# Complication Pattern of Sacral Primary Tumor Resection: A Study on the Risk Factors of Surgical Site Infection and Bowel or Bladder Dysfunction and Their Associations with Length of Hospital Stay

Kristof Koch<sup>1,2</sup>, Peter Pal Varga<sup>1</sup>, Marton Ronai<sup>1</sup>, Istvan Klemencsics<sup>1,3</sup>, Zsolt Szoverfi<sup>1,3</sup>, Aron Lazary<sup>1,3</sup>

<sup>1</sup>National Center for Spinal Disorders, Buda Health Center, Budapest, Hungary <sup>2</sup>School of PhD Studies, Semmelweis University, Budapest, Hungary <sup>3</sup>Department of Spine Surgery, Department of Orthopaedics, Semmelweis University, Budapest, Hungary

Study Design: Retrospective open cohort study.

**Purpose:** The current study aimed to explore the pattern of complications after primary sacral tumor resection, to investigate the possible effect of several perioperative parameters on the development of complications, and to identify which complications are associated with the length of hospital stay (LOS).

**Overview of Literature:** Primary sacral tumor (pST) resection is associated with a high complication rate. However, the number of studies on these complications and their effect on LOS is limited.

**Methods:** The clinical data of 140 patients with pST surgeries and 106 subsequent patients with local recurrence surgeries in four subgroups (index surgery, local recurrence surgery, malignant, and benign tumor) were prospectively collected and analyzed. The prognostic value of several perioperative factors on the development of surgical site infection (SSI), bowel and bladder dysfunction (BBD), and LOS was investigated using the logistic and linear regression models.

**Results**: The overall complication rates were 61.2% after index surgeries and 50.9% after local recurrence surgeries. The most frequent complications were SSI, vegetative dysfunction, urinary tract infections, and neurological deterioration. Age >55 years, malignant tumors, and red blood cell transfusion had a predictive effect on the development of SSI in the logistic model (p<0.01,  $R^2$ =0.43). Bilateral S2 or S3 resection commonly caused postoperative BBD (chi-square test=62.5, degrees of freedom=4, p<0.001). In the multiple linear regression model, wound dehiscence, BBD, systemic and urinary tract infection, cerebrospinal fluid leak, and neurologic deterioration were associated with a significantly long LOS (p<0.01,  $R^2$ =0.62).

**Conclusions:** Surgical resection of pSTs has a high complication rate. Its common complications are SSI and BBD, both of which can have a significant influence on global therapeutic outcome. Malignant tumor diagnosis, old age, and red blood cell transfusion can remarkably increase the risk of SSI. Further, the development of BBD is significantly associated with the number of resected sacral nerve roots. By decreasing perioperative complications, LOS can decrease significantly.

Keywords: Sacrum; Primary sacral tumor; Surgical wound infection; Bladder bowel dysfunction; Risk factors

Received Mar 9, 2023; Revised Jun 2, 2023; Accepted Jun 18, 2023 Corresponding author: Aron Lazary National Center for Spinal Disorders, Buda Health Center, Kiralyhago str. 1-3, Budapest 1126, Hungary Tel: +36-1-887-7900, Fax: +36-1-887-7987, E-mail: aron.lazary@bhc.hu



## Introduction

The surgical management of primary sacral tumors (pSTs) is challenging due to a complicated regional anatomy and limited experience attributed to the rarity of these pathologies [1,2]. pSTs are insidious and can have large volumes before diagnosis, thereby invading the sacral bone, nerve roots, retroperitoneal structures, and posterior soft-tissue elements. Due to the high ratio of aggressive benign (Enneking S3) and low-grade malignant (Enneking IA/B) lesions in this region, the management options are limited. To effectively treat these conditions, surgical resection, enbloc resection in cases of malignant lesions, and even total sacrectomy in some cases are required [3]. Heavy particle radiotherapy is a promising alternative; however, owing to its lack of accessibility, it is not widely used [4]. The local recurrence rate after surgery, even in the case of tumorfree margins, is relatively high [5]. Further, surgical resection for local recurrence can be demanding. In both cases, the complication rate after pST surgery is high at 6%-75% [6]. However, only a few studies have reported the factors causing complications [7-10]. Similarly, some studies have evaluated the effect of complications on length of hospital stay (LOS) after sacral tumor surgeries [11-13], despite the fact that this is one of the most important inpatient indicators of surgical success.

The current study aimed to explore the complication pattern of pST surgeries in different patient subpopulations, to investigate the possible effect of several perioperative parameters on the development of perioperative complications, and to identify which complications are associated with the length of LOS.

## **Materials and Methods**

#### 1. Study design

This retrospective open cohort study was conducted at an international spine tumor referral center. The inclusion criteria were patients diagnosed with pST who underwent surgical resection between January 1995 and December 2019 and those who developed local recurrences after the index surgeries who were treated surgically. The exclusion criteria were patients with secondary sacral tumors, lymphoma, or myeloma, those who only underwent biopsy, and those who had insufficient clinical data. Data collection was performed in a prospective, standardized man-



Fig. 1. Study cohort. pST, primary sacral tumor; LR, local recurrence.

ner (institutional tumor registry) (Fig. 1).

