Kyphoplasty in the treatment of occult and non-occult metastatic vertebral tumors

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Abstract
To compare the efficacy and safety of kyphoplasty (KP) in the treatment of occult metastatic vertebral tumors (OMVT) and non-occult metastatic vertebral tumors (MVT).

From January 2013 to December 2017, 65 cases of occult metastatic vertebral tumors and 82 cases of metastatic vertebral tumors were selected and divided into 2 groups. After KP, they were followed up by a year of outpatient visits and telephone calls. The visual analogue scale (VAS) and Oswestry disability index (ODI) scores, the amount of bone cement injected, the change of vertebral height and the incidence of complications were recorded, compared and analyzed by SPSS software. t test was used to compare the differences between the same group of patients at different times and between the 2 groups of patients.

In the OMVT group, the operation time was 24.52 ± 4.24 minutes, the fluoroscopy time was 10.18 ± 1.53 minutes and the volume of bone cement was 3.62 ± 0.93 ml. The VAS score decreased from 7.26 ± 0.08 preoperatively to 2.77 ± 0.93 postoperatively (P < .01). The ODI score decreased from 64.89 ± 9.05 preoperatively to 25.82 ± 4.63 postoperatively (P < .01). In the MVT group, the operation time was 26.63 ± 4.61 minutes, the fluoroscopy time was 11.04 ± 2.15 minutes and the volume of bone cement was 4.09 ± 1.10 ml. The VAS score decreased from 7.73 ± 0.94 preoperatively to 3.22 ± 0.80 postoperatively (P < .01). The ODI score decreased from 69.20 ± 7.14 preoperatively to 28.02 ± 4.40 postoperatively (P < .01). The vertebral height of MVT patients was significantly improved after operation (P < .01), but there was no difference in OMVT patients (P > .05).

Occult metastatic vertebral tumors can be detected by Magnetic Resonance Imaging (MRI), and KP may be more effective and safer in the treatment of OMVT.

Abbreviations: KP = kyphoplasty, MRI = Magnetic Resonance Imaging, MVT = non-occult metastatic vertebral tumor, ODI = Oswestry disability index, OMVT = occult metastatic vertebral tumor, VAS = visual analogue scale.

Keywords: efficacy, kyphoplasty, metastatic vertebral tumor, occult, safety

1. Introduction
With the advancement of medical technology, the survival time of cancer patients is prolonged, and the number of advanced patients with primary tumors metastasis to the spine is also increasing.[1] Hu et al describe the epidemiological characteristics of patients with spinal metastases in China between 2007 and 2019. The spine is a common site of tumor metastasis, accounting for about 30% of bone metastases, and 40% to 70% of patients with advanced tumors eventually metastasize to the spine, which show that it is necessary for society and people to actively treat tumors.[2] 80% of spinal metastases involve the vertebral body and the incidence of thoracic vertebrae is the highest, followed by lumbar vertebrae and cervical vertebrae.[3,4] Tumor transferred to the spine can cause severe pain, vertebral fracture and spinal cord compression.[5] Previous studies have shown that at least a quarter of patients who die of cancer suffer from occult spinal diseases.[6] X-ray is usually the first method of imaging examination for patients with spinal diseases. The destruction and collapse of the vertebral body invaded by the tumors can be observed by X-ray examination.[7] However, when there is no obvious bone destruction, deformation and collapse in the early stage of metastatic vertebral body, X-ray examination is meaningless.[8] Magnetic Resonance Imaging (MRI) has high sensitivity and specificity in the diagnosis of metastatic vertebral tumors, and usually shows low signal in T1 sequence and high signal in T2 sequence.[9,10]

Kyphoplasty (KP) is commonly used in the palliative treatment of metastatic vertebral tumors and needs to restore vertebral...
height through an inflatable balloon and reduce the pressure of cement injection and the risk of leakage.\textsuperscript{11,12} From the perspective of elderly patients in the United States, KP are cost-effective compared with non-surgical treatment, even for the oldest patients.\textsuperscript{13} There is little discussion on the treatment of occult metastatic vertebral tumor (OMVT) in clinic, because OMVT patients are not easily detected in time. Therefore, we evaluated the safety and efficacy of KP in the treatment of OMVT and MVT respectively to determine whether it is safer and more effective to detect OMVT patients through MRI in time and treat them with KP.

