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**Interventional endoscopic ultrasound for a symptomatic pseudocyst secondary to gastric heterotopic pancreas**

Jin HB *et al*. Heterotopic pancreas causing gastric outlet obstruction

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**Abstract**

Heterotopic pancreas (HP) is a relatively uncommon entity that is defined as pancreatic tissue without a true anatomical or vascular connection to the pancreas. HP does not cause symptoms in most cases but can occasionally produce various manifestations, including nausea, vomiting, abdominal pain, and even heterotopic pancreatitis. Here, we report an unusual case in which heterotopic pancreatitis complicated by the formation of a pseudocyst that caused gastric outlet obstruction was diagnosed based on serum hyperamylasemia and findings from endoscopic ultrasonography (EUS)-guided fine needle aspiration (EUS-FNA) cytology. EUS-guided single pigtail stent insertion was successfully performed for recurrent gastric outlet obstruction. The patient has remained healthy and symptom-free during 4 years of surveillance. In the context of relevant literature, the described case is a rare case of HP complicated by a pseudocyst treated *via* EUS-FNA and stent insertion.

**Key words:** heterotopic pancreas; pseudocyst; gastric outlet obstruction; endoscopic ultrasound-guided fine needle aspiration

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**Core tips:** Heterotopic pancreas (HP), which is defined as the atypical presence of pancreatic tissue without an anatomical or vascular connection to the pancreas, is relatively rare. As an intra- or submucosal lesion, HP is typically asymptomatic and found incidentally. We report a case involving a 40-year-old male with gastric outlet obstruction secondary to pseudocyst formation in HP.

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**INTRODUCTION**

Heterotopic pancreas (HP) is defined as the presence of pancreatic tissue in an abnormal location without anatomical or vascular connections with the orthotopic pancreas. HP was first described by Jean-Schultz in 1729 and was confirmed histologically by Klobb[[1](file:///E:\\==科室工作相关资料==\\==投稿==\\金航斌\\1\\34943_修改版本.docx" \l "_ENREF_1" \o "Ulrych, 2015 #37)]. Other terms for HP, such as ectopic pancreas, aberrant pancreas and accessory pancreas, have been used in the literature[[2](file:///E:\==科室工作相关资料==\==投稿==\金航斌\1\34943_修改版本.docx#_ENREF_2)]. In most cases, HP is asymptomatic and found incidentally[[1](file:///E:\\==科室工作相关资料==\\==投稿==\\金航斌\\1\\34943_修改版本.docx" \l "_ENREF_1" \o "Ulrych, 2015 #37)], but it can occasionally produce non-specific symptoms that depend on its location, its size and types of complications. Here, we describe a case involving a 40-year-old male with gastric outlet obstruction secondary to heterotopic pancreatitis with pseudocyst formation.

**CASE REPORT**

A 40-year-old male was admitted to a community hospital for upper abdominal pain accompanied by nausea and vomiting. He also complained of abdominal distention and moderate fever (38.3 ℃). Physical examination revealed minimal tenderness in the epigastric region without a palpable mass or peritoneal signs, and the patient’s vital signs were unremarkable. Routine blood tests showed significant elevation of the leukocyte count and neutrophil ratio, and biochemical examination revealed serum amylase of 460 U/L and urine amylase of 3500 U/L. Transabdominal ultrasound was initially performed when the patient was diagnosed with pancreatitis, but no gallbladder stones or common bile duct stones were detected. The patient was treated for acute pancreatitis; his treatment included fasting, injection of a proton pump inhibitor (PPI) and somatostatin. All symptoms disappeared 3 days later but recurred after the patient ingested fluids. Subsequently, the patient was transferred to our hospital for further diagnosis. A computed tomography (CT) scan of the abdomen revealed manifestations of a normal pancreas and an ill-defined enhancing mass at the gastric antrum that was causing circumferential narrowing of the pyloric channel (Figure 1A). In an esophagogastroduodenoscopy (EGD) examination, a sub-epithelial lesion without overlying umbilication or dimpling was found in the prepyloric region (Figure 1B). The pyloric channel was only wide enough to allow for the passage of an Olympus Q260 endoscope (OD 9.2 mm), but the duodenum was normal. Radial endoscopic ultrasonography (EUS), which was used as an additional diagnostic tool, confirmed the presence of a cystic lesion measuring 21 mm × 25 mm (Figure 1C) originating from the third layer with anechoic contents and debris. Subsequently, EUS-guided fine needle aspiration (EUS-FNA) was successfully performed (Figure 1D), and a total of 25 ml of clear fluid was aspirated to the greatest possible extent with negative pressure. Cystic fluid analysis revealed an amylase level of 83770 U/L and a carcinoembryonic antigen level of 3.7 µg/L; a cytologic examination of smears revealed clusters of benign-appearing ducts and small acini mixed with inflammatory cells (Figure 2).

