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Management of Pyogenic Spondylodiscitis Following Nonspinal Surgeries: A Tertiary Care Center Experience

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ABSTRACT

Background: The literature is lacking on the incidence and management of pyogenic spondylodiscitis (PS) following routine elective surgical procedures. This study aimed to analyze the presentation and treatment outcome in patients with PS following nonspinal surgeries at a tertiary care center with a minimum follow-up of 12 months.

Methods: The demographic, clinical-radiologic features, and treatment outcomes in 40 patients with a diagnosis of PS following nonspinal surgical procedures were retrospectively reviewed and analyzed.

Results: The mean age at presentation was 36.4 ± 11.8 years, with 80% of patients being female. The common surgical procedures associated with PS were cesarean delivery (30%), gastric sleeve surgery (12.5%), and dilatation and curettage (12.5%). The tissue biopsy culture was positive in 82.5% of patients. A total of 26 patients (65%) were treated with conservative management and 14 patients (35%) were treated surgically. The mean pretreatment Core Outcome Measure Index score significantly decreased at 12 months (P < .0001) after treatment. The mean pretreatment erythrocyte sedimentation rate (P < .0001) and C-reactive protein (P < .0001) levels significantly decreased at 12 months after treatment.

Conclusions: With most patients with PS following nonspinal surgeries treated with conservative management, excellent clinical outcomes were achieved in all patients at 12 months after treatment. The diagnosis of PS should be considered in patients presenting with low back pain (LBP) with a recent history of undergoing a nonspinal surgical procedure. Patients who undergo surgical procedures are an important "at-risk" patient population, and early diagnosis and treatment can help achieve excellent clinical outcomes. Further studies are required to determine risk factors and possible perioperative precautions that can be taken to prevent PS in patients who undergo nonspine surgeries.

Level of Evidence: 3.

Clinical Relevance: Pyogenic spondylodiscitis should be suspected in patients presenting with LBP after a recent non-spinal surgical procedure. Early diagnosis and treatment can help achieve excellent clinical outcomes in these patients.

Research Article

Keywords: pyogenic spondylodiscitis, spinal infections, outcome, infection, surgery

INTRODUCTION

The incidence of pyogenic (nontubercular) spondylodiscitis (PS) has been reported to be up to 5.8 cases per 100 000 person-years, with an average adjusted annual increase of 7.0%.¹ With the increase in the number of the elderly in the general population and the increased prevalence of comorbidities like diabetes, malignancies, and end-stage renal disease, the incidence of PS has risen over the years.^{1,2} Although PS following spinal procedures such as surgery or injections, secondary to immunecompromise, or due to hematogenous spread from an infective focus elsewhere in the body is well reported,^{3–5} PS secondary to elective surgical procedures is rare. Few case reports of PS secondary to genitourinary surgical procedures, such as sacrohysteropexy,⁶ rectopexy,⁷ and transrectal prostate biopsy,⁸ or following major surgeries, such as abdominal aortic aneurysm repair⁹ and lung transplantation,¹⁰ have been reported previously. Surgical procedures are known to cause dysregulation of the immuneinflammatory response and reduce resistance to infection postoperatively.¹¹ Furthermore, gynecologic and genitourinary surgical procedures are known to cause postoperative bacteremia despite prophylactic antibiotics, increasing the risk of bone and joint infection following these procedures.^{12,13} However, the literature is lacking on PS following routine elective surgical procedures, such as casar-

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ean delivery, ligament reconstruction, and bariatric surgery, and the treatment outcome of such patients. Hence, this study aimed to analyze the presentation and treatment outcome in patients with PS following nonspinal surgical procedures with a minimum posttreatment follow-up of 12 months.

METHODS

Study Population

We retrospectively reviewed the records of patients who were treated for PS at the orthopaedic spine surgery department of a tertiary care center between September 2016 and April 2020. The inclusion criterion was all patients with a diagnosis of PS after evaluation (clinical, radiologic, and blood investigations) for low back pain (LBP). The exclusion criteria were PS following spine surgery or patients with a history of spine surgery, history of receiving epidural injection or block for LBP, patients with tubercular spondylodiscitis or spontaneous PS, and patients who did not have a minimum follow-up of 12 months. This study was approved by an institutional ethics committee and has been performed in accordance with the ethical standards as laid down in the 1964 Declaration of Helsinki and its later amendments or comparable ethical standards.

