Synchronous Bone Metastasis From Multiple Myeloma and Prostate Adenocarcinoma as Initial Presentation of Coexistent Malignancies

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The radiographic appearance of bone metastases is usually determined by tumor histology and can be osteolytic, osteoblastic, or mixed. We present a patient with coexistent bone metastasis from multiple myeloma and prostate adenocarcinoma who exhibited synchronous bone involvement of both histologies within the same bone lesion, a rare phenomenon that has not been previously reported and led to atypical radiographic findings. The radiograph of a 71-year-old man with thigh swelling and pain demonstrated a lytic femoral lesion. Magnetic resonance imaging (MRI) confirmed a destructive process, but showed coexistent metaphyseal sclerosis. Multiple myeloma was suspected by demonstration of monoclonal gammopathy and confirmed by computed tomography (CT)-guided biopsy. Incidentally, CT demonstrated areas of sclerosis corresponding to T2 hypointensity on MRI. Further studies revealed osteoblastic spinal metastasis, prostate enhancement on CT and prostate-specific antigen (PSA) level of 90 ng/mL, concerning for concomitant prostate neoplasm. After endoprosthetic reconstruction, pathology of the femur identified both plasma cell neoplasm and metastatic prostate adenocarcinoma. An association between prostate cancer and multiple myeloma is hypothesized due to tumor microenvironment similarities and possible common genetic variations, however, coexisting bone metastases have never been reported. This unusual finding explains the discrepant imaging features in our patient and is evidenced that certain clinical situations merit contemplation of atypical presentations of common malignancies even if this leads to additional testing.

Keywords: bone metastasis, radiographic imaging, synchronous malignancies, multiple myeloma, prostate cancer

INTRODUCTION

Metastases are the most common type of malignancy affecting the bones (1). Bone metastases are classified as osteoblastic, osteolytic, or mixed depending on their radiographic appearance, which in turn is frequently determined by the underlying tumor histology. Osteolytic lesions are radiographically lucent and are characterized by osteoclast-mediated bone destruction. These
lesions are commonly present in multiple myeloma, renal cell carcinoma, thyroid cancer, and melanoma. Osteoblastic lesions are radiographically dense and sclerotic, and are typically present in prostate adenocarcinoma, small cell lung cancer, Hodgkin’s lymphoma, and carcinoid tumors. Mixed lesions are both osteolytic and osteoblastic and are typical of gastrointestinal tumors or squamous cell carcinomas (2).

Coexistent malignancies are usually associated with common genetic, infectious, environmental, or occupational factors (3), but some synchronous malignancies may be coincidental. A potential association between prostate adenocarcinoma and multiple myeloma has been proposed but not confirmed (4, 5), with only a few reported cases. We present a patient with atypical radiographic findings caused by synchronous occurrence of multiple myeloma and prostate adenocarcinoma metastasis within the same bone lesion. To our knowledge, this is the first case reported of this unusual phenomenon which in retrospect highlights the utility of proper radiographic imaging assessment and interpretation for cancer patients.

BACKGROUND

A 71-year-old man was evaluated in orthopedics clinic for progressive left thigh swelling and pain leading to impaired ambulation for 3 months. Radiographs demonstrated a large lytic lesion of the distal femur with cortical destruction (Figure 1). Urgent evaluation with magnetic resonance imaging (MRI) confirmed a 10 cm destructive distal metadiaphyseal lesion in the left femoral diaphysis as well as coexistent central areas of T2 hypointensity suggestive of metaphyseal sclerosis which were not appreciated radiographically (Figure 2). Multiple myeloma was clinically suspected on the basis of normocytic anemia, increased serum creatinine, and elevated globulin levels. Serum protein electrophoresis showed an IgA-kappa monoclonal M-spike and percutaneous computed tomography (CT)-guided biopsy of the femoral lesion was consistent with plasma cell neoplasm affecting the bone.

Review of the axial CT images of the femur obtained to localize the lesion for biopsy demonstrated a predominantly lytic lesion and also small areas of sclerosis (Figure 3A) corresponding to the areas of T2 hypointensity on MRI (Figure 2B). Given the atypical MRI findings of sclerotic appearance on T2, a CT of the chest, abdomen, and pelvis was performed and demonstrated diffuse osteoblastic metastatic lesions, including multiple sclerotic lesions throughout the spine (Figure 3B). Pelvic images also demonstrated asymmetric enhancement of the left anterior prostate and prominent retroperitoneal and pelvic lymph nodes, which were concerning for concomitant metastatic prostate neoplasm. Serum prostate-specific antigen (PSA) level was 90 ng/mL.

