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Effects of anti-osteoporosis treatment in elderly patients with osteoporosis and lumbar discectomy and fusion

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Abstract.
BACKGROUND: The effect of anti-osteoporosis treatment in elderly patients with osteoporosis and lumbar discectomy and fusion (LIF) for lumbar degenerative diseases is not well known.
OBJECTIVE: This study aimed to evaluate the effect of perioperative anti-osteoporosis treatment in the patients with osteoporosis and LIF.
METHODS: From January to December 2022, patients were divided into three groups according to the inclusive criteria: the normal group (Group A), the osteopenia group (Group B) and the osteoporosis group (Group C). Quantitative computed tomography (QCT), height of the intervertebral space (HIS), segmental sagittal angle (SSA), visual analogue scale (VAS) score and Oswestry Disability Index (ODI) were compared between the groups at the follow-up time. The serum Ca^{2+}, osteocalcin (OC), propeptide of type I procollagen (PINP) C-terminal cross-linking telopeptide of type I collagen (β-CTX) and 25-OH vitamin D (25-OH V_D) levels were compared between the groups at the time of follow-up. Interbody fusion was graded on the X-ray and CT images at the follow-up time.
RESULTS: There were 165 patients in this study. There were significant differences in the mean age, mean score, HIS and SSA between the groups at the different follow-up times. There were significant differences in the concentrations of serum Ca^{2+}, OC, β-CTX, 25-OH V_D and PINP at the sixth month after surgery between the groups. There were significant differences in the concentrations of serum Ca^{2+}, β-CTX and 25-OH V_D between the pre-surgery and at six months after surgery in Group B and β-CTX and 25-OH V_D in Group C. There was a significant difference in the degree of fusion between Group B and C ($\chi^2 = 5.6243$, $P < 0.05$).
CONCLUSION: In elderly patients with LIF and osteoporosis, anti-osteoporosis therapy could reduce bone resorption and thus facilitate fusion. Anti-osteoporosis medication tends to enhance radiological, functional, and fusion short-term outcomes.

Keywords: Elderly patients, degenerative lumbar disease, lumbar discectomy and fusion, osteoporosis, anti-osteoporosis

1. Introduction

Posterior lumbar interbody fusion (PLIF) and transforaminal lumbar interbody fusion (TLIF) with segmental pedicle screws are common procedures for treating lumbar degenerative disorders that relieve pain and enhance function [1,2]. However, as more people experience spinal fusion, the number of fusion-related problems has risen [3,4]. There have been reports of adjacent vertebral body fractures caused by rigid pedicle screw fixation [5,6]. The reason for this is that arthrodesis may modify spinal kinematics, resulting in relative hypermobility of spinal segments next to the fused level [7,

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8]. This alteration in segmental spinal biomechanics could lead to a higher rate of vertebral compression fractures (VCFs). Especially among elderly patients, osteoporosis further increases fracture risk.

After spinal fusion surgery, rigid spinal instrumentation can cause device-related osteoporosis of the fused segment, as well as a decrease in the bone mineral density (BMD) of adjacent vertebrae, which can contribute to adjacent level VCFs [9,10,11]. Moreover, osteoporosis increases the risk of fractures in elderly individuals. The decrease in BMD could be due to immobilization or to changes in biomechanics caused by arthrodesis. However, the mechanism of BMD loss after spinal fusion surgery, is yet unknown. Zoledronic acid (ZA) has been shown to be an effective osteoporosis treatment in postmenopausal women in several studies, as it significantly reduces the risk of vertebral, hip, and other fractures [12]. Additionally perioperative ZA treatment may offer protection against a significant decrease in the BMD of cephalad vertebrae after spinal fusion surgery among postmenopausal women with osteoporosis [13].

Denosumab is a biological product that blocks the RANK ligand (RANKL)-mediated activation of osteoclasts. It has been demonstrated to be useful in treating osteoporosis [14], with effectiveness comparable to that of bisphosphonate [15]. However, to our knowledge, no previous study has evaluated the effect of denosumab on the change in the BMD of the adjacent vertebral body after instrumented intervertebral fusion, fusion time, or operation. As a result, the goal of this study was to determine whether denosumab can increase the BMD of the adjacent vertebral body after spinal surgery while also improving the fusion time and outcome of the operative segment.

2. Methods

2.1. Study design

This study was conducted from January to December 2022. The study protocol was approved by the Ethics Committee of Tianjin First Central Hospital (approval ID: 2022N122KY).

Written informed consent was obtained from all study participants and confidentiality of information was assured.

