Case Report

Pulmonary hamartoma as a differential diagnosis of breast cancer metastasis: case report*

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Abstract

We present the case of a 60-year-old female patient who had been in menopause for 14 years and presented a pulmonary nodule on chest X-ray diagnosed in the postoperative follow-up evaluation of breast cancer. The patient had a history of mastectomy and ipsilateral axillary lymphadenectomy for invasive ductal breast carcinoma, as well as of hormone therapy, chemotherapy, and adjuvant radiotherapy. After thoracoscopic nodulectomy, the frozen section analysis revealed a pulmonary hamartoma. Recent studies show that 75% of patients who undergo surgery for pulmonary nodules after a curative mastectomy for breast cancer present lung metastases, 11.5% present primary lung cancer, and 13.5% present benign lesions, including hamartoma.

Keywords: Hamartoma; Breast neoplasms; Lung neoplasms; Neoplasm metastasis.

Introduction

Pulmonary metastasis is a common occurrence in patients with breast cancer, (1,2) who are usually treated with systemic chemotherapy. (1) Some clinical studies suggest that resection of pulmonary metastases (metastasectomy) can be a treatment option. (2-5) However, the role of surgery for metastases originating from breast cancer has yet to be fully determined. (6)

Pulmonary nodules that appear in patients who underwent mastectomy for breast cancer are usually pulmonary metastases. However, it is not uncommon to find metachronous lung tumors as well as some benign diseases, among which pulmonary hamartoma stands out. ⁽⁶⁾

Case report

A 60-year-old Caucasian female who had been in menopause for 14 years was referred to the Department of Oncology due to the presence of a pulmonary nodule on chest X-ray (Figure 1) diagnosed in the postoperative follow-up evaluation of breast cancer.

This was a patient with a personal history of right total mastectomy and ipsilateral axillary lymphadenectomy for invasive ductal breast carcinoma (diagnosed by core biopsy) performed 3 years prior. The lesion removed was a nuclear grade II, stage IIB (T2N1Mx) tumor measuring 45 mm along its longest axis. The patient also had a history of hormone therapy (20 mg/day of tamoxifen) and chemotherapy (cyclo-

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Figure 1 – Simple anteroposterior chest X-ray revealing the absence of the right breast and also revealing a peripheral nodular parenchymal lesion in the upper third of the right hemithorax (arrow).

phosphamide + methotrexate + 5-fluorouracil: 4 neoadjuvant cycles and 6 adjuvant cycles), as well as of adjuvant radiotherapy (total dose of 50 Grays for 5 weeks).

She reported no pulmonary symptoms, fever, or weight loss.

The patient then underwent computed tomography of the chest, which revealed a pleural, parenchymal pulmonary nodule measuring approximately 30×25 mm (Figure 2) in the right lung. Therefore, a hypothetical diagnosis of pulmonary metastasis of the breast carcinoma previously removed was made, and biopsy was indicated for diagnostic confirmation.

The patient was then submitted to video-assisted thoracoscopic nodulectomy. The frozen section analysis revealed that it was a benign tumor (probably a pulmonary hamartoma), and the procedure was then finished.

The anatomopathological examination revealed that it was a non-capsulated, regular lesion measuring 30×25 mm and presenting a firm and fibroelastic consistency. Microscopy (Figure 3) revealed vessels, adipose tissue, cartilage, well-differentiated (but disorganized) bronchial structures, and highly vascularized fibrovascular stroma with

interposed embryonic bronchi, compatible with pulmonary hamartoma.

The patient recovered well and was discharged on postoperative day 2. She is now under annual outpatient follow-up treatment in the Department of Oncology, having been on hormone therapy with tamoxifen for 5 years, and has presented no signs of tumor recurrence to date.

Discussion

Pulmonary hamartoma accounts for 77% of benign lung tumors⁽¹⁾ and for 4% of all solitary lung nodules.⁽⁷⁾ This lesion has been described as a benign neoplasia of fibrous connective tissue of the bronchus surrounded by respiratory epithelium, almost always containing cartilage and fat tissue, not respecting the usual histological distribution of the lung.⁽²⁾

Ninety per cent of the hamartomas manifest as a solitary peripheral mass, $^{(7)}$ and rarely occur in the form of multiple lesions. $^{(3)}$ They are more common in adults, and their incidence is twice as high in males than in females, $^{(1)}$ with a mean population growth of 3.2 ± 2.6 mm/year. $^{(8)}$

Radiographically, the peripheral lesion is typically found in the lung bases and appears as a well-defined, homogeneous mass; sometimes, the margins can be lobulated or bosselated. The usual size ranges from 1 to 2 cm, although larger lesions can also be observed. There can be calcifications, usually diffuse in form, in up to 30% of the cases. Some authors have also demonstrated the presence of fat tissue in 50% of the hamartomas evaluated by computed tomography.

There is the occurrence of malignancy in hamartomas. Some researchers⁽⁵⁾ have found that the incidence of bronchial carcinoma is 6.3-fold higher in patients with hamartoma than in a normal population, suggesting the presence of an etiologic relationship.