The patient was allowed to ingest low-fat fluids 5 days after the procedure, but the aforementioned symptoms immediately re-emerged, and serum amylase increased to 512 U/L. A narrowed pyloric channel was observed in a second EGD examination. The patient refused surgical treatment. Balloon dilation, bougie dilation and needle-knife puncture of the cyst were not initially selected. In accordance with an endoscopic drainage procedure used for pancreatic pseudocysts, we successfully performed cyst cavity (20 mm × 23 mm) puncture, wire exchange, and the gradual insertion of a single pigtail stent (5 Fr × 4 cm) using a 19-gauge needle (EchoTip Ultra, ECHO-19; Cook Endoscopy, Winston-Salem, NC, United States) (Figure 1E and F). Informed consent was obtained from the patient and his family prior to this endoscopic procedure. No abnormal signs or symptoms were noted during or immediately after the procedure. Shortly thereafter, the patient’s abdominal pain and vomiting subsided, and his leukocyte count and serum amylase returned to normal. The patient was discharged on the sixth day after the second drainage procedure.

Five months later, the patient returned for a follow-up examination without any complaints. EGD surveillance revealed no stent at the original position, and radial EUS confirmed the presence of a smaller anechoic cystic lesion measuring 3 mm × 5 mm (Figure 3A and B). The stent had completely passed, as confirmed by abdominal radiography (Figure 4). The patient has remained healthy and symptom-free during 4 years of follow-up.

**DISCUSSION**

In autopsy studies, HP has generally had an incidence ranging from 0.5%-13.0% and been more common in 30- to 50-year-old subjects, with male predominance[[3](file:///E:\==科室工作相关资料==\==投稿==\金航斌\1\34943_修改版本.docx#_ENREF_3)]. HP is most frequently located in the stomach (25%-38%), followed by the duodenum (17%-36%) and the jejunum (15%-21.7%); it is rarely found in the esophagus, common bile duct, spleen, mesentery, mediastinum or fallopian tube[[4](file:///E:\==科室工作相关资料==\==投稿==\金航斌\1\34943_修改版本.docx#_ENREF_4)]. In the gastrointestinal tract, HP is most frequently located only in the submucosa (54%) and is less frequently found in both the submucosa and muscularis propria (23%), only the muscularis propria (8%), the subserosa (11%), and the whole wall (4%)[1].

HP has been classified into 4 types by Heinrich and Gaspar-Fuentes[5]. Type I HP shows the presence of ducts, acini and islets of Langerhans cells; Type II HP shows only ducts and acini; Type III HP has only ducts; and Type IV HP has only islets. Most cases of HP are Type Ⅱ. The pathogenesis of HP remains unclear, although the two most widely accepted hypotheses are the misplacement and metaplasia theories. The misplacement theory proposes that ectopic pancreatic tissue develops from misplacement during foregut rotation, whereas the metaplasia theory suggests that heterotopia arises from a pancreatic metaplasia of the endoderm that migrates to the submucosa during embryogenesis[[6](file:///E:\==科室工作相关资料==\==投稿==\金航斌\1\34943_修改版本.docx#_ENREF_6)].

