Case Report

A rare collision tumor of gastrointestinal stromal tumor of the stomach and pancreatic ductal adenocarcinoma

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Abstract: Collision tumors involving GIST are very rare. Herein, we report a collision tumor consisting of gastric gastrointestinal stromal tumor (GIST) and pancreatic ductal adenocarcinoma. The patient was a 76-year-old Chinese male, presenting with abdominal distension, haematemesis and melena. Both the CT scan images and the gross appearance demonstrated special features indicating the possibility of a tumor with two different elements, which were once ignored by the initial diagnosticians. Microscopic and immunohistochemical features identified a collision tumor involving GIST and pancreatic ductal adenocarcinoma. To our best, this is the first such a case reported in the English literature.

Keywords: Collision tumor, pancreatic ductal adenocarcinoma, GIST, diagnostic pitfall

Introduction

Gastrointestinal stromal tumors (GISTs) are the most common mesenchymal tumors of the gastrointestinal tract. Although synchronous or metachronous occurrence of separate GIST and other tumors is not uncommon [1-15], with a few cases of synchronous GIST and pancreatic neoplasms [1, 2, 9-15], collision tumors involving GIST are very rare [1, 16-22]. Herein, we report a collision tumor of gastric GIST and pancreatic ductal adenocarcinoma (PDAC), which provided diagnostic pitfalls due to its rarity and the morphologic properties of the two components involved in it.

Case presentation

A 76-year-old Chinese male presented with abdominal distension for one month and haematemesis with melena for two days. Physical examination revealed an upper abdominal mass. No significant past medical history was noted. The upper gastrointestinal endoscopy showed a raised nodular mass at the fundus of the stomach. Biopsy was performed, but no evidence of neoplasm was obtained. Abdominal CT scan demonstrated a large ill-defined inhomogeneous mass located in the body of the

pancreas (**Figure 1**). The patient underwent a palliative tumor resection with distal pancreatectomy, partial gastrectomy and splenectomy in May 2013.

Grossly, there was a bulky, heterogeneous mass up to 10 cm involving both the pancreas and the stomach. The portion involving the pancreas was a poorly defined, infiltrative mass with a solid and firm appearance, and the cut surface was grayish yellow mixed with white in color. The portion attaching to the gastric wall by a narrow pedicle was a well-circumscribed 6 cm nodule with a fleshy, pink-tan cut surface (Figure 2). These two portions merged together into a single mass, and no definite boundary was seen.

Microscopically, the pancreatic part of the tumor revealed a diffuse infiltration of haphazardly arranged dysplasia glands involving the parenchyma of the pancreas with a desmoplastic reaction of the spindle-shaped myofibroblastic cells, suggestive of classic conventional PDAC (Figure 3A). However, the gastric part was composed of fasciculated arranged spindle cells, which exhibited minimal atypia with delicate cytoplasmic processes and fine chromatin. The spindle cells demonstrated a syncytial

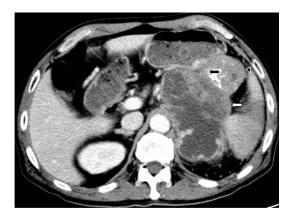


Figure 1. Contrast-enhanced CT scan showed a mass arising from the body and tail of the pancreas (white arrow). Note the relatively well-defined nodule (black arrowhead) with multiple calcifications, which adhered closely to the stomach. The boundary between the mass and the nodule could be delineated (black arrow).

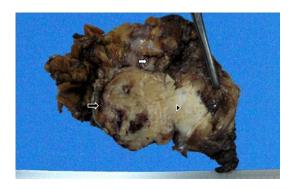


Figure 2. The collision tumor consisting of PDAC (arrowhead) and GIST (black arrow), which was attaching to the gastric wall by a narrow pedicle (white arrow).

appearance. Mitoses of these spindle cells numbered 3 per 5 mm². Fibrosis and calcification were seen in some areas. Notably, in the intermingled part of the mass, the malignant glands were found infiltrating into the spindle cells of the gastric part (Figure 3B and 3C). Immunohistochemically, the adenocarcinoma cells were positive for cytokeratin AE1/AE3, while the reactive myofibroblastic cells were only positive for smooth muscle actin. Meanwhile, the spindle cells of the gastric part were positive for CD117 (Figure 3D), CD34, smooth muscle actin, and negative for DOG-1, desmin, S-100 protein and cytokeratin AE1/ AE3. Molecular genetic test was performed on the gastric spindle-cell portion and KIT exon 11 mutation (Q556_V559delinsH) was observed.

