Acinic Cell Carcinoma of the Salivary Gland with Metastatic Spread to the Pancreas

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Key Words
Acinic cell carcinoma · Pancreas · Metastasis · Salivary cancer

Abstract
Metastatic disease to the pancreas is rare among solid tumors and has not been well described for salivary cancers. We report a patient who developed an isolated metastatic lesion in the pancreas from acinic cell carcinoma of the salivary gland, presenting as acute pancreatitis.

Introduction
Salivary gland carcinomas are relatively uncommon, accounting for approximately 6% of all malignancies of the head and neck and <1% of all newly diagnosed cancers in the United States annually [1]. The World Health Organization describes 24 distinct malignant epithelial tumors of salivary origin [2]. Mucoepidermoid carcinoma, adenoid cystic carcinoma, and adenocarcinoma account for over 75% of all salivary gland carcinomas, as described by the Memorial Sloan-Kettering experience [3]. Acinic cell carcinoma (AcCC) is a less common subtype comprising 6–10% of all salivary gland tumors [3, 4]. AcCCs are typically well differentiated and rarely develop distant metastases. We report a case of locally recurrent AcCC that developed an isolated metastasis to the pancreas.
Case Presentation

In 2005, a 46-year-old woman presented with several years of progressive left jaw pain and swelling and ultimately developed partial left-sided facial paralysis. In August 2005, she underwent a left total parotidectomy with partial sacrifice of the left facial nerve. Histopathologic examination of the deep left parotid gland showed a thinly encapsulated low-grade salivary gland neoplasm. Tumor cells exhibited round to oval nuclei with inconspicuous nucleoli and basophilic granular cytoplasm that is characteristic of well-differentiated AcCC. Surgical margins were negative. In April 2008, the patient developed a local recurrence in the left neck, confirmed by fine needle aspiration. On May 29, 2008, she underwent surgical excision of the left neck mass with negative margins and one sampled lymph node negative for disease. No other treatment modalities were given. Two years later, the patient developed a second local recurrence, and in June 2010, a 5 × 3-mm mass consistent with recurrent AcCC was removed from the posterior triangle. In October 2010, a surveillance PET/CT scan revealed a 1-cm hypermetabolic nodule between the left styloid process and the tip of the mastoid. Fine-needle aspiration findings were consistent with a third local recurrence of AcCC. In January 2011, the patient had a left posterior neck exploration surgery with left neck dissection. Pathology revealed a single focus of recurrent disease in a lymph node measuring 2.6 cm. Surgical margins were negative but narrow at <0.1 cm. Three other lymph nodes sampled were negative for metastatic disease. The patient did not receive any adjuvant therapy and was placed on expectant observation.

In August 2011, the patient presented to our institution for a second opinion regarding management of a fourth local recurrence presenting as an enlarging, asymptomatic neck mass below the left mandible. On physical exam, a single 1-cm non-tender nodule was palpable just below the mandible, overlying the left sternocleidomastoid muscle. She had no other palpable masses or lymphadenopathy present. An MRI scan of the face and neck revealed a 1.3-cm mass in the previous operative bed in addition to a 1.4-cm enhancing mass at the skull base. CT of the chest, abdomen, and pelvis showed no evidence of distant metastatic disease. On August 29, 2011, the patient had a wide local excision of a locally recurrent AcCC measuring 1.5 × 1.3 × 1.1 cm (fig. 1a). Margins were negative. The second skull base mass seen on imaging was not amenable to surgical resection. The patient subsequently received concurrent chemoradiation therapy to the left neck from October through December 2011 with a total dose of 7,000 cGy in 35 fractions with weekly cisplatin. A surveillance PET/CT scan performed in March 2012 showed no evidence of disease.

In August 2012, the patient was admitted to a local hospital for epigastric pain, nausea, and vomiting, and was diagnosed with acute pancreatitis. An outside MRI scan of the abdomen revealed a mass in the head of the pancreas (fig. 2). Esophagogastroduodenoscopy and an endoscopic ultrasound with biopsy of the mass were performed, and pathology revealed a well-differentiated neoplasm consistent with metastatic AcCC of salivary gland origin. A PET/CT scan showed minimal fluorodeoxyglucose avidity corresponding with the pancreatic mass but no other areas of active disease and no evidence of local recurrence in the head and neck. On September 21, 2012, the patient underwent a laparoscopic metastasectomy. Microscopic examination revealed a 3.0-cm lobulated neoplasm set within a bed of desmoplastic stroma of the pancreatic neck (fig. 1b). These tumor cells were morphologically compatible with the excised primary and recurrent tumors resected. Moreover, an immunohistochemical stain for trypsin (Meridian Life Science, Inc., Memphis, Tenn., USA) was negative, essentially ruling out the possibility of acinar cell carcinoma of the pancreas and confirming the diagnosis of metastatic AcCC to the pancreas. Perineural invasion was present. Surgical margins were negative. Multiple (24) regional lymph nodes were negative
for metastatic disease. The patient recovered well and resumed routine surveillance. To date, she has had no evidence of disease progression.

**Discussion**

To our knowledge, this is the first report of a pancreas metastasis from AcCC of the salivary gland. Despite multimodality treatment, salivary gland malignancies can recur loco-regionally or distantly. For slower-growing histologies such as AcCC, recurrences can occur after a delayed length of time, not uncommonly past the standard five-year follow-up period [5]. Therefore, lifelong surveillance and follow-up is recommended. Distant metastatic disease may occur in salivary gland malignancies, with the best data coming from studies of adenoid cystic carcinoma. The most common site of distant metastasis is the lung [6], followed by bone and liver [7, 8]. Other reported metastatic sites include the cerebrum, thyroid, and spleen [7, 8]. A report from 1979 describes a case of adenoid cystic carcinoma metastasizing to the skin [9]. Overall, metastases to the pancreas are uncommon in solid tumors, with autopsy studies showing pancreatic involvement in widely metastatic disease in 3–12% of all patients with diffuse metastatic disease [10]. The most common solid tumors to metastasize to the pancreas include carcinomas of the kidney, lung, breast, colon, stomach, esophagus, and melanoma [11]. Metastatic spread to the pancreas has not been well described for salivary cancers. As chemotherapy plays a limited role in the treatment of salivary cancers, even in the palliative setting, surgery is often a mainstay of treatment. Given the isolated metastatic site and the prolonged natural history of AcCC, surgical resection was a good therapeutic option for our patient.

**Conclusion**

Distant metastatic disease can occur years after the initial diagnosis of AcCC. Although rare, metastatic spread to the pancreas can occur and should be kept in the differential diagnosis as clinically indicated.

**Disclosure Statement**

The authors declare that there are no conflicts of interest.

**References**

Geiger et al.: Acinic Cell Carcinoma of the Salivary Gland with Metastatic Spread to the Pancreas

Fig. 1. a Intermediate-power examination of the recurrent salivary gland tumor in the left neck (H&E. ×100). The tumor cells are relatively monotonous, show round to oval nuclei, and basophilic granular cytoplasm, characteristic of well-differentiated AcCC. b Intermediate-power examination of the metastatic salivary gland tumor in the head of the pancreas (H&E. ×100). Similar to the primary salivary gland tumor, the tumor cells are relatively monotonous and exhibit basophilic granular cytoplasm that is characteristic of well-differentiated AcCC.

Fig. 2. An MRI scan with and without gadolinium with the arrow indicating a 9-mm hypoenhancing lesion in the pancreas.