Lung adenocarcinoma isolated metastasizing to pancreas a case report and review

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Case report

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Abstract

BACKGROUND: Lung cancers are prone to metastasis. It is commonly associated with metastases in the brain, bone, and liver. However, isolated pancreatic metastasis are extremely rare in lung cancer patients.

CASE SUMMARY: A Chinese man underwent pulmonary lobectomy for adenocarcinoma 6 years ago. He was referred to our department because postoperative reexamination revealed space occupying lesions in the neck of the pancreas. Abdomen CT showed a space occupying lesion in the neck of the pancreas, and no abnormality was found in any other organ. Therefore, it was considered to pancreatic carcinoma. Pancreatectomy of the pancreatic body and tail, as well as splenectomy, were performed, and he was given four courses of pemetrexed and Carboplatin chemotherapy after one month of operation. Then the gefitinib was given. A year after surgery, the patient survived without tumor.

CONCLUSION: Secondary pancreatic tumors are rare in clinical practice. To our knowledge, this is the first report of isolated pancreatic metastasis of lung cancer in China and it is the longer tumor-free survival in patients with isolated pancreatic metastasis of lung cancer. However, the value of surgical treatment for isolated metastasis of lung cancer needs further discussion.

Background

Lung cancer has the highest morbidity and mortality among malignant diseases in the world; and adenocarcinoma of the lung, which is a common type of lung cancer, has particularly poor sensitivity to radiotherapy and chemotherapy. Lung cancer is commonly associated with metastases in the brain, bone, and liver, and other organs. Pancreatic metastasis in lung cancer patients, however, is very rare, with published probabilities in the range of 6–11%. Pancreatic metastases, which are often found in patients with metastatic clear cell carcinoma of the kidney, colorectal cancer, and sarcomas, are rarely isolated, but rather are usually accompanied by metastasis of other organs. The probability of solitary pancreatic-occupying lung cancer metastasis has been reported to be only 0.3–3.0%.

Here, we report the case of a man who underwent pulmonary lobectomy for adenocarcinoma of the lung and then resection of pparent metastases occupying the pancreatic neck and body 6 years later. A brief review of the relevant literature follows.

Case Presentation

Six years ago, a 56-year-old Chinese male patient was admitted to the First Hospital of Jilin University for a pulmonary space-occupying lesion revealed by computerized tomography (CT). No obvious abnormalities were apparent in a physical examination and the patient did not have cough, expectoration, hemoptysis, or any other apparent discomfort. The patient had no family history of respiratory or digestive tract diseases. He was not a smoker or alcohol abuser. Liver function tests at the time of
admission showed the following: aspartate aminotransferase, 29 U/L; alanine aminotransferase, 32 U/L, alkaline phosphatase, 12,348 U/L, total protein, 76.4 g/L; albumin, 46.6 g/L; total bilirubin; 16.5 μmol/L; direct bilirubin, 4.0 μmol/L. Meanwhile, we obtained the following routine blood test results: leukocytes, 5.72 × 10⁹/L, red blood cells 5.43 × 10¹²/L, hemoglobin: 166 g/L, platelet: 253 × 10⁹/L. A chest computerized tomography(CT) with secondary enhancement demonstrated ground glass density nodule (approximate size, 2.1 × 2.2 cm) in the upper lobe of the left lung with irregular edges, burrs, lobes, and vacuoles. Imaging-based differential diagnosis led us to exclude alveolar carcinoma as vitreous nodules in the upper lobe of the left lung.

The patient underwent thoracoscopic upper lobe resection of the left lung. Postoperative pathology showed infiltrating adenocarcinoma with medium differentiation(Figure 1A). Most of the abnormal tissue (~90%) showed an acinar growth pattern, whereas some showed a micropapillary growth pattern (10%). The tumor volume was about 2.0 × 1.5 × 1.2 cm. No metastases were found in the visceral pleura, bronchiol walls, bronchotomy margins, vascular incision margins, vasculature, nerves, or para-bronchogenic lymph nodes. Tumor-node-metastasis staging was T1N0M0, with a pathological stage of 1A.

A regular scheduled follow-up re-examination 6 years after the lung tumor resection in January of 2018 revealed untreated pancreatic neck lesions. A subsequent planned re-examination revealed that the mass in the pancreatic neck had increased in volume(Figure 2). The patient was admitted to the First Hospital of Jilin University inshowed secondary distal pancreatic atrophy and pancreatic duct dilation. Based on these imaging findings, we made a preoperative working diagnosis of space-occupying lesions of the pancreatic neck, secondary distal pancreatic contraction, and expansion of the pancreatic duct; primary pancreatic cancer could not be ruled out preoperatively. No metastases were observed in head and chest CT scans. The apparent mass were treated surgically.

Following distal pancreatectomy and splenectomy, the patient's postoperative recovery went well, with no complications (e.g. hemorrhage, pancreatic leakage, abdominal infection). In postoperative pathology, the mass measured 2.3 × 2.0 × 1.8 cm, was pale yellow color, and a firm, tough consistency.postoperative pathological findings showed adenocarcinoma infiltration in the pancreatic tissue,including Napsin A (Figure 1B) and thyroid transcription factor 1 (TTF-1) (Figure 1C) immunopositivity.In light of the patient's medical history, these postoperative pathological findings were demonstrative of metastatic lung adenocarcinoma infiltration of pancreatic tissue(Figure 1D). Gene analysis of the mass revealed a deletion mutation in exon 19 of EGFR. Based on the above findings, we diagnosed the patient with isolated pancreatic metastasis of lung cancer.