2. Material and methods

2.1. Patients

This study was approved by the Ethics Committee of the First Affiliated Hospital of Suzhou University. From January 2013 to December 2017, patients with metastatic vertebral tumors in our hospital were screened. Most patients suffer from severe low back pain.

Inclusion criteria-All cases met the following criteria:\textsuperscript{14}

Metastatic vertebral tumors, diagnosed by a multidisciplinary team, which included an experienced radiologist, orthopedic surgeon and oncologist; the ability to maintain prone position for at least 2 hours; a Karnofsky performance score $>60$; according to the evaluation of oncologists, the expected survival time was longer than 1 year, less than or equal to 2 cases of metastasis, no other serious diseases were found. All follow-up patients were transferred to the oncology department for anti-cancer treatment after KP.

Exclusion criteria-Only one of the following criteria needed to be satisfied: Infections, psychiatric disorders, coagulation disorders, and other systemic diseases; spinal cord compression.

2.2. Surgical operation

The whole operation was performed by the professional spine surgeons. The patients under anesthesia lay prone on the operating table. Pedicle puncture guided by C-arm fluoroscopy. When the puncture needle entered the internal edge of pedicle under frontal fluoroscopy and the puncture needle entered the posterior edge of vertebral body under lateral fluoroscopy, the process indicated that the puncture was successful. After successful puncture, the core of the needle was pulled out. Then the cannula was fixed and used as the working channel, and the pneumatic balloon was inserted. Finally, the bone cement was injected slowly. The operation time, fluoroscopy time and the amount of cement injection were recorded during the operation. After completion of the KP, the patients were monitored for 8 hours postoperatively.

2.3. Clinical evaluation

The anterior and middle vertebral height were measured by X-ray lateral films from case data and outpatient reexamination preoperatively, 3 days and 1 year after operation. All patients were surveyed by questionnaires in hospitals or telephone interviews. Visual analogue scale (VAS) was used to evaluate the degree of back pain. The pain increased with the increase of the numerical value through the horizontal line from 0 to 10. Oswestry disability index (ODI) score was used to assess patients’ quality of life from different aspects. The higher the score, the more severe the dysfunction.\textsuperscript{11}

2.4. Complication

Postoperative complications such as fever, infection, pulmonary embolism and death were recorded, and cement leakage rate was evaluated by X-ray examination.

![Figure 1. Location of occult metastatic vertebral tumors (n=86) and metastatic vertebral tumors (n=104).](image-url)
2.5. Statistical analysis

The mean deviation and standard deviation of anterior and middle vertebral height, VAS and ODI scores were calculated and analyzed by SPSS software (SPSS 19.0, USA). The basic characteristics and results evaluation parameters of the 2 groups of data were compared by t test of group design data. When \( P < .05 \), the difference is statistically significant.

3. Results

3.1. Patients

From January 2013 to December 2017, patients with metastatic vertebral tumors received kyphoplasty in our hospital were 257. Patients who met the criteria were followed up after adequate evaluation. We performed a 1-year follow-up for the 147/172 patients. 10 patients died in 1 year because of underlying disease and 15 patients lost follow-up. 65 patients with OMVT received 86 KP and 82 patients with OMVT received 104 KP (Supplementary Fig. 1, http://links.lww.com/MD/E340). The metastatic vertebrae treated with KP ranged from T6 to L5 in Figure 1. General characteristics are compared of the patients of OMVT in Table 1. A representative case of a male patient with L3 OMVT who received KP treatment in Figures 2 and 3.

