HOT in Madrid simply due to the altitude effect. A somewhat similar situation has been demonstrated in critical patients with regards to the PaO$_2$/FiO$_2$ ratio. Pérez Padilla has clearly observed that the patients have a lower shunt and presumably a lower degree of lung injury than residents at sea level when meeting the criteria for progressive adult respiratory distress syndrome with a PaO$_2$/FiO$_2$ ratio of 200 mmHg. This author suggests keeping in mind the effect of altitude when comparing these patients in published studies as there can be important differences in the results obtained.

The effect of acclimatization or adaptation to heights of humans is well known. There are two fundamental mechanisms developed by subjects living at moderate altitudes, these being the increase in ventilation and the decrease in plasma volume, with consequent hemococoncentration. The increase in red blood cells maintains the total oxygen content of the blood and guarantees the transport of oxygen to the tissue. What is attractive is the hypothesis that the absolute PaO$_2$ value in a COPD patient would not be the best indicator for whether to receive home oxygen therapy. Instead, the degree of efficacy of the compensatory mechanisms of the patient and his/her degree of adaptation to the altitude in which he/she lives would be a better indication. It is necessary to develop studies that more thoroughly analyze these physiological adaptation mechanisms in order to establish dynamic criteria for considering HOT.

Currently, HOT is prescribed by extrapolating the criteria established at sea level to the entire population. Thomas Petty said on an occasion: "If we applied the HOT criteria in Denver (Colorado – 1,609 m above sea level), we would have to give O$_2$ to the entire population." Does this make sense? We should reconsider HOT criteria according to altitude.

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Salvador Díaz-Lobato, Sagrario Mayoralas Alises

Corresponding author.
E-mail address: sdiazlobato@gmail.com (S. Díaz-Lobato).

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Skeletal Muscle Metastasis as Initial Presentation of Non-Small-Cell Lung Carcinoma

Metástasis en músculo esquelético como presentación inicial de un carcinoma no microcítico de pulmón

Dear Editor:

Although muscle tissue makes up more than 50% of the total body mass, metastatic extension to the skeletal muscle is an exceptional event in solid organ neoplasms, with an accumulated incidence of less than 1.5% in recent series, and it is usually limited to advanced phases of the disease. Due to their clinical and radiological appearance, skeletal muscle metastases (SMM) are similar to soft tissue sarcomas, entities with greater prevalence in which case surgical resection offers potentially curative results. This differential diagnosis takes on special relevance if the SMM is the initial manifestation of a primary tumor that had been clinically silent up until that moment, a situation that has been infrequently reported in the literature.

Our patient is a 69-year-old male ex-smoker, whose medical history included the presence of arterial hypertension, dyslipidemia and polymyalgia rheumatica. The patient reported the appearance of a tumor on the left thigh, with slow progressive growth and accompanied by continuous dull pain. With the exception of an unquantified weight loss, the patient denied any other systemic symptoms. After several weeks, he also reported the appearance of a second smaller lesion on his right calf. The physical examination showed that the patient was in a state of good general health, with no findings on the cardiopulmonary auscultation or palpable lymphadenopathies. On the outer side of the left thigh, there was a stone-like mass that was not painful to touch (10 cm × 5 cm), with scarce mobility on deeper planes and no signs of local inflammation, which was covered with skin that was intact. A second lesion (2 cm × 1 cm) with similar characteristics was on the right gastrocnemius. Magnetic resonance imaging (MRI) with a contrast revealed a large, very heterogeneous mass that was sarcomatous in appearance on the vastus externus muscle of the left thigh (Fig. 1), and a smaller-sized mass in the lateral gastrocnemius of the right leg. A cytology obtained from this latter lesion, using fine-needle aspiration (FNA), was consistent with a poorly differentiated carcinoma. Later, an incisional biopsy of the thigh mass confirmed infiltration by an epidermoid carcinoma with complex epithelial phenotype in the immunohistochemistry study (positivity for high molecular weight keratins, pankeratins AE1–AE3, thrombomodulin, carcinoembryonic antigen and p63). The study was completed with a thoracoabdominal computed tomography study that revealed multiple bilateral pulmonary nodules indicative of metastasis, among which there was a larger lesion in the right upper vertex (3 cm × 3.2 cm) with spiculated contours that was identified as a probable primary tumor. By means of transthoracic FNA, a sample was obtained, whose cytological study was compatible with a scarcely differentiated carcinoma. With the determination of the lesion as a stage IV non-small-cell lung cancer
(NSLC), the patient underwent palliative chemotherapy (cisplatin and docetaxel). Despite the initial decrease in size of the SMM after 6 cycles of treatment, the tumor progression was confirmed 10 months later in the lungs, brain and liver. At that moment, we opted for symptom management.

Various mechanisms have been invoked to justify the apparent resistance of the muscles to the metastatic infiltration, among which there are mechanical factors (tissue pressure), metabolic factors (production of lactic acid and oxygen free radicals, local pH) and immunological factors. The primary tumor that is most frequently involved in the majority of series is bronchogenic carcinoma, followed by gastrointestinal and urothelial neoplasms. From a clinical standpoint, SMMs usually present as nodules or painful masses with signs of local inflammation, and among their most frequent locations are the lower extremities, the chest wall and the paravertebral musculature. MRI is the radiological technique of choice, even though its typical findings (tissue hyperintensity in the T2 sequences, signal reduction in T1 sequences and irregular captation of gadolinium) are relatively unspecific. Therefore, when using this diagnostic approach, it is crucial to also have a histological exam, which, on occasions, should be completed with immunohistochemistry techniques given the limited differentiation that the SMM can exhibit. An exhaustive review of the literature identified 114 cases of NSLC associated with SMM, with a mean survival of 6 months; the existence of a single metastasis and its metachronous presentation with the primary tumor defined a subgroup of patients with better prognoses. In this as well as in other series, the initial presentation of NSLC in the form of SMM in the lower extremities, with no evidence of previous dissemination to other anatomically more predictable locations, is very infrequent. This present case illustrates the need, in cases with similar scenarios, to carry out a complete extension study with histological determination which can exclude other more prevalent situations, particularly the possibility of a soft tissue sarcoma with lung metastasis, given the therapeutic implications of this differential diagnosis.

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Mario Fernández-Ruiz*, Juan Vila-Santos, Juan Manuel Guerra-Vales

**Servicio de Medicina Interna, Hospital Universitario 12 de Octubre, Madrid, Spain**

*Corresponding author.

E-mail address: mario_fdezruiz@yahoo.es (M. Fernández-Ruiz).