Primary Pulmonary Paraganglioma
A Case Report and Review of Literature

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Abstract: Primary pulmonary paraganglioma is a rare disease. We report a case of a 37-year-old female patient with space-occupying lesions in the right lower pulmonary lobe during a routine examination without any symptoms. The patient underwent video-assisted thoracoscopic surgery (VATS) resection of the right middle lobe and dissection of hilar and mediastinal lymph nodes under general anesthesia. She recovered without recrudescence. Preoperative diagnosis is difficult. Accurate diagnosis requires pathological examination, and immunohistochemical test is particularly important. Complete resection is the first treatment option for solitary primary pulmonary paraganglioma; however, VATS is a better technique. Given the high local control rates and few complications of radiotherapy, it is considered as a standard treatment.

(Introduction)

Paraganglioma is a rare disease that affects 2 to 5 million per year. A limited number of cases and sample clinical studies have been reported about primary pulmonary paraganglioma. In this article, we report 1 case of this disease and introduce its diagnosis and treatment.

CASE PRESENTATION

Primary pulmonary paraganglioma is a rare disease and seldom reported. A 37-year-old Chinese stay-at-home woman was treated in our department. This patient had space-occupying lesions in the right lower pulmonary lobe during a routine examination without any symptoms. Her past medical history was unremarkable, and no similar cases were found in her family. Her physical examination did not reveal any abnormalities. Chest computed tomography (CT) showed a round-like soft tissue of approximately 3 cm × 2.8 cm in size. The CT values of the plain scan and enhancement scan were 44 and 245 Hu, respectively. No enlarged lymph nodes were observed at the bilateral hilar and mediastinal lymph nodes or mediastina (Figure 1). Imaging examination showed a tumor with a rich blood supply beside the hilus pulmonis of the right middle lobe. The round-like and hard lesion was approximately 4 cm × 3 cm × 3 cm in size. The lesion also showed a relatively clear boundary, without evident membrane indentation, and was combined with the right middle pulmonary atelectasis and consolidation. No abnormality was found in the other parts. Pathological diagnosis for intraoperative frozen section indicated benign tumor. Postoperative pathological diagnosis showed the following. First, the right middle lung tumor cells were fusiform and acinar and, between cell nests were thin vascular net. Moreover, the tumor cells had abundant granular, pink cytoplasm, and a few tumor giant cells and karyokinesis sign were observed. Interstitial fibrous tissue hyperplasia and hyaline degeneration were found in some areas. Immunohistochemistry showed that the tumor was paraganglioma. Second, no tumor was found at the incisional margin of bronchus. Third, the sending sample of lymph showed reactive hyperplasia. Immunohistochemical test results (Figure 2) revealed chromogranin A(+), synaptophysin(+), CD56(+), Ki-67 index of approximately 2%, creatine kinase (CK) (+), Vimentin(−), LCA(−), CK7(−), CK5/6(−), thyroid transcription factor(−), P63(−), human melanoma Black 45 (−), CD34 (vessel+), CD31 (vessel+), S100 (lesion+), Desmin(−), and MyoD1(−). The postoperative recovery of the patient was good, and the patient did not complain of obvious discomfort. After surgery, the case was followed until September 2014, and the tumor did not recrudesce.

DISCUSSION

Paraganglioma originates from parasympathetic ganglion. This disease mainly occurs at body parts with rich paragangliomas, such as head, neck, mediastinum, adrenal gland, and posterior peritoneum, even bladder, duodenum, and thyroid as reported. Paraganglioma is a rare disease that affects 2 to 5 people per million per year. Primary pulmonary paraganglioma is even rarer. Paragangliomas are slowly growing tumors, present as painless masses, and have a culture doubling time of approximately 42 years. The tumors are potentially low-grade malignant. However, invasive biological behaviors have been reported; these behaviors can cause adjacent lymph node metastasis, such as mediastinal lymph node.
of the primary pulmonary paragangliomas do not show evident clinical symptoms. Few of these symptoms are cough and chest pain. The susceptible population is unknown. Other pulmonary benign tumors and special infections (eg, tuberculoma and inflammatory pseudotumor) are hard to distinguish through imaging examination. Therefore, preoperative diagnosis is difficult, and accurate diagnosis requires pathological examination. Pathologically, primary pulmonary paraganglioma should be identified from carcinoids and other diseases. Hence, immunohistochemical test is particularly important. Several authors summarized the genetic and clinical appearances of paragangliomas, attempting to identify the mutation of genes and determine the algorithm about genetic test and aiming an accurate treatment and better prognosis.

FIGURE 1. The chest computed tomography (CT) showed a round-like soft tissue of approximately 3 cm × 2.8 cm in size. The CT values of the plain scan and enhancement scan were 44 and 245 Hu, respectively. No enlarged lymph nodes were observed at the bilateral hilus pulmonis or mediastina.