### HP is typically asymptomatic; however, depending on its location and size, HP can occasionally cause clinical symptoms, the most common of which are vomiting and abdominal pain. Peptic ulceration and gastrointestinal bleeding are rare presentations, as are malignant transformation, pancreatitis, pseudocyst formation, gastric outlet obstruction, jejunal intussusception, and ileus[[3](file:///E:\==科室工作相关资料==\==投稿==\金航斌\1\34943_修改版本.docx#_ENREF_3),7-10]. A thorough search of the English literature revealed 5 cases of gastric heterotopic pancreatitis in the last 20 years, including our case[11-14] (Table 1). These cases involved 3 male patients and 2 female patients. The median age of the patients was 32 years. Epigastric pain was reported in 4 cases, and vomiting occurred in 3 cases. In our case, the patient presented with recurrent epigastric pain and vomiting, which were mainly induced by gastric outlet obstruction induced by heterotopic pancreatitis. The symptoms of pain and fever in our case might have been attributable to inflammation or chemical irritation of the involved tissue and enzyme secretion.

It is typically complex and challenging to preoperatively diagnose HP due to this condition’s varied presentations. Gastric HP is difficult to radiologically diagnose; however, the double contrast barium meal technique may characteristically show a focally raised mucosal area with associated superficial ulceration within the gastric antrum[15]. CT can localize lesions with normal pancreatic tissue but cannot distinguish HP from other submucosal tumors[1[6](file:///E:\==科室工作相关资料==\==投稿==\金航斌\1\34943_修改版本.docx#_ENREF_7)]. Upper endoscopy generally reveals a broad-based, umbilicated, submucosal lesion, with superficial biopsies being non-diagnostic in the vast majority of cases[1[7](file:///E:\==科室工作相关资料==\==投稿==\金航斌\1\34943_修改版本.docx#_ENREF_8)]. EUS has been reported to be helpful in diagnosing HP; EUS-FNA in particular is especially effective for confirming a diagnosis of HP, with a reported sensitivity of 80%-100%[1[8](file:///E:\==科室工作相关资料==\==投稿==\金航斌\1\34943_修改版本.docx#_ENREF_9),19]. A novel sampling method using submucosal endoscopy with a mucosal flap was recently reported by Kobara *et al*[20]. Submucosal dissection allows for the collection of tissue samples even from small lesions (< 2 cm in diameter) under direct endoscopic view. This method was used to eventually prove that a gastrointestinal submucosal tumor (SMT) was a gastric duplication cyst with HP and an ectopic submucosal gland[21]. This approach can be used for preoperative diagnosis of an SMT, albeit with certain limitations.

Symptomatic HP should be respected. However, the optimal management of asymptomatic HP remains debatable. Surgical resection has been reported in most HP cases[11-14]. In the aforementioned cases (Table 1), four patients underwent partial resection of the stomach because this approach was preferable for the preservation of gastric function. The prognosis after surgery was generally favorable, and complete symptom remission was achieved in these cases. Endoscopic submucosal dissection (ESD) and endoscopic mucosal resection (EMR) have also been reported for symptomatic HP; these approaches offer the possibility of localized treatment for HP with relatively low mortality and morbidity rates[22]. Zhong *et al*[23] reported 60 cases of gastric HP that were treated using ESD or EMR; in particular, 14 patients underwent EMR, and 46 patients underwent ESD. Complications occurred in 12 patients (20%), including arterial bleeding in 9 patients and pneumoperitoneum in 3 patients; all of these patients were treated conservatively. However, it has been reported that in total, 23% of gastric HP are located in the muscularis propria, the subserosa, or the whole wall; the technical difficulty and complication rates of endoscopic procedures may be significantly increased for HP at these locations[1]. Recently, EUS has been increasingly used for pancreatic fluid collection, biliary and pancreatic duct drainage, gallbladder drainage, and other applications[24]. The roles of interventional EUS continue to expand, and growing global experience supports the feasibility and efficacy of this approach[25,26]. In our case, an endoscopic view of the lesion did not reveal the typical umbilicated indentation that has been recognized as characteristic of a central ductal orifice. The absence of communication between the glandular epithelium and the gastric lumen can be attributed to the retention of exocrine secretions that subsequently led to recurrent episodes of acute obstructive pancreatitis, the formation of a retention pseudocyst and eventually recurrent gastric outlet obstruction. EUS-guided stenting was used to relieve the gastric outlet obstruction by creating an artificial fistula in the pseudocyst. Consequently, this stenting not only rapidly relieved clinical symptoms but also resulted in the long term in the conversion of symptomatic HP to asymptomatic HP that no longer required surgical treatment. To our knowledge, no previous reports have described heterotopic pancreatitis treated using interventional EUS.

