

Bronchiectasis: Treatment of Breathing Difficulties

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Introduction

Bronchiectasis is a disease that contains a substance that enlarges bronchi and bronchioles, decreasing resistance in the respiratory airway and increasing airflow to the lungs to make breathing easier. Bronchiectasis may be originated naturally within the body or they may be used for the treatment of breathing difficulties. They are useful in obstructive lung disease such as asthma and in chronic obstructive pulmonary disease. The impact on healthcare systems is substantial [1]. A recent multicentre European study of patients with bronchiectasis identified an annual exacerbation frequency per patient per year, with a hospitalisation rate over few years follow-up. Bronchiectasis has a clear attributable mortality. In the largest cohort study reported to date, half of the patients died from respiratory causes, with around one-quarter dying from cardiovascular diseases. Loebinger provided long-term data on mortality by following up a cohort of patients first recruited for the validation of the St. Georges Respiratory Questionnaire. These patients were followed up for years. In a prospective cohort analysis of patients in secondary care in Belgium, Goeminne found that deaths were respiratory related and remaining were cardiovascular. Therefore, it is clear, at least in secondary care bronchiectasis cohorts, that patients experience a high rate of exacerbations, hospital admissions and attributable mortality, emphasising the need for high-quality specialised care for these patients. The pathophysiology of bronchiectasis and the goals of treatment our understanding of the pathophysiology of bronchiectasis is limited, in part because of the lack of representative experimental models. Airway inflammation in bronchiectasis is dominated by neutrophils, driven by high concentrations of neutrophil chemo-attractants such as interleukin and leukotriene [2]. Airway bacterial colonisation occurs because of impaired mucociliary clearance and because of failure of neutrophil opsonophagocytic killing. Since neutrophils from bronchiectasis patients are believed to be normal prior to their arrival in the airway, it is likely that the airway inflammatory milieu itself impairs bacterial clearance. Work over several decades has implicated neutrophil elastase in this process.

The effects of elastase on airway epithelial cells includes slowing of ciliary beat frequency and promotion of mucus hypersecretion while impairment of opsonophagocytosis occurs at multiple levels, through cleavage of opsonins from the bacterial surface and cleavage of the neutrophil surface receptors FcγRIIIb and CD11b. Alpha defensins released from neutrophil granules also suppress phagocytic responses. Other mechanisms of immune dysfunction include failure of clearance of apoptotic cells and T cell infiltration, with recent evidence pointing to an important role of T cells.

Discussion

Nevertheless, much more work is needed to unravel the

complexities of the host response in bronchiectasis. Significant recent advances in our understanding of bronchiectasis have arisen through rRNA sequencing technologies which allow a comprehensive analysis of polymicrobial bacterial communities in the lung. Such technologies have clearly disproven the previous teaching that the healthy airway is sterile [3]. Studies in bronchiectasis reveal colonisation with familiar pathogens such as *Haemophilus* sp., *Pseudomonas aeruginosa* and

break the cycle. As with other respiratory diseases, patients with bronchiectasis should be encouraged to stop smoking. Vaccination against influenza and pneumococcal disease is also recommended as for other chronic respiratory disorders although there are no specific data in bronchiectasis about its impact. Bronchiectasis represents the final common pathway of a number of diseases, many of which require specific treatment. Host-infectious bronchiectasis is often used as a diagnostic label for patients with a history of severe or childhood respiratory infections, affecting patients. There is little evidence so far that they represent a distinct phenotype from idiopathic bronchiectasis and some cases may represent recall bias [5]. Less data on aetiology is available outside the UK, but data from Italy and Belgium suggested a spectrum similar to the UK with perhaps fewer patients with allergic broncho-pulmonary aspergillosis and more with chronic obstructive pulmonary disease. Data from the USA clearly demonstrate more bronchiectasis due to non-tuberculous Mycobacteria in some centres, and a report by patients identified aetiology in few of cases. The BTS guidelines recommend testing for underlying causes including measurement of immunoglobulin, testing to exclude ABPA and specific antibody responses to pneumococcal and Haemophilus vaccination. Sputum culture to exclude NTM and measurement of autoantibodies are also suggested. Testing for CF is recommended for patients with recurrent P. aeruginosa and Staphylococcus aureus isolation, or upper lobe predominant disease irrespective of age. Additional testing is recommended in specific circumstances. COPD appears to be a very common aetiology, with bronchiectasis reported in up to 20% of patients with moderate-to-severe COPD. Bronchiectasis also appears relatively common in patients meeting the diagnostic criteria for asthma. Focal bronchiectasis may be associated with bronchial obstruction. Gastro-oesophageal reflux frequently co-exists with bronchiectasis and has been suggested as an aetiological factor in some cases. Immunoglobulin replacement, steroids and anti-fungal for ABPA, treatment for NTM and of CF all represent opportunities to specifically treat the underlying cause and so systematic testing of all