Based on the aforementioned findings, the final diagnosis was a collision tumor of an advanced moderately differentiated PDAC and a low-risk gastric GIST [23].

The patient died 3 months after the surgery.

Discussion

Collision tumors consisting of GIST and other neoplasms have rarely been reported. To our best knowledge, there were less than 20 cases reported in the English literature [1, 16-22]. The most common cases reported were gastric tumors involving GIST and adenocarcinoma [16-19], while rare cases included GIST with inflammatory myofibroblastic tumor in a single gastric polypoid mass [20], gastric tumor involving collision of GIST and angiosarcoma [21], and rectal GIST involving prostatic adenocarcinoma [22]. Some other reported collision tumors involving GIST were actually composed of a gastric GIST and a closely associated tumor without intermingling with each other [24-27]. The current case is the first published example describing a collision tumor involving gastric GIST and PDAC.

Collision tumors could increase the complexity of the diagnosis and treatment process. Due to the rarity, it's unlikely for diagnosticians to be aware of a collision tumor initially. As in this case, neither the radiological report nor the initial macroscopical description of the tumor gave any suggestion to the possibility of a collision tumor at the very beginning. When a diagnosis of collision tumor was proposed based on the histology, both the CT scan and the gross specimen were reviewed. Actually, both of them did suggest a possible collision tumor when examined closely (Figures 1 and 2). In addition, the histological features of the two components in the current case may be confusing and also contribute to the diagnostic pitfalls. As we know, the extensive desmoplastic reaction of myofibroblasts is a property for most PDACs. So, when sampling was insufficient or only a biopsy was performed, the spindle tumor cells of the GIST here might be easily interpreted as the florid desmoplastic stroma of the PDAC. On the other hand, the small cluster of well differentiated carcinoma glands infiltrating into the GIST might be interpreted as non-neoplastic glands entrapped in the GIST as well [16]. Either situation could lead to a missed diagnosis of

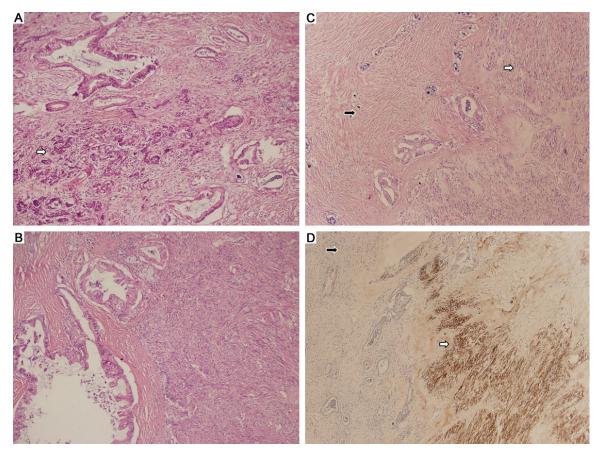


Figure 3. A. The PDAC with desmoplastic stromal reaction in the parenchyma of pancreas (white arrow) (\times 100). B and C. At the merging interface of collision tumor. The adenocarcinoma glands were infiltrating into the GIST (\times 100). C and D. The tumor cells (white arrows) of GIST were strongly positive for CD117, while the reactive myofibroblastic cells (black arrows) were negative (\times 100).

one of the components of the collision tumor. Besides, there are also some other uncommon lesions which demonstrate a biphasic histological feature and should be considered in the differential diagnostic list, including carcinosarcoma, pancreatic mucinous cystic neoplasm with sarcomatous stroma [28] and primary phylodes tumor of the pancreas [29]. Therefore, sufficient sampling, attentive observation of the morphologic traits, essential differential immunohistochemical markers, and careful clinicopathological correlation should be required and would be much helpful in confirming the diagnosis.

