



The role of lumbosacral paraspinal muscle degeneration and low vertebral bone mineral density on distal instrumentation-related problems following long-instrumented spinal fusion for degenerative lumbar scoliosis: a retrospective cohort study

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Background: This study aimed to confirm the role of paraspinal muscle degeneration and low vertebral bone mineral density (vBMD) of the lumbosacral region in the development of distal instrumentation-related problems (DIPs) in degenerative lumbar scoliosis (DLS) patients undergoing long-instrumented spinal fusion.

Methods: From 2013 to 2019, 125 DLS patients with 24-month follow-up after long-instrumented spinal fusion in Beijing Chao-Yang Hospital were retrospectively recruited and divided into DIP and non-DIP groups. Demographic characteristics, surgical data, and radiographic parameters were statistically compared between the groups. Degeneration of the paraspinal muscle was evaluated using the relative gross cross-sectional area (rGCSA), relative functional cross-sectional area (rFCSA), ratio of the rFCSA to rGCSA, gross muscle-fat index, and functional muscle-fat index of the multifidus (MF), erector spinae (ES), paraspinal extensor muscle (PSE), and psoas major determined by preoperative magnetic resonance imaging (MRI). The vBMD of the lumbosacral region and lower instrumented vertebra (LIV) was assessed using Hounsfield unit (HU) values determined by computed tomography (CT) scans. The DeLong test was performed to select MRI and CT scan variables. Multivariable logistic regression analysis was applied to determine the independent predictive factors of DIPs.

Results: The incidence of DIPs was 16.0% (20/105). There were no significant differences in demographic characteristics or surgical data between the groups. The rFCSAs of the MF (65.74±21.51 vs. 92.37±21.68; P<0.001), ES (82.67±21.44 vs. 111.48±24.21; P<0.001) and PSE (144.31±36.12 vs. 208.48±41.57; P<0.001) and the HU values of the lumbosacral region (103.80±22.64 vs. 132.19±19.17; P<0.001) and LIV (111.70±23.23 vs. 128.69±20.70; P=0.005) were significantly lower in the DIP group. Significantly less preoperative pelvic tilt and greater postoperative lumbosacral lordosis and sagittal vertical axis (SVA) values were observed in the DIP group. The rFCSA of the PSE, the HU value of the lumbosacral region, and the postoperative SVA value were detected as independent predictive factors of DIPs.

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Conclusions: Lower muscularity of the PSE, a lower vBMD of the lumbosacral region, and postoperative sagittal malalignment were independent predictive factors of DIPs. Surgeons should emphasize the preoperative evaluation of paraspinal muscle and bone mass in DLS patients.

Keywords: Lumbosacral paraspinal muscle degeneration; fatty infiltration; low vertebral bone mineral density; degenerative lumbar scoliosis; distal instrumentation-related problems

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Introduction

Degenerative lumbar scoliosis (DLS) is a kind of adult spinal deformity defined as a lumbar Cobb angle greater than 10° (1,2). The prevalence of DLS ranges from 7.5% to 68.0% worldwide and continues to increase with the aging of the population (3). Severe low back and leg pain could be caused by lateral olisthesis-induced or rotatory subluxation-induced foraminal stenosis, persistently impacting patients' daily activities (4-7). If relief is not achieved with conservative treatment, decompression and restoration of spinal alignment by long-instrumented spinal fusion can be considered.

A high complication rate of long-instrumented spinal fusion for DLS has been reported, and the most common complication is related to instrumentation, with an incidence ranging from 12.4% to 31.7% (8-10). Distal instrumentation-related problems (DIPs) are a series of complications related to the lower instrumented vertebra (LIV), including distal junctional kyphosis (11), screw loosening or breakage (12), fracture of the LIV (13), and segmental instability (14), which can significantly impact postoperative health-related quality of life (HRQoL) (15). For patients with severe neurological symptoms caused by DIPs, revision surgery may be needed. The development of DIPs is multifactorial, and various predictive factors of DIPs have been reported, including age, preoperative sagittal malalignment, high body mass index (BMI), and LIV selection (11,16).

Recently, the role of paraspinal muscle degeneration and low vertebral bone mineral density (vBMD), evaluated by Hounsfield unit (HU) values determined by computed tomography (CT) scans, has been increasingly investigated in patients with DLS (3,17-21). A lower muscularity of the paraspinal muscle is associated with persistent low back pain, the loss of lumbar lordosis, disability, and proximal junctional mechanical complications (22,23). Lower HU

values of the vertebral body and pedicle trajectory could induce a high risk of stress fracture and implant failure in patients who undergo lumbar fusion (24). However, the effect of muscularity and the vBMD of the lumbosacral region on the development of DIPs in patients with DLS is still unknown.

This study aimed to confirm the relationship among paraspinal muscle degeneration, a low vBMD of the lumbosacral region, and DIPs in patients who undergo long-instrumented spinal fusion for DLS. We present this article in accordance with the STROBE reporting checklist (available at <https://qims.amegroups.com/article/view/10.21037/qims-22-1394/rc>).

Methods

Study design and participants

This was a single-center retrospective cohort study. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by the Research Ethics Committee of Beijing Chao-Yang Hospital (No. 2021-09-07-10), and the need for individual consent for this retrospective analysis was waived. From January 2013 to December 2019, patients who underwent posterior instrumentation and fusion for DLS with a minimum 24-month follow-up in Beijing Chao-Yang Hospital were retrospectively recruited. The follow-up procedures included physical examinations and standing full-length spine radiographs at 3, 6, 12, and 24 months postoperatively. The inclusion criteria were as follows: (I) an age >50 years old at the time of surgery; (II) a Cobb angle of scoliosis $>10^\circ$; (III) instrumented levels ≥ 4 ; (IV) LIV at or above S1; (V) upper instrumented vertebra (UIV) at or above L2; and (VI) a duration of follow-up ≥ 24 months. Patients with other etiologies of spinal deformity, tumors, infections, a history of spinal surgery,

and nonambulatory status were excluded from this study. In our hospital, spinopelvic fixation is not routinely performed in DLS patients. Considering that the placement of pelvic screws would significantly influence the biomechanical characteristics of the spine and that extensive dissection would directly reduce the function of the paraspinal muscle, to eliminate the impact of these factors on the outcomes, patients undergoing spinopelvic fixation were also excluded. Patients with a concomitant diagnosis of osteoporosis began antiosteoporosis therapy after discharge from the hospital.

Diagnostic criteria of distal instrumentation-related problems

The included patients were divided into two groups based on the occurrence of DIPs during the follow-up period. The diagnostic criteria for DIPs were as follows: (I) distal junctional kyphosis, which indicated that the distal junctional angle (formed by the superior endplate of the LIV and the inferior endplate of the adjacent distal vertebra) was at least 10° at the last follow-up and at least 10° greater than the preoperative measurement; (II) screw loosening, pull-out, or breakage at the LIV; (III) fracture of the LIV; (IV) segmental instability at the distal level of the LIV; and (V) the requirement for revision surgery due to severe clinical symptoms caused by instrument failure of the LIV (16,25). Patients who met one of the above diagnostic criteria were classified into the DIP group; otherwise, they were included in the non-DIP group.

