

CASE REPORT

ALLERGIC BRONCHOPULMONARY ASPERGILLOSIS WITHOUT BRONCHIAL ASTHMA: A RARE OCCURRENCE

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ABSTRACT

Allergic bronchopulmonary aspergillosis (ABPA) is a complex clinical entity resulting from an allergic immune response to *Aspergillus* species and most often occurs in patients with asthma and cystic fibrosis (CF). ABPA is rarely observed in the absence of asthma which is in fact the principal criterion for its diagnosis. We report a case of a 31-yr-old young man who had all other criteria except Bronchial asthma and was diagnosed as a case of ABPA.

Key Words: ABPA, Allergic bronchopulmonary Aspergillosis, Allergic lung disease.

INTRODUCTION:

Allergic Bronchopulmonary Aspergillosis (ABPA) is an allergic lung disease caused by hypersensitivity reactions to antigens of *Aspergillus* species, which is a genus of fungi with worldwide distribution¹. Diagnostic criteria for ABPA include the presence of bronchial asthma, immediate skin test reactivity to *Aspergillus fumigatus*, elevated total and *A.fumigatus* specific IgE levels, pulmonary infiltrates (transient or fixed), central bronchiectasis, peripheral blood eosinophilia and presence of precipitins against *Aspergillus* antigen². Bronchial Asthma is considered to be essential for diagnosis and is thought to play an important role in the development of the disease³. Majority of the patients of ABPA have asthma or cystic fibrosis, however there are several case reports of ABPA without coexisting bronchial asthma or cystic fibrosis⁴⁻⁸.

CASE REPORT:

A 31-yr-old man visited our hospital for evaluation of chronic cough which had persisted for eight months with minimal expectoration coupled with an intermittent low-grade fever and myalgia. He had been taking treatment from private doctors and now had been advised to take anti tubercular treatment. He came to us for confirmation of his diagnosis of pulmonary tuberculosis. The patient reported no history of wheezing or dyspnea nor did he have any personal or family history of atopy. The patient did not smoke and had no previous history of diseases like bronchial asthma or pulmonary tuberculosis. His chest was clear on auscultation. Other physical examination showed no abnormal findings.

On investigations the patient had a total leukocyte count of 8,300 cells/mm³ with 47% neutrophils, 21% lymphocytes and 32% eosinophils. His sputum & sputum collected by induction method both were negative for AFB. Intradermal tests with *Aspergillus fumigatus* spp elicited strong type I hypersensitivity reaction. His total serum IgE level estimated using fully automated chemiluminiscence system was 11583 IU/mL(N~ 100 IU/ml). Specific IgE against *A.fumigatus* was >100 KU/L(N~0.24KU/L) .

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His specific IgG levels against *Aspergillus* were 233 IU/ml (N~40 IU/ml). His sputum cultures yielded no growth of *Aspergillus* species. Spirometry was normal, with a forced vital capacity (FVC) of 2.97 L (82% of predicted), a forced expiratory volume in 1 sec (FEV₁) of 2.52 L (85% of predicted) and an FEV₁/FVC ratio of 85%, there was no bronchodilatation after administering a bronchodilator. Post exercise Spirometry was also normal. The methacholine challenge test was not performed as patient didn't agree for the same.

The patient's chest radiograph revealed a heterogenous opacity in the left upper lobe with bronchiectatic changes in both lung fields (Fig I). A chest computed tomography scan displayed consolidation in the left upper lobe, central bronchiectasis and mucous impacted dilated bronchi in bilateral upper lobes (Fig II). Air space nodules with few of them forming tree in bud pattern were also seen in posterior segment of right upper lobe and anterior basal segment of left upper lobe.

A diagnosis of ABPA was thus made as the patient fulfilled majority of the criteria given by Rosenberg et al¹. He was treated with prednisolone 0.75 mg/kg for 6 weeks, 0.5 mg/kg for next 6 weeks, and then tapered by 5 mg every 6 week to continue for a total duration of 9 months⁹. The patient showed marked improvement in his condition and there was marked clearing of his X-ray after 2 months of treatment (Fig III). His total serum IgE after two months of treatment decreased significantly to 4784 IU/ml. The patient is still visiting this hospital for follow up every month. No symptoms or signs of asthma have appeared till now.