Patient Evaluation

All patients were evaluated with a thorough history and clinical examination at presentation. Demographic data, including sex, age, body mass index, comorbidities, details of past surgeries, or procedures, were collected from all patients. The algorithm proposed by Duarte and Vaccaro¹⁴ was followed in the current study to diagnose and treat PS, which was based on clinical, radiographic, and culture findings. The clinical diagnosis of PS was suspected when patients had symptoms such as nonspecific spinal pain not relieved with rest or pain medication, painful spinal range of motion, neurologic symptoms, such as sensory or motor deficits, associated fever (when present), and presence of associated comorbidities, such as diabetes and prolonged corticosteroid intake.^{2,15} Patients with suspected PS underwent blood investigation to include erythrocyte sedimentation rate (ESR) and C-reactive protein (CRP) and magnetic resonance imaging (MRI) with contrast. In all patients with MRI evidence of vertebral infection (T1-weighted images showing decreased signal intensity in the vertebral body and disk and loss of end plates, and T2-weighted images showing increased signal intensity of the disk and vertebral body),^{16,17} subsequent blood culture and biopsy and culture of intradiscal tissue sample were performed. All patients also underwent a polymerase chain reaction test of the tissue sample to rule out the diagnosis of tuberculosis. The severity of PS was classified into 3 grades (grade I, II, or III) based on the clinical-radiologic scoring system proposed by Homagk et al.¹⁸ For the lumbar spine fluoroscopy-guided biopsies, and for cervical spine computed tomography (CT)–guided biopsies, were performed to collect tissue samples for culture.

Treatment

Conservative treatment of patients included bed rest for an initial 1 week, rigid bracing for 12 weeks, and antibiotic therapy for 12 weeks. For patients with positive biopsy cultures, intravenous (IV) antibiotic treatment was administered for the first 6 weeks followed by oral antibiotics for the next 6 weeks with the 2 most sensitive antibiotics according to culture sensitivity results. Intravenous and oral vancomycin at 15 to 20 mg/kg BID and IV ceftriaxone 1 g BID was the combination commonly used in these patients. For patients with negative biopsy culture, IV and oral antibiotics were administered for 12 weeks based on the blood culture (if positive) or empirically based on broad Gram-positive, Gram-negative, and anaerobic antibacterial coverage. Intravenous ciprofloxacin 500 mg BID, IV ceftriaxone 1 g BID, and oral levofloxacin 500 mg OD was the combination commonly used in these patients. Surgical management was indicated in patients with neurologic deficit or patients with grade II or III PS with significant end plate destruction and features of instability on MRI during presentation, and nonresponse or failure of conservative treatment at the end of 6 weeks, especially in the presence of a negative biopsy culture. Surgical procedures performed as treatment included debridement with posterior pedicle screws fixation, or 360° fusion in 2 stages (anterior followed by posterior), anterior cervical discectomy and fusion, or anterior cervical corpectomy and fusion (ACCF). All patients who underwent surgical management received 6 weeks of IV antibiotics followed by 6 weeks of oral antibiotics based on intraoperative tissue biopsy culture or empirical broad-spectrum antibiotics. Intravenous and oral vancomycin at 15 to 20 mg/kg BID and IV ceftriaxone 1 g BID was the combination commonly used in these patients. Response to treatment was monitored based on improving clinical symptoms and decreasing ESR and CRP levels during the follow-up period. Additional plain spine x-rays were done in all surgical patients to evaluate improvement during the follow-up period.

Outcome Measures

All patients were followed up at 6 weeks, 3 months, 6 months, and 12 months after treatment. Clinical improvement was measured using the Core Outcome Measure Index (COMI) Score, which evaluated pain, function, generic health status or well-being, disability, and satisfaction,¹⁹ and ESR and CRP levels at each follow-up visit.

