Pneumothorax after transbronchial needle biopsy

Tatjana Boskovic1, Milos Stojanovic1, Jelena Stantic2, Slobodanka Pena Karan1, Gordana Vujsinovic1, Dragan Dragisic1, Konstantinos Zarogoulidis3, Ioanna Kougioumtzi4, Georgios Dryllis5, Ioannis Kioumis3, Georgia Pitsiou3, Nikolaos Machairiotis6, Nikolaos Katsikogiannis4, Antonis Papaioannour4, Athanasios Madesis7, Konstantinos Diplaris7, Theodoros Karaikos7, Bojan Zarić8, Perin Branislav8, Paul Zarogoulidis3

1Institute for Pulmonary Diseases of Vojvodina, Center for Radiology, Faculty of Medicine, University of Novi Sad, Sremska Kamenica, Vojvodina, Serbia; 2Institute for Pulmonary Diseases of Vojvodina, Clinic for Thoracic Oncology, Faculty of Medicine, University of Novi Sad, Sremska Kamenica, Vojvodina, Serbia; 3Pulmonary Department, “G. Papanikolaou” General Hospital, Aristotle University of Thessaloniki, Thessaloniki, Greece; 4Surgery Department, University General Hospital of Alexandroupolis, Democritus University of Thrace, Alexandroupolis, Greece; 5Hematology Department, “Laiko” University General Hospital, Athens, Greece; 6Obstetric - Gynecology Department, “Thriassio” General Hospital of Athens, Athens, Greece; 7Thoracic Surgery Department, “G. Papanikolaou” General Hospital, Aristotle University of Thessaloniki, Thessaloniki, Greece; 8Institute for Pulmonary Diseases of Vojvodina, Clinic for Thoracic Oncology, Faculty of Medicine, University of Novi Sad, Novi Sad, Serbia

Correspondence to: Paul Zarogoulidis, MD, PhD. Pulmonary Department, “G. Papanikolaou” General Hospital, Aristotle University of Thessaloniki, Thessaloniki, Greece. Email: pzarog@hotmail.com.

Abstract: Currently there several diagnostic techniques that are used by radiologists and pulmonary physicians for lung cancer diagnostics. In several cases pneumothorax (PNTX) is induced and immediate action is needed. Both radiologists and pulmonary physicians can insert a chest tube for symptom relief. However, only pulmonary physicians and thoracic surgeons can provide a permanent solution for the patient. The final solution would be for a patient to undergo surgery for a final solution. In our current work we will provide all those diagnostic cases where PNTX is induced and treatment from the point of view of expert radiologists and pulmonary physicians.

Keywords: Pneumothorax (PNTX); bronchoscopy; transbronchial needle aspiration; endobronchial ultrasound (EBUS)

doi: 10.3978/j.issn.2072-1439.2014.08.37

View this article at: http://dx.doi.org/10.3978/j.issn.2072-1439.2014.08.37

Introduction

Bronchoscopy was initially developed in 1895 for the purpose of removing foreign bodies from the main stem bronchi, when Gustav Killian removed a piece of bone from the right main-stem bronchus of a 36-year-old man. In the year 1968, Shigeto Ikeda introduced flexible fiberoptic bronchoscopy for clinical use. It became more used that rigid bronchoscopy and it was confirmed in 1980’s, that flexible bronchoscopy with topical anesthesia was safer than rigid bronchoscopy (1,2). Although new studies suggest that the annual referral rate for rigid bronchoscopy has been rising since 2007, which demonstrates the varying diagnostic and therapeutic modalities available and highlight the favorable morbidity rates and 100% diagnostic rates for this safe procedure, this book chapter will focus on flexible bronchoscopy transbronchial biopsy, its associated complications—primarily pneumothorax (PNTX), and management strategies (3,4).

