CLINICO-PATHOLOGICAL PROFILE AND SURGICAL OUTCOME IN VERTEBRAL COLLAPSE OF NEOPlastic ORIGIN

Kapil Jain¹, Abhinav Jain², Jayesh Sardhara³, Amit Singh⁴, Arun Kumar Srivastava⁵

¹Consultant, Department of Neurosurgery, Max Super Speciality Hospital, Saket, New Delhi.
²Assistant Professor, Department of Radiodiagnosis, Hamdard Institute of Medical Sciences and Research & HAHC Hospital, New Delhi.
³Senior Resident, Department of Neurosurgery, Sanjay Gandhi Postgraduate Institute of Medical Sciences, Lucknow.
⁴Associate Professor, Department of Neurosurgery, Sanjay Gandhi Postgraduate Institute of Medical Sciences, Lucknow.

ABSTRACT

BACKGROUND
Data addressing the clinical/histopathological profile and post-operative neurological outcome in patients of neoplastic vertebral collapse is scarce. This study was done with an aim to learn about these facts in patients of vertebral collapse of neoplastic origin from our region.

METHODS
It was an historical retrospective cum prospective study in which 56 patients operated for neoplastic vertebral collapse were included from historical case records. After pre-operative evaluation and recording of data, spinal cord decompression and tumour resection with/without instrumentation had been done depending on the individual case. All surgical details, intra-operative and post-operative events were also recorded. Biopsy of the surgical specimen was considered gold standard to define etiology. Patients were given perioperative adjuvant therapies as per the recommended published protocols. Neurological status was reassessed after a minimum follow-up of 6 months. Descriptive statistics were applied on the data.

RESULTS
Most common primary tumour was plasmacytoma and most common primary site for spinal metastasis was thyroid. Thoracic spine was most commonly involved. Posterolateral approach was used in 59% of the primary tumours and 69% of metastasis. Instrumentation was done in 31% and 33% patients of primary and secondary tumours respectively. There was no intra-operative mortality in the primary tumour group, but one post-operative mortality. In the metastasis group there were 2 intra-operative deaths, but no post-operative mortality. Improvement in Frankel grade was seen on follow-up in 68% primary tumours and 81% patients of metastasis. Subjective pain relief was present in 86% and 100% patients of primary and secondary tumours respectively.

CONCLUSIONS
The neurological outcome of neoplastic vertebral collapse was good with multimodality therapeutic approach approaching re-establishment of ambulation in 44-63% patients.

KEYWORDS
Neoplastic, Neurological Deficit, Pain Relief, Vertebral Collapse.


INTRODUCTION
Spine is unique among the other parts of skeletal system due to its load-bearing and nerve-protecting function. Loss of its structural integrity as a result of primary or metastatic tumours may cause severe pain and/or paralysis. These symptoms impair the ambulatory ability of the patients and worsen their quality of life, thus requiring surgical interventions.

Metastatic tumours are far more common than the primaries of spine. In patients with metastatic disease, 30-70% has involvement of spine.[1-4] With an increase in survival of cancer patients with multimodality therapeutic approach, the need for management of symptomatic spinal metastasis has also increased.

To the best of our knowledge, there is no published Indian data in the indexed medical literature with reference to the clinical and histopathological profile including the neurological outcome after surgery in neoplastic vertebral collapse. In this communication, we present our experience with vertebral collapse of neoplastic aetiology and their outcome following neurosurgical treatment.

METHODS
In this historical retrospective cum prospective study, 56 patients of vertebral collapse of neoplastic aetiology presenting in the Department of Neurosurgery were included. Post-traumatic and post-infectious causes were excluded. These patients had been operated over an eight-year period (January 2000 - December 2008). Data of patients was collected from inpatient and pre-operative outpatient file records, discharge summaries and follow-up outpatient records.
The presenting complaints, duration of symptoms, detailed clinical neurological assessment and radiological findings (level of involvement, number of vertebrae involved) were recorded for all study subjects. In patients with metastatic disease, site of primary tumour was searched as per the published recommended guidelines. Pre/post-operative investigations including chest X-ray, full skeletal survey, sonography (USG) abdomen, urine and serum electrophoresis for M-bands, prostate specific antigen, thyroid scanning gynaecological examination including PAP smear, bone marrow biopsy, bone scan and local site biopsy were performed as and where indicated.