DISCUSSION:

ABPA is a hypersensitivity disorder induced by *Aspergillus* species colonizing the lung cavity predominantly in patients with asthma¹⁰. It is the most frequently recognized manifestation of allergic aspergillosis occurring worldwide. ABPA was first reported in the United Kingdom in 1952¹¹ and since then has been reported worldwide. From India the first 3 cases were reported in 1971¹². Although Rosenberg-Patterson criteria^{1,13} are most often used for the diagnosis, there is still no consensus on the number of criteria needed to diagnose ABPA and patients in different stages may not fulfill these criteria¹⁴.

Despite the fact that ABPA is usually seen in association with bronchial asthma, ABPA has also be known to be an important complication of pulmonary disease associated with cystic fibrosis. ABPA has been variously reported in 1 to 15% of CF patients. The diagnosis of ABPA in CF is more complicated and disagreement exists in the literature regarding the diagnostic criteria. The difficulty lies in the fact that the usual criteria for ABPA and the common signs and symptoms of CF overlap.¹⁵

Bronchial asthma has classically been considered an essential diagnostic criterion for ABPA and has also been believed to play a crucial role in its development¹. It is considered that role of bronchial asthma in the development of ABPA is to make the expectoration of *Aspergillus* difficult and colonization easy by the viscous sputum⁵. ABPA is rarely thought of in the absence of asthma as this is the first criterion for diagnosis. However several cases without bronchial asthma have been reported⁴⁻⁸. Glancy et al⁴ reported that 11 out of 42 patients with ABPA had no evidence of bronchial asthma. A

systematic MEDLINE search performed by Agarwal et al⁷ for the occurrence of ABPA without bronchial asthma, were able to record 36 cases reported across the globe.

Because of the absence of bronchial asthma, these cases are often mistaken initially for other pulmonary disorders like bronchogenic carcinoma^{6, 16-17} or pulmonary tuberculosis⁷. Furthermore, the remarkable radiologic similarity to pulmonary tuberculosis has important clinical implications in high tuberculosis prevalent areas¹⁸.

Absence of bronchial asthma in our case was a reason for the delay in diagnosis of ABPA. Peripheral blood eosinophilia prompted further investigations which clinched the diagnosis of ABPA without bronchial asthma. The patient showed marked improvement with steroids and is symptom free for past one year. It has been stated that asthma may not be the initial criteria but may appear in the later stages of the disease¹⁹. Absence of asthma can be a good prognostic factor showing the early stages of the disease²⁰.

To conclude, we suggest that although ABPA most commonly occurs in patients with pre-existing bronchial asthma, a high index of suspicion should be maintained in the absence of asthma. ABPA without clinical asthma can and does pose diagnostic difficulties as in our case. Thus ABPA should be kept as a diagnostic possibility in patients with radiographic abnormalities and peripheral eosinophilia but no history or symptoms related to bronchial asthma.

REFERENCES:

1. Rosenberg M, Patterson R, Mintzer R, Cooper BJ, Roberts M, Harus K. Clinical and immunologic criteria for the diagnosis of allergic bronchopulmonary aspergillosis. *Ann Intern Med* 1977;86:405-14.
2. Aggarwal R, Gupta D, Aggarwal AN, Saxena AK, Chakrabarti A, Jindal SK. Clinical significance of hyperattenuating mucoid impaction in allergic bronchopulmonary aspergillosis, an analysis of 155 patients. *Chest* 2007;132:1183-90.
3. Slavin RG, Fischer VW, Levine EA, Tsai CC, Winzenburger P. A primate model of Allergic Bronchopulmonary Aspergillosis. *Int Arch Allergy Appl Immunol* 1978;56:325-33.
4. Galancy JJ, Elder JL, McAleer R. Allergic Bronchopulmonary fungal disease without clinical Asthma. *Thorax* 1981;36:345-9.
5. Hoshino H, Tagaki S, Kon H, Shibusu T, Takabatake H, Fujita A et al. Allergic bronchopulmonary aspergillosis due to *Aspergillus niger* without bronchial asthma. *Respiration*. 1999;66(4):369-72.
6. Shah A, Maurya V, Panjabi C, Khanna P. Allergic bronchopulmonary aspergillosis without clinical asthma caused by *Aspergillus niger*. *Allergy* 2004;59:236-7.
7. Agarwal R, Aggarwal AN, Gupta D, Bal A, Das A. Case report: A rare cause of miliary nodules -- allergic bronchopulmonary aspergillosis. *Br J Radiol*. 2009;82(980): 151-4.
8. Koh WJ, Han J, Kim TS, Lee KS, Jang HW, Kwon OJ. Allergic bronchopulmonary aspergillosis coupled with broncholithiasis in a non-asthmatic patient. *J Korean Med Sci*. 2007;22(2):365-8.

9. Agarwal R, Gupta D, Aggarwal AN, Behera D, Jindal SK. Allergic bronchopulmonary aspergillosis: lessons from 126 patients attending a chest clinic in north India. *Chest*. 2006 Aug;130(2):442-8.
10. Kumar R, Chugh T, Gaur SN. Allergic Bronchopulmonary Aspergillosis - A review. *Indian J Allergy Asthma Immunol* 2003;17(2):55-66.
11. Hinson KF, Moon AJ, Plummer NS. Bronchopulmonary aspergillosis: A review and report of eight new cases. *Thorax* 1995;7:313-33.
12. Shah JR. Allergic Bronchopulmonary Aspergillosis. *J Assoc Phys India* 1971;19:835-41
13. Greenberger PA, Patterson R, Ghory A, Arkins JA, Walsh T, Graves T et al. Late sequelae of allergic bronchopulmonary aspergillosis. *J Allergy Clin Immunol* 1980;66(4):327-35.
14. Bedi RS, Bedi GK. Allergic Bronchopulmonary Aspergillosis: Indian scenario. *Lung India* 2007;24:156-61.
15. Nepomuceno IB, Esrig S, Moss RB. Allergic Bronchopulmonary Aspergillosis in Cystic Fibrosis. Role of Atopy and Response to Itraconazole. *Chest* 1999;115:364-70.
16. Berkin KE, Vernon DR, Kerr JW. Lung collapse caused by allergic bronchopulmonary aspergillosis in non-asthmatic patients. *BMJ (Clin Res Ed)* 1982;285(6341):552-3.
17. Bondue B, Rimmelink M, Gevenois PA, Yernault JC, De Vuyst PA pulmonary cavitated mass complicating long-standing allergic bronchopulmonary aspergillosis. *Respir Med Extra* 2005; 1:39-42.
18. Shah A, Panjabi C. Allergic bronchopulmonary aspergillosis: a review of a disease with a worldwide distribution. *J Asthma* 2002;39:273-89.
19. Greenberger PA, Patterson R. Allergic Bronchopulmonary Aspergillosis and the evaluation of the patient with asthma. *J Allergy Clin Immunol* 1988;81:646-50.
20. Kumar R. Mild moderate and severe forms of Allergic Bronchopulmonary Aspergillosis: A Clinical and Serologic evaluation. *Chest* 2003;124:890-2.

Legends

Fig I: Xray chest showing heterogenous opacity in the left upper zone and bronchiectatic changes in both lung fields.

Fig II: CT chest showing consolidation in the left upper lobe, central bronchiectasis and mucous impacted dilated bronchi in bilateral upper lobes.

Fig III: Xray chest after 8 weeks showing marked clearing of the lesions.
Allergic bronchopulmonary aspergillosis (ABPA) is a complicating factor of cystic fibrosis which can result in a devastating combination as lung disease progresses. The overlap between the signs and symptoms of the two conditions makes diagnosis problematic.