Data collection

Demographic information was collected, including sex, age, BMI, and comorbidities. Surgical data, including the location of the UIV and LIV, levels of instrumentation, levels of interbody fusion, osteotomy status, rod material, and rod diameters, were recorded. For patients with LIV at S1, the performance of L5-S1 interbody fusion was also recorded. Full-length spine standing radiographs were obtained preoperatively and ≤ 2 weeks postoperatively. Radiographic parameters, including Cobb angle values, lumbar lordosis (LL, L1-S1), lumbosacral lordosis (LSL, L4-S1), sagittal vertical axis (SVA), T1 pelvic angle (TPA), pelvic tilt (PT), sacral slope (SS), pelvic incidence (PI), and incidence of PI-LL mismatch, were measured by two senior spinal surgeons with the Picture Archiving and Communication System (PACS). PI-LL mismatch indicated that the absolute value of PI minus LL was greater than

10° . Two spine surgeons independently collected these data from April 2022 to September 2022.

Paraspinal muscle evaluation by magnetic resonance imaging

Preoperative magnetic resonance imaging (MRI) of the lumbar spine was performed using Signa HDxt 3.0 T (General Electric Company), with a slice thickness of 3 mm. The slicing plane was parallel to the inferior endplate of the vertebral body. Axial T2-weighted images at the inferior vertebral endplate level of L4 and L5 were exported as Digital Imaging and Communications in Medicine (DICOM) data and analyzed using ImageJ software 1.52k (National Institutes of Health, USA).

The mean values of the cross-sectional area (CSA) and signal intensity (SI) of the multifidus (MF), erector spinae (ES), paraspinal extensor muscle (PSE), and psoas major (PS) were measured (*Figure 1A,1B*). The PSE indicated the combination of the MF and ES. The mean values of both the CSA and SI at L4 and L5 were calculated to reflect the muscularity and fatty infiltration of the paraspinal muscle at the lumbosacral region. To decrease the bias caused by patient stature, the relative cross-sectional area (rCSA), which indicated the ratio of the CSA of the muscle to that of the vertebral body, was applied. The gross cross-sectional area (GCSA) included the CSA of lean muscle, intramuscular fat, and soft tissue, while the functional cross-sectional area (FCSA) solely indicated the CSA of lean muscle (3). Then, the relative gross cross-sectional area (rGCSA) and relative functional cross-sectional area (rFCSA) were calculated, and their values were multiplied by 100 and used for statistical analysis (20). The ratio of the rFCSA to the rGCSA (F/G ratio) was also calculated. The muscle-fat index (MFI) indicated the ratio of the SI of muscle to that of subcutaneous fat (26). The MFIs of gross and functional muscle were defined as the gross MFI (GMFI) and functional MFI (FMFI), respectively.

Vertebral bone mineral density evaluation by CT scan

Preoperative lumbar spine CT scans were performed in the supine position with the following parameters: 320 mAs, 120 kVp, and a 5-mm thickness. The slicing plane was parallel to the endplates of the vertebral body on the coronal and sagittal planes, through the midpoint of the posterior edge of the spinal canal and the anterior edge of the vertebral body on the axial plane. Four images were obtained for each

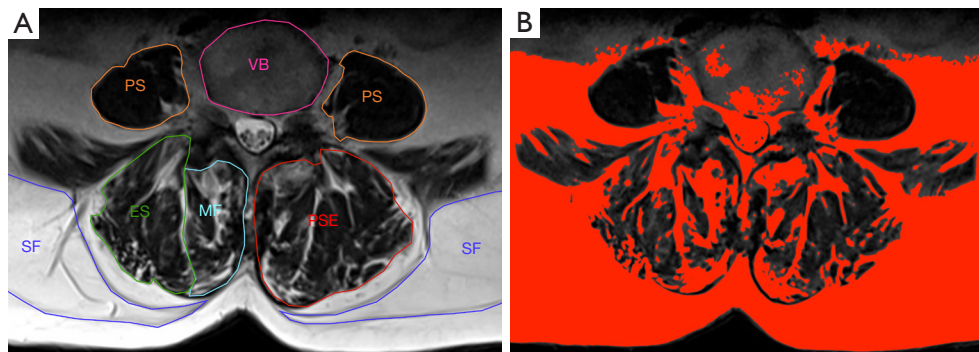


Figure 1 Measurement of paraspinal muscle parameters. (A) Preoperative lumbar spine magnetic resonance imaging; (B) ImageJ software. The red region indicates intramuscular fat, soft tissue, and subcutaneous fat through the threshold technique. MF, multifidus; ES, erector spinae; PSE, paraspinal extensor muscle; PS, psoas major; VB, vertebral body; SF, subcutaneous fat.

vertebra: for L4 and L5, the sagittal image at the midsagittal plane and three axial images (immediately inferior to the superior endplate, in the middle of the vertebral body, and immediately superior to the inferior endplate) were selected (21); for S1, the sagittal image at the midsagittal plane and three consecutive axial images immediately inferior to the superior endplate were selected (27). The region of interest (ROI) on each image was circular or oval and as large as possible, excluding the cortical margins, osteosclerotic areas, and osteophytes to prevent volume averaging (*Figure 2A-2F*). The HU value of each ROI was measured using OsiriX 13.0.1 software (Pixmeo Sarl, Switzerland). The mean HU value of the four ROIs was calculated as the HU value of the vertebral body. Then, the mean HU values of L4, L5, and S1 were calculated to reflect the vBMD of the lumbosacral region. The HU value of the LIV was also measured. The relationship between the HU values and dual-energy X-ray absorptiometry (DXA)-assessed BMD was evaluated by Pearson correlation analysis.

It should be noted that the imaging examinations were performed by the radiologic technologists who were duty that day. They all received the same training and obtained the same qualifications and certificates, and their operations were consistent. The radiologic technologists were blinded to the grouping of the patients and the grouping information of the MRI and CT images.

Statistical analysis

All statistical analyses were performed utilizing SPSS 25.0 (Chicago, IL, USA) and MedCalc 19.1.3 (Ostend, Belgium). A two-sided P value of less than 0.05 was considered

statistically significant. The Shapiro-Wilk test was performed to determine whether continuous variables had a normal distribution. Continuous variables with a normal distribution are presented as the mean \pm standard deviation; otherwise, the median and interquartile range are used. The counts and percentages are presented for categorical variables. For the comparison of continuous variables between the DIP and non-DIP groups, the independent-sample *t*-test or nonparametric Mann-Whitney U test was performed; for categorical variables, the Pearson chi-square test or Fisher's exact test was applied. Since all the patients included in this study were from our hospital, their data were available in our quality-controlled medical record system and the PACS. Therefore, there were no missing data.

The receiver operating characteristic curve was constructed using each MRI and CT scan parameter, and the best cutoff value was determined. The area under the curve (AUC) of each parameter was calculated. For each MRI parameter of each muscle, the variable with the maximum AUC was selected for the subsequent analysis. Then, the DeLong test was applied to select the MRI parameter and CT scan parameter with the most discriminatory ability for predicting DIPs.

