Spontaneous regression of pulmonary alveolar proteinosis: a case report

Regressão espontânea de proteinose alveolar pulmonar: relato de caso

Rodrigo Canellas de Souza¹, Daniel Kanaan², Pedro Henrique Rodrigues Martins³, Guilherme Abdalla Gavinho Vianna⁴, Viviane Brandão Amorim⁴, Edson Marchiori⁵

INTRODUCTION

Pulmonary alveolar proteinosis (PAP) is characterized by alveolar filling with protein material. Such material can be detected by periodic acid-Schiff staining (PAS) and has high lipid content. PAP is most commonly diagnosed in adult individuals between 20 and 50 years of age, but it is also reported in other age groups.

The most accepted hypothesis is that PAP results from an abnormal production, altered metabolism or depuration of pulmonary surfactant by type II alveolar cells and macrophages. Most PAP cases are idiopathic, although in some cases the condition results from exposure to silica (silico-proteinosis), association with hematologic diseases (for example: lymphoma and leukemia) or with infection by the human immunodeficiency virus.

CASE REPORT

A 21-year-old female presenting with a six-month history of progressive dyspnea, dry cough and weight loss of approximately 6 kg. The patient did not report fever or chest pain and did not present a history of smoking. Physical examination demonstrated normal heart and lung auscultation, with no cyanosis or digital clubbing. Blood count and serum biochemistry test results were also normal.

Chest radiography revealed symmetric reticular perihilar infiltrate, with confluence areas in the right lower lobe (Figure 1).

Canellas R et al.  Spontaneous regression of pulmonary alveolar proteinosis


ground glass attenuation and inter and intralobular septal thickening, characterizing the crazy-paving pattern, with areas of focal sparing (Figure 2). The patient was submitted to transbronchial lung biopsy, and the histopathological analysis identified alveolar filling with amorphous granular eosinophilic PAS-positive material, which preserved the alveolar septal architecture. Such characteristics were compatible with PAP diagnosis. The patient was followed-up on an outpatient basis, with no specific treatment and two years later presented significant improvement of clinical symptoms and pulmonary opacities at HRCT (Figure 3).

**DISCUSSION**

PAP is a diffuse pulmonary disease characterized by accumulation of amorphous lipoproteinaceous, PAS-positive material in the distal air spaces\(^1,3\). Typically, there is little or no pulmonary inflammation and the adjacent pulmonary architecture is preserved\(^1,3\). The following three forms of this condition are recognized: congenital, secondary and acquired PAP.

The congenital form occurs in the neonatal period and seems to be either a result of mutations in genes that encode the surfactant, the macrophage colony stimulating factor receptor and granulocytes (GM-CSF), or the result of a defect in the plasma membrane of the cationic amino acid transporter (known as lysinuriic protein intolerance)\(^4\).

The secondary form of PAP develops in the adulthood, generally as a result of exposure to high dust levels (for example: silica, aluminum and titanium), malignant hematologic diseases or after allogeneic bone marrow transplant\(^4\). In Brazil, silicoproteinosis is the most common form of secondary alveolar proteinosis\(^5,7\).

The acquired form is associated with a high prevalence of anti-GM-CSF auto-antibodies and is the most common form of PAP\(^4\).

Patients with idiopathic or secondary PAP present with moderate and nonspecific respiratory symptoms such as progressive dyspnea which may occur for months or years, besides dry or minimally productive cough. Less common signs and symptoms include weight loss, fatigue, low fever, chest pain and hemoptysis\(^3,8\). Physical examination may reveal crepitations, clubbing or cyanosis\(^8\). The mean age at PAP diagnosis is 40 ± 13 years; and a strong association with smoking has been observed\(^3,8\).

Patients with PAP present greater predisposition to pneumonic infection. Infec-

---

**Figure 2.** HRCT at two different levels showing the crazy paving pattern. Areas of ground-glass attenuation and intralobular septal thickening are observed in association with areas of focal sparing.

**Figure 3.** HRCT at the same levels as those on Figure 2. Images acquired obtained two years after the diagnosis showing marked improvement of the lesions.
tious pneumonias in PAP patients are many times opportunistic infections\(^9\).

The diagnosis of PAP usually requires chest radiography, although it is not conclusive for such diagnosis. Radiography can detect bilateral, symmetrical and central opacities\(^1(1,8)\). Additionally, many times there is a notable disparity between the clinical symptoms of PAP and radiographic changes (“clinical-radiological dissociation”)\(^8\). HRCT provides more anatomical details and information on the disease extent. The crazy paving pattern, which corresponds to the septal thickening overlapping ground-glass attenuation areas is the most frequent tomographic finding associated with PAP\(^10,11\). Regions with crazy paving pattern are typically generalized and bilateral, many times sparing well delimited areas or even an entire pulmonary lobe\(^12\).

Although the crazy paving pattern is frequently detected at HRCT in PAP patients, such finding is also observed in several infectious, hemorrhagic, neoplastic, idiopathic disorders, as well as in pulmonary edema\(^11,13\).

In summary, the present report describes the case of a previously healthy, young, female patient with no history of heart problems or occupational disease, presenting crazy paving pattern at HRCT. The diagnosis of pulmonary alveolar proteinosis was confirmed by biopsy. Two years after diagnosis, the patient is in good clinical conditions, with marked improvement of the pulmonary opacities.

REFERENCES