Endobronchial aspergilloma: A case report and literature review

DONGDONG HUANG^{1*}, BING LI^{2*}, HAIQING CHU², ZHEMIN ZHANG², QIUHONG SUN², LAN ZHAO², LIYUN XU², LI SHEN², TAO GUI², HUIKANG XIE³ and JUN ZHANG⁴

¹Department of Clinical Medicine, Tongji University School of Medicine, Shanghai 200092; Departments of
 ²Respiratory Medicine and ³Pathology, Shanghai Pulmonary Hospital, Tongji University School of Medicine, Shanghai 200433, P.R. China; ⁴Division of Hematology, Oncology and Blood & Marrow Transplantation, Department of Internal Medicine, Holden Comprehensive Cancer Center, University of Iowa Carver College of Medicine, Iowa, IA 52242, USA

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Abstract. The present study aimed to investigate the clinical and radiological characteristics in addition to the bronchoscopic appearance in patients with endobronchial aspergilloma (EBA). Clinical and radiological characteristics were analyzed alongside the bronchoscopic appearance in 17 patients with EBA diagnosed by bronchoscopy with histological examination. The present study assessed the relevant literature and 13 males and 4 females were included in the comparison, with a median age of 59. Associated diseases included 8 previous diagnoses of pulmonary tuberculosis (47.6%), 4 previous diagnoses lung cancer (23.5%), 1 pulmonary resection (5.9%) and 1 bronchial foreign body (5.9%). The primary symptom was hemoptysis (9/17, 53%). Chest computed tomography (CT) indicated a markedly higher incidence of aspergillosis lesion in

Correspondence to: Professor Haiqing Chu or Professor Zhemin Zhang, Department of Respiratory Medicine, Shanghai Pulmonary Hospital, Tongji University School of Medicine, 507 Zhengmin Road, Shanghai 200433, P.R. China

E-mail: chu_haiqing@126.com E-mail: zhemindoc@163.com

*Contributed equally

Abbreviations: EBA, endobronchial aspergilloma; CT, computer tomography; IPA, invasive pulmonary aspergillosis; CAN, chronic necrotizing aspergillosis; ABPA, allergic bronchopulmonary aspergillosis; SIA, semi-invasive aspergillosis; FDG, ¹⁸F-deoxyglucose; FOB, fiberoptic bronchoscopy; DCMP, dilated cardiomyopathy; HT, hypertension; NSCLC, non-small cell lung cancer; ET, essential thrombocytosis; SDH, subarachnoid hemorrhage; ICH, intracranial hemorrhage; TB, tuberculosis; BW, bronchial washing; MTB, Mycobacterium tuberculosis; IV, intravenous; GGO, ground glass opacity; LLL, left lower lobe; LUL, left upper lobe; RML, right middle lobe; RUL, right upper lobe; RBI, right bronchus intermedius

Key words: fungi, endobronchial aspergilloma, bronchoscopy

the left lung (13/17; 76.5%) compared with the right lung (4/17; 23.5%). CT manifestation included space occupying disease in 10 patients (58.8%), aspergilloma in 3 patients (17.6%), pneumonic consolidation in 2 patients (11.8%) and ground glass opacity in 1 patient (5.9%). Bronchoscopy examination identified masses in all 17 patients' bronchial lumen and 15 patients had endobronchial obstruction by necrotic material. The case presented in the current study demonstrated the merits of combining bronchosopic intervention with voriconazole. The dominant symptom of EBA was hemoptysis. Chest CT demonstrated that aspergillosis lesions were more frequently identified in the left lung compared with the right. EBA often occurs in individuals with underlying lung diseases, which cause lumen structural change or bronchial obstruction. EBA may be clearly diagnosed by bronchoscopy biopsy, although the potential for a co-exististing tumor requires consideration. Bronchoscopic intervention and anti-fungal therapy may have an advantage in the effective treatment of patients with EBA.

