014	Retro, Cohort	Student's t test, Wilcoxon rank-sum test, chi-square, Fisher's exact test	III	165	NS	NS	60.75	Approach (posterior), surgical duration (>208 min), ASA (>3)	Decompression, fusion
Kerwin et al <sup>37</sup> 2008	Retro, Matched cohort	Student's t test	III	16812	NS	Spinal fracture	54	47	Decompression, fusion
Kim et al <sup>46</sup> 2014	Retro, Cohort	Multivariate logistic regression	III	4588	NS	NSQIP database, focused on surgical duration	—	Time to surgery	Stabilization
							—	Surgical duration	NS (excluded trauma)
							—	—	Fusion (single-level)

(continued)

**Table I.** (continued)

Author	Study Design	Group Demographics (Overall)			Group Demographics (Control)–If Applicable (Infected)			Group Demographics (Control)–If Applicable (Infected)			Non-significant Variables	Spinal levels	Instrumentation?	Indication for Surgery	Surgical Procedure			
		Number of Patients	Mean Age (y)	Special Characteristics	Number of Patients	Mean Age (y)	Significant Variables	Mean Age (y)	Significant Variables	Mean Age (y)								
Kim et al <sup>39</sup> 2017	Retro, Cohort	Multivariate logistic regression, single variate t test	III	1831	NS			30	63.7	1801	63.6	Surgical duration	Gend, local bone irrigation, intradiscal irrigation	L	P	Fusion (PLIF)		
Klekamp et al <sup>40</sup> 1999	Retro, CS	Chi-square, Fisher's exact test	III	2614	NS								DM, weight, gender, trauma, inpatient status, smoking, UTI, age, cholesterol, albumin, total protein, ESR, triglycerides	C/T/L	NS	Some	NS	
Klemencics et al <sup>20</sup> 2016	Pro, Cohort	Multivariate logistic regression	II	1030	50			37		993			Age, BMI, DM, CAD, arrhythmia, chronic liver disease, autoimmune disease	SL, instrumentation	L	P	Y	Degenerative
Kouroumbelis et al <sup>21</sup> 2011	Retro, Cohort	Multivariate logistic regression	II	3218	56.9			86	60				Gender (female), DM, osteoporosis, CAD, number of comorbidities, obesity, number of personnel in operating room, dural tear, EBL (>500 cm <sup>3</sup> )	L	P	Y	Decompression, fusion	
Kudo et al <sup>41</sup> 2016	Retro, Cohort	Chi-square and Mann-Whitney U	III	105	64.4	Infection based on CRP		35	65.9 ± 16.9	70	63.6 ± 14.2	Surgical duration	Age, smoking, HTN, cholesterol, OSA, CHF, RA, number of comorbidities, number of surgeries, number of residents or fellows, surgical duration, number of drains, LOS, revision	C/T/L	NS	NS	NS	
Kukkula et al <sup>42</sup> 2015	Retro, Cohort	Multivariate logistic regression	III	266439	55.6								Age, gender, BMI, smoking, EOH, DM, EBL, instrumentation, preoperative total lymphocyte count, preoperative transferrin, preoperative prealbumin, preoperative retino binding protein	C/T/L	NS	NS	NS	
Kumar et al <sup>43</sup> 2015	Retro, Cohort	Multivariate logistic regression	III	98	60.1	Spinal metastases	17		81				Emergency surgery, timing of surgery (after day of incident in emergency cases)	—	L	A/P comb	Some	Degenerative, deformity, metasteses, trauma
Kurz et al <sup>44</sup> 2012	Retro, Cohort	Kaplan-Meier survival analysis, Cox regression	III	15069	NS	Medicare database							Number of levels operated (≥7), albumin (low), neurologic disability (read)	C/T/L	NS	NS	NS	Fusion
Lee et al <sup>45</sup> 2014	Retro, Cohort	Pearson chi-square, Fisher's exact test, multivariate logistic regression	III	1532	49.5	Spine End Result Registry (SEER)		66					Age, obesity, Charlston comorbidity index, socioeconomic status, revision, number of levels used, approach	SL, DM, CHF	Age, gender, RA, trauma, BMI	NS	NS	Spinal metastases
Lee et al <sup>46</sup> 2016	Retro, Cohort	Multivariate logistic regression	III	149	53.5 ± 15.8								Maximum fat thickness (T12-L5, operated levels or L4), prior surgery	C/T/L	NS	NS	NS	Decompression, fusion
Li et al <sup>47</sup> 2013	Retro, Cohort	Multivariate logistic regression	III	387	46.4	Sacral tumors							Age, gender, DM, preoperative albumin, prior sacral tumor resection, tumor size, histopathological diagnosis, blood control method, incision type (Y vs 2-way), proximal sacral segment resected, instrumentation, EBL	A/P comb	Some	Primary tumor	Tumor resection	

Table 1. (continued)

Author	Study Design	Group Demographics (Overall)			Group Demographics (Infected)			Group Demographics (Control) – If Applicable			Non-significant Variables	Indication for Surgery	Surgical Procedure	
		Number of Patients	Mean Age (y)	Special Characteristics	Number of Patients	Mean Age (y)	Significant Variables	Number of Patients	Mean Age (y)	Significant Variables				
Lieber et al <sup>50</sup> 2016	Multivariate logistic regression	III	60/179	57.1	NSQIP database	1110	59.069	Gender (female), inpatient, BMI, preoperative steroid use, anemia, ASA (>2), surgical duration	Instrumentation, bone graft use, transfusion, DHI, function status, COD, disseminated cancer, weight loss, preoperative transfusion, dialysis, deformity, hematoctit	C/T/L	A/P/ lateral/ comb	Some	NS (excluded trauma)	
Lim et al <sup>51</sup> 2014	Retro, Cohort	Chi-square	III	3353	NSQIP database	86		Obesity, ASA (>2), surgical duration (>6 hours)		L	A/P/ lateral/ comb	Y	NS (excluded trauma)	
Loijon et al <sup>52</sup> 2012	Pro, Cohort	Univariate, Fisher's exact test/Wilcoxon test	IV	169	50.0 ± 20.1	Spinal trauma			Age, ASA DM, surgical duration (>3 hours), time from injury to surgery (>3 days), number of levels fused, EBL (>600 cm <sup>3</sup> ), urinary catheter (>2 days)	Gender, BMI, smoking EtOH, antidiabetic agent/anticoagulant use, spinal region of trauma, neurologic impairment, surgical time of day (day vs night), approach, MIS, intraoperative transfusion, bedrest duration, drain	C/T/L	A/P/ comb	Y	Trauma
Manso et al <sup>53</sup> 2014	Retro, Cohort	Multivariate logistic regression	III	1532									Decompression, fusion, stabilization, deformity correction, tumor resection	
Maragakis et al <sup>54</sup> 2009	Retro, CCS	Multivariate logistic regression	III										Decompression, fusion	
Marguez-Lara et al <sup>55</sup> 2014	Retro, Cohort	Chi-square, Student's t test	IV	24/96	NS	Focused on BMI								
Martin et al <sup>56</sup> 2016	Retro, Cohort	Multivariate logistic regression	III	35/77		Focused on smoking								
Mehta et al <sup>57</sup> 2012	Retro, Cohort	Student's t test, Wilcoxon signed-rank test, chi-square, logistic regression	III	298										
Murphy et al <sup>58</sup> 2017	Retro, Cohort	Multivariate logistic regression	III	87/44	65	Focused on age								
Nordin et al <sup>59</sup> 1995	Retro, Case series	None	IV	11	30	Tracheostomy in SCI (quadriplegia)								
Ogihara et al <sup>60</sup> 2015	Pro, Cohort	Fisher's exact test, Wilcoxon signed-rank test, multivariate logistic regression	III	2736										

(continued)

Table I. (continued)