This study was approved by the Scientific and Research Ethics Committee of the National Medical Research Council (30851-2/2016/EKU). The need for patient informed consent was not required owing the nature of the study and the use of deidentified data.

#### 2. Data collection

The following data were collected from the pre- and postoperative inpatient and outpatient clinical records: age, histological diagnosis, detailed medical history, body mass index (BMI), tumor volume, operating room (OR) time, surgical margins, use of implants, nerve resection, volume of blood loss, perioperative transfusion, length of intensive care unit stay, LOS, complications, and local recurrence. LOS was defined as the period between hospital admission and discharge to the patient's home or physical rehabilitation care. The patients remained in the hospital until the treating physicians were certain that no additional surgical care was required to address early postoperative complications. All complications were treated by the same team.

The imaging (computed tomography scan and magnetic resonance imaging) results and histological diagnosis were used to determine localization and tumor volume. The height, width, and depth of the tumor were recorded. The volume was calculated using the formula for an ellipsoid mass (volume= $\pi/6 \times$ height×width×depth) [6]. The detailed medical history of the patients was standardized

using the Charlson comorbidity index.

In addition to categorizing resection (wide-marginal enbloc resection versus intralesional surgery), the oncological outcome of surgery was determined via histopathologic examination (where R0 indicates tumor-free margins; R1, microscopic residuum; and R2, macroscopic tumor remnant). In case of total sacrectomy or if there was SI joint involvement, lumbopelvic reconstruction was performed using different constructs. Sacral resection was categorized according to the Fourney's classification system [14]. In case of midline tumors, low sacral resection was performed at the S4 level and below, midsacral resection at the S3 level and below, and high sacral resection at the S2 level and below. Total sacrectomy was defined as bilateral resection at the S1-S5 levels. Eccentric lesions were resected via sacroiliac joint excision and hemisacrectomy. Nerve resection was categorized as no nerve resection or resection below the S3, bilateral S2, unilateral S2, bilateral S3, and unilateral S3 nerve roots. Perioperative transfusion was assessed based on the number of packed red blood cells (RBCs) transfused. All complications occurred intraoperatively and during the hospital stay and follow-up period. Followup data were obtained via either the direct examination of the patients or telephone interviews and data collection from electronic medical reports. Further, governmental vital statistics databases were accessed. The study data were collected and managed using the institutional REDCap electronic data capture system [15].

#### 3. Statistical analysis

To enhance interpretability, data on index surgeries, local recurrence surgeries, and benign and malignant tumors were analyzed separately, thereby creating four subgroups: index surgery benign tumor (IB), index surgery malignant tumor (IM), local recurrence surgery benign tumor (LRB), and local recurrence malignant surgery (LRM). Index surgery was defined as the first surgery aiming at tumor resection. Biopsy or other previous diagnostic techniques were not considered. The differences between subgroups in the demographic analysis were analyzed using the chi-square test and analysis of variance with the Bonferroni post hoc test. The effects of several perioperative variables on the development of surgical site infection (SSI) were analyzed using univariate and multivariate logistic regression models. All variables identified previously in the literature and those with clinical plausibility were assessed using univariate analysis. A pvalue of <0.1 was considered for a backward multiple regression model. The probability of logistic regression was used as a test variable in the receiver operating characteristic curve analysis to calculate the c-index of the model. The effect of nerve root resection on the development of bowel and bladder dysfunction (BBD) was analyzed using the chi-square test. A linear regression model was used to assess the effect of complications on LOS. The endpoint (LOS in days) was normalized using logarithmic transformation. All complications, other than mortality, cardiac arrest (which decreased the LOS), pressure sores (a complication of LOS), and dural tear (similar to cerebrospinal fluid [CSF] leak), were included in the analysis. Statistical analyses were performed using the IBM SPSS software ver. 27.0 (IBM Corp., Armonk, NY, USA).

## Results

During the study period, 140 patients underwent primary sacral tumor resection. Most patients had a malignant sacral tumor (n=103, 73.5%) (Table 1). Local recurrence

Table 1. Primary sacral tumor histological diagnoses

Variable	No. (%)				
Benign (n=37)					
Schwannoma	15 (41)				
Giant cell tumor	8 (22)				
Osteoblastoma	5 (14)				
Hemangioma	2 (5)				
Osteochondroma	2 (5)				
Fibrosus dysplasia	1 (3)				
Hemangioblastoma	1 (3)				
Myxopapillary ependymoma	1 (3)				
Osteoid osteoma	1 (3)				
Malignant (n=103)					
Chordoma	57 (55)				
Chondrosarcoma	18 (17)				
Ewing sarcoma	8 (8)				
Osteosarcoma	5 (5)				
Fibrosarcoma	2 (2)				
Malignant peripheral nerve sheath tumor	2 (2)				
Myofibrosarcoma	2 (2)				
Neuroblastoma	1 (1)				
Other <sup>a)</sup>	8 (8)				

<sup>a)</sup>Ganglioneuroma, leiomyosarcoma, neurofibroma, other malignant soft tissue tumor, pleomorphic sarcoma, and synovial sarcoma.

