Pancreatic Squamous Cell Carcinoma

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Abstract
Pancreatic squamous cell carcinoma (SCC) is a rare event. Here, we present a 56-year-old man with pancreatic SCC. Imaging methods demonstrated a hypodense mass at the head and trunk of the pancreas. Also, some lymphadenopathy has been seen around the pancreas and para-aorta. The mass created pressure and encasement on the celiac trunk, portal vein, and arteries of the liver and spleen. Endoscopic ultrasound showed a mass lesion of 45–37 mm (mixed echoic) at the trunk of the pancreas. Histological examination of the endoscopic ultrasound-guided fine needle aspiration specimen confirmed the diagnosis of SCC. The disease is highly aggressive, most often locally advanced or metastatic at diagnosis, and poorly responsive to treatment. It also has generally poor survival rates.

Introduction

The majority of pancreatic carcinomas are primary tumors. Lately, expansion of imaging techniques led to more consideration for tumors of the pancreas [1]. Normally, the pancreas lacks squamous epithelium. During periods of inflammation such as in pancreatitis and squamous cell carcinoma (SCC) these cells can be observed [2]. The pancreas neoplasms are...
classified as endocrine or non-endocrine. Based on the location of the tumor, non-endocrine neoplasms can have five different sources: acinar, connective tissue, ductal, mixed cell type, and uncertain origin [3]. Also, the pancreatic ductal malignancy can be subdivided into adenocarcinoma and SCC. SCC of the pancreas is a controversial entity of uncertain origin [4]. Pancreatic SCC is a primary rare malignancy [5, 6]. It accounts for approximately 0.5–2% of all malignant pancreatic tumors [7]. Because it is an infrequent disorder, a diagnosis of primary pancreatic SCC is made only after excluding other neoplastic lesions of the pancreas that contain squamous epithelial components [2]. The main differential diagnoses of primary SCC of the pancreas are adenosquamous carcinoma (ASC) – another rare primary tumor of the pancreas – and metastatic SCC from another primary site [8]. Here, we describe a patient with pancreatic SCC along with the clinicopathological features.

**Case Report**

Three months ago, a 56-year-old man with epigastric and right upper quadrant pain was admitted to a hospital. The disease was diagnosed as gallstones and cholecystectomy was done for the patient. After surgery, not only did the symptoms resolve but also loss of appetite and weight (15%) were added to the clinical signs. The patient was referred to our hospital because of abdominal pain. Subsequently, physical examination was done. Initial laboratory evaluation revealed moderate elevations of amylase 215 IU/L (normal up to 100 IU/L) and lipase 130 IU/L (normal up to 60 IU/L) along with mild anemia (Hb 11.5 g/dL). Blood samples taken at the time of admission and clinical parameters were as follows: BUN 13 mg/dL (normal range 8–20 mg/dL); Cr 1.5 mg/dL (normal range for male 0.6–1.3 mg/dL); uric acid 3.3 mg/dL (normal range for male 3.6–7.7 mg/dL); TG 88 mg/dL (normal range up to 150 mg/dL); cholesterol 124 mg/dL (normal range 85–10.5 mg/dL); ph 3.8 mg/dL (normal range 2.7–5 mg/dL); Na 133 mEq/L (normal range 135–145 mEq/L); K 4.2 (normal range 3.5–5 mEq/L); AST 17 IU/L (normal <40); ALT 11 IU/L (normal <40); ALP 193; LDH 301 IU/L (normal 225–450 IU/L); amylase 88 IU/L (normal up to 100 IU/L); lipase 66 IU/L (normal up to 60 IU/L); total protein 6.2 IU/L (normal 6.6–8.3 IU/L); albumin 3.9 g/dL (normal 3.5–5.2 g/dL); total bilirubin 0.4 mg/dL (normal 0.3–2 mg/dL); direct bilirubin 0.2 mg/dL (normal <1 mg/dL); WBC 6.1 × 10^3/μL; RBC 4.5 × 10^6/μL; HGB 11.5 g/dL (normal 13–16 g/dL); HCT 33.9% (normal 39–48%); MCV 79.8 fl (normal 80–100); MCH 27.1 pg (normal 26–34.4); MCHC 33.9 g/dL (normal 31.8–36.3 g/dL); PLT 399 × 10^3/μL (normal 150–400 × 10^3/μL).