2.2. Study patients

The medical records of consecutive patients admitted for lumbar degenerative diseases (LDDs), including lumbar disc herniation (LDH) with instability, lumbar spinal stenosis (LSS), degenerative spondylolisthesis (DSI), isthmic spondylolisthesis (IS) and degenerative scoliosis (DS) to the hospital from January to December 2022 were collected.

Inclusion criteria: 1) X-ray, computed tomography (CT) and quantitative computed tomography (QCT, Siemens Edge, the measuring software was QCT Pro from Midways, www.QCT.com) and magnetic resonance imaging (MRI) results were obtained; 2) Serum Ca^{2+}, osteocalcin (OC), propeptide of type I procollagen (PINP) C-terminal cross-linking telopeptide of type I collagen (β-CTX) and 25-OH Vitamin D (25-OH V_D) results were obtained; 3) other blood test results were normal; 4) patients underwent PLIF or TLIF; 5) the follow-up results were intact; 6) no operative complications; 7) osteoporosis was defined by BMD < 80 mg/cm^3 and T score < -2.5, osteopenia was defined by 80 mg/cm^3 $<$ BMD $<$ 120 mg/cm^3 and $-2.5 <$ T score < -1, and normal defined by BMD $>$ 120 mg/cm^3 and T score > -1.0 were on QCT measures obtained before surgery; 8) patients underwent follow-up QCT evaluation at 6–12 months after surgery.

Exclusion criteria: 1) required intervertebral fusion of > 3 levels; 2) had a prior history of lumbar surgery and required lumbar surgery during the follow-up period; 3) had no lumbar disorders that may affect bone metabolism, such as cancer, infection or trauma; 4) those with osteoporosis secondary to various metabolic diseases, such as thyroid or parathyroid disease, and other endocrine diseases; 5) required use of a lumbar brace for > 2 months after surgery; 6) had any disability that limited the patient's walking ability for > 1 week during the follow-up period; and 7) smoked and(or) had sarcopenia. The patients were divided into three groups: Group A included normal patients; Group B included patients with osteopenia and Group C included patients with osteoporosis.

2.3. Surgical procedure

All patients underwent PLIF or TLIF. A posterior midline incision was made and the posterior elements of the spine were exposed. After pedicle screw insertion, a medial facetectomy was performed. Then the nerve root was mobilized medially to access the disc space. In the TLIF procedure, the superior and inferior articular processes of the facet joint are resected and the disc in the neuroforamen is exposed. After removal of the disc and scraping of the endplates, the anterior part of the disc space is packed with autologous bone chips. The

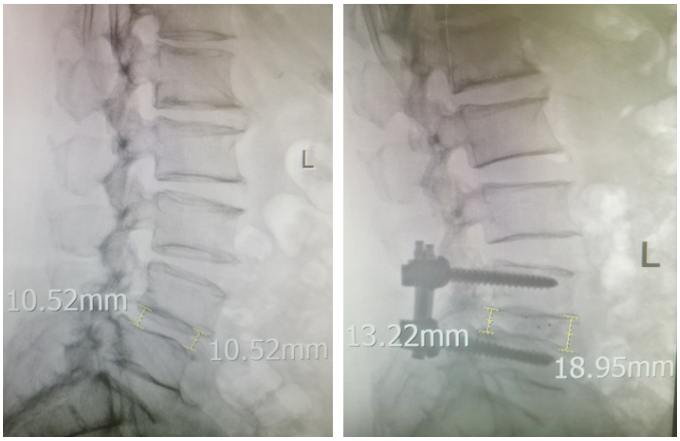

Fig. 1. HIS before and after surgery.

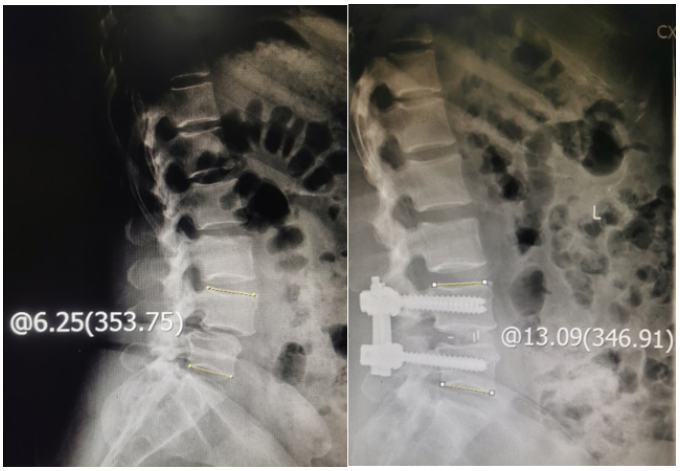

Fig. 2. SSA before and after surgery.

disc space is measured by trial insertion of variously sized spacers and a sizeable cage is inserted. All patients were asked to use a lumbar brace while walking for 4–6 weeks after surgery.

