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Case Report

Small Bowel Intussusception Secondary to Metastatic Spindle Cell Sarcoma: A Case Report and Review of the Literature

Tina W Wong^{1*}, Debra A Wong², Hashem Ayyad³, David Row¹, Ronald A Gagliano¹, and James Mankin¹

¹Department of Surgery, Creighton University School of Medicine Phoenix, USA

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Abstract

Background: Intussusception is an uncommon diagnosis in adults. It is characterized by an abnormal luminal "lead point" that precipitates bowel invagination, resulting in obstructive symptoms that warrants further investigation. Both benign and malignant tumors have been reported to cause small bowel intussusception, with primary small bowel malignancies occurring more commonly. We present an unusual case of a middle-aged gentleman with small bowel intussusception and obstruction caused by a metastatic spindle cell sarcoma and a review of the available literature.

Case Presentation: A 50-year-old male patient with a history of recurrent spindle cell sarcoma of the scalp treated 3 years prior presented with abdominal pain and bloating. Diagnostic workup revealed two segments of small bowel intussusception and obstruction as well as several lung nodules. The patient underwent exploratory laparotomy for resection and primary anastomosis of the intussuscepted segments of small bowel. Histologic analysis of the surgical specimens confirmed metastatic spindle cell sarcoma. After completion of full metastatic workup, patient was counseled on the diagnosis and offered palliative chemotherapy while awaiting results from tumor molecular profiling to guide future treatment and clinical trial options.

Conclusion: Intussusception in the adult patient should be carefully evaluated, with a high suspicion for malignancy as the underlying cause, especially in those with a personal history of cancer.

Keywords: Intussusception; Sarcoma; Spindle Cell

List of Abbreviations: CT: Computed Tomography; MRI: Magnetic Resonance Imaging; PET: Positron Emission Tomography; FDG: Fluorodeoxyglucose; NET: Neuroendocrine Tumor; GIST: Gastrointestinal Stromal Tumor; UPS: Undifferentiated Pleomorphic Sarcoma; RMS: Rhabdomyosarcoma; LS: Liposarcoma; AS: Angiosarcoma; ES: Ewing Sarcoma; FS: Fibrosarcoma; OS: Osteosarcoma; CS: Chondrosarcoma; IDCS: Interdigitating Dendritic Cell Sarcoma; ASPS: Alveolar Soft Part Sarcoma; LMS: Leiomyosarcoma; MFH: Malignant Fibrous Histiocytoma; LCS: Lung Carcinosarcoma; MS: Melanosarcoma; KS: Kaposi Sarcoma; RCS: Round-Celled Sarcoma; SCS: Spindle Cell Sarcoma; H&E: Hematoxylin And Eosin

Introduction

Intussusception of the small or large intestine is defined by the telescoping of a segment of bowel on itself, with the "donor" invaginated segment termed "intussusceptum" and the "recipient" segment termed "intussuscipien" [1]. The majority of intussusception is associated with a lead point from which the telescoping begins, such as enlarged mesenteric lymph nodes or tumor. While intussusception is a common phenomenon in the pediatric population and most often of benign etiology, its occurrence in the adult may be the first sign of malignancy and warrants further investigation [1-3]. Here we describe a

²Department of Oncology, University of Arizona Cancer Center, USA

³Department of Pathology, St. Joseph's Hospital and Medical Center, USA

^{*}Corresponding author: Tina W Wong, Department of Surgery, Creighton University School of Medicine Phoenix, Maricopa Medical Center, 2601 E Roosevelt Street, Phoenix, AZ 85008, USA

case of a recurrent spindle cell sarcoma with metastases to the gastrointestinal tract, resulting in jejunal intussusception 3 years after initial treatment of the primary lesion.

