

# Report of a lung carcinoma extended to the left atrium through pulmonary vein

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**Abstract:** Lung cancers may extend along or grow through the pulmonary veins to invade or lie within the left atrium (LA). A 62-year-old man, previously healthy, presented with 1-month ventilatory-independent right hemithorax back pain, dry cough and large effort dyspnea. He also referred weight loss of 12 kg in 10 months and denied hemoptysis. As antecedents, he smoked for 40 years and moderate daily alcoholism. On physical examination, the patient was in good general condition, hydrated and regular respiration at rest [blood pressure (BP) =120/80 mmHg; heart rate (HR) =90 bpm; respiratory rate (RR) =16 rpm]. Cardiac auscultation revealed two standard rhythmic sounds without murmurs. Pulmonary auscultation revealed a slightly diminished vesicular murmur in the lower 1/3 of the right hemithorax without adventitious noises. Chest radiography showed a mass over the right lower lung. A CT scan confirmed the radiography image with the mass extending along the right inferior pulmonary vein and a tumor in the LA. Transthoracic and transesophageal echocardiography revealed large mass within the LA (occupying almost the entire cavity), measuring about 10 cm × 3 cm at its largest diameter, prolapsing into the left ventricle. Bronchoscopy, head CT scan, and whole-body bone scintigraphy investigation did not show any distant metastasis. The patient was successfully operated removing the intracardiac and inferior pulmonary vein tumor with the aid of cardiopulmonary bypass, followed by a right inferior lobectomy carried out after 25 days. After 30 days from surgery presented seizures associated a brain metastasis evidenced by CT when adjuvant radio and chemotherapy was started. During the next 90 days, the clinical conditions worsened, and the patient died 4 months after the surgical treatment. The case report has two primary justifications, even considering the poor outcome: (I) rarity and (II) the possibility of the surgical treatment.

**Keywords:** Lung cancer; pulmonary vein; left atrium (LA); cardiothoracic surgery

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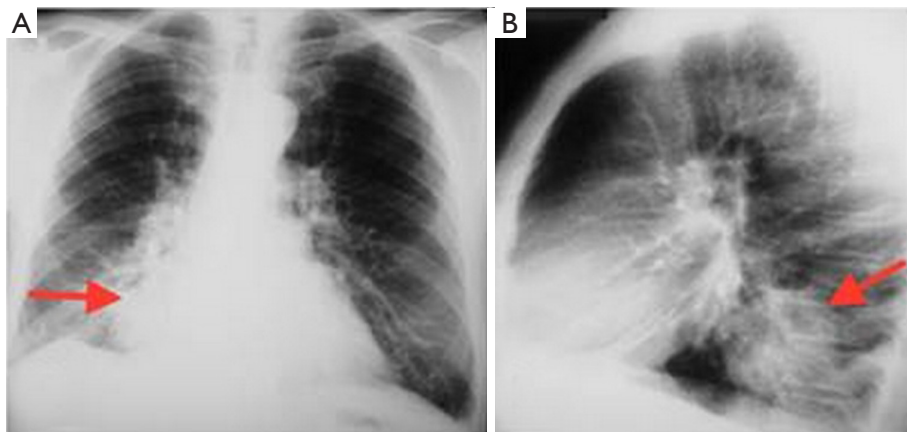
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## Introduction

Although relatively rare, the extension of the pulmonary neoplasm to the left atrium (LA) has been well documented, particularly in patients with primary lung cancer (1,2). In a previous review of 215 lung cancer patients studied by gadolinium-enhanced 3D magnetic resonance angiography, an involvement of the proximal portion of the pulmonary veins and an extension into the LA were found in 9 (4.2%) and 2 (0.9%) patients, respectively (3). Similarly,

a more recent retrospective analysis of 4,668 patients who underwent surgery for lung cancer found pathological evidence of pulmonary vein and LA involvement in 34 (0.7%) and 25 (0.5%) subjects, respectively (4).

Tumors that directly invade the LA belong to T4 of the TNM classification, suggesting that the tumor is inoperable. However, lately, the indication of surgical resection of lung cancers has been expanded to some T4 patients, and long-term survival has been reported. Primary lung carcinomas can invade the LA through or along the pulmonary veins.