PDACs are much more aggressive than GISTs, with the mean survival time of 3-5 months for untreated patients. So, it was the PDAC in this case that determined the prognosis of the patient.

In summary, we report a rare collision tumor of gastric GIST and PDAC, which presented with

special radiological and macroscopic features once ignored by the diagnosticians. The final diagnosis has been based on the careful review of the clinical, histopathological, immunohistochemical and molecular features of the tumor.

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Disclosure of conflict of interest

None.

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References

- [1] Agaimy A, Wunsch PH, Sobin LH, Lasota J and Miettinen M. Occurrence of other malignancies in patients with gastrointestinal stromal tumors. Semin Diagn Pathol 2006; 23: 120-129.
- [2] Pandurengan RK, Dumont AG, Araujo DM, Ludwig JA, Ravi V, Patel S, Garber J, Benjamin RS, Strom SS and Trent JC. Survival of patients with multiple primary malignancies: a study of 783 patients with gastrointestinal stromal tumor. Ann Oncol 2010; 21: 2107-2111.
- [3] Shen C, Chen H, Yin Y, Chen J, Han L, Zhang B, Chen Z and Chen J. Synchronous occurrence of gastrointestinal stromal tumors and other digestive tract malignancies in the elderly. Oncotarget 2015; 6: 8397-8406.
- [4] Goncalves R, Linhares E, Albagli R, Valadao M, Vilhena B, Romano S and Ferreira CG. Occurrence of other tumors in patients with GIST. Surg Oncol 2010; 19: e140-143.
- [5] Adim SB, Filiz G, Kanat O and Yerci O. Simultaneous occurrence of synchronous and metachronous tumors with gastrointestinal stromal tumors. Bratisl Lek Listy 2011; 112: 623-625.
- [6] Sevinc A, Seker M, Bilici A, Ozdemir NY, Yildiz R, Ustaalioglu BO, Kalender ME, Dane F, Karaca H, Gemici C, Gumus M and Buyukberber S. Co-existence of gastrointestinal stromal tumors with other primary neoplasms. Hepatogastroenterology 2011; 58: 824-830.
- [7] Giuliani J and Bonetti A. The occurrence of gastrointestinal stromal tumors and second malignancies. J Gastrointest Cancer 2015; 46: 408-412.
- [8] Wronski M, Ziarkiewicz-Wroblewska B, Gornicka B, Cebulski W, Slodkowski M, Wasiutynski A and Krasnodebski IW. Synchronous occurrence of gastrointestinal stromal tumors and other primary gastrointestinal neoplasms. World J Gastroenterol 2006; 12: 5360-5362.
- [9] Liu YJ, Yang Z, Hao LS, Xia L, Jia QB and Wu XT. Synchronous incidental gastrointestinal stromal and epithelial malignant tumors. World J Gastroenterol 2009; 15: 2027-2031.
- [10] Liszka L, Zielinska-Pajak E, Pajak J, Golka D and Huszno J. Coexistence of gastrointestinal stromal tumors with other neoplasms. J Gastroenterol 2007; 42: 641-649.
- [11] Arnogiannaki N, Martzoukou I, Kountourakis P, Dimitriadis E, Papathanasaki A, Nastoulis E, Gazalidou M, Fida A, Apostolikas N and Agnantis NJ. Synchronous presentation of GISTs and other primary neoplasms: a single center experience. In Vivo 2010; 24: 109-115.
- [12] Dasanu CA, Mesologites T and Trikudanathan G. Synchronous tumors: adenosquamous car-

