CT fluoroscopy-guided percutaneous vertebroplasty in patients with multiple myeloma: analysis of technical results from 44 sessions with 67 vertebrae treated

Christoph Trumm, Tobias Jakobs, Anne Pahl, Robert Stahl, Thomas Helmbberger, Philipp Paprottka, Maximilian Reiser, Ralf-Thorsten Hoffmann

PURPOSE
This study aimed to assess the results of computed tomography (CT) fluoroscopy-guided vertebroplasty in patients with multiple myeloma, focusing on the frequency and clinical impact of polymethylmethacrylate (PMMA) leaks.

MATERIALS AND METHODS
From December 2001 to August 2008, 39 patients (17 females, 22 males; mean age, 65±7 years) with multiple myeloma suffering from painful spinal osteolyses underwent vertebroplasty. A total of 67 vertebrae were treated in 44 sessions under CT fluoroscopy (single-slice, 4-row CT, and 16-row CT). In the planning CT scan, osteolytic destruction (i.e., none, ≤25%, ≤50%, ≤75%, or ≤100%) was assessed regarding the vertebral cross-sectional area and adjacent soft tissues in the CT fluoroscopic cross-sectional image and of the needle inclination. Intradiscal, intraspinal, paravertebral, and intercostovertebral/posterior canal, and the outer circumference. CT performed after vertebroplasty was used to detect local PMMA leaks. Patient charts were retrospectively reviewed with special respect to peri and postinterventional adverse events. Clinical outcomes were assessed on a visual analog scale (VAS) 24 hours before, 24 hours after, and 6 months after vertebroplasty.

RESULTS
Overall, 37.3%, 12.0%, and 6.0% of vertebrae showed at least 50% osteolytic involvement of the cross-sectional area, the cortical border of the spinal canal, and outer vertebral cortex, respectively. Intradiscal, intraspinal, paravertebral, and intercostovertebral/posterior leaks were seen in 21.6%, 35.1%, 43.3%, and 0% of vertebrae, respectively. The ratio of basivertebral to segmental venous leaks was 16.2%/40.3%. No major complications occurred. The mean VAS score decreased significantly (P < 0.05) from 6.4 at 24 hours before vertebroplasty to 3.2 at a mean follow-up of 9.0 months.

CONCLUSION
Vertebroplasty in multiple myeloma can be performed safely under CT fluoroscopy, even with substantial destruction of the vertebral cross-sectional area or cortical bone. A high clinical success rate was achieved, regardless of whether PMMA leaks were present.

Key words: • multiple myeloma • vertebroplasty • fluoroscopy • computed tomography

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compression fractures. Moreover, we wanted to assess the clinical success rate of vertebroplasty with regard to pain reduction, patient comfort, and the ability to reduce the use of analgesic agents.

Materials and methods

Patient selection

Percutaneous vertebroplasty is an accepted and widely used method to stabilize fractured vertebral bodies. Our institutional ethics board did not demand approval of this retrospective study regarding the review of patient charts and the assessment of clinical outcomes after vertebroplasty using a standardized questionnaire. The principles of the Declaration of Helsinki were followed. All patients were referred by the local oncology department. The diagnosis of myeloma had been made using standard clinical criteria (15).

Clinical indications for vertebroplasty were regularly confirmed by an interdisciplinary team of oncologists, radiation oncologists, spinal surgeons, and interventional radiologists prior to the intervention. A biopsy of the spinal segment to be treated with vertebroplasty had not generally been performed prior to vertebroplasty, and biopsy results were not analyzed separately.

Each patient underwent a physical examination by the referring clinician to identify the painful or fractured vertebral level(s). Informed consent by the patient or his/her legal guardian to undergo vertebroplasty was obtained 24 hours before and directly prior to the intervention after an extensive explanation of the method and its potential complications as well as alternative treatments.

Between December 2001 and August 2008, 224 patients with tumor involvement of the spine underwent vertebroplasty, 39 of whom (17 females, 22 males) suffered from painful spinal osteolyses and compression fractures due to MM. The mean age of the patients was 65±7 years, with a range from 51 to 82 years. All patients referred for vertebroplasty suffered from severe back pain refractory to conservative analgesic therapy, either alone or in combination with chemotherapy and/or radiation therapy.