Case Presentation

A 50-year-old man presented to the Emergency Department with acute-onset severe left-sided abdominal pain and associated bloating. On examination, his abdomen was moderately distended and tender to palpation in the setting of normal vital signs. The patient's past medical history was significant for spindle cell sarcoma of the frontal scalp diagnosed and excised three years prior. Pathology of the excised scalp mass was consistent with high-grade spindle cell sarcoma with a high mitotic rate and tumor extension to <1mm from the surgical margin. The patient thus underwent re-resection of the close margin followed by adjuvant radiation

therapy. Two years later, the patient was noted to have a progressive growth around the surgical scar. Excisional biopsy confirmed recurrent spindle cell sarcoma with positive margins. Subsequent re-excision was performed without further adjuvant treatment. In the Emergency Department, a complete blood count and metabolic panel were normal. A computed tomography (CT) of the chest, abdomen, and pelvis with contrast revealed two segments of jejunal intussusception resulting in small bowel obstruction, in addition to several lung nodules concerning for metastatic disease (Figure 1). Surgical consultation was obtained, and the patient was taken for exploratory laparotomy and small bowel resection with primary anastomoses of the two separate segments of intussuscepted small bowel. The patient's hospital course was uncomplicated, and he was discharged home on postoperative day 3.

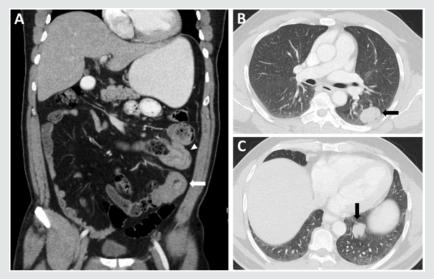


Figure 1: Computed tomography of the chest, abdomen, and pelvis with contrast demonstrates small bowel intussusception and lung nodules. (A) Two segments of jejunum with radiographic finding of invagination (white arrowhead) and target sign (white arrow), indicative of intussusception. Distended stomach and moderately dilated small bowel are seen proximal to the intussusception without passage of oral contrast through this region, consistent with small bowel obstruction. (B-C) Solid pulmonary nodules with irregular borders (black arrows) within the left chest, concerning for metastatic disease.

At his 2-week follow-up in surgery clinic, the patient was found to be recovering well. By that time, surgical pathology was finalized, after thorough analysis and expert consultation. The resected proximal small bowel revealed a 4.5 cm neoplasm and a second neoplasm also measuring 4.5 cm was seen in the distal small bowel. Both tumors were composed of moderately atypical spindle cells arranged in intersecting fascicles, with focally myxoid and collagenous stroma (Figure 2). Mitotic activity was greater than 30/HPF. Immunohistochemistry was positive for CD34 and CD99, and negative for pancytokeratin, S100, CD117, DOG1, desmin, SOX10, calretinin, BCL-2, HMB45, smooth muscle actin, STAT6, ALK-1, and ERG. MDM2 was positive only in scattered nuclei. Ki-67 proliferation index was 15-20 percent. Collectively, these findings were consistent with recurrent metastatic high-grade

(FNCLCC grade 3) malignant spindle cell sarcoma. Consultation with medical oncology was obtained. The patient underwent staging imaging with a brain magnetic resonance imaging (MRI) with and without gadolinium, which showed no locally recurrent or intracranial disease. Whole body positron emission tomography (PET)-CT revealed several fluorodeoxyglucose (FDG)-avid lesions in the left lung and chest wall as well as multiple periportal and pericaval enlarged lymph nodes (Figure 3). CT-guided biopsy of an intrathoracic lesion confirmed metastatic spindle cell sarcoma (Figure 4). Next-generation sequencing tumor molecular profiling is being pursued to help guide future treatment strategies and clinical trial options. The patient, who maintains an excellent performance status, has been counseled on palliative doxorubicin and ifosfamide-based chemotherapy, which he is eager to commence.

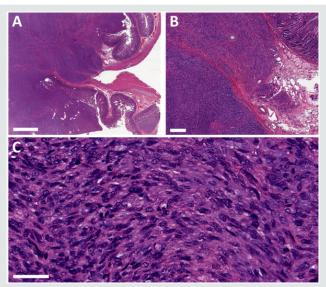


Figure 2: Histologic examination of surgical specimen reveals metastatic spindle cell sarcoma to the small bowel. Formalin-fixed paraffin-embedded tissue was sectioned and stained with hematoxylin and eosin (H&E), demonstrating atypical spindle cells with focally myxoid and collagenous stroma. (A) Low magnification, scale bar = 3 mm. (B) Medium magnification, scale bar = $600 \mu m$.

