Correlation of Sagittal Imbalance and Recollapse after Percutaneous Vertebroplasty for Thoracolumbar Osteoporotic Vertebral Compression Fracture: A Multivariate Study of Risk Factors

Whoan Jeang Kim, Sang Beom Ma, Hyun Min Shin, Dae Geon Song, Jae Won Lee, Shan Haw Chang, Kun Young Park, Won Sik Choy, Tae Ho Oh

Department of Orthopedic Surgery, Eulji University School of Medicine, Daejeon, Korea

Study Design: A retrospective case control study.

Purpose: This study aimed to assess the clinical significance of sagittal balance for predicting and managing the recollapse of cemented vertebra following percutaneous vertebroplasty (PVP) in patients with thoracolumbar osteoporotic vertebral fracture (OVF).

Overview of Literature: Recently, the recollapse of cemented vertebra following PVP for OVF has been reported. Although the risk factors for recollapse have been determined, the association between sagittal spinopelvic parameters and sagittal imbalance with recollapse has not been established.

Methods: Ambulatory patients who underwent single-level PVP for thoracolumbar OVF with a follow-up of at least 24 months were retrospectively reviewed. The patients were divided into two groups depending on the presence of symptomatic recollapse at the cemented vertebra: (1) recollapsed (RC) group and (2) noncollapsed (NC) group. The patient characteristics and radiographic measurements associated with sagittal imbalance were analyzed at each follow-up visit.

Results: Overall, 134 patients (RC group, n=28; NC group, n=106) were enrolled. The mean fracture-free interval was 3.2 months (range, 1.2–25.1 months). The multivariate binary logistic regression analysis identified low bone mineral density (p=0.047), degree of dynamic mobility within the vertebra (p=0.025), and sagittal imbalance as significant risk factors for recollapse (p=0.013; odds ratio, 5.405). The progression of sagittal imbalance and thoracolumbar kyphosis (T10–L2) was more significant in the RC and sagittal imbalance groups than in the NC group (both p<0.001).

Conclusions: Sagittal imbalance, lower bone mineral density, and dynamic mobility within the vertebra are associated with the recollapse of cemented vertebrae following PVP. Sagittal imbalance, rather than local kyphosis or thoracolumbar kyphosis, is particularly significant in that it results in more progressive collapse and sagittal deformity and is accompanied by substantial back pain and neurological deficits. Therefore, a stricter and more active management, including anti-osteoporosis medication, is required for the treatment of OVF with sagittal imbalance of the spine.

Keywords: Osteoporosis; Compression fractures; Sagittal imbalance; Thoracolumbar kyphosis; Vertebroplasty
Introduction

Osteoporotic vertebral compression fracture (OVF) is one of the most common complications of osteoporosis and has become more important as populations continue to age worldwide [1]. OVF results in substantial back pain and disability despite the use of analgesic medications. Moreover, it may lead to prolonged immobilization, which is associated with increased mortality in the elderly [2].

Percutaneous vertebroplasty (PVP) with polymethyl methacrylate is widely used to relieve back pain and reduce the need for prolonged immobilization in patients with symptomatic OVF [3-5]. Nevertheless, the recollapse of cemented vertebrae or cement loosening seldom occurs and is accompanied by the recurrence of back pain following cement augmentation [6-9]. Previous studies have identified the risk factors for the recollapse of the vertebra, namely, low bone mineral density (BMD), older age, intradiscal cement leakage, presence of multiple preexisting vertebral fractures, intravertebral vacuum cleft (IVC) sign, and fracture at the thoracolumbar junction [6,7,10-17].

The recollapse of cemented vertebrae should be carefully evaluated as it is often accompanied by an adjacent new vertebral fracture or instability, which potentially leads to further progression of kyphosis with sagittal imbalance (SI), especially at the thoracolumbar junction [17]. Several studies have indicated an association between an imbalanced spine and an increased kyphotic angle and further collapse of the affected vertebrae [10,18,19]. Moreover, when the kyphosis progresses with an accompanying neurological deficit or SI, revision surgery that involves anterior–posterior fusion should be performed [20,21] (Fig. 1).

For these reasons, the prediction of further progression of thoracolumbar kyphosis associated with SI facilitates the selection of appropriate treatment. Although some studies have demonstrated the recollapse of cemented vertebrae, the associations of sagittal parameters with recollapse following PVP in patients with thoracolumbar OVF have not been clearly elucidated.