Introduction

Aspergillus spp. are a ubiquitous fungus in the environment, which are easily identified in soil, water and various types of decomposing organic matter (1). Inhalation of Aspergillus spores causes pulmonary aspergillosis, which is the most common form of aspergillus filamentous fungal infection, and is often diagnosed in patients with immune deficiencies or systemic diseases (2). Aspergillosis is currently divided into the following categories (3): i) Invasive pulmonary aspergillosis (IPA), a severe disease and major cause of mortality in severely immunocompromised patients; ii) chronic necrotizing aspergillosis (CNA), presents as a locally invasive disease and is observed primarily in patients who are mildly immunocompromised or have chronic lung disease; and iii) allergic bronchopulmonary aspergillosis (ABPA), a non-invasive hypersensitivity pulmonary disease caused by Aspergillus. Aspergilloma is a fungus ball that develops in a pre-existing cavity in lung parenchyma, which may be easily detected by X-ray and computed tomography (CT) scans (3). In addition, a novel category of aspergillosis with semi-invasive form (SIA) has been described (4).

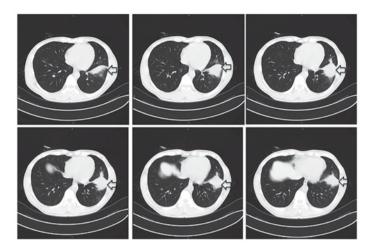


Figure 1. Chest computed tomography scan of a 48-year-old male patient revealed a spiculated mass and exudation in left lower lobe (arrow). The patient initially presented with a recurrent fever, cough, expectoration and bloody sputum lasting 3 months.

Endobronchial aspergilloma (EBA) is a rare disease, described as a massive non-invasive overgrowth of the aspergillus species in bronchial lumen, causing space-occupying lesions (5,6). It is often incidentally diagnosed during bronchoscopy to evaluate the cause of hemoptysis (5,6), and may be considered as another form of pulmonary aspergillosis. At present, there are very few reports regarding the characteristics of EBA. The present study aimed to investigate the clinical and radiological characteristics in addition to the bronchoscopic appearance in patients with EBA through a representative case from Shanghai Pulmonary Hospital affiliated to Tongji University (Shanghai, China) and a review of the current literature.

Case report

Study design. The present study reports a case of EBA with no prior treatment that was diagnosed at Shanghai Pulmonary Hospital affiliated to Tongji University, informed consent was also obtained from the patient. A retrospective analysis of the clinical and radiological characteristics and bronchoscopic results was also completed by combining this case with 16 other previously reported EBA cases (5-10). All patients we subjected to a chest CT, bronchoscopy and histological examination. The tissue used for histological examination was obtained via bronchoscopy biopsy of the mass in the trachea, avoiding necrotic tissue or secondary infection areas, with a volume of at least 0.5x0.5x0.2 cm and had a fresh non-necrotic color. The tissue was fixed using 10% neutral formaldehyde for 24 h at room temperature, routinely selected, embedded, sliced in continuous sections with a thickness of 4 µm and hematoxylin-eosin (HE) stained. The test results and basic clinical data were collected. Primary evidence to confirm EBA were as follows: i) Presence of bronchoscopy-visible intraluminal mass and necrotic tissue; ii) characteristic appearance of Aspergillus species under microscopic examination, such as septate fungal hyphae bearing acute branching angles; iii) positive result of Aspergillus culture; and iv) following the elimination of other possibilities, such as tumors, foreign bodies, parasites or mucus by evaluating the patient's personal history, performing bronchoscopy and pathology examinations.

Representative case. A 48-year-old male was admitted to Shanghai Pulmonary Hospital affiliated to Tongji University due to a recurrent fever, cough, expectoration and bloody sputum lasting 3 months. The patient had a 20-year history of smoking, and a 4-year history of diabetes and was undergoing diet therapy. On admission, he presented with normal vital signs. Physical examination of the chest revealed no evident crackles or wheezing. Complete blood count results were normal. Routine blood chemistry and tumor markers were all within normal limits. Mycodextranase and latex agglutination tests from the blood were negative. Human immunodeficiency virus antibody test was negative. Chemical luminescence of syphilis test was negative.

