Nonstructural versus structural bone grafting in the treatment of lumbar tuberculosis: a retrospective cohort study

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Research article

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Abstract

[Background] Various bone grafting have been reported to achieve anterior column support and bone fusion after one stage posterior debridement in the treatment of lumbar tuberculosis. However, nonstructural bone grafting has been rarely studied. This study aimed to compare the efficacy of nonstructural bone grafting versus structural bone grafting in the treatment of lumbar tuberculosis.

[Patients and methods] We retrospectively reviewed patients with nonstructural (n=27) and structural (n=22) bone grafting after single-stage posterior debridement and instrumentation for lumbar tuberculosis with at least 24 months of follow-up. Plain radiographs, magnetic resonance imaging, computed tomography scans, and health-related outcomes (e.g. Visual Analog Scale for back pain, the Oswestry Disability Index) were collected before surgery and at follow-up.

[Results] Both nonstructural and structural bone grafting were associated with significant improvements in the quality of life parameters, the laboratory tests and the Cobb angle of local kyphosis. A slight loss of Cobb angle correction was observed in two groups. Nonstructural bone grafting resulted in the significantly less operation duration, blood loss and lower bone fusion rate than structural bone grafting group. There were three complications in the nonstructural bone grafting and four complications in the structural bone grafting group, with no significant difference between two groups.

[Conclusions] Nonstructural bone grafting can achieve anterior column support with less surgical trauma and simpler surgical procedure than structural bone grafting, but the lower rate of grafted bone fusion should be taken into consideration when choosing nonstructural bone grafting.

Background

Spinal tuberculous (STB) is the most common type of extra-pulmonary tuberculosis[1], and accounts for nearly 50% of skeletal tuberculosis. STB is prone to occur at the lower thoracic and lumbar spine, and leads to vertebral collapse, kyphosis and even paralysis [2-3]. The goals of spinal tuberculosis surgery are to debride lesions, decompress spinal cord and reconstruct spinal stability. Nowadays the one stage posterior surgery is widely used for the treatment of thoracolumbar TB [4-5]. Posterior fixation played a significant role in the correction of kyphosis deformity and prevention of correction loss [6-7]. Anterior bone grafting requires the removal of the affected vertebral bone and intervertebral discs, which may be not necessary as reported by some studies [8-10]. Many types of bone grafting have been reported for successful usage, such as iliac bone, titanium mesh cage filled with bone granules, rib bone and so on[11]. To the best of our knowledge, there are few studies about the nonstructural bone grafting for lumbar TB.

Our previous study demonstrated that nonstructural bone graft achieved similar curative effects as structural bone graft for single segment thoracic tuberculosis and has the advantages of less surgical trauma and shorter bone graft fusion time[12]. The lumbar spine sustains more body weight than thoracic spine and is devoid of the protection of rib cage. This study aimed to investigate the anterior
column support and bone fusion of nonstructural bone graft after one stage posterior debridement in the treatment of lumbar TB. We have reported the successful use of nonstructural bone graft for lumbar STB[13-14], but these studies did not set control and the conclusions may be biased. Liu compared the clinical efficacy of one-stage posterior debridement with nonstructural autogenous bone graft versus anterior debridement, structural bone graft combined with posterior instrumentation [15]. The results showed the former method had a significantly shorter operation time, lower blood loss, shorter hospital stay and less hospitalization cost, except the bone graft fusion time. However, the conclusion may be biased because the surgical approaches used in the two groups were quite different.

The aim of this retrospective study was to compare the clinical and radiological outcomes of nonstructural versus structural bone grafts in combination with one stage posterior debridement and instrumentation for lumbar TB.

