CASE REPORT

Heterogeneous osteoblastic activity in the right ischium of unclear etiology seen on NaF18-PET/CT

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Abstract
We present the case of a 54 year old white male with a history of lumbar spondylolisthesis and hip arthralgia, which prompted a hip X-ray. The X-ray showed hip arthritis as well as a sclerotic lesion in the right femur concerning for possible metastasis. He was then referred for a NaF18-PET/CT bone scan to exclude metastatic bone disease. The NaF18-PET/CT study did not show evidence of osteoblastic activity in the sclerotic right femoral lesion, but showed an incidental finding of an ill-defined intense and heterogeneous osteoblastic activity affecting the entire right ischium concerning for malignancy due to the level of activity and heterogeneity. This lesion was followed up with MRI imaging. MRI imaging has been stable and the final diagnosis is favored to be chronic avulsion injury. If a case such as ours is encountered in everyday practice, the possibility of a benign traumatic etiology should be given high consideration.

Key words
Unclear etiology, NaF18-PET/CT, Bone scan, Osteoblastic lesion

1 Introduction
We present a case of single ill-defined area of heterogeneous high osteoblastic activity in the right ischium associated with slight cortical thinning of unclear etiology seen on NaF18-PET/CT bone scan in a patient with hip pain. NaF18-PET/CT bone scan is more sensitive than Tc99m bone scan in detecting benign and malignant bone lesions [1]. This type of bone scan can produce high quality images of the skeleton due to the fact that 18F has high avidity for bone which results in rapid bone uptake, creating high bone-to-background ratio in a short time. Thus, it is very suitable for skeletal exams to rule out bone pathologies [2]. The differential diagnoses of the initial findings included benign or malignant primary bone neoplasm, early Paget's disease, osteomyelitis or avulsion injury.

Bone is a common site of metastasis for carcinomas of the prostate, breast, lung, kidney, bladder, thyroid, lymphomas and sarcomas. Prostate, breast, and lung primaries account for 80% of all bone metastases. Metastatic tumors of bone are more common than primary bone tumors. Tumors usually spread to bone hematogenously. Bone metastases may be asymptomatic or may produce pain, swelling, nerve root or spinal cord compression, pathologic fracture, or myelophthisis (replacement of the marrow) [3].
The four most common malignant primary bone tumors are osteosarcoma, chondrosarcoma, Ewing's sarcoma, and malignant fibrous histiocytoma and they tend to be heterogeneous and very osteoblastic. Rare malignant primary bone tumors include chordoma, malignant giant cell tumor, adamantinoma, and hemangioendothelioma. Benign primary bone tumors include enchondroma, osteochondroma, chondroblastoma, chondromyxoid fibroma, osteoid osteoma, osteoblastoma, fibroma, desmoplastic fibroma, and hemangioma. Benign primary bone tumors may show some osteoblastic cortical reaction of the bone [3].

Paget disease has a prevalence of 1%-2% in the United States and it affects men more commonly than women. It is usually diagnosed in patients over age 40 years and its prevalence doubles with each decade thereafter, reaching an incidence of about 10% after age 80 [4]. It is usually discovered incidentally during radiology imaging or because of incidental discovery of elevation in serum alkaline phosphatase. The cause of Paget disease is unknown. However, there is often a genetic component since about 15%-25% of affected patients have a first-degree relative with the disease. The finding of virus-like particles in affected bones also led to the suggestion that Paget's disease may represent a virus infection of bone [5]. Paget's disease is manifested by overactive osteoclastic bone resorption followed by a compensatory increase in osteoblastic new bone formation. Paget disease is often mild and asymptomatic. Only about 27% of affected individuals are symptomatic at the time of diagnosis [5]. Bone pain may be the first symptom. It can involve just one bone (monostotic) or multiple bones (polystotic). The skeletal sites most commonly involved are the pelvis, vertebral bodies, skull, femur, and tibia. Patients with limited monostotic involvement may have serum alkaline phosphatase levels within the normal range [6]. Paget's disease is associated with an increased risk (2%-5%) of malignant neoplasms in the involved bones—most often osteosarcoma, with fibrosarcoma and chondrosarcoma occurring less commonly [6].

In the United States, osteomyelitis affects 0.1%-1.8% of the healthy adult population [7]. With the aging of the population and the epidemics of obesity and diabetes, this prevalence can rise. Staphylococcus aureus is the most likely bacterial pathogen [7]. Osteomyelitis of the pelvis can result from direct trauma, post-operative infection, decubitus ulcers, or hematogenous dissemination [7]. Lastly, external trauma or falling in sitting position and forceful contraction of hamstrings can cause avulsion fracture of the ischium [8].