Logistic regression analysis was performed to explore the potential predictive factors of DIPs. Demographic, radiographic, and surgical variables were first selected using univariate analysis. Significant factors and previously selected MRI/CT scan parameters were included in the multivariable logistic regression analysis to determine the independent predictive factors of DIPs. The odds ratio (OR) and 95% confidence interval (CI) were used.

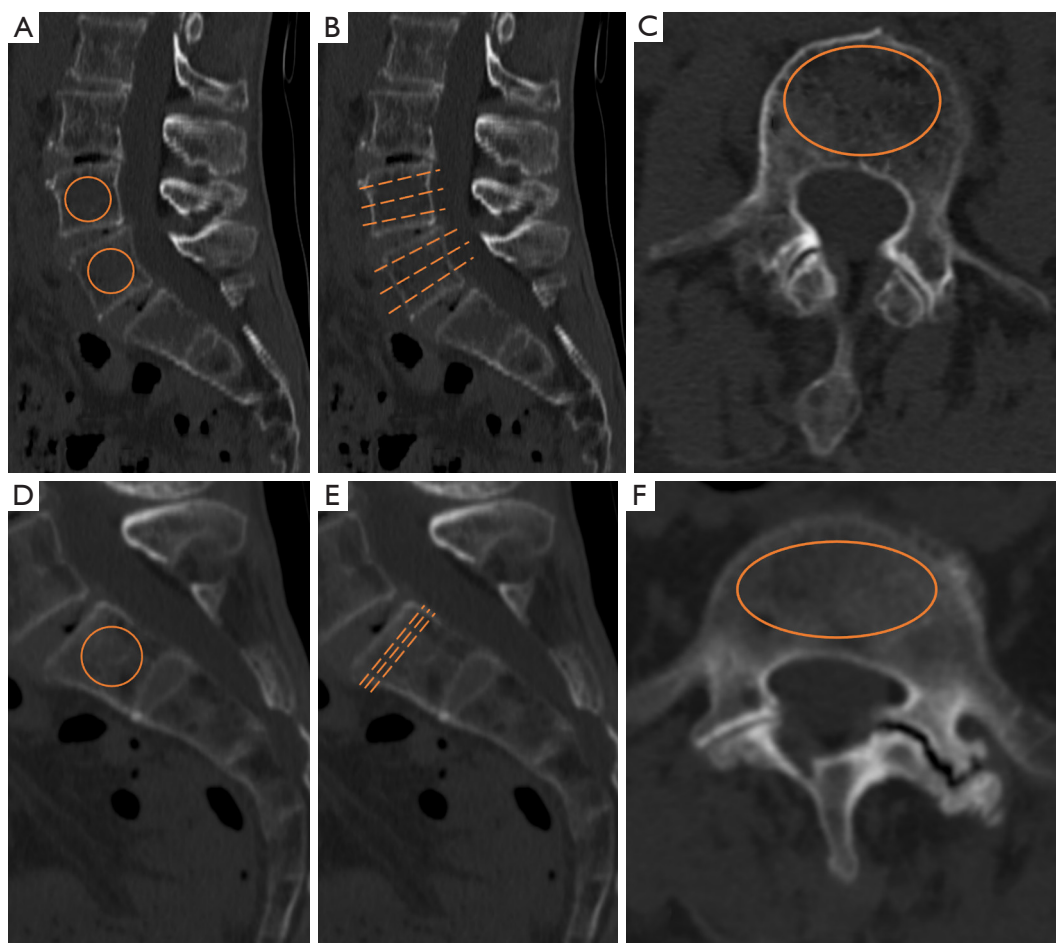


Figure 2 Measurement of HU values on a preoperative lumbar spine CT scan. The orange lines indicate the slicing planes. The orange circles indicate the ROI. (A) The ROI at the midsagittal plane image for L4 and L5; (B) The slicing planes at the midsagittal plane for L4 and L5; (C) The ROI at the axial plane image for L4 and L5; (D) The ROI at the midsagittal plane image for S1; (E) The slicing planes at the midsagittal plane for S1; (F) The ROI at the axial plane image for S1. ROI, region of interest; HU, Hounsfield unit.

Results

Demographic characteristics and surgical data

One hundred fifty-seven patients were screened, and 125 patients (70 females, 55 males) who met the inclusion and exclusion criteria were recruited for this study (Figure 3). The mean age at surgery was 66.57 ± 5.92 years, and the mean duration of follow-up was 27.23 ± 3.32 months. The incidence of DIPs was 16.0% (20/125). The demographic information and surgical data are summarized in Table 1. There were no significant differences in age ($P=0.71$), BMI ($P=0.71$), hypertension ($P=0.21$), chronic obstructive pulmonary disease ($P=0.51$), coronary artery disease ($P=0.63$), diabetes mellitus ($P=0.22$), the UIV ($P=0.73$) and LIV locations ($P=0.54$), the levels of instrumentation

($P=0.23$), the levels of interbody fusion ($P=0.83$), the performance of L5-S1 interbody fusion (for patients with LIV at S1) ($P=0.89$), osteotomy status ($P=0.37$), the rod material ($P=0.64$), or the rod diameter ($P=0.30$) between the DIP and the non-DIP groups. There were no patients who underwent three-column osteotomy (3-CO). As multiple-rod constructs were not routinely performed for DLS patients before June 2021 in our hospital, two-rod constructs were used for all recruited patients in the current study.

Degeneration of the paraspinal muscle of the lumbosacral region by MRI parameters

