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Review

SURGICAL TREATMENT OF METASTATIC DISEASE OF THE VERTEBRAL COLUMN

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SUMMARY – Spinal metastatic disease is a rather common occurrence and definitely warrants attention and treatment due to the high likelihood of leaving cancer patients severely disabled in their final months of life. Recent developments in the understanding of the behavior of different tumor types, as well as advances in surgical treatment, are allowing for the evolution of treatment algorithms, especially when surgical treatment is to be considered. This paper gives an overview of the decision-making process and the array of surgical options currently available.

Key words: Spine – surgery; Spine – pathology; Spinal neoplasms, secondary; Neoplasm metastasis – surgery; Palliative care – methods; Algorithms; Decision support techniques

Introduction

The purpose of this article is to give an overview of the approach to treating spinal column metastases in cancer patients. This is a complex field which requires thorough understanding of all treatment modalities available and decisions must be guided by a rational framework based on a number of parameters.

Spinal metastases are by no means an uncommon phenomenon, as virtually any malignant tumor type has the potential to metastasize to bone, with the spine being the predominant bony site¹.

Since some form of bony metastasis is almost to be expected in advanced stage malignant disease, it may be discovered through targeted screening using any of a number of diagnostic modalities (CT/PET-CT, MRI, bone scan). However, in case of an unknown primary tumor, symptoms from the metastases may be the first signs of any illness. In this case, there are many 'red flags' which have been described to help guide the need for spinal malignancy screening². Although none of these red flags except for prior history of cancer carried much weight as a warning sign, careful use of combinations of red flags can be of benefit (age over 50, pain not relieved after a month of rest, thoracic pain which is worse at night, unexplained weight loss, generally poor state of health, etc.)².

It must be borne in mind that the goals of treatment of spinal metastases differ from the goals of treating primary tumors considering that treatment is seldom curative. In metastatic cases, surgeons strive to improve the overall quality of life in a patient otherwise undergoing treatment for their primary tumor by decreasing pain, relieving neurological impairment, providing stability, and increasing patient independence¹.

Non-surgical treatment includes chemotherapy, radiotherapy, hormonal therapy (for example, in breast and prostate tumors), angiography and embolization (in highly vascular metastases, including renal cell,

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Classification	Neurologic symptom and degree of bone destruction
Class 1	Minimal neurology
Class 2	Involvement of bone without collapse or instability and minimal neurology
Class 3	Major neurologic impairment without spinal instability
Class 4	Vertebral collapse and instability, without major neurologic impairment
Class 5	Vertebral collapse and instability with major neurologic impairment
Treatment principle	
Class 1, 2	Chemotherapy, hormonal therapy, radiotherapy
Class 3	Radiotherapy with corticosteroid treatment
Class 4, 5	Surgery

Table 1. Harrington's classification and treatment principle system

Table 2. Revised Tokuhashi scoring system for treatment decision guidance based on expected outcome

Characteristic	Score		
General condition (performance status)		77 . 10	
Poor (PS 10 ~ 40%)		Total Score	Conservative treatment
Moderate (PS 50 ~ 70%)	1	0 - 8	Conservative treatment
Good (PS 80 ~ 100%)	2	Predicted prognosis	Palliative surgery
No. of extraspinal bone metastases foci		9 - 11	0,
≥ 3	0	Predicted prognosis	 Single lesion No metastases to the major
1 ~ 2	1	6 months ≤	internal organs
0	2	12 - 15	Excisional surgery
No. of metastases in the vertebral body		Predicted prognosis 1 year ≤	
≥ 3	0		
2	1		
0	2		
Metastases to the major internal organs			
Unremovable	0		
Removable	1		
No metastases	2		
Primary site of the cancer			
Lung, osteosarcoma, stomach, bladder, esophagus, pancreas	0		
Liver, gallbladder, unidentified	1		
Others	2		
Kidney, uterus	3		
Rectum	4		
Thyroid, breast, prostate, carcinoid tumor	5		
Palsy			
Complete (Frankel A, B)	0		
Incomplete (Frankel C, D)	1		
None (Frankel E)	2		
Criteria of predicted prognosis: Total Score			
$(TS) 0 \sim 8 \Rightarrow 6 mo; TS 9 \sim 11 \le 6 mo; TS 12 \sim 15 \le 1 yr$			