He was given four courses of pemetrexed and carboplatin chemotherapy after one month of operation. Then the gefitinib was given.A year after the operation, the patient underwent a full body examination. Abdominal CT images(Fig3A,3B), Head(Fig3C) and chest(Fig3E) , with and without enhancement, conducted showed no abnormalities. Meanwhile a systemic bone(Fig3D) scan also showed no abnormalities.
Discussion And Conclusions

Here, we reported the case of a lung cancer patient whose isolated Pancreatic metastasis could be resected surgically. Due to the rarity of lung metastases in the pancreas and the extreme rarity of isolated pancreatic metastases, there is limited representation of this clinical presentation and its treatment in the literature. Lee et al. [8] reported that surgical resection of secondary pancreatic tumors can improve the postoperative survival rate of patients, depending on the tumor’s pathological type, whether there has been vascular invasion, and the marginal state of the primary tumor. In a multi-center study, Moussa and colleagues showed that surgical treatment of pancreatic metastases of renal or colorectal primary cancers tumor can improve patient survival [9]. Similarly, Zerbi et al. [10] reported that surgical treatment for pancreatic metastasis of renal cancer can prolong 5-year survival, and Aimoto et al. [11] reported a case of a patient with isolated pancreatic metastasis of renal cancer with a successful surgical treatment. Because isolated pancreatic metastasis of lung cancer is rare, and isolated pancreatic metastasis of lung cancer in particular is extremely rare, there are few cases of surgical resection for isolated pancreatic metastasis of lung cancer, and the value of surgical resection has not been clarified. To our knowledge, this is the first report of isolated pancreatic metastasis of lung cancer in China. Igai and colleagues [12] reported the case of a 67-year-old male patient who underwent pancreaticoduodenal resection of a metastasis from a lung cancer tumor that had been removed 5 years prior and remained free of recurrence at his 6-month postoperative follow-up. Watanabe et al. [13] reported the case of 64-year-old male patient who underwent right pulmonary resection due to adenocarcinoma of the right lung and then, 1 year later, had a pancreatic tail mass resected. Mori et al. [14] reported the case of a patient who underwent lobectomy for lung cancer, carboplatin-paclitaxel chemotherapy for 6 months, and subsequent successful surgical resection of an isolated pancreatic metastasis followed by postoperative chemotherapy, with no recurrence at the 6 month-follow-up examination. Despite these encouraging case reports, however, Reddy et al. reported a mean survival time of only 8.75 months for pancreatic secondary tumors, with a 0% 5-year survival rate [7]. However, this Chinese male patient who with isolated pancreatic metastases of lung cancer underwent surgical treatment, but still no tumor recurrence was found six months later, so the value of surgical treatment for isolated metastasis of lung cancer needs further discussion. The indications for surgical treatment of metastatic pancreatic tumors are unclear. Wilson et al. [15], who suggested that surgical removal of isolated pancreatic metastases may extend patient survival to some extent, urged clinicians to be attentive to the following two considerations before pursuing surgical resection: 1) the type of primary disease; and 2) whether the isolated distant metastasis has occurred after a long disease-free period. Notwithstanding, it is very difficult to diagnose pancreatic metastases by preoperative imaging. Because pancreatic metastatic tumors and primary pancreatic tumors are similar in size, location, and number of tumors [16], imaging alone cannot distinguish between them. Differential diagnosis between the two may require endoscopic ultrasonography-guided puncture biopsy [17].
In conclusion, although pancreatic metastases are rare, oncologists should be mindful of the possibility of isolated metastasis in the pancreas. Further studies are needed to clarify whether surgical treatment of isolated pancreatic metastases of lung cancer improve patient survival as has been shown for renal and colorectal cancer patients.

**Abbreviations**

CT = computerized tomography. EGFR=epidermal growth factor receptor.

**Declarations**

Ethics approval and consent to participate
The study was approved by the The First Hospital Of Jilin University Institutional Review Board for human studies.

Consent for publication
Written informed consent was obtained from the patient for publication of this case report and accompanying images.

Availability of data and materials
All data generated or analyzed during this study are included in this published article.

Competing interests
The authors declare that they have no competing interests

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Authors' contributions
Shupeng Wang, Kaimeng Liu, Ludong Tan and Feng Jia reviewed the literature and contributed to manuscript drafting.

Yu Gu, Yahui Liu were responsible for the revision of the manuscript for important intellectual content.

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Figures
**Figure 1**

1A: Histology and immunohistochemistry demonstrating lung infiltrating adenocarcinoma. 1B: H-E stained Napsin A of section near tissue pancreas was immunopositivity in panel. 1C: TTF-1 of section near tissue pancreas was immunopositivity. It showed the adenocarcinoma component. 1D: H-E stained section showed the arrow head indicated the component of adenocarcinoma. All photomicrographs, 200x.
Figure 2

Enhanced axial abdominal CT image series showing a nodular low-density shadow in the pancreatic head and neck. Secondary distal pancreatic atrophy can be seen and dilatation of the pancreatic duct can be seen in panel.
Figure 3

Abdominal CT (Fig3A, Fig3B), chest CT (Fig3D), head CT (Fig3C) and bone scanning (Fig3E) with no signs of metastasis.