(LR) occurred after 46 index surgeries. In total, 106 LR surgeries were performed during this period.

#### 1. Data on demographic and surgical characteristics

Table 2 shows the demographic and clinical characteristics of the four subgroups. The patients were homogeneously distributed in the subgroups according to sex, with a ratio of approximately 1:1. There was a statistically significant difference in terms of age distribution between malignant and benign cases, with a median age of 52 years and 57.5 years in the IM, LRM groups and 33.5 years and 30 years in the IB, LRB groups, respectively. The IM subgroup had the largest tumor volume, with a median of 250 mL (range, 1–6,240 mL). This value was significantly different from that of the IB and LRM groups. Similarly, the IM group (24.3%) commonly received neoadjuvant chemo- and radiotherapy compared with the IB group. Most patients were

Table 2. Demographic and surgical characteristics of the patients

Characteristic	Index surg	ery (N=140)	LR surgery (N=106)		
	Benign (n=37)	Malignant (n=103)	Benign (n=9)	Malignant (n=97)	
Demographic characteristics					
Gender					
Male	21	58	4	50	
Female	16	45	5	47	
Age (yr)	33.5 (13–82)	52 (12-84) <sup>a)</sup>	30 (20–70)	57.5 (16–84) <sup>a)</sup>	
Body mass index (kg/m <sup>2</sup> )	25 (11–32)	25 (15–41)	24 (19–32)	25 (18–36)	
Charlson score	2 (2–6)	2 (2–7)	2 (2–3)	2 (2–10)	
Volume (mL)	62 (1–1,429)	250 (1–6,240) <sup>a)</sup>	63 (2–666)	93 (5–3,204) <sup>b)</sup>	
Preoperative oncotherapy	2 (5.4) <sup>a)</sup>	25 (24.3) <sup>a)</sup>	1 (11.1)	9 (9.3)	
Surgical characteristics					
Total sacrectomy	0 <sup>a)</sup>	10 (9.7) <sup>a)</sup>	0	0	
High sacrectomy	3 (8.1) <sup>a)</sup>	15 (14.5) <sup>a)</sup>	0	0	
Mid sacrectomy	3 (8.1%) <sup>a)</sup>	13 (12.6) <sup>a)</sup>	1 (11.1) <sup>a)</sup>	2 (2.1) <sup>a)</sup>	
Low sacrectomy	2 (5.4%) <sup>a)</sup>	13 (12.6) <sup>a)</sup>	1 (11.1) <sup>a)</sup>	2 (2.1) <sup>a)</sup>	
Hemisacrectomy	2 (5.4%) <sup>a)</sup>	16 (15.5) <sup>a)</sup>	0	1 (1.0)	
Resection type					
En bloc	19	55	2	35	
Intralesional	18	48	7	62	
Resection margin					
RO	19	55	2	35	
R1	14	39	4	25	
R2	4	9	3	37	
Implant	3 (8.1) <sup>a)</sup>	25 (24.3) <sup>a)</sup>	1 (11.1)	8 (8.2)	
Reconstructive plastic procedure	2 (5.4) <sup>a)</sup>	23 (22.3) <sup>a),b)</sup>	1 (11.1)	4 (4.1) <sup>b)</sup>	
Operating room time (min)	110 (30–510) <sup>a)</sup>	200 (25–910) <sup>a),b)</sup>	175 (50–490)	110 (20–540) <sup>b)</sup>	
Blood loss (mL)	800 (50-11,700)	1,650 (50–12,200) <sup>a),b)</sup>	1,200 (100–6,000) <sup>a)</sup>	500 (10–4,250) <sup>a),b)</sup>	
Transfusion (RBC unit)	0 (0–17) <sup>a)</sup>	2 (0-20) <sup>a),b)</sup>	1 (0–21)	0 (0–12) <sup>b)</sup>	
Intensive care unit (day)	0 (0–13) <sup>a)</sup>	3 (0-34) <sup>a),b)</sup>	3 (0–19)	1 (0–17 <sup>)b)</sup>	
Length of hospital stay (day)	11 (3-70) <sup>a)</sup>	19 (3–180) <sup>a),b)</sup>	12 (4–67)	11 (3–141) <sup>b)</sup>	

Values are presented as number, median (min-max), or number (%).

LR, local recurrence; RBC, red blood cell.

a<sup>I</sup>Shows the significant difference (p<0.05) between benign and malignant subgroups. <sup>bI</sup>Shows the significant difference (p<0.05) between the index and LR surgery.

# ASIAN SPINE JOURNAL

slightly overweight, with a median BMI of  $25.0 \text{ kg/m}^2$  and a relatively low median Charlson comorbidity index at 2.

In total, 10 sacrectomy procedures were performed in malignant cases (Table 2). The IM group commonly underwent wide resections. Meanwhile, the LRM group frequently had intralesional resection. Implants were used in 24.3% and 8.1% of malignant and benign index surgeries. Approximately 12.2% of surgeries involved a complex plastic closure, and there was a significant difference in the need for reconstructive plastic procedures between the IM, IB, and LRM groups. The IM group had a significantly high median volume of blood loss (1,650 mL), operating room time (200 minutes), and median number of pack RBCs transfused (2 packs). Similarly, among all groups, the IM group had the highest median length of intensive care unit stay and LOS (3 days and 19 days, respectively).