According to the Society of Interventional Radiology (SIR) quality improvement guidelines for percutaneous vertebroplasty (which are in accordance with the Cardiovascular and Interventional Radiological Society of Europe quality assurance guidelines for percutaneous vertebroplasty) (16), radiculopathy in excess of vertebral pain (caused by a compressive syndrome unrelated to vertebral collapse), asymptomatic retropulsion of a fracture fragment causing significant spinal canal compromise, and asymptomatic tumor extension into the epidural space were regarded as relative contraindications to vertebroplasty (17). Absolute contraindications were patients improving on analgesic therapy, myelopathy in patients with spinal canal compromise due to retropulsion of bone fragments or a tumor, active local or systemic infections, uncorrectable coagulopathy, and an allergy to bone cement or opacification agents (17).

Imaging workup and guidance

Before vertebroplasty, previous cross-sectional images not older than two weeks, such as those from CT, magnetic resonance imaging (MRI) or positron emission tomography (PET)/CT, were checked for all patients by a team of interventional radiologists. Additionally, all patients underwent in-house MRI (T1-weighted sequence ± contrast, short tau inversion recovery sequence, axial and sagittal orientation) prior to vertebroplasty to assess the pattern of bone marrow involvement and vertebral and paravertebral tumor involvement and to detect the level(s) of vertebral edema in cases of multiple VCFs detected on CT (18).

Each patient underwent a pre and postinterventional CT scan (Siemens Somatom Plus 4, Sensation 4 or 16 MDCT, Siemens Medical Solutions, Forchheim, Germany) of the involved spine level, including at least two vertebral body heights above and below the osteolytic/fractured vertebral body. The CT scan involved 3-mm slices and coronal and sagittal multiplanar reconstructions (MPRs) to visualize the extent of vertebral compression and bone destruction, to detect possible involvement of the posterior wall of the vertebral body, and to plan the needle trajectory for vertebroplasty. Using the postinterventional CT scan, the distribution of polymethylmethacrylate (PMMA) in the vertebral body and cement leaks were analyzed.

For CT guidance, a single-detector Siemens Somatom Plus 4 scanner (120 kV, 15 to 25 mA, 4×2.5 mm collimation, 3-mm slices), a Siemens Sensation 4 (120 kV, 15 to 25 mA, 4×1 mm collimation, 3-mm slices) or a Sensation 16 MDCT (120 kV, 15 to 25 mA, 12×0.75 mm collimation, 3-mm slices) with CT fluoroscopy was used (CARE Vision CT, Siemens Medical Solutions). CT fluoroscopy was performed with angular beam modulation (HandCare™, Siemens Medical Solutions), i.e., the X-ray beam was turned off within a 120° angle sector above the patient to decrease radiation exposure to the interventional radiologist and patient (19). The entire vertebroplasty procedure, including needle placement and PMMA injection, was performed under CT fluoroscopic guidance only. The needle was inserted under intermittent single-shot CT fluoroscopic acquisitions. Continuous CT fluoroscopy was only used during the PMMA injection. The needle tip and the adjacent cement distribution within the vertebral body were monitored in real time on a ceiling-mounted, in-room monitor (256×256 imaging matrix interpolated to 1024×1024 for display) by moving the table along the z-axis in incremental steps of 1.0 mm using a control panel attached to the CT table. Beginning in February 2006, CARE View, with a synchronous display of three adjacent slices on the in-room monitor, was employed. Precautions with respect to the radiation protection of the operator during CT fluoroscopy included aprons, thyroid shields and eyeglasses of 0.5-mm lead equivalent. An additional shield was put onto the lower half of the patient before sterile draping to reduce the amount of scattered radiation.