**Figure 1** Chest radiography was showing a mass (arrows) over the right lower lung in the topography of the inferior pulmonary vein (A,B).

The tumor extension through the pulmonary veins is few reported with scarce publications on the specialized literature (3). Therefore, the unusual primary lung carcinoma invasion of LA through pulmonary veins justifies this case report (5,6).

### Case presentation

A 62-year-old man, previously healthy, presented with 1-month ventilatory-independent right hemithorax back pain, dry cough and large effort dyspnea. He also referred weight loss of 12 kg in 10 months and denied hemoptysis. As antecedents, he smoked 1–2 packs/year and moderate daily alcoholism. On physical examination, the patient was in good general condition, hydrated and regular respiration at rest [blood pressure (BP) =120/80 mmHg; heart rate (HR) =90 bpm; respiratory rate (RR) =16 rpm], with cardiac auscultation presenting two phonetically normal rhythmic sounds without murmurs. Pulmonary auscultation revealed a slightly diminished vesicular murmur in the lower 1/3 of the right hemithorax without adventitious noises. Chest radiography showed a mass over the right lower lung (Figure 1). A CT scan confirmed the radiography image with the mass extending along the right inferior pulmonary vein and a tumor in the LA (Figure 2). Transthoracic and transesophageal echocardiography revealed large mass within the LA (occupying almost the entire cavity), prolapsing into the left ventricle (Figure 3). Bronchoscopy, head CT scan, and whole-body bone scintigraphy, as staging investigation, did not show any distant metastasis. The patient was successfully operated removing the

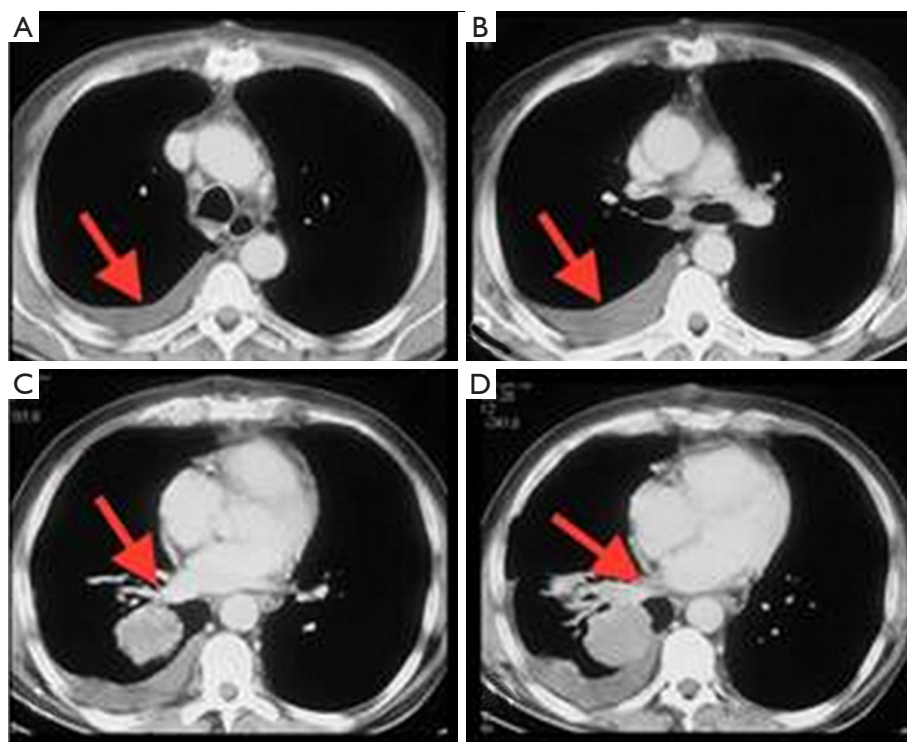
intracardiac and inferior pulmonary vein tumor with the aid of cardiopulmonary bypass (Figure 4), followed by a right inferior lobectomy carried out after 25 days (Figure 5).

After 30 days from surgery presented seizures associated a brain metastasis evidenced by CT when adjuvant radio and chemotherapy was started (Figure 6). During the next 90 days the clinical conditions worsening and the patients died 4 months after the surgical treatment.