Statistical Analysis

Pretreatment and at 6 weeks, 3 months, 6 months, and 12 months posttreatment ESR and CRP levels and COMI scores were compared to determine the effectiveness of treatment. Subgroup analysis was performed to compare pretreatment and posttreatment (at 12 months) COMI, ESR, and CRP values in the subgroup treated conservatively versus the subgroup treated surgically. All categoric variables were compared using Fisher test and continuous variables were compared using t test. A Pearson correlation (between 2 continuous variables) and a Spearman ρ correlation (between 1 continuous and 1 categoric variable) analyses were performed to determine correlation between posttreatment COMI score (at 12 months) and pretreatment factors, such as age, sex, site of lesion (lumbar and nonlumbar), duration of symptoms, MRI grade, COMI, ESR, and CRP values. A P value <.05 was considered statistically significant. Statistical analysis was performed using the GraphPad QuickCalcs online statistical analysis tool (GraphPad Software, San Diego, California).

RESULTS

Patient Characteristics

During the study period, a total of 70 patients received a diagnosis of PS. Of these 70 patients, 19 patients were excluded because of postspinal surgery PS, and 11 patients were excluded because of spontaneous (nonsurgical, spinal or nonspinal) PS. Table 1. Demographic data of the study population.

Parameters	Value
No. of patients	40
Age, mean \pm SD (95% CI), y	$36.4 \pm 11.8 (32.6-40.1)$
Sex, n (%)	
Male	8 (20)
Female	32 (80)
Clinical presentation, No. (%)	
Back pain	23 (57.5)
Back pain with radiculopathy	5 (12.5)
Neck pain	10 (25)
Neck pain with radiculopathy	2 (5)
Comorbidities, n (%)	
Diabetes	4 (10)
Hypertension	5 (12.5)
Urinary tract infection	1 (2.5)
Steroid intake	2 (5)
Smoking	3 (7.5)
Clinicoradiologic severity of PS, n (%)	
Grade I	28 (70)
Grade II	5 (12.5)
Grade III	7 (17.5)
Duration of presenting symptoms, mean ± SD (95% confidence interval), days	$15.2 \pm 10.4 (11.8-18.5)$

Abbreviation: PS, pyogenic spondylodiscitis.

Hence, 40 patients of PS that occurred in patients who underwent nonspinal surgery were included and analyzed for this study. The demographic details of the study population are summarized in Table. 1. The mean age at presentation was $36.4 \pm$ 11.8 years, with 80% of patients being female. The mean onset of spine pain after nonspinal surgery was 15.2 ± 10.4 days (range, 5-60 days), with 57.5% of patients presenting with low back pain and 17.5% presenting with sensory and/or motor deficit in the upper or lower limbs (Table 1). Comorbidities were present in 37.5% (15 of 40) of the patients (Table 1).

All patients received general anesthesia, and none of the patients reported any postoperative surgical site infection following their surgical procedure. The common surgical procedures associated with spondylodiscitis were cesarean delivery (30%), gastric sleeve surgery (12.5%), and dilatation and curettage (12.5%; Table 2). The lumbar spine was the most commonly affected region (77.5%), with a single level affected in 95% of patients and the L3 to L4 (23%) and L4 to L5 (23%) levels most commonly affected (Table 3). Based on the scoring system proposed by Homagk et al,¹⁸ there were 28 patients (70%) with grade I PS, 5 patients (12.5%) with grade II PS, and 7 patients (17.5%) with grade III PS (Table 1 and Figure 1). The blood culture was positive in 42.5% of patients, and biopsy culture was positive in 82.5% of patients (Table 4). The common microorganism isolated was Staphylococ-

Table 2. Details of previous surgery done in the study population.

Type of Surgery	Surgical Wound Type	n (%)
Rhinoplasty	Clean contaminated	2 (5)
Laparoscopic cholecystectomy	Clean contaminated	4 (10)
Cesarean section	Clean	12 (30)
Abdominal hernia repair	Clean	1 (2.5)
Syme amputation of the left foot	Contaminated	1 (2.5)
Decompressive drilling of the hip joint	Clean	1 (2.5)
Knee ACL reconstruction	Clean	1 (2.5)
Gastric sleeve surgery	Clean contaminated	5 (12.5)
Diagnostic laparoscopy	Clean	2 (5)
Transvaginal biopsy	Clean contaminated	2 (5)
Breast lump excision	Clean	1 (2.5)
Dilatation and curettage	Contaminated	5 (12.5)
Tonsillectomy	Clean contaminated	2 (5)
Transurethral endoscopic surgery for BPH	Contaminated	1 (2.5)

Abbreviations: ACL, anterior cruciate ligament; BPH, benign prostate hyperplasia.

cus aureus in 20% of patients with positive blood culture, and *S aureus* in 30% of patients and *Escherichia coli* in 20% of patients with positive tissue biopsy culture (Table 4).