An intial CT scan of the chest revealed a spiculated mass with exudation in the left lower lobe (Fig. 1). ¹⁸F- deoxyglucose (FDG) dual-head coincidence single-photon emission computerized tomography (DHTC) indicated that the maximum tumor to non-tumor ratio was 11.63 and therefore, the possibility of lung cancer was not eliminated (Fig. 2). Initial fiberoptic bronchoscopy (FOB) examination (11) revealed a white necrotic mass in the medial anterior basal segment of left lower lobe, which completely obstructed the subsegmental bronchus (Fig. 3). Removal of the mass was attempted but was not successful as it was impacted in the narrow lumen. Notably, histopathological examination performed using HE staining of the bronchoscopic biopsied specimen was indicated to be negative. FOB was performed a second time 2 days later and a purulent material partially obstructing the lumen was identified at the same site. The lumen was finally unblocked following repetitive clearing of the necrotic material and sputum aspiration. The clearing work performed with a BF-1T260 bendable bronchoscopy (Olympus, Ishikawa, Japan) and biopsy forceps (Aohua, Shanghai, China), to cut and remove the tissue, 10 min a time for a total of 3 times. Histopathological examination of the bronchoscopic biopsied specimen indicated the presence of necrotic tissue and Aspergillus. Periodic acid-Schiff staining was performed and samples were exposed to 0.5% periodic acid for 5 min, Schiffs' reagent for 15 min and then washed twice using 0.5% sodium metabisulfite for 1 min each time before bright green staining was performed for 10 sec at room temperature. Furthermore, hexamine silver staining

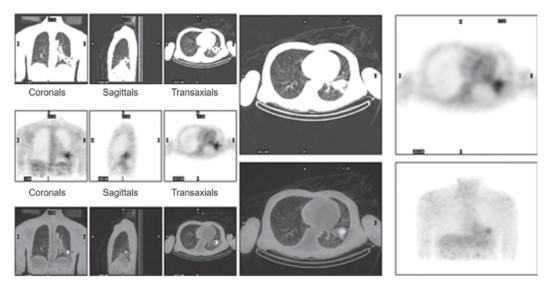


Figure 2. ¹⁸F-deoxyglucose dual-head coincidence single-photon emission computerized tomography scan indicated that the maximum tumor/non-tumor was 11.63 and therefore, lung cancer should not be excluded.

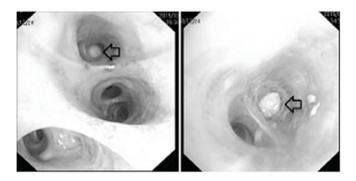


Figure 3. Fiberoptic bronchoscopy revealed a white necrotic mass in the medial anterior basal segment of left lower lobe, which completely obstructed the subsegmental bronchus (arrow).

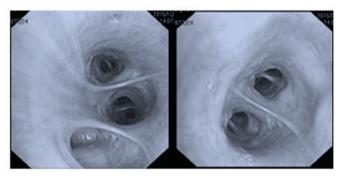


Figure 5. Following fiberoptic bronchoscopy, the patient was administered interventional voriconazole therapy for 1 month. A second bronchoscope assessment demonstrated that the lumen was completely unobstructed.

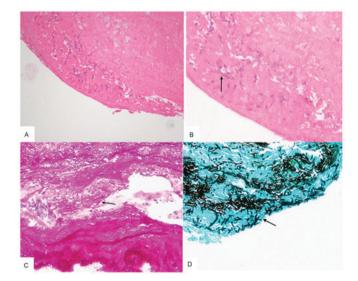


Figure 4. Histopathological examination of bronchoscopic biopsied specimen was completed using haematoxylin and eosin staining (A) at a magnification of x100 where hyphae like structure were observed, (B) and a magnification of x200 where sections of necrotic tissue and visible fungus hyphae with branch and sub-section were observed (black arrow). (C) Periodic acid-Schiff staining allowed the observation of red hyphae (black arrow) and (D) hexamine silver staining indicated the presence of black hyphae (black arrow).

was conducted and samples were stained using silver solution for 1.5 h at 65°C first, then exposed to 0.25% chloride for 2 min, 3% sodium thiosulfate for 1 min and 0.1% bright green staining for 1 min at room temperature. Periodic acid-Schiff and hexamine silver staining results were positive (Fig. 4). The patient was then treated with oral voriconazole (Pfizer, New York, United States) 200 mg once a day for 1 month. The lumen was reassessed with the bronchoscope and was observed to be completely unobstructed (Fig. 5). A repeat chest CT demonstrated that the left lung lesion was largely absorbed (Fig. 6). In addition, the patient's symptoms were noticeably relieved, no fever was observed, bloody sputum was no longer observed 3 days post interventional therapy. Complete blood count and routine blood chemistry markers were maintained at normal levels.