2.4. Imaging parameters

HIS was measured via X-ray before surgery, after surgery and at the last follow-up (Fig. 1). The HIS was the average of the anterior and posterior HISs at the same intervertebral space. SSA was measured via X-ray before surgery, after surgery and at the last follow-up (Fig. 2).

2.5. Bone metabolic markers

Serum Ca^{2+}, OC, PINP β-CTX and 25-OH V_D were tested before surgery and at the sixth month after surgery.

2.6. Anti-osteoporosis treatment

The anti-osteoporosis treatments used were 1200 mg/day oral calcium, 800IU/day activated vitamin D and 60 mg subcutaneous injection of denosumab every 6 months for the patients in Group C. Denosumab (American Amgen Inc.) was injected on the sixth day after surgery when the incision was not swollen and exuding. The treatment for patients in Group B was 1200 mg/day oral calcium and 800IU/day activated vitamin D.

2.7. Clinical treatment assessment

The VAS and ODI scores were assessed before the operation, after the operation and at the last follow-up. Interbody fusion was graded on the X-ray by the

Table 1
Demographic characteristic of 165 patients with LIF (n-165)

Characteristics	Group A ($n = 52$)	Group B ($n = 62$)	Group C ($n = 51$)	P
Mean age(yr)	50.7 ± 7.3	66.0 ± 4.5	69.1 ± 6.4	< 0.01
Sex, M/F	13/39	12/50	2/49	
Follow-up duration (month)	10.7 ± 0.2	10.8 ± 0.3	10.8 ± 0.2	> 0.05
Fusion level				
1 level	30	25	19	
2 levels	20	25	20	
3 levels	2	12	12	
Total	76	111	95	
Mean level	1.5 ± 0.6	1.8 ± 0.7	1.9 ± 0.8	< 0.05

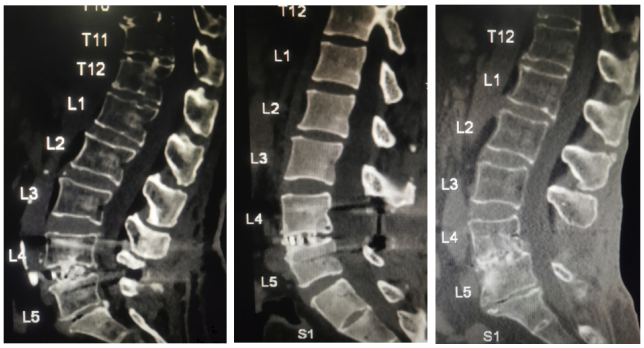

Fig. 3. According to Mayer's research, Grade 3, Grade 2 and Grade 1 were idectified at the sixth month after surgery (from left to right).

method of Brantigan and Steffee [16] as modified to describe the Fraser definition of locked pseudarthrosis (BSF scale) [17] at the third month and last follow-up after surgery. Interbody fusion by CT was performed as described by Mayer et al. [18] at the sixth month after surgery (Fig. 3). According to the CT results of Mayer et al. the definite fusion is Grade 1 or Grade 2.

2.8. Statistical analysis

Variables were compared among the three groups using one-way ANOVA and t tests for several parameters preoperatively and at the sixth month after surgery. P value < 0.05 was considered to indicate statistical significance. All statistical analyses were performed using SPSS software version 11.0 (SPSS Inc., Chicago, IL, USA).

3. Results

3.1. Demography

There were 165 patients in this study including 52 patients in Group A, 62 in Group B and 51 in Group C. Table 1 displays the mean age, follow-up duration, and sex and fusion level of the different groups. There were significant differences in the mean age ($P < 0.01$) and mean level ($P < 0.05$), but not at the follow-up.

3.2. Imaging parameters

Table 2 displays the preoperative, postoperative and last follow-up HIS and SSA. There were significant differences in the HIS among the three groups ($P < 0.05$) and among the different follow-up times ($P < 0.01$). Table 3 displays the QCT and T score results before the operation and at the sixth month after surgery. There were significant differences in the BMD and T score among the three groups at the same follow-up time ($P < 0.01$) and between the Group B ($P < 0.05$) and C ($P < 0.05$) at the different follow-up times.