There was no significant difference in the rGCSA of each

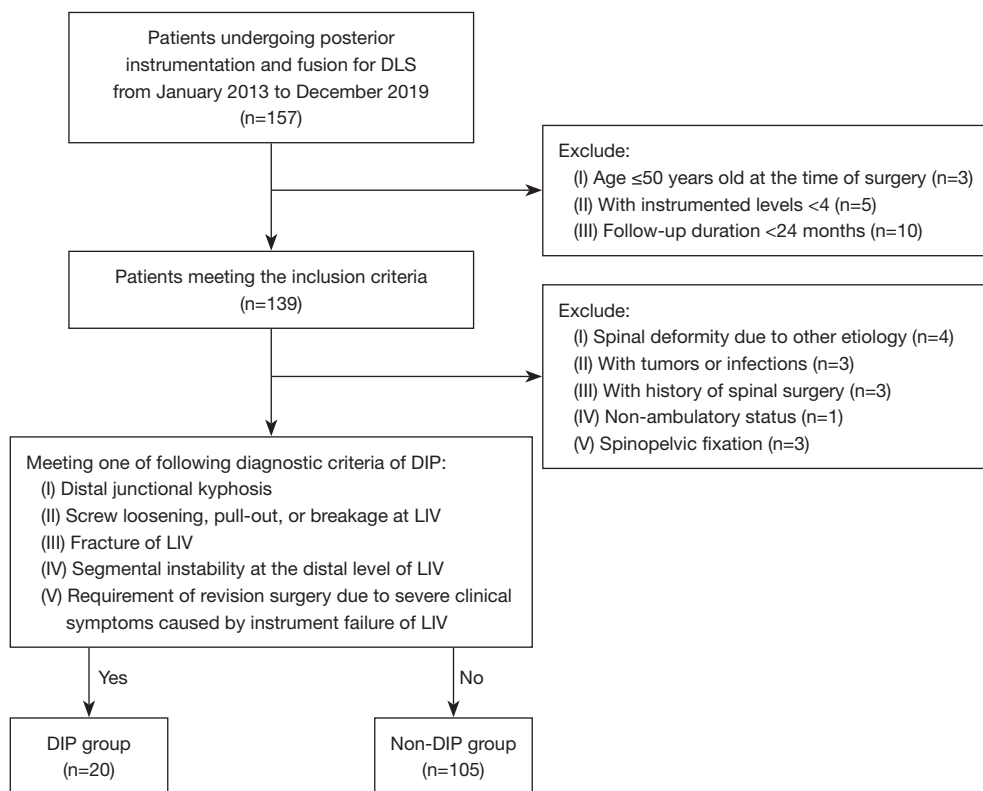


Figure 3 Flow chart of patient inclusion and exclusion. DLS, degenerative lumbar scoliosis; DIPs, distal instrumentation-related problems; LIV, lower instrumented vertebra.

muscle between the DIP and non-DIP groups. Additionally, none of the MRI parameters of the PS were significantly different between the groups. However, the rFCSA and F/G ratio of the MF, ES, and PSE were significantly smaller or lower, while the GMFI and FMFI were significantly greater in the DIP group (*Table 2* and *Figure 4A-4E*). Among the MRI parameters, the maximum AUC was observed in the rGCSA for the ES (0.528), rFCSA for the PSE (0.917), F/G ratio for the PSE (0.833), GMFI for the MF (0.784), and FMFI for the MF (0.805) (*Figure 5A-5E*). The DeLong test revealed that the AUC of the rFCSA for the PSE was significantly greater than that of the other four parameters (*Figure 6*). The cutoff value was 154.26, with a sensitivity and specificity of 88.6% and 85.0%, respectively.

Vertebral bone mineral density of the lumbosacral region by CT scan parameters

The HU value of the lumbosacral region (103.80 ± 22.64 vs. 132.19 ± 19.17 ; 95% CI: -37.064 to -19.717 ; $P < 0.001$) and LIV (111.70 ± 23.23 vs. 128.69 ± 20.70 ; 95% CI: -32.154

to -8.583 ; $P = 0.005$) were significantly lower in the DIP group than in the non-DIP group (*Table 3*). The Pearson correlation analysis revealed a significantly positive relationship between DXA-assessed BMD and the HU value of the lumbosacral region ($r = 0.832$, $P < 0.001$) or the HU value of the LIV ($r = 0.745$, $P = 0.005$). According to the DXA outcome, the incidence of osteoporosis was significantly higher in the DIP group than in the non-DIP group (85.0% vs. 61.0%, $P = 0.04$). The DeLong test revealed that the AUC of the HU value of the lumbosacral region was significantly greater than that of the LIV (0.858 vs. 0.756; 95% CI: 0.016 to 0.204; $P = 0.04$), with cutoff values of 112.03 and 101.39, respectively.

Radiographic parameters

There were no significant differences between the groups in the preoperative Cobb angle, LL, LSL, SVA, TPA, or SS. However, the preoperative PT was significantly lower in the DIP group than in the non-DIP group (19.86 ± 9.33 vs. 24.56 ± 9.54 ; 95% CI: 0.056 to 9.358; $P = 0.05$). For

Table 1 Comparison of demographic information and surgical data between the DIP and non-DIP groups

Variable	DIP group (n=20)	Non-DIP group (n=105)	95% CI	P value
Demographic Information				
Age (years)	66.20±6.71	66.74±5.79	-2.327 to 3.413	0.71
Sex				0.38
Male	7 (35.0)	48 (45.7)		
Female	13 (65.0)	57 (54.3)		
BMI (kg/m ²)	25.82±2.88	25.57±2.17	-1.655 to 1.146	0.71
Comorbidities				
Hypertension	9 (45.0)	63 (60.0)		0.21
COPD	4 (20.0)	15 (14.3)		0.51
CAD	7 (35.0)	31 (29.5)		0.63
Diabetes mellitus	2 (10.0)	23 (21.9)		0.22
Surgical Data				
Location of the UIV				0.73
At L2	11 (55.0)	67 (63.8)		
Between T11 and L1	2 (10.0)	7 (6.7)		
Above/at T10	7 (35.0)	31 (29.5)		
Location of the LIV				0.54
Above/at L5	14 (70.0)	66 (62.9)		
At S1	6 (30.0)	39 (37.1)		
Level of instrumentation	5.75±1.98	5.43±1.95	-1.471 to 0.628	0.23
Interbody fusion				0.62
No	6 (30.0)	26 (24.8)		
Yes	14 (70.0)	79 (75.2)		
Level of interbody fusion	1.95±1.19	2.06±1.04	-0.459 to 0.569	0.83
Osteotomy status				0.37
No	14 (70.0)	83 (79.0)		
Yes	6 (30.0)	22 (21.0)		
Rod material				0.64
Titanium alloy	18 (90.0)	98 (93.3)		
Cobalt chrome	2 (10.0)	7 (6.7)		
Rod diameter				0.30
5.5 mm	12 (60.0)	75 (71.4)		
6.0 mm	8 (40.0)	30 (28.6)		

Data are presented as mean ± standard deviation or n (%). BMI, body mass index; UIV, upper instrumented vertebra; LIV, lower instrumented vertebra; DIPs, distal instrumentation-related problems; COPD, chronic obstructive pulmonary disease; CAD, coronary artery disease; CI, confidence interval.

Table 2 Comparison of paraspinal muscle degeneration of the lumbosacral region between groups and the AUC of each parameter