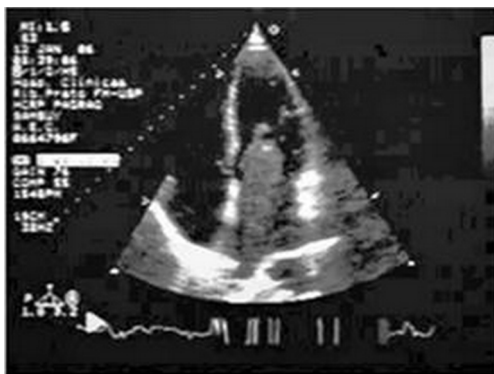
The morphological and immunohistochemically findings confirmed a diagnosis of large cell neuroendocrine carcinoma of the lung. The immunohistochemistry study confirmed the neuroendocrine differentiation of the tumor cells with positivity for chromogranin A, synaptophysin and neuro-specific enolase (NSE). On microscopic examination, the tumor had an organoid pattern composed of nests of large, pleomorphic cells with ovoid nuclei, prominent nucleoli and clear cytoplasm (Figure 6A). There were extensive areas of necrosis (Figure 6B). The immunohistochemistry study revealed positivity of the neoplastic cells for AE1/AE3 (focal), 35 $\beta$ H11, CAM5.2 (focal), CEA (focal), chromogranin A (focal), synaptophysin (Figure 6C), NSE, and negativity for LCA, 34 $\beta$ E12, CK7, CK20, TTF-1. The proliferation index (Ki-67) was approximately 40% (Figure 6D).

### Discussion

Left atrial tumor direct extension via pulmonary veins is uncommon and the literature search shows more cases involving sarcoma than direct lung extension of bronchogenic carcinoma (7). Guha and colleagues (2011) presenting an important left atrial extension of lung



**Figure 2** Chest tomography. (A,B) Chest tomography with contrast showing the presence of pleural effusion without the pathological reinforcement of the parietal pleura (arrows); (C) lower right lobe lung mass in close contact with the inferior pulmonary vein (arrow); (D) the tumor is seen invading the inferior pulmonary vein and prolapsing into the left atrium but without signs of invasion of the left atrial wall (arrow).



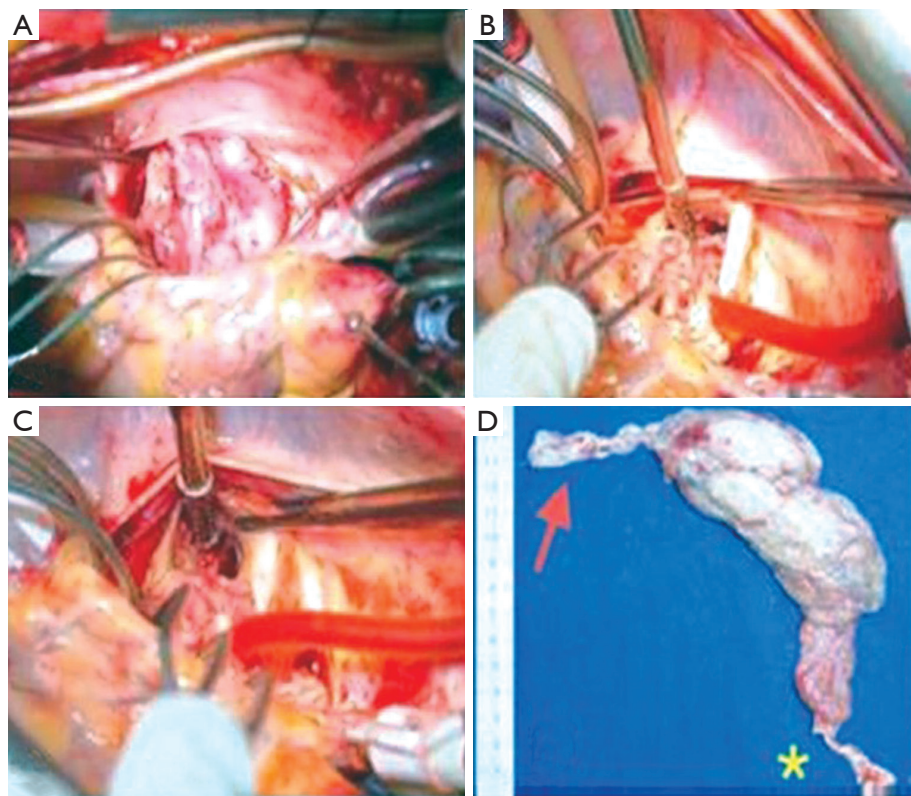
**Figure 3** Transthoracic echocardiography apical four-chamber view showing pulmonary vein left atrial invasion by a large mass, with protrusion through the mitral valve into the left ventricle during diastole.