**Methods**

**Patients**

This study was approved by the Institutional Review Board of our hospital and the informed consent was obtained from all individual participants included in the study. The methods used in this study were conducted in accordance with the Declaration of Helsinki. From January 2015 to July 2017, 49 patients with the diagnosis of lumbar TB underwent surgery at our spine surgery center. The selection criteria were as follows: 1) adult lumbar TB; 2) the number of destructed vertebral body less or equal to three; 3) local spinal instability due to obvious bony destruction of the vertebral bodies; 4) follow-up time was beyond 24 months. The exclusion criteria included: 1) the requirement of osteotomy for severe pott’s deformity; 2) a history of lumbar spine surgeries by posterior approach. Patients with progressive neurologic deficit, persistent pain due to instability and kyphosis were the candidates for surgery.

Patients were divided into two groups due to the different methods of bone grafting. In group A, 27 patients received autogenous bone granules for grafting while 22 patients obtained titanium mesh filled with autogenous bone in group B after the similar procedure of posterior debridement and instrumentation. The following criteria should be satisfied when choosing nonstructural bone grafting: (1) bone defect ≥1/2 height of the vertebral body; (2) the local Cobb angle ≥20°; (3) at least one side of intact pedicle to ensure placing a pedicel screw; (4) at least one side of intact lateral wall of the vertebral body. In addition, if newly formed bone bridged the lesion vertebra, nonstructural bone grafting is also chosen. Of the 49 patients, 16 males and 11 females cases were included in group A, aged from 21 to 71 years old; while there were 12 males and 10 females cases in group B, aged from 26 to 69 years old. In group A, 8 patients were complicated with psoas major or iliac fossa abscess, while psoas major or iliac fossa abscess was present in 7 patients of group B.

**Preoperative preparation**
All patients obtained HRZE regimen for anti-TB treatment (isoniazid 300mg, rifampicin 450mg, ethambutol 1500mg and pyrazinamide 750 mg) at least 3 weeks before surgery. After admission to hospital, all patients underwent routine laboratory testing for excluding surgical contraindication. According to the program of enhanced rehabilitation after surgery (ERAS), a preoperative dietary regimen of high energy and protein was administered in the patients if needed. Surgery was performed when the patients resumed normal appetite without low fever or night sweat, and anemia was improved as hemoglobin≥ 100g/L. Erythrocyte sedimentation rate (ESR) and c-reactive protein (CRP) were lower than 80mm/h and 30mg/L, respectively. However, if the patients showed a rapid deterioration of neurological function, the surgery was performed immediately.

**Operative procedure**

In group A, patients were placed in prone position after general endotracheal anesthesia, and a longitudinal midline incision was made. Then the musculus sacrospinalis was stripped off from the lamina; while the facet joints at contralateral side were exposed through Wiltse approach for instrumentation. Pedicle screws were placed into one to two normal vertebrae superior and inferior the lesion segments. If the pedicle screw can be successfully placed into the lesion vertebrae, whatever unilaterally or bilaterally, upper and lower fixation of one segment was employed; if not, upper and lower fixation of two segments was conducted. One rod was temporarily fixed at the none-debridement side to maintain the spinal stability. Then a unilateral laminectomy was performed to clear off the caseous necrotic tissue, pus, granulation tissue, sequestrum and necrotic disc. The paravertebral or psoas abscess was drained off using a soft catheter tube to insert into the abscess cavity. Another 3cm long incision along the posterior superior iliac spine was made to expose the iliac crest and the cancellous bone was harvested for bone grafting (Fig.1); while at the lower lumbar spine the process was done in the same incision. After flushing by saline, 450 mg of isoniazid and 1.0 g of streptomycin powder were placed at the lesion site for local anti-tuberculosis treatment. Then the autogenous bone particles were placed into the intervertebral space and packed tightly. During the whole procedure the spinous process and ligament complex were kept intact, and the neuro elements were protected meticulously (Fig.1). Finally, the rods were locked, and then the incision closed. The pathological materials taken from the lesion site were sent for culture and histopathologic examination.

The similar procedure was applied in group B. Differently, the musculus sacrospinalis was stripped off from the lamina at both side, and posterior ligament complex, spinal process and vertebral lamina were removed to expose the neuro elements. Then pediculectomy was performed to remove the lesion and a titanium mesh filled with autogenous bone granules was implanted after distracting the vertebral interspace for kyphosis correction.