Variable	DIP group (n=20)	Non-DIP group (n=105)	¹ 95% CI	¹ P value	AUC	² 95% CI	² P value
rGCSA							
MF	125.17±26.58	124.45±25.77	-12.549 to 11.979	0.96	0.507	0.367 to 0.646	>0.99
ES	152.30±27.53	155.21±29.63	-17.895 to 12.081	0.70	0.528	0.395 to 0.661	0.69
PSE	272.32±49.09	283.67±48.78	-26.771 to 20.387	0.79	0.511	0.378 to 0.644	0.87
PS	146.52±30.36	151.11±33.27	-21.573 to 12.395	0.59	0.510	0.382 to 0.637	0.89
rFCSA							
MF	65.74±21.51	92.37±21.68	-37.091 to -16.177	<0.001	0.823	0.710 to 0.936	<0.001
ES	82.67±21.44	111.48±24.21	-39.734 to -17.869	<0.001	0.827	0.721 to 0.934	<0.001
PSE	144.31±36.12	208.48±41.57	-82.807 to -57.453	<0.001	0.917	0.865 to 0.968	<0.001
PS	122.86±26.78	129.30±29.84	-21.762 to 8.885	0.41	0.519	0.394 to 0.643	0.79
F/G ratio							
MF	0.55±0.13	0.72±0.10	-0.188 to -0.041	0.004	0.773	0.653 to 0.893	<0.001
ES	0.56±0.14	0.70±0.11	-0.219 to -0.112	0.002	0.751	0.614 to 0.888	<0.001
PSE	0.56±0.15	0.75±0.12	-0.232 to -0.134	<0.001	0.833	0.710 to 0.955	<0.001
PS	0.83±0.05	0.87±0.04	-0.034 to 0.003	0.11	0.635	0.507 to 0.762	0.06
GMFI							
MF	0.50±0.16	0.35±0.13	0.091 to 0.220	<0.001	0.784	0.671 to 0.898	<0.001
ES	0.48±0.18	0.31±0.14	0.096 to 0.235	0.002	0.760	0.633 to 0.886	<0.001
PSE	0.51±0.21	0.34±0.11	0.107 to 0.241	0.002	0.732	0.583 to 0.881	0.001
PS	0.16±0.04	0.15±0.04	-0.009 to 0.029	0.29	0.588	0.447 to 0.730	0.21
FMFI							
MF	0.33±0.11	0.22±0.08	0.067 to 0.150	<0.001	0.805	0.696 to 0.914	<0.001
ES	0.35±0.13	0.23±0.10	0.076 to 0.177	<0.001	0.797	0.680 to 0.915	<0.001
PSE	0.34±0.12	0.21±0.08	0.086 to 0.168	<0.001	0.788	0.652 to 0.925	<0.001
PS	0.12±0.02	0.11±0.03	-0.002 to 0.024	0.10	0.625	0.493 to 0.757	0.08

Data are presented as mean ± standard deviation. ¹, independent-sample *t*-test; ², receiver operating characteristic curve analysis. rGCSA, relative gross cross-sectional area; rFCSA, relative functional cross-sectional area; F/G ratio, the ratio of the rFCSA to the rGCSA; GMFI, gross muscle-fat index; FMFI, functional muscle-fat index; MF, multifidus; ES, erector spinae; PSE, paraspinal extensor muscle; PS, psoas major; AUC, area under the curve; DIPs, distal instrumentation-related problems; CI, confidence interval.

postoperative radiographic parameters, there were no significant differences in the Cobb angle, LL, TPA, PT, SS, PI-LL, or the incidence of PI-LL mismatch between the groups. The postoperative LSL (33.63±9.58 *vs.* 28.48±9.41; 95% CI: -9.705 to -0.591; *P*=0.03) and SVA [38.84±17.83 *vs.* 25.30 (12.00, 33.27), *P*<0.001] were significantly greater in the DIP group than in the non-DIP group (Table 4). No significant difference in PI was detected between the groups (47.06±8.50 *vs.* 48.47±11.40; 95% CI: -3.903 to 6.722;

P=0.60). According to the above results, the preoperative PT, postoperative LSL, and postoperative SVA were selected for further multivariable logistic regression analysis.

Independent predictive factors of DIPs by multivariable logistic regression analysis

Based on the results of the previous analysis, the preoperative PT, the postoperative LSL, the postoperative

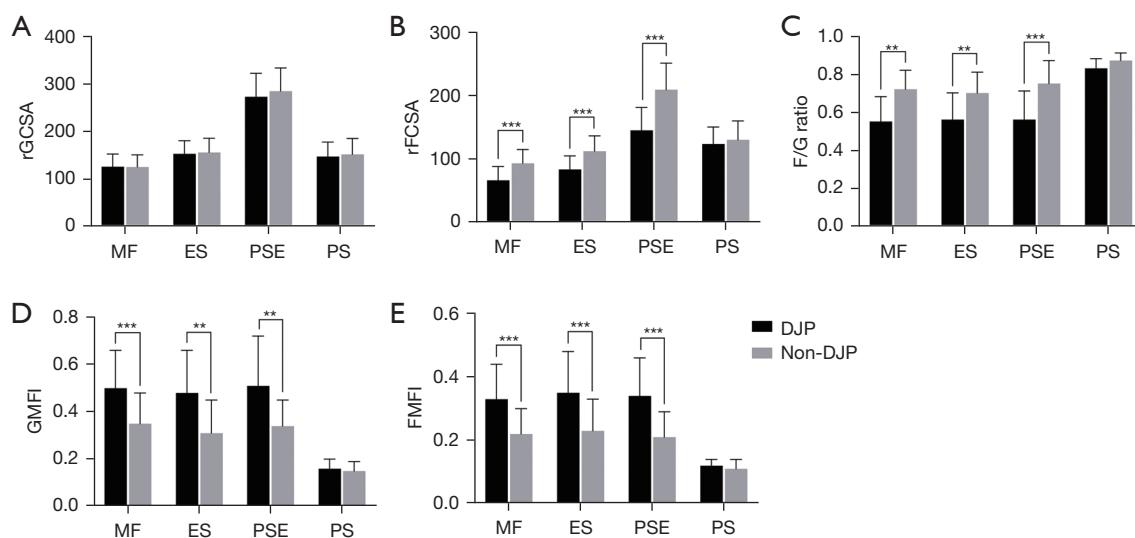


Figure 4 Differences in the rGCSA, rFCSA, F/G ratio, GMFI and FMFI of each muscle regarding the development of DIPs. (A) rGCSA; (B) rFCSA; (C) F/G ratio; (D) GMFI; (E) FMFI. **, P<0.01; ***, P<0.001; rGCSA, relative gross cross-sectional area; rFCSA, relative functional cross-sectional area; F/G ratio, the ratio of the rFCSA to the rGCSA; GMFI, gross muscle-fat index; FMFI, functional muscle-fat index; DIPs, distal instrumentation-related problems; MF, multifidus; ES, erector spinae; PSE, paraspinal extensor muscle; PS, psoas major.

