Squamous Cell Carcinoma of the Pancreas with Massive Invasion of the Retroperitoneum

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A 79-year-old woman with a rare form of pancreatic carcinoma with massive invasion of the retroperitoneum presented with upper abdominal pain and vomiting. Although examination (computed tomography, barium enema, upper gastrointestinal series) suggested peritonitis carcinomatosa due to pancreatic cancer, a primary lesion of the pancreas was not confirmed by endoscopic retrograde pancreatography. Autopsy ultimately revealed a small tumor (5×8 mm) of the uncinate process of the pancreas near the duodenum with peritonitis carcinomatosa. Microscopically, the tumor and its metastasis consisted of poorly differentiated squamous cell carcinoma without adenocarcinomatous change, a rare form of pancreatic tumor.

Key words: pancreatic cancer, peritonitis carcinomatosa

Introduction

Squamous cell carcinoma is a rare form of pancreatic cancer that is derived from ductal cells, with most studies citing an incidence of about 0.5–2.0% (1–5). We report an unusual case of squamous cell carcinoma of the pancreas; a small tumor in the uncinate process close to the third portion of the duodenum had massively invaded the retroperitoneal region.

Case Report

A 79-year-old Japanese woman was referred to us with upper abdominal pain, vomiting, and a slight fever. She reported having epigastric discomfort and vomiting after meals intermittently for the previous 4 months. Her history revealed a cholecystectomy and choledocho-duodenostomy performed for gallstones 24 years earlier. She did not smoke or drink alcohol. Physical examination revealed a firm tender mass deep in the left upper quadrant of the abdomen. Ascites was absent. The remainder of the physical examination, including pelvic and rectal examination, was negative. Laboratory data revealed an increase in the white blood cell count to 10,100/mm³ and elevated serum levels of aspartate aminotransferase (50 U/l), alanine aminotransferase (94 U/l) and alkaline phosphatase (1,777 U/l). The serum level of elastase-1 was slightly increased, but the serum amylase level was normal. Tumor marker levels were normal except for Dupan-2. Serum levels of electrolytes and of blood urea nitrogen (BUN) were abnormal due to severe vomiting (Table 1). A chest X-ray showed no abnormality.

Ultrasonography showed mobile high-echogenic spots in the liver which suggested pneumobilia, but there was no tumor in the upper left quadrant of the abdomen. Computed tomography (CT) of the abdomen showed an irregularity around the third portion of the duodenum. The left kidney was surrounded by a thick area of low density (Fig. 1). An upper gastrointestinal series showed the presence of choledocho-duodenal anastomosis and a narrowed and irregular stenosis of the third portion of the duodenum (Fig. 2). Endoscopic retrograde pancreatography (ERP) revealed no abnormality of the main pancreatic duct nor branches of the uncinate process of the head of the pancreas (Fig. 3). Endoscopic findings showed marked stenosis in the third portion of the duodenum, but no malignant change was observed on the mucosal surface of that region. The catheter was easily passed through the stenotic area (Fig. 4) and no malignancy was apparent in the biopsy specimens obtained from that area. Barium enema showed segmental narrowing with spiculation in the transverse and descending colon (Fig. 5). Even no abnormality was revealed on ERP, the clinical picture was considered consistent with a pancreatic carcinoma.
Table 1. Laboratory Data on Admission

