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# A Lesion Mimicking Malignancy: Granulomatous Orchitis

#### Abstract

Granulomatous orchitis is a rare disease with mixed chronic and granulomatous inflammation. Various factors cause granulomatous inflammation in the testicles, and it may accompany malignancies. Radiologically, granulomatous orchitis can be confused with malignancy. This paper presents the differential diagnoses of granulomatous orchitis. A 52-year-old man with testicular swelling was referred to the urology clinic. A palpable mass was detected by physical examination. Ultrasonography revealed a solid, heterogeneous 2.5 cm mass suspected as a malignancy in the right testes. The patient underwent elective orchiectomy. Microscopically, the testicle showed necrotising granulomatous inflammation accompanied by an inflammatory infiltrate that destroyed the testicular structure and caused rete testis hyperplasia. Radiologically, granulomatous orchitis can be confused with malignancy. Sufficient samples should be taken to exclude malignancy. Idiopathic granulomatous orchitis can be diagnosed after excluding all possible causes of granulomatous orchitis.

Keywords: Granulomatous orchitis, malignancy, testicle

#### Introduction

Granulomatous orchitis is a rare disease with mixed chronic and granulomatous inflammation. Tuberculosis, brucellosis, actinomycosis, and sarcoidosis are known aetiological factors. Idiopathic granulomatous orchitis (IGO), an inflammatory condition of the testis of unknown aetiology, is rarely encountered (1). Since the ultrasonographic image demonstrated diffuse hypoechoic or focal hypoechoic areas, it is confused with malignancy. Therefore, patients often undergo orchiectomy, and IGO is diagnosed only through histopathological examination (2).

## **Case Presentation**

A 52-year-old man with testicular swelling was referred to the urology clinic. A palpable mass was detected by physical examination. Ultrasonography revealed a solid heterogeneous mass suspected as a malignancy in the right testes. On biochemical examination, beta human chorionic gonadotropin, alpha-fetoprotein and lactate dehydrogenase levels were within their normal range. The patient underwent elective orchiectomy. On macroscopic examination, the right testicle contained an exophytic mass which, upon dissection, demonstrated a necrotic nodule measuring 2.5×1.5×1.5 cm³ (Figure 1A). The whole lesion was examined. On microscopic examination, the

testicle showed multiple necrotising granulomas and rete testis hyperplasia (Figure 1B). There was granulomatous inflammation accompanied by an inflammatory infiltrate consisting of plasma cells and lymphocytes that destroyed the testicular structure (Figure 2A, B), which was accompanied with rete testis hyperplasia (Figure 3). Microorganisms were not seen in periodic acid-Schiff (PAS) and Ziehl-Neelsen (EZN) staining. Burned-out/regressed germ cell tumours were excluded with immunohistochemical study. The histopathological diagnosis was chronic necrotising granulomatous orchitis. The patient underwent clinical, serological and radiological tests after pathological diagnosis, but no potential factor was detected.

## Discussion

IGO most commonly occurs at age 50-70 years, although cases have been reported in all ages (2). IGO can be acute or chronic. In the acute form, patients present with sudden onset of pain, while in the chronic form, they present with unilateral scrotal swelling. Patients can also present with fever, haematuria, dysuria and hydrocele (3).

The aetiology of IGO is not known exactly, but it is thought to be related to trauma. Additionally, autoimmune diseases and infections have been associated with IGO (2). IGO can clinically mimic malignancy. Testicular trauma, surgery, history

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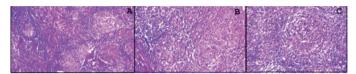
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**Figure 1.** A: Macroscopic view of the testicle which demonstrated a necrotic nodule. B: Microscopic view of the normal testicular tissue on the right and lesion area on the left (haemotoxylin and eosin staining, ×40)



**Figure 2.** A: Multiple necrotising granulomas (hematoxylin and eosin staining (HE),  $\times$ 100). B: Multiple granuloma structures (HE,  $\times$ 200). C: Granuloma structure accompanied by an inflammatory infiltrate (HE,  $\times$ 200)

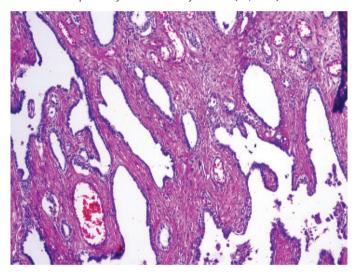


Figure 3. Rete testis haemotoxylin (HE, ×100)

of tuberculosis and urinary tract and epididymis infections may raise clinical suspicion for IGO (4).

Distinguishing IGO from malignancy is extremely challenging because of the destruction of the testicular tissue, as shown in ultrasonography. Therefore, orchiectomy is often performed (2). On histopathological examination, the granulomatous inflammatory infiltrate consisting of plasma cells and lymphocytes invade the seminiferous tubules and germ cells degenerate. Fibrosis develops in the later period (5).

Inflammatory and infectious factors should be considered in the differential diagnosis. The most common cause of granulomatous inflammation is tuberculosis. The presence of granulomas with caseification necrosis accompanied by Langhans-type giant cells and the presence of microorganisms in the acid-fast histochemical staining suggest tuberculosis (6). Other factors in the differential diagnosis are syphilis, leprosy, brucella and fungal infections. In these factors, the

specific morphological appearance of granulomas; presence of microorganisms in histochemical stains such as Warthin-Starry, EZN and PAS; and clinical laboratory findings are helpful in the diagnosis (7). Sarcoidosis that rarely involves the genitourinary system should be also considered. Non-caseating granulomas, minimal lymphocytic infiltration, Schauman basophilic, asteroid bodies and systemic involvement in patients support the diagnosis of sarcoidosis (8). Granulomatous seminoma on histopathological examination should also be considered in the differential diagnosis. The presence of intratubular germ cell neoplasia, immunohistochemical presence of a tumour such as OCT3/4 and CD117 and placental alkaline phosphatase are also helpful in the differential diagnosis (9). Malignant lymphoma is another neoplasia that should be included in the differential diagnosis. In IGO, lymphoma is excluded if the inflammatory infiltrate is not monotonous and shows polyclonal pattern on immunohistochemical staining and partially preserved tubular structure of the testis (10).

In the present case, malignancy was considered following clinical and ultrasonographic examinations. Histopathologically, the granuloma structure specific to infectious agents was not observed, and organisms were not detected histochemically. On morphological examination, burned-out/regressed germ cell tumour was not considered and was supported by immunohistochemical studies. Thus, the present case was evaluated as IGO after all other causes were excluded. Moreover, rete testis hyperplasia was present. Rete testis hyperplasia is a non-neoplastic lesion that mimics a malignancy of unknown aetiology (11). The accompanying rete testis hyperplasia in our case may be due to degeneration resulting from granulomatous inflammation.

Radiologically, granulomatous orchitis can be confused with malignancy. Thus, sufficient samples should be taken to exclude malignancy. IGO can be diagnosed after excluding all possible causes of granulomatous orchitis.

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## **Ethics**

**Informed Consent:** The patient provided written informed consent which includes the case details and agreed to the publication of histopathological images.

Peer-review: Externally peer-reviewed.

## **Authorship Contributions**

Concept: Ç.Ö., Design: Ç.Ö., E.P., Data Collection or Processing: Ç.Ö., T.B.Ş., Analysis or Interpretation: T.B.Ş., Literature Search: Ç.Ö., E.P., Writing: Ç.Ö., E.P., T.B.Ş.

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