Collapsed vertebra (X-ray and MRI based) was defined as 1) Vertebra with fracture of end plates adjacent to the osteolytic lesion and 2) Vertebra with reduction of vertebral body height because of pathological fracture of the anterior and/or lateral cortex of vertebral body [Figure 1]. Vertebral height was considered to be reduced when the height of affected vertebral body was less than 90% of the estimated original height. This was calculated from an average of the corresponding measurements at adjacent unaffected levels above and below the involved vertebra.[6]

Pre-operative morbidity was scored objectively in form of Frankel grade.[6]

Frankel Classification
- Complete lesion (Paraplegia).
- Only sensory function present.
- Motor function present, but of no practical use (Non-ambulatory).
- Motor function present, sufficient to allow walking (Ambulatory).
- No neurologic signs or symptoms.

Patients underwent spinal cord decompression and tumour resection with/without instrumentation depending on the need of individual case, location, maximum bulk of the tumour, neuraxial compression and the decision of the operating surgeon. Surgical approach adopted along with mention regarding instrumentation (if done) was also noted down. Biopsy of the surgical specimen was considered gold standard for defining the aetiology of the collapse. Intra- and post-operative complications were recorded.

Patients were given post-operative adjuvant therapies as per the standard guidelines in the form of radiotherapy, chemotherapy and radioiodine ablation in the departments of radio-oncology, haematology and nuclear medicine respectively. Patients with minimum follow-up of 6 months were included in the follow-up analysis. Frankel score was assessed again at the time of last follow-up. Motor status was considered improved when the patient upgraded by one/more Frankel grade. Pain was considered improved when either it absolutely disappeared or decreased as per the patient’s verbal statement. Objective pain scales were not used. Descriptive statistics were applied on the data.

RESULTS
Out of total 56 included cases of vertebral collapse, histopathology of the surgical specimen suggested 27 (48.2%) primary tumours and 29 (51.8%) secondary/metastases.

Most common primary tumour causing vertebral collapse was Plasmacytoma (12/27, i.e. 44.4%) followed by Ewing sarcoma (4/27, i.e. 14.8%). Other aetiologies seen in the primary tumour group were osteoscleroma (3), lymphoma (3), chordoma (1), aneurysmal bone cyst (1), haemangioma (1), round cell tumour (1) and giant cell tumour (1) [Table 1].
As per the historical records of metastatic vertebral collapse, primary site for the spinal metastasis could not be identified in 44.8% (12/29) cases. Most common primary site in spinal metastasis was thyroid (follicular carcinoma) in 34.5% (10/29) cases. Other primary sites identified were liver (2), prostate (2), kidney (1), ovary (1) and cervix (1) [Table 1].

In the primary tumour group, the mean age at presentation was 45.4±16.6 years. Male:female ratio was 4.4:1. Local pain was the most common presenting complaint in all (100%) patients. Along with local pain, 25.9% (7/27) patients had radicular pain also. Limb weakness was also present in all (100%) patients (Frankel grade A-D). Overall, 77.8% (21/27) patients were non-ambulatory (Frankel grade A-C). Sensory deficits were present in 74.1% (20/27) and bladder involvement was present in 48.3% (13/27) patients [Table 2].

In the metastasis group, the mean age at presentation was 52.2±11.5 years. Male:female ratio was 1.2:1. Local pain was the most common presenting complaint in this group also. Approximately, 10% patients had radicular pain also in association with local pain. Limb weakness was present in all (100%) patients. Overall, 86.2% (25/29) patients were non-ambulatory. Sensory deficits were present in 93.1% (27/29) and bladder involvement was present in 48.3% (14/29) patients [Table 2].