In this study, we evaluated the risk factors for recollapse following PVP in patients with thoracolumbar OVF and its association with sagittal spinal parameters. We hypothesized that a spine with SI and thoracolumbar kyphosis would be at a higher risk of recollapse following PVP.

Materials and Methods

This study was approved by our Institutional Review Board. Data from all patients treated during the study period was available for review and analysis. All the study participants provided informed consent prior to data collection.

In this study, we retrospectively reviewed patients who underwent single-level PVP for OVF from January 2010 to October 2017. The study inclusion criteria are as follows: (1) ambulatory with senile or postmenopausal osteoporosis; (2) recent diagnosis of OVF; (3) a BMD T-score below –2.5; (4) follow-up of more than 24 months; and (5) fracture at the thoracolumbar junction. Moreover, the exclusion criteria are as follows: (1) a neurological deficit associated with vertebral fracture, (2) another pathologic condition as a cause of osteoporosis, (3) a history of spine surgery, or (4) a history of high-energy trauma, including falls from height and motor vehicle collisions.

The lumbar spine BMD values were measured in the lateral side of the lumbar body from L2 to L4 with the use of dual X-ray absorptiometry (Lunar Prodigy Advance; GE Healthcare, Madison, WI, USA), and the corresponding T-score was calculated.

In the present study, the “recollapse” of a cemented vertebra was classified as follows: (1) new-onset back pain (a Visual Analog Scale [VAS] score of >5 on a 0–10-point
scale); (2) resolution of back pain after the initial PVP for at least 4 weeks; (3) a reduction of more than 10% in cemented vertebral height based on the whole-spine lateral standing radiographs or computed tomography scans compared with that documented immediately following the initial PVP; and (4) a change in the signal intensity of the vertebral body based on magnetic resonance imaging, which was performed on those who were suggestive of recollapse (Fig. 2).

The patients were divided into two groups: (1) recollapsed group (RC group) and (2) noncollapsed group (NC group). Any new fractures of the vertebrae adjacent to the cemented vertebra were identified.

PVP was initially indicated for patients with painful OVF who were aged ≥80 years or had the IVC sign. Conservative treatment was indicated for patients younger than 80 years and did not have the IVC sign. PVP was performed if persistent pain or progression of a fracture was identified after 2 weeks of nonoperative treatment. The subcutaneous injection of teriparatide or oral bisphosphonate was prescribed at the time of identification of OVF for all patients. A thoracolumbosacral orthotic brace was utilized in all patients for at least 2 months after the procedure. PVP was performed using a transpedicular approach under the guidance of C-arm fluoroscopy. Poly-methyl methacrylate (Exolent Spine; Elmdown, Milano, Italy) was injected into the cleft site or fracture site of the collapsed vertebral body while checking for cement leakage through C-arm fluoroscopy, and the amount of cement injected for each patient was retrospectively reviewed. The patients were scheduled for follow-up in our clinics at 1, 3, 6, 12, and 24 months postoperatively.

1. Clinical evaluation

Pain was evaluated based on the VAS score before and after surgery and at each follow-up visit. A higher VAS score indicates more painful symptoms. Moreover, we reviewed the patients’ characteristics, including age, sex, body mass index, BMD, and previous OVF.

2. Radiographic evaluation

Simple anteroposterior radiographs of the thoracolumbar junction and whole-spine lateral standing radiographs with the inclusion of the femoral head were obtained in all patients. By applying Cobb’s method, we measured the spinopelvic parameters, including the sagittal vertical axis (SVA) and thoracolumbar kyphosis angle (TLKA; T10–L2), in whole-spine lateral standing radiographs. The pa-
tients with an SVA of over 5 cm on a preoperative whole-spine lateral view were classified as the preoperative SI group and the others as the sagittal balance (SB) group.

We measured the local Cobb angle between the upper and lower endplates of the affected vertebra on both standing and supine lateral radiographs and calculated the degree of dynamic mobility within the vertebra by subtracting the local Cobb angle on a lateral standing radiograph from that on a simple lateral radiograph [6,22].

The cemented vertebral restoration ratio after the procedure was calculated as the ratio of the average height of the fractured vertebra subtracted from the average height of the cemented vertebra immediately after surgery on a simple lateral radiograph to the average height of the fractured vertebra.