Procedure

All procedures were performed by one of the board-certified authors or by residents under their supervision who were experienced in CT-guided interventions. The vertebroplasty needles (OptiMed Medical Devices, Ettingen, Germany) utilized were 15, 13, or 10 gauge and had a length of 10 or 15 cm. In the cervical spine, an anterolateral approach (patient in the supine position) was generally used, while an intercostovertebral approach (patient in the prone position) was used in the thoracic spine. A transpedicular access
was chosen in the thoracic spine only when the pedicular diameter was large enough. In the lumbar and sacral spine, a 10 G needle with transpedicular or posterolateral (in cases of insufficient pedicle diameter or pedicle osteolysis) access was applied in most cases.

All patients received preinterventional antimicrobial antibiotic with 2 g of mezlocillin (Baypen®, Bayer AG, Leverkusen, Germany). During the procedure, each patient was monitored with pulse oxymetry. After sterile draping and disinfection of the skin overlying the treated vertebral body, local anesthesia with 10–20 mL of 2% mepivacaine hydrochloride (Scandicain®, AstraZeneca GmbH, Wedel, Germany) was applied to the soft tissue and the periosteum along the access route. Additional conscious sedation with midazolam hydrochloride (5 mg of intravenous Dormicum®, Ratiopharm GmbH, Ulm, Germany) and piritramide (15 mg of intravenous Dipidolor®, Janssen-Cilag GmbH, Neuss, Germany) was administered in six cases of additional radiofrequency ablation before vertebroplasty.

A small skin incision was made, and the vertebroplasty needle was advanced with a sterile hammer through the cortical bone under intermittent single-shot CT fluoroscopic control. The direction of the needle was additionally guided by turning the beveled tip in 90° steps. The insertion was stopped when the needle tip reached the anterior third of the vertebral body or the center of the osteolysis. Three types of bone cement were used during the study period: Osteopal 40, Osteopal V, and Bioment Bone Cement V (Biomet Deutschland GmbH, Berlin, Germany). After the needle positioning, the PMMA was mixed in a sterile bowl according to the manufacturer’s guidelines. To reduce the risk of extravertebral leakage and venous extravasation, the cement injection was started with the application device (Optimed Gangi Cemento-Re Gun, OptiMed Medical Devices) after another 60–90 s when the PMMA was in its pasty polymerization phase. The PMMA was slowly injected by the operator under intermittent CT fluoroscopy using sequential withdrawing of the needle and incremental table movement along the z-axis to assess the localization and direction of cement distribution within the vertebral body. In cases of initial cement extravasation to the adjacent disc spaces or to the soft tissue structures on the real-time sectional CT fluoroscopic images, the injection was immediately stopped by depressurizing the application screw. If the leakage persisted, the needle tip was repositioned by turning the handle bar in 90° steps or by sequentially withdrawing the needle. The cement injection was stopped when at least one- to two-thirds of the osteolytic lesion was filled. In cases of two or three vertebrae treated in one session, the vertebra with major cortical destruction was treated last to ensure a pasty consistency of the cement. The stylet of the needle was finally inserted again to empty the needle cavity completely, and then both were removed by slowly turning the handle bar. Then, a CT scan of the treated vertebral level was immediately performed for documentation of the cement distribution. After the intervention, all patients were sent back to the ward for further clinical monitoring.

**Technical outcomes**

In a retrospective analysis, two interventional radiologists who were experienced in performing CT-guided vertebroplasty (RTH and CT) retrospectively evaluated pre and postinterventional CT scans, as well as the perinterventional CT fluoroscopic datasets acquired in the 44 vertebroplasty sessions for the purpose of obtaining a consensus regarding the degree of vertebral osteolytic involvement, and the types of cement leakage.

**Vertebral osteolytic destruction**

The degree of osteolytic destruction was quantified in the preinterventional CT dataset by visual estimation (Fig. 1) of the following:

1. The osteolytic destruction of the vertebral cross-sectional area, including the vertebral body and the pedicles
   (a. 0%–25%, b. 26%–50%, c. 51%–75%, d. 76%–100%)

2. The cortical border of the posterior wall and spinal canal
   (a. 0%, b. 1%–25%, c. 26%–50%, d. 51%–75%, e. 76%–100%)

3. The cortical border of the outer vertebral body
   (a. 0%, b. 1%–25%, c. 26%–50%, d. 51%–75%, e. 76%–100%)

4. The presence of pathologic fractures was evaluated on both axial images and multiplanar reconstructions.
Vertebroplasty procedure

The type of needle access to the vertebral body (anterolateral, intercostovertebral, transpedicular, or posterolateral) was evaluated in the perinterventionally acquired CT fluoroscopic datasets. The amount of PMMA (mL) applied was taken from the report that documented the procedure.