**Postoperative management**

Antibiotics were used to prevent infection in 3 days and the same regimen of anti-TB drugs were prescribed to all patients and continued for 12 to 18 months. The drainage tube at the debridement side was kept for 7 days to inject with isoniazid 450 mg and streptomycin 1.0g every day for local anti-TB treatment.
medication; whereas the drainage tube at contralateral side was pulled out within 48 hours. Blood routine test, ESR and CRP, liver and kidney function were periodically checked. A dietary of protein and energy rich was administered during the whole therapeutic period. All patients were encouraged for mobilization under the protection of brace in 3 days postoperatively for quicker rehabilitation.

**Follow-up index and Statistical analysis**

The operation time, blood loss and duration of hospital stay were recorded. The visual analogue scores (VAS) of back pain and Oswestry disability index (ODI) were evaluated at the time points of pre-operation, 1 week after operation, and last follow-up. X-ray films were taken regularly after operation and 1, 3, 6 and 12 months postoperatively and each year thereafter for each patient. The state of internal fixation and sagittal Cobb angle change were examined based on X-rays. While the computed tomography (CT) scan were examined at the first six-month postoperatively, and then taken periodically in every 6 months till intervertebral bone fusion was confirmed. Bone fusion was defined on thin-cut CT scan according to the standards as Burkus reported [16].

**Statistics**

Measurement data were recorded as mean +/- standard deviation. The independent-sample t test was used to compare the data between preoperative and postoperative measurements, and the data between two groups. While differences in bone fusion rate and number of complications between two groups were evaluated using Fisher exact test. Statistical significance was considered as P value < 0.05.

**Results**

**Clinical outcomes**

Patients in both groups were followed up for 24 to 48 months, with an average of 34.3 months. No significant difference of preoperative clinical data was found between the two groups. As shown in Table 1, the operation duration of group A was 4.1 ± 0.7 hours, significantly shorter than 4.8 ± 0.8 hours of group B (P = 0.000). The blood loss in group A was statistically lower than that of group B (335 ± 75 ml vs. 462 ± 120 ml, P = 0.000). The VAS scores and ODI index did not present a significant difference between the two groups before surgery and improved significantly after surgery and at last follow-up in both groups, and no difference was observed between the two groups. After surgery, a significant improvement of ESR and CRP was found at the final follow-up, and no statistical difference was found between the two groups (P=0.382 and 0.058 respectively).

**Tab.1 Comparison of clinical and radiological data between two groups**
| Measurements | Group A (27) | Group B (22) | P Value  | P Value  |
|--------------|--------------|--------------|----------|----------|
| Sex (male / female) | 16/11 | 12/10 | 0.746 | 0.746 |
| Age (X±S, year) | 52.6±13.7 | 49.5±10.5 | 0.387 | 0.387 |
| Operation duration (min) | 4.1±0.7 | 4.8±0.8 | 0.000 | 0.000 |
| Blood loss (ml) | 335±75 | 462±120 | 0.000 | 0.000 |
| Hospital stay | 14.3±4.5 | 14.6±4.4 | 0.841 | 0.841 |
| VAS score | | | | |
| Preoperative | 3.8±1.2 | 3.9±1.4 | 0.895 | 0.895 |
| Postoperative | 2.5±0.9 | 2.2±0.9 | 0.334 | 0.334 |
| Last follow-up | 0.9±0.8 | 1.1±0.9 | 0.522 | 0.522 |
| P value* | 0.000 | 0.000 | 0.000 | 0.000 |
| ODI (% ) | | | | |
| Preoperative | 38.1±11.9 | 41.2±14.0 | 0.405 | 0.405 |
| Postoperative | 31.6±7.8 | 31.5±8.3 | 0.953 | 0.953 |
| Last follow-up | 15.0±5.2 | 16.7±6.6 | 0.347 | 0.347 |
| P value* | 0.000 | 0.000 | 0.000 | 0.000 |
| ESR (mm/h) | | | | |
| Preoperative | 58.4±17.4 | 59.1±19.6 | 0.891 | 0.891 |
| Postoperative | 41.4±15.4 | 41.6±13.3 | 0.949 | 0.949 |
| Last follow-up | 14.6±6.7 | 16.3±6.6 | 0.382 | 0.382 |
| P value* | 0.000 | 0.000 | 0.000 | 0.000 |
| CRP (mg/L) | | | | |
| Preoperative | 30.5±11.4 | 32.5±16.5 | 0.619 | 0.619 |
| Postoperative | 17.9±7.3 | 20.8±8.7 | 0.211 | 0.211 |
| Last follow-up | 6.2±2.9 | 8.3±4.2 | 0.058 | 0.058 |
| P value* | 0.000 | 0.000 | 0.000 | 0.000 |
| Bone fusion rate | | | | |
| 6-month | 4/27 | 7/22 | 0.185 | 0.185 |
| 12-month | 12/27 | 20/22 | 0.001 | 0.001 |
| Last follow-up | 20/27 | 22/22 | 0.012 | 0.012 |
Radiological evaluation