Radiographic measurements were performed using an image analysis software (Maroview, version 5.4; Marotech, Seoul, Korea) on the hospital’s picture archiving and communication system. Measurements were obtained twice for each investigator, and the mean values were calculated. Reliability was classified as small (0–0.24), low (0.25–0.49), medium (0.50–0.69), excellent (0.70–0.89), and best (0.90–1.0), depending on the coincidence correlation coefficient in the group. The intraobserver reliability was estimated to be 0.92.

3. Statistical analysis

The independent paired t-test and chi-square test were employed to evaluate the differences in patient characteristics and radiographic measurements between the RC and NC groups. A multivariate logistic regression analysis was then conducted to identify the risk factors associated with a recollapsed vertebra after cement augmentation. Repeated measures analysis of variance was conducted to compare the progression of thoracolumbar kyphosis and SI recorded at each follow-up visit between the two study groups. All statistical analyses were conducted using the

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>RC group (n=28)</th>
<th>NC group (n=106)</th>
<th>p-value</th>
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<td>Patients’ characteristics</td>
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<tr>
<td>Age (yr)</td>
<td>79.4±7.8</td>
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<td>22.7±3.9</td>
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<td>Bone mineral density (T-score)</td>
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<td>-2.8±0.7</td>
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<td>Previous osteoporotic vertebral fracture</td>
<td>14 (50)</td>
<td>19 (18)</td>
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</tr>
<tr>
<td>Average time to recollapse (mo)</td>
<td>3.2 (1.2–25.1)</td>
<td>-</td>
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<td>Radiographic measurements</td>
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<td>Degrees of mobility (°)</td>
<td>6.3±5.2</td>
<td>2.6±1.8</td>
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<td>Intravertebral vacuum cleft</td>
<td>17 (61)</td>
<td>30 (28)</td>
<td>0.002*</td>
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<td>Sagittal imbalance</td>
<td>20 (71)</td>
<td>10 (9)</td>
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<td>Local Cobb angle (°)</td>
<td>14.3±8.1</td>
<td>15.4±6.5</td>
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<td>Thoracolumbar kyphosis angle (°)</td>
<td>33.6±10.8</td>
<td>21.7±14.2</td>
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<td>Sagittal vertical axis (mm)</td>
<td>10.8±5.8</td>
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<tr>
<td>Adjacent vertebral fractures</td>
<td>13 (46)</td>
<td>11 (10)</td>
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<tr>
<td>Intervertebral disc cement leakage</td>
<td>7 (25)</td>
<td>20 (19)</td>
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<td>Restoration ratio</td>
<td>0.20±0.21</td>
<td>0.16±0.17</td>
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<td>Average amount of cement injected (mL)</td>
<td>4.2 (3.0–6.5)</td>
<td>4.1 (2.8–5.5)</td>
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Values are presented as mean±standard deviation, number of patients (%), or mean (range), unless otherwise stated.

RC group, recollapsed group; NC group, noncollapsed group.

*p<0.05 (using an independent-sample Student t-test and chi-square test).
IBM SPSS ver. 20.0 (IBM Corp., Armonk, NY, USA). A p-value of <0.05 was considered statistically significant.

**Results**

Overall, 122 of the 256 patients who underwent PVP for thoracolumbar OVF during the study period did not meet the study’s eligibility criteria, leaving 28 patients (21%) for inclusion in the RC group and 106 in the NC group.

Table 1 presents the patient characteristics and radiographic measurements. The BMD was significantly lower in the RC group than in the NC group (p=0.000). Previous OVF was more common in the RC group (p=0.001). The dynamic mobility (indicated by the difference in the Cobb angle between supine and standing radiographs) was greater and the IVC sign was more frequent in the RC group (p=0.000 and p=0.002, respectively). SI was predominantly found in the RC group with increased TLKA and SVA (p=0.000, p=0.002, and p=0.021, respectively). Fractures of adjacent vertebra identified during follow-up were more common in the RC group (p=0.000). No significant between-group difference in the local Cobb angle, likelihood of leakage of cement into the intervertebral disc, or restoration rate following PVP was observed. The average amount of cement injected during the procedure also showed no differences between the two groups.

Table 2 presents the results of the multivariate binary logistic regression analysis of possible risk factors for recollapse after cement augmentation. Lower BMD (p=0.047), degree of dynamic mobility (p=0.025), and SI (p=0.013; odds ratio, 5.405) were statistically significant risk factors associated with the recollapse of the vertebrae after the procedure.