3.3. Bone metabolic markers

Table 4 shows the serum Ca^{2+}, OC, PINP, β-CTX and 25-OH V_D levels before surgery and at the sixth month after surgery in the three groups. There were significant differences in the concentrations of serum Ca^{2+}, OC, β-CTX, 25-OH V_D and PINP at six months after surgery among the three groups. Between preoperation and six months after surgery, there were significant differences in the concentrations of serum Ca^{2+}, β-CTX and 25-OH V_D in Group B, β-CTX and 25-OH V_D in Group C.

Table 2
Comparison of image parameters between different groups ($n = 165$)

	Group A ($n = 52$)	Group B ($n = 62$)	Group C ($n = 51$)	P
Preoperative HIS (mm)	7.9 ± 2.0	8.9 ± 2.5	9.0 ± 2.7	< 0.05
Postoperative HIS (mm)	10.9 ± 1.8	11.5 ± 1.9	11.9 ± 2.2	< 0.05
Last follow-up HIS (mm)	10.5 ± 1.8	10.9 ± 1.7	11.8 ± 2.3	< 0.01
P	< 0.01	< 0.01	< 0.01	
Preoperative SSA (°)	15.6 ± 7.7	16.3 ± 9.2	17.4 ± 7.3	> 0.05
Postoperative SSA (°)	18.8 ± 8.0	20.5 ± 8.5	21.3 ± 7.8	> 0.05
Last follow-up SSA (°)	18.7 ± 5.0	19.8 ± 9.3	21.3 ± 8.5	> 0.05
P	< 0.05	< 0.01	< 0.01	

HIS: Height of intervertebral space; SSA: segmental sagittal angle.

Table 3
Comparison of QCT results between different groups ($n = 165$)

	Group A ($n = 52$)	Group B ($n = 62$)	Group C ($n = 51$)	P
BMD (g/cm^3)				
Pre-operation	138.8 ± 9.9	108.2 ± 18.5	64.7 ± 18.8	< 0.01
At the sixth month	137.5 ± 9.2	99.1 ± 17.9	72.1 ± 11.2	< 0.01
P	> 0.05	< 0.05	< 0.05	
T score				
Pre-operation	1.3 ± 0.1	-2.1 ± 0.3	-2.9 ± 0.3	< 0.01
At the sixth month	1.3 ± 0.1	-2.2 ± 0.2	-2.7 ± 0.2	< 0.01
P	> 0.05	< 0.05	< 0.01	

BMD: Bone mineral density.

Table 4
Comparison of bone metabolic markers results between different groups ($n = 165$)

	Group A ($n = 52$)	Group B ($n = 62$)	Group C ($n = 51$)	P
Serum Ca^{2+} (mmol/L)				
Preoperative	2.3 ± 0.1	2.3 ± 0.1	2.2 ± 0.1	< 0.01
At the sixth month after surgery	2.3 ± 0.1	2.2 ± 0.1	2.2 ± 0.2	< 0.01
P	> 0.05	< 0.01	> 0.05	
OC (ng/ml)				
Preoperative	21.8 ± 3.7	18.3 ± 4.1	16.3 ± 9.5	< 0.01
At the sixth month after surgery	22.9 ± 3.9	18.1 ± 3.3	17.8 ± 8.2	< 0.01
P	> 0.05	> 0.05	> 0.05	
PINP (ng/ml)				
Preoperative	42.3 ± 6.2	39.1 ± 10.8	40.7 ± 20.7	> 0.05
At the sixth month after surgery	43.5 ± 6.8	37.7 ± 9.9	41.8 ± 18.8	< 0.05
P	> 0.05	> 0.05	> 0.05	
$\beta-$CTX (ng/ml)				
Preoperative	0.3 ± 0.1	0.5 ± 0.2	0.6 ± 0.3	< 0.01
At the sixth month after surgery	0.3 ± 0.1	0.6 ± 0.1	0.5 ± 0.1	< 0.01
P	> 0.05	< 0.01	< 0.05	
25-OH V_D (ng/ml)				
Preoperative	37.8 ± 4.2	15.8 ± 6.6	16.7 ± 7.1	< 0.01
At the sixth month after surgery	36.9 ± 3.9	18.3 ± 5.8	27.3 ± 2.1	< 0.01
P	> 0.05	< 0.05	< 0.01	

OC: Osteocalcin; PINP: Propeptide of type I procollagen; β-CTX: C-terminal cross-linking telopeptide of type I collagen; 25-OH V_D: 25-OH Vitamin D.