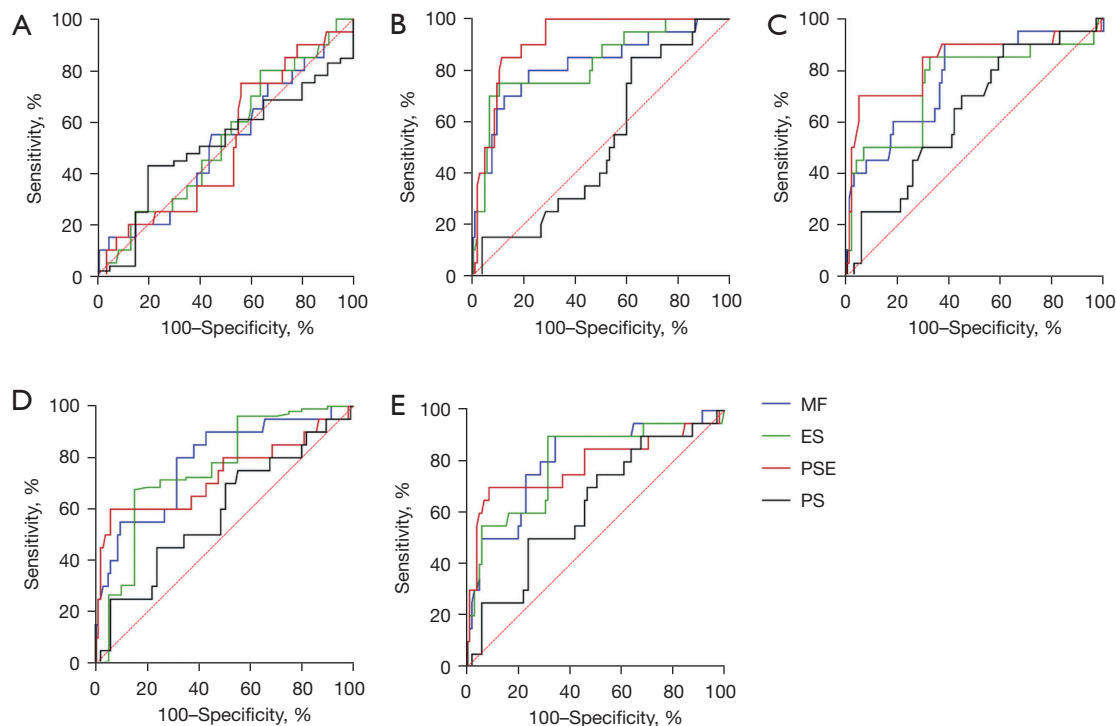


Figure 5 The ROC of the rGCSA, rFCSA, F/G ratio, GMFI and FMFI of each muscle. (A) rGCSA; (B) rFCSA; (C) F/G ratio; (D) GMFI; (E) FMFI. rGCSA, relative gross cross-sectional area; rFCSA, relative functional cross-sectional area; F/G ratio, the ratio of the rFCSA to the rGCSA; GMFI, gross muscle-fat index; FMFI, functional muscle-fat index; ROC, receiver operating characteristic; MF, multifidus; ES, erector spinae; PSE, paraspinal extensor muscle; PS, psoas major.

SVA, the rFCSA of the PSE, and the HU value of the lumbosacral region remained as covariates for inclusion in multivariable logistic regression analysis. In the multivariable model, the rFCSA of the PSE (OR 0.958; 95% CI: 0.936 to 0.981, $P < 0.001$), the HU value of the lumbosacral region (OR 0.941; 95% CI: 0.906 to 0.977, $P = 0.002$), and the postoperative SVA (OR 1.047; 95% CI: 1.003 to 1.093, $P = 0.04$) were detected as independent predictive factors of DIPs (Table 5). An illustrative case is demonstrated in Figure 7A-7S.

Discussion

Owing to the accumulating evidence indicating that the restoration of spinal alignment could better improve HRQoL for DLS patients, an increasing number of long-instrumented spinal fusions have been performed in recent decades (28-30). Proximal junctional kyphosis/failure, which

is a common mechanical complication that occurs at the proximal end of instrumentation, has been widely reported by previous studies (31-33). Low muscularity and vBMD at the UIV have been identified as predictive factors for this complication (21,34). Similarly, DIPs include mechanical complications at the distal end of instrumentation, which could cause sagittal decompensation, neurological symptoms and revision requirements, having a devastating impact on patients' HRQoL (13,35). However, studies focused on DIPs in DLS patients are still limited, and the role of paraspinal muscle degeneration and a low vBMD of the lumbosacral region is poorly understood. The current study observed DIPs in 20 patients, with an incidence of 16.0%. We found that a preoperative low lean muscle mass, a low vBMD, and postoperative sagittal malalignment could independently increase the risk of DIPs in DLS patients after long-instrumented spinal fusion.

The paraspinal muscle plays an important role in maintaining spinal stability and alignment (36). The paraspinal muscle of the lumbosacral region could support the spine, protecting against and distributing stress on the instrumentation. The stress on the instrumentation of the LIV is the strongest because it is located at the transition region between the instrumented and mobile spine, which has contrary biomechanical characteristics. In long-instrumented spinal fusion for DLS, the LIV is commonly at L5 or S1; however, fixation in the lumbosacral junction continues to be a challenge due to its unique biomechanical forces, anatomy, and sacral cancellous nature (37,38). The degeneration of the paraspinal muscle, which is presented as a decreasing CSA and high FI, would impact its strength and function, resulting in more biomechanical stress on the instrumentation of the LIV (39). Additionally, the extensive dissection of the PSE during surgery would increase the

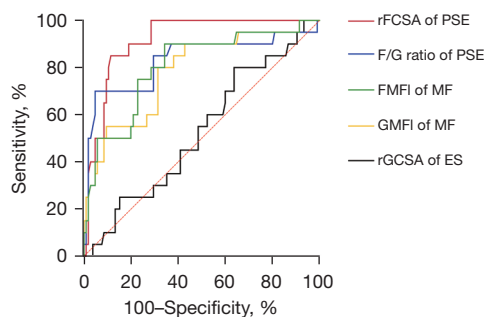


Figure 6 Results of the DeLong test. rGCSA, relative gross cross-sectional area; rFCSA, relative functional cross-sectional area; F/G ratio, the ratio of the rFCSA to the rGCSA; GMFI, gross muscle-fat index; FMFI, functional muscle-fat index; MF, multifidus; ES, erector spinae; PSE, paraspinal extensor muscle.

Table 3 Comparison of the vertebral bone mineral density of the lumbosacral region and LIV between the groups, the AUC of each parameter, and the outcome of DeLong test

Variable	DIP group (n=20)	Non-DIP group (n=105)	¹ 95% CI	¹ P value	AUC	² 95% CI	² P value	AUC difference	³ 95% CI	³ P value
HU value										
Lumbosacral region	103.80±22.64	132.19±19.17	-37.064 to -19.717	<0.001	0.858	0.749 to 0.966	<0.001	0.102	0.016 to 0.204	0.04
LIV	111.70±23.23	128.69±20.70	-32.154 to -8.583	0.005	0.756	0.641 to 0.872	<0.001			

Data are presented as mean ± standard deviation. ¹, independent-sample *t*-test; ², receiver operating characteristic curve analysis; ³, DeLong test. HU, Hounsfield unit; LIV, lower instrumented vertebra; AUC, area under the curve; DIP, distal instrumentation-related problem; CI, confidence interval.

Table 4 Comparison of radiographic parameters between the DIP and Non-DIP groups

Variable	DIP group (n=20)	Non-DIP group (n=105)	95% CI	P value
Cobb angle (°)				
Preoperative	19.07±4.53	21.26±5.01	-0.990 to 5.384	0.18
Postoperative	8.93±4.58	10.06±4.92	-2.694 to 5.555	0.49
LL (°)				
Preoperative	30.84±15.78	29.60±14.18	-8.205 to 5.738	0.73
Postoperative	45.01±14.52	41.42±12.70	-9.868 to 2.688	0.26
LSL (°)				
Preoperative	29.13±10.82	26.69±11.73	-8.040 to 3.157	0.39
Postoperative	33.63±9.58	28.48±9.41	-9.705 to -0.591	0.03
SVA (°)				
Preoperative	47.69±21.58	36.65±23.99	-22.449 to 0.377	0.06
Postoperative	38.84±17.83	25.30 (12.00, 33.27)		<0.001
TPA (°)				
Preoperative	20.28±10.84	23.27±10.22	-1.994 to 7.971	0.24
Postoperative	14.94±10.04	18.58±9.38	-0.943 to 8.217	0.12
PT (°)				
Preoperative	19.86±9.33	24.56±9.54	0.056 to 9.358	0.05
Postoperative	22.91±8.84	22.70±9.25	-4.608 to 4.185	0.92
SS (°)				
Preoperative	27.21±10.79	23.91±9.90	-8.149 to 1.554	0.18
Postoperative	24.15±10.44	25.77±10.84	-3.584 to 6.827	0.54
PI (°)				
Preoperative	47.06±8.50	48.47±11.40	-3.903 to 6.722	0.60
Postoperative PI-LL (°)	3.63±7.95	3.90 (-0.55, 12.65)		0.42
PI-LL mismatch	4/20 (25.0%)	32/105 (30.5%)		0.31

