Small Cell Breast Cancer with Lung Metastases

Carcinoma microcítico de mama con afectación pulmonar

Dear Editor,

We report the case of a 58-year-old patient with a significant history of smoking and previous pituitary microadenoma. In August 2014, she consulted due to the appearance of a hard, rapidly growing mass in her right breast detected by self-examination. A screening mammogram 6 months previously had been negative. She underwent mammogram/ultrasound, which showed a dense 18-mm nodule with undefined, spiculated borders in the lower external quadrant, BI-RADS category 5, with no axillary lymphadenopathies. Biopsy of the nodule showed a high-grade neuroendocrine carcinoma with no expression of hormone receptors or HER2 onco-protein, and a cell proliferation index (Ki67) greater than 80%. Given the presence of a rare, highly undifferentiated tumor, we decided to complete the study with imaging tests to rule out distant disease and confirm the mammary origin of the lesion. Computed tomography (CT) confirmed the presence of the breast nodule and showed mediastinal lymphadenopathies in the pathological range (Fig. 1), and multiple millimetric pulmonary nodules, septal thickening, and nodules suggestive of carcinomatous lymphangitis. These findings were confirmed on positron emission tomography, and other disease foci were ruled out. Endobronchial ultrasound was performed, with aspiration of the mediastinal lymphadenopathies, which again showed the presence of a high-grade TTF1-negative neuroendocrine carcinoma with a Ki67 of 98%.

In view of the unusual nature of the case, it was discussed in the multidisciplinary session, and the specimens obtained from both sites were compared by the pathology lab. Similar morphology was observed: small and medium-sized cells with minimal cytoplasm, rounded nuclei with glandular chromatin and occasional nucleoli, arranged in the form of solid nests, with no glandular or squamous identification. The immunohistochemical profile showed expression in both cases of cytokeratin 19, synaptophysin, and chromogranin. After review of the various tests and the peculiarity of the case, with the presence of synchronous high-grade neuroendocrine tumors, 2 possible scenarios were considered: the presence of primary lung cancer with secondary breast involvement or a primary breast tumor with lung involvement. The first situation was ruled out when no lung mass was observed in the imaging tests, whereas there was clear evidence of the mammary nodule. Moreover, TTF1 negativity in the specimens supported this hypothesis, leading us to think that the most likely scenario was primary breast tumor with secondary pulmonary metastases. With this in mind, we decided to start polychemotherapy for metastatic disease based on a platinum-etoposide doublet, and a total of 6 cycles were administered. In a post-treatment follow-up with CT and PET-CT, complete morphological and metabolic response was observed, both in the breast and in the lymph nodes. Radiation therapy was subsequently given to the breast and the mediastinum, followed by prophylactic cranial irradiation. However, despite the good initial response, cerebral and systemic recurrence were observed 4 months later. The patient did not respond to the different treatment regimens, and died soon after.

Breast cancer is the most common malignant tumor worldwide, and the most predominant histopathological variety is infiltrating ductal carcinoma. One unusual variety is neuroendocrine carcinoma, described in 1977 to refer to a group of neoplasms that, while not demonstrating neuroendocrine-secreting granules, may come from argyrophilic cells originating in the neural crest that might have migrated to the mammary ducts. This histology was included in the WHO classification in 2003, and 3 variants are distinguished: solid, small cell and large cell. These tumors can be low and high grade which, like hormone receptor and HER2/neu onco-protein expression, has a prognostic and predictive impact. The presence of a small cell component generally determines the biologic aggressivity of the tumor. Immunohistochemistry is important for diagnosis, and includes specific markers such as cytokeratin, neuron-specific enolase, synaptophysin, serotonin, bombesin, and chromogranin, among others. In our case, these techniques showed the tumor was positive for synaptophysin and chromogranin, and negative for breast tumor markers (estrogen and progesterone hormone markers, and c-erb-2).

As the natural history of this tumor is similar to that of other equivalent lung masses that have a tendency toward early metastasis, the therapeutic approach is also similar, given the lack of prospective clinical trials in this very special scenario. In the case of extensive disease, the only possible treatment modality is palliative chemotherapy, perhaps with the addition of radiation therapy on an individualized basis, as received by our patient. In extrapulmonary neuroendocrine tumors, cerebral involvement is less common, so brain irradiation has less potential benefit. Prognosis in extrapulmonary tumors differs according to the series, as some authors show it to be similar or even better than in small cell lung cancer, while others claim otherwise. According to the available data, mean survival in limited and extended

Fig. 1. CT slice of breast lesion and mediastinal disease.

---

Please cite this article as: Cruz Castellanos P, Quintana L, de Castro J. Carcinoma microcítico de mama con afectación pulmonar. Arch Bronconeumol. 2018;54:586–587.
Primary Endobronchial Tumors in Childhood

Tumores endobronquiales primarios en edad pediátrica

Dear Editor,

Primary lung tumors are extremely rare in childhood, although the exact incidence is unknown.1–4 They are classified into 2 broad groups according to the degree of malignancy:5–7 benign tumors include inflammatory pseudotumor and hamartoma, whereas malignant tumors (accounting for approximately 70–75% of cases) include primary tumors and metastases, usually from extracranial tumors.5 Primary tumors include bronchial carcinoid tumors (the most common primary lung cancer), followed by mucoepidermoid tumors, bronchogenic tumors, pleuropulmonary blastoma, and lymphoma.1,2,5

Diagnosis is based on imaging and pathology studies, but often delayed by the low incidence of this type of disease.2,5

We performed a retrospective analysis of the cases of endobronchial tumors diagnosed in our hospital between 2000 and 2017. We found 4 patients, 3 girls and 1 boy, aged from 3 months to 12 years, with a diagnosis of endobronchial tumor. We describe each case briefly below.

Clinical case 1: a 9-year-old girl under investigation for an image of persistent consolidation/atelectasis in the right lower lobe (RLL). Clinical picture: 2-month history of intermittent febrile syndrome, cough, and weight loss. The initial diagnosis was pneumonia, and antibiotic treatment was administered, with no response. Personal history: episodic asthma with sensitization to dust mites and pollens and 3 previous episodes of pneumonia. Physical examination: hypoventilation in lower third of the right hemithorax. Additional tests: Complete blood count, biochemistry, hepatic and renal profile normal; ESR 113 mm/h; Mantoux 0 × 0 mm; Sweat test negative; α-1 antitrypsin normal. AP chest X-ray (Fig. 1A) consolidation in right lung base. Lung CT (Fig. 1B): consolidation in RLL, atelectasis in RLL associated with small right posterior pleural effusion, and right paratracheal, subcarinal, and hilar lymphadenopathies; urinary catecholamines normal. Fiberoptic bronchoscopy: tumor mass at entrance to bronchus of the right basal segments, almost entirely obstructing the bronchial lumen. Pathology report: bronchial carcinoid tumor. Right middle and lower lobectomy was performed with subsequent favorable progress.

Clinical case 2: a 12-year-old boy under investigation for massive atelectasis of the left lung. Clinical picture: 2–3 month history of dry cough, vomiting, and intermittent fever. He received 2 cycles of antibiotic treatment with transient improvement. Examination: hypoventilation in left hemithorax. Additional tests: complete blood count, biochemistry, hepatic and renal profile normal; ESR 94 mm/h; LDH 1207 IU/l; enolase, catecholamines, bone marrow aspiration and biopsy normal. AP chest X-ray (Fig. 1C) and chest CT (Fig. 1D): massive atelectasis of left lung. Abdominal CT: right intrarenal mass. Fiberoptic bronchoscopy: left endobronchial mass. Pathology: CD45 and CD20–positive lymphocytic cells, consistent with B cell non-Hodgkin’s lymphoma (NHL). FNA of right renal tumor: B cell non-Hodgkin lymphoma. Given these results, the patient was diagnosed with Stage III NHL, primary location unclear. He was treated with chemotherapy and remains disease-free after 5 years.

Clinical case 3: 10-year-old girl, asymptomatic, with no significant history, referred due to contact with tuberculosis. Additional tests: Mantoux 0 × 0 mm. Pulmonary CT: occupation of airspace in basal and lateral segments of the RLL with a tree-in-bud image, suggestive of infectious bronchiolitis. In view of this situation, anti-tuberculosis treatment was administered for 6 months. Second Mantoux test negative. Follow-up lung CT (Fig. 1E): hyperdense punctiform image of polypoid appearance, partially occupying the lumen of the bronchus of the left basal segment. Fiberoptic bronchoscopy (Fig. 1F): 2 reddish, pedunculated, well-defined masses in the RLL bronchus, that were resected. Pathology report: inflammatory polyp with no granulomas; culture and PCR of bronchoalveolar lavage and sputum negative. Favorable progress.

Clinical case 4: 3-month-old infant who had been rejecting feeds for 10 days. No history of interest. Physical examination: hypoventilation in left hemithorax. Additional tests: chest X-ray (Fig. 1G) showing air trapping in left lung with no significant shift of mediastinum or contralateral lung. Lung CT (Fig. 1H): stenosis of left

References


Patricia Cruz Castellanos, a, * Laura Quintana, b Javier de Castro c

a Servicio de Oncología Médica, Hospital Universitario La Paz, Madrid, Spain
b Servicio de Oncología Médica, Hospital San Pedro de Alcántara, Cáceres, Spain

corresponding author.
E-mail address: cruz.patricia@hotmail.com (P. Cruz Castellanos).

1579-2129/ © 2018 Published by Elsevier España, S.L.U. on behalf of SEPAR.