#### 2. Complication rate

In our cohort, pST surgeries were characterized by a high overall complication rate at 56.5%. In total, 85 (61.2%) and 54 (50.9%) patients developed complications after index surgeries and LR surgeries, respectively. The number of complications per surgery ranged from 0 to 9, with at

#### Table 3. The distribution of complications between different groups

Variable	Overall	Index surg	Index surgery (N=140)		ery (N=106)
	Uverali	Benign (n=37)	Malignant (n=103)	Benign (n=9)	Malignant (n=97)
Intraoperative complications					
Dural tear	11 (4.47)	3 (8.1)	5 (4.9)	-	3 (3.1)
Cerebrospinal fluid leak	4 (1.6)	1 (2.7)	3 (2.9)	-	-
Massive blood loss	31 (12.6)	3 (8.1)	12 (11.7)	6 (66.7)	10 (10.3)
Vascular injury	10 (4.1)	1 (2.7)	5 (4.9)	-	4 (4.1)
Visceral injury	4 (1.6)	-	4 (3.9)	-	-
Nerve root injury	15 (6.1)	-	8 (7.8)	-	7 (7.2)
Neurologic complication					
Neurological deterioration	15 (6.1)	1 (2.7)	7 (6.8)	-	7 (7.2)
Postoperative neuropathy	7 (2.8)	2 (5.4)	2 (1.9)	1 (11.1)	2 (2.1)
Bowel and bladder dysfunction	40 (16.2)	4 (10.8)	31 (30.1)	-	5 (5.1)
Septic complications					
Surgical site infection	62 (25.2)	2 (5.4)	42 (40.8)	1 (11.1)	17 (17.5)
Wound dehiscence	14 (5.7)	2 (5.4)	8 (7.8)	1 (11.1)	3 (3.1)
Systemic infection	22 (8.9)	2 (5.4)	13 (12.6)	-	7 (7.2)
Clostridium infection	4 (1.6)	1 (2.7)	3 (2.9)	-	-
Urinary tract infection	12 (4.8)	3 (8.1)	4 (3.9)	-	5 (5.2)
Other complications					
Deep vein thrombosis	8 (3.3)	1 (2.7)	3 (2.9)	-	4 (4.1)
Pressure sores	3 (1.2)	-	2 (1.9)	1 (11.1)	-
Hematoma	6 (2.4)	-	3 (2.9)	2 (22.2)	1 (1.0)
Intestinal obstruction/colostomy	5 (2.0)	-	2 (1.9)	1 (11.1)	2 (2.1)
Construct failure	3 (1.2)	-	1 (4.0) <sup>a)</sup>	-	2 (25.0)
Death	12 (4.8)	-	7 (6.8)	-	5 (5.2)
Other <sup>b)</sup>	16 (6.5)	2 (5.4)	10 (9.7)	-	4 (4.1)

Values are presented as number (%).

LR, local recurrence.

<sup>a</sup>)Shows the percentage distribution of all the implant-use surgeries in the same subgroups. <sup>b</sup>Abdominal pain, pneumonia, hearth failure, delirium, diarrhea, intraabdominal bleeding, lower extremity arterial occlusion, peritonitis, pleural effusion, rectal prolapse, renal failure, vaginal infection, anesthesia-related, colitis, herpes zoster, and pulmonary effusion. least one complication in 50 cases, 2 in 36 cases, 3 in 17 cases, 4 in 14 cases, and 5 in eight cases, and from 6 to 9 in 14 cases. Table 3 shows the distribution of complications among the IM, IB, LRM, and LRB groups. SSI and BBD were the most frequent complications, followed by a significantly high volume of blood loss and systemic infections. We lost 12 patients (4.8%) in the perioperative period (n=7 in the IM group and n=5 in the LRM group).

#### 3. Surgical site infection

Wound complications were identified after 76 surgeries (30.9%) (wound dehiscence in 14 cases [5.7%] and SSI in 62 cases [25.2%]). SSI commonly occurred in the IM and LRM groups (42 [40.8%] versus 17 [17.5%]). Approximately 66.7% of patients in the IM group, 58.8% in the LRM group, and 100% in the LRB, SSI groups required one or more debridement surgeries. Systemic infection occurred due to SSI in 26.2% of patients in the IM group and 41.2% of patients in the LRM, SSI groups. Six patients died due to septic complications in the IM group and three in the LRM group (Table 4).