Literature comparison. By combining the present case with similar cases identified in the literature, a retrospective analysis of the clinical and radiological characteristics in addition to the bronchoscopic appearance was performed. The key word 'endobronchial aspergilloma' was searched in the PubMed NCBI database (ncbi.nlm.nih.gov/pubmed?holding=icnsibslib), by excluding cases without histopathological confirmation and

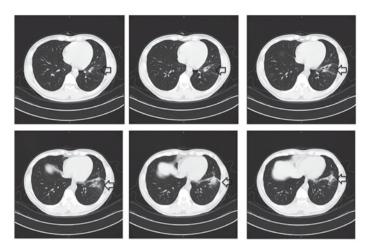


Figure 6. Following fiberoptic bronchoscopy, the patient was administered interventional voriconazole therapy for 1 month. A follow-up computed tomography scan demonstrated that the left lung lesions was absorbed.

simple Aspergilloma. In total, 17 patients meeting the EBA diagnosis criteria were included (including the case reported in the current study). The number of males (13/17; 76.5%) was notably higher, compared with women (4/17; 23.5%), and ages ranged from 33 to 82 years old (median age, 59). Of the 17 patients, 9 (52.9%) were smokers. Diagnoses of previous lung diseases included pulmonary tuberculosis (n=8; 47.6%), lung cancer (n=4; 23.5%), pulmonary resection (early stage lung cancer; n=1; 5.9%) and foreign body of the right main bronchus (n=1; 5.9%). Notably, 2 patients had previously been diagnosed with endobronchial lung cancer covered by aspergillosis (patients 13 and 14). A total of 8 patients had non-pulmonary comorbidities, including dilated cardiomyopathy (n=1), primary thrombocytosis (n=1), intracerebral hemorrhage (n=1), chronic heart failure (n=2), subarachnoid hemorrhage (n=1), hypertension (n=1), rheumatoid arthritis (n=1), hepatichemangima (n=1) and diabetes (n=4). None of the patients exhibited evidence of severe immune deficiency. Primary clinical symptoms were hemoptysis (9/17; 53%), cough (7/17; 41.2%) and dyspnea (3/17; 17.6%). Microbiology examination was performed in the bronchial lavage fluid of 12 patients, among whom only 2 exhibited aspergilloma, and 5 patients were treated with antifungal drugs (unknown) but the treatment response was not clear, with the exception for the present patient. Clinical characteristics of the 17 patients with EBA are summarized in Table I.

X-ray examination was performed for 13 patients, and all but 1 exhibited abnormal manifestations. The abnormal region was most often in the upper lobe (9/13; 69.2%), than in the left (8/13; 61.5%). Primary manifestations were pulmonary fibrotic changes along with lung volume loss, similar to old pulmonary tuberculosis. Mass or nodular shadows were also detected as presentations.

CT scanning indicated that Aspergillus lesions were more likely to be identified in the left lung (13/17; 76.5%) than in the right (4/17; 23.5%). Lesions were most common in the left upper lobe (6/17; 35.3%) and left lower lobe (6/17; 35.3%) on the CT scan. The types of Aspergillus identified by chest CT included: Space-occupying lesions (n=10; 58.8%), aspergilloma (n=3; 17.6%), pneumonic consolidation (n=2; 11.8%), and ground glass opacity (n=1; 5.9%). CT results

indicated a mass with a multi-lobulated contour in 3 patients (patients 1, 10 and 15), and an endobronchial lesion, such as a foreign body or broncholithiasis, was suspected in 4 patients (patients 2, 8, 16 and 17). Multiple calcified nodules and fibrotic changes as sequelae of previous pulmonary tuberculosis were noted in 6 patients (patients 4-9).

Bronchoscopy examination identified a mass in the bronchial lumen of all 17 patients and 15 patients presented with endobronchial obstruction by necrotic material. The abnormal regions under bronchoscopy were consistent with the CT locations. Table II summarizes the radiological characteristics and the bronchoscopic appearance in 17 patients with EBA.