carcinoma simulating myocardial infarction highlighted a worldwide scarcity of reported cases. Very few of such cases have been reported worldwide (8). A left atrial mass as a manifestation of lung carcinoma was reported by researchers

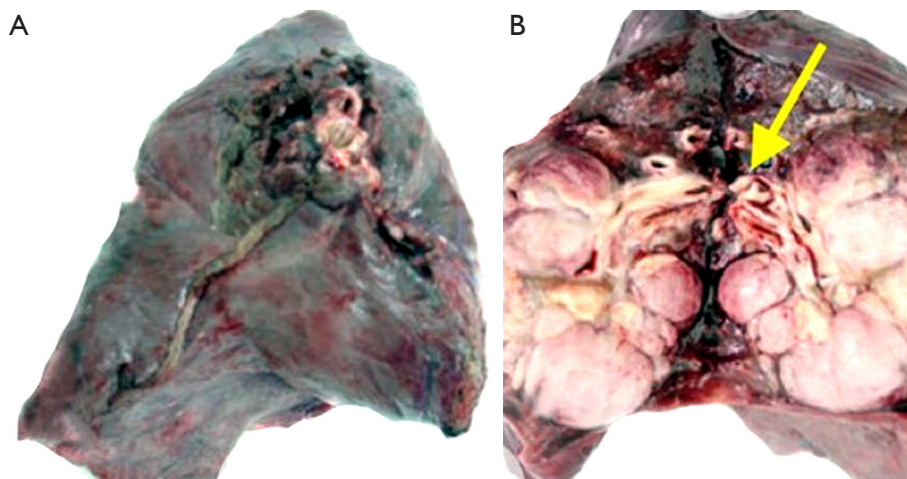
in Australia (9) and Memphis (10). Bronchogenic carcinoma mimicking left atrial myxoma was reported from Bombay, India (7). Left atrial invasion by lung carcinoma through a pulmonary vein was visualized by Lestuzzi *et al.* from Italy (11) by Desai *et al.* from Pittsburgh (12) and also by Watanabe and Kubo (6).

There were no clear patterns regarding the histological type or tumor location in these cases. On the other hand, all the tumors showed rapid growth and comprised a large mass in the lung parenchyma. Especially the cardiac and intravascular portions of each tumor were less differentiated and more necrotic than the primary focus in the lung. Radical surgery using a cardiopulmonary bypass for lung cancer patients with such intracardiac involvement is useful for improvement of the performance status and reduces the risk of sudden death due to cardiac failure or tumor emboli (13).

According to Ballo and colleagues, the evolution towards cerebral stroke has rarely been reported. They described an atypical case of hematogenous metastatic invasion of the LA from pulmonary neoplasm extension presenting as an