3.4. Fusion assessment

Table 5 shows the X-ray results for the BSF at the third month and last follow-up after surgery and Mayer Grade results on the CT scan at the sixth month after surgery. According to the BSF scale, the prevalence of BSF-3 was 94.2% in Group A, 90.3% in Group B and 92.3% in Group C. Grade 1 and Grade 2 were defined according to Mayer's grade and were 96.2% in Group A, 64.5% in Group B and 78.4% in Group C. There was a significant difference in the percentage of patients with definite fusion between Groups B and C ($\chi^2 = 5.6243$, $P < 0.05$). The X-ray results were the same result at the last follow-up. The definite fusion in the

Table 5
Comparison of interbody fusion results between different groups ($n = 165$)

	Group A ($n = 52$)	Group B ($n = 62$)	Group C ($n = 51$)	P
X-ray assessment				
Third month after surgery				
BSF-1		6	3	
BSF-2	35	48	41	
BSF-3	17	8	7	
Last follow-up after surgery				
BSF-1		3	1	
BSF-2	3	3	2	
BSF-3	49	56	48	
CT assessment				
Sixth month after surgery				
Grade 1	47	21	19	
Grade 2	3	19	24	
Grade 3	2	16	6	
Grade 4		6	2	
VAS				
Pre-operation	6.8 ± 0.3	6.8 ± 0.2	6.9 ± 0.3	> 0.05
Post-operation	2.2 ± 0.2	3.3 ± 0.3	3.4 ± 0.2	< 0.01
Last follow-up	0.8 ± 0.1	1.3 ± 0.4	1.1 ± 0.1	< 0.01
P	< 0.01	< 0.01	< 0.01	
ODI				
Pre-operation	66.2 ± 8.5	69.2 ± 10.3	70.1 ± 11.0	> 0.05
Post-operation	32.9 ± 7.1	35.8 ± 7.1	36.2 ± 8.3	> 0.05
Last follow-up	8.3 ± 3.7	11.7 ± 6.1	10.9 ± 5.8	< 0.01
P	< 0.01	< 0.01	< 0.01	

VAS: visual analogue scale; ODI: Oswestry Disability Index.

Table 6
Comparison of interbody fusion results of the different diseases according to CT assessment ($n = 165$)

	LDH	LSS	DSI	IS	DS
Grade 1	66	11	2	7	1
Grade 2	24	14	1	6	1
Grade 3	2	3	7	5	6
Grade 4			2	2	4

LDH: lumbar disc herniation with instability; LSS: lumbar spinal stenosis; DSI: degenerative spondylolisthesis, IS: isthmic spondylolisthesis; DS: degenerative scoliosis.

different diseases was LDH, followed by LSS, and the fusion rates were lower for DSI and DS (Table 6).

3.5. Clinical treatment assessment

The VAS and ODI scores before surgery, after surgery and at the last follow-up are displayed in the Table 5. There were significant differences in VAS and ODI scores between the same group at the different follow-up times and among the three groups at the last follow-up time. At the sixth month after surgery, Fig. 4 shows a typical case in Group A, Fig. 5 displays a typical case in Group B and Fig. 6 shows a typical case in Group C. Figure 7 displays the screw pedicle cut out and fusion failure at the third month after surgery in Group B.

4. Discussion

Osteoporosis is a silent killer of human health and a major public health concern. In 2006 there were approximately 70 million osteoporotic patients and more than 200 million patients with osteopenia in China [19]. According to previous research, Asians have a lower bone mass than Caucasians and Afro-Caribbeans [20]. LDH, LSS, DSI, IS and DS are common LDDs that occur mostly in elderly patients and need LIF for treatment. In elderly individuals, the osteoporosis and LDD often coexist. Osteoporosis, a degenerative disease characterized by bone loss and structural deterioration, is the most common challenge of LIF. The main cause of osteoporosis in women is menopause, while aging is main cause in men, and is also the main cause of osteopenia.