Author	Study Design	Analysis	Group Demographics (Overall)			Group Demographics (Control)—If Applicable (Infected)			Number of Patients	Mean Age (y)	Special Characteristics	Number of Patients	Mean Age (y)	Significant Variables	Non-significant Variables	Surgical Procedure	Indication for Surgery
			Level of Evidence	Number of Patients	Mean Age (y)	Group Demographics (Control)—If Applicable (Infected)	Number of Patients	Mean Age (y)									
Ohya et al <sup>61</sup> 2017	Retro, Cohort	Multivariate logistic regression	III	47.252	65.4	Iapane Diagnos Procedure Combination Database, focused on effect of month on surgery	438	46.814							Month of surgery (timing when medical staff rotate, only in academic hospital)	C/T/L	A/P comb
Oichi et al <sup>62</sup> 2017	Retro, Matched-pair cohort	Multivariate logistic regression	III	6921		Focused on Parkinson's disease									Parkinson's disease	—	Y
Ojo et al <sup>63</sup> 2016	Retro, Cross-section	Fisher's exact test	IV	62	44.2				10						DM, surgical region (cervical), procedure type (laminctomy and fixation); surgical duration	C/T/L	A/P comb
Olsen et al <sup>64</sup> 2003	Retro, CCS	Uni-/multivariate logistic regression	III	219					41	54.3	178	52.9			Ostomy, TB, anemia, diagnosis, instrumentation	C/T/L	P
Olsen et al <sup>65</sup> 2008	Retro, CCS	Multivariate logistic regression	III	273	52.4				46		227				Postoperative fecal incontinence, approach (posterior), tumor resection, obesity (mobid)	A/P comb	Y
Ones et al <sup>66</sup> 2011	Retro, Cohort	t test, chi-square	III	678		Nonsacral tumor (primary or metastases)	65	52.1	61.3	47.4					DM, timing of prophylactic antibiotics ( $\geq$ hour before surgery), preoperative glucose ( $>200$ ), obesity, number of residents ( $\geq$ )	NS (included C)	Y
Pull ter Gunne et al <sup>67</sup> 2009	Retro, Cohort	Cochran/Mantel-Haenszel, chi-square, multivariate	III	3174	55.6 ± 15.5										Prior surgery, preoperative radiation, any comorbidity, number of surgical teams involved ( $>$ ), complex plastics closure, LOS, hospital acquired infection	C/T/LUS (LS junction, not primary SJ)	Some
Pull ter Gunne et al <sup>68</sup> 2010 (deformity)	Retro, Cohort	Chi-square, multivariate	III	830	55.4 ± 16.1	Adult spinal deformity									Obesity, approach (not anterior), DM, prior SS, EBL ( $> 1$ ), surgical duration ( $>2$ hours)	A/P comb	Some
Pull ter Gunne et al <sup>69</sup> 2010 (osteotomy)	Retro, Cohort	Multivariate logistic regression	III	363	55.8	Types of osteotomies	20		343		VCR, obesity				Gender, DM, NSAID use, HTN, other cardiovascular pathology, smoking, preoperative protein, preoperative albumin, prior surgery, number of levels fused, approach, procedure type, surgical region, surgical duration, EBL, number of attending surgeons	C/T/LUS	A/P comb
Raidif et al <sup>68</sup> 2013	Retro, Cohort	Student's t test	III	791	53.7	Focused on anesthesia ready time	276	58.4			Anesthesia ready time ( $> 1$ hour)	—			Obesity, history of SS	C/T/L	A/P comb
															Gender, DM, NSAID use, HTN, other cardiovascular pathology, smoking, preoperative protein, preoperative albumin, prior surgery, number of levels fused, approach, procedure type, surgical region, surgical duration, EBL, number of attending surgeons	—	Y
															Deformity correction, stenosis, tumor/metastases, arthritis.	Deformity correction and fusion	Deformity correction and fusion
																	NS (excluded infection)
																	NS (excluded infection)

(continued)

Table 1. (continued)

Author	Study Design	Analysis	Level of Evidence	Group Demographics (Overall) (Infected)	Group Demographics (Control) If Applicable	Group Demographics (Control) If Applicable	Number of Patients	Mean Age (y)	Special Characteristics	Number of Patients	Mean Age (y)	Significant Variables	Non-significant Variables	Surgical Procedure
Ramos et al <sup>70</sup> 2016	Retro, Cohort	Unadjusted and adjusted logistic regression analysis	IV	668	63.9	Also included arthroplasty patients; review only looking at spine cohort; Focused on S aureus colonization	10	—	—	—	—	—	—	NS
Rao et al <sup>71</sup> 2011	Retro, CCS	Uni-/multivariate logistic regression	III	—	—	—	57	55 ± 15	18	57 ± 15	BMI, gender (male), drain duration	—	—	NS
Rechinc et al <sup>72</sup> 2001	Retro, Case series	Not listed	IV	117	NS	Thoracolumbar fracture	12	—	—	—	—	—	—	Decompression, stabilization
Rodgers et al <sup>73</sup> 2010	Retro, Cohort	Multivariate logistic regression, Student's t test, chi-square	III	600	61.4	Focused on XLIF procedure	—	—	—	—	Open surgery (vs XLIF)	—	—	Debridement, stabilization
Ruggieri et al <sup>74</sup> 2012	Retro, Case series	Kaplan-Meier survival analysis, log-rank test	IV	82	47	Primary sacral tumors	—	—	—	—	Procedure type (intralesional vs marginal vs wide resection), surgical duration	—	—	Debridement, stabilization
Saeedina et al <sup>75</sup> 2015	Retro, Cohort	Chi-square, ANOVA, multivariate regression	III	978	46	—	27	—	—	—	Age, level of resection (proximal vs distal), location of prior treatment (same institution vs other institution), tumor volume, neurological status (bowel/bladder continence)	—	—	Tumor debulking or resection
Silver et al <sup>76</sup> 2017	Retro, Case-control cohort	Chi-square, multivariate logistic regression	III	—	—	—	951	—	—	—	Age, gender, BMI, myopathy, NDU, approach, revision	—	—	Debridement, stabilization
Sarale et al <sup>77</sup> 2013	Retro, Cohort	Chi-square, Mann-Whitney U	III	110	—	—	32	—	—	—	Age, gender, BMI, surgical duration, comorbidities	—	—	Debridement, stabilization
Schimminel et al <sup>78</sup> 2010	Retro, Cohort	Uni-/multivariate logistic regression	III	1568	—	—	11	—	—	—	DM, procedure	—	—	Debridement, stabilization
Schoenfeld et al <sup>79</sup> 2013	Retro, Cohort	Uni-/multivariate logistic regression	III	5887	55.9	NSQIP database	36	—	—	135	Prior surgery, number of levels operated DM, smoking	—	—	Debridement, stabilization
											BMI, resident involvement, ASA (>2), surgical duration	—	—	Debridement, stabilization

(continued)

Table I. (continued)

Author	Study Design	Group Demographics (Overall)				Group Demographics (Infected)				Group Demographics (Control)—If Applicable				Non-significant Variables	Spinal levels	Approach	Instrumentation?	Indication for Surgery	Surgical Procedure	
		Number of Patients	Level of Evidence	Mean Age (y)	Special Characteristics	Number of Patients	Mean Age (y)	Number of Patients	Mean Age (y)	Significant Variables	Mean Age (y)	Number of Patients	Mean Age (y)							
Schwarzkopf et al <sup>39</sup> 2010	Multiple logistic regression	III			Focus on blood transfusion	61	56	71	53	BMI, transfusion				Gender, smoking, EtOH, DM, HTN, steroid use	T/L	?	Some	NS	Discectomy, decompression, fusion	
Schubba et al <sup>40</sup> 2008	Retro, Cohort	IV	Univariate, Fisher's exact test, multivariate	46	Sacral tumors					Prior lumbarosacral surgery, number of surgeons				Preoperative albumin, EBL, CSF leak, DM, instrumentation, laminectomies, obesity, plastic surgery closure, prior radiation, gender, smoking, age, bowel-bladder dysfunction, complex tissue reconstruction	S	A/P comb	Some	Primary tumor	Tumor resection	
Sebastian et al <sup>41</sup> 2016	Retro, Cohort	III	Student's t test, chi-square/Fisher's exact test, multivariate	5441	59 ± 13.6	NSQIP database	160	56.9 ± 12.2		BMI (>35 kg/m <sup>2</sup> ), chronic opioid use, surgical duration (>197 min)				Type of posterior surgery, DM, smoking, resident involvement, paralysis	C	P	Some	NS	Decompression, fusion, laminoplasty	
Shousha et al <sup>42</sup> 2014	Retro, Cohort	NS	Chi-square	139	53.6	Transoral approach for upper cervical spine				Indication (rheumatologic or tumor cases higher risk)				Age, gender, presence of metal implant	C	Transoral	Some	NS	Odontoidectomy, fusion, tumor resection	
Singh et al <sup>43</sup> 2017	Retro, Cohort	III	Chi-square	88 540	Lumbar epidural steroid injection prior to surgery	1411	87 129		Lumbar epidural steroid injection (within 3 mo) prior to surgery				Age, gender, presence of metal implant	—	L	P	Y	Degenerative	Fusion	
Skovrij et al <sup>44</sup> 2015	Retro, Cohort	III	Chi-square	517	51.8	Adult scoliosis, focused on surgeon experience				Less surgeon experience				Age, gender, presence of metal implant	—	T/L	NS	Y	Deformity	Deformity correction and fusion
Stambough et al <sup>45</sup> 1992	Retro, Case series	NS		IV			19	44		Malnutrition, trauma, UTI (all based on % of patients with these risk factors, no formal analysis)				No formal analysis	CT/L	NS	Some	NS	Trauma, degenerative, deformity, tumor, disc disease	Decompression, fusion, deformity correction
Sigita et al <sup>46</sup> 2016	Retro, Cohort	III	Mann-Whitney U, chi-square	279	63	Spinal metastases with intraoperative radiation	41	62	238	64	Katagiri/Tokuhashi's prognostic score, postoperative Frankel score, preoperative radiation, and postoperative performance				Surgical duration, EBL	NS	P	Y	Spinal metastases	Decompression/fusion, radiation
Temple et al <sup>47</sup> 2015	Retro, Case series	None		IV			83	56		Serum prealbumin below normal range (no formal analysis)				No formal analysis	CT/US	NS	Some	NS	Degenerative, trauma, tumor, deformity, hematoma, syringomyelia	Fusion, decompression, shunt
Tominaga et al <sup>48</sup> 2016	Retro, Cohort	III	Mann-Whitney U and Fisher's, multiple logistic regression	805	59		14	57.5	81	59	Surgical duration, ASA class 3, instrumentation, surgical region (thoracic in non-instrumented cases)				Two stage, revision, DM, smoking, BM, anemia, preoperative UTI, number of levels, incision length, number of personnel or surgeons	A/P comb	Y	Degenerative, infection, tumor, scoliosis	Decompression, fusion, deformity shunt	
Veeravagu et al <sup>49</sup> 2009	Retro, Cohort	III	Uni-/multivariate logistic regression	24 774	NS	Veterans Affairs' NSQIP database	752			DM, ASA (>2), weight loss, dependent functional status, intraoperative transfusion, cancer, fusion/instrumentation, surgical duration (>3 hours)				Age, gender, race, emergency surgery, bleeding disorder, smoking, EtOH, WBC, creatinine	CT/L	NS	Some	NS (excluded trauma)	Decompression, fusion, instrumentation	
Watanabe et al <sup>50</sup> 2010	Retro, Cohort	III	Uni-/multivariate logistic regression	223	53	Focused on effect of intraoperative irrigation	14	49	209	53	DM, trauma				Surgical duration, EBL, instrumentation, gender, age, smoking, obesity	CT/L	A/P comb	Y	Trama, tumor, degenerative, deformity	Decompression, fusion, deformity