Nearly 500,000 bronchoscopies are done each year in the United States. The number and complexity of procedures that can be performed in the bronchoscopy unit is increasing. These procedures carry inherent risks, and patient safety is of paramount concern. Flexible bronchoscopy is an invasive procedure that is utilized to visualize the nasal passages, pharynx, larynx, vocal cords,
and tracheal bronchial tree. It is utilized for both the diagnosis and treatment of lung disorders. The procedure may be performed in an endoscopy suite, the operating room, the emergency department, a radiology suite, or at the bedside in the ICU. Flexible bronchoscopy has revolutionized the practice of pulmonary medicine, enhanced our understanding of pulmonary disease, and has evolved into the most commonly used invasive diagnostic as well as therapeutic procedure. Bronchoscopy is widely carried out today by pulmonologists, thoracic surgeons, critical care specialists, otolaryngologists, anesthesiologists and pediatric pulmonologists (1,5).

Many interventional bronchology procedures for the diagnosis and treatment of lung illnesses can be performed on an outpatient basis or during a short hospital stay. The problem most responsible for complicating outpatient management, after needle biopsy was performed, is not the presence of the PNTX per se, but an increase in the size of the PNTX that requires chest tube placement and patient hospitalization. The risks and benefits of the procedure and knowledge of the wishes of the patient will enable the management decision to be tailored to the needs of the individual (6,7).

**Bronchoscopic lung biopsy**

Lung biopsy is a relatively frequently performed procedure with considerable benefit for patient management but it may, on rare occasions, give complications or even result in the death of the patient. It is a multidisciplinary procedure involving respiratory physicians, bronchologists, radiologists and surgeons with an interest in chest diseases. It is integral in the diagnosis and treatment of many thoracic diseases, and is an important alternative to more invasive surgical procedures.

Classification of lung biopsies may be done according to the method of access (percutaneously, bronchoscopically, open operation) or by the reason for biopsy (sampling of diffuse lung disease or obtaining tissue from a mass when malignancy is suspected) (8,9).

Biopsy via a bronchoscope is useful for proximal endobronchial lesions but is unable to access more peripheral lesions. Transbronchial biopsy is generally done with two main types of forceps; cup forceps (without teeth) and the alligator forceps (saw-toothed). It is noted that the alligator-type forceps are being used less than the cup, as the teeth tend to tear tissue and cause hemoptysis (1).

Biopsy of diffuse infiltrates is more likely to provide diagnosis than small peripheral nodules. Overall yield from bronchoscopic lung biopsy is reported to be 72%. The American Thoracic Society/European Respiratory Society Consensus recommends bronchoscopic lung biopsy not for the diagnosis of idiopathic interstitial pneumonias but for the exclusion of sarcoidosis, infections and lymphangitis carcinomatosis (10,11). Transbronchial biopsy of diffuse lung disease may be assisted by some imaging guidance. High resolution computed tomography (CT) might be helpful in deciding which areas should be targeted to improve diagnostic yield and areas that should be avoided, such as bullae or vascular abnormalities. Because it does not cross the pleura, PNTX is much less common than in percutaneous biopsy (12).

**Technique**

Ideally, the technique must not only be able to diagnose malignancy but also to make a definite diagnosis if the lesion is benign. transbronchial needle biopsy (TBNB) usually begins with review of the chest radiograph and, in most instances, is greatly facilitated by a CT scan. Knowledge of the anatomy is critical for selecting the proper anatomic location for the needle aspiration or biopsy. This is true for selecting the proper location of a peripheral lesion that is to be sampled. For peripheral lesions, fluoroscopy is used to localize the lesion (12).

At minimum, the equipment needed is a bronchoscope, light source, cytology brushes, biopsy forceps, needle aspiration catheters, suction apparatus, supplemental oxygen, fluoroscopy (C-arm), pulse oximetry, sphygmanometer, and equipment for resuscitation including an endotracheal tube. A video monitor is a useful accessory, but not required. Fluoroscopy may be needed to facilitate certain transbronchial biopsy procedures (Figure 1).

Both therapeutic and diagnostic procedures can be performed during flexible bronchoscopy. Depending on the indication, the following diagnostic procedures can be performed: BAL, endobronchial or transbronchial biopsies, cytologic wash or brush, and TBNA, endobronchial ultrasound (EBUS), and autofluorescence bronchoscopy. Therapeutic procedures and selected stent placement can all be accomplished through flexible bronchoscopy (5).