To restore spinal stability and decompress nerves in elderly patients with LDD and osteoporosis, there have been some reports about the use of cement-augmented pedicle screw fixation to restore spinal stability [21, 22], but these complications may be fatal [23,24]. One study revealed that the strong pedicle screw fixation could be achieved by increasing the BMD around the screw and prevent fatal complication, this treatment involved anti-osteoporosis treatment during the spinal surgery and the follow-up period [25]. In our study, one patient with osteopenia, spondylolisthesis and lumbar

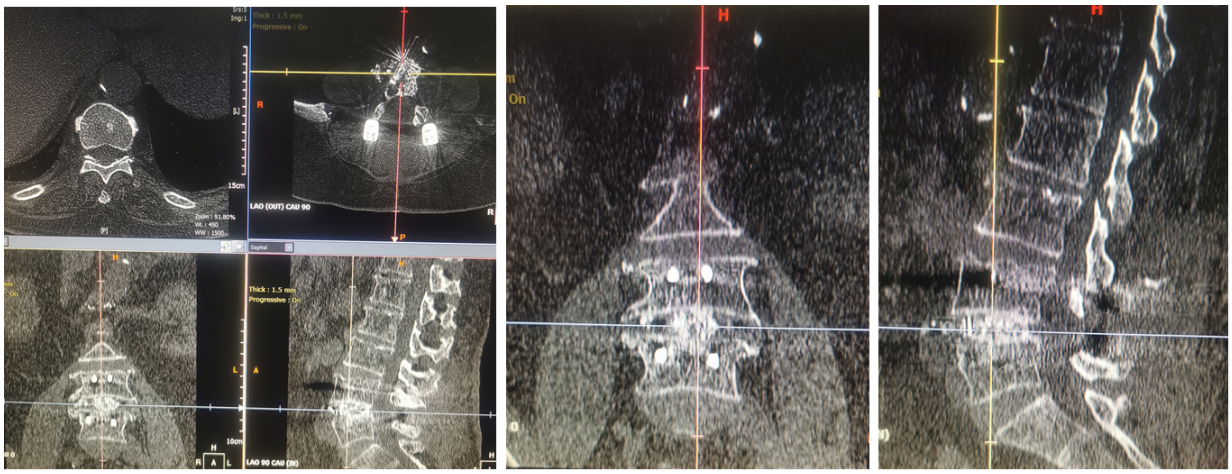

Fig. 4. Female patient, 52 yrs, lumbar spondylolisthesis and lumbar disc herniation before surgery, and noramal patient. PLIF was carried out. At the sixth month after surgery the interbody fusion was Grade 1 according to Mayer's research. The trabeculae were filled with interbody space.

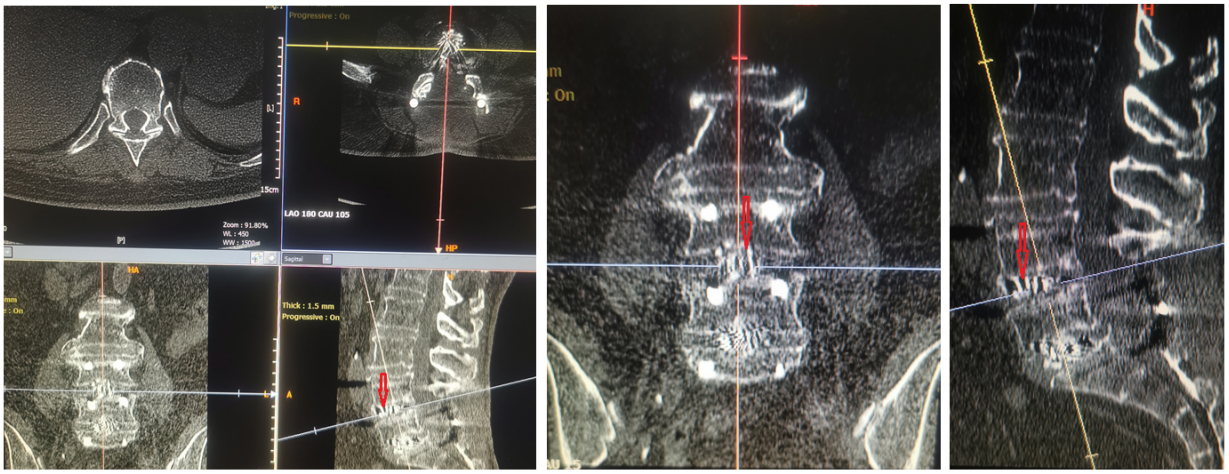

Fig. 5. Female patient, 63 years old, with lumbar spondylolisthesis and lumbar disc herniation before surgery; osteoponia; PLIF; oral calcium 1200mg/day; activated vitamin D 800IU/day before and after sugery. At the sixth month after surgery, the interbody fusion was Grade 3 according to Mayer's research. The red arrows show the unfused areas; and the trabeculae exhibited a palisade pattern.

disc herniation underwent PLF, but at the third month after surgery, the screw pedicle was cut out, the reduction effect on lumbar spondylolisthesis was lost, and fusion failed. This patient had no other uncomfortable symptoms except for mild low back pain and was still being followed up. Anti-osteoporosis agents have been shown to result in strong pedicle screw fixation with an increase in the BMD around the screw, but there have been few reports about the effectiveness of anti-osteoporosis agents for fusion of LIF. Our study focused on the short-term effect of LIF fusion during the spinal surgery in combination with denosumab for preventing osteoporosis.

