

Online Submissions: http://www.wjgnet.com/esps/ bpgoffice@wjgnet.com doi:10.5492/wjccm.v3.i1.8 World J Crit Care Med 2014 February 4; 3(1): 8-14 ISSN 2220-3141 (online) © 2014 Baishideng Publishing Group Co., Limited. All rights reserved.

EDITORIAL

# Iatrogenic pneumothorax related to mechanical ventilation

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 Received: June 25, 2013
 Revised: October 30, 2013

 Accepted: November 18, 2013
 Published online: February 4, 2014

## Abstract

Pneumothorax is a potentially lethal complication associated with mechanical ventilation. Most of the patients with pneumothorax from mechanical ventilation have underlying lung diseases; pneumothorax is rare in intubated patients with normal lungs. Tension pneumothorax is more common in ventilated patients with prompt recognition and treatment of pneumothorax being important to minimize morbidity and mortality. Underlying lung diseases are associated with ventilatorrelated pneumothorax with pneumothoraces occurring most commonly during the early phase of mechanical ventilation. The diagnosis of pneumothorax in critical illness is established from the patients' history, physical examination and radiological investigation, although the appearances of a pneumothorax on a supine radiograph may be different from the classic appearance on an erect radiograph. For this reason, ultrasonography is beneficial for excluding the diagnosis of pneumothorax. