A heterotopic pancreas is defined as pancreatic tissue that is lying outside its normal location and it lacks any anatomic or vascular connections with the pancreas. It is also described as pancreatic heterotopia, a term coined by Barbosa et al. (1) in 1946. The frequency of heterotopic pancreas has been estimated to be 1 case per 500 explorations of the upper abdomen or 0.6–13.7% of autopsies (2). Heterotopic pancreatic tissue may be found anywhere along the alimentary tract including the stomach, duodenum, small intestine, Meckel’s diverticulum and the biliary tract, and even in the lungs, umbilicus or fallopian tubes. Yet the most common site is the stomach and the pyloric canal. This lesion is usually asymptomatic; however, some patients have epigastric pain, upper gastrointestinal bleeding and occasional gastric outlet obstruction. In a few cases, the reported complications of heterotopic pancreas are pancreatitis, pseudocyst, cyst formation, insulinoma, adenoma and malignant transformation (3), but the exact rate of complications via a pathologic process hasn’t been identified and malignant transformation is a rare finding. Furthermore, islet cell tumor arising from heterotopic pancreas in the duodenum is extremely rare; we found only one well documented case in our review of the literature, and this was a pathologic case report (4). In this report, we describe a case of islet cell tumor arising from a heterotopic pancreas in the duodenal wall.

**Index words** : Heterotopic pancreas  
Islet cell tumor  
Duodenum

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**Case Report**

A 77-year-old woman presented to us with a poor oral intake, epigastric pain and vomiting. She was suffering from known hypertension and the rest of her medical history was unremarkable. On the routine blood tests, mild hypokalemia was noted, which was probably due to her vomiting. The other laboratory findings were normal. Endoscopy revealed a round polypoid mass in the second portion of the duodenum. Endoscopic ultrasono-
gram (EUS) revealed a mass occupying the mucosal and submucosal layers, and it showed mixed echogenicity with a central necrotic portion (Fig. 1A). Abdominal computed tomography (CT) was done. On the CT scan, a well-defined, polypoid mass was observed with intraluminal protrusion into the second portion of the duodenum (Fig. 1B), but there were no signs of bowel obstruction. The mass was well and homogeneously enhanced in the arterial and venous phases, at 125-130 HU (Fig. 1C). On the barium study, about a 2.5×2 cm sized ovoid filling defect was noted in the second portion of the duodenum and this was sharply outlined with barium. The mass showed an abrupt and slightly obtuse angulation with the duodenal wall. The overlying mucosal

Fig. 1. A. The endoscopic ultrasonogram shows a well-defined mass occupying the mucosal and submucosal layers. The mass reveals mixed echogenicity with a central necrotic portion.
B, C. On the axial CT images, the mass shows an intraluminal location in the second portion of the duodenum and it is well enhanced (HU 125-130) with relative homogeneity [solid arrows].
D. The barium study shows an ovoid filling defect with smooth surface, sharp margination and an obtuse angulation with the duodenal wall [solid arrows].
surface of the mass was intact and smoothly marginated [Fig. 1D]. Here too, there was no evidence of passage disturbance. We suspected that the mass was a duodenal polyp or a submucosal tumor such as gastrointestinal stromal tumor. The endoscopic biopsy showed chronic nonspecific inflammation and there were no tumor cells found. Duodenotomy with polypectomy was performed. At surgery, a pedunculated mass was removed from the posterior wall of the second portion of the duodenum. The specimen consisted of a round mass of polypoid soft tissue that measured 1.8×1.5×1 cm. The mucosal and submucosal layers were covering it. The cut section of the specimen showed a relatively defined, yellowish solid submucosal tumor that measured 1.8×1.2×1 cm. Microscopically, the mass had infiltrated the mucosa and submucosa layers and it had extended to the proper muscle. The low magnification photomicrograph demonstrated a solid islet cell tumor located in the more superficial portion as well as a heterotopic pancreas in deep muscular layer [Fig. 1E]. The high magnification photomicrograph showed uniform round tumor cells in a trabecular or acinar arrangement with

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**Fig. 1.**
E. Low magnification photomicrograph of the mass demonstrates a solid islet cell tumor located at the more superficial portion of the heterotopic pancreas in the deep muscular layer of the duodenum (solid arrows).
F. High magnification photomicrograph shows the uniform round tumor cells in a trabecular or acinar arrangement with amyloid deposition (asterisks).
G. The tumor cells are diffusely stained with anti-chromogranin A antibody with a strong intensity.
H. Electron micrograph shows the secretory granules in the tumor cell cytoplasm and the amyloid fibrils (asterisks).
amyloid deposition (Fig. 1F). The tumor cells were diffusely stained with anti-chromogranin A antibody with a strong intensity. This suggested that the tissue consisted of chromogranin A-positive neuroendocrine cell nests (Fig. 1G). Also, the sections showed amyloid deposition in the tumorous stroma. Amyloid was identified in the tumor focus where the cells were stained for anti-insulin antibody. Several studies have revealed that insulinomas commonly express islet amyloid polypeptide (IAPP) and in approximately 5% of these cases, IAPP may be precipitated as amyloid in the tumor stroma (1). Electron microscopy was carried out and the further histologic examination of the resected tumor demonstrated many closely packed round to ovoid cells with abundant cytoplasmic organelles. Many membrane-bound neurosecretory granules and a disordered meshwork of nonbranching fibrils were noted in the cytoplasm. Fibrillar depositions were also noted in the stroma (Fig. 1H). These features were all compatible to islet cell tumor with amyloid deposition arising from a heterotopic pancreas.

**Discussion**

Heterotopic pancreas is thought to arise at the time of embryonic development and during fusion of the pancreatic buds. The alimentary tract is in close proximity to the developing pancreas; thus, tissue can become implanted in the bowel wall and then it’s carried to its final location (5). The aberrant tissue consists of endoderm that is composed of all the cell types normally found in the pancreas, and it often includes islets of Langerhans. Consequently, pancreatitis with fat necrosis, islet cell tumors with hyperinsulinism and pancreatic carcinoma with metastases may occur in heterotopic pancreas (6). Symptoms rarely come into existence with the presence of heterotopic pancreatic tissue alone. This is usually an asymptomatic condition that is found incidentally during laparotomy or at autopsy in the stomach, duodenum and small intestine, and it is even found in Meckel’s diverticulum or in the biliary tract. Heterotopic pancreatic tissue is most commonly found in the gastric antrum along the greater curvature. In our patient, the tumor was located in the duodenum, a second most common location for heterotopic pancreas. The frequency of heterotopic pancreas has been estimated to be 1 case per 500 explorations of the upper abdomen or 0.6-13.7% of autopsies (1). It is generally diagnosed with the development of complications such as hemorrhage, obstruction and malignant transformation (7). However, malignant transformation in heterotopic pancreatic tissue is a rare finding, and a review of literature revealed only 15 well-documented cases of carcinoma arising in heterotopic pancreas (2). Islet cell tumor arising from heterotopic pancreas is also extremely rare. We found only one report of a case of an islet cell tumor arising from a heterotopic pancreas in our review of literature. Tolentino et al (4) reported one case in 2003 and that was a pathologic report.

The radiologic diagnosis of a pancreatic tumor is usually made on the basis of the typical location and appearance, which is that of a small umbilicated lesion in a submucosal mass (3). However, this appearance may be a nonspecific finding for diagnosis. That is, other submucosal tumors such as gastrointestinal stromal tumor, lymphoma, carcinoid or adenomatous polyp can reveal as being small umbilicated lesions such as an overlying mucosal ulceration. Besides, it is more difficult to distinguish islet tumor originating from a heterotopic pancreatic tissue from the other submucosal tumors, and there are no reported specific findings about islet cell tumor arising from heterotopic pancreas. In our case, on the initial EUS and barium study, we suspected a submucosal tumor such as gastrointestinal stromal cell tumor or duodenal polyp. Furthermore, CT was not helpful for the diagnosis. The CT findings were merely a well defined, intraluminal mass in the duodenum. If there is one thing to be considered, it is that the mass was enhanced at 125-130 Housfield units (HU), and that was higher than the HUs of the normal enhancing pancreas, which showed an average of 100 HU in the venous phase of our case. Typically, but not always, islet tumor in a normal pancreas shows intense enhancement in the arterial and venous phase.

Islet cell tumors are classified as functioning if they produce symptoms related to excessive hormone production; intractable hypoglycemia, low blood levels of glucose and high circulating plasma insulin (8). Nonfunctioning tumors are those pancreatic tumors with endocrine differentiation in the absence of a clinical syndrome related to hormone production. In our case, GI symptoms such as vomiting and epigastric pain were presented, but the blood hormone levels were normal. On the pathology, the uniform round tumor cells that were diffusely stained with anti-chromogranin A, which suggested that this tissue had differentiated into chromogranin A-positive neuroendocrine cells, and they were located in the more superficial portion of the mass.
and in deep muscular layer where the heterotrophic pancreas was based. Also, the sections showed amyloid deposition in the tumorous stroma. Amyloid was identified in the tumor focus where the cells were stained for anti-insulin antibody. So our case was a symptomatic nonfunctioning islet cell tumor arising from a heterotopic pancreas in the duodenal wall.

Although malignant transformation of heterotopic pancreas, including islet cell tumor, is extremely rare, it remains an important consideration in the differential diagnosis for duodenal submucosal masses. We have demonstrated the radiologic appearance and clinical-pathologic findings of a rare case of islet cell tumor arising from a heterotopic pancreas in the duodenal wall.

References
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