The patient finally underwent distal femur resection with endoprosthetic reconstruction. Pathology examination of the distal femur identified both plasma cell neoplasm and metastatic

**FIGURE 1** | Frontal radiograph of the left femur demonstrates a large lytic lesion of the distal diaphysis with wide zone of transition and aggressive-appearing periosteal reaction at the medial margin.

**FIGURE 2** | (A) Coronal T1 magnetic resonance imaging (MRI) image of the left femur demonstrates T1 hypointense lesions (starts) with normal intervening bone marrow (arrow). (B) Sagittal T1 fat-saturated MRI image of the dominant distal femur lesion demonstrated intermediate T1 hyperintensity with areas of marked low signal intensity centrally (arrows), suggestive of areas of sclerosis not appreciated radiographically.
FIGURE 3 | (A) Axial computed tomography (CT) image of the femur obtained to localize the lesion for CT-guided biopsy demonstrates a destructive lytic lesion with cortical breakthrough anteriorly (arrow) and areas of bone sclerosis (arrowheads). (B) Sagittal image of a CT of the chest, abdomen, and pelvis demonstrates multiple sclerotic lesions in the spine, including within T6, T8, T10, L2, and L3 vertebrae (arrows).

FIGURE 4 | (A) Excised left femur with hemorrhagic tumor involving the diaphysis (gross specimen), (B) prostatic specific membrane antigen expression in prostate adenocarcinoma (left) and negative in multiple myeloma (right), and (C) coexisting metastatic prostate adenocarcinoma and multiple myeloma in same histologic section; note the gland forming adenocarcinoma (left) and sheets of plasma cell infiltration (right).

The incidence of simultaneous diagnosis of prostate cancer and lymphoid malignancies is reported to be approximately 1.2% (6), but synchronous occurrence of prostate adenocarcinoma and multiple myeloma is only reported in a few cases (3, 4, 6). Coexistence of plasma cell neoplasm and metastatic prostate carcinoma in the bone marrow has been described (6, 7), but synchronous skeletal metastases from these malignancies affecting adenocarcinoma of prostate origin within the same metastatic lesion (Figure 4). The final diagnoses were stage IV hormone-naïve prostate adenocarcinoma and ISS stage III IgA-kappa multiple myeloma. Treatment was started with bicalutamide and leuprolide for androgen deprivation therapy (ADT), which achieved PSA suppression after 3 months. Myeloma was simultaneously treated with five 28-day cycles of cyclophosphamide, bortezomib, and dexamethasone followed by autologous hematopoietic stem cell transplantation with melphalan conditioning, with the patient achieving a very good partial response (VGPR). Currently, at 2 years of follow-up, he remains on leuprolide for ADT with continued suppression of PSA and is on maintenance lenalidomide with persistent VGPR from myeloma. No bone-specific events have occurred and his bone pain is well controlled.

DISCUSSION

The incidence of simultaneous diagnosis of prostate cancer and lymphoid malignancies is reported to be approximately 1.2% (6), but synchronous occurrence of prostate adenocarcinoma and multiple myeloma is only reported in a few cases (3, 4, 6). Coexistence of plasma cell neoplasm and metastatic prostate carcinoma in the bone marrow has been described (6, 7), but synchronous skeletal metastases from these malignancies affecting...
synchronously, despite initial pathological confirmation of one of them. After surgical excision, the unusual finding of prostate adenocarcinoma and multiple myeloma within the same bone specimen explained the discrepant imaging features of the distal femur lesion, having features of both osteolytic and osteoblastic (marked hypointensity on T1, and areas of sclerosis on CT) lesion.

**CONCLUDING REMARKS**

Understanding and distinguishing the characteristic appearance of multiple myeloma and prostate cancer skeletal metastasis guided appropriate diagnostic testing and ultimately allowed for proper treatment in this patient who continues to respond well to cancer therapy to this day. Although cost-conscious avoidance of excessive testing is usually the preferred approach in oncological clinical practice, specific clinical situations, such as discordant imaging and atypical presentations of common malignancies, warrant consideration of unusual presentations, even if it requires additional testing, and especially, when it may impact treatment.

**ETHICS STATEMENT**

This work was exempted from ethics approval by Montefiore Einstein Center for Bioethics because it constitutes a purely descriptive case report. The report does not involve any risk to the participating patient, including no foreseeable risk, harm, or discomfort. Written informed consent was granted by the patient reported in this work for the publication of this case report and radiographic images.

**AUTHOR CONTRIBUTIONS**

DH and BG conceived the presentation of this case and summarized the findings. JSh reviewed all clinical information regarding multiple myeloma. AS reviewed all clinical information regarding prostate cancer. SG-S performed the review and description of radiographic studies. JSa verified and described the pathological findings. All authors discussed the results and contributed to the final manuscript.

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Conflict of Interest Statement: The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.