The effects of several perioperative variables (malignant/ benign histology, BMI, age, previous sacral tumor surgery, Charlson comorbidity index, operating room time, tumor volume, volume of blood loss, transfusion, surgical margins, implant use, preoperative oncotherapy, dural tear, urinary tract infection, and postoperative BBD) on the development of SSI were analyzed using univariate logistic regression models (Table 5). Based on these factors, histological diagnosis, age, implant use, volume of blood loss, tumor volume, BMI, postoperative BBD, and transfusion were entered into a backward multiple logistic regression model. In the final model, age (odds ratio, 3.0), transfusion (odds ratio, 1.3), and malignant histology (odds ratio,

 $\ensuremath{\textbf{Table 4}}\xspace.$  The distribution of SSI and related complications in the different groups

	Index sur	Index surgery (N=44) LR surger		
Variable	Benign (n=2)	Malignant (n=42)	Benign (n=1)	Malignant (n=17)
Duration of SSI (day)	30 (5–51)	39 (25–316)	34 (34–34)	30 (15–127)
Systemic infection	1 (50.0)	11 (26.2)	-	7 (41.2)
Surgery for SSI	-	28 (66.7)	1 (100.0)	10 (58.8)
Death	-	6 (14.3)	-	3 (17.6)

Values are presented as median (min-max) or number (%). SSI, surgical site infection; LR, local recurrence.

Table 5. Univariate logistic regression analysis of perioperative factors on SSI in index surgeries

Variable	B±SE	Wald	OR (95% CI)	<i>p</i> -value
Body mass index	0.060±0.033	3.417	1.062 (0.996–1.32)	0.065
Age >55 yr	1.004±0.300	11.201	2.729 (1.516–4.914)	0.001
Gender	0.136±0.291	0.218	1.146 (0.647–2.027)	0.641
Previous sacral tumor surgery	0.140±0.092	2.292	0.870 (0.726–1.042)	0.130
Malignancy	1.504±0.546	7.601	4.500 (1.545–13.109)	0.006
Charlson score	0.115±0.136	0.709	1.121 (0.859–1.464)	0.400
Operating room time (hr)	0.314±0.059	28.567	1.368 (1.220–1.535)	<0.001
Tumor volume (L)	0.649±0.236	7.586	1.914 (1.206–3.039)	0.006
Blood loss (L)	0.524±0.099	28.184	1.689 (1.392–2.050)	<0.001
Transfusion	0.258±0.039	44.465	1.294 (1.200–1.396)	<0.001
Resection type	0.096±0.292	0.107	0.909 (0.513–1.609)	0.743
Resection margin	0.174±0.229	0.577	0.840 (0.537–1.316)	0.448
Implant use	1.484±0.371	16.009	0.227 (0.110-0.469)	<0.001
Preoperative oncotherapy	0.105±0.414	0.065	0.900 (0.400–2.027)	0.799
Dural tear	0.911±0.624	2.130	2.486 (0.732-8.455)	0.144
Urinary tract infection	0.751±0.605	1.542	2.119 (0.648–6.929)	0.214
Postoperative BBD	0.846±0.298	8.078	2.329 (1.300–4.173)	0.004

Statistically significant results are marked in bold.

SSI, surgical site infection; SE, standard error; OR, odds ratio; CI, confidence interval; BBD, bowel and bladder dysfunction.

4.5) were significant risk factors for SSI (model: p<0.01, chi-square test=85.97,  $R^2$ =0.43, c-index=0.863).

#### **4. BBD**

Of 140 index patients, 12 (8.5%) had preoperative BBD, and only two of them recovered after surgery. Among 128 patients without preoperative BBD symptoms before index surgery, 35 (27.3%) developed BBD after surgery (Table 6). The development of postoperative BBD was strongly associated with the level of nerve root resection (chi-square test=62.5, df=4, p<0.001). In total, 6 (6.9%) of 87 patients who had no nerve root resection or only below the S3 level developed postoperative BBD. Moreover, 25 patients underwent bilateral S2 resection, and only eight patients had unilateral S2 loss. In both groups, approximately 75% of patients developed postoperative BBD. The incidence rate of postoperative BBD in patients who underwent bilateral S3 resection was 60%, and unilateral S3 resection caused BBD in 33% of cases.

#### 5. Length of hospital stay

The median LOSs were 14 (3-180) days in the whole co-

#### Table 7. Multiple linear regression analysis of complications on LOS

hort, 19 (3–180) days in the IM subgroup, and 11 (3–70) days in the IB subgroup (p<0.05). The median LOSs were 12 (4–67) days among patients who underwent local recurrence surgery in the LRB subgroup and 11 (1–141) days among those in the LRM subgroup. In the multiple linear regression model, SSI, wound dehiscence, BBD, systemic infection, urinary tract infection, CSF leak, and neurological deterioration had a significant effect on LOS. Approximately 62% of the total LOS variance was ex-

 
 Table 6. The effect of different nerve resections on BBD among patients with or without preoperative BBD

Variable	No preoperative BBD (N=128)			
Vallable	No postop BBD (n=93)	Postop BBD (n=35)		
Both S2	6 (24.0)	19 (76.0)		
Unilateral S2	2 (25.0)	6 (75.0)		
Both S3	2 (40.0)	3 (60.0)		
Unilateral S3	2 (66.7)	1 (33.3)		
No nerve resection or below S3	81 (93.1)	6 (6.9)		

Values are presented as number (%). The percentage distribution calculated by rows according to the level of nerve root resection (chi-square=62.5, df=4,  $\rho$ <0.001).

BBD, bowel and bladder dysfunction; Postop, postoperative; df, degrees of freedom.