- cinoma of pancreas and GIST of stomach. J Gastrointest Cancer 2011; 42: 186-189.
- [13] He JJ, Ding KF, Zheng L, Xu JH, Li J, Wu YL, Sun LF, Zhou DE and Zheng S. Adenosquamous carcinoma of the uncinate process of the pancreas with synchronous gastrointestinal stromal tumor of the stomach: Case report and review of the literature. Oncol Lett 2012; 4: 1191-1194.
- [14] Alabraba E, Bramhall S, O'Sullivan B, Mahon B and Taniere P. Pancreatic insulinoma co-existing with gastric GIST in the absence of neurofibromatosis-1. World J Surg Oncol 2009; 7: 18.
- [15] Tavares AB, Viveiros FA, Cidade CN and Maciel J. Gastric GIST with synchronous neuroendocrine tumour of the pancreas in a patient without neurofibromatosis type 1. BMJ Case Rep 2012; 2012.
- [16] Kleist B, Lasota J and Miettinen M. Gastrointestinal stromal tumor and gastric adenocarcinoma collision tumors. Hum Pathol 2010; 41: 1034-1039.
- [17] Liu SW, Chen GH and Hsieh PP. Collision tumor of the stomach: a case report of mixed gastrointestinal stromal tumor and adenocarcinoma. J Clin Gastroenterol 2002; 35: 332-334.
- [18] Bi R, Sheng W and Wang J. Collision tumor of the stomach: gastric adenocarcinoma intermixed with gastrointestinal stromal tumor. Pathol Int 2009; 59: 880-883.
- [19] Katsoulis IE, Bossi M, Richman PI and Livingstone JI. Collision of adenocarcinoma and gastrointestinal stromal tumour (GIST) in the stomach: report of a case. Int Semin Surg Oncol 2007; 4: 2.
- [20] Shin HC, Gu MJ, Kim SW, Kim JW and Choi JH. Coexistence of gastrointestinal stromal tumor and inflammatory myofibroblastic tumor of the stomach presenting as a collision tumor: first case report and literature review. Diagn Pathol 2015; 10: 181.
- [21] Adhikari M, Wu ML and Zhao X. Gastrointestinal stromal tumor colliding with angiosarcoma. Int J Surg Pathol 2006; 14: 252-256.
- [22] Macias-Garcia L, De la Hoz-Herazo H, Robles-Frias A, Pareja-Megia MJ, Lopez-Garrido J and Lopez JI. Collision tumour involving a rectal gastrointestinal stromal tumour with invasion of the prostate and a prostatic adenocarcinoma. Diagn Pathol 2012; 7: 150.
- [23] Miettinen M and Lasota J. Gastrointestinal stromal tumors: review on morphology, molecular pathology, prognosis, and differential diagnosis. Arch Pathol Lab Med 2006; 130: 1466-1478.
- [24] Ozgun YM, Ergul E, Sisman IC and Kusdemir A. Gastric adenocarcinoma and GIST (collision tumors) of the stomach presenting with perforation; first report. Bratisl Lek Listy 2009; 110: 504-505.

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- [25] Qian T, Gao F, Chen MZ, Meng FH, Li XJ, Liu YJ and Yin HB. Collision tumor of the esophagus: report of a case with mixed squamous cell carcinoma and gastrointestinal stromal tumor. Int J Clin Exp Pathol 2014; 7: 1206-1211.
- [26] Toyoda A, Komaba A, Yoshizumi H, Hanaoka R, Sakuma S, Ichinohe A, Kawana H and Harigaya K. Collision of advanced gastric adenocarcinoma and gastrointestinal stromal tumour: a case report. BMJ Case Rep 2009; 2009.
- [27] Idema DL, Daryanani D, Sterk LM and Klaase JM. Collision tumor of the stomach: a case of an adenocarcinoma and a gastrointestinal stromal tumor. Case Rep Gastroenterol 2008; 2: 456-460.
- [28] Wayne M, Gur D, Ascunce G, Abodessa B and Ghali V. Pancreatic mucinous cystic neoplasm with sarcomatous stroma metastasizing to liver: a case report and review of literature. World J Surg Oncol 2013; 11: 100.
- [29] Hirabayashi K, Fujihira T, Oyamada H, Serizawa A, Yamashita T, Tobita K, Imaizumi T, Kajiwara H, Nakamura N and Osamura RY. First case of primary phyllodes tumor of the pancreas: case report and findings of immunohistochemical and ultrastructural studies. Virchows Arch 2010; 456: 587-593.