The initial diagnosis was unclear and the patient underwent computed tomography (CT) of the abdomen for further evaluation (Fig. 1). The CT showed the presence of a hypodense mass at the head and trunk of the pancreas. Also, some lymphadenopathy has been seen around the pancreas and para-aorta. The mass created a pressure and encasement on the celiac trunk, portal vein, and arteries of the liver and spleen. For further evaluation, endoscopic ultrasound (EUS)-guided fine needle aspiration (FNA) was performed (Fig. 2). EUS showed a mass lesion of 4.5–37 mm (mixed echoic) at the trunk of the pancreas invading the splenic vein and artery. The pancreatic duct was 7 mm at the tail of the pancreas. FNA samples from the pancreas were sent for pathological examination (Fig. 3).

Pathological finding indicated sheets of malignant cells with round to oval nuclei with thick nuclear membrane. Occasional prominent nucleoli and high N/C ratio cytoplasm were eosinophilic granular and moderate to abundant with rather well-defined borders. The chromatin pattern was coarse and the mitotic figures were easy to find. Cytologic study was
positive for SCC and further confirmed by immunohistochemistry. The immunohistochemical studies (Fig. 4) were performed using antihuman antibodies against the following markers: unlike CEA, which was negative in tumor cells, CK5/6 and P63 were strongly positive in tumor cells.

The definitive diagnosis of pancreatic SCC was justified by immunohistochemical and pathological study; thus, the patient was sent for chemotherapy. The patient received chemotherapy with gemzar and oxaliplatin (GEMOX at doses of 1,000 and 100 mg/m²) for 4 cycles. He had no response to treatment and chemotherapy was changed to weekly irinotecan (140 mg weekly). Four months after chemotherapy, clinical signs continued and abdominal CT was done again (Fig. 5). Although chemotherapy was done for the patient, the CT image revealed a development in SCC of the pancreas.

Discussion

Pancreatic SCC is a rare exocrine pancreatic carcinoma [9]. There are different reports for the incidence of SCC, ranging from 0.5 to 5% [10]. The histogenesis of pancreatic SCC is vague, but there are some hypotheses: (1) a bipotential primitive cell capable of differentiating into either squamous or glandular carcinoma undergoes malignant change; (2) squamous metaplasia of the ductal epithelium undergoes a malignant transformation; (3) preexisting adenocarcinoma undergoes squamous change; or (4) an atypical squamous cell undergoes a malignant change [11, 12]. The prominent feature of pancreatic SCC is similar to ductal adenocarcinoma including upper abdominal and back pain, anorexia, weight loss, nausea, fatigue, vomiting, and obstructive jaundice [13]. Local lymph node involvement and liver metastases are common with this tumor [14]. At present, a tumor blush in angiography and endoscopic retrograde cholangiopancreatography or enhancement of the tumor on contrast CT can be useful in SCC diagnosis [15–17]. However, EUS-FNA has been increasingly accepted for the identification of pancreatic malignancies with a high sensitivity and specificity [18]. The disease has a poor prognosis and the survival from the time of diagnosis is similar to that of pancreatic adenocarcinoma [19]. Although different therapeutic methods have been used in the handling of pancreatic SCC, which include surgical resection, chemotherapeutic regimens, and radiotherapy, none have been proven effective [20].

Statement of Ethics

All procedures and data gathering were performed by the patient’s informed consent.

Disclosure Statement

The authors declare that they have no conflict of interest.
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**Fig. 1.** CT of the abdomen using pancreatic protocol.

**Fig. 2.** EUS image of the pancreas with squamous cell carcinoma.
Fig. 3. The neoplasm is composed of many malignant cells with round to oval nuclei (HE). a Sheets of malignant cells with round to oval nuclei with thick nuclear membrane. b Prominent nucleoli and high N/C ratio cytoplasm which are eosinophilic granular and moderate to abundant with rather well-defined borders.

Fig. 4. The CEA marker (a) is negative, the CK5/6 (b) and P63 (c) markers are positive.

Fig. 5. Abdominal axial CT scan depicting a developed pancreatic SCC.