Testicular Metastasis of Prostate Cancer: A Case Report

Ayumu Kusaka  Takuya Koie  Hayato Yamamoto  Itsuto Hamano  Takahiro Yoneyama  Yasuhiro Hashimoto  Chikara Ohyama  Yuki Tobisawa  Tohru Yoneyama
Department of Urology, Hirosaki University Graduate School of Medicine, Hirosaki, Japan

Key Words
Prostate cancer  ·  Testicular metastasis  ·  Orchiectomy

Abstract
The incidence of secondary neoplasms of the testis during autopsies is approximately 2.5%. Although most secondary testicular metastases are due to prostate cancer, only a few patients with prostate cancer have clinically manifested testicular metastasis. We report the case of a prostate cancer patient with testicular metastasis who was diagnosed after the presence of a palpable mass in the right testis. A 56-year-old Japanese male presented to our hospital with an elevated serum prostate-specific antigen (PSA) level of 137 ng/ml. He was diagnosed with stage IV (T3N1M1b) prostate cancer and received androgen deprivation therapy, followed by various hormonal manipulations. His serum PSA level was undetectable for 1 year. No distant metastases were detected during imaging examinations. He received radiation therapy; however, his serum PSA level increased gradually. Four months later, he presented with right testicular swelling. Computed tomography revealed a heterogenous mass in the right testis and a right high inguinal orchiectomy was performed. Histopathological analysis showed that the right testis was infiltrated with metastatic adenocarcinoma with a Gleason score of 8. This is a rare case of right testicular metastasis in a patient with prostate cancer. Testicular metastasis of prostate cancer can be aggressive and metastasize.

Introduction
The incidence of secondary neoplasms of the testis during autopsies is approximately 2.5%, including nonneoplastic deaths [1]. Most secondary testicular metastases originate from distant primary sites, of which the lung, prostate, and gastrointestinal tract are
common [2]. Testicular metastases occur in up to 4% of all prostate cancer (PCa) cases and are often detected incidentally after orchiectomy treatment for advanced disease [3]. Generally, PCa metastases to the pelvic lymph nodes, bones, lungs, and liver occur frequently in the advanced stages; however, few of these patients have clinically manifested testicular metastasis [4]. We report the case of a patient with PCa with testicular metastasis who was diagnosed based on the presence of a palpable mass in the right testis.

Case Presentation

A 56-year-old Japanese male presented to our hospital with an elevated serum prostate-specific antigen (PSA) level of 137 ng/ml. On digital examination, the prostate was palpable with hard consistency on both lobes. He underwent a transrectal ultrasound-guided systematic sextant biopsy. Histological examination revealed adenocarcinoma of the prostate with a Gleason score of 9 (fig. 1). Computed tomography (CT) showed multiple lymph node enlargements from the pelvis to the bifurcation of the left renal artery. Bone scan imaging showed multiple skeletal metastases in the left ilium and ischium.

Written informed consent was obtained from the patient for the publication of this case report and the use of the accompanying images. A copy of the written consent is available for review by the Editor-in-Chief of this journal.

The patient was diagnosed with stage IV (T3N1M1b) PCa. He received androgen deprivation therapy (ADT) for 4 years, followed by luteinizing, hormone-releasing hormone and estramustine for 1 year. His serum PSA level was undetectable for 1 year, and no lymph node enlargement and no distant or skeletal metastases were visualized using imaging examinations. He received radiation therapy to the prostate (74 Gy/37 fractions). Although he underwent ADT, his serum PSA level increased gradually. He presented with a right testicular swelling 4 months later. CT revealed a heterogeneous mass of $5 \times 4$ cm in the right testis (fig. 2). Laboratory results for $\alpha$-fetoprotein, human chorionic gonadotropin, and lactate dehydrogenase were negative, and his serum PSA level had increased to 4.94 ng/ml. A right high inguinal orchiectomy was performed. Macroscopic examination of the resected testis revealed a solid yellowish-white tumor of $5 \times 4$ cm. Histopathological analysis showed that the right testis was infiltrated by metastatic adenocarcinoma with a Gleason score of 8 (fig. 3). Immunohistochemical examination revealed that tumor cells were diffusely positive for PSA and $\alpha$-methylacyl CoA racemase (P504S). The patient was diagnosed with right testicular metastasis of PCa.

The patient has been asymptomatic and disease-free with a serum PSA level that decreased to an undetectable level 1 year after surgery.

Discussion

With the exception of the infiltration of leukemia and lymphoma, secondary neoplasms of the testis are rare, with a reported incidence of 0.02–2.5% [1, 2, 5, 6]. The most common primary site of testicular metastases is the prostate (15%), followed by the lung, melanomas, skin, colon, and kidney [7]. However, most testicular metastases of PCa were identified after the examination of large numbers of testes from patients with tumors who had undergone therapeutic orchiectomies [8, 9]. Nevertheless testicular metastases of PCa are rare. Bubendorf et al. [7] studied a cohort of 1,589 patients with PCa and reported that 35% had
hematogenous metastases most frequently involving bone (90%), lung (46%), and liver (25%); metastases in the testes were found in 0.5% (n = 3).

The proposed mechanisms for the spread of PCa to the testis include: (i) retrograde venous extension or embolism; (ii) arterial embolism; (iii) lymphatic extension, and (iv) endocanaliculal spread [6, 7]. The involvement of the prostatic urethra in PCa increases the risk of testicular metastases [3]. In our case, retrograde venous extension might have caused testicular metastasis due to embolism after radiation therapy.

The histological features of testicular metastasis are similar to those of primary PCa [10]. Most patients with PCa with lymph node involvement or skeletal metastases undergo ADT for several years. In previous studies, the interval from diagnosis of PCa to testicular metastasis ranged from 2.5 to 15 years [11–13]. Therefore, the histological pattern of testicular metastasis might change to another subtype of PCa such as small cell carcinoma [3]. Tu et al. [3] reported that the pathological features of testicular metastases resembled ductal or endometrioid adenocarcinoma of the prostate and that tumors contained tall columnar cells in a papillary, cribriform, and/or solid pattern. Therefore, a ductal prostate adenocarcinoma that metastasizes to the testis might be an admixture of different tumor types arising from specific prostatic stem cells [3].

In patients with testicular metastasis of PCa, survival after diagnosis is usually <1 year [14]. Lu et al. [15] found that the mean survival period after orchiectomy was 12.8 months in patients with PCa and 7.4 months in those with other forms of cancer. Nevertheless, as in the current case, the patient’s serum PSA can remain undetectable, but testicular metastasis can behave aggressively and further metastasize.

**Disclosure Statement**

The authors declare that there are no conflicts of interest regarding the publication of this paper.

**References**

Fig. 1. HE-stained section of the primary tumor. The pathological findings indicate prostate cancer with a Gleason score of 9. ×100.

Fig. 2. Abdominal enhanced CT. Abdominal CT shows a heterogeneous tumor (size, 5 × 4 cm) in the right testis (arrowheads).

**Fig. 3.** HE-stained section. The pathological findings indicated prostate cancer with a Gleason score of 8. ×40.