Discussion

Aspergilloma often occurs in lung parenchyma or the lung cavity (12). EBA is a rare space-occupying disease of the endobronchial fungi, and it may not belong to the current classification of Aspergillus spp. (IPA, CNA, ABPA and SIA). It is a substantive or cavitary lesion caused by an overgrowth of aspergilloma with a non-invasive form in endobronchial areas through inhalation and fumigation, which is always limited to the main bronchus (13). EBA may present either as the sole manifestation of pulmonary aspergillosis or with other forms of pulmonary aspergillosis. CNA and ABPA may be associated with the EBA or the visible aspergilloma inside the lung cavity, demonstrated by bronchoscopy. Among 10 cases of EBA reported by Ma *et al* (6), 2 patients had lung parenchymal lesions with EBA in the lung cavity.

It is not easy to directly diagnose EBA by imaging or other noninvasive tests. In the present study, all patients were diagnosed incidentally when having a bronchoscopy and this was confirmed by histological examination and culture. Only 3 patients were suspected to have Aspergillus on the basis of CT results.

Hemoptysis (9/17; 53%) and cough (7/17; 41.2%) were the most common symptoms of EBA, and hemoptysis specifically, may be a result of the Aspergillus colonization, which makes the diagnosis via FOB important (6). The current study suggested the detection rate of the microbiology examination using bronchial lavage fluid was low (16.7%); therefore,

Table I. Clinical characteristics of 17 patients with endobronchial aspergilloma.

(Refs.)	(9)	(9)	(9)	(9)	(9)	(9)	(9)	(9)	(9)	(9)	(5)	(8)	(7)		(7)	The present study	(6)	(10)
Treatment	Itraconazole via IV and oral for 4 weeks	1	Oral itraconazole for 3 weeks	1	ı	ı	ı	ı	ı	1	Voriconazole for a number of months	ı	Voriconazole,	not clear	Itraconazole,	None	None	None
Microbiologic examination of BW	Aspergillus spp. No MTB isolated	Not completed	Not completed	No MTB isolated	No MTB isolated	No MTB isolated	No MTB isolated	No MTB isolated	No Aspergillus spp. isolated No MTB isolated	Not completed	Not completed	Not completed	No fungal organisms	or acid-fast bacilli were identified in sputum cultures.	Sputum culture grew not clear	No Aspergillus spp. isolated No MTB isolated	No Aspergillus spp. isolated No MTB isolated	No Aspergillus spp. isolated No MTB isolated
Smoking status	Ex	NS	CS	CS		CS	NS	Ex	Ex	CS	ı	NS	ı		CS A fumigatus	Ex	NS	NS
History of pulmonary TB	1	ı	2 months	25 years	30 years	6 years	30 years	7 years	10 years	ı	6 months	None	I		None	None	None	None
Underlying disease	DCMP	HT	NSCLC	1	I	I	ET, hepatichemangima	I	SDH, ICH	NSCLC	CHF, RA	None	NSCLC, pneumonia	and bronchitis	NSCLC, COPD, CHF, Diabetes, gastroesophageal reflux disease	Diabetes	Diabetes	Pneumothorax surgery
Symptom	Dyspnea, cough	Cough, sputum, dyspnea	Cough, sputum	Hemoptysis	Hemoptysis	Hemoptysis, Dyspnea	Cough, sputum	Hemoptysis	Cough, sputum	Hemoptysis, cough	No	Hemoptysis	Shortness of breath,	hemoptysis	Shortness of breath	Fever, cough, expectoration and blood-tinged sputum	Fever and chills	Blood-tinged sputum, chest pains
Age	75	53	70	70	51	46	57	36	92	50	49	59	99		82	48	75	33
Sex	M	Ţ	Σ	Σ	\boxtimes	\mathbb{Z}	M	\mathbb{Z}	\boxtimes	\mathbb{Z}	\boxtimes	\boxtimes	Ц		Г	\mathbb{M}	ഥ	M
No.	1	2	8	4	5	9	7	8	6	10	11	12	13		41	15	16	17

F, female; M, male; TB, tuberculosis; BW, bronchial washing; DCMP, dilated cardiomyopathy; Ex, ex-smoker; HT, hypertension; IV, intravenous; NS, never smoked; NSCLC, non-small cell lung cancer; CS, current smoker; MTB, Mycobacterium tuberculosis; ET, essential thrombocytosis; SDH, subarachnoid hemorrhage; ICH, intracranial hemorrhage; CHF, chronic heart failure; RA, rheumatoid arthritis.