(continued)

**Table 1.** (continued)

Author	Study Design	Analysis	Group Demographics (Overall)			Group Demographics (Control) = I-Aplicable (Infected)			Non-significant Variables			Indication for Surgery	Surgical Procedure
			Number of Patients	Mean Age (y)	Special Characteristics	Number of Patients	Mean Age (y)	Significant Variables	Spinal levels	Approach	Instrumentation?		
Weinstein et al <sup>32</sup> 2000	Retro, Case series	None	IV			46	57.2					C/L	A/P
Wimmer et al <sup>32</sup> 1998	Retro, Cohort	F test, paired Wilcoxon	III	850	Included some pediatric patients	22		Type of surgery (based on overall rate of infection, not statistically challenged)				NS	A/P
Woods et al <sup>33</sup> 2013	Retro, CCS	Conditional logistic regression	III		Focused on perioperative transfusions	56	61 ± 12	91	60 ± 148	Perioperative transfusion	—	L	A/P
Yang et al <sup>34</sup> 2016	Retro, Cohort	Pearson chi-square	III	1831	Patients n=65, lumbar epidural steroid injection prior to surgery	1%		Lumbar epidural steroid injection (within 3 months) prior to OR				L	P
												N	Degenerative Decompression

Abbreviations: ACDf, anterior cervical discectomy and fusion; ASA, American Society of Anesthesiologists class; BMI, body mass index; CAD, coronary artery disease; CCS, case-controlled study; CHF, congestive heart failure; COPD, chronic obstructive pulmonary disease; CSF, cerebrospinal fluid; CVA, cerebrovascular accident; DM, diabetes mellitus; DVT, deep vein thrombosis; EBL, estimated blood loss; EtOH, alcohol use; GCS, Glasgow Coma Scale; HTN, hypertension; IA, inflammatory arthropathy; IVDU, intravenous drug use; LOS, length of stay; MI, myocardial infarction; MIS, minimally invasive surgery; MUST, Malnutrition Universal Screening Tool score; NNIS, National Nosocomial Infection Surveillance index; NS, not specified; NSAID, nonsteroidal anti-inflammatory drug; NSQIP, National Surgical Quality Improvement Program; CSA, obstructive sleep apnea; PE, pulmonary embolus; Pro, prospective; PVD, peripheral vascular disease; RA, rheumatoid arthritis; Retro, retrospective; SCI, spinal cord injury; SI, surgical invasiveness index; UTI, urinary tract infection; XLIF, extreme lateral interbody fusion. C, cervical; T, thoracic; L, lumbar; A, anterior; P, posterior; Comb, A/P combined.

**Table 2.** Patient-Associated Variables and Association With Surgical Site Infection by Study.

(continued)

**Table 2.** (continued)

(continued)

**Table 2.** (continued)

**Abbreviations:** Y, yes—association found; C, conditional—association under certain conditions; N, no association found; BMI, body mass index; COPD, chronic obstructive pulmonary disease.

**Table 3.** Diagnosis-Associated Variables and Association With Surgical Site Infection by Study.

Author	Deformity	Intra- vs Extradural	Tumor Size	Tumor Histopathological Diagnosis	Primary vs Metastatic Tumor
Abdul-Jabbar et al <sup>4</sup> 2012				Y	
Aoude et al <sup>5</sup> 2016					
Asomugha et al <sup>6</sup> 2016					
Atkinson et al <sup>7</sup> 2016					
Babu et al <sup>8</sup> 2012					
Barnes et al <sup>9</sup> 2012					
Berney et al <sup>10</sup> 2008					
Blam et al <sup>11</sup> 2003					
Bohl et al <sup>12</sup> 2016					
Boston et al <sup>13</sup> 2009					
Browne et al <sup>14</sup> 2007					
Chaichana et al <sup>15</sup> 2014					
Chen et al <sup>16</sup> 2009					
Chen et al <sup>17</sup> 2011					
Cizik et al <sup>3</sup> 2012					
De La Garza Ramos et al <sup>18</sup> 2015					
De La Garza Ramos et al <sup>19</sup> 2016					
De La Garza Ramos et al <sup>20</sup> 2017	C				
Demura et al <sup>21</sup> 2009					
Dubory et al <sup>22</sup> 2015					
Ee et al <sup>23</sup> 2014					
Fang et al <sup>24</sup> 2005					
Fehlings et al <sup>25</sup> 2012					
Fisahn et al <sup>26</sup> 2017					
Glassman et al <sup>27</sup> 2016					
Golinvaux et al <sup>28</sup> 2014					
Gruskay et al <sup>29</sup> 2012					
Haddad et al <sup>30</sup> 2016					
Hayashi et al <sup>31</sup> 2015				N	
Hijas-Gomez et al <sup>32</sup> 2017					
Hikata et al <sup>33</sup> 2014					
Jalai et al <sup>34</sup> 2016					
Kanafani et al <sup>35</sup> 2009					
Keam et al <sup>36</sup> 2014					
Kerwin et al <sup>37</sup> 2008					
Kim et al <sup>38</sup> 2014					
Kim et al <sup>39</sup> 2015					
Klekamp et al <sup>40</sup> 1999					
Klemencsics et al <sup>41</sup> 2016					
Koutsoumbelis et al <sup>42</sup> 2011					
Kudo et al <sup>43</sup> 2016					
Kukreja et al <sup>44</sup> 2015					
Kumar et al <sup>45</sup> 2015					
Kurtz et al <sup>46</sup> 2012					
Lee et al <sup>47</sup> 2014					
Lee et al <sup>48</sup> 2016					
Li et al <sup>49</sup> 2013				N	N
Lieber et al <sup>50</sup> 2016	Y				

(continued)

**Table 3.** (continued)

Author	Deformity	Intra- vs Extradural	Tumor Size	Tumor Histopathological Diagnosis	Primary vs Metastatic Tumor
Lim et al <sup>51</sup> 2014					
Lonjon et al <sup>52</sup> 2012					
Manoso et al <sup>53</sup> 2014					
Maragakis et al <sup>54</sup> 2009					
Marquez-Lara et al <sup>55</sup> 2014					
Martin et al <sup>56</sup> 2016					
Mehta et al <sup>57</sup> 2012					
Murphy et al <sup>58</sup> 2017					
Northrup et al <sup>59</sup> 1995					
Ogihara et al <sup>60</sup> 2015					
Ohya et al <sup>61</sup> 2017					
Oichi et al <sup>62</sup> 2017					
Ojo et al <sup>63</sup> 2016					
Olsen et al <sup>64</sup> 2003					
Olsen et al <sup>65</sup> 2008					
Omeis et al <sup>66</sup> 2011		N			N
Pull ter Gunne et al <sup>1</sup> 2009					
Pull ter Gunne et al <sup>67</sup> 2010 (deformity)					
Pull ter Gunne et al <sup>68</sup> 2010 (osteotomy)					
Radcliff et al <sup>69</sup> 2013					
Ramos et al <sup>70</sup> 2016					
Rao et al <sup>71</sup> 2011					
Rechtine et al <sup>72</sup> 2001					
Rodgers et al <sup>73</sup> 2010					
Ruggieri et al <sup>74</sup> 2012			N		
Saeedinia et al <sup>75</sup> 2015					
Salvetti et al <sup>76</sup> 2017					
Satake et al <sup>77</sup> 2013					
Schimmel et al <sup>2</sup> 2010					
Schoenfeld et al <sup>78</sup> 2013					
Schwarzkopf et al <sup>79</sup> 2010					
Sciubba et al <sup>80</sup> 2008					
Sebastian et al <sup>81</sup> 2016					
Shousha et al <sup>82</sup> 2014					
Singla et al <sup>83</sup> 2017					
Skovrlj et al <sup>84</sup> 2015					
Stambough et al <sup>85</sup> 1992					
Sugita et al <sup>86</sup> 2016					
Tempel et al <sup>87</sup> 2015					
Tominaga et al <sup>88</sup> 2016					
Veeravagu et al <sup>89</sup> 2009					
Watanabe et al <sup>90</sup> 2010					
Weinstein et al <sup>91</sup> 2000					
Wimmer et al <sup>92</sup> 1998					
Woods et al <sup>93</sup> 2013					
Yang et al <sup>94</sup> 2016					

Abbreviations: Y, yes—association found; C, conditional—association under certain conditions; N, no association found.