**Indications**

The first indication for fiberoptic bronchoscopy is identification of an indeterminate lung lesion identified on the chest X-ray. When transbronchial biopsy is added to the care
Flexible bronchoscopy, the most common indications are those conditions in which histopathology is important in therapeutic decision making, such as lung transplantation and rare or unusual parenchymal lung diseases. Diagnostic and staging information in the presence of malignancy in mediastinal lymph nodes, are also indications for TBNB.

Most contraindications to flexible bronchoscopy are relative rather than absolute they include: uncooperative patient, recent myocardial infarction, tracheal obstruction or stenosis, moderate to severe hypoxemia or any degree of hypercarbia, uremia and pulmonary hypertension (possible serious hemorrhage after biopsy), unstable asthma, bleeding disorders, immunosuppression, respiratory failure or insufficiency, cardiac arrhythmias. Special attention must be paid to respiratory and bleeding status. In unstable patients or prolonged procedures, rigid bronchoscopy may be preferred (1,5).

**Safety of TBNB in patients with COPD and pulmonary hypertension**

Fiberoptic bronchoscopy is being used increasingly as a diagnostic and research tool in patients with COPD to assess airway pathology and the effects of treatment. Previous reports have shown bronchoscopy and bronchial biopsy procedures to be safe in asthmatic patients, but there is little safety data specific to COPD. Guidelines suggest that bronchoscopy may be safer in patients with COPD than with asthma because of lower levels of bronchial hyperresponsiveness. Fiberoptic bronchoscopy is a relatively invasive procedure for investigating patients with COPD, but it allows for the detection of coexisting endobronchial pathology and allows multiple samples of both tissue and fluid phase material to be obtained for research or diagnostic purposes (13,14).

Patients with pulmonary hypertension (PH) are considered to be at risk for complications associated with flexible bronchoscopy. Although previous reports suggest that transbronchial biopsies increase the risk for hemorrhage in this population, review of patients with diagnosis of PH who underwent flexible bronchoscopy at the Cleveland Clinic between 2002 and 2005. Total of 90 patients, showed that flexible bronchoscopy can be performed safely in patients with mild to moderate PH. Transbronchial biopsies were not associated with worsening hypoxemia or an increased risk of hemorrhage. Prospective studies with hemodynamic measurements are necessary to confirm these findings (15).

**Safety of new bronchoscopic modalities**

In 2007, a randomized trial study was performed on 120...
patients, to examine influence of new bronchoscopic modalities on incidence of pneumothoraces, using EBUS and electromagnetic navigation bronchoscopy (ENB). These two modalities have increased the diagnostic yield of bronchoscopic diagnosis. All procedures were performed via flexible bronchoscopy and transbronchial forceps biopsies were obtained without fluoroscopic guidance. The PNTX rates ranged from 5% to 8%, with no significant differences between the groups. Four cases were treated with chest drains. No cases of bleeding that required therapeutic interventions, were recorded. It was concluded that combined EBUS and ENB improved the diagnostic yield of flexible bronchoscopy without compromising safety (16).

**Complications**

Diagnostic flexible bronchoscopy with TBNB is usually very safe procedure as long as some basic precautions are taken. Two most common complications are bleeding and PNTX. Major complications such as bleeding, respiratory depression, cardiorespiratory arrest, arrhythmia, pneumomediastinum and PNTX occur in <1% of cases. Hemorrhage is more likely in uraemic or immunosuppressed patients. Significant bleeding rarely occurs even after a major vessel puncture. Fever and bacteremia have been reported following the procedure, although this may be related to the bronchoscopic procedure itself rather than this specific technique. Mortality is rare, with a reported death rate of 0% to 0.04% in >68,000 procedures. Therefore, caution should be taken and healthcare providers must be prepared for these emergencies (1,5,7,17). According to recent nationwide survey from Japan, the complication rate after forceps biopsy for a peripheral pulmonary lesion was 1.79%. In comparison, in USA a review of 173 procedures from university hospital has reported a complication rate of 6.8% after TBB (18,19).