Thoracic spine was the most common site of collapse in 77.8% (21/27) cases of primary tumour. Similarly, in the metastatic group also, the thoracic spine was most common site of collapse in 82.7% (24/29) cases. None of the patients had C1/C2 or atlanto-axial joint involvement [Table 3].

As a special mention about 12 patients of plasmacytoma, all had solitary lesion and the pre-operative diagnosis was available, which was done on the basis of criteria laid by International Myeloma Working Group, 2003.[7,8] None of the patients had evidence of underlying systemic multiple myeloma. Although radiotherapy alone may be the treatment of choice in solitary plasmacytoma, surgery is required in a few patients with vertebral instability or in patients with radio-resistance.[7-10] All these 12 patients of solitary plasmacytoma were operated, because they all had accompanying vertebral collapse with spinal instability and progressive neurological deficits. Post-operative radiotherapy was given as per the standard protocols.[6,11-13] There is insufficient data for the role of adjuvant chemotherapy in solitary plasmacytoma and thus it was not given to these patients.

Surgical details, complications, mean follow-up period and outcome in terms of improvement/deterioration in motor status and pain were recorded [Table 4]. Postero-lateral approach was more commonly used in our series, 59.3% (16/27) in the primary tumour group and 69.0% (20/29) in the secondary tumour group. Overall, the instrumentation was done in 33.3% (9/27) of the primary tumour patients and 31.0% (9/29) of the secondary tumour patients. There was no intra-operative mortality in the primary group, but the post-operative mortality was 3.7%
(1/27). In the metastatic group, the intra-operative mortality was higher (2/29=10.7%), because of uncontrollable bleeding (1 patient renal cell carcinoma and 1 follicular carcinoma of thyroid); there was no post-operative mortality.

In the primary tumour group, instrumentation was done in 9 patients (5 plasmacytoma, 2 Ewing’s sarcoma, 1 aneurysmal bone cyst and 1 giant cell tumour). Among these 9 patients undergoing instrumentation, there was no intra- or post-operative mortality or post-operative complication. On follow-up (mean period 26.7±31.0 months), 85.7% (6/7) had motor improvement and 100% had pain improvement. One patient which did not show motor improvement was plasmacytoma of thoracic region operated from anterior approach. In the 18 non-instrumented patients of primary tumour group, 78.6% (11/14) patients improved in their Frankel grade and all (100%) showed pain improvement as seen over a follow-up period of 28.7±25.2 months.

Regarding metastases group, 8 patients had instrumentation and none of the instrumented patients suffered mortality or any post-operative complication. Seven patients could be followed for a period of 11.6±4.2 months and 71.4% (5/7) showed motor improvement. Pain improvement was also present in 71.4% (5/7). Out of 21 non-instrumented patients, follow-up was available for 15 patients for a period of 21.3±20.0 months. There was motor improvement in 66.7% (10/15) patients and pain relief in 93.3% (14/15) patients.

Overall, patients of both primary and secondary groups had good neurological outcome in terms of motor improvement, which was 81.0% (17/21) in the primary group and 68.2% (15/22) in the secondary group. Pain relief was present in all 100% (21/21) patients from the primary group and 86.4% (19/22) of the patients with secondaries. Plasmacytoma patients, where half (6/12) were operated by anterior approach and half (6/12) by postero-lateral approach, 77.8% (7/9) had motor improvement and all 100% (9/9) had pain improvement. Among the follicular carcinoma of thyroid, 80.0% (8/10) were approached postero-laterally and 20.0% (2/10) anteriorly. Motor improvement was present in 75.0% (6/8) patients and all 100% (8/8) had pain relief.