thyroid, melanoma, giant cell tumors, and hepatocellular carcinomas), and symptom management. Radiotherapy is often used with or without surgery and must be considered where histopathology of the primary tumor is one of known radiosensitivity (breast, prostate, Wilm's tumor, Ewing's sarcoma, lymphoma, seminoma, multiple myeloma, medulloblastoma, etc.), although even in relatively radioresistant histologies (lung, renal, etc.) pain relief may be achievable³. Chemotherapy has its role in chemosensitive tumors (lymphomas, Ewing's, Wilm's, rhabdomyosarcoma, teratoma, etc.).

Surgery can also have an important role in cases in which it is deemed feasible. Surgical treatment is, of course, not without its own risks to the patient and benefits of the selected treatment modality must always outweigh the risks to the patient. It has long been held as general consensus that minimal life expectancy of 3 to 6 months is the minimum requirement for considering a major surgery⁴. However, in shorter life expectancies, palliative surgical procedures (kyphoplasty, vertebroplasty, placement of a morphine pump) could be beneficial notwithstanding the morbidity they present⁵. In order to help systematize the decision-making process, several classification scoring systems, as well as treatment algorithms have been developed which take into account parameters such as primary tumor histopathology, severity of impairment, overall wellbeing, and likelihood to receive any benefit from surgery.

Harrington has proposed the treatment algorithm shown in Table 1 solely based on the amount of vertebral destruction and instability^{6,7}. Enneking, Tomita, and Weinstein/Boriani/Biagini describe patterns of tumor spread throughout a vertebra8. The Tokuhashi scoring system shown in Table 2 is one of the commonly used tools for treatment guidance^{9,10}. It takes into account the general state of the patient as evaluated by the Karnofsky performance score (shown in Table 3), the number of local (spinal), extraspinal and visceral metastases, tumor histology, and neurological status according to Frankel (summarized in Table 4). However, the modified Bauer score has been shown to be a simpler system because it considers only the parameters which have been shown to mostly influence prognosis (shown in Table 5)¹¹. It is reasonable to consider combining several systems in order to synthesize final decision. For example, by using the Harrington principles, the Spinal Instability Neoplastic Score (SINS) (Table 6), Tomita type¹², and Tokuhashi score together, one can more confidently determine the best course of action for a given patient.

According to histologic subtypes, tumors can be classified into (1) slow growing (breast, prostate, carcinoid, thyroid), (2) moderately growing (kidney, uterus), and (3) rapidly growing (lung, liver, stomach, esophagus, pancreas, sarcomas, bladder) tumors¹³.

Table 3. Karnofsky performance score

Score	Description
100	Normal; no complaints; no evidence of disease
90	Able to carry on normal activity; minor signs or symptoms of disease
80	Normal activity with effort; some signs or synptoms of disease
70	Cares for self; unable to carry on normal activity or to do active work
60	Requires occasional assistance, but is able to
50	care for most of his personal needs Requires considerable assistance and frequent medical care
40	Disabled; requires special care and assistance
30	Severely disabled; hospital admission is
	indicated although death not imminent
20	Very sick; hospital admission necessary; active
	supportive treatment necessary
10	Moribund; fatal processes progressing rapidly
0	Dead

Table 4. Frankel classification of patients according to initial neurological function

A – Absence of motor or sensory function below
the level of the lesion
B – Absence of motor function, but with some
degree of sensitivity preserved below the level
of the lesion
C – Some degree of motor function but without
practical usefulness
D – Useful motor function below the level
of the lesion
E – Normal sensory and motor function, although
there may be some abnormality of reflexes