Tables 3 and 4 present the progression of thoracolumbar kyphosis and SI in the two groups (except in patients who underwent surgical correction and fusion during follow-up) (Fig. 3). The progression of sagittal spinal deformity,
including thoracolumbar kyphosis and positive SI, was identified in both groups and was more prominent and aggressive in the RC and SI groups \((p=0.000\) for both).

**Discussion**

PVP has been frequently used for OVF in elderly patients with osteoporosis and achieves substantial relief of pain and improvement in the quality of life [23]. Recently, the recollapse of the cemented vertebra has been demonstrated as a complication following cement augmentation [16,24,25]. One study identified the radiographic loss of height in the cemented vertebra in 63% of 98 patients during an average of 27 months of follow-up [26]. Another study found a symptomatic re-fracture of the cemented vertebra in 16% of 356 patients, with 52% of the re-fractures occurring within a month and 32% within 3–5 months [25]. In another study, the mean prevalence of re-fracture involving adjacent vertebrae or vertebrae remote to the cemented vertebrae (which depends on the duration of follow-up and study design) was 21.28% (ranging from 6.81% to 37.95%) [27]. In the present study, the recollapse of the cemented vertebra was identified in 28 (21%) of 134 patients during a 2-year follow-up. The patients in the RC group were prescribed with anti-osteoporosis medication, mainly subcutaneous injection of teriparatide, and were educated about the importance of strict compliance. Repeat PVP was sometimes performed for substantial back pain or accompanying adjacent fractures. Two patients of the RC group developed severe back pain, worsened SI, multiple adjacent fractures, and progressive neurological deficits and finally underwent surgical correction (Fig. 4).

Previous studies have described several risk factors for recollapse following cement augmentation for OVF [10,19,24,28]. The patient-related risk factors for the re-fracture of cemented vertebrae or new fracture of an adjacent vertebra include excessive body weight, the use of steroids, advanced age, and poor compliance with anti-
osteooporosis medication [7,11,15,24]. The technical factors associated with recollapse include a larger or smaller volume of injected cement and performing percutaneous balloon kyphoplasty instead of PVP [7,10]. Statistically significant radiographic risk factors have also been identified, including higher local kyphosis, intravertebral instability, and presence of the IVC sign with pseudarthrosis [6,7,10,13]. Furthermore, strong evidence has indicated that low BMD, leakage of intervertebral disc cement, fractures at the thoracolumbar junction, higher restoration rate of the vertebral height, and preexisting vertebral fractures are general risk factors [17].

In the present study, we identified three risk factors (lower BMD, dynamic mobility, and SI) for recollapse following PVP for thoracolumbar OVF. We had anticipated that the IVC sign would also be a significant risk factor as intravertebral osteonecrosis and a defect in the cancellous component of the bone would be related to gradual deformation [13]. However, this sign was not a statistically significant risk factor for recollapse. This finding suggests that significant dynamic instability accompanied by the IVC sign would be clinically important in terms of further collapse of a cemented vertebra, which is also associated with more osteoporotic bone. SI, which has not been evaluated before, would be a clinically significant factor when managing thoracolumbar OVF associated with further recollapse. This finding indicates that progressive thoracolumbar kyphosis with insufficient spinopelvic compensation may lead to stooping and further SI, which aggravates weight loading and stresses on the anterior–superior portion of the cemented vertebra or adjacent vertebrae at the thoracolumbar junction [6,29].

In the present study, the progression of the average TLKA following vertebroplasty was identified over time and was more prominent in patients with SI or a recollapsed vertebra. We demonstrated that SI was a clinically significant factor predicting further collapse and spinal deformity for patients with a recollapsed vertebra. The statistical significance of the relationship between SI and recollapse indicates the need for sagittal spinal parameters in the management of OVF, especially at the thoracolumbar junction. Therefore, when managing such patients, a whole-spine lateral standing radiograph should be checked for the presence of SI with spinopelvic decompensation and dynamic mobility within the vertebra.

In the present study, some variables were not significantly correlated with recollapse. For example, TLKA was not a risk factor for recollapse despite the difference in the average value between the two study groups, which indicates that the TLKA would not correlate quantitatively with the risk of recollapse. We believed that higher thoracolumbar kyphosis or local kyphosis would lead to vertebral recollapse following PVP. However, the degree of thoracolumbar kyphosis or local kyphosis was not clinically important. Instead, the presence of SI was a clinically significant risk factor associated with further collapse and the progression of sagittal deformity of the spine. Adjacent vertebral fractures were more common in the RC group. However, their correlation with recollapse was not statistically significant. Considering the reports indicating that adjacent fractures are associated with intradiscal cement leakage or greater restoration of the vertebral height, which was not different between our two groups, it is possible that the biomechanics of cemented vertebrae that recollapse are different from those of adjacent vertebrae that develop fractures [29,30].