**Table 4.** Surgery-Associated Variables and Association With Surgical Site Infection by Study.

Author																			
Abdul-Jabbar et al <sup>4</sup> 2012																			
Aoudé et al <sup>5</sup> 2016																			
Asomugha et al <sup>6</sup> 2016																			
Atkinson et al <sup>7</sup> 2016																			
Babu et al <sup>8</sup> 2012																			
Barnes et al <sup>9</sup> 2012																			
Bernay et al <sup>10</sup> 2008																			
Bian et al <sup>11</sup> 2003																			
Bohl et al <sup>12</sup> 2016																			
Boston et al <sup>13</sup> 2009																			
Browne et al <sup>14</sup> 2007																			
Chaiachana et al <sup>15</sup> 2014																			
Chen et al <sup>16</sup> 2009																			
Chen et al <sup>17</sup> 2011																			
Cizik et al <sup>3</sup> 2012																			
De La Garza Ramos et al <sup>18</sup> 2015																			
De La Garza Ramos et al <sup>19</sup> 2016																			
De La Garza Ramos et al <sup>20</sup> 2017																			
Demura et al <sup>21</sup> 2009																			
Dubory et al <sup>22</sup> 2015																			
Ee et al <sup>23</sup> 2014																			
Fang et al <sup>24</sup> 2005																			
Fehlings et al <sup>25</sup> 2012																			
Fisahn et al <sup>26</sup> 2017																			
Glassman et al <sup>27</sup> 2016																			
Golinvaux et al <sup>28</sup> 2014																			
Gruskay et al <sup>29</sup> 2012																			
Haddad et al <sup>30</sup> 2016																			
Hayashi et al <sup>31</sup> 2015																			
Hijas-Gomez et al <sup>32</sup> 2017																			
Hikata et al <sup>33</sup> 2014																			

(continued)

**Table 4** (continued)

Author	Antibiotic Timing/Redosing	Approach	Bone Graft	Case Order	Cervical Collar	Complex Closure	Delay to Surgery	Dural Tear/CSF Leak	Early Tracheostomy	EBL	Emergency Surgery	Epidural Steroid	Incision Length	Instrumentation	Intraoperative Temperature	Length of Stay	Number of Levels Operated/Fused	Number of Staff	Number of Surgical Teams	Open vs MIS	Preoperative Admission	Procedure Type	Resident-Fellow	Revision	Surgical Invasiveness	Surgical Region	Transfusion	U/I
Jalai et al <sup>34</sup> 2016																												
Kanafani et al <sup>35</sup> 2009																												
Kean et al <sup>36</sup> 2014																												
Kerwin et al <sup>37</sup> 2008																												
Kim et al <sup>38</sup> 2014																												
Kim et al <sup>39</sup> 2015																												
Klekamp et al <sup>40</sup> 1999																												
Klemencsics <sup>41</sup> et al 2016																												
Koutsoumbelis et al <sup>42</sup> 2011																												
Kudo et al <sup>43</sup> 2016																												
Kukreja et al <sup>44</sup> 2015																												
Kumar et al <sup>45</sup> 2015																												
Kurtz et al <sup>46</sup> 2012																												
Lee et al <sup>47</sup> 2014																												
Lee et al <sup>48</sup> 2016																												
Li et al <sup>49</sup> 2013																												
Lieber et al <sup>50</sup> 2016																												
Lin et al <sup>51</sup> 2014																												
Lonion et al <sup>52</sup> 2012																												
Manoso et al <sup>53</sup> 2014																												
Margaklis et al <sup>54</sup> 2009																												
Marquez-Lara et al <sup>55</sup> 2014																												
Martin et al <sup>56</sup> 2016																												
Mehta et al <sup>57</sup> 2012																												
Murphy et al <sup>58</sup> 2017																												
Northrup et al <sup>59</sup> 1995																												
Ogihara et al <sup>60</sup> 2015																												
Ohyu et al <sup>61</sup> 2017																												
Oichi et al <sup>62</sup> 2017																												
Ojo et al <sup>63</sup> 2016																												
Olsen et al <sup>64</sup> 2003																												
Olsen et al <sup>65</sup> 2008																												

(continued)

Table 4. (continued)

Author	Antibiotic Timing/Redosing	Approach	Bone Graft	Case Order	Cervical Collar	Complex Closure	Delay to Surgery	Dural Tear/CSF Leak	Early Tracheostomy	Instrumentation	Length of Stay	Number of Levels	Number of Staff	Open vs MIS	Preoperative Admission	Procedure Type	Resident-Fellow	Review	Surgical Durations	Surgical Invasiveness	Surgical Region	Transfusion	UTI	
Onnis et al <sup>66</sup> 2011																								
Pull ter Gunne et al <sup>67</sup> 2010 (deformity)																								
Pull ter Gunne et al <sup>68</sup> 2010 (osteotomy)																								
Raddcliff et al <sup>69</sup> 2013																								
Ramos et al <sup>70</sup> 2016																								
Rao et al <sup>71</sup> 2011																								
Rechtine et al <sup>72</sup> 2001																								
Rodgers et al <sup>73</sup> 2010																								
Ruggieri et al <sup>74</sup> 2012																								
Saeednia et al <sup>75</sup> 2015																								
Salvetti et al <sup>76</sup> 2017																								
Satake et al <sup>77</sup> 2013																								
Schimmele et al <sup>2</sup> 2010																								
Schoenfeld et al <sup>78</sup> 2013																								
Schwarzkopf <sup>79</sup> et al 2010																								
Scubba et al <sup>80</sup> 2008																								
Sebastian et al <sup>81</sup> 2016																								
Shousha et al <sup>82</sup> 2014																								
Singla et al <sup>83</sup> 2017																								
Skovrlj et al <sup>84</sup> 2015																								
Stambaugh et al <sup>85</sup> 1992																								
Sugita et al <sup>86</sup> 2016																								
Tempel et al <sup>87</sup> 2015																								
Tominaga et al <sup>88</sup> 2016																								
Veeravagu et al <sup>89</sup> 2009																								
Watanabe et al <sup>90</sup> 2010																								
Weinstein et al <sup>91</sup> 2000																								
Wimmer et al <sup>92</sup> 1998																								
Woods et al <sup>93</sup> 2013																								
Yang et al <sup>94</sup> 2016																								

Abbreviations: Y, yes—association found; C, conditional—association under certain conditions; N, no association found; CSF, cerebrospinal fluid; EBL, estimated blood loss; MIS, minimally invasive surgery; UTI, urinary tract infection.

## Nutrition

There are several serum markers such as transferrin, prealbumin, albumin, total lymphocyte count that can be measured for early detection of nutritional deficits. Bohl et al<sup>12</sup> performed a retrospective review of the ACS-NSQIP database and found the overall prevalence of hypoalbuminemia (defined as <3.5 g/dL) as 4.8% in patients who underwent posterior lumbar fusion of 1 to 3 levels. The authors found patients with preoperative hypoalbuminemia had a higher risk of wound dehiscence, SSI and urinary tract infection. Furthermore, those patients also had longer inpatient stay and a higher risk of unplanned hospital readmission within 30 days of surgery. Chen et al<sup>17</sup> found that hypoalbuminemia was an independent risk factor for SSI in a cohort of patients who underwent sacral chordoma resection.

While albumin has been routinely used as a surrogate marker for nutritional status, recent studies have shown that prealbumin (half-life of 2 days) may also be used to assess a patient's nutritional status in the perioperative period. Salvetti et al<sup>76</sup> found that preoperative prealbumin level of <20 mg/dL had higher risk of developing SSI with adjusted hazard ratio of 2.12. This collection of literature would suggest that for the reduction of SSI, it is advisable to assess nutritional status preoperatively by checking prealbumin, albumin and total lymphocyte count. Nutritional supplementation may be considered if the patient is malnourished and undergoing complex surgical reconstruction.

## Smoking

Nicotine leads to peripheral vasoconstriction and tissue hypoxia and results in impaired local angiogenesis and epithelialization. Smoking leads to decreased wound collagen production in *in vitro* and in animal studies. Martin et al<sup>56</sup> in 2016 found that active smokers are at a significantly higher risk of SSI compared with former smokers. That study from the ACS-NSQIP database, of patients who underwent elective lumbar surgery, categorized patients into: never smoked, former smoker (quit 12 months ago) and active smoker. Active smokers had a significantly higher risk of SSI compared with nonsmokers. Former smoker had an increased risk, but it was not significantly different from nonsmokers. Pack years of 1 to 20 and 20 to 40 were both found to have increased risk for SSI.

## Obesity/Body Mass Index

Much has been studied about the relationship between obesity/body mass index (BMI) and SSI. Cizik et al<sup>3</sup> performed a retrospective review of all patients who had spine surgery at a single institution and found that BMI >35 kg/m<sup>2</sup> was an independent risk factor for increased risk of SSI. In a retrospective cohort review, De la Garza-Ramos et al<sup>18</sup> found that obesity (BMI >30 kg/m<sup>2</sup>) resulted in an increased risk of SSI (risk ratio 3.11) in patients who underwent one to three level lumbar fusion surgery. Marquez-Lara et al<sup>55</sup> also found that BMI >30 kg/m<sup>2</sup> (class I obesity) had increased risk of

superficial wound infection. Furthermore, Mehta et al<sup>57</sup> found that body mass distribution, in particular increased skin to lamina distance and subcutaneous fat thickness, are independent risk factors for SSI. This study may indicate that although higher BMI is an independent risk factor associated with increased SSI, in patients with higher muscle mass, BMI may not be the most accurate variable to predict postoperative SSI. Lee et al<sup>48</sup> found that for every 1-mm of thickness in subcutaneous fat there was 6% increase in risk of SSI. Patients with at least 50 mm of posterior lumbar fat thickness had 4-fold increase in risk of SSI compared to those with less than 50 mm.

## Surgery-Associated Risk Factors

### Timing and Duration of Surgery

Most studies have found no significant association between "emergency surgery" and SSI.<sup>7,21,31,52,54,60,71,89</sup> Three studies have shown that increased duration from time of injury or admission to time of surgery was associated with increased risk of SSI.<sup>11,44,52</sup> Lonjon et al<sup>52</sup> found no association between the risk of SSI and surgery being done at night or after-hours.