2 Case presentation

Our patient is a 54 year old white male with a history of hip arthralgia, lumbar spondylolithesis, obesity, hypertension and hyperlipidemia. He suffered a back injury from a fall about 30 years. He was complaining of persisting right hip pain, unrelieved by naproxen or tramadol. The patient stated that the pain started 2 years ago while playing handball. The pain was dull to stabbing in nature. At the beginning, he was unable to walk even a few blocks but the pain had improved gradually since then. No tenderness in the right hip area was noted in the physical exam. His primary care physician ordered an X-ray study. The X-ray image revealed an oval or flame like sclerotic lesion in the intertrochanteric portion of the right femur (see Figure 1) suspicious for metastatic disease. Consequently, the patient had a NaF-18 PET/CT whole body bone scan to rule out any bone metastases or other bone problems. The bone scan showed an incidental finding of intense heterogeneous osteoblastic activity in the right ischium associated with a rarefied cortex and areas of thin linear cortical interruption suspicious for primary bone neoplasm, Paget’s disease, osteomyelitis in the right clinical setting or trauma. As a consequence he had a MRI exam 1 month later. Alkaline phosphatase (ALP) level was 42 U/L (normal 40-125 U/L). No other lab abnormalities were noted. There were no significant changes in the follow up MRI 3 months later. Orthopedic consult recommended a third MRI study in the next 3 months.
Figure 1. X-Ray image of the right hip showing normal appearing ischium and a sclerotic lesion in the intertrochanteric portion of the right femur, adjacent to the lesser trochanter (arrow), which prompted a further evaluation by a NaF-PET/CT scan.

3 Discussion

The NaF-18 PET/CT whole body bone scan (see Figure 2) excluded widespread metastatic bone disease but the single finding had a broad differential diagnoses, including malignant primary bony lesion, benign primary bony lesion, early Paget's disease, osteomyelitis, infiltrative bone lesion and chronic avulsion injury at the hamstring attachment. Focal benign bone lesions usually show mild osteoblastic activity in the bone scans and tend to be nonspecific. In general, secondary metastasis tend to show up as multiple lesions on bone scans and solitary bony lesions are usually benign [9].

The solitary lesion on right ischium was stable on the two follow-up MRIs (see Figure 3a~3b), and the possibility of primary or secondary bone tumor was ruled out.

Monostotic Paget's disease involving the pelvis represents 22% of all Paget's disease cases [10]. Elevated ALP is characteristic of Paget’s disease but the rise in ALP can take decades to develop [11]. A new follow-up lab test again showed no elevation of ALP. Bone scans are helpful in detecting early Pagetic lesions even before any radiologic changes are apparent. Paget’s disease tend to have very intense osteoblastic activity in metabolic imaging of the bone, usually with effect of bone expansion and even deformity. Although the intense osteoblastic activity tends to be very diffuse and homogeneous with Tc99m- MDP-bone scan imaging, it appears more patchy and heterogeneous with NaF-PET/CT imaging.

Both osteomyelitis and Paget's disease can have increased uptake on bone scans due to high perfusion to the affected areas [10, 12]. Osteomyelitis was ruled out on the basis of absence of fever and elevated WBC, and no history of intervention.
in this area. Infiltrative bone lesions can have many causes and they can range from benign to malignant in nature [13], but there are usually accompanying marrow abnormalities. The bone marrow appears normal on the two MRI studies (see Figure 3a and 3b), and infiltrative process was ruled out.

**Figure 3.** (a) Follow up T2 weighted MRI study one month later showed slight cortical thinning without focal mass or marrow abnormality in the right ischium. (b) Follow up MRI 3 months after the last one showed no interval changes.

Finally, considering prior history of trauma, avulsion injury at muscle insertion was favored. Avulsion injuries of the ischium are associated with local tenderness and difficulty to ambulate [14]. Our patient had chronic pain in the right hip area and he had difficulty walking right after the injury. Chronic avulsion fractures can have prominent reactions on MRI and heterotrophic bone formations that can look like tumoral growths [15]. It is then expected to present marked osteoblastic osseous reaction in NaF-PET/CT metabolic bone imaging. Furthermore, not all avulsion injuries will have osseous displacement [16] like the present case where only cortical thinning was observed. There may be clear signs of soft tissue injury on MRI such as a tendon tear. However, no soft tissue injury was apparent in the two MRI studies (see Figure 3a and 3b). If a case such as ours is encountered in everyday practice, the possibility of a benign traumatic etiology should be given high consideration.

**References**