A strong correlation was observed between OVF at the thoracolumbar junction and further progression of thoracolumbar kyphosis, which could be aggravated by the collapse of a vertebral body and multiple adjacent fractures as well as the severity of osteoporosis and could result in neurological and functional deficits [1]. For these reasons, prediction and prevention of the progression of kyphotic deformity or gradual SI would be necessary to preserve physiologic functions and decrease the mortality associated with OVF and other compromising conditions. A comprehensive understanding of SI and careful assessment of sagittal spinopelvic parameters on whole-spine lateral standing radiographs would be essential for the prediction of the prognosis and decision-making with regard to further treatment for such patients.

Patients who have undergone PVP a few weeks earlier should be carefully evaluated for the recollapse of the cemented vertebra; any pain usually occurs at the same site. Therefore, it may be difficult to diagnose a new fracture. The recollapse is usually associated with lower BMD and is often accompanied by an adjacent new vertebral fracture, which potentially leads to further progression of kyphosis, especially at the thoracolumbar junction [27]. In the present study, the average TLKA following vertebroplasty progressed over time, more prominently in patients with SI or a recollapsed vertebra. Therefore, we assume that the recollapse of the cemented vertebra accompanied by lower BMD or adjacent vertebral fractures would result...
in the progression of SI and that the SI would aggravate the collapse of the affected vertebra or lead to new adjacent vertebral fractures. Patients with SI along with a greater progression of thoracolumbar kyphosis should be managed more carefully, given that an imbalanced spine may lead to decreased quality of life and occasionally require surgical correction. During the study period, two of the 28 patients in the RC group and none of the NC group underwent additional surgeries. The two patients demonstrated a significant progression of SI and substantial back pain. After the corrective surgery, both patients showed satisfactory clinical outcomes and correction of the SI.

From a clinical point of view, corrective surgery for SI should not always be performed on patients with osteoporosis vertebral fractures with spinal imbalance. Conservative treatment with strict management for osteoporosis and careful monitoring for the progression of SB using whole-spine lateral standing radiographs and dynamic X-rays should be performed. The corrective surgery and stabilization for osteoporotic fractures with SI should be considered when the progression of substantial pain or spinal imbalance or neurologic deficits is identified.

Unlike earlier studies, we evaluated the association between SI and thoracolumbar kyphosis and recollapse following PVP. Moreover, we demonstrated the clinical significance of SB for managing the recollapsed vertebra. This study may be helpful in the understanding of recollapse despite cement augmentation and in predicting the likelihood of a need for further surgical correction.

However, this study has some limitations. First, it had a retrospective design and was conducted at a single center. Second, some variables, including the amount of cement inserted, compliance with anti-osteoporosis treatment, and individual ambulatory status, could not be evaluated by a retrospective review of medical records. Third, our stringent inclusion and exclusion criteria could have introduced a degree of selection bias. Fourth, in the present study, only vertebroplasty was employed for cement augmentation and investigated, which was mainly due to the surgeon’s preference. Finally, various factors including back pain could affect the difficulties of proper measurements of the sagittal spinal parameters based on the whole-spine lateral standing radiographs. Future research with a larger study population along with a longer-term follow-up may better identify further correlations, allowing better prediction of the requirement for further corrective surgery.

**Conclusions**

In conclusion, SI, dynamic mobility within the vertebra, and lower BMD are risk factors for the recollapse of a vertebra following PVP for thoracolumbar OVF. SI was a significant predictor for further collapse of a cemented vertebra and the progression of sagittal deformity in patients with a recollapsed vertebra following PVP for thoracolumbar OVF. The degrees of thoracolumbar kyphosis and local kyphosis were not significant factors for predicting further collapse or the progression of spinal deformity. Therefore, spinopelvic parameters, rather than kyphosis angle, should be carefully evaluated based on the whole-spine lateral standing radiographs for thoracolumbar OVF at each follow-up visit. More careful and active management, including patient education regarding strict adherence to anti-osteoporosis medication and counseling with regard to the potential need for re-PVP and revision surgery, is required for patients with underlying thoracolumbar kyphosis or SI.

**Conflict of Interest**

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

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