Types of cement leakage

According to the compartment involved, the following types of cement leakage were differentiated:
- intradiscal (upper or lower intervertebral space),
- intraspinal (including nonvascular soft-tissue leaks through the cortical bone and vascular leaks into the basivertebral vein),
- paravertebral (including nonvascular leaks through cortical defects into the surrounding soft tissue and vascular leaks into the segmental veins), and
- intercostovertebral/posterolateral (along the needle access path).

Vascular leaks were assessed separately if a basivertebral or segmental vein could be clearly identified.

In cases of several leakage types occurring at the same vertebral level, every leak location was counted as a separate event.

Clinical outcomes and adverse events

Twenty-four hours before and after the procedure, all patients were asked to answer a standardized questionnaire assessing the following items:
1. Pain level on a 10-point (0 [no pain]–10 [maximum pain intensity]) visual analog scale (VAS)
2. Type and dose of analgesic agents.

Six months after the procedure, all patients were sent the same questionnaire to assess any changes in analgesic agent use, subjective changes in the severity of pain (better, no change, or worse) and quantitative VAS scores. Clinical complications that occurred during the procedure or that had been reported by the referring clinicians within 24 hours after the procedure were documented in the final report. Additionally, clinical patient charts were retrospectively reviewed with respect to adverse events potentially related to the procedure that occurred more than 24 hours after the vertebroplasty.

Statistical analysis

The data analysis was performed using a commercially available software (Statistical Package for Social Sciences, version 18, SPSS Inc., Chicago, Illinois, USA). P values of less than 0.05 were considered statistically significant. A Wilcoxon rank sum test was used for comparisons of VAS pain scores 24 hours before vertebroplasty and after the follow-up period.

Results

The total number of differently treated vertebrae in 44 sessions was 67 (one level, 25; two levels, 15; three levels, 4 sessions), including 3 cervical, 32 thoracic, 27 lumbar and 5 sacral vertebrae. All 67 vertebrae were treated successfully in the first session (Fig. 2). The access route was anterolateral
in 3, intercostovertebral in 30, transpedicular in 28, and posterolateral in 6 vertebrae. For the purpose of tumor-debulking, six patients underwent additional radiofrequency ablation of the spinal segment and were consecutively treated with vertebroplasty (three thoracic and three lumbar vertebrae).

**Vertebral osteolytic destruction**

Table 1 shows the absolute number and percentage of treated vertebrae that are characterized by a 0%, 1%–25%, 26%–50%, 51%–75%, or 76%–100% ratio of destructed bone of 1. the cross-sectional area, 2. the cortex of the posterior wall, and 3. the outer cortical circumference. A total of 25 vertebrae (37.3%) were characterized by an osteolytic destruction of more than 50% of the cross-sectional area. The proportion of vertebrae with an osteolytic involvement of the posterior wall was 49.3% (33 vertebrae), with 8 vertebrae (12%) showing more than 50% osteolytic posterior wall destruction. Thirty-four (50.7%) and 18 (26.8%) of the 67 vertebrae showed neither cortical bone destruction of the posterior wall nor destruction of the outer vertebral circumference, respectively. Pathologic fractures were present in 37 of the 67 vertebrae (55.2%).

**PMMA filling volume**

The mean filling amount of PMMA was 2 mL in the cervical (n=3), 3.5 mL in the thoracic (n=32), 3.9 mL in the lumbar (n=27), and 5 mL in the sacral spine (n=5).