Fig. 1: T2 Weighted MRI Sagittal View showing Collapsed D4 Vertebra

### Table 1: Histopathological Diagnosis in Neoplastic Vertebral Collapse

<table>
<thead>
<tr>
<th>Histopathology of Primary Spinal Tumours - n (%)</th>
<th>Primary Sites for Metastasis Spine - n (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Solitary Plasmacytoma - 12 (44.4)</td>
<td>Thyroid (Follicular Carcinoma) - 10 (34.5)</td>
</tr>
<tr>
<td>Ewing Sarcoma - 4 (14.8)</td>
<td>Liver - 2 (6.9)</td>
</tr>
<tr>
<td>Osteoclastoma - 3 (11.1)</td>
<td>Prostate - 2 (6.9)</td>
</tr>
<tr>
<td>Lymphoma - 3 (11.1)</td>
<td>Kidney - 1 (3.4)</td>
</tr>
<tr>
<td>Chordoma - 1 (3.7)</td>
<td>Ovary - 1 (3.4)</td>
</tr>
<tr>
<td>Aneurysmal bone cyst - 1 (3.7)</td>
<td>Cervix - 1 (3.4)</td>
</tr>
<tr>
<td>Haemangioma - 1 (3.7)</td>
<td>Unknown - 12 (41.4)</td>
</tr>
<tr>
<td>Round cell tumour - 1 (3.7)</td>
<td></td>
</tr>
<tr>
<td>Giant cell tumour - 1 (3.7)</td>
<td></td>
</tr>
</tbody>
</table>

### Table 2: Clinical Profile of Patients with Neoplastic Vertebral Collapse

<table>
<thead>
<tr>
<th>Duration of symptoms</th>
<th>Motor weakness present (Frankel Grade A-D)</th>
<th>Non-ambulatory (Frankel Grade A-C)</th>
<th>Pain +</th>
<th>Sensory deficit +</th>
<th>Bladder involvement +</th>
</tr>
</thead>
<tbody>
<tr>
<td>Total number of cases</td>
<td>27</td>
<td>29</td>
<td>21/27 (77.8%)</td>
<td>20/27 (74.1%)</td>
<td>20/27 (74.1%)</td>
</tr>
<tr>
<td>Mean age (in years)</td>
<td>45.4±16.65 (7-74)</td>
<td>52.21±11.5 (32-77)</td>
<td>27/29 (93.1%)</td>
<td>7/27 (25.9%)</td>
<td>13/27 (48.1%)</td>
</tr>
<tr>
<td>Sex ratio (Male:Female)</td>
<td>4:1</td>
<td>1.2:1</td>
<td>26/29 (90.7%)</td>
<td>0/29 (0.0%)</td>
<td>14/29 (48.3%)</td>
</tr>
<tr>
<td>&lt;1 month</td>
<td>5/27 (18.5%)</td>
<td>6/29 (20.7%)</td>
<td>20/27 (74.1%)</td>
<td>0/27 (0.0%)</td>
<td>7/29 (24.1%)</td>
</tr>
<tr>
<td>1-6 months</td>
<td>12/27 (44.4%)</td>
<td>16/29 (55.2%)</td>
<td>7/27 (25.9%)</td>
<td>27/29 (93.1%)</td>
<td></td>
</tr>
<tr>
<td>&gt;6 months</td>
<td>10/27 (37.0%)</td>
<td>7/29 (24.1%)</td>
<td>13/27 (48.1%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Local site only</td>
<td>20/27 (74.1%)</td>
<td>0/27 (0.0%)</td>
<td>27/29 (93.1%)</td>
<td>13/27 (48.1%)</td>
<td>14/29 (48.3%)</td>
</tr>
<tr>
<td>Radicular only</td>
<td>0/27 (0.0%)</td>
<td>0/29 (0.0%)</td>
<td>14/29 (48.3%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Both</td>
<td>7/27 (25.9%)</td>
<td>3/29 (10.3%)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

Vertebral column is the most common osseous site for secondaries. Conversely, primary spinal tumours are relatively rare comprising only 10% or less of all tumours to the spine.[2] In our series of operated neoplastic vertebral collapse, there was almost equal distribution of primary and secondary spinal tumours. This is contradictory to other reports mentioning secondaries being far more frequent. This discrepancy could have been because of the fact that we had selectively included patients with collapse, which required surgery whereas other reports consist of all spinal tumour patients either with or without collapse.