A total of 26 patients (65%) were treated with conservative management, and 14 patients (35%) were treated surgically. Three patients were initially treated conservatively and were subsequently treated with surgery (debridement and posterior instrumentation). A total of 10 patients underwent debridement and posterior instrumentation, 2 pa-

Table 3. Magnetic resonance imaging findings in the study population.

Parameter	n (%)
Spinal regions involved	
Lumbar	31 (77.5)
Cervical	8 (20)
Thoracic	1 (2.5)
Patients with the number of spinal levels involved	
1 level	38 (95)
2 levels	1 (2.5)
3 levels	1 (2.5)
Specific spinal levels involved	
C4-C5	4 (9)
C5-C6	5 (11.5)
C6-C7	1 (2.5)
D7-D8	2 (5)
D12-L1	2 (5)
L1-L2	1 (2.5)
L2-L3	3 (7)
L3-L4	10 (23)
L4-L5	10 (23)
L5-S1	5 (11.5)

tients underwent anterior cervical discectomy and fusion, and 1 patient each underwent ACCF (Figure 2) and 360° fusion.

Treatment Outcomes

The mean pretreatment COMI score of 7.3 ± 0.7 significantly decreased to 6.1 ± 0.9 (P < .0001) at 6 weeks, to 2.5 ± 1.5 (P < .0001) at 3 months, to 0.5 ± 0.6 (P < .0001) at 6 months, and to 0.2 ± 0.2 (P < .0001) at 12 months after treatment (Figure 3). The mean pretreatment ESR level significantly



Figure 1. Example of grade I pyogenic spondylodiscitis (PS) at C3 to C4 level (arrow) on magnetic resonance imaging (MRI; left), grade II PS at L4 to L5 level (arrow) on MRI (center), and grade III PS at L3 to L4 level (arrow) on computed tomography scan (right).



Figure 2. Preoperative cervical spine magnetic resonance image of a patient with grade III pyogenic spondylodiscitis at the C4 to C6 levels (left) and cervical spine lateral radiograph of the same patient at 12 months after anterior cervical corpectomy and fusion surgery (right).

decreased at 6 weeks (P = .02), 3 months (P < .0001), 6 months (P < .0001), and 12 months (P < .0001) after treatment (Figure 4). Similarly, the mean CRP value significantly decreased at 6 weeks (P < .0001), 3 months (P < .0001), 6 months (P < .0001), and 1 year (P < .0001) after treatment when

Table 4. Details of culture findings in the study population.

Culture Findings	n (%)
Blood culture	
No growth	23 (57.5)
S aureus	8 (20)
MRSA	2 (5)
E coli	4 (10)
Burkholderia cepacia	1 (2.5)
Proteus sp	1 (2.5)
Klebsiella pneumoniae	1 (2.5)
Tissue biopsy culture	
S aureus	12 (30)
E coli	8 (20)
No growth	7 (17.5)
MRŠA	5 (12.5)
B cepacia	3 (7.5)
Proteus sp	3 (7.5)
K pneumoniae	1 (2.5)
Brucella sp	1 (2.5)

Abbreviations: MRSA, methicillin-resistant S aureus.

compared to the pretreatment value (Figure 4). There was no significant difference between post-treatment COMI, ESR, and CRP values at 12 months when the subgroup treated conservatively was compared to the subgroup treated surgically (Table 5).

Correlation Analysis

There was a weak but significant correlation between the site of lesion (r = -0.31, P = .04) and posttreatment COMI value at 12 months. However,



Figure 3. Mean Core Outcome Measure Index (COMI) scores at 6 weeks, 3 months, 6 months, and 12 months after treatment.



Figure 4. Mean erythrocyte sedimentation rate (ESR) and C-reactive protein (CRP) levels at 6 weeks, 3 months, 6 months, and 12 months after treatment.

no correlation was found between age (r = -0.11, P = .49), sex (r = 0.17, P = .27), duration of symptoms (r = -0.00, P = 1.00), MRI grade (r = 0.00, P = .97), pretreatment COMI (r = -0.10, P = .53), ESR (r = -0.11, P = .49), CRP (r = 0.06, P = .71) values and posttreatment COMI value at 12 months.