In this study, the HIS and SSA were greater at the last follow-up than at the pre-operation, which demonstrated that the LIF could increase the HIS and improve lumbar lordosis. In the osteopenia group the HIS and SSA at the last follow-up were lower than those at the post-operation, but not in the osteoporosis group. According to some studies patients with low lumbar BMD may have a relatively high incidence of interbody fusion cage-related complications; for example, sinking of the interbody fusion cage may lead to gradual narrowing of the HIS, which affects anterior support of the spine and prevents successful fusion [26]. Several researchers have shown that BMD is related to the stable failure load of the endplate and cage [27,28]. Addition-

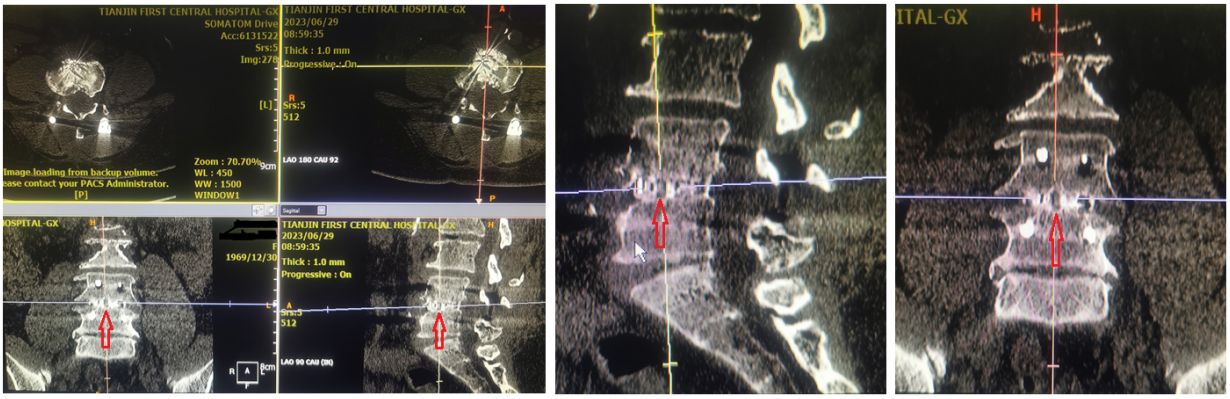

Fig. 6. Female patient, 66 years old, with lumbar spondylolisthesis and lumbar disc herniation before surgery; osteoporosis; PLIF; oral calcium 1200mg/day; and activated vitamin D 800IU/day before and after surgery. Denosumab was injected at the sixth day after surgery. At the sixth month after surgery, the interbody fusion was Grade 2 according to Mayer's research. The red arrows show the unfused area, which was just a small area, and the other areas were fused.

Fig. 7. Female patient, 67 yrs, with lumbar spondylolisthesis and lumbar disc herniation before surgery; osteoponia; PLIF; oral calcium 1200mg/day; and activated vitamin D 800IU/day before and after sugery. At the third month after surgery, the screw pedicle was cut out; and fusion failure occurred.

ally the correlation between the BMD and subsidence was very weak, and patients with a score $< -3{:}0$ had an increased risk of subsidence [29]. Cho et al. reported that the sedimentation rate was higher in the patients with the T score $\leqslant -2.5$ than in the patients with the T score $\geqslant -1.0$ [30]. In our study, the subsidence of the HIS in the osteopenia group was greater than that in the osteoporosis group because anti-osteoporosis treatment was used in the osteoporosis group.

In our study the serum Ca^{2+} and 25-OH V_D concentrations were greater at six months after surgery than before surgery in the osteopenia and osteoporosis groups, which proves the 1200 mg/day calcium and 00IU/day activated vitamin D 8 was necessary for patients with osteopenia or osteoporosis. There was little change in the bone formation markers including the OC and PINP in the osteopenia and osteoporosis groups, but there was a high variability in the bone resorption marker, which is the β-CTX index in the osteopenia and osteoporosis groups. The β-CTX increased at the sixth month after surgery in the osteopenia group, but decreased in the osteoporosis group due to the use of anti-osteoporosis

drugs. This finding also proved that denosumab could inhibit the activity of osteoclasts.