FIGURE 2. First, the (right middle) lung tumor cells were fasciculate and acinar form, and between cell nests were thin vascular net. Moreover, the tumor cells had abundant granular, pink cytoplasm, and a few tumor giant cells and karyokinesis sign were observed. Interstitial fibrous tissue hyperplasia and hyaline degeneration were found in some areas. Immunohistochemistry showed that the tumor was paraganglioma. Second, no tumor was found at the incisal margin of bronchus. Third, the sending sample of lymph showed reactive hyperplasia. Immunohistochemical test results (A-hematoxylin and original magnification 100×). Immunohistochemical test results: synaptophysin (syn)(+) (B-original magnification 400×), chromogranin A (CgA)(+) (C-original magnification 400×), CD56(+) (D-original magnification 400×), Ki-67 index about (2%) (E-original magnification 400×), and Vimentin(−) (F-original magnification 400×).
In this case, the patient refused the fibrobronchoscopic examination based on the following reasons. First, according to the imaging examination before surgery, the tumor had rich blood supply; therefore, the patient is at a high risk of bleeding in the process of fibrobronchoscopic forceps biopsy. Second, completely resection of the tumor should be conducted before operation.

Complete resection is the first treatment option for solitary primary pulmonary paraganglioma. However, the postoperative complication rate for such benign tumor is too high. Thoracoscopic surgery becomes the standard method of early lung cancer treatment. VATS allows excellent exposure of the operating field of vision and shows fine structure of lesions. Furthermore, VATS causes less surgical injury and postoperative pain than thoracotomy. Considering the reduced postoperative complications, conduciveness to postoperative recovery, and patient’s aesthetic requirements, we preferred the VAST operation. Based on the condition during the operation, we have an option to continue VATS or perform open thoracotomy. In this case, the postoperative recovery was smooth without any complications.

During the surgery, the hilar and mediastinal lymph nodes were enlarged with a hard texture. To distinguish from malignant lymph node enlargement, we performed lymph node dissection.

Many authors proposed that paraganglioma is sensitive to radiation therapy, without serious side effects. Given the high local control rates and few complications, some authors consider radiotherapy as a first-choice treatment for paraganglioma. The effects of external beam radiation therapy have been evaluated. However, external beam radiation therapy can only be considered when local disease progresses. The use of the chemotherapy regimen is not unanimously approved. In some metastasis cases, sorafenib use has been reported. Paraganglioma needs to be differentiated from the following kinds of diseases, including lung cancer, lung tuberculosis, inflammatory pseudotumor, hamartoma, lung carcinoid tumor and lung metastasis tumor, etc.

CT and magnetic resonance imaging are the important imaging examination methods of positioning and qualitative diagnosis of paraganglioma. CT scan of pulmonary paraganglioma shows isodensity or slightly lower density. Enhanced scan shows significant enhancement. T1W1 of magnetic resonance imaging shows isodensity and low signal. T2W1 shows medium, high, or uneven mixed signal. DWI shows high signal. Enhanced scan shows that the parenchyma of the tumor is partially obviously enhanced. The occupied effect of the tumor causes the corresponding clinical symptoms.

Lung Cancer
Derived from bronchial epithelial cells and appears to invasive growth. Imaging examination shows that the tumor invades bronchi, blood vessels, adjacent tissues, and organs. The obstructive changes appear. Pulmonary paraganglioma is mainly located around blood vessels and nerve. Tumor is closely related to pulmonary blood vessels and nerves, rather than bronchia. This is the key point to differentiate pulmonary paraganglioma from lung cancer.

Lung Metastasis Tumor
Multiple nodules in bilateral lungs with clear edge and uniform density, mainly in medium, lower, and peripheral lung field. The hilar nodes are enlarged. There is a history of primary malignant tumor, which can be differentiated by CT multidimensional reconstruction.

Pulmonary Tuberculosis
Pulmonary tuberculosis is mainly appeared in the dorsal pulmonary segment with uniform density and smooth boundary. The long rough burrs can be observed in some patients; enhanced scan shows the nonenhanced or circular enhanced focus. The caseous necrosis can be observed in the center of focus. The satellite foci are observed around the focus. The pleura is thickened. The hilar and mediastinal lymph nodes are not enlarged. The focus is not slowly absorbed or not absorbed after antituberculosis treatment.

Pulmonary Inflammatory Pseudotumor
It is located around the lung field, mainly confined to a leaf. The shape is irregular. The focus often has the wide fundus and adheres on the pleura. The adjacent pleura is thickened. Enhanced scanning shows high density and homogeneous enhancement. The lesion can be shrank after antiinflammatory treatment.

Hamartoma
The tumor components are complex. Most are carcinoid malformation caused by abnormal development of normal tissues. A few is mesenchymal tumors. Fat and calcification are the manifestation of most hamartomas. Popcorn-like calcification is common in pulmonary hamartoma.

Lung Carcinoid
The histopathologic characteristics include typical neuroendocrine morphology with positive cytokeratin, the expressions of chromogranin A, synaptophysin, CD56, and other neuroendocrine factors. The expression of Ki-67 index is helpful for differential diagnosis. The feature of imaging examination is absent. The diagnosis needs to be clarified by the combination of clinical manifestations and pathological examination.

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