Data are presented as mean ± standard deviation, median (interquartile range), or n (%). DIP, distal instrumentation-related problem; CI, confidence interval; LL, lumbar lordosis; LSL, lumbosacral lordosis; SVA, sagittal vertical axis; TPA, T1 pelvic angle; PT, pelvic tilt; SS, sacral slope; PI, pelvic incidence.

flexional moment of the distal uninstrumented level adjacent to the LIV. Therefore, for DLS patients who undergo long-instrumented spinal fusion, the effect of paraspinal muscle degeneration on the incidence of DIPs is more significant.

In the current study, significantly smaller rFCSA and higher MFI values of the MF, ES, and PSE were observed in patients with the development of DIPs. This finding was similar to the results of Yuan *et al.* (3) and Leng *et al.* (12), who reported the association between extensor muscle degeneration and instrumentation-related complications

in patients with DLS. According to the literature, lean muscle mass could indirectly reflect muscle strength; therefore, the rFCSA was more sensitive than the rGCSA or MFI in predicting the prognosis of DLS patients (40,41). The results of the Delong test in the current study were consistent with those of previous studies. We found that the AUC of the rFCSA of the PSE was significantly greater than that of other MRI parameters, and this variable was identified as an independent predictive factor of DIPs by multivariable logistic regression analysis. There is no doubt

Table 5 Multivariate logistic regression of the independent predictors of DIPs

Variables	Odds ratio	95% CI		P value
		Lower limit	Upper limit	
Preoperative PT	0.942	0.867	1.022	0.15
Postoperative LSL	1.051	0.963	1.146	0.27
Postoperative SVA	1.047	1.003	1.093	0.04
rFCSA of PSE	0.958	0.936	0.981	<0.001
HU value of lumbosacral region	0.941	0.906	0.977	0.002

DIPs, distal instrumentation-related problems; CI, confidence interval; PT, pelvic tilt; LSL, lumbosacral lordosis; SVA, sagittal vertical axis; rFCSA, relative functional cross-sectional area; PSE, paraspinal extensor muscle; HU, Hounsfield unit.

that posterior spine surgery causes PSE atrophy due to iatrogenic denervation, ischemia, and thermal damage (42). Therefore, patients with low preoperative muscularity are more susceptible to DIPs. For these patients, some strategies could be considered to reduce the risk. Multiple-rod constructs have been recommended to reduce the concentrated stress on the instrumentation of L5 or S1 (9). Avoiding extensive musculoligamentous dissection is also of paramount importance. As there was no significant association between the muscularity of the psoas and DIPs, a more minimally invasive lateral surgical approach combined with posterior percutaneous instrumentation could be performed to preserve the PSE at the distal level (43).

Spinopelvic fixation is another important protective procedure for decreasing the risk of distal failure in patients undergoing long spinal fusion surgery (13,25). Many cohort studies of patients undergoing spinopelvic fixation using S2-alar-iliac (S2AI) screws or iliac screws have been reported (44-48). However, in our hospital, the indications for spinopelvic fixation only include neuromuscular scoliosis, 3-CO at the lumbar spine, and revisions for lumbosacral failure. Most of the DLS patients in our hospital were elderly and had comorbidities (e.g., hypertension, chronic obstructive pulmonary disease, coronary artery disease, and diabetes mellitus). The spinal deformity severity in these patients was relatively mild (usually a Cobb angle <20° and an SVA <50 mm), mainly presenting as lateral spondylolisthesis, axial rotation, and anterior translation. The primary complaints of these patients were usually low back or leg pain rather than spinal deformity. To solve their problems with a lower risk of postoperative

systemic complications and shorter time for surgery as well as anesthesia, we rarely perform radical 3-CO for these patients, and the UIV is commonly at the upper lumbar vertebra (L2) or lower thoracic region. Additionally, the performance of spinopelvic fixation may impact their HRQoL due to extensive dissection and sacroiliac joint violation (49,50). Therefore, spinopelvic fixation is not routinely performed for these patients. In the current study, none of the recruited patients had a history of spinal surgery, and only low-grade posterior column osteotomy (PCO) was performed. As a result, instrumentation was stopped at or above S1 for all recruited patients. The findings of this study may provide surgeons with an important caution that low preoperative paraspinal muscularity should also be an indication for spinopelvic fixation.

After the placement of instrumentation, additional compression loadings are exerted on the LIV. If the structural rigidity of the vertebra cannot adequately tolerate the alteration of compressive forces, bony failure occurs, leading to DIPs (17). Therefore, evaluation of vBMD is important before correction surgery for DLS. Although DXA scans are the gold standard for measuring BMD, they may not be reliable for evaluating cancellous bone in patients with spinal deformities, especially those with DLS (51). The HU value measured by CT scan continues to gain recognition because it could be a proxy for standard BMD and reflect bone strength (52,53). In the current study, the HU values of the lumbosacral region and LIV were significantly lower in patients with DIPs. The vertebra with a lower HU value usually had a lower Young's modulus, which indicated susceptibility to deformation when extra stress was exerted (54). The endplate of this vertebra may collapse, and the bone-screw interface could be loosened. The DeLong test showed that the HU values of the lumbosacral region had more predictive value than those of the LIV, indicating that surgeons should pay more attention to the deterioration of bone strength at the lower lumbar and lumbosacral junctions before surgery. The optimal cutoff value of this parameter was 112.03, which maximizes sensitivity (93.3%) and specificity (80.0%) in predicting DIPs. These data are paramount to preoperative optimization for DLS patients to pursue satisfactory outcomes. For patients with suspiciously low HU values of the lumbosacral region, DXA is recommended to evaluate their standard BMD, and surgery should be postponed until after 3-6 months of anti-osteoporotic medication therapy, such as teriparatide. During surgery, bone cement-augmented screws could be applied to reinforce the lumbosacral region and minimize