The VAS and ODI scores were improved at the last follow-up compared to the pre-operation. At six months after surgery the rate of definite fusion in the osteoporosis group were greater than that in the osteopenia group. The incidence of osteopenia and osteoporosis was lower than the normal group. According to these studies, the fusion rates were different after treatment with different anti-osteoporosis drugs; for example, one year after surgery, the fusion rate was 95% in the alendronate group [31], but the rate of bone fusion tended to be greater in the weekly teriparatide group than in the bisphosphonate group at the sixth month after surgery (46.8% vs. 32.7%) [32]. One study showed that alendronate does not influence the fusion process in osteoporotic patients, and in alendronate group, fusion was achieved in 66.7% of patients compared to 73.9% of patients in control group (no medication) [33]. The fusion rate was greater in the osteoporosis group than in the osteopenia group, but this difference was detected at only the sixth month according to the QCT. There were three typical cases from three groups: in the normal group the trabeculae fill the intervertebral space and fusion was complete at the sixth month after surgery; but in the osteopenia group the shape of trabeculae was like palisade, there was no trabeculae growth at some areas and lack of bone support, fusion effect was poor at the sixth month after surgery; in the osteoporosis group there was also lack of trabeculae at some areas which were smaller than in the osteopenia group and no shape of palisade area, and the fusion effect was better than the osteopenia group and worse than normal group.

Denosumab is an anti-resorptive agent with a novel mechanism of action [34,35] and a fully human monoclonal antibody that binds RANKL, preventing RANKL from activating RANK, its receptor on the osteoclast surface [35]. Reducing RANK-RANKL binding inhibits osteoclast formation, function, and survival, which results in a decrease in bone resorption and an increase in bone mass [35,36,37]. The mechanism of action of denosumab is different from that of bisphosphonates. The mechanism of action of bisphosphonates includes: 1) a strong affinity for bone embedding in the bone mineral, and crossing cell membranes when osteoclasts resorb the bone matrix; 2) clearance from the circulation via renal excretion or adsorption to bone minerals, but bone-associated drugs must first be released by osteoclast-mediated bone resorption; and 3) residue in the bone for a period of weeks to years. The mechanism of action of denosumab includes: 1) not embedding within bone tissue, binding to RANKL in the extracellular fluid and circulation, and inhibiting osteoclast formation, function and survival; 2) clearing from the bloodstream through the reticuloendothelial system; and 3) maintaining a half-life of approximately 26 days and not inducing the formation of neutralizing antibodies.

The follow-up time varied according to the detection type. Bone metabolic markers were detected at the sixth month after surgery due to injection of denosumab every 6 months. X-ray imaging was carried out at the third and sixth month to assess the degree of facet joint fusion. QCT was assessed at the sixth month to determine the degree of interbody fusion. The detections were carried out as little as possible to avoid unnecessary radiation and pain, except for necessary tests.

This study has several limitations. First, the follow-up time in this study was relatively short, but we think that follow-up time was long enough for patients to preliminarily recover, at which time all patients needed to complete interbody fusion after surgery. However, a prospective study with a larger sample size and longer term follow up should be conducted in the future. Second, this was a single-center-study, and the sample size was small. A multicenter study will be ideal. In addition, the study lacked detailed classification of patients with osteoporosis due to deficiencies in related detection methods.

5. Conclusions

In elderly patients with LIF and osteoporosis anti-osteoporosis therapy could reduce bone resorption and thus facilitate fusion. Anti-osteoporosis medication tends to enhance radiological, functional, and fusion short-term outcomes. The anti-osteoporosis treatment should also be advocated for patients with LIF and osteopenia.

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Author contributions

T.S.: initial idea, wrote and revised manuscript and acquired funding. F.Y.S.: data collection, editing and

statistical analysis. X.Q.: methodology. T.Z.: data collection. T.D.H.: data collection. W.L.J.: data collection. Q.Z.: data collection. All authors read and approved the final manuscript.

Conflict of interest

The authors declare that they have no conflict of interest.

Data availability

The datasets used and/or analyzed in this study are available from the corresponding author on reasonable request.

Ethics approval

Ethical approval was obtained from the Committee for Medical Ethics of Tianjin First Central Hospital (2022N122KY).

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Informed consent

Written informed consent was obtained from all study participants and confidentiality of information was assured.

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