Respiratory bronchiolitis associated with interstitial lung disease

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Case report

A 52 year-old man, who had been smoking about 20 cigarettes per day for thirty years, worked for about 40 years in mechanical workshops on the maintenance staff repairing diesel engines and pneumatic brakes. The inherent risks in this kind of work are exposure to different toxic substances, many of which are considered irritating to the respiratory apparatus: acid fumes (electric battery), exhaust fumes, powders from braking systems, asbestos fibres, exhalation from oil and oily liquids. The man suffered for years from a persistent cough in the mornings and light exertional dyspnoea. In May 2000 he underwent a chest X-ray for the Department of Occupational Medicine, which evidenced reticular micro-nodular interstitial lung disease and in June 2000 he was sent to the Pulmonary Department of the Hospital of Sarzana, La Spezia, Italy. On physical examination, the chest, abdomen and lymph node examinations were negative.

Initial laboratory values showed a white blood cell count of 22,100/mcI with 60% neutrophils and 29.7% lymphocytes; blood platelets, 1,180,000; LDH, 992 (normal 230/460 IU/L); the other routine tests as well as the oncological markers were normal. A serial chest X-ray showed diffuse reticulo-nodular interstitial pattern.

A high resolution CT scan (HRCT) showed the presence of reticular thickening of the bronchiolar interstitium and of several small blisters of emphysema scattered mainly on the upper parts of the lungs on both sides (fig. 1). There was also the presence of a few subpleural micro-nodules.

The ECG and cardiac ultrasonography were normal.

The pulmonary function tests, lung volumes and CO diffusion capacity were normal. A bronchostimulation test with methacoline was negative. Arterial blood gas values were as follows: pH 7.39; pO₂ 9.2 kPa; pCO₂ 6.3 kPa and arterial oxygen saturation, 93.1%.

The bronchoscopy showed the tracheal and first bronchial tracts explorable without significant abnormalities. The transbronchial biopsies showed biopsy flaps of pulmonary parenchyma with slight sectorial thickening.

The result of BAL analysis was: macrophages 97%, neutrophils 2%, lymphocytes 1% with CD4/CD8 ratio 0.62. Surgical biopsy (lingula) was made in Video-assisted thoracoscopy (VATS) showing sections of pulmonary parenchyma (fig. 2) with focal thickening of the septa. In the peribronchial areas there was a slight inflammatory lymphohistiocyte infiltration and in the alveolar spaces the presence of pigmented macrophages. The staining for the Langherans cells shows a few elements in the peribronchial areas. The other ab-
Respiratory bronchiolitis associated with interstitial lung disease (RB-ILD) is a rare entity, more common in smokers, that should be distinguished from the other interstitial lung diseases and in particular from Desquamative Interstitial Pneumonia (DIP), although the two conditions share a similar histopathological pattern. In 1974 Niewoehner and al [1] described RB in an autopsy study of cigarette smokers who had died from a disease that was not associated with their lungs but was associated with light symptoms and clinical or functional abnormalities.

In 1986 Myers and al [2] described six patients who had clinical and/or radiographic symptoms of interstitial lung diseases, with RB as the only histopathological lesion in the biopsy specimens, suggesting that RB-ILD was the cause of the clinical findings.

A recent review of ten cases [3] showed an equal gender distribution with an age range of 36-65 years; nine patients out of ten were smokers (smoking from 3 to 80 packs per year). The non-smoker had been exposed to the fumes of solder flux in her employment. Symptoms were non specific with dyspnea in 70% of the cases, cough in 30%, haemoptysis in 20%; the physical examination showed basal inspiratory crackles in 60 per cent of the cases and digital clubbing in 20%; the pulmonary function tests showed both restrictive and obstructive patterns with severe reductions in the carbon monoxide transfer factor in seven out of ten patients.

Chest X-rays showed reticular or reticulonodular shadowing in 50% of the cases, a ground glass pattern in 30% and a normal pattern in 25%; CT scans showed a patchy ground glass pattern in
75%, a emphysematous pattern in 37.5%, dilatation of the airways in 25%, bronchial thickening in 12.5% and basal “honeycombing” in 12.5%.

Histologically, respiratory bronchiolitis superficially resembles DIP in that there is an accumulation of intra-alveolar macrophages associated with mild interstitial thickening [4]. It differs from DIP, however, in that the process is patchy rather than diffuse, and follows a strictly peribronchiolar distribution; the macrophages are present mainly within respiratory bronchioles and alveolar ducts. They may extend into adjacent alveolar spaces and focally are present in the interstitium. The macrophages have abundant cytoplasm that contains fine granular yellow-brown pigmention. Moreover peribronchiolar fibrosis with mild peribronchiolar chronic inflammation and centrolobular emphysema [1-4] are present. Ultrastructural analysis showed several needle-shaped inclusions in the lysosomes and cytoplasmic phagolysosomes of the macrophages in the cell cytoplasm that could be aluminium silicate caused by cigarette smoking [3]. RB-ILD does not have a typical functional clinical or radiographic expression and the diagnosis can be made through a histopathological examination. The evolution is benign in 70 per cent of the cases with stabilisation or improvement of the clinical picture if the patient abstains from smoking; cortisone-based, immunodepressive therapies do not seem to affect the evolution of the disease [2]. RB-ILD is clearly connected with tobacco smoking and has been inserted in the “smoking related interstitial lung disease” together with DIP and cell histiocytosis of Langerhans [2, 4]. There are no data in the literature which associate BR-ILD with blood diseases.

This case of RB-ILD shows the importance, in the case of interstitial lung diseases, of histopathological diagnosis of tissue samples so as to have a nosographic definition, a targeted therapy and a correct prognosis. The case also shows that tobacco smoking is not only the cause of Chronic Obstructive Pulmonary Disease and lung cancer but also of lung interstitial diseases which can revert if the patient quits smoking. We confirm that the exposure to irritating fumes, especially in the work environment, may contribute to the pathogenesis of RB-ILD.

References