A large number of studies have found that increased operative time increases the risk of SSI,\* with a smaller number of contradicting studies.<sup>6,11,16,23,27,35,48,67,71,86</sup> Several studies used a cutoff of surgical duration in determining an association with SSI, although this varies between papers, ranging anywhere from 100 minutes to 5 hours,<sup>13,24</sup> and no conclusions can be made with regards to a specific duration as an inflection point in the risk for SSI.

### Surgical Approach, Procedure, and Invasiveness

**Surgical Approach: Anterior, Posterior, or Combined.** If one considers studies that evaluate only cervical<sup>25,30,34</sup> or only lumbar procedures<sup>2,46</sup> separately, or separately analyzed approach in each spinal level subgroup,<sup>28,64</sup> most find an association between approach and SSI. In all studies with either cervical only groups or cervical subanalysis,<sup>25,28,30,34,64</sup> a posterior approach is consistently reported as a risk factor for SSI as compared with an anterior approach. Of those examining lumbar procedures,<sup>2,28,46,64</sup> for the most part, a combined anterior-posterior or posterior only approach was a risk factor for SSI as compared with anterior approach. Only 1 study had a thoracic subgroup analysis for approach, with Olsen et al<sup>64</sup> finding a posteriorly only approach to be associated with SSI as compared anterior alone. For the most part, those studies that have not found an association<sup>11,22,52,65,68,71,75,77</sup> have included a combination of cervical, thoracic, and lumbar procedures, which may confound the significance of approach given that the relative risk of an anterior versus posterior approach is different at various spinal levels. In those studies showing approach to be a risk factor for SSI,<sup>1,4,25,28,30,31,34,46,54,64</sup> the general trend is for a combined anterior-posterior approach to

\*References 1, 4, 13, 19, 22, 28, 31, 34, 38, 39, 43, 51, 52, 54, 60, 63, 78, 81, 92.

have the highest risk for SSI, followed by a posterior approach, with the anterior approach often reducing the risk for SSI.

**Minimally Invasive Versus Open Surgery.** Both Ee et al<sup>23</sup> and Rodgers et al<sup>73</sup> found that open surgery was associated with a higher risk of SSI as compared to MIS techniques (procedures performed through a tubular retractor system or extreme lateral interbody fusion (XLIF)) in elective lumbar spine surgery. Dubory et al<sup>22</sup> and Lonjon et al<sup>52</sup> found no such difference in SSI rates in spinal trauma. It should be noted the latter studies come from the same group, one of two that utilized only univariate analysis, and the type of MIS technique used was not defined, making it difficult to compare these results with those of either Ee et al<sup>23</sup> or Rodgers et al<sup>73</sup>.

**Surgical "Invasiveness".** Surgical invasiveness can be considered a composite of a number of variables as previously described, including number of levels operated on, the type of procedure performed at each level, and approach used. To allow comparison of the invasiveness of disparate spinal procedures, a surgical invasiveness index (SII) was developed by Mirza et al.<sup>95</sup> This index is a composite score based on the number of vertebral levels operated on, the type of intervention on each vertebra—decompression, fusion, instrumentation—as well as the approach used at each level, and has been validated against both blood loss and surgical duration. Of the 4 studies that evaluated SII as a variable with regards to SSI, 3 found that an increase in SII was associated with SSI.<sup>3,48,53</sup> However, Klemencsics et al<sup>1</sup> found no such association. This may be related to the populations and procedure types studied, as Klemencsics et al<sup>1</sup> looked at elective routine degenerative lumbar procedures, with a maximal SII of 15, while the other 3 studies looked across a broad range of surgery types using large databases and presumed higher maximal SII scores.<sup>3,48,53</sup> If this is the case, the association between SII and SSI may only exist in the upper range of the SII.

### Perioperative Interventions

**Tracheostomy.** Despite theoretical concerns, all 3 studies evaluating the potential of cross-contamination, have found no increased SSI risk for early tracheostomy (either pre- or post-operatively) in anterior cervical spine surgery. Babu et al<sup>8</sup> and Berney et al<sup>10</sup> found a low rate of SSI with early tracheostomy after anterior cervical stabilization for acute cervical trauma with spinal cord injury. Northrup et al,<sup>59</sup> in a review of 11 spinal cord injury patients, concluded that an existing tracheostomy was not a risk factor for SSI for subsequent anterior cervical spine stabilization.

**Cervical Orthosis.** Barnes et al<sup>9</sup> reported that the use of a Philadelphia collar for a minimum of 48 hours postoperatively increased the rate of SSI in posterior cervical spine surgery. This is in keeping with the known effects of pressure on skin and soft tissue from cervical orthoses.<sup>96</sup>

**Blood Transfusion.** Transfusion is an independent risk factor for SSI in other surgical specialties,<sup>97,39,98</sup> and it has been strongly suggested to similarly be a risk factor in adult spine surgery. There exists some conflict in the literature to date, with a majority of studies finding a significant increase in SSI associated with transfusion,<sup>4,5,28,61,79,89,93</sup> but others finding it not to be of significance.<sup>22,31,33,46,52,54,71</sup> However, of those studies that have focused on the implications of blood transfusion in adult spine surgery,<sup>5,28,79,93</sup> all 4 have shown transfusion to be an independent risk factor for SSI. The association of transfusion with SSI has been thought to be a result of transfusion-related immunomodulation (TRIM), a phenomenon whereby antigens in blood products may result in T-cell unresponsiveness and subsequent immunosuppression.<sup>99</sup> Bacterial contamination of blood products are another potential explanation for the effects of transfusion on SSI.<sup>100</sup>

Urinary tract infection (UTI) has been investigated as a possible source and hence risk factor for SSI,<sup>88,101</sup> and presence of a catheter is a well-established risk for UTI.<sup>102</sup> However, there has been limited study into urinary catheters as an independent risk factor for SSI in spine surgery, with both articles on this topic coming from the same group.<sup>22,52</sup> While Dubory et al<sup>22</sup> found that presence of a bladder catheter was not a significant risk for SSI after multivariate analysis, Lonjon et al<sup>52</sup> did find that a prolonged duration of catheterization greater than five days was associated with SSI after univariate analysis, although no multivariate analysis was performed. Based on these results, limited if any conclusion about urinary catheterization and SSI can be made.

Radiation is known to have deleterious effects on tissue, both in short-term effects on wound healing such as skin breakdown, lower tensile strength, and delayed healing rates from damage to epithelial cells and fibroblasts,<sup>103</sup> and in long-term effects on soft tissue resulting in fibrosis, poor vascularity, and a higher propensity to go onto atrophy or necrosis.<sup>104</sup> As such, preoperative radiation, whether recent or remote, has been regarded as a substantial risk factor for SSI. In nonsacral tumors, 3 studies focused on risk factors for SSI in spinal metastases or primary spinal tumors found preoperative radiation to be a significant risk for SSI.<sup>21,66,86</sup> In primary sacral tumors, the results have been more mixed, with 2 studies suggesting no significant association between previous radiation and SSI<sup>17,80</sup> against 1 study finding previous radiation to be a risk factor.<sup>49</sup> This is unsurprising, given the complexities of sacral tumor resection, higher infection rates, and smaller case numbers within each study by which to find association.

Evidence from a single controlled-cohort study suggests that use of epidural steroid paste in lumbar decompression is a risk factor for SSI, with the rate of SSI in the steroid paste group being 5.83% as compared to 1.11% in the control group.<sup>5</sup> Two studies from the same institution have shown preoperative lumbar epidural injections, if within 3 months of surgery, can also be a risk factor for SSI in both lumbar decompression<sup>94</sup> and lumbar fusion<sup>83</sup> surgery.

## Surgical Team

Only 1 study has looked at surgeon experience in relation to SSI, with Skovrlj et al<sup>84</sup> finding that in adult scoliosis surgery, candidate members as compared with active members for the Scoliosis Research Society had a 2-fold increase in the rate of superficial, though not deep, SSI which was statistically significant. In regards to the effect of resident involvement and experience, 3 studies looking at different aspects of this have found an association with SSI.<sup>61,65,78</sup> Looking at seasonal variation in the risk of reoperation for SSI, Ohya et al.<sup>61</sup> found that April, during which medical staff turnover in Japan, was associated with the highest rate of SSI and reoperation for the same in academic centers while no such seasonal variation occurred in nonacademic hospitals, suggesting that the influx in new and henceforth inexperienced staff may be a contributor to this result. More directly, Schoenfeld et al.<sup>78</sup> found that resident involvement was an independent risk factor for SSI even after multivariate analysis encompassing procedure time and patient comorbidity, while Olsen et al.<sup>65</sup> found that the participation of 2 or more residents increased the risk of SSI although the latter assumed this to be a proxy for surgical complexity rather than a result of resident involvement. Koutsoumbelis et al,<sup>42</sup> however, found no significant association between number of residents and fellows and SSI and Sebastian et al<sup>81</sup> found no association between resident involvement and SSI. As such, it remains unclear as to the effect of residents on SSI.

The number of surgeons involved in spine surgery does not appear to be a significant risk factor, with 3 studies,<sup>23,67,88</sup> finding no significant association between number of scrubbed or senior surgeons and SSI. However, Sciubba et al.<sup>80</sup> found a larger number of surgeons to be associated with SSI in sacral tumor resection, where a multidisciplinary surgical team is often required. Koutsoumbelis et al<sup>42</sup> found that the overall number of personnel may be a risk if 10 or more personnel are present in the operating room. Operating room traffic and the number of personnel both have been linked to an increase in airborne contaminants<sup>103</sup> and could thereby increase the risk of contamination of the surgical wound.

