Gallbladder Metastasis from Renal Cell Carcinoma

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Key Words
Gallbladder metastasis · Renal cell carcinoma · Sunitinib

Abstract
A 73-year-old female was operated with radical nephrectomy and cholecystectomy for renal cell carcinoma and suspected gallstones after 9 courses of sunitinib treatment. Gallbladder specimen showed gallbladder metastasis originating from the renal cell carcinoma. Gallbladder metastasis from renal cell carcinoma is rare. Here, we discuss a case of gallbladder metastasis from renal cell carcinoma.

Introduction
Renal cell carcinoma is a tumor that often spreads to organs with a large blood flow volume, such as the lungs, the liver, and bones. Reports of clinical cases of metastasis to the gallbladder are rare, and in many of these reported cases, the metastasis is found by chance during autopsy. We treated a case of gallbladder metastasis of renal cell carcinoma and, in this study, we report this case in addition to several bibliographical considerations.

Case Report
The patient was a 73-year-old female with a chief complaint of gross hematuria. She had no past history in particular.
Current Medical History

In May 2008, the patient noticed blood in her urine and visited our department. A computed tomography (CT) showed renal cell carcinoma with multiple lung metastasis and invasion to the inferior vena cava, resulting in a diagnosis of clinical stage T3bN0M1. She received 9 courses of sunitinib treatment: 1 course of 50 mg/body from July 2008 to August 2008, 1 course of 37.5 mg/body from August 2008 to September 2008, 7 courses of 25 mg/body from September 2008 to August 2009. Considering the reduction of tumor thrombosis and good control of lung metastases, she was admitted to our department for surgery.

Laboratory Data at the Time of Admission

Hematological Values. RBC 283/μl, WBC 2,600/μl, Hb 9.5 g/dl, Plt 11.0 × 10⁴/μl.

Biochemistry. AST 75 IU/l, ALT 20 IU/l, ALP 263 IU/l, LDH 475 IU/l, BUN 11 mg/dl, Cre 0.93 mg/dl, CRP 0.81 g/dl.

Urinary Analysis. Within normal limits.

18F-Fluorodeoxyglucose Positron Emission Tomography (PET)/CT Findings

18F-fluorodeoxyglucose (FDG) accumulations with a low level of accumulation in the center were observed in the upper pole of the right kidney. These accumulations extended from the renal hilus toward the inferior vena cava and the SUVmax was 5.8. Multiple negative nodules were observed on both sides, and these were believed to be lung metastases of the renal carcinoma.

Specimens Removed

The removed kidney was 827 g in weight and 110 × 100 × 90 mm in size. A yellowish white to wine-colored tumor was observed along with a clear cell renal carcinoma 4 × 3 mm in size within the kidney on the same side. The removed gallbladder was 105 × 60 × 20 mm in size, and a polypoid nodule 10 × 8 mm in size was observed in the neck of the gallbladder (fig. 1).

Histopathological Findings

The renal carcinoma within the removed kidney was of a clear cell type, and invasion into the right vein was observed (fig. 2). With regard to the polypoid nodule in the gallbladder, an alveolar composed of clear tumor cells accompanied by abundant angiogenesis was observed. The histological type coincides with that of the removed kidney (i.e., clear cell carcinoma) (fig. 3).

Discussion

Distant metastases of renal cell carcinoma discovered during autopsy are mostly found in the lungs, the liver, and bones. Metastasis to the gallbladder is very rare and is found in only 0.4–0.6% of specimens [1, 2]. Besides, malignant melanomas are the most common cause of metastatic tumors of the gallbladder, and metastases of lung, renal, pancreatic, and colorectal cancers have been reported [1, 3]. However, the number of clinical cases reported so far does not exceed 20. Furthermore, there have only been 10 cases, including this one, in which the metastasis did not involve any organs other than the gallbladder [4–11]. In all cases, the histological type was clear cell carcinoma.

With respect to the mechanisms of secondary tumors of the gallbladder, Willis [1] grouped the processes of metastasis to the gallbladder into 2 categories including direct invasion of the tumor and invasion of the tumor into the blood vessel capillaries, saying that the latter process is comparatively rare. The gallbladder metastasis occurred...
simultaneously in half of the cases and recurred metachronously in half of the cases. Another characteristic of these cases is the discrepancy of 19 months to 27 years for the time between the resection of the primary tumor and the reappearance of the tumor cells.

According to Takebayashi et al. [12], the features to be noted in preoperative diagnoses of metastatic gallbladder carcinoma with the use of diagnostic imaging include: (1) the appearance of the carcinoma on CT images and in angiography as a polypoid lesion representing significant tumor staining; and (2) the appearance of an echo-bright area on the surface of the tumor indicating a submucosal tumor on ultrasound images. However, they reported that it is extremely difficult to distinguish cases of primary and metastatic gallbladder carcinoma. In our present case, PET/CT scans were performed before surgery. There having been no reports on PET/CT scans used to detect gallbladder metastasis of renal cell carcinoma, this report is the first such report. The PET/CT images indicated a tumor mass within the gallbladder wall, but the FDG images were not very clear.

In cases of renal carcinoma in which a gallbladder mass is observed simultaneously or metachronously, although it is difficult to obtain a preoperative diagnosis of gallbladder metastasis, the possibility of gallbladder metastasis should be taken into account.

Fig. 1. Gallbladder specimen.
**Fig. 2.** Renal. Hematoxilin-Eosin (HE) staining (×400).

**Fig. 3.** Gallbladder. Hematoxilin-Eosin (HE) staining (×400).
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