<table>
<thead>
<tr>
<th></th>
<th>Blood chemistry</th>
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<tbody>
<tr>
<td>Urinalysis</td>
<td></td>
<td>Not particular</td>
</tr>
<tr>
<td>Stool occult blood</td>
<td></td>
<td>(+)</td>
</tr>
<tr>
<td>Peripheral blood</td>
<td></td>
<td></td>
</tr>
<tr>
<td>White blood cell count</td>
<td></td>
<td>10,100/mm$^3$</td>
</tr>
<tr>
<td>Red blood cell count</td>
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<td>481×10$^4$/mm$^3$</td>
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<tr>
<td>Hemoglobin</td>
<td></td>
<td>13.9 g/dl</td>
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<tr>
<td>Hematocrit</td>
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<td>41.4%</td>
</tr>
<tr>
<td>Platelet</td>
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<td>34.1×10$^4$/mm$^3$</td>
</tr>
<tr>
<td>Serology</td>
<td></td>
<td></td>
</tr>
<tr>
<td>C reactive protein</td>
<td></td>
<td>7.4 mg/dl</td>
</tr>
<tr>
<td>Rheumatoid arthritis factor</td>
<td></td>
<td>(-)</td>
</tr>
<tr>
<td>Treponema pallidum</td>
<td></td>
<td>(-)</td>
</tr>
<tr>
<td>Hemagglutination test</td>
<td></td>
<td>(-)</td>
</tr>
<tr>
<td>Hepatitis B virus surface antigen</td>
<td></td>
<td>(-)</td>
</tr>
<tr>
<td>Hepatitis C virus antibody</td>
<td></td>
<td>(-)</td>
</tr>
<tr>
<td>Tumor marker</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Carcinoembryonic antigen</td>
<td></td>
<td>2.3 ng/ml</td>
</tr>
<tr>
<td>Carcinogenic antigen 19-9</td>
<td></td>
<td>&lt;10 U/ml</td>
</tr>
<tr>
<td>Arphafetoprotein</td>
<td></td>
<td>5 ng/ml</td>
</tr>
<tr>
<td>DUPAN-2</td>
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<td>2,000 U/ml</td>
</tr>
<tr>
<td>Elastase-1</td>
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<td>480 ng/ml</td>
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<tr>
<td>Chest X-P</td>
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<td>Not particular</td>
</tr>
<tr>
<td>Electrocardiogram</td>
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<td>Not particular</td>
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<td></td>
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</tbody>
</table>

**Urinalysis**
- Not particular

**Stool occult blood**
- (+)

**Peripheral blood**
- White blood cell count: 10,100/mm$^3$
- Red blood cell count: 481×10$^4$/mm$^3$
- Hemoglobin: 13.9 g/dl
- Hematocrit: 41.4%
- Platelet: 34.1×10$^4$/mm$^3$

**Serology**
- C reactive protein: 7.4 mg/dl
- Rheumatoid arthritis factor: (-)
- Treponema pallidum: (-)
- Hemagglutination test: (-)
- Hepatitis B virus surface antigen: (-)
- Hepatitis C virus antibody: (-)

**Tumor marker**
- Carcinoembryonic antigen: 2.3 ng/ml
- Carcinogenic antigen 19-9: <10 U/ml
- Arphafetoprotein: 5 ng/ml
- DUPAN-2: 2,000 U/ml
- Elastase-1: 480 ng/ml
- Chest X-P: Not particular
- Electrocardiogram: Not particular

**Blood chemistry**
- Na: 130 mEq/l
- K: 3.3 mEq/l
- Cl: 79 mEq/l
- Ca: 8.7 mg/dl
- BUN: 40 mg/dl
- Creatinin: 1.4 mg/dl
- Uric acid: 11.7 mg/dl
- Total protein: 7.1 g/dl
- Albumin: 3.8 g/dl
- Total bilirubin: 1.1 mg/dl
- Direct bilirubin: 0.8 mg/dl
- Alkaline phosphatase: 1,777 U/l
- γ-glutamyl transpeptidase: 699 U/l
- Choline esterase: 245 U/l
- Aspartate aminotranspeptidase: 50 U/l
- Alanine aminotranspeptidase: 94 U/l
- Lactate dehydrogenase: 120 U/l
- Creatinin phosphokinase: 16 U/l
- Amylase: 259 U/l
- Total cholesterol: 186 mg/dl
- Triglyceride: 124 mg/dl
- Glucose: 93 mg/dl
- Total bile acid: 4.7 μmol/l

**Chest X-P**
- Not particular

**Electrocardiogram**
- Not particular

**Fig. 1.** Abdominal CT. Abdominal contrast CT showed irregularity around the third portion of the duodenum and an area of low density around the left kidney.