The effects of involvement of more than one surgical team on SSI is not well studied and is confounded by the fact the presence of additional surgical teams may imply greater surgical complexity and therefore potential risk for infection. Blam et al<sup>11</sup> found that the combined involvement of both orthopedic and neurosurgical teams had a reduced rate of SSI as compared with orthopedics alone, with a trend toward the same as compared with neurosurgery alone, despite the greater operating room traffic involved although no clear explanation could be had for this effect. On the other hand, Rao et al<sup>71</sup> found no significant association between involvement of both services as compared with either orthopedics or neurosurgery alone. Involvement of more than 1 surgical team was found by Omeis et al<sup>66</sup> to be a risk for SSI in spinal tumors. However, in most cases this was due to involvement of plastic surgery and the requirement of a complex soft tissue reconstruction with its

attendant risks with regard to infection, confounding the effect on SSI. In the case of sacral tumors, Sciubba et al<sup>80</sup> found no statistically significant association between having a plastic surgeon for closure and SSI.

## Intraoperative Concerns and Complications

Increased intraoperative blood loss has not been clearly shown to be a risk factor for SSI, with a number of studies on either side of whether an association exists or not.<sup>†</sup> It is difficult to separate blood loss from other confounding variables such as surgical duration, invasiveness, as well as the need for transfusion. Only 3 studies reporting on intraoperative blood loss also reported on transfusion, with one showing an independent association between each and SSI,<sup>93</sup> one showing no association between either and SSI,<sup>22</sup> and one showing an association between blood loss but not transfusion and SSI.<sup>52</sup> Enough contradiction exists to preclude any conclusions with regard to blood loss as a possible risk factor.

Intraoperative hypothermia has been viewed as a potential risk factor for SSI due to its induction of vasoconstriction and its negative effects on oxygenation, neutrophil function, and wound healing.<sup>105</sup> However, intraoperative temperature has not been found to be a risk factor so far for SSI in spine surgery, with all three studies including this variable demonstrating no significant association between intraoperative temperature and SSI.<sup>54,64,71</sup> In the lone study examining the effect of intraoperative inspired oxygen, Maragakis et al<sup>54</sup> found that intraoperative administration of fractionated inspired oxygen less than 50% was an independent risk factor for SSI, even after adjusting for other variables. The authors suggested that its effects may be related to the role of oxygen in the bactericidal process of leukocytes.

The argument behind a potential association between intraoperative dural tear and SSI is based on the longer surgical time required to repair a dural tear, as well as the risk of persistent cerebrospinal fluid leakage compromising wound healing. However, no clear relationship between intraoperative dural tear and SSI has been found. Three studies demonstrated no association between dural tear and spinal SSI,<sup>54,60,64</sup> in contrast to a single study finding dural tear to be associated with an increased risk of SSI.<sup>42</sup> In sacral tumors, no definitive association can be made between CSF leak and SSI, as the 2 studies found opposing results.<sup>49,80</sup>

## Discussion

SSIs are associated with many risk factors that can be patient or surgically related. Our review was able to identify important modifiable and nonmodifiable risk factors that can be essential in surgical planning and discussion with patients.

<sup>†</sup>References 1, 6, 11, 16, 22, 33, 42, 43, 49, 52, 67, 80, 86, 92, 93.

Factor	Conclusion
<i>Patient-associated factors</i>	
Age	In general, the literature suggests a mixed finding of association between age and SSI.
Diabetes mellitus (DM)	In general, the literature suggests a strong association between DM/A1c and SSI.
General comorbidities	In general, the literature has mixed finding of specific comorbid conditions in association of SSI. There is evidence to suggest higher number of comorbidities is associated with SSI.
Nutrition	In general, the literature suggests malnutrition is associated with SSI.
Smoking	In general, the literature has mixed results of association between smoking and SSI. More recent evidence would suggest there is correlation between the two.
Obesity/Body mass index	In general, the literature suggests a strong association between obesity and SSI.
<i>Surgery-associated factors</i>	
Time and duration of surgery	In general, the literature is mixed, with conflicting results, making it difficult to firmly establish an association.
Surgical approach/ Invasiveness	In general, the literature is mixed with general trend indicating combined approach have highest incidence of SSI, followed by posterior approach. There is strong evidence increased invasiveness is associated with SSI.
Perioperative interventions	Preoperative radiation and postoperative blood transfusion have strong association with SSI. There is mixed evidence of UTI/ urinary catheter in association of SSI.
Surgical team	In general, there is mixed evidence of resident fellow involvement, number of surgeons and SSI, unable to establish an association.
Intraoperative concerns and complications	There is mixed evidence of intraoperative blood loss, dural tear, hypothermia and SSI, no established association can be made.

### Declaration of Conflicting Interests

The author(s) declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

### Funding

The author(s) disclosed receipt of the following financial support for the research, authorship, and/or publication of this article: This Supplement was supported by funding from AOSSpine North America.

### References

1. Pull ter Gunne AF, Cohen DB. Incidence, prevalence, and analysis of risk factors for surgical site infection following adult spinal surgery. *Spine (Phila Pa 1976)*. 2009;34:1422-1428.
2. Schimmel JJP, Horsting PP, de Kleuver M, Wonders G, van Limbeek J. Risk factors for deep surgical site infections after spinal fusion. *Eur Spine J*. 2010;19:1711-1719.
3. Cizik AM, Lee MJ, Martin BI, et al. Using the Spine Surgical Invasiveness Index to identify risk of surgical site infection. *J Bone Joint Surg Am*. 2012;94:335-342.
4. Abdul-Jabbar A, Takemoto S, Weber MH, et al. Surgical site infection in spinal surgery: description of surgical and patient-based risk factors for postoperative infection using administrative claims data. *Spine (Phila Pa 1976)*. 2012;37:1340-1345.
5. Aoude A, Nooh A, Fortin M, et al. Incidence, predictors, and postoperative complications of blood transfusion in thoracic and lumbar fusion surgery: an analysis of 13 695 patients from the American College of Surgeons National Surgical Quality Improvement Program database. *Global Spine J*. 2016;6:756-764.
6. Asomugha EU, Miller BS, McLain RF. Surgical site infections in posterior lumbar surgery: a controlled-cohort study of epidural steroid paste. *Spine (Phila Pa 1976)*. 2017;42:63-69.
7. Atkinson RA, Stephenson J, Jones A, Ousey KJ. An assessment of key risk factors for surgical site infection in patients undergoing surgery for spinal metastases. *J Wound Care*. 2016;25(suppl 9): S30-S34.
8. Babu R, Owens TR, Thomas S, et al. Timing of tracheostomy after anterior cervical spinal fixation. *J Trauma Acute Care Surg*. 2013;74:961-966.
9. Barnes M, Liew S. The incidence of infection after posterior cervical spine surgery: a 10 year review. *Global Spine J*. 2012; 2:3-6.
10. Berney S, Opdam H, Bellomo R, et al. An assessment of early tracheostomy after anterior cervical stabilization in patients with acute cervical spine trauma. *J Trauma*. 2008;64:749-753.
11. Blam OG, Vaccaro AR, Vanichkachorn JS, et al. Risk factors for surgical site infection in the patient with spinal injury. *Spine (Phila Pa 1976)*. 2003;28:1475-1480.
12. Bohl DD, Shen MR, Mayo BC, et al. Malnutrition predicts infectious and wound complications following posterior lumbar spinal fusion. *Spine (Phila Pa 1976)*. 2016;41:1693-1699.
13. Boston KM, Baraniuk S, O'Heron S, Murray KO. Risk factors for spinal surgical site infection, Houston, Texas. *Infect Control Hosp Epidemiol*. 2009;30:884-889.
14. Browne JA, Cook C, Pietrobon R, Bethel MA, Richardson WJ. Diabetes and early postoperative outcomes following lumbar fusion. *Spine (Phila Pa 1976)*. 2007;32:2214-2219.
15. Chaichana KL, Bydon M, Santiago-Dieppa DR, et al. Risk of infection following posterior instrumented lumbar fusion for degenerative spine disease in 817 consecutive cases. *J Neurosurg Spine*. 2014;20:45-52.
16. Chen S, Anderson MV, Cheng MK, Wongworawat MD. Diabetes associated with increased surgical site infections in spinal arthrodesis. *Clin Orthop Relat Res*. 2009;467:1670-1673.
17. Chen KW, Yang HL, Lu J, et al. Risk factors for postoperative wound infections of sacral chordoma after surgical excision. *J Spinal Disord Tech*. 2011;24:230-234.
18. De la Garza-Ramos R, Bydon M, Abt NB, et al. The impact of obesity on short- and long-term outcomes after lumbar fusion. *Spine (Phila Pa 1976)*. 2014;40:56-61.
19. De la Garza-Ramos R, Abt NB, Kerezoudis P, et al. Deep-wound and organ-space infection after surgery for degenerative spine

