Probe-based confocal laser endomicroscopy in diagnosis of desquamative interstitial pneumonia in nonsmoker

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Abstract

Desquamative interstitial pneumonia is a rare disease that is predominantly associated with smoking. Surgical biopsy is usually recommended for the diagnosis of this disorder. Probe-based confocal laser endomicroscopy (pCLE) is a new method of minimally invasive in vivo microscopic image of airways and alveoli. We use this method for desquamative interstitial pneumonia in nonsmoker. The pCLE image shows thickened intra-alveolar septae and clusters of autofluorescent cells within the alveoli, which is unusual for non-smoking patients. We think this pattern can be used for differential diagnosis of desquamative interstitial pneumonia in nonsmokers.

Introduction

Desquamative interstitial pneumonia (DIP) is a rare disease from the group of idiopathic interstitial pneumonias [1]. This disorder was first described by Liebow in 1965 and called DIP because of the belief that the main histological feature was the desquamation of epithelial cells into alveoli [2]. Nowadays it is characterized by the accumulation of numerous pigmented macrophages within most of the distal airspace of the lung and, sometimes, the presence of giant cells [3]. DIP is usually associated with tobacco smoke and is classified as smoking-related idiopathic interstitial pneumonia [1]. But it can also occur in patients exposed to certain inhaled toxins (occupational exposure) and drugs, or in certain viral illnesses and autoimmune diseases [3]. According to scientific literature, approximately 10-42% of patients with DIP are nonsmokers [3]. Differential diagnosis of DIP is difficult and transbronchial lung biopsy has a low diagnostic yield, thus surgical biopsy is required to make a confident diagnosis [4]. However, surgical lung biopsy could have complications such as prolonged air leak, acute exacerbation or even death especially in patients with immunocompromised status, mechanical ventilation dependence, or severe respiratory dysfunction [5,6]. Probe-based confocal laser endomicroscopy (pCLE) is a new method of minimally invasive in vivo morphological diagnosis of airways and alveoli (so called “optical biopsy”) [7]. It is useful for a diagnosis of rare lung diseases such as, for example, alveolar proteinosis and metastatic pulmonary calcification [8,9].

We report a case of a diagnosis of DIP in nonsmoker using confocal laser endomicroscopy.

Case Report

A 59-year-old female was admitted to the Center of Thoracic Surgery of the “St. Petersburg Research Institute of Phthisiopulmonology” with complaints of shortness of breath with minimal exertion, moderate weakness, cough, fever by evening. From the anamnesis it is known that similar symptoms appeared for the first time in 2014, and since then she was treated with a diagnosis of pneumonia in several hospitals. Also it was known that the patient was a nonsmoker, who has been...
The patient completed a full range of examinations. Blood analysis showed mild signs of inflammation. Microbiology of sputum showed no bacteria. Spirometrical indicators were within normal limits. Computer tomography (CT) scan demonstrated multiple mosaic ground glass opacities and lymphadenopathy (Figure 1). By CT data, it was mostly resembled hypersensitivity pneumonitis, nonspecific interstitial pneumonia or alveolar proteinosis. Standard bronchoscopy revealed some signs of bronchitis and atrophy. To clarify the diagnosis, it was decided to perform a probe-based confocal laser endomicroscopy by Cellvizio system (Mauna Kea Technologies, Paris, France) and transbronchial biopsy.

The pCLE was performed during the bronchoscopy under the local anesthesia of 2% lidocaine. Probe Alveoflex was inserted through the working channel of the endoscope into the distal airways of the affected segments. In all areas studied, the pCLE image showed thickened intra-alveolar septae and a large number of blending autofluorescent cells (average diameter 20 µm) in alveoli (Figure 2). The pCLE image after bronchoalveolar lavage in one of the pathological zones did not show any changes. Transbronchial biopsy was obtained from an area of the largest accumulation of autofluorescence cells by pCLE data.

Histological examination of samples of transbronchial biopsy revealed focal accumulation of macrophages and lymphocytes on the background of moderate fibrosis. Unfortunately, tissue samples were too small to determine the uniformity of lung tissue involvement in the pathological process. The obtained histological picture could also correspond to respiratory bronchiolitis-interstitial lung disease, organizing pneumonia and other interstitial pneumonias. In this regard thoracoscopic lung biopsy was obtained. Histological examination of surgical biopsy samples showed picture typical for desquamative interstitial pneumonia: multiple clusters of macrophages and alveolar cells type II in the alveolar space, uniform thickening of the interalveolar septae (Figure 3). There was no dusty pigment in macrophages, which is usual for smokers. Diagnosis of desquamative interstitial pneumonia was established by multidisciplinary consensus of specialists by the data of all examinations. After that, the patient was prescribed the correct treatment.

Discussion

Due to difficulties in diagnosis of DIP surgical lung biopsy is usually recommended [4]. The pCLE is a minimally invasive method of morphological diagnosis of lung tissue that could give some additional information about diagnosis. It is known that macrophages are visualized by pCLE mostly due to accumulation of tobacco-tar-induced fluorescence [7]. But there are some conditions when we can see highly autofluorescent cells in nonsmokers by pCLE, for example amiodarone-related pneumonia or acute lung allograft rejection [10,11]. Most authors supposed that these cells are so called activated macrophages or/and alveolar type II cells [11].

We used pCLE in nonsmoking patient with DIP and found clusters of autofluorescent cells within the alveoli in all studied areas. At the same time, the samples obtained during transbronchial biopsy also contained focal accumulation of macrophages. But specimens
were too small to determine how uniform the presented changes were. A larger thoracoscopic biopsy sample proved the uniformity of pathological changes in lung tissue.

Thus, we proved the correlation between the pCLE image and the histopathological picture in DIP. We found only one research in scientific literature with pCLE image of DIP without reference about smoking status of a patient [12]. Some authors have proven that the hypercellular pattern (presence of a large number of autofluorescent cells) is less frequent in chronic fibrosing pneumonias [13]. But we found that DIP is characterized by such pattern, and think that it can be used in the differential diagnosis. Perhaps pCLE can also contribute to the correct interpretation of the histological pattern of transbronchial biopsy samples. Of course it is only one case and more research is needed to confirm this. To sum up, we think that the presence of autofluorescent cells in pCLE image in DIP in nonsmokers could give some additional information for differential diagnosis of interstitial lung diseases.

References