As the films shown in Fig 2, 3, 4, the TB lesions were cured in all patients, but different intervertebral bone fusion rate was demonstrated based on CT scan between the two groups. As shown in Table 1, the bone fusion rate at 6-month postoperatively was 4/27 in group A and 7/22 in group B, with no statistical difference (P=0.185). While at 12-month postoperatively and last follow-up, the fusion rate was significantly lower in the nonstructural group. The preoperative kyphotic Cobb angle was significantly improved after surgery in both groups. Despite a slight loss of Cobb angle was present at the last follow-up, no statistical difference was found between the two groups (P=0.190). No related complications such as fixation failure or spinal deformity occurred.

Complications

The incidence of complications between the two groups was not significantly different (P=0.685). In group A, the TB recurred in one case with internal fixation loosening and kyphotic deformation of the lumbar spine as the patient didn’t regularly take the drugs. After restart of anti-TB medication and external fixation with a rigid brace, the disease was cured, and intervertebral bony fusion was achieved with local kyphosis left. Cold abscess recurred in 2 cases but disappeared after CT guided puncture for abscess drainage. No implant failures were found, or no additional surgeries were undergone in the cases of incomplete grafted bone union during the follow-up.

In group B, cerebrospinal fluid leakage occurred in 2 cases, which were healed via periodically clipping drainage tube for 1 week. One case complained weakness and neuralgia of unilateral lower extremity after awaking from general anesthesia and recovered after the treatment with medication in three months. The wound nonunion happened in 1 case and was healed after dress change lasting for one month.

Discussion

| Cobb angle | Preoperative | 5.2±8.9 | 6.2±9.3 | 0.690 |
|------------|--------------|---------|---------|-------|
|            | Postoperative | -4.0±5.3 | -6.6±6.1 | 0.062 |
|            | Last follow-up | -3.6±4.5 | -6.1±3.3 | 0.032 |
|            | P value* | 0.807 | 0.000 |
| Cobb angle loss° | 1.3±1.9 | 0.6±1.1 | 0.153 |
| Complications | 3/27 | 4/22 | 0.685 |

*measurements between the preoperative postoperative and last follow-up
This study compared the clinical effect of nonstructural autograft with titanium mesh filled up with autograft for bone graft and fusion in combination with posterior debridement and internal fixation for mono-segment lumbar TB. The results demonstrated that the nonstructural autograft was equivalent to titanium mesh with autogenous bone for spinal reconstruction but had the advantages of diminished operation time and surgical trauma, except for lower rate of grafted bone fusion. Compared with titanium mesh cage, the removal of spinal process and pedicle is not needed for the implantation of bone granules in the lumbar spine, which was associated with decreased surgical trauma and better maintenance of the spinal stability. Moreover, implantation of bone granules into the intervertebral space is much easier than placing a titanium mesh cage during the posterior procedure, in avoidance of interrupting neurological elements.

