

## Fatigue as a presenting symptom of multiple myeloma

### Multipl miyelomda başvuru semptomu olarak halsizlik

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#### ABSTRACT

A 73-year-old man, without any significant previous history, presented to the Internal Medicine clinic with a one-year history of fatigue and weakness. As demonstrated in our case, the diagnosis of multiple myeloma (MM) presenting as fatigue requires a high degree of clinical suspicion.

**Keywords:** Fatigue, multiple myeloma, presenting signs, weakness.

#### ÖZ

Daha önce klinik öz geçmişi olmayan 73 yaşında bir erkek hasta son bir yıldır yorgunluk ve halsizlik öyküsü ile İç Hastalıkları Kliniğine başvurdu. Olgumuzda olduğu gibi, halsizlik olarak ortaya çıkan multipl miyelom (MM) tanısı yüksek derecede klinik şüphe gerektirir.

**Anahtar sözcükler:** Yorgunluk, multipl miyelom, başvuru semptomlar, halsizlik.

Multiple myeloma (MM) is a neoplastic plasma cell disease estimated to constitute approximately 1.6% of all cancers and 10% of hematological malignancies.<sup>[1]</sup> Patients usually present with non-specific symptoms such as bone pain, fatigue, weakness, and recurrent infections, while many patients present with laboratory abnormalities such as anemia, renal failure, and high erythrocyte sedimentation rate or globulin levels. In the suspicion of MM, the diagnostic workup should include erythrocyte sedimentation rate, serum creatinine, calcium, immunoglobulin levels, proteinuria in 24-hour urine, skeletal survey, and bone marrow assessment.<sup>[1]</sup> In this report, we present a case who was hospitalized to investigate the etiology of fatigue and weakness and subsequently diagnosed as MM with the presence of severe anemia, hypercalcemia, albumin/globulin reversal, creatinine increase, prominent punched-out lytic

lesions in skull X-ray and 50% plasma cell in bone marrow aspiration. The patient was transferred to the Hematology clinic.

#### CASE REPORT

A 73-year-old male patient with known diagnoses of hypertension and coronary artery disease was admitted to the Internal Medicine clinic with a one-year history of progressive weakness. On physical examination, he was conscious, oriented, and cooperative. The rest of the physical examination was normal. Laboratory tests were as follows: white blood cell (WBC): 2,500/mm<sup>3</sup>, hemoglobin: 7.8 g/dL, platelet: 160,000/mm<sup>3</sup>, alanine aminotransferase (ALT): 16 U/L, aspartate aminotransferase (AST): 47 U/L, lactate dehydrogenase (LDH): 625 U/L, urea: 56 mg/dL, creatinine: 1.4 mg/dL, glomerular

**Received:** June 24, 2019 **Accepted:** August 27, 2019 **Published online:** October 31, 2019

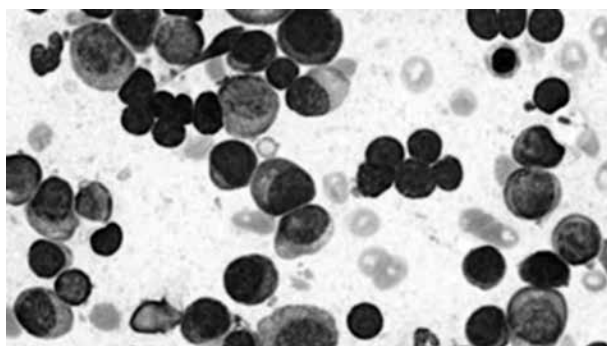
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#### Cite this article as:

Türk U, Dalda Y, Olgun A, Akar H. Fatigue as a presenting symptom of multiple myeloma. FNG & Demirođlu Bilim Tıp Dergisi 2019;5(3):149-151.



**Figure 1.** Multiple punched-out lytic lesions in skull X-ray.



**Figure 2.** Bone marrow aspiration showed 50% plasma cell (Giemsa  $\times 100$ ).

filtration rate (GFR): 52.80 mL/min/1.73m<sup>2</sup>, total protein: 10.3 g/dL, albumin: 2 g/dL, globulin: 8.3 g/dL, calcium: 12 mg/dL, immunoglobulin (IgA): 6610 mg/dL, IgG: 376 mg/dL, IgM: 21.1 mg/dL. Other laboratory values were within normal limits. Multiple punched-out lytic lesions were observed in the patient's skull X-ray (Figure 1). Bone marrow aspiration and biopsy showed 50% plasma cells in the aspiration of the patient (Figure 2). The patient was transferred to the hematology unit with the diagnosis of MM.

## DISCUSSION

Since the symptoms that bring the MM patient to the physician are usually non-specific, early diagnosis may be a diagnostic challenge in primary care. Fatigue, weakness, back pain, weigh loss, or abnormal laboratory results are red flag warnings of MM. Multiple myeloma has several symptoms similar to common patient complaints, which is why the disease may typically undiagnosed until it reaches the

advanced stage.<sup>[2]</sup> Back and lower back pain may be deformative and discogenic but may also be a sign of MM. Pain may be caused by a rheumatic pathology, but also from a large myelomatic osteolytic lesion that is prone to spontaneous fracture.<sup>[2]</sup> Multiple myeloma is among the many causes of anemia and renal failure of unknown origin. Multiple myeloma should be considered in the differential diagnosis when investigating the causes of poor immunity, frequent infections, and hypercalcemia.<sup>[2]</sup> Fatigue, weakness, drowsiness, and confusion can also be attributed to the prominent increase in the concentration of serum globulin (hyperviscosity), calcium levels, and anemia associated with MM.<sup>[2,3]</sup> Most anemic patients with MM suffer from moderate to severe fatigue.<sup>[4]</sup> Replenishment of hemoglobin levels leads to improvement in fatigue and quality of life. Goldschmidt et al.<sup>[5]</sup> reported that 58% of MM patients had back pain and 34% had complaints of fatigue or weight loss. In a study by Sultan et al.,<sup>[6]</sup> the most common presenting complaints were fatigue (81.9%), back pain (80.3%), and bone pain (67.2%). Fuchs et al.<sup>[7]</sup> suggested that an active investigation of paraproteinemia is indicated in cases of end-organ damage such as anemia, renal failure, recurrent infections, or prolonged fatigue. As MM has many different manifestations, it may sometimes present as severe fatigue and weakness. The workup of a patient with long standing weakness and fatigue of unknown cause should include immunofixation electrophoreses, free light chain assay, or bone marrow evaluation.<sup>[8]</sup>

As demonstrated in our case, long standing progressive fatigue and weakness together with anemia, decrease in GFR, albumin/globulin reversal, hypercalcemia and lytic lesion on direct X-ray should be considered red flag warnings for MM.

Multiple myeloma presenting as prolonged fatigue and weakness requires a high degree of suspicion to avoid delay in diagnosis. Multiple myeloma should always be kept in mind in patients presenting with prolonged weakness and fatigue, exhaustion, deep anemia, hypercalcemia, and renal failure.<sup>[9]</sup> Patients with long standing progressive fatigue and weakness of unknown etiology should be screened for possible plasma cell dyscrasia.

**Declaration of conflicting interests**

The authors declared no conflicts of interest with respect to the authorship and/or publication of this article.

**Funding**

The authors received no financial support for the research and/or authorship of this article.

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