### Table 1. Distribution of osteolytic destruction in treated vertebrae

<table>
<thead>
<tr>
<th>Osteolytic destruction</th>
<th>Incidence</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cross-sectional area (%)</td>
<td></td>
</tr>
<tr>
<td>0–25</td>
<td>4 (6%)</td>
</tr>
<tr>
<td>26–50</td>
<td>38 (56.7%)</td>
</tr>
<tr>
<td>51–75</td>
<td>15 (22.4%)</td>
</tr>
<tr>
<td>76–100</td>
<td>10 (14.9%)</td>
</tr>
<tr>
<td>Cortical bone of posterior wall/spinal canal (%)</td>
<td></td>
</tr>
<tr>
<td>0</td>
<td>34 (50.7%)</td>
</tr>
<tr>
<td>1–25</td>
<td>15 (22.4%)</td>
</tr>
<tr>
<td>26–50</td>
<td>10 (14.9%)</td>
</tr>
<tr>
<td>51–75</td>
<td>5 (7.5%)</td>
</tr>
<tr>
<td>76–100</td>
<td>3 (4.5%)</td>
</tr>
<tr>
<td>Cortical bone of outer vertebral body (%)</td>
<td></td>
</tr>
<tr>
<td>0</td>
<td>18 (26.8%)</td>
</tr>
<tr>
<td>1–25</td>
<td>28 (41.8%)</td>
</tr>
<tr>
<td>26–50</td>
<td>17 (25.4%)</td>
</tr>
<tr>
<td>51–75</td>
<td>2 (3%)</td>
</tr>
<tr>
<td>76–100</td>
<td>2 (3%)</td>
</tr>
</tbody>
</table>

### Table 2. Distribution of treated vertebrae (n=67) and number of vertebrae showing PMMA leaks per spinal section

<table>
<thead>
<tr>
<th>Spinal section</th>
<th>Treated vertebrae</th>
<th>Vertebrae with leaks</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cervical</td>
<td>3</td>
<td>0</td>
</tr>
<tr>
<td>Thoracic</td>
<td>32</td>
<td>20</td>
</tr>
<tr>
<td>Lumbar</td>
<td>27</td>
<td>12</td>
</tr>
<tr>
<td>Sacral</td>
<td>5</td>
<td>2</td>
</tr>
<tr>
<td>Total</td>
<td>67 (100%)</td>
<td>34 (50.8%)</td>
</tr>
</tbody>
</table>

### Cement leaks

A total of 33 (49.2%) of the 67 vertebrae treated presented with no cement leaks (Table 2). In the remaining 34 vertebrae (50.8%), 37 leaks were detected (Fig. 3). Most leaks occurred in the thoracic (56.8%) or lumbar (37.8%) spine (Table 3). Cement leaks were observed within the paravertebral soft tissues (43.3%), in the intraspinal/epidural space (35.1%), and within the neighboring intervertebral discs (21.6%). A pathologic fracture was present in all vertebrae showing intradiscal leaks, and most pathologic fractures (62.5%) were seen in the thoracic spine (cervical, 1; thoracic, 20; lumbar, 15; sacral, 1). A segmental vein was clearly identified in 15 of 16 paravertebral leaks (93.8%), and a basivertebral vein was identified in 6 of 13 (46.2%) intraspinal leaks. No leaks were observed in the intercostovertebral/posterolateral region.

Correlating the presence of PMMA leaks to the degree of osteolytic destruction, most vertebrae showing leaks were either in the category of an osteolytic involvement of 26%–50% of the cross-sectional area (n=19) or outer cortical border (n=15) or were in the category with no osteolytic involvement of the posterior wall (n=17) (Fig. 4).

### Semiquantitative/subjective outcome analysis

Six months after vertebroplasty, 11 of 39 patients (28.2%) did not answer the questionnaire (n=8) or were lost to follow-up due to death (n=3). On a 10-point VAS, the mean pain level showed a significant decrease from 6.4 (24 hours before vertebroplasty) to 3.9 (24 hours after vertebroplasty; \(P = 0.030\)) (n=28). A significant reduction of the VAS score to 3.2 (\(P = 0.009\)) was observed after the follow-up period (mean follow-up of 9.0 months) (Fig. 5).