Table 5. Modified Bauer score

Score 1 point for each of the following positive
prognostic factors:
No visceral metastases
• No lung cancer
• Primary tumor * breast, kidney, lymphoma,
multiple myeloma
• One solitary skeletal metastasis
Suggestions for treatment
• Score 0–1: Supportive care, no surgery
• Score 2: Short term palliation, dorsal surgery
• Score 3–4: Middle term local control,
ventral-dorsal surgery

Table 6. The Spinal Instability Neoplastic Score (SINS): a
score of 0-6 is considered stable, 7-12 potentially unstable,
and 13–18 unstable

	1		
SINS	Description	Score	
component	onent		
Location	Junctional (occiput-C2,	3	
	C7-T2, T11-L1, L5-S1)		
	Mobile spine (C3-C6, L2-L4)	2	
	Semi-rigid (T3-T10)	1	
	Rigid (S2-S5)	0	
Pain	Yes	3	
	Occasional pain but not mechanical	1	
	Pain-free lesion	0	
Bone lesion	Lytic	2	
	Mixed (lytic/blastic)	1	
	Blastic	0	
Radiographic	Subluxation/translation	4	
spinal	Kyphosis/scoliosis	2	
alignment	Normal	0	
Vertebral body	50% collapse	3	
collapse	50% collapse	2	
	No collapse with 50% body	1	
	involved		
	None of the above	0	
Posterolateral	Bilateral	3	
involvement	Unilateral	1	
of spinal elements	None of the above	0	

Knowing the biological behavior of certain subtypes can give one a general sense of urgency required for the overall workup and treatment duration. In all cases, it is prudent to engage a team of spinal surgeons as soon as possible to optimize outcome when surgical intervention is feasible.

Surgical Techniques

Before the advent of surgical treatment for metastatic spine disease, the standard treatment options were radiotherapy and corticosteroid application. Surgical treatment for spinal metastases first received confirmation of efficacy by Patchell *et al.*, who were the first to demonstrate in a randomized prospective study

that patients that were surgically decompressed clearly showed better results than those undergoing only palliative radiotherapy without decompression¹⁴. Since then, various criteria have been developed for the selection of patients that would benefit most from surgical treatment^{9,13}. The degree of spinal cord compression has been better defined¹⁵, a framework has been established for considering spinal instability brought about by malignant disease¹⁶, and most recently, the required extent of treatment has been better defined¹⁷. Thus, spinal surgery has established its place within the complex, multidisciplinary and palliative approach to patients with spinal metastatic disease. The goals of surgical treatment of spinal metastases are local control of tumor size, pain reduction, maintaining motor function and sphincter control, prevention of deformity, and upgrading the quality of remaining lifetime.

However, before deciding on surgical treatment of spinal metastatic disease, it is imperative to know the histology of the specific tumor and the overall spread of the malignant disease, as these are the most important predictors of survival. Therefore, biopsy should be the first step to surgical treatment. Biopsy can be performed percutaneously by needle in local or general anesthesia and under control of an image intensifier or by computerized tomography (Fig. 1). If the result of such a biopsy should turn out negative for malignant cells and strong suspicion of malignancy remains, then open biopsy should be the next step.

Upon deciding on the need of surgical treatment, it may be necessary to perform preoperative embolization of the tumor in order to reduce intraoperative bleeding. This is especially important for histologic types which are prone to excess bleeding (for example, renal cell or thyroid cancer), as seen in Figure 2.

The spinal surgeon has an array of treatment options that differ in invasiveness and how radical the tumor resection should be¹⁷. The most radical operation is *en bloc* resection of a tumor according to the principles of treatment of primary spinal tumors¹². They are divided into 3 types of radical resections: vertebrectomy, sagittal resection of a vertebra, and total resection of posterior elements⁸. The indication for such a procedure would be a controllable histologic type of tumor, the presence of a metastasis that is technically resectable with a surrounding layer of healthy tissue, the lack of other distant metastases, and a generally good overall state of the patient (Fig. 3). The complex anatomy of the surrounding structures and the central placement of neural elements within the spinal canal pose a unique challenge to performing radical resections of spinal tumors. That is why it is usually only possible to perform a marginal or intralesional resection, or reduction of the tumor along