The objective of surgical treatment for STB is debridement, decompression, prevention or correction of deformity, bone grafting to achieve solid fusion and stable internal fixation[17]. To date, however, the surgical approach hasn't reached a consensus and has evolved for decades. The anterior approach enables to direct removal of the TB lesions, but the risks of injuring great vessels are relatively large especially at lower lumbar spine [18]. Combined anterior and posterior surgery improves the effects of debridement and spinal reconstruction, but the surgical trauma is enlarged [19]. The single posterior surgery become popular as the advantages of less surgical trauma, easier kyphosis correction, stable fixation and simple techniques [20-22].

The typical STB affects intervertebral disc and adjacent vertebral bodies, and their destruction often results in spinal instability and kyphosis. Based on the three-column theory of Denis [23] and load-sharing principles [24], anterior column reconstruction by a structural bone grafting was mandatory after removing the affected vertebral bones and intervertebral discs. Many types of bone grafts have been used, such as tricortical iliac bone, titanium mesh cage filled with bone granules, rib bone and so on [10, 25-26]. The autogenous strut harvested from iliac was frequently used to repair bone defects due to good osteogenesis, bone induction, bone conductibility and biocompatibility [27]. However, harvesting a big buck of bone may increase the complications of donor site including chronic pain and infection [28]. Harvesting spongy bone granules be achieved by a tiny skin incision (2 centimeters long) and fenestration of the posterior iliac crest, avoiding position change during the procedure and decreasing the donor site complications.

Titanium mesh cage used in our study was filled with resected cancellous bone from vertebral arch or even the iliac crest using the same procedure. Although several studies reported that titanium mesh cage exhibited an important potential in reliable spinal reconstruction, high bone fusion, sufficient sagittal profile maintenance and lower implant-related problems[29], placing a titanium mesh cage from posterior approach in the lumbar spine increased the operation duration and risk of injuring nerve roots[30]. The operation time was longer, and the intraoperative blood loss was more in the titanium mesh group in our study, as placing a titanium mesh required extensive bone removal during the operation, resulting in increased surgical time and intraoperative bleeding. Moreover, the postoperative VAS scores of back
pains were much higher because of extensive stripping of paravertebral muscle, sacrifice of posterior spinal structures and over-traction of nerve roots.

The most concern about nonstructural bone grafting is the failure of support and breakage of instrumentation as it seems to violate the rule of anterior column reconstruction. Nevertheless, several studies have reported the success of using nonstructural bone graft for lumbar STB [13-15]. Our previous study also demonstrated the nonstructural bone graft had comparable bony fusion rate and even shorter fusion time than titanium cage containing bone granules in single segment thoracic tuberculosis[12]. In the present study, however, the cancellous bone granules didn’t exhibit the similar bone graft fusion compared with titanium mesh cage contained autogenous bone. We think the reasons for failure of graft bone fusion in some cases may be insufficient volume of bone graft and resorption of grafted bone in the early stage. Owing to the preservation of the posterior structure and reliable pedicle rod fixation, complications like internal fixation failure or lumbar STB recurrence were not present in nonstructural bone graft group except one case who didn’t regularly take the drugs.

In the study, we found that the loss of Cobb angle correction was present in both groups. However, the Cobb angle loss in both groups is slight and the lumbar curvature are well maintained from post-operation to last follow-up, with no significant difference. However, the Cobb angle correction of nonstructural bone group was smaller than that of structural bone group, as the loose structure of granular bone could not bear the same supporting force as titanium mesh. To correct kyphosis through placement of contour rods and distracting intervertebral space, anterior support is necessary. In this case, structural bone grafts such as titanium mesh should be chosen instead of nonstructural bone grafts. Therefore, the essentials for success of nonstructural bone grafting are: 1) the posterior spinous process and ligamental complex should be retained during the operation to preserve the local stability. 2) during the procedure of debridement, the sclerotic bone and bone bridge between the anterior edge of vertebral bodies is preferred to be preserved. As more bony structure of vertebrae reserved, the volume of bone grafting needed is decreased and the demand for structural bone reconstruction could be avoided. 3) a rigid pedicle screw fixation should be guaranteed, without distraction of the vertebral interspace.