In summary, this case demonstrates the use of a novel endoscopic approach to successfully manage a gastric outlet obstruction secondary to heterotopic pancreatitis with pseudocyst formation. Based on this case, we suggest that HP should be considered in the differential diagnosis of a submucosal gastric mass. EUS-guided intervention can be beneficial for both diagnosing and providing symptomatic relief of heterotopic pancreatitis with pseudocyst formation, and endoscopic treatment should be recommended in selected cases.

**COMMENTS**

***Case characteristics***

A 40-year-old male complained of upper abdominal pain, nausea and vomiting.

***Clinical diagnosis***

Heterotopic pancreatitis with pseudocyst formation located in the prepyloric region causing gastric outlet obstruction.

***Differential diagnosis***

Hyperplastic polyp, gastric adenoma, gastrointestinal stromal tumor (GIST), lymphoma.

***Laboratory diagnosis***

Laboratory workup revealed significant elevation of the leukocyte count and neutrophil ratio, serum amylase of 460 U/L and urine amylase of 3500 U/L.

***Imaging diagnosis***

The abdominal computed tomography scan showed an ill-defined enhancing mass at the gastric antrum. Endoscopic ultrasound (EUS) revealed a cystic lesion measuring 21 mm × 25 mm originating from the third layer with anechoic contents and debris.

***Pathological diagnosis***

Histology revealed heterotopic pancreatic tissue in the prepyloric mucosa with clusters of benign-appearing ducts and small acini mixed with inflammatory cells.

***Treatment***

EUS-guided stenting.

***Related reports***

A total of 4 cases of gastric heterotopic pancreatitis have been reported in the literature. However, there is the first report of heterotopic pancreatitis with pseudocyst formation treated using interventional EUS.

***Experiences and lessons***

Heterotopic pancreas (HP) is a rare entity, and should be considered in the differential diagnosis of a submucosal gastric mass. Although surgical resection has been reported in most symptomatic HP cases, interventional EUS can sometimes be an effective minimally invasive treatment procedure for HP.

***Peer-review***

The authors describe an interesting and rare case of heterotopic pancreas complicated with pseudocyst. Heterotopic pancreatitis with pseudocyst formation has been reported previously, but this case is unique in that endoscopic drainage was performed which has not been previously described to our knowledge.