3.2. Surgical operation

In OMVT patients, the operation time for every vertebral body was 24.52 ± 4.24 minutes, and the fluoroscopy time was 10.18 ± 1.53 minutes. The amount of cement injected was 3.62 ± 0.93 ml. In MVT patients, the operation time for each vertebral body was 26.63 ± 4.61 minutes, and the fluoroscopy time was 11.04 ± 2.15 minutes. The amount of injected cement is 4.09 ± 1.10 ml. The operation time, fluoroscopy time and the amount of bone cement injection in the 2 groups had statistical significance \( (P < .01) \).

3.3. Clinical evaluation

In the OMVT group, The VAS score decreased from 7.26 ± 0.10 preoperatively to 2.77 ± 0.93 postoperatively \( (P < .01) \), and remained at 2.65 ± 0.89 1 year after operation. The ODI score decreased from 64.89 ± 9.05 preoperatively to 25.82 ± 4.63 postoperatively \( (P < .01) \), and remained at 29.49 ± 6.96 1 year after operation. The vertebral height of OMVT patients preoperatively and postoperatively had no difference \( (P > .05) \) (Table 2).

In the MVT group, The VAS score decreased from 7.73 ± 0.94 preoperatively to 3.22 ± 0.80 postoperatively \( (P < .01) \), and remained at 2.61 ± 0.84 1 year after operation. The ODI score decreased from 69.20 ± 7.14 preoperatively to 28.02 ± 4.40 postoperatively \( (P < .01) \), and remained at 28.37 ± 5.53 1 year after operation. The vertebral height of MVT patients improved significantly compared with that preoperatively \( (P < .01) \) (Table 2).

![Figure 2](http://www.md-journal.com)
3.4. Complication

By X-ray examination postoperatively, there was no leakage of bone cement in 65 OMVT patients receiving 86 KP treatments, but in 82 MVT patients receiving 104 KP treatments, 5 of them had leakage of bone cement.

4. Discussions

Jae Hwan-cho et al have found that early vertebral metastases are asymptomatic and only occur when the tumor invades the nerve or develops into a pathological fracture. Studies have shown that at least 30% of bone destruction or vertebral body deformation and collapse, X-ray can detect diseased tissue. However, MRI has a good imaging effect on soft tissues such as spinal cord and tumor. Therefore, X-rays are often prone to missed diagnosis for OMVT, and MRI is an important method for diagnosing OMVT. Treatment options for patients with metastatic vertebral tumors are usually palliative, usually for the purpose of improving quality of life and reducing or eliminating pain. Traditional treatments include painkillers, bed rest, radiation therapy, chemotherapy, and surgery. Although these treatments are palliative treatment, patients are generally faced with the dependence of painkillers, poor therapeutic effects of radiotherapy and chemotherapy, and inability to tolerate open surgery. With the development of minimally invasive surgery, it provides a new way to solve the above problems. Some studies have shown that KP can significantly reduce pain in patients with...
metastatic vertebral tumors, prevent vertebral collapse and improve quality of life compared with non-surgical treatment.[20] The purpose of our study was to describe our experience and assess the safety and effectiveness of KP in the treatment of OMVT and MVT. In 65 patients with OMVT and 82 patients with MVT, we found that VAS and ODI scores were significantly improved after KP treatment. We found that there were significant differences in VAS and ODI between OMVT patients and MVT patients preoperatively, suggesting that patients with MVT suffered more severe pain and lower quality of life than those with OMVT. In OMVT patients, the vertebral height remained unchanged after KP, while in MVT patients, the vertebral height improved significantly after KP. In 1 year after operation, there was no significant difference in vertebral height between the 2 groups. The results showed that KP was effective in the management of OMVT and MVT, and early MRI examination and timely KP treatment can help OMVT patients to suffer from less pain, avoid developing into MVT and improve their quality of life. The exact mechanism of pain relief in these operations remains controversial. It is likely that the pain is relieved by stabilizing the fracture with cement. Another explanation is that monomer toxicity and exothermic aggregation of bone cement lead to necrosis of pain receptors in bone.[21]