Table II. Radiographic and bronchoscopy findings in 17 patients with endobronchial aspergilloma.

No.	Chest X-ray	Chest CT	Bronchoscopy
-	Round shaped mass lesion, LUL	Well-marginated and multi-lobulating mass in LUL, non-enhanced low density mass.	Protruding whitish mass in LUL apicoposterior segment
2	No definite lesion	Small sized high-density lesion suggesting foreign body or broncholith in RBI.	Foreign body adjacent granulation tissue covered with necrotic tissue in RBI.
3	RUL lobectomy state	RUL lobectomy state. No parenchymal lesion.	Whitish and necrotic tissue in anastomosis site of RUL.
4	Sequelae of TB, LUL	Sequelae of pulmonary TB in LUL with endobronchial lesion in LUL apical segment, focal GGO in LUL.	Small whitish mass in LUL upper division.
W	Sequelae of TB, both upper lobes	Soft tissue mass, low density and non-enhanced, with focal calcification causing obstruction of LUL apical segment.	Whitish mass-like lesion in LUL.
9	Sequelae of TB of both lungs	Sequelae of pulmonary TB in both upper lobes, cavitary lesion in RUL and RML, fungus ball in LUL. Ground glass opacity in LLL superior segment.	Large yellowish-necrotic mass obstructing LUL apicoposteriorsegment.
7	Sequelae of TB, LUL	Pneumonic consolidation in lingular division of LUL.	Mucus like yellowish mass in lingular division of LUL
∞	Sequelae of TB and Fungus ball formation, LUL	Severe emphysematous change in the two lungs. Fungus ball in LUL and aspirated blood in the two lower lung fields.	Whitish to yellow necrotic mass in LUL.
6	Sequelae of TB in RUL and pneumonic consolidation in LLL	Severe fibrotic change and volume loss in right lung and pneumonic consolidation in LLL.	Mass with yellowish and dark brownish exudates in LUL apicoposterior segment.
10	Mass like shadow in left hilum	Low-density and lobulating mass in LLL causing obstruction of LLL superior segment.	Protruding mass with yellow necrotic material in LLL superior segment.
11		Cavitary lesion with a suggestive image of a fungus ball, in the lingular division of the LUL.	A large yellowish hard mass with almost complete obstruction of the lower segment of the lingular division of the LUL,
12	Nodular opacity in left hilar field	Chest revealed a 2.4x1.8 cm spiculated mass and surround small nodular opacities in superior segment of left lower lobe.	Irregular mass-like, brownish material, which completely obstructed the subsegmental bron chus and a foreign body in superior segmental bronchus of left lower lobe.
13		Left pulmonary infrahilar soft tissue mass with an endobronchial component extending into the left lower lobe.	Friable, white mass almost completely obstructing patients left lower lobar bronchus.
14	ı	Right main bronchial mass was identified, causing collapse of the right upper lobe. Furthermore, a number of smaller nodules were observed in the lung parenchyma bilaterally.	Numerous fragments of carcinoid in part covered by fibrin, necrotic debris, and fungi, the latter consistent with aspergillus.

lower lobe; RML, right middle lobe.

glass opacity; LLL, left

, computer tomography; LUL, left upper lobe; RBI, right bronchus intermedius; RUL, right upper lobe; TB, tuberculosis; GGO, ground

CT.

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Bronchoscopy	lesion with post-obstructive Mass blocking the left basal segment was removed by grasping forceps. The mass was attached firmly to the bronchus and slight bleeding occurred at the removal site. The mass had an irregular yellow and black surface and broke apart easily	ely
Chest CT	A 1.0x0.5-cm calcified endobronchial lesion with post-obstructive pneumopathy in the left lower bronchus.	Further evaluation of the nodular density, a 1.5x2.5-cm relatively well-defined, oval shaped mass at the left lower lobe near the origin of the laterobasal segmental bronchus.
Chest X-ray	Diffuse consolidation in the left lower lobe	III-defined nodular density; at the left infrahilar area
No.	16	17

histopathological examination was required. None of the 17 cases were reported to have exhibited complications or adverse effect following biopsy.