- disease: an analysis from 2006-2012. *Neurol Res.* 2016;38:117-123.
- 20. De la Garza-Ramos R, Nakhla J, Nasser R, Bhashyam N, Kinon MD, Yassari R. Case-control study of risk factors for surgical site infection after three-column osteotomy for spine deformity [published online June 2, 2017]. *Turk Neurosurg.* doi:10.5137/1019-5149.JTN.20372-17.0
  - 21. Demura S, Kawahara N, Murakami H, et al. Surgical site infection in spinal metastasis: risk factors and countermeasures. *Spine (Phila Pa 1976).* 2009;34:635-639.
  - 22. Dubory A, Giorgi H, Walter A, et al. Surgical-site infection in spinal injury: incidence and risk factors in a prospective cohort of 518 patients. *Eur Spine J.* 2015;24:543-554.
  - 23. Ee WW, Lau WL, Yeo W, Von Bing Y, Yue WM. Does minimally invasive surgery have a lower risk of surgical site infections compared with open spinal surgery? *Clin Orthop Relat Res.* 2014;472:1718-1724.
  - 24. Fang A, Hu SS, Endres N, Bradford DS. Risk factors for infection after spinal surgery. *Spine (Phila Pa 1976).* 2005;30:1460-1465.
  - 25. Fehlings MG, Smith JS, Kopjar B, et al. Perioperative and delayed complications associated with the surgical treatment of cervical spondylotic myelopathy based on 302 patients from the AO/Spine North America Cervical Spondylotic Myelopathy Study. *J Neurosurg Spine.* 2012;16:425-432.
  - 26. Fisahn C, Jeyamohan S, Norell DC, et al. Association between allogeneic blood transfusion and postoperative infection in major spine surgery. *Clin Spine Surg.* 2017;30:E988-E992.
  - 27. Glassman S, Carreon LY, Andersen M, et al. Predictors of hospital readmission and surgical site infection in the United States, Denmark, and Japan: is risk stratification a universal language? *Spine (Phila Pa 1976).* 2017;42:1311-1315.
  - 28. Golinvaux NS, Varthi AG, Bohl DD, Basques BA, Grauer JN. Complication rates following elective lumbar fusion in patients with diabetes: insulin dependence makes the difference. *Spine (Phila Pa 1976).* 2014;39:1809-1816.
  - 29. Gruskay J, Kepler C, Smith J, Radcliff K, Vaccaro A. Is surgical case order associated with increased infection rate after spine surgery? *Spine (Phila Pa 1976).* 2012;37:1170-1174.
  - 30. Haddad S, Milhouse PW, Maltenfort M, Restrepo C, Kepler CK, Vaccaro AR. Diagnosis and neurologic status as predictors of surgical site infection in primary cervical spine surgery. *Spine J.* 2016;16:632-642.
  - 31. Hayashi H, Murakami H, Demura S, et al. Surgical site infection after total en bloc spondylectomy: risk factors and the preventive new technology. *Spine J.* 2015;15:132-137.
  - 32. Hijas-Gomez AL, Egea-Gamez RM, Martinez-Martin J, Gonzalez-Diaz R, Losada-Vinas JJ, Rodriguez-Caravaca G. Surgical wound infection rates and risk factors in spinal fusion in a university teaching hospital in Madrid, Spain. *Spine (Phila Pa 1976).* 2017;42:748-754.
  - 33. Hikata T, Iwanami A, Hosogane N, et al. High preoperative hemoglobin A1c is a risk factor for surgical site infection after posterior thoracic and lumbar spinal instrumentation surgery. *J Orthop Sci.* 2014;19:223-228.
  - 34. Jalai CM, Worley N, Poorman GW, Cruz DL, Vira S, Passias PG. Surgical site infections following operative management of cervical spondylotic myelopathy: prevalence, predictors of occurrence, and influence on peri-operative outcomes. *Eur Spine J.* 2016; 25:1891-1896.
  - 35. Kanafani ZA, Dakdouki GK, El-Dbouni O, Bawwab T, Kanj SS. Surgical site infections following spinal surgery at a tertiary care center in Lebanon: incidence, microbiology, and risk factors. *Scand J Infect Dis.* 2006;38:589-592.
  - 36. Keam J, Bilsky MH, Laufer I, et al. No association between excessive wound complications and preoperative high-dose, hypofractionated, image-guided radiation therapy for spine metastasis. *J Neurosurg Spine.* 2014;20:411-420.
  - 37. Kerwin AJ, Griffen MM, Tepas JJ 3rd, Schinco MA, Devin T, Frykberg ER. Best practice determination of timing of spinal fracture fixation as defined by analysis of the National Trauma Data Bank. *J Trauma.* 2008;65:824-831.
  - 38. Kim BD, Hsu WK, De Oliveira GS Jr, Saha S, Kim JY. Operative duration as an independent risk factor for postoperative complications in single-level lumbar fusion: an analysis of 4588 surgical cases. *Spine (Phila Pa 1976).* 2014;39:510-520.
  - 39. Kim JL, Park JH, Han SB, Cho IY, Jang KM. Allogeneic blood transfusion is a significant risk factors for surgical-site infection following total hip and knee arthroplasty: a meta-analysis. *J Arthroplasty.* 2017;32:320-325. doi:10.1016/j.arth.2016.08.026
  - 40. Klekamp J, Spengler DM, McNamara MJ, Haas DW. Risk factors associated with methicillin-resistant staphylococcal wound infection after spinal surgery. *J Spinal Disord.* 1999;12:187-191.
  - 41. Klemencsics I, Lazary A, Szoverfi Z, Bozsodi A, Eltes P, Varga PP. Risk factors for surgical site infection in elective routine degenerative lumbar surgeries. *Spine J.* 2016;16:1377-1383.
  - 42. Koutsoumbelis S, Hughes AP, Girardi FP, et al. Risk factors for postoperative infection following posterior lumbar instrumented arthrodesis. *J Bone Joint Surg Am.* 2011;93:1627-1633.
  - 43. Kudo D, Miyakoshi N, Hongo M, et al. Relationship between preoperative serum rapid turnover proteins and early-stage surgical wound infection after spine surgery. *Eur Spine J.* 2017;26:3156-3161.
  - 44. Kukreja S, Ambekar S, Ahmed OI, Mengher RP, Sin AH, Nanda A. Impact of elective versus emergent admission on perioperative complications and resource utilization in lumbar fusion. *Clin Neurol Neurosurg.* 2015;136:52-60.
  - 45. Kumar S, van Popta D, Rodrigues-Pinto R, et al. Risk factors for wound infection in surgery for spinal metastasis. *Eur Spine J.* 2015;24:528-532.
  - 46. Kurtz SM, Lau E, Ong KL, et al. Infection risk for primary and revision instrumented lumbar spine fusion in the Medicare population. *J Neurosurg Spine.* 2012;17:342-347.
  - 47. Lee MJ, Cizik AM, Hamilton D, Chapman JR. Predicting surgical site infection after spine surgery: a validated model using a prospective surgical registry. *Spine J.* 2014;14:2112-2117.
  - 48. Lee JJ, Odeh KI, Holcombe SA, et al. Fat thickness as a risk factor for infection in lumbar spine surgery. *Orthopedics.* 2016;39:e1124-e1128.
  - 49. Li D, Guo W, Qu H, et al. Experience with wound complications after surgery for sacral tumors. *Eur Spine J.* 2013;22:2069-2076.