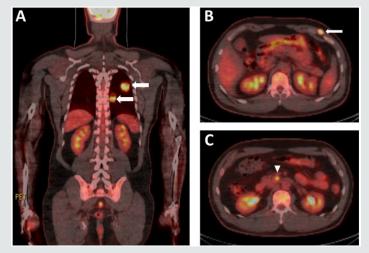


Figure 3: Whole body PET-CT reveals FDG-avid lesions in the left lung (A; thick white arrows) and left chest wall (B; thin white arrow) as well as FDG-avid enlarged pericaval lymph node (C; white arrowhead), suggestive of metastatic disease.

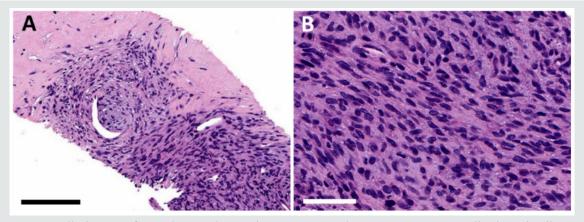


Figure 4: Core needle biopsy of intrathoracic lesion demonstrates malignant appearing spindle-shaped cells, confirming metastatic spindle cell sarcoma. Formalin-fixed paraffin-embedded tissue was sectioned and stained with H&E. (A) Medium magnification, scale bar = $150 \, \mu m$. (B) High magnification, scale bar = $50 \, \mu m$.

Discussion

Intussusception is an uncommon diagnosis in adults. Although physiologic intussusception can occur with hyperdynamic peristalsis of the intestines, such as in the setting of gastroenteritis, when combined with obstructive symptoms, intussusception in an adult patient suggests an ominous etiology, and a pathologic lead point is associated 70-90% of the time [1,2]. Management of intussusception in adults differs from that in the pediatric population. Specifically, in adults, surgical resection of the involved bowel is recommended rather than simple reduction (either via pneumatic/hydrostatic enema or operatively) [3]. Clinical suspicion for a neoplastic process was high in our patient, given his history of soft tissue sarcoma of the scalp and the concurrent finding of lung nodules on his imaging. Our patient thus underwent laparotomy and multi-segment small bowel resection, and pathologic analysis of the surgical specimen confirmed recurrent metastatic spindle cell sarcoma. Spindle cell sarcoma is a subset of soft tissue sarcoma that encompasses a myriad of histologic subtypes. While all subtypes are of mesenchymal origin, more than 50 subtypes of soft tissue sarcomas exist, each with variable degree of differentiation, recurrence rates, and metastatic potential. In general, sarcomas can be classified based on the cell-lineage they resemble (e.g., smooth muscle, fibroblast, adipocyte, bone, etc.). Not uncommonly, however, the irregularity and degree of dedifferentiation of the tumor cells make determination of cell-lineage impossible; these tumors are thus named based on their morphologic appearance [4]. Spindle cell sarcoma, for example, is characterized by its long, narrow, spindle-like appearance microscopically.

Soft tissue sarcomas, including spindle cell sarcomas, are typically diagnosed at an early stage, with less than 15% with metastatic disease at presentation [5]. However, given the aggressive nature of these tumors, approximately one-third of patients with localized soft tissue sarcoma will develop metastatic disease within 5 years, despite adequate initial treatment. Overall survival in patients with soft tissue sarcoma correlates with the

stage of the disease, with an approximately 60% 5-year survival rate for localized disease but only 15% for those with distant metastases [6]. Lung is the primary site for distant metastases, but other common sites of metastases include bone, liver, brain, and lymph nodes [5]. The small intestine is an extremely rare location for sarcoma metastasis. Review of the literature reveals only 46 case reports of various subtypes of bone or soft tissue sarcoma with metastases to the small bowel. Of these, 30 patients presented with small bowel intussusception (with or without obstruction) secondary to the metastatic lesion [7-36]. Other clinical findings that led to the diagnosis of metastatic sarcoma have included obstruction without intussusception (n=6) [37-42] gastrointestinal bleeding (n=10) [11-46], and bowel perforation (n=6) [37-51]. In one report, metastatic sarcoma to small bowel was found incidentally on imaging in an asymptomatic patient [52]. The most common types of sarcoma reported to have metastasized to the small bowel include osteosarcoma (n=12), angiosarcoma (n=5), rhabdomyosarcoma (n=5), liposarcoma (n=4), and leiomyosarcoma (n=4) (Table 1). Overall, neoplasms of the small bowel are rare, with the majority being benign lesions (e.g. lipoma, leiomyoma, fibroma, hamartoma, or ectopic gastric or pancreatic tissue). Of all the cancers of the gastrointestinal tract, those found within the small bowel account for only 1-3% [53,54]. Malignancies of the small bowel are most often primary (e.g. adenocarcinoma, gastrointestinal stromal tumors (GIST), neuroendocrine tumor (NET), or non-Hodgkin's lymphoma) [55]; however, metastases from extra-abdominal sites have also been reported. Melanoma represents the most common metastatic tumor to the small bowel, accounting for over 50% [53]. Other cancers with a propensity for small bowel metastases include invasive lobular carcinoma of the breast, adenocarcinoma of the lung, and clear cell carcinoma of the kidney. As demonstrated in our case, metastases from a primary cutaneous spindle cell sarcoma to the small bowel can occur and cause intussusception, but our literature review shows that it is an exceedingly rare problem.