DISCUSSION

The current case series aimed to analyze the presentation and treatment outcome in patients with PS following nonspinal surgical procedures with a minimum posttreatment follow-up of 12 months. The most common surgical procedure associated with PS was a cesarean delivery, the most common pathogen responsible was *S aureus*, and the lumbar region was the most commonly affected area in this case series. Furthermore, with most patients treated

with conservative management (65%), excellent clinical outcomes were achieved in all patients, as indicated by sequential improvement in COMI scores and ESR and CRP levels at 6 weeks, 3 months, 6 months, and 12 months after treatment.

Pyogenic spondylodiscitis following nonspinal surgery is a rare condition with few cases published in the literature.^{6–10} Although PS following nonspinal surgery is typically associated with procedures that involve entry into the gastrointestinal or genitourinary tract or in immunocompromised patients or patients with serious comorbidities,^{6–10} the current case series included patients who had undergone a wide variety of surgical procedures, including procedures that are classified as class I or "clean" surgical wound (Table 2) based on the US Centers for Disease Control and Prevention classi-

Table 5. Comparison of Core Outcome Measure Index (COMI), erythrocyte sedimentation rate (ESR), and C-reactive protein (CRP) values between conservative and surgical treatment subgroups.^a

Parameters	Conservative Treatment	Surgical Treatment	P Value
No. of patients	26	14	
Pretreatment COMI score	$7.4 \pm 0.6 (7.1-7.6)$	$6.9 \pm 0.8 \ (6.4-7.3)$.03
Pretreatment ESR level	$64.3 \pm 27.7 (53.1-75.4)$	69.9 ± 33 (50.8-88.9)	.57
Pretreatment CRP level	35.5 ± 21 (27.0-43.9)	$41.6 \pm 17.8 (31.3 - 51.8)$.36
Posttreatment COMI score at 12 mo	$0.3 \pm 0.2 (0.2 - 0.3)$	$0.2 \pm 0.2 (0.0-0.3)$.13
Posttreatment ESR level at 12 mo	19.4 ± 7.5 (16.3-22.4)	20.4 ± 5.2 (17.3-23.4)	.65
Posttreatment CRP level at 12 mo	$2.8 \pm 1.5 (2.1-3.4)$	$2.5 \pm 0.5 (2.2-2.7)$.47

^aAll data presented as mean \pm standard deviation (95% confidence interval); P < .05 is considered statistically significant (bold).

fication of surgical wounds.²⁰ Most patients in the current series were female (male to female sex ratio of 1:4), with most of them undergoing gynecologic, obstetric, and gastrointestinal procedures. Furthermore, the median age of patients in the current series was 34 years (range, 18-65 years), with most (82.5%) of the patients between ages 20 and 50 years. This is contrary to the findings of a previous report on PS by Pola et al,²¹ where the median age was 67 years and most of the patients were male (male to female sex ratio of 1.8:1). This could be explained by the fact that the underlying cause of PS in the study by Pola et al²¹ was secondary to hematogenous spread in patients with comorbidities or patients who were immunocompromised, or following spinal procedures.

The median duration for presentation after the onset of symptoms was 13 days (range, 5-60 days), and most of the patients in the current series presented with LBP (57.5%), 25% of patients presented with neck pain with cervical spine involvement, and 17.5% of patients had a neurologic impairment. Pola et al²¹ in an analysis of 250 patients with PS reported a median duration for presentation after the onset of symptoms of 30 days (range, 15-63 days), with most of the patients presenting with LBP (93.2%) and 23.5% of patients having a neurologic impairment. The greater duration of presentation, predominant lumbar spine affection, and a greater percentage of patients with neurologic impairment could be due to 23.4% of their cases occurring secondary to spine procedures.²¹ In the current series, although blood culture was positive in 42.5% of patients, tissue biopsy culture isolated the causative organism of PS in 82.5% of patients, with S aureus the most common pathogen isolated in 30% of tissue biopsy culture (Table 4). These findings were similar to results published by Sur et al,²² where tissue culture biopsy was negative in 16.5% of patients and the most common isolated organism was S aureus in patients with spontaneous PS.