To the best of our knowledge, this is the first study comparing nonstructural bone with structural bone grafting in combination with posterior debridement and instrumentation for lumbar STB, and the results from long follow-up time are good. However, the study still has some limitations. Firstly, this is a retrospective study of a small-scale of patients from a single center of institution. Second, the patients are not chosen randomly, so the selection bias is inevitable.

**Conclusions**

In combination with single-stage posterior debridement and pedicle screws fixation, nonstructural bone grafting is associated with reduced surgical trauma, simplified surgical procedure and decreased intraoperative hemorrhage compared to structural bone grafting in the treatment of lumbar STB, but
showed the lower rate of bone fusion. The nonstructural bone grafting is better suitable for the cases with less bony defect after debridement and slight local kyphosis requiring no deformity correction.

**List Of Abbreviations**

**TB**: tuberculosis  
**STB**: spinal tuberculosis  
**ERAS**: enhanced rehabilitation after surgery  
**ESR**: erythrocyte sedimentation rate  
**CRP**: c-reactive protein  
**CT**: computed tomography  
**VAS**: visual analogue scores  
**ODI**: Oswestry disability index

**Declarations**

**Ethics approval and consent to participate** This study was approved by the Institutional Review Board of the First Affiliated Hospital of Chongqing Medical University and the informed consent was obtained from all individual participants included in the study. Serial number of the project was 2019 Ke Yan Lun Li (No. 2019-123).

**Consent to publish** Yes.

**Availability of data and materials** All data generated or analyzed during this study are included in this published article and its supplementary information files.

**Competing interests** The authors declare that they have no competing interests.

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**Authors’ Contributions** ZZH collected and interpreted the patient data and was a major contributor in writing the manuscript. LJX and PQQ analyzed the data and made the statistical analysis. ZY, LW and DX made the patients follow-up work and analyzed the radiological films of the patients. OYS devised the research protocol and performed the operations.

All authors read and approved the final manuscript.

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Figures
Figure 1

The picture taken during the operation shows the incision for exposing posterior structure and for iliac bone harvesting.

Figure 2

A 25 years old male with L4-5 TB reconstructed by none-structural bone a, b. Pre-OP sagittal and axial CT films demonstrate the destructed vertebral body and the iliac fossa abscess. c, d. The immediate post-OP X-ray films shows the internal fixation. e, f. Post-OP CT scan films demonstrate solid intervertebral bone fusion at 24 months’ follow-up.

Figure 3

A 39 years old male presented with L2-3 TB reconstructed by titanium mesh cage. a, b, c. Pre-OP X-ray and MRI films show the vertebral and disc destruction of the spine and epidural abscess; d, e. The immediate post-OP X-ray films showed pedicle screws fixation with anterior spine reconstruction using titanium mesh cage. f, g. The CT films at 24-month show solid intervertebral bone fusion. h, i. The X-ray films at 24-month post-OP show good spinal alignment after removing the instrumentation.
A 24 years old male with L1-3 TB reconstructed by non-structural bone a, b, c. Pre-OP sagittal CT and MRI films demonstrate the destructed vertebral bodies and local kyphosis. d, e. The immediate post-OP X-ray films show the reconstruction of the lumbar alignment with internal fixation. f. Six-month follow-up CT scan films demonstrate the graft bone fusion in not good. g. Two-year follow-up CT scan films show the intervertebral bone fusion as the formation of bone bridge and the lumbar alignment is maintained in spite of the bad graft bone fusion.