<u>Table 1</u>: Review of the literature – case reports of metastatic bone or soft tissue sarcoma to the small bowel.

	UPS	RMS	LS	AS	ES	FS	os	CS	IDCS	ASPS	LMS	MFH	LCS	MS	KS	RCS	SCS
Abbo et al. [13]							Х										
Avagnina et al. [41]		х															
Benis et al. [38]				Х													
Bustinza-Linares et al. [14]								X									
Chandramohan et al. [22]							x										
Chiang et al. [44]											Х						
Costa et al. [46]							Х										
Eng et al. [30]													X				
Esfehani et al. [11]						х											
Fleetwood et al. [43]				Х													
Ganguli et al. [26]							Х										

Gerst et al. [29]											Х					
Goldberg et al. [35]							х									
Gorman et al. [28]												Х				
Gys et al. [9]		х														
Horiuchi et al. [19]							Х									
Hsu et al. [45]				X												
Hung et al. [24]							Х									
Ise et al. [49]		Х														
Kamo et al. [20]				X												
Kanoh et al. [27]												Х				
Kehoe et al. [34]								Х								
Kuwabara et al. [12]							Х									
Lee et al. [15]										Х						
Mahamid et al. [47]									х							
Michael [42]													X			
Mitchell et al. [50]														х		
Monjazeb et al. [21]			х													
Mozes et al. [32]							х									
Najawa et al.			х													
Okada et al. [25]											Х					
Oldfield et al. 36						х										
Panizo-Santos et al. [39]							х									
Plestina et al. [37]	х															
Rowe et al. [52]			х													
Ruffolo et al. [48]				X												
Sabel et al. [18]										Х						
Shibata et al. [17]			х													
Sondenaa et al. [31]											Х					
Sun et al. [7]		х														
Tan et al. [10]					х											
Waring [51]															х	
Webster et al. [33]							х									
Weiss et al. [16]																х
Wootton-Gorges et al. [23]							х									
Xi et al. [8]		x														

UPS: Undifferentiated Pleomorphic Sarcoma; RMS: Rhabdomyosarcoma; LS: Liposarcoma; AS: Angiosarcoma; ES: Ewing Sarcoma; FS: Fibrosarcoma; OS: Osteosarcoma; CS: Chondrosarcoma; IDCS: Interdigitating Dendritic Cell Sarcoma; ASPS: Alveolar Soft Part Sarcoma; LMS: Leiomyosarcoma; MFH: Malignant Fibrous Histiocytoma; LCS: Lung Carcinosarcoma; MS: Melanosarcoma; KS: Kaposi Sarcoma; RCS: Round-Celled Sarcoma; SCS: Spindle Cell Sarcoma

Conclusion

This is an unusual case of an adult patient with metastatic spindle cell sarcoma to the small bowel who presented with intussusception and bowel obstruction. The case highlights the importance of careful evaluation and workup of intussusception in the adult patient population. The clinician must have a high index of suspicion for underlying malignancy, and the appropriate surgical management should be undertaken with this in mind. Close attention should be paid to patients with a personal history or extensive family history of malignancy. As illustrated in this case,

expert pathologic analysis is integral to establishing an accurate diagnosis and prognosis. Early involvement of the appropriate specialists is recommended whenever cancer is suspected, to help achieve the best possible outcomes for patients via a multidisciplinary approach.

Acknowledgement

Not applicable.

Conflict of Interest

The authors declare that they have no competing interests.

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