The leakage of bone cement is still the main complication of KP. It has been reported that the leakage rate of bone cement in the treatment of metastatic spinal tumors with KP is 7%. [14] According to our experience, good surgical techniques can reduce some leakage caused by vertebral wall defects. Cement injection must be done slowly and carefully. Once a cement leak is detected, the operator can adjust the direction of the needle or stop the injection immediately. When cement reaches the edge of the vertebral body, the injection process stops. In our retrospective study, 65 patients with OMVT underwent 86 KP operations without leakage, and 82 patients with MVT received 104 KP operations, 5 of which had leakage. First, MVT patients have more cement volume than OMVT patients, so the risk of leakage is higher. In addition, compared with OMVT patients whose shape of the vertebral body is good, the vertebral body destruction of MVT patients is obvious, and even the vertebral body wall is broken, which is considered to increase the risk of bone cement leakage during surgery.[22,23] Finally, because of the invasion of tumor tissue, the strength of vertebral body decreases. In the process of KP, the balloon destroys the vertebral bone when restoring the height of vertebral body, and it is easy to cause leakage.[24]

This study has some limitations. First, the study was retrospective and produced less evidence than prospective studies. Secondly, our study only included patients receiving KP, without control or alternative treatment, such as vertebroplasty. Further prospective studies are needed to evaluate the efficacy and safety of KP therapy. Finally, we studied the efficacy of KP 1 year after surgery. Further studies are needed to determine the safety and effectiveness of KP in the treatment of OMVT and MVT.

5. Conclusion
In order to reduce the pain and improve the quality of life of patients with advanced cancer, patients with back pain, especially those with a history of cancer, even if the X-ray results are normal, also need to be examined by MRI for OMVT. Compared with MVT, KP may be more effective and safer in the treatment of OMVT.

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Table 2

| Parameters                  | OMVT       | MVT       | P       |
|-----------------------------|------------|-----------|---------|
| Kyphoplasty operation       |            |           |         |
| Operation time per vertebrae (minutes) | 24.52 ±4.24 | 26.63 ±4.61 | .005    |
| Fluoroscopy time per vertebrae (minutes) | 10.18 ±1.53 | 11.04 ±2.15 | .008    |
| Injected cement volume (ml) | 3.62 ±0.93 | 4.03 ±1.10 | .007    |
| VAS                         |            |           |         |
| Preoperatively              | 7.26 ±1.08 | 7.73 ±0.94 | .006    |
| 3 days postoperatively      | 2.77 ±0.93 | 3.22 ±0.80 | .002    |
| 3 months postoperatively    | 2.72 ±0.78 | 2.66 ±0.74 | .61     |
| 1 year postoperatively      | 2.65 ±0.99 | 2.61 ±0.64 | .80     |
| ODI                         |            |           |         |
| Preoperatively              | 64.98 ±9.05 | 69.20 ±7.14 | .002    |
| 3 days postoperatively      | 25.82 ±4.63 | 28.02 ±4.40 | .004    |
| 3 months postoperatively    | 25.77 ±4.90 | 26.51 ±5.11 | .37     |
| 1 year postoperatively      | 24.99 ±6.96 | 28.37 ±5.53 | .80     |
| Anterior vertebral height   |            |           |         |
| Preoperatively              | 26.25 ±4.11 | 18.09 ±4.33 | .000    |
| 3 days postoperatively      | 26.62 ±5.17 | 25.79 ±3.93 | .13     |
| 1 year postoperatively      | 25.89 ±4.25 | 28.04 ±3.75 | .79     |
| Middle vertebral height     |            |           |         |
| Preoperatively              | 26.04 ±3.72 | 17.72 ±4.53 | .000    |
| 3 days postoperatively      | 25.58 ±4.05 | 25.22 ±2.98 | .49     |
| 1 year postoperatively      | 25.81 ±3.76 | 25.66 ±3.25 | .75     |

* P < .01 compared to preoperative value.
† P > .05 compared to preoperative value.
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