**Fig. 2.** Upper gastrointestinal series. An irregular stenosis of the third portion of the duodenum was found.

arising in the uncinate process and massively invading the retroperitoneum. The patient secreted 1,000 ml/day of gastric juice and bile through a nasogastric tube, thus she was scheduled for a gastro-jejunalostomy. However, as severe malignant invasion of the mesentery was found, only a gastrostomy could be performed. Examination of the biopsy specimen obtained from the mesentery intraoperatively revealed a squamous cell carcinoma. The patient subsequently developed numerous complications, including electrolyte disturbances from a persistent loss of gastric juice and bile, acute renal failure, and ultimately, heart failure. She died on the 44th hospital day.

Autopsy revealed a marked thickening of the mesentery and the retroperitoneum associated with progressive peritonitis carcinomatosa. No tumor was observed macroscopically in the uncinate process of the pancreas. The cut surface of the uncinate process, including the stenotic lesion of the duodenum, showed a small (5×8 mm) whitish tumor very close to the markedly thickened duodenal wall (Fig. 6). The tumor was present at the
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Fig. 3. Endoscopic retrograde pancreatography. Pancreatogram demonstrated the absence of abnormal findings on the main pancreatic duct and branches of the uncinate process.

Fig. 4. Radiography of the duodenum. Basket catheter seems to pass easily through the stenosis of the third portion of the duodenum.

Fig. 5. Radiography of the colon. X-ray following barium enema showed irregular narrowing of the transverse and descending colon.

Fig. 6. The cut surface of the uncinate process of the pancreas. A small tumor (5x8 mm) was found close to the duodenal wall.

Discussion

It was difficult to definitively diagnose this case as pancreatic carcinoma before autopsy. Although the clinical picture and results of the various examinations all suggested peritonitis carcinomatosa due to pancreatic cancer, the ERP revealed no abnormality in the pancreas. Evidence obtained at autopsy was consistent with a diagnosis of primary squamous cell carcinoma of the pancreas.

edge of the pancreatic tissue and had disrupted the muscular layer of the duodenum (Fig. 7). Histological examination revealed a poorly differentiated squamous cell carcinoma in the tumor and its metastasis (Fig. 8). Adenocarcinomatous component was not present. Malignancy was not found in the biliary tract, lung, urinary tract, ovary, uterus, or the other organs examined.
Squamous cell carcinoma is a relatively rare form of carcinoma of the pancreas. Of the 1300 cases of pancreatic cancers observed at autopsy in a survey in Japan in 1992, only 9 cases (0.7%) were squamous cell carcinomas (1). No cases of squamous cell carcinoma were found in the 1211 cases compiled by Saitou from registries for pancreatic cancer in Japan (1993) (2). Several reports cited an incidence of 0.5–1.9% (3–5). It is difficult to obtain a histological diagnosis of squamous cell carcinoma prior to either operation or autopsy. Sprayregen et al reported that hypervascularity and tumor brush in angiography are marked findings that distinguish squamous cell carcinoma from adenocarcinoma, in which the encasement and occlusion of arteries are generally observed (6). Sajjou et al reported that hypervascularity and tumor brush in angiography are marked findings that distinguish squamous cell carcinoma from adenocarcinoma, in which the encasement and occlusion of arteries are generally observed (6).

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Although the histogenesis of a squamous cell carcinoma of the pancreas, whether alone or combined with adenocarcinoma, is not known, at least four theories have been suggested (12–14); 1) a primitive cell capable of differentiating into either squamous or glandular carcinoma undergoes malignant change; 2) a preexisting adenocarcinoma undergoes squamous change; 3) a squamous metaplasia of the ductal epithelium undergoes a malignant formation; or 4) an aberrant squamous cell undergoes a malignant change. The tumor in this case was considered to be derived from aberrant squamous cells in the pancreas, because no metaplastic changes of the ductal cells were observed in the field near the carcinoma and no adenomatous changes were seen in either the carcinoma or the invading lesions.

References