**Bleeding**

The reported incidence of bleeding after TBNB has varied from 0% to 26% in different series (20). The risk of bleeding is significantly higher in patients with underlying renal insufficiency (12,13). A major bleeding complication after transbronchial biopsy is unnerving but the bleeding can be controlled in the majority of cases in the bronchoscopy room without adverse patient outcome. The most effective way to control the bleeding is to maintain the bronchoscope in wedged position into bleeding segment till a blood clot is formed (21,22).

**Pneumothorax**

Definition of PNTX is presence of gas in the pleural space. By cause, it can be spontaneous, traumatic and iatrogenic. Occult pneumothoraces are a relatively recent radiological phenomenon. They are defined as pneumothoraces detected with thoracic or abdominal CT that were not diagnosed on preceding supine anteroposterior chest radiography. However, most pneumothoraces are iatrogenic and caused by a physician during surgery, central line placement, lung biopsy or bronchoscopy. PNTX is a rare complication of bronchoscopy with transbronchial biopsy (23-25).

Acutely symptomatic pneumothoraces may develop at the time of the transbronchial biopsy procedure and require immediate drainage. Smaller or better tolerated pneumothoraces will be detected on post bronchoscopy chest radiographs. Acute presentation is usually with acute ipsilateral chest pain and dyspnoea. Clinical findings may be minimal or may include diminished breath sounds and mediastinal shift. In a tension PNTX the patient may become tachycardic and hypotensive and develop cyanosis. Monitoring of oxygen saturation is advised, together with the administration of oxygen as necessary. In an acutely unwell patient a chest radiograph or CT scan can be used to identify whether symptoms relate to pulmonary haemorrhage or PNTX (12).

**Timing of chest radiography**

Several studies have shown that most significant pneumothoraces will be detected on a chest radiograph performed 1 hour after the procedure, although they may not be visible on radiographs taken immediately after the procedure (26-28). Occasional delayed pneumothoraces have been reported more than 24 hours after biopsy, despite the absence of a PNTX on chest radiographs taken 4 hours after biopsy (29,30).

The need for routine chest X-ray after fiberoptic bronchoscopy was advised but never fully implemented, so a very recent study on 454 patients was done in 2013 to assess the incidence of post bronchoscopy PNTX and to determine the need for routine post- fiberoptic bronchoscopy X-rays. Of 454 total fiberoptic bronchoscopies, only 1 case (0.22%) resulted in iatrogenic PNTX which was diagnosed clinically, confirmed with chest X-ray and required thoracotomy. From the data obtained it was concluded that routine chest X-ray after fiberoptic bronchoscopy may not be cost-effective or even medically necessary in patients without clinical evidence...
of PNTX (12) (Figure 2).

PNTX is reported in literature in 1-6% of patients after TBNB. Mild to moderate PNTX can cause disproportionate symptoms due to underlying pulmonary limitation in some patients undergoing flexible bronchoscopy. Appropriate fluoroscopic guidance during TBNB reduces the risk of PNTX. Chest roentgenogram is recommended to be performed after ½ to 1 h after completion of biopsies, when the PNTX is clinically suspected despite normal findings on immediate post-bronchoscopy fluoroscopy. Fluoroscopic examination detects PNTX immediately after the biopsies, but in some cases slowly developing delayed PNTX can manifest several hours after completion of procedure. If no symptoms develop for 4 hours after completion of procedure, a clinically significant PNTX is unlikely (22,31,32).

Failure to control coughing during transbronchial biopsy greatly increases the risk of PNTX. Patients receiving positive pressure ventilation are also more likely to develop PNTX after TBNB. Risk may also be higher in presence of bullous emphysema and in patients with pneumocystis pneumonia. There are potential advantages of fluoroscopy when performing transbronchial biopsies in subjects with localized peripheral lesions. Severity of symptoms and the extent on chest roentgenogram dictate the management of PNTX after TBNB (17,33).

A large study was published in 2011 by Garcha, et al. of incidence and management of PNTX post-flexible bronchoscopy (FB). Out of 2,365 patients who underwent FB, 14 developed PNTX. Majority of the patients (93%) who developed PNTX had undergone transbronchial biopsies. Average number of transbronchial biopsy procedures was 5. Incidence of PNTX was 0.006%, with a mortality rate of 0.0004%. This complication rate reported was much lower than in previously published studies. Lung nodules and lung infiltrates were the most common indication for FB. Majority of patients were asymptomatic and PNTX was detected on routine fluoroscopy post procedure. 21% developed chest pain post procedure leading to the diagnosis. All of the patients required a tube thoracostomy for management of PNTX and were admitted to hospital for a short stay and there was one death directly related to development of tension PNTX. Although TBNB is assumed to be primary risk for PNTX, the majority of patients had other associated peripheral lung instrumentation. In new advanced techniques that were employed (ENB and EBUS), only 2 had pneumothoraces, representing 7% of the cases after ENB and EBUS (34).

In addition, two large retrospective studies of over 4,000 cases, using bronchoscopy with transbronchial biopsies, the latter in lung transplant recipients, showed no deaths and overall major and minor complication rates of 0.5% and 0.8%, respectively (19,35). Somewhat higher incidence of PNTX was reported to be 14% in a group of patients having transbronchial biopsies, while being mechanically ventilated (36).

A retrospective study published in 2004. In Chest journal, reviewed all patients with iatrogenic PNTX during 4-year period. Out of 86 patients, the most common cause reported to be central venous catheter placement, then thoracentesis, transthoracic needle aspiration biopsy (ten patients) and only one patient with PNTX after TBNB (37).

Management options

Where a PNTX is detected following a transbronchial biopsy procedure, the management options include observation, aspiration, or drain insertion. This decision will be affected by factors such as the size of PNTX, co-existent lung pathology such as emphysema affecting respiratory reserve, and severity of symptoms. BTS guidelines on the management of PNTX suggest initial treatment by aspiration, with subsequent drainage if a leak and significant PNTX persist (38). Supplemental oxygen and observation in the hospital is sufficient in most cases. Moderately symptomatic patients with significant PNTX may be managed with Heimlich’s valve placed in the bronchoscopy suite. These patients can be discharged with Heimlich’s valve after 4-6 h of observation, if repeat chest roentgenogram shows no further increase in PNTX. Patients who develop severe symptoms or tension PNTX,
and those who fail to show resolution of PNTX with Heimlich’s valve require chest tube placement. Chest tube should also be placed without further delay when PNTX develops on mechanical ventilation (22). In the UK most clinicians attach drains to an underwater seal, but the Heimlich one way flutter valve is an alternative. This valve allows prolonged drainage for a PNTX and outpatient management. If the PNTX continues to enlarge or the patient develops surgical emphysema, the flutter valve can be replaced by a system attached to an underwater seal (39). A major PNTX requiring drainage is reported to be in 3.5% of patients undergoing bronchoscopy (40-50) from which transbronchial biopsy specimens were taken (51-62).

Conclusions
Transbronchial biopsy is an essential skill for every bronchoscopist. TBNB is usually performed in outpatient setting under conscious sedation. If successfully performed, it may spare patients additional, more invasive procedures or surgery. Hemoptysis and PNTX are the two leading complications of TBNB, occurring in less than 2% of cases. PNTX is one of the major complications of FB, which can be life threatening if not managed in timely fashion. A major PNTX requiring drainage reported to be in 3.5% of patients undergoing bronchoscopy from which transbronchial biopsy specimens were taken. All bronchoscopists must develop proficiency in performing TBNB and managing complications that can arise after the procedure.

Acknowledgements

Disclosure: The authors declare no conflict of interest.

References
30. Sun SW, Zabaneh RN, Carrey Z. Incidence of Pneumothorax after Fiberoptic Bronchoscopy (FOB) in Community-Based Hospital; Are Routine Post-