Twenty patients (71.4%) reported a reduction of pain, four patients (14.3%) reported no change in pain, and four patients (14.3%) reported an increase in pain due to a progressive course of the disease. Ten patients (35.7%) were able to stop taking analgesic medication, and eight patients (28.6%) were able to either reduce the dose of their analgesic medication by at least 50% or replace opioids with non-narcotic drugs, such as non-steroidal anti-inflammatory drugs (NSAIDs) or metamizol. Six patients (21.4%) had not changed their medication, and four...
patients (14.3%) had increased their use of analgesic drugs or had changed to opioids at the time of the survey.

Most patients (n=18; 64.3%) were satisfied with the procedure, while the remaining patients (n=10; 35.7%) complained about discomfort during the treatment under local anesthesia. The main points of criticism were being placed in the prone position, the duration of the procedure, and the discomfort during needle insertion.

**Adverse events**

No major adverse events (i.e., neurological symptoms requiring surgical decompression, pulmonary embolism, or death) according to the SIR definition (minor hospitalization...
less than 48 hours, prolonged hospitalizations longer than 48 hours for major therapy, permanent adverse sequelae, or deaths) were observed after dismissal of the patients to the ward or were documented in the clinical records of the referring clinicians 6 months after the procedures (17). Moreover, no cases of infection within the treated spinal segments were found after the follow-up period. In patients with obvious local cement extravasation detected on the postinterventional CT scan, no chest pain, dyspnea or significant decrease of blood oxygen saturation was observed by the operators perinterventionally or immediately after the procedure.

After the application of the local anesthetic drug, one patient developed a generalized seizure that was successfully managed by the anesthesiologist on call. The vertebroplasty was successfully performed one week later under general anesthesia.

Two patients treated at the first lumbar and first sacral vertebral level, respectively, showed prolonged bleeding at the site of the skin incision and needle entry; they were successfully treated with manual compression and an additional single-knot suture.

Discussion

Percutaneous vertebroplasty has been described as a safe and effective minimally invasive method for the treatment of tumoral involvement of the spine, resulting in rapid and continuing clinical improvements in pain and patient mobility (5, 6, 20, 21). Several reports have focused on the use of vertebroplasty, particularly in the treatment of osteolyces and compression fractures in myeloma patients (22–30). Conservative pain management with bed rest, bracing, bisphosphonates and analgesic drugs is often not sufficient to control the acute and chronic spinal pain accompanying pathologic fractures in myeloma patients. While the stabilizing effect of radiotherapy with

Table 3. Different PMMA leakage types per spinal section

<table>
<thead>
<tr>
<th></th>
<th>Cervical</th>
<th>Thorac</th>
<th>Lumbar</th>
<th>Sacral</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>Intradiscal</td>
<td>0</td>
<td>5</td>
<td>3</td>
<td>0</td>
<td>8 (21.6%)</td>
</tr>
<tr>
<td>Intraspinal</td>
<td>0</td>
<td>8</td>
<td>4</td>
<td>1</td>
<td>13 (35.1%)</td>
</tr>
<tr>
<td>Paravertebral</td>
<td>0</td>
<td>8</td>
<td>7</td>
<td>1</td>
<td>16 (43.3%)</td>
</tr>
<tr>
<td>Intercostovertebral/posterolateral</td>
<td>0</td>
<td>0</td>
<td>0</td>
<td>0</td>
<td>0</td>
</tr>
<tr>
<td>Total</td>
<td>0</td>
<td>21</td>
<td>14</td>
<td>2</td>
<td>37 (100%)</td>
</tr>
<tr>
<td>Venous</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Basivertebral vein</td>
<td>0</td>
<td>4</td>
<td>2</td>
<td>0</td>
<td>6 (16.2%)</td>
</tr>
<tr>
<td>Segmental vein</td>
<td>0</td>
<td>8</td>
<td>7</td>
<td>0</td>
<td>15 (40.5%)</td>
</tr>
</tbody>
</table>

Intraspinal leaks included nonvascular soft-tissue leaks through the cortical bone and vascular leaks into the basivertebral vein. Paravertebral leaks included nonvascular leaks into the surrounding soft tissue and vascular leaks into the segmental vein. Venous leaks were separately assessed if a vein could be clearly identified. In cases of several leakage types occurring at the same vertebral level, each leak location was counted as a separate event.
The aforementioned cases did not show an amount of 4 mL PMMA injected, overall leakage rate of 21% and a mean of 14% of the treated vertebrae. Given an and epidural disease was present in patients suffering from MM of the spine. The primary intentions of the procedures were the relief of back pain refractory to NSAIDs, opioid medication, and radiation therapy, as well as the stabilization of malignant VCFs or extensive osteolyses.