Variable	B±SE	<i>t</i> -value	Significance	Tolerance	VIF
Surgical site infection	1.09±0.10	10.89	<0.01	0.68	1.46
Wound dehiscence	0.91±0.17	5.44	<0.01	0.87	1.15
Bowel and bladder dysfunction	0.49±0.10	4.74	<0.01	0.75	1.32
Systemic infection	0.52±0.16	3.35	<0.01	0.66	1.50
Urinary tract infection	0.42±0.18	2.22	0.02	0.86	1.16
Cerebrospinal fluid leak	0.64±0.32	2.01	0.04	0.81	1.12
Neurologic deterioration	0.37±0.16	2.27	0.02	0.85	1.17
Hematoma	0.48±0.30	1.62	0.10	0.61	1.61
Nerve root injury	0.06±0.17	0.39	0.69	0.79	1.13
Deep vein thrombosis	0.24±0.22	1.08	0.28	0.87	1.15
lleus	0.22±0.32	0.68	0.49	0.66	1.51
Vascular injury	-0.03±0.21	-0.14	0.89	0.77	1.23
Construct failure	-0.45±0.36	-1.26	0.21	0.84	1.19
Clostridium infection	-0.09±0.32	-0.28	0.78	0.81	1.24
Massive blood loss	0.01±0.14	0.01	0.99	0.66	1.52
Visceral injury	-0.38±0.31	-0.01	0.22	0.85	1.17
Postoperative neuropathy	-0.34±0.24	-1.43	0.16	0.86	1.16

Statistically significant results are marked in bold. Approximately 62% of the total LOS variance was explained by the final mode ( $\rho$ <0.01,  $R^2$ =0.62). LOS, length of hospital stay; SE, standard error; VIF, variance inflation factor.

plained by the final mode (p<0.01,  $R^2$ =0.62) (Table 7).

## **Discussion**

As pSTs are rare and histologically heterogeneous, only a few studies have described the complication patterns of sacral tumor resections in different patient subpopulations, including those who underwent index versus LR surgeries and those with benign, malignant tumors. In our cohort comprising 140 patients with surgically treated pSTs and 106 subsequent patients with LR surgeries, the overall complication rates were 61.2% after index surgeries and 50.9% after local recurrences. The overall complication rate was higher in malignant pST surgeries in our cohort, which was consistent with the results of other studies. Angelini et al. showed that the complication rate can reach 80% in sacral chordoma resections [9]. However, it can be significantly low (17%) in patients with benign lesions [16].

Index surgeries are associated with a higher complication rate in our cohort. One of the reasons can be surgical invasiveness because wide *en-bloc* resection is the gold standard surgical therapy for malignant sacral tumors [1,3,6]. Further, there is a cautious follow-up protocol after index sacral tumor surgery at our institution. Hence, local recurrences are identified at an earlier phase. This notion is supported by data showing that the average tumor volume is smaller in cases of LR surgeries, which assumes less nerve, vessel involvement and smaller surgical wound, which can result in a lower risk of complications.

The most common complications were SSI and BBD. The incidence of SSI in our overall cohort was 25.2%, which is in the lower third of the published values (14.8%-44%) [6,8,12,17,18]. Malignancy was a significant predictor for developing SSI even in the multivariate model in our analysis and in other studies [8,19]. SSI and its predictors were analyzed in other studies that showed a high incidence rate but on a variable group of predictors. Li et al. [20] reported 29.2% of cases of wound-related complications in a large sacral tumor cohort. Results showed that previous irradiation, rectal injury, longer duration of surgery, and CFS leakage were significantly associated with SSI development. Chen et al. [21] conducted a retrospective analysis to assess the SSI rate after sacral chordoma resection. Of 45 patients, 16 (35.6%) developed postoperative wound infections. Significant risk factors were low albumin levels (<3.0 g/dL), long surgery dura-

tion (>6 hours), and previous history of surgery. Similarly, Sciubba et al. [17] reported an SSI rate of 39% in 46 patients with surgically treated malignant sacral tumors. In their cohort, previous history of surgery, number of surgeons involved in the surgery, old age, presence of complex soft-tissue reconstruction, and BBD were associated with a higher risk of SSI. In contrast, Marmouset et al. [22] did not find an association between bowel incontinence and SSI after en-bloc sacrectomy. In our cohort, BBD was associated with SSI in the univariate analysis. However, it lost its significance in the multivariate model. SSI after spinal surgical procedures is generally more frequent in elderly individuals [22-24], which is in accordance with our results, where age >55 years has resulted in a 3-fold risk of having an SSI after sPT surgeries. sPT resection is commonly characterized by a significant volume of blood loss, and patients often require RBC transfusion during the intraoperative and/or postoperative periods. A previous study showed the immunosuppressive effect of homologous blood transfusion [25]. De la Garza Ramos et al. [26] reported that transfusion after metastatic spinal tumor surgery significantly increased the overall complication rate. Results showed a similar prognostic effect of transfusion on the development of SSI in our pST cohort, and the association of which remained significant in the multivariate model. Nakai et al. [27] emphasized the role of autologous blood transfusion in sacral tumor resection, which is the mechanism underlying the association.