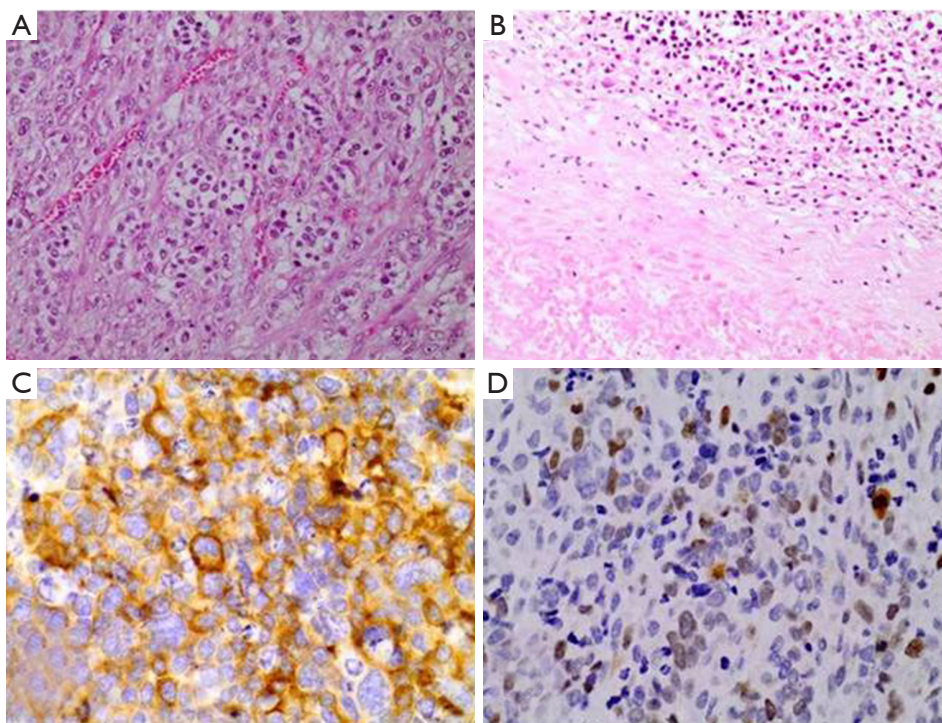


**Figure 4** Surgical time with extracorporeal circulation. (A) Left atrium opening and tumor exposure; (B) there was no direct invasion of the atrial wall by the tumor allowing easy detachment; (C) identification of the ostium of the right inferior pulmonary vein at its entry into the left atrium and showing that tumor prolapsed through the distal portion of the vein into the atrium and without invasion of the walls; (D) tumor resected in its free portion within the inferior pulmonary vein (arrow) and the left atrium (asterisk).



**Figure 5** Macroscopy of the right lung lower lobe. (A) Resected right lower lobe without cardiopulmonary bypass; (B) longitudinally sectioned lung mass showing an involvement of the inferior pulmonary vein in its intrapulmonary portion and which was sectioned very close to its insertion in the left atrium ostium (arrow) where there was no involvement of its wall but the only intravascular extension of the tumor into the left atrium.





**Figure 6** Histopathological study. (A) Organoid aspect of the tumor with nesting of tumor cells (H&E, 200×); (B) foci of necrosis (inferior left) were frequently seen (H&E, 200×); (C) the tumor cells presented positivity for synaptophysin (immunostaining, 400×); (D) the proliferation index (Ki-67) was around 40% (immunostaining, 400×).

ipsilateral stroke (14). In the present case, the intracardiac and the pulmonary masses were similar on the microscopic examination. The tumor showed an organoid pattern and was composed of large cells with ovoid nuclei, dispersed chromatin, and prominent nucleoli. There were extensive necrotic areas, and mitotic figures were frequent. No areas with squamous or glandular differentiation were found, ruling out a combined tumor (large cell neuroendocrine carcinoma and squamous cell carcinoma or large cell neuroendocrine carcinoma and adenocarcinoma). Even though extremely rare, the intracardiac extension of lung large cell neuroendocrine has been previously reported in the literature (15).

As pointed in the introduction of this text, the case report has two primary justifications, even considering the poor outcome: (I) rarity and (II) the possibility of the surgical treatment of a tumor belong to T4 of the TNM classification. It is possible to speculate that cerebral metastases, directly related to poor clinical outcome, predate the cardiac surgery. This consideration is based on the surgical observation that the cardiac mass was removed *en bloc* without fragmentation.

Finally, two points deserve criticism: (I) the patient received brain CT to check metastasis and brain MRI with contrast is the standard image to check brain metastasis as the initial staging of lung cancer, and (II) even considering the evolution towards cerebral stroke has been rarely reported. In this case report, we describe an atypical case of haematogenous metastatic invasion of the LA from pulmonary neoplasm extension presenting as an ipsilateral stroke whose ASCO classification changed during the clinical management. The option of intraoperative radiotherapy for such a trouble case we did not carry out it because, unfortunately, our hospital does not have technical conditions to do it.

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### Footnote

*Conflicts of Interest:* The authors have no conflicts of interest to declare.

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