Chew et al. (31) recently published a meta-analysis of 30 studies that included 987 patients who underwent vertebroplasty due to spinal metastases and myeloma. Several studies have demonstrated that an applied cement volume of greater than 4 mL correlates with an increased rate of complications (4.2%–13.5%) (32, 33). All authors reported small-volume local, intradiscal or paravertebral cement leaks, but none of them was asymptomatic. In a patient group with tumoral posterior wall involvement of the treated vertebrae, no increase in the complication rate was observed.

In a further meta-analysis of 69 studies by Hulme et al. (34), PMMA leaks without clinical consequences were also frequently observed (27 vertebroplasty studies; 41% vertebral leakage rate), while only 2.6% of the treated vertebrae (3.9% of patients) among 31 studies resulted in clinical complications.

Tancioni et al. (28) reviewed 28 vertebral levels in a cohort of MM patients treated at their institution. In 10 cases, vertebroplasty was performed at the time of diagnosis (six patients after chemotherapy and four before systemic therapy), and vertebroplasty was performed following disease progression in four patients (two patients were treated after radiotherapy received for vertebral body fracture). Fracture or substitution of part of the posterior wall or pedicular tract was seen in 28%, and epidural disease was present in 14% of the treated vertebrae. Given an overall leakage rate of 21% and a median amount of 4 mL PMMA injected, the aforementioned cases did not show any cement leaks.

Bartolozzi et al. (24) reported 19 procedures (10 kyphoplasties, 9 vertebroplasties) performed in 14 patients (13 patients in Durie-Salmon stage IIIA, 1 patient in stage IIA) with MM. Vertebroplasty was chosen in collapsed vertebral bodies in cases of wide destruction of the posterior wall or in cases of increased bone density. Kyphoplasty was chosen in cases of a high risk of cement leakage due to vertebral instability. No complications were observed in either patient group. The authors concluded that both techniques can be safely and effectively performed in every phase of the disease, independent of concomitant chemotherapy.

Ramos et al. (26) performed 19 vertebroplasty procedures in 12 consecutive myeloma patients. Despite a leakage rate of 84%, no patient developed neurological or other clinical symptoms.

It is commonly known that the rate of cement leakage detected depends on the utilized imaging modality. Therefore, when comparing leakage rates, the method of leak detection should always be mentioned. Schmidt et al. (35) compared the sensitivity of intraoperative fluoroscopy and postoperative radiography in the detection of cement leakage on postoperatively-acquired CT scans. With only lateral or biplane (anteroposterior and lateral) radiograms, the authors could only detect 34% or 48% of the leaks verified on CT, respectively. Due to the concavity and overprojection of the posterior wall in the thoracolumbar spine, the majority of leaks into the basivertebral vein (50% to 60%) were not detected with biplane radiography. Using intraoperative fluoroscopy, the detection rate was even lower (21%) due to the inferior image resolution.

In our patient group, local PMMA leaks were detected in 34 of 67 treated vertebrae (50.8% leakage rate). Given an injected cement volume ranging between 2 mL in the cervical and 5 mL in the sacral spine, 37.3% of the treated vertebrae were characterized by osteolytic destruction of more than 50% of the cross-sectional area and by osteolytic involvement of the posterior wall in 49.3% of the treated vertebrae. Aside from the aforementioned periprocedural adverse events that could be successfully managed, there were no major adverse events observed in our patient group. As our results and those of other authors showed, the number and volume of extrasosseous cement leakages alone did not correlate with clinical complications (10). In our study, the amount of PMMA leakage was not calculated but was very small in most cases.