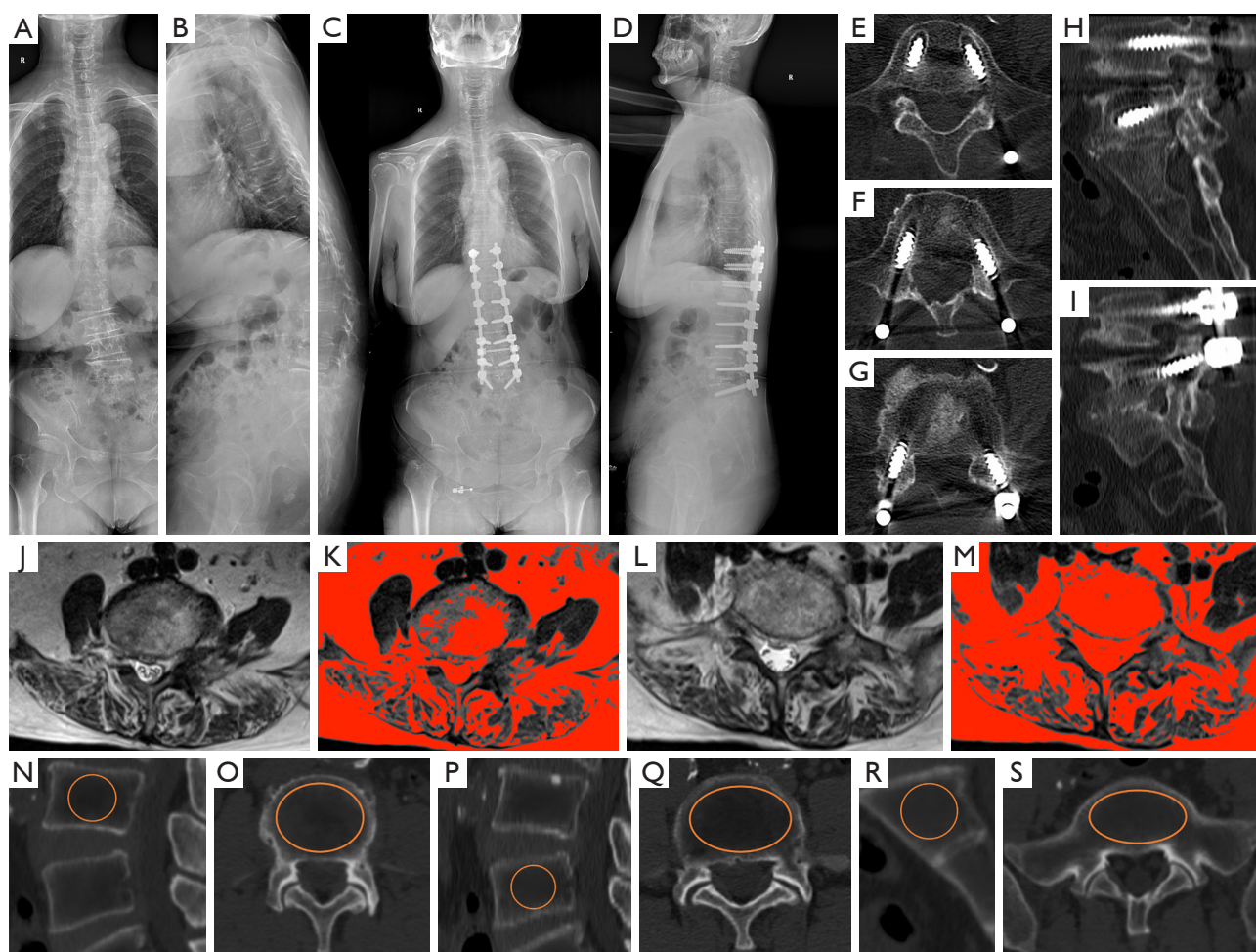


Figure 7 An illustrative case. A 70-year-old female with degenerative lumbar scoliosis and kyphosis was admitted to the hospital due to low back pain, leg pain, and intermittent claudication. She underwent long-instrumented spinal fusion from T10-L5, and she began antiosteoporosis therapy after discharge from the hospital. Two years after surgery, this patient complained of recurrent severe low back pain. The lumbar spine CT scan showed pedicle screw loosening at the LIV. The preoperative lumbar spine MRI showed low muscularity of the paraspinal muscle. The rFCSA for the PSE was 137.40. The preoperative lumbar spine CT scan showed low bone mass of the vertebra. The HU values of the lumbosacral region and LIV were 96.23 and 90.01, respectively. (A-D) Preoperative and two-year postoperative standing full-length spine radiographs; (E-I) two-year postoperative lumbar spine CT scan of the LIV; (J-M) measurement of the paraspinal muscle using preoperative lumbar spinal MRI at L4 and L5; (N-S) measurement of the HU values of the lumbosacral region and LIV using preoperative lumbar spine CT scans at L4, L5, and S1. The orange circles indicate the region of interest. CT, computed tomography; LIV, lower instrumented vertebra; MRI, magnetic resonance imaging; rFCSA, relative functional cross-sectional area; PSE, paraspinal extensor muscle; HU, Hounsfield unit.

the risk of DIPs (55). For patients with instrumentation ending at S1, L5-S1 anterior column support should also be considered, which could enhance the stability of the lumbosacral junction and decrease the mechanical complication rate in this region (56).

Biomechanical studies revealed that an anterior shift of

the gravity center relative to the LIV can cause additional loadings and stress on the endplate and instrumentation, increasing the risk of instrumentation failure (57,58). Additionally, it has been reported that paraspinal muscularity in the lower lumbar spine can only maintain local alignment rather than global balance (20). Consistent

with previous studies, the current study revealed that a larger postoperative SVA was an independent predictive factor of DIPs. For elderly patients with DLS, the flexibility of the spine is always rigid and has limited compensatory ability for postoperative sagittal malalignment. Radical posterior 3-CO, such as pedicle subtraction osteotomy (PSO), is usually needed to better restore the SVA during surgery. However, applying high-grade posterior osteotomy is also a risk factor for mechanical complications (59). Therefore, the anterior column realignment (ACR) procedure could be considered an alternative, performed through an anterior or lateral approach with no requirement for paraspinal muscle dissection (60). For DLS patients with severe preoperative sagittal imbalance, ACR combined with only PCO was adequate to restore sagittal alignment, which was as effective as PSO.

Limitations

Several limitations should be noted in this study. First, muscularity was only evaluated by CSA or MFI values measured by MRI, while ultrasound shear-wave elastography may be a better tool to directly assess paraspinal muscle function (61). Second, muscle strength and daily activities, which are important components in spinal mechanics, were not assessed in this study according to the guidance from the Asian Working Group for Sarcopenia (62). Third, posterior spine surgery commonly injures the paraspinal muscle; however, postoperative lumbar MRI is not routinely performed to evaluate the association between DIPs and the paraspinal muscle in postoperative variations. Last, the sample size was still limited, and the duration of follow-up was not long enough, which may hinder this study from detecting other risk factors.

Conclusions

The incidence of DIPs in patients who underwent long-instrumented spinal fusion for DLS was 16.0%. Lower muscularity of the PSE and vBMD of the lumbosacral region were independent predictive factors of the development of DIPs. Higher postoperative SVA values were also independently associated with this complication. Surgeons should pay attention to the preoperative evaluation of the paraspinal muscle and bone mass for patients with DLS, and targeted perioperative strategies should be considered.

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Footnote

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Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at <https://qims.amegroups.com/article/view/10.21037/qims-22-1394/coif>). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work by ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by the Research Ethics Committee of Beijing Chao-Yang Hospital (No. 2021-09-07-10), and the need for individual consent for this retrospective analysis was waived.

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