The incidence of BBD (16.2%) in this research is lower than that in other studies, which is 38%-78% depending on the composition of the study cohort [6,8,12,17,18]. The results of published data about the association between the level of nerve root resections and the risk for BBD vary. Our results support the findings of Fourney et al. [14] who reported that patients with amputations distal to the S3 levels only generally experienced minor or no deficit, with the preservation of sphincter function in most cases. The S2 roots are often considered vital for bowel, urinary, and sexual functions, and several authors have published that the preservation of the bilateral S2 roots decreases the chance of BBD. However, a significant group of patients developed BBD after unilateral S2 or bilateral S3 root resections. This result is not in accordance with previous findings [28,29]. However, it supports the results of a systematic review conducted by Zoccali et al. [28]. Accordingly, in a case series of 50 sacrectomy procedures, Guo et al. [12] found a close association between resection of the S3 nerve roots and the development of BBD. Our study supports their findings. However, we must emphasize the fact that even unilateral S3 resection can lead to BBD in a third of these cases.

LOS is among the few parameters widely used to characterize the complexity, severity, and economic consequences of a medical condition or treatment. However, only few papers have published this parameter in case of pST surgeries. The median LOS in our study was shorter than that of other publications. Moreover, SSI, wound dehiscence, BBD, systemic and urinary tract infections, CSF leak, and neurologic deterioration had a prolonged effect on LOS after sacral tumor resections. In case of 75 index (53.5%) and 32 LR (30.2%) surgeries, at least one of these LOS prolonging complications occurred. Guo et al. [12] found that the median LOSs after sacrectomy procedures were 48.5 days in case of colostomy and 18.5 days in participants without colostomy. Moreover, patients who underwent surgery using a myocutaneous flap and those with BBD had a longer LOS [12]. Feghali et al. [13] showed that iliac vessel involvement, larger tumor volume, staged procedure, and S1 nerve root sacrifice were independent predictors of prolonged LOS after sacral tumor surgery.

The current study had some strengths and limitations. In terms of limitations, previous studies on this topic are related to the heterogeneous histological distribution of tumors and different treatment principles according to the Enneking's criteria. Further, the data collection had a relatively long-time frame. However, the principles of surgical treatment for pSTs have not changed during this period. Finally, an external model validation was not performed. Regarding the study's strengths, a relatively large patient population was included, and prospectively collected data were mainly analyzed. The high number of cases allowed us to analyze the effect of tumor dignity on the complication pattern, thereby emphasizing that a malignant histology is an independent risk factor of SSI in pSTs. Nevertheless, further studies on the risk factors of various complications should be performed using a large multicentric dataset by incorporating other important outcome variables, such as quality of life.

## Conclusions

To the best of our knowledge, to date, this is the largest study focusing on complication patterns after primary

sacral tumor surgeries. The complication rates were high at 61.2% after index surgeries and 50.9% after LR surgeries. Malignant tumor diagnosis, age >55 years, and number of pack RBCs transfused had a predictive effect on the emergence of SSI, which is among the most common types of complications. Bilateral S2 or S3 nerve root resection commonly resulted in postoperative BBD. However, this severe complication was also frequently observed in unilateral S2 resection and occurred in cases of unilateral S3 nerve root resection. SSI, wound dehiscence, BBD, systemic and urinary tract infection, CSF leak, and neurological deterioration are associated with an increased risk of prolonged LOS. Based on our findings, SSI and BBD are the most common complications of pST surgeries. Moreover, malignant tumor diagnosis, old age, and RBC transfusion remarkably increased the risk of SSI, and LOS can decrease significantly by decreasing perioperative complications.

## **Conflict of Interest**

No potential conflict of interest relevant to this article was reported.

## Funding

This work was supported by the UNKP-21-5 New National Excellence Program of the Ministry for Innovation and Technology from the source of the National Research, Development, and Innovation Fund as well as by the by the János Bolyai Research Scholarship of the Hungarian Academy of Sciences.

## ORCID

Kristof Koch: https://orcid.org/0000-0003-4847-8647 Peter Pal Varga: https://orcid.org/0000-0002-3485-9396 Marton Ronai: https://orcid.org/0009-0009-2169-7832 Istvan Klemencsics: https://orcid.org/0000-0001-9168-2833 Zsolt Szoverfi: https://orcid.org/0000-0003-3535-7395 Aron Lazary: https://orcid.org/0000-0002-0157-719X

## **Author Contributions**

All authors contributed to the study conception and design. Material preparation, data collection, and analysis were performed by KK. The first draft of the manuscript was written by KK, PPV, and ZS, and all authors commented on previous versions of the manuscript. All authors read and approved the final manuscript. All the authors agree to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