While osteolytic destruction of the posterior vertebral body wall and significant epidural tumor extension have been considered as relative contraindications for vertebroplasty (16), CT fluoroscopy enables the operator to detect initial cement extravasation through the posterior wall; therefore, this imaging modality might increase the safety of the vertebroplasty procedure, especially in those cases. According to our experience, with the use of a tube current-time product for CT fluoroscopic acquisitions between 15 to 25 mAs, a sufficient visualization of the posterior wall during needle insertion and cement injection is possible, and relevant cement extravasation or dislocation of fragments can be avoided. In combination with careful needle repositioning, initial leaks can be stopped if the injection is paused for at least 20–30 s, until the viscosity of the PMMA has increased.

The mechanism of pain reduction following PMMA injection during vertebroplasty or kyphoplasty is still unclear. The analgesic effect is mainly attributed to stabilization that prevents both microscopic movements and macroscopic collapse due to osteolytic destruction (36). Kaufmann et al. (37) and Cotten et al. (38) could not find a significant association between the amount of injected PMMA and the analgesic effect because even small cement volumes lead to partial or complete pain relief. Other hypotheses regarding the underlying mechanism include the destruction of nerve endings through the exothermic reaction of PMMA during polymerization and its additional cytotoxic effects (39). These results are in accordance with our observations because even patients with a small vertebral filling volume of 1–2 mL of PMMA (13 of 67 vertebrae; 19.4%) in the thoracic and lumbar spine have reported substantial pain relief.

McDonald et al. (22) presented clinical outcome data from 67 MM patients undergoing vertebroplasty over a 7-year period. They reported a statistically significant improvement on the
Rolland-Morris Disability Questionnaire, as well as in the VAS values that their patients documented immediately after the procedure and during the follow-up period of up to 1 year. In particular, the authors observed a similar short- and long-term reduction in scores when comparing their results in myeloma patients to earlier analog measures of pain at rest and during activity in patients with osteoporotic fractures (40), suggesting a similar prognostic outcome in both patient groups. Our retrospective assessment of pain levels showed comparable, significant mid-term (mean follow-up of 9.0 months) clinical success in the majority of the myeloma patients regarding both the semiquantitative (VAS score) and subjective (reduction of pain; use of analgesic agents) parameters that were assessed.

In addition to evaluating the mid-term analgesic effect after vertebroplasty (as attributed by the patients to the procedure) and the changes in the use of analgesic medication with respect to dose or strength, we did not separately assess the presence or duration of chemo- or radiation therapy administered prior to or after the vertebroplasty procedure. The patients were usually referred to our department due to intractable back pain, despite the above-mentioned, non-invasive, palliative treatment regimens, or due to painful, pathologic compression fractures. After the treatment, systemic chemotherapy and/or local radiotherapy—if conformable with the general patient condition—were continued. As a consequence, an interference with the analgesic effect of vertebroplasty cannot be excluded in most cases.

Regarding the assessment of the clinical outcomes with the questionnaires, another drawback of this study is the moderate response rate of only 28 of 39 patients who completed the last questionnaire after 6 months.

In our patient group with osteolyses and VCFs due to MM of the spine, primary technical success, in terms of the PMMA injection during the first session, was reached in all vertebrae, and significant clinical success was maintained after a 9-month follow-up period. No clinical or neurological sequelae were evident during the follow-up period. Given an experienced operator, CT fluoroscopy provides excellent visualization of the posterior wall and paravertebral soft tissues during needle placement and PMMA injection. This process enables palliative treatment to be given to myeloma patients, even with substantial osteolytic vertebral destruction.

**Conflict of interest disclosure**

The authors declared no conflicts of interest.

**References**


28. Tancioni F, Lorenzetti M, Navarria P, et al. Percutaneous vertebroplasty in multiple myeloma patients documented immediately after the procedure and during the follow-up of 9.0 months. No clinical or neurological sequelae were evident during the follow-up period. Given an experienced operator, CT fluoroscopy provides excellent visualization of the posterior wall and paravertebral soft tissues during needle placement and PMMA injection. This process enables palliative treatment to be given to myeloma patients, even with substantial osteolytic vertebral destruction.

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