In the previously published reports - most common primary benign tumours were haemangioma, osteoblastoma and osteochondroma. Plasmacytoma and Ewing sarcoma were the most common primary malignant tumours of spine.[1,11,12,14-22]

Among the metastatic tumours of spine - lungs, breast, prostate and kidneys were the most important primary sites.[11,15,18,23] In the index study, Plasmacytoma was the most common primary tumour and thyroid was the most common primary site for spinal metastasis. This difference in aetiologies could be due to: (a) The pattern for the aetiology of vertebral tumours is different in our region with a different genetic and environmental background in contrast to the Western countries; (b) Certain aetiologies of primary vertebral tumours like plasmacytoma and secondary vertebral tumours like follicular carcinoma of the thyroid are more aggressive in causing vertebral damage and ultimately collapse; (c) It could be due to the referral bias also to our tertiary care hospital.

There was a male sex predilection for the primary spinal tumours (male:female ratio 4.4:1). This is in agreement with the previous reports.12,24,25 In both primary and metastatic tumour groups, thoracic spine was most common site of involvement in 74% and 83% cases respectively. This fact also agree with the previously published data,11,26 and can be explained by the fact that the thoracic spine is the longest segment of vertebral column and thus the mathematical probability of involvement is maximum there.

In the last 2 decades, more effective neurosurgical techniques have been developed to provide dorsal and ventral spinal stabilisation. This has allowed the surgeons to become more aggressive with spinal cord decompression and tumour resection. In general, the surgical approach is dictated by the segment of spine involved (Cervical, thoracic, lumbar, sacral), location of the tumour in spinal segment (Anterior, posterior, right, left or circumferential to the neural elements), nature of the tumour, type of spinal reconstruction or stabilisation required after tumour resection. Anterior approach helps to achieve better decompression in metastatic tumours. Vertebral body resection for the tumour is now most commonly performed via anterior approach followed by vertebral column reconstruction and anterolateral plating. Posterior stabilisation with instrumentation is indicated in those with significant damage to posterior elements, significant kyphosis, thoracolumbar junction lesions, 2 or more adjacent vertebrectomies.[22,27]

Recent literature has revealed that the surgical outcome has improved with advances in the surgical techniques for spinal metastasis. However, it is difficult to generalise because of the diversity of surgical approaches,
decompressive and reconstructive techniques, patient variables and tumour pathology. Most series report that more than 50% patients of spinal metastasis receive significant neurological recovery and more than 90% patients receive pain relief.[28-30] Even the more recent studies[13,31-34] support the fact that surgical treatment improves outcome in patients with spinal metastasis. Hirabayashi et al[31] reported that the ambulatory status improved from 38% in the pre-operative period to 71% in the post-operative period and there was neurological improvement in about 50% of the patients. Same findings were supported by other authors also, neurological improvement in up to 60% of the patients.[13,32-34] Pain relief (Complete/partial) was seen in 56-96% of the patients.[13,31-34] Major complications including pneumothorax, sepsis, wound infection, wound dehiscence, pulmonary embolism and hardware failure, and minor complications including urinary infection, pulmonary infection and CSF leaks were reported in 11-32% cases.[13,31-34]

To the best of our knowledge, there is no published Indian literature describing the neurological outcomes after surgery in neoplastic vertebral collapse. In the present study, we found neurological improvement (Upgradation of Frankel score by one/more) in 81% and 68% patients of vertebral collapse with primary or metastatic aetiology respectively. In the primary tumour group 24% patients were ambulatory in the pre-operative period, 71% were ambulatory in the post-operative period and 63% non-ambulatory patients achieved ambulation with the treatment. In the metastasis group 13% patients were ambulatory in the pre-operative period, 55% were ambulatory in the post-operative period and 44% non-ambulatory patients became ambulatory after treatment. Pain relief (Complete/partial) was present in all of the primary tumour patients and 86% of the metastasis patients.

REFERENCES