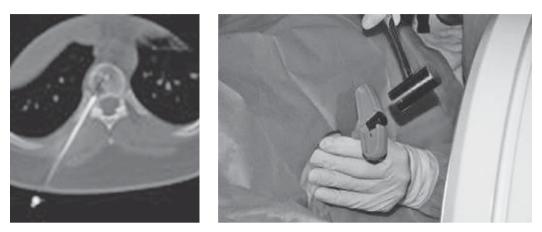


Fig. 1. Computerized tomography guided percutaneous transpedicular placement of a biopsy needle in a thoracic vertebral body.

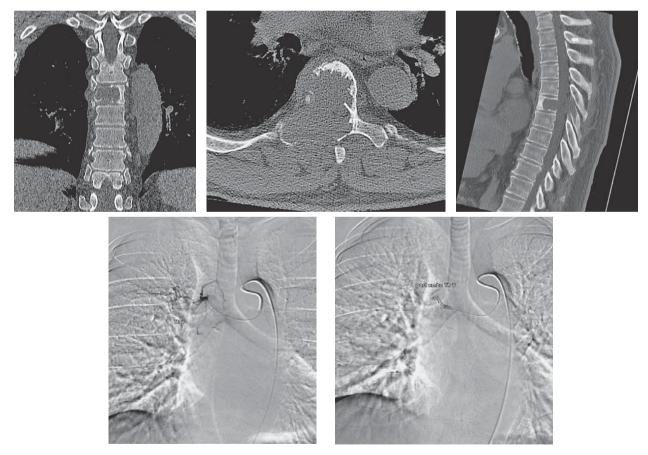


Fig. 2. Computerized tomography images in 3 planes of a highly vascularized thyroid cancer spinal metastasis, followed by pre- and post-embolization fluoroscopy images of the same case. Note the instillation of contrast medium in the pre-embolization image, followed by complete blockage of the feeding vessel in the post-embolization image.

with stabilization of the spinal column. These procedures are less radical than *en bloc* resections, although they may yield virtually identical results if stereotactic radiosurgery (which has significantly altered the treatment approach to patients with spinal metastases) is additionally performed¹⁹.

The goal of modern surgery in metastatic spine disease is to ensure sufficient distancing of the tumor from the spinal cord (separational surgery) and to provide stable fixation of the spinal column to allow for safe application of radiotherapy on the whole tumor volume.

In the case of spinal instability and great pain, which would hinder upright posture and/or ambulation in the patient, there are several ways in which stabilization can be achieved. Stabilization is indicated when there is a clear risk of instability according to the SINS score, re-



Fig. 3. Magnetic resonance images of a solitary Th12 metastasis of a primary breast cancer, confined to the vertebral body with slight bulging into the epidural space with immediate postoperative plain film radiographs after en bloc vertebrectomy of the above Th12 metastasis with anterior support with a graft-filled mesh cage and posterior transpedicular fixation.

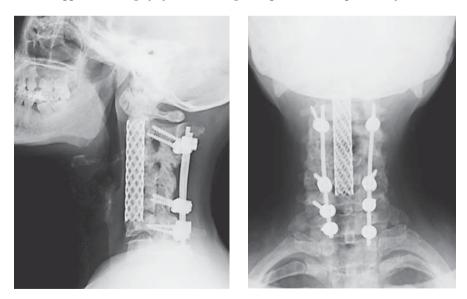


Fig. 4. Plain film radiographic demonstration of typical examples of combined anterior and posterior stabilization using interbody cages filled with graft material anteriorly and lateral mass screws and rods, as in the above cervical segments.

gardless of histologic tumor type and overall predicted survival, provided that the general state of the patient allows for surgery to be performed, some examples of which can be seen in Figure 4¹⁴. Stabilization methods include open anterior and posterior stabilization, percutaneous stabilization methods, and vertebroplasty or kyphoplasty²⁰. Minimally invasive methods for stabilization of the vertebral body are particularly convenient as palliative techniques for patients who otherwise would not be candidates for radical surgery, such as patients with multilevel involvement (Fig. 5)²¹. Percutaneous stabilization of the spine is indicated in patients in whom there is clear instability without significant spinal cord compression (Fig. 6)²².