Impaired immune function and certain comorbidities, such as active tuberculosis or diabetes, are important factors for fungal infection. Although there were no reports regarding the patients' immune status in the current study, it was revealed that of the 17 patients, 12 (70.6%) had previously been diagnosed with lung diseases, including pulmonary tuberculosis and lung cancer, and 8 patients (47.6%) had non-pulmonary comorbidities, such as dilated cardiomyopathy, essential thrombocytosis, cerebrovascular accident, hypertension and diabetes.

In patients with normal immune function, the formation of aspergilloma was considered to require a nest structural change that induces the airflow stasis, which helps Aspergillus in the airflow to colonize (6). In the present study, 8 patients with old pulmonary tuberculosis were included, and they exhibited destructive and fibrotic changes of the parenchyma, which potentially resulted in airway stenosis and obstruction. In addition, the majority of those who had no history of tuberculosis presented with a narrow or completely blocked lumen caused by lung cancer, lung resection, foreign bodies or granulation tissue, which was confirmed through imaging and bronchoscopy. Therefore, this change of the bronchial lumen may be considered to be an important factor to potentially induce airflow stasis and Aspergillosis colonization in bronchioles, in addition to the formation of aspergilloma. Lesions were also identified in the left lung more often than in the right, and it is suggested that this is because the left lung is anatomically closer to the heart and aorta. In addition, the left main bronchus is longer and narrower than the right side, thus it is easier to induce airflow stasis in the bronchioles of the left lung, causing EBA (13).

Notably, in the present study, 4 patients had lung cancer, of which 1 underwent surgery, 1 was affected by aspergillus (6), and the other 2 reported by Nilsson *et al* (7) had endobronchial tumor covered by aspergilloma. In Nilsson's report, the fungal hyphae were identified via histopathology and the patients were subsequently treated with anti-fungal drugs (unknown). However, in both the reported cases, the mass kept growing and finally the tumor was identified through further biopsy. There was no clear evidence to link Aspergillus and cancer development in the literature (7) and it is rarely reported that fungal infection may co-exist with solid types of cancer in immunocompetent patients (14,15). Malignant tumor may be a factor for local fungal infection including EBA.

The optimal treatment of EBA has not yet been established. The short-term follow-up in the current study confirmed that through interventional treatment using bronchoscopy and systemic antifungal therapy, the prognosis of EBA is excellent. Nevertheless, Jung *et al* (8) argued that for clinically asymptomatic patients, there is no requirement for treatment, and in cases of mild hemoptysis such as in the present patient's case, medical therapy with bed rest, humidified oxygen, cough suppressants and postural drainage is sufficient (8). To the best of our knowledge, there is no consistent evidence that aspergilloma responds to systemic administration of antifungal agents (12). In certain reports, surgical treatment was a good alternative (16-18), but the pulmonary function and tolerance of patients should be considered. It is suggested that

selecting appropriate treatments according to different conditions of patients is reasonable, and the combination of direct bronchoscopic intervention with antifungal therapy may be advantageous.

In conclusion, the dominant symptom for EBA was identified to be hemoptysis in the present analysis. Chest CT demonstrated that aspergillosis lesions were more frequently identified in the left lung compared with the right. It was also revealed that endobronchial aspergilloma often occurs in individuals with comorbidities that may potentially impair immune function and underlying lung diseases that cause lumen structural change or bronchial obstruction. Previous studies have provided sufficient evidence that endobronchial aspergilloma may be clearly diagnosed by bronchoscopy biopsy, but the potential for the co-exististing of tumors requires consideration (5-10). Finally, based on clinical experience, the combination of bronchosopic intervention and anti-fungal therapy may be an effective treatment for patients with EBA.