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Grade D (Fair): 0

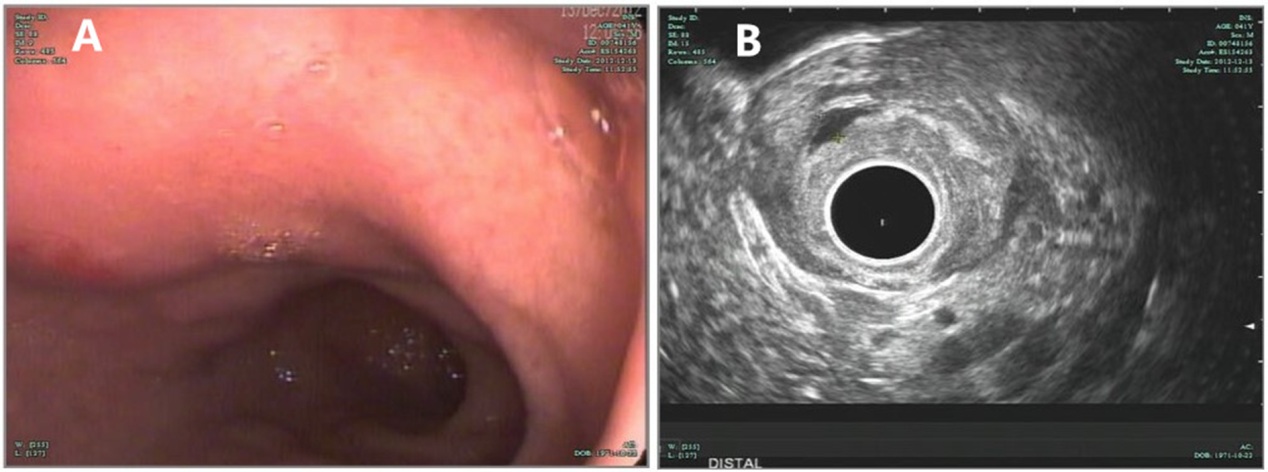
Grade E (Poor): 0



**Figure 1 Computed tomography scan and endoscopic treatment of the lesion.** A: Contrast-enhanced CT revealed manifestations of a normal pancreas and an ill-defined enhancing mass (arrows) at the gastric antrum that caused circumferential narrowing of the pyloric channel; B: EGD showed a narrowed pyloric channel with normal mucosa; C: Radial EUS imaging confirmed the presence of a cystic lesion measuring 21 mm × 25 mm that originated from the third layer with anechoic contents and debris; D: EUS-guided fine needle aspiration, D: Cyst cavity radiography and guide-wire exchange; E: Successful insertion of a single pigtail stent (5 Fr × 4 cm). CT: computed tomography; EUS: endoscopic ultrasonography; EGD: esophagogastroduodenoscopy .



**Figure 2** **Cytologic examination with HE staining (low-power field).** A cytologic examination of smears revealed clusters of benign-appearing ducts and small acini mixed with inflammatory cells.

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**Figure 3 esophagogastroduodenoscopy and endoscopic ultrasonography surveillance 5 mo later.** A: EGD surveillance 5 mo after stent insertion; B: Radial EUS confirmed the presence of a small anechoic cystic lesion measuring 3 mm × 5 mm. EUS: endoscopic ultrasonography; EGD: esophagogastroduodenoscopy .



**Figure 4 Abdominal radiograph.** A normal abdominal radiograph confirmed excretion of the stent without stent impaction.

**Table 1 Reported cases of gastric heterotopic pancreatitis**

|  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| **Author** | **Year** | **Age/ gender** | **Symptom(s)** | **Size (mm)** | **Location** | **Serum amylase (IU/L)** | **Preoperative diagnosis** | **Diagnostic method** | **Therapy** |
| Matsushita | 1997 | 33/M | Epigastric pain | 20 | N/A | 59 | SMT | Surgery | Enucleation |
| Hirasaki | 2005 | 32/M | Epigastric pain | 35 | Angulus | 262 | SMT | Surgery | Partial resection |
| Matissek | 2012 | 15/F | Vomiting | 30 | Pylorus | 144 | SMT | Surgery | Gastroduodenostomy |
| Matsumoto | 2012 | 21/F | Epigastric pain and vomiting | 26 | Antrum | 122 | HP | EUS-FNA | Laparoscopic partial resection |
| Our case | 2017 | 40/M | Epigastric pain and vomiting | 25 | Pylorus | 460 | HP | EUS-FNA | EUS-guided stenting |

EUS: endoscopic ultrasonography; FNA: fine needle aspiration; HP: heterotopic pancreas; N/A: not available; SMT: submucosal tumor.