In the case that the expected remaining lifespan of the patient is under 3 months and there are multiple metastatic changes along the spinal column and unbearable pain that is unresponsive to high doses of opioid analgesics, it is recommended to install an intrathecal opioid pump to be able to control pain with much lower doses of opioids and fewer complications (Fig. 7).

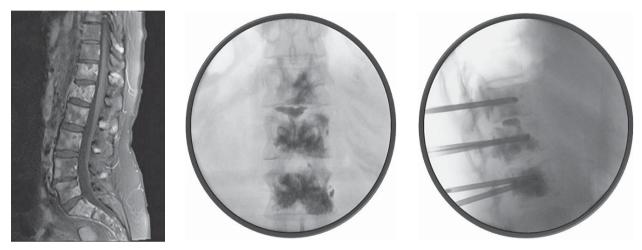


Fig. 5. Magnetic resonance image of multilevel metastatic involvement of the spinal column with corresponding intraoperative fluoroscopic images of percutaneous vertebral body augmentation (vertebroplasty) in the same patient.

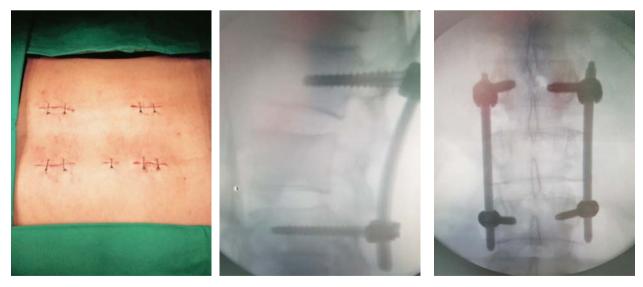


Fig. 6. Postoperative view of the skin incisions after minimally invasive (percutaneous) fixation of the lumbar spine with corresponding intraoperative fluoroscopic sagittal and coronal views.



Fig. 7. Placement of an intrathecal pump device for administration of opioids.

Conclusion

Patients with spine metastases frequently present with complex diagnostic and therapeutic challenges requiring an integrated multi-step approach and multidisciplinary care, including a team of surgeons and oncologists. In managing these patients, every effort should be made to provide an early diagnosis and appropriate treatment for preservation of neurological function and quality of remaining life. Unfortunately, diagnosis is frequently not established until significant neurological deficit is present, by which time functional recovery may be unlikely. Newer surgical techniques that address decompression, spinal stabilization, with or without tumor resection, and minimally invasive pain relieving strategies must be taken into consideration in surgical candidates. Studies have shown clear benefit of surgical management in terms of maintaining ambulation and sphincter control, as well as in providing pain relief and prolonging patient independence. Although surgical intervention will not decrease mortality, it should nonetheless be strongly considered in patients that fulfill the established criteria for a reasonable outcome.

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Sažetak

KIRURŠKO LIJEČENJE METASTAZA KRALJEŽNICE

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Metastatska bolest kralježnice je dosta učestala i zasigurno zaslužuje pozornost i liječenje zbog visoke vjerojatnosti da bi inače bolesnici s karcinomom ostali oduzeti u svojim posljednjim mjesecima života. Novija postignuća i shvaćanja ponašanja raznih tipova tumora te unaprjeđenja u kirurškim tehnikama omogućuju evoluciju algoritama, pogotovo kada se kirurško liječenje razmatra. Ovaj rad daje pregled postupka donošenja odluka i ističe razne kirurške mogućnosti koje su trenutno dostupne.

Ključne riječi: Kralježnica – kirurgija; Kralježnica – patologija; Kralježnica, tumori, sekundarni; Metastaze – kirurgija; Palijativna skrb – metode; Algoritmi; Odlučivanje, tehnike potpore