Most patients (65%) in the current series were treated conservatively, whereas 35% were treated with surgery. The clinical outcome measured by mean COMI score showed significant improvement during 3, 6, and 12 months after treatment. These results are similar to the results reported by Pandita et al,⁵ who reported significant improvement in the Oswestry Disability Index score in 11 patients with subacute, spontaneous PS at the end of 1 year where

54% of patients underwent surgical treatment. There was a significant improvement in blood indicators of infection (ESR and CRP) at 3, 6, and 12 months of treatment in the current series (Figure 2). This was similar to improvements in ESR and CRP levels reported in previous studies following the treatment of PS.^{4,5,21} Furthermore, there was no significant difference in treatment outcomes in terms of the COMI, ESR, and CRP values at 12 months between the conservatively treated and surgically treated subgroups, indicating that conservative treatment had success rates similar to those of surgical treatment in patients with PS following nonspinal surgeries. These findings were similar to those in the study by Tschugg et al,⁴ who reported no significant difference in terms of numeric rating scale for pain, ESR, and CRP values at 12 months between subroups treated conservatively or surgically in patients with both primary acquired and postspinal surgery PS. Among pretreatment factors, such as age, sex, site of lesion (lumbar and nonlumbar), duration of symptoms, MRI grade, COMI, ESR, and CRP values, our study demonstrated a significant weak correlation between mean COMI score at 12 months and the site of lesion. Hence, patients with lumbar spine lesion were associated with lower COMI scores and better clinical outcome at 12 months. This could be because cervical spondylodiscitic lesions have been reported to be associated with more destructive disease and a higher risk of undergoing surgical treatment.²³

In the current series, most patients underwent abdominal, gynecologic, or obstetric surgeries (77.5%; Table 2). Apart from reducing systemic immune resistance to infection postoperatively,¹¹ gynecologic and genitourinary surgical procedures are known to cause postoperative bacteremia despite prophylactic antibiotics, increasing the risk of bone and joint infection.^{12,13} Furthermore, pregnancy and laparoscopic surgery are associated with an increase in intra-abdominal pressure (IAP). An increase in IAP can lead to increased pressure in the venous system of the spine via valveless intraabdominal lumbar veins and increased intrathoracic pressure, causing a decrease in venous drainage from the central nervous system via the jugular system.²³ Hence, a combination of an altered immune response, postoperative bacteremia, and alteration in venous pressure gradient with the rise in IA,P which may promote the venous spread of infection, increases the risk of spinal infection in patients who have undergone nonspinal surgical procedures.^{11–14,24}

Our study has a few limitations. The treatment outcomes achieved in the current series at the end of 12 months need further validation in a larger study population. Furthermore, we did not aim to investigate the risk factors for developing PS following nonspinal surgery because of the small numbers in the current series. Most patients in the current series underwent surgeries with "clean" and "clean-contaminated" surgical wound categories (Table 2), received adequate perioperative antibiotic prophylaxis, and did not report surgical site infection in the postoperative period, and most of the patients had no comorbidities. This indicates that the general health and nutrition of the patient may have played an important role in increasing their risk for developing PS after their primary nonspinal surgery. A study by Lalueza et al²⁵ investigated risk factors for metastatic osteoarticular infections after S aureus bacteremia and reported that patients with osteoporosis and preexisting osteoarticular disease were at high risk for developing a metastatic osteoarticular infection. Hence, risk factors that may put patients who are undergoing nonspinal surgery at risk for PS need further investigation with a larger multicenter study. To the best of our knowledge, the current study is the largest series of patients with PS developed following nonspinal surgery with treatment outcomes reported for 12 months. This series highlights an important "at-risk" patient population who can develop PS.

CONCLUSION

With most patients with PS following nonspinal surgeries treated with conservative management, excellent clinical outcomes were achieved in all patients at 12 months after treatment. Pyogenic spondylodiscitis following nonspinal surgery requires a high degree of suspicion for early diagnosis and treatment. The diagnosis of PS should be considered in patients presenting with LBP with a recent history of undergoing a nonspinal surgical procedure. Early diagnosis and treatment can help achieve excellent clinical outcomes in these patients. Further studies are required to determine risk factors and possible perioperative precautions that can be taken to prevent PS in patients who undergo nonspinal surgeries.

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