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References

- 1. Silva EF, Barbosa Mde P, Oliveira MA, Martins RR and Fontinele e Silva J: Chronic necrotizing pulmonary aspergillosis. J Bras Pneumol 35: 95-98, 2009 (In English, Portuguese).
- Hospenthal DR, Kwon-Chung KJ and Bennett JE: Concentrations of airborne aspergillus compared to the incidence of invasive aspergillosis: Lack of correlation. Med Mycol 36: 165-168, 1998.
- 3. Zmeili OS and Soubani AO: Pulmonary aspergillosis: A clinical update. QJM 100: 317-334, 2007.
- Chabi ML, Goracci A, Roche N, Paugam A, Lupo A and Revel MP: Pulmonary aspergillosis. Diagn Interv Imaging 96: 435-442, 2015.

- Araújo D, Figueiredo M and Monteiro P: Endobronchial aspergilloma: An unusual presentation of pulmonary aspergillosis. Rev Port Pneumol (2006) 22: 61-62, 2016.
- Ma JE, Yun EY, Kim YE, Lee GD, Cho YJ, Jeong YY, Jeon KN, Jang IS, Kim HC, Lee JD and Hwang YS: Endobronchial aspergilloma: Report of 10 cases and literature review. Yonsei Med J 52: 787-792, 2011.
- Nilsson JR, Restrepo CS and Jagirdar J: Two cases of endobronchial carcinoid masked by superimposed aspergillosis: A review of the literature of primary lung cancers associated with aspergillus. Ann Diagn Pathol 17: 131-136, 2013.
- Jung SW, Kim MW, Cho SK, Kim HU, Lee DC, Yoon BK, Jeong JP and Ko YC: A case of Endobronchial aspergilloma associated with foreign body in immunocompetent patient without underlying lung disease. Tuberc Respir Dis (Seoul) 74: 231-234, 2013.
- Yeo CD, Baeg MK and Kim JW: A case of endobronchial aspergilloma presenting as a broncholith. Am J Med Sci 343: 501-503, 2012.
- Kim JS, Rhee Y, Kang SM, Ko WK, Kim YS, Lee JG, Park JM, Kim SK, Kim SK, Lee WY and Chang J: A case of endobronchial aspergilloma. Yonsei Med J 41: 422-425, 2000.
- 11. British Thoracic Society Bronchoscopy Guidelines Committee, a Subcommittee of the Standards of Crae Committee of the British Thoracic Society: British thoracic society guidelines on diagnostic flexible bronchoscopy. Thorax 56 (Suppl. 1): i1-i21 2001
- nostic flexible bronchoscopy. Thorax 56 (Suppl I): i1-i21, 2001.

 12. Patterson KC and Strek ME: Diagnosis and treatment of pulmonary aspergillosis syndromes. Chest 146: 1358-1368, 2014.
- nary aspergillosis syndromes. Chest 146: 1358-1368, 2014.

 13. Soubani AO and Chandrasekar PH: The clinical spectrum of pulmonary aspergillosis. Chest 121: 1988-1999, 2002.
- 14. Tashiro T, Izumikawa K, Tashiro M, Takazono T, Morinaga Y, Yamamoto K, Imamura Y, Miyazaki T, Seki M, Kakeya H, *et al:* Diagnostic significance of aspergillus species isolated from respiratory samples in an adult pneumology ward. Med Mycol 49: 581-587, 2011.
- 15. Ohmagari N, Raad II, Hachem R and Kontoyiannis DP: Invasive aspergillosis in patients with solid tumors. Cancer 101: 2300-2302, 2004.
- Stevens DA, Kan VL, Judson MA, Morrison VA, Dummer S, Denning DW, Bennett JE, Walsh TJ, Patterson TF and Pankey GA: Practice guidelines for diseases caused by Aspergillus. Infectious diseases society of america. Clin Infect Dis 30: 696-709, 2000.
- 17. Sagan D and Goździuk K: Surgery for pulmonary aspergilloma in immunocompetent patients: No benefit from adjuvant antifungal pharmacotherapy. Ann Thorac Surg 89: 1603-1610, 2010.
- fungal pharmacotherapy. Ann Thorac Surg 89: 1603-1610, 2010.

 18. Lee JG, Lee CY, Park IK, Kim DJ, Chang J, Kim SK and Chung KY: Pulmonary aspergilloma: Analysis of prognosis in relation to symptoms and treatment. J Thorac Cardiovasc Surg 138: 820-825, 2009.