It is common for confusion to occur when pulmonary nodules are found in the follow-up evaluation of patients who underwent mastectomy.

According to a recent study, (6) the pathologic diagnoses of pulmonary nodules demonstrated that 75% of the patients who underwent surgery for pulmonary nodules after a curative mastectomy for breast cancer presented lung metastases, 11.5% presented primary lung cancer (adenocarcinoma and

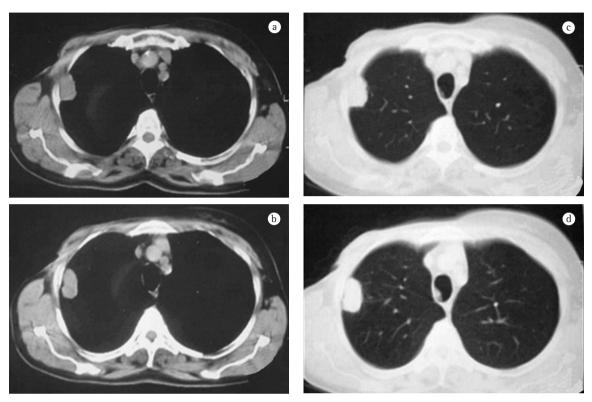


Figure 2 – Computed tomography of the chest (a/b: mediastinal window; c/d: pulmonary window) revealing a pleural, parenchymal lesion in the posterior segment (segment II) of the upper lobe of the right lung, with possible involvement of the thoracic wall, measuring approximately 30×25 mm.

squamous cell carcinoma), and 13.5% presented benign lesions (pulmonary tuberculosis, sclerosing hemangioma, typical organized pneumonia), 28.6% of which were pulmonary hamartomas.

Therefore, the pulmonary nodules that appear in patients submitted to mastectomy for breast cancer are not always pulmonary metastases, and the confirmation of the histopathological diagnosis is of fundamental importance in order to define the treatment.⁽¹¹⁾

Bronchoscopy with biopsy is indicated in endobronchial lesions and also in patients with pulmonary symptoms, such as cough, hemoptysis, recurrent pulmonary infections, or atelectasis. Percutaneous transthoracic aspiration biopsy can diagnose 85% of the hamartomas close to the thoracic wall, differentiating them from nodules of other etiologies, such as breast cancer lung metastasis. When cartilage fragments are present on cytology, the diagnosis is highly suggestive of hamartoma. If a diagnosis has not been made, since percutaneous biopsy is not

always possible due to the stiffness of the tumor, enucleation or resection via open thoracotomy or video-assisted resection are indicated. (9,11,13)

Since breast cancer has a slow progression, ⁽¹⁴⁾ the mean time between mastectomy and identification of pulmonary nodules is approximately 60 months, this time being greater in cases of primary malignant tumors of the lung (mean of 272 months) and benign diseases of other etiologies (mean of 132 months), including hamartoma. ⁽⁶⁾ In addition, the incidence of multiple pulmonary nodules is significantly more frequent in patients with metastatic breast cancer.

The literature has yet to establish whether it is possible to predict the diagnosis of pulmonary metastasis in multiple pulmonary nodules that are revealed in the early follow-up evaluation (within less than five years) of patients who underwent mastectomy for breast cancer. (15)

Recently, other ways to differentiate primary malignant cancers or benign diseases from pulmo-

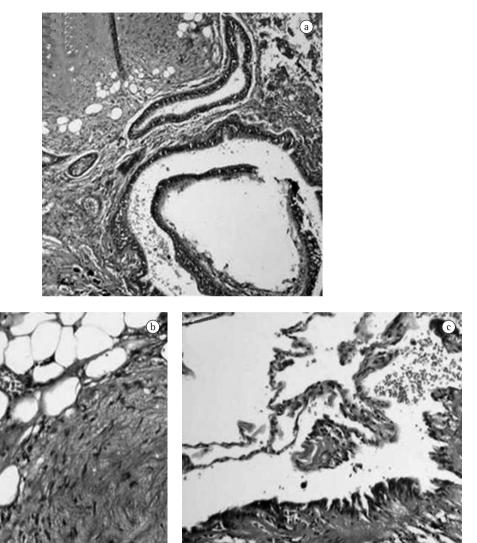


Figure 3 - Photomicrograph showing the presence of adipose tissue, vessels, hyaline cartilage, well-differentiated bronchial structures, and embryonic bronchi permeating myxomatous stroma - Pulmonary hamartoma (hematoxylin and eosin: $\times 100$ magnification in a, $\times 200$ in b, and $\times 400$ in c).

nary metastases have been studied. Positron emission computed tomography using glucose marked with fluoride 18, which is an imaging method with high specificity for malignant diseases, [16] can be cited as an example. Another equally important ally is immunohistochemical study, which can be performed in biopsy specimens and also differentiates those three entities; thyroid transcription factor 1 monoclonal antibody, which is one of the markers of primary lung carcinoma, is worthy of mention. [17]

Therefore, pulmonary nodules in patients with breast cancer should only be treated after histopathological confirmation, and surgical resection should be considered both as a diagnostic and as a therapeutic option.

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