## References

- 1. Fisher CG, Goldschlager T, Boriani S, et al. A novel scientific model for rare and often neglected neoplastic conditions. Evid Based Spine Care J 2013;4:160-2.
- Varga PP, Szoverfi Z, Lazary A. Surgical treatment of primary malignant tumors of the sacrum. Neurol Res 2014;36:577-87.
- Dea N, Fisher CG, Reynolds JJ, et al. Current treatment strategy for newly diagnosed chordoma of the mobile spine and sacrum: results of an international survey. J Neurosurg Spine 2018;30:119-25.
- 4. Bible JE, Mirza M, Knaub MA. Blood-loss management in spine surgery. J Am Acad Orthop Surg 2018;26:35-44.
- Varga PP, Szoverfi Z, Fisher CG, et al. Surgical treatment of sacral chordoma: prognostic variables for local recurrence and overall survival. Eur Spine J 2015;24:1092-101.
- Verlaan JJ, Kuperus JS, Slooff WB, Hennipman A, Oner FC. Complications, secondary interventions and long term morbidity after en bloc sacrectomy. Eur Spine J 2015;24:2209-19.
- Dolan RT, Butler JS, Wilson-MacDonald J, et al. Quality of life and surgical outcomes after soft-tissue reconstruction of complex oncologic defects of the spine and sacrum. J Bone Joint Surg Am 2016;98:117-26.
- 8. Ruggieri P, Angelini A, Pala E, Mercuri M. Infections in surgery of primary tumors of the sacrum. Spine (Phila Pa 1976) 2012;37:420-8.
- Angelini A, Pala E, Calabro T, Maraldi M, Ruggieri P. Prognostic factors in surgical resection of sacral chordoma. J Surg Oncol 2015;112:344-51.
- Arikan M, Togral G, Hasturk AE, Aktas E, Gungor S. Management and retrospective analysis of primary and metastatic sacral tumors and infections: evaluation with 73 cases. Eklem Hastalik Cerrahisi 2014;25:126-32.
- 11. Gao S, Zheng Y, Liu X, Tian Z, Zhao Y. Effect of early

fasting and total parenteral nutrition support on the healing of incision and nutritional status in patients after sacrectomy. Orthop Traumatol Surg Res 2018;104:539-44.

- Guo Y, Palmer JL, Shen L, et al. Bowel and bladder continence, wound healing, and functional outcomes in patients who underwent sacrectomy. J Neurosurg Spine 2005;3:106-10.
- Feghali J, Pennington Z, Hung B, et al. Sacrectomy for sacral tumors: perioperative outcomes in a large-volume comprehensive cancer center. Spine J 2021;21:1908-19.
- Fourney DR, Rhines LD, Hentschel SJ, et al. En bloc resection of primary sacral tumors: classification of surgical approaches and outcome. J Neurosurg Spine 2005;3:111-22.
- 15. Harris PA, Taylor R, Thielke R, Payne J, Gonzalez N, Conde JG. Research electronic data capture (RED-Cap): a metadata-driven methodology and workflow process for providing translational research informatics support. J Biomed Inform 2009;42:377-81.
- Zang J, Guo W, Yang Y, Wei R. Surgical treatment of giant benign sacral neurogenic tumors using the posterior-only approach. Clin Neurol Neurosurg 2019;185:105483.
- Sciubba DM, Nelson C, Gok B, et al. Evaluation of factors associated with postoperative infection following sacral tumor resection. J Neurosurg Spine 2008;9:593-9.
- Moran D, Zadnik PL, Taylor T, et al. Maintenance of bowel, bladder, and motor functions after sacrectomy. Spine J 2015;15:222-9.
- Ruggieri P, Angelini A, Ussia G, Montalti M, Mercuri M. Surgical margins and local control in resection of sacral chordomas. Clin Orthop Relat Res 2010;468:2939-47.
- 20. Li D, Guo W, Qu H, et al. Experience with wound complications after surgery for sacral tumors. Eur Spine J 2013;22:2069-76.
- 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;24:230-4.
- 22. Marmouset D, Haseny B, Dukan R, et al. Characteristics, survivals and risk factors of surgical site infections after en bloc sacrectomy for primary malignant sacral tumors at a single center. Orthop Traumatol Surg Res 2022;108:103197.

# ASIAN SPINE JOURNAL

- 23. 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;16:1377-83.
- 24. Fang A, Hu SS, Endres N, Bradford DS. Risk factors for infection after spinal surgery. Spine (Phila Pa 1976) 2005;30:1460-5.
- 25. Mezrow CK, Bergstein I, Tartter PI. Postoperative infections following autologous and homologous blood transfusions. Transfusion 1992;32:27-30.
- 26. De la Garza Ramos R, Gelfand Y, Benton JA, et al. Rates, risk factors, and complications of red blood cell transfusion in metastatic spinal tumor surgery: an analysis of a prospective multicenter surgical database. World Neurosurg 2020;139:e308-15.
- Nakai S, Yoshizawa H, Kobayashi S, Naga K, Ichinose H. Role of autologous blood transfusion in sacral tumor resection: patient selection and recovery after surgery and blood donation. J Orthop Sci 2000;5:321-7.
- Zoccali C, Skoch J, Patel AS, Walter CM, Maykowski P, Baaj AA. Residual neurological function after sacral root resection during en-bloc sacrectomy: a systematic review. Eur Spine J 2016;25:3925-31.
- 29. Gunterberg B, Kewenter J, Petersen I, Stener B. Anorectal function after major resections of the sacrum with bilateral or unilateral sacrifice of sacral nerves. Br J Surg 1976;63:546-54.