Case presentation

A 52-year-old African American man, with a history of sickle cell trait, alpha-thalassemia, gynecomastia and nephrolithiasis presented to the emergency department of an urban, tertiary-care medical center with worsening back pain. The patient stated that 3 weeks prior he experienced minor trauma while driving a bus and has since had severe pain in his lumbar region that was unresponsive to over-the-counter analgesics. The patient denied further symptoms, including weight loss, fever, night sweats, abdominal pain, dyspepsia, nausea, vomiting, hematochezia, and melena. Of note, the patient had a family history significant for a first-degree relative with multiple myeloma (MM) and another with breast cancer. On physical exam, the patient was tender to palpation in the lower thoracic and lumbar spine. The patient had no saddle anesthesia or focal neurologic deficits, and the abdominal exam was benign. Initial laboratory evaluation revealed a white blood cell (WBC) count of 13.1 K/UL, a hemoglobin of 10.2 G/DL with an MCV of 65, and a platelet count of 184 K/UL. The creatinine was elevated to 1.47 mg/DL and the serum calcium was 10.8 mg/DL. Liver function tests (LFT) were normal. Computed tomography imaging of the chest, abdomen, and pelvis revealed a compression fracture of the L1 vertebral body as well as lytic lucencies in the T5 and T9 vertebral bodies and a prominent lymph node in the right upper quadrant, and a 1.8 cm hypoattenuating lesion in the left lobe of the thyroid gland.

The patient was admitted to the medical ward for further evaluation of the lytic lesions, as well as pain management. The presenting labwork, lytic lesions, and family history prompted an initial evaluation for MM. A serum protein

Bone marrow infiltration as the initial presentation of gastric signet ring cell adenocarcinoma

Christopher Dittus¹, Hannah Mathew², Anita Malek¹, Andreea Negroiu¹

¹Hematology-Oncology Department, Boston Medical Center, 820 Harrison Avenue, FGH Building, 1st Floor, Boston, MA 02118, USA; ²Internal Medicine Department, Boston Medical Center, 72 East Concord Street, Evans 124 Boston, MA 02118, USA; ³Pathology Department, Boston Medical Center, 670 Albany Street, Boston, MA 02118, USA

Correspondence to: Christopher Dittus, D.O., M.P.H. Boston Medical Center, 820 Harrison Avenue, FGH Building, 1st Floor, Boston, MA 02118, USA. Email: Christopher.dittus@bmc.org.

Abstract: This case report describes a 52-year-old African American man who initially presented with worsening back pain. The patient was found to have lytic lucencies in the T5 and T9 vertebral bodies and a subsequent bone marrow biopsy revealed an extensive infiltrate of signet ring cells. These findings prompted a workup for a gastrointestinal malignancy, and upper endoscopy revealed a mass in the gastric pylorus. A biopsy of this mass was positive for signet ring cell adenocarcinoma. This case is significant for two reasons. First, it highlights the importance of a broad differential diagnosis when approaching a patient with lytic bone lesions. Second, bone marrow involvement is more common in patients with diffuse type gastric cancer and occurs in particularly young patients. The increasing incidence of diffuse type gastric adenocarcinoma means bone marrow metastases will likely play a greater role in the presentation and management of gastric cancer.

Keywords: Bone marrow infiltration; gastric cancer; signet ring cell; bone marrow metastasis; lytic bone metastasis; stomach cancer; diffuse type

Submitted Jul 08, 2014. Accepted for publication Jul 14, 2014.
doi: 10.3978/j.issn.2078-6891.2014.050
View this article at: http://dx.doi.org/10.3978/j.issn.2078-6891.2014.050
electrophoresis, serum immunofixation, and serum free light chain assay returned positive for a monoclonal IgG kappa paraprotein. A subsequent bone marrow evaluation did not yield an aspirate (“dry tap”), but a biopsy specimen was obtained. The pathologic review of this specimen was significant for marked hypercellularity (90%), which was due to an extensive infiltrate of signet ring cells accounting for 90% of the hypercellularity (Figure 2). Myeloid and erythroid elements were markedly decreased and megakaryocytes were nearly absent. A mild plasmacytosis was present with small pockets of mature plasma cells. Immunohistochemistry staining of the signet ring cells was positive for PAS, AE1/AE3, Cam5.2 (Figure 3A), CD138, and cyclin D1. Staining with CDx2 was positive, indicating a gastrointestinal origin for the signet ring cells (Figure 3B). Subsequent staining with CK7 was positive (Figure 3C),

Figure 1 Sagittal view of lytic spine lesions in T5 and T9 vertebrae (A), and axial view of lytic spine lesion at T9 vertebrae (B).

Figure 2 Bone marrow biopsy under 10x magnification (A) and 20x magnification (B) showing abundance of signet ring cells.

Figure 3 Immunohistochemistry on bone marrow biopsy showing positive staining of signet ring cells for CAM5.2 (A), CDx2 (B), and CK7 (C) (10x magnification).
Gastric cancer has been broadly divided into intestinal-type and diffuse-type since this was initially described almost 50 years ago (4). The histologic shift from intestinal type gastric cancer to diffuse type will have important clinical implications. Diffuse type gastric cancer occurs at a younger age, and is more advanced at presentation, particularly when compared to well or moderately differentiated intestinal type gastric cancer (5). The traditional risk factors for stomach cancer do not play as great a role in diffuse type gastric cancer, and studies have described the importance of familial syndromes such as hereditary diffuse gastric cancer (6). This is of particular importance to our case because our patient had the signet ring cell type of gastric adenocarcinoma, which is a type of diffuse gastric cancer. Our patient presents with younger age, which is a hallmark of the diffuse type, as was our patient’s advanced disease at diagnosis. The unique characteristic of our case is that our patient presented with only back pain, denying all gastrointestinal complaints. It was through the evaluation of our patient’s back pain that the patient was ultimately diagnosed with signet ring cell gastric adenocarcinoma.

The occurrence of bone marrow metastasis has been documented in gastric cancer, although this is rarely the initial presentation. If bone marrow involvement is discovered, it is usually during the workup for metastatic disease. Interestingly, our patient only had mild microcytic anemia, which was likely preceding his gastric cancer and related to his alpha-thalassemia. Our patient did not have leukopenia or thrombocytopenia, the latter of which was determined to be indicative of bone marrow involvement in at least one study (7). When bone marrow metastasis occurs, it is more commonly a signet ring cell subtype of gastric carcinoma and occurs in younger patients (8). The prognosis for bone marrow involvement with gastric adenocarcinoma is abysmal, with patient’s living an average of 44 days from the time of documented bone marrow involvement (7).

This case is significant for two reasons. First, it highlights the importance of a broad differential diagnosis when approaching a patient with lytic bone lesions. Second, bone marrow involvement is more common in patients with diffuse type gastric adenocarcinoma and occurs in particularly young patients. The increasing incidence of diffuse type gastric adenocarcinoma means bone marrow metastases will likely play a greater role in the presentation and management of gastric cancer.

Acknowledgements

Disclosure: The authors declare no conflict of interest.

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Cite this article as: Dittus C, Mathew H, Malek A, Negroiu A. Bone marrow infiltration as the initial presentation of gastric signet ring cell adenocarcinoma. J Gastrointest Oncol 2014;5(6):E113-E116. doi: 10.3978/j.issn.2078-6891.2014.050