50. Lieber B, Han BJ, Strom RG, et al. Preoperative predictors of spinal infection within the National Surgical Quality Inpatient Database. *World Neurosurg.* 2016;89:517-524.
51. Lim S, Edelstein AI, Patel AA, Kim BD, Kim JY. Risk factors for postoperative infections following single level lumbar fusion surgery. *Spine (Phila Pa 1976).* 2018;43:215-222.
52. Lonjon G, Dauzac C, Fourniols E, Guigui P, Bonnomet F, Bonneville P; French Orthopaedic Surgery Traumatology Society. Early surgical site infections in adult spinal trauma: a prospective, multicentre study of infection rates and risk factors. *Orthop Traumatol Surg Res.* 2012;98:788-794.
53. Manoso MW, Cizik AM, Bransford RJ, Bellabarba C, Chapman J, Lee MJ. Medicaid status is associated with higher surgical site infection rates after spine surgery. *Spine (Phila Pa 1976).* 2014; 39:1707-1713.
54. Maragakis LL, Cosgrove SE, Martinez EA, Tucker MG, Cohen DB, Perl TM. Intraoperative fraction of inspired oxygen is a modifiable risk factor for surgical site infection after spinal surgery. *Anesthesiology.* 2009;110:556-562.
55. Marquez-Lara A, Nandyala SV, Sankaranarayanan S, Noureldin M, Singh K. Body mass index as a predictor of complications and mortality after lumbar spine surgery. *Spine (Phila Pa 1976).* 2014;39:798-804.
56. Martin CT, Gao Y, Duchman KR, Pugely AJ. The impact of current smoking and smoking cessation on short-term morbidity risk after lumbar spine surgery. *Spine (Phila Pa 1976).* 2016;41: 577-584.
57. Mehta AO, Babu R, Karikari IO, et al. The distribution of body mass as a significant risk factor for lumbar spinal fusion post-operative infections. *Spine (Phila Pa 1976).* 2012;37:1652-1656.
58. Murphy ME, Gilder H, Maloney PR, et al. Lumbar decompression in the elderly: increased age as a risk factor for complications and nonhome discharge. *J Neurosurg Spine.* 2017;26:353-362.
59. Northrup BE, Vaccaro AR, Rosen JE, Balderson RA, Cotler JM. Occurrence of infection in anterior cervical fusion for spinal cord injury after tracheostomy. *Spine (Phila Pa 1976).* 1995;20: 2449-2453.
60. Ogiwara S, Yamazaki T, Maruyama T, et al. Prospective multi-center surveillance and risk factor analysis of deep surgical site infection after posterior thoracic and/or lumbar spinal surgery in adults. *J Orthop Sci.* 2015;20:71-77.
61. Ohya J, Chikuda H, Oichi T, et al. Seasonal variations in the risk of reoperation for surgical site infection following elective spinal fusion surgery: a retrospective study using the Japanese Diagnosis Procedure Combination database. *Spine (Phila Pa 1976).* 2017; 42:1068-1079.
62. Oichi T, Chikuda H, Ohya J, et al. Mortality and morbidity after spinal surgery in patients with Parkinson's disease: a retrospective matched-pair cohort study. *Spine J.* 2017;17:531-537.
63. Ojo OA, Owolabi BS, Oseni AW, Kanu OO, Bankole OB. Surgical site infection in posterior spine surgery. *Niger J Clin Pract.* 2016;19:821-826.
64. Olsen MA, Mayfield J, Lauryssen C, et al. Risk factors for surgical site infection in spinal surgery. *J Neurosurg Spine.* 2003;98: 149-155.
65. Olsen MA, Nepple JJ, Riew KD, et al. Risk factors for surgical site infection following orthopaedic spinal operations. *J Bone Joint Surg Am.* 2008. 90:62-69.
66. Omeis IA, Dhir M, Sciubba DM, et al. Postoperative surgical site infections in patients undergoing spinal tumor surgery. *Spine (Phila Pa 1976).* 2011;36:1410-1419.
67. Pull ter Gunne AF, van Laarhoven CJHM, Cohen DB. Surgical site infection after osteotomy of the adult spine: does type of osteotomy matter? *Spine J.* 2010;10:410-416.
68. Pull ter Gunne AF, van Laarhoven CJHM, Cohen DB. Incidence of surgical site infection following adult spinal deformity surgery: an analysis of patient risk. *Eur Spine J.* 2010;19:982-988.
69. Radcliff KE, Rasouli MR, Neusner A, et al. Preoperative delay of more than 1 hour increases the risk of surgical site infection. *Spine (Phila Pa 1976).* 2013;38:1318-1323.
70. Ramos N, Stachel A, Phillips M, Vigdorchik J, Slover J, Bosco JA. Prior *Staphylococcus aureus* nasal colonization: a risk factor for surgical site infections following decolonization. *J Am Acad Orthop Surg.* 2016;24:880-885.
71. Rao SB, Vasquez G, Harrop J, et al. Risk factors for surgical site infections following spinal fusion procedures: a case-control study. *Clin Infect Dis.* 2011;53:686-692.
72. Rechtine GR, Bono PL, Cahill D, Boileau MJ, Chrin AM. Postoperative wound infection after instrumentation of thoracic and lumbar fractures. *J Orthop Trauma.* 2001;15:566-569.
73. Rodgers WB, Gerber EJ, Patterson J. Intraoperative and early postoperative complications in extreme lateral interbody fusion. *Spine (Phila Pa 1976).* 2010;36:26-33.
74. Ruggieri P, Angelini A, Pala E, Mercuri M. Infections in surgery of primary tumors of the sacrum. *Spine (Phila Pa 1976).* 2012;37: 420-428.
75. Saeednia S, Nouri M, Azarhomayoun A, et al. The incidence and risk factors for surgical site infection after clean spinal operations: a prospective cohort study and review of the literature. *Surg Neurol Int.* 2015;6:154. doi:10.4103/2152-7806.166194
76. Salvetti DJ, Tempel ZJ, Gandhoke GS, et al. Preoperative prealbumin level as a risk factor for surgical site infection following elective spine surgery. *Surg Neurol Int.* 2015;6(suppl 19): S500-S503. doi:10.4103/2152-7806.166893
77. Satake K, Kanemura T, Matsumoto A, Yamaguchi H, Ishikawa Y. Predisposing factors for surgical site infection of spinal instrumentation surgery for diabetes patients. *Eur Spine J.* 2013;22: 1854-1858.
78. Schoenfeld AJ, Carey PA, Cleveland AW, Bader JO, Bono CM. Patient factors, comorbidities, and surgical characteristics that increase mortality and complication risk after spinal arthrodesis: a prognostic study based on 5887 patients. *Spine J.* 2013;13:1171-1179.
79. Schwarzkopf R, Chung C, Park JJ, Walsh M, Spivak JM, Steiger D. Effects of perioperative blood product use on surgical site infection following thoracic and lumbar spinal surgery. *Spine (Phila Pa 1976).* 2010;35:340-346.
80. Sciubba DM, Nelson C, Gok B, et al. Evaluation of factors associated with postoperative infection following sacral tumor resection. *J Neurosurg Spine.* 2008;9:593-599.
81. Sebastian A, Huddleston P 3rd, Kakar S, Habermann E, Wagle Z, Nassr A. Risk factors for surgical site infection after posterior

- cervical spine surgery: an analysis of 5441 patients from the ACS NSQIP 2005-2012. *Spine J.* 2016;16:504-509.
82. Shousha M, Mosafer A, Boehm H. Infection rate after transoral approach for the upper cervical spine. *Spine (Phila Pa 1976).* 2014;39:1578-1583.
  83. Singla A, Yang S, Werner BC, et al. The impact of preoperative epidural injections on postoperative infection in lumbar fusion surgery. *J Neurosurg Spine.* 2017;26:645-649.
  84. Skovrlj B, Cho SK, Caridi JM, Bridwell KH, Lenke LG, Kim YJ. Association between surgeon experience and complication rates in adult scoliosis surgery: a review of 5117 cases from the Scoliosis Research Society database 2004-2007. *Spine (Phila Pa 1976).* 2015;40:1200-1205.
  85. Stambough JL, Beringer D. Postoperative wound infections complicating adult spine surgery. *J Spinal Disord.* 1992;5:277-285.
  86. Sugita S, Hozumi T, Yamakawa K, Goto T, Kondo T. Risk factors for surgical site infection after posterior fixation surgery and intraoperative radiotherapy for spinal metastases. *Eur Spine J.* 2016;25:1034-1038.
  87. Tempel Z, Grandhi R, Maserati M, et al. Prealbumin as a serum biomarker of impaired perioperative nutritional status and risk for surgical site infection after spine surgery. *J Neurol Surg A Cent Eur Neurosurg.* 2015;76:138-143.
  88. Tominaga H, Setoguchi T, Ishidou Y, Nagano S, Yamamoto T, Komiya S. Risk factors for surgical site infection and urinary tract infection after spine surgery. *Eur Spine J.* 2016;25:3908-3915.
  89. Veeravagu A, Patil CG, Shivanand PL, Boakye M. Risk factors for postoperative spinal wound infections after spinal decompression and fusion surgeries. *Spine (Phila Pa 1976).* 2009;34: 1869-1872.
  90. Watanabe M, Sakai D, Matsuyama D, Yamamoto Y, Sato M, Mochida J. Risk factors for surgical site infection following spine surgery: efficacy of intraoperative saline irrigation. *J Neurosurg Spine.* 2010;12:540-546.
  91. Weinstein MA, McCabe JP, Cammisa FP Jr. Postoperative spinal wound infection: a review of 2391 consecutive index procedures. *J Spinal Disord.* 2000;13:422-426.
  92. Wimmer C, Gluch H, Franzreb M, Ogon M. Predisposing factors for infection in spine surgery a survey of 850 spinal procedures. *J Spinal Disord.* 1998;11:124-128.
  93. Woods BI, Rosario BL, Chen A, et al. The association between perioperative allogeneic transfusion volume and postoperative infection in patients following lumbar spine surgery. *J Bone Joint Surg Am.* 2013;95:2105-2110.
  94. Yang S, Werner BC, Cancienne JM, et al. Preoperative epidural injections are associated with increased risk of infection after single-level lumbar decompression. *Spine J.* 2016;16:191-196.
  95. Mirza SK, Deyo RA, Heagerty PJ, et al. Development of an index to characterize the “invasiveness” of spine surgery: validation by comparison to blood loss and operative time. *Spine (Phila Pa 1976).* 2008;33:2651-2661.
  96. Ham WHW, Schoonhoven L, Schuurmans MJ, Leenen LPH. Pressure ulcers, indentation marks and pain from cervical spine immobilization with extrication collars and headblocks: an observational study. *Injury.* 2016;47:1924-1931.
  97. Horvath KA, Acker MA, Chang H, et al. Blood transfusion and infection after cardiac surgery. *Ann Thorac Surg.* 2013;95: 2194-2201.
  98. Oliveira RA, Turrini RNT, Poveda VB. Risk factors for development of surgical site infections among liver transplantation recipients: an integrative literature review. *Am J Infect Control.* 2018;46:88-93. doi:10.1016/j.ajic.2017.05.021
  99. Raghavan M, Marik PE. Anemia, allogenic blood transfusion, and immunomodulation in the critically ill. *Chest.* 2005;127: 295-307.
  100. Vamvakas EC, Blajchman MA. Transfusion-related mortality: the ongoing risks of allogeneic blood transfusion and the available strategies for their prevention. *Blood.* 2009;113:3406-3417.
  101. Nunez-Pereira S, Pellise F, Rodriguez-Pardo D, et al. Individualized antibiotic prophylaxis reduces surgical site infections by gram-negative bacteria in instrumented spinal surgery. *Eur Spine J.* 2011;20(suppl 3):397-402.
  102. Chenoweth CE, Saint S. Urinary tract infections. *Infect Dis Clin North Am.* 2016;30:869-885.
  103. Andersson AE, Bergh I, Karlsson J, Eriksson BI, Nilsson K. Traffic flow in the operating room: an explorative and descriptive study on air quality during orthopedic trauma implant surgery. *Am J Infect Control.* 2012. 40:750-755.
  104. Straub JM, New J, Hamilton CD, Lominska C, Shnayder Y, Thomas SM. Radiation-induced fibrosis: mechanisms and implications for therapy. *J Cancer Res Clin Oncol.* 2015;141: 1984-1994.
  105. Reynolds L, Beckmann J, Kurz A. Perioperative complications of hypothermia. *Best Pract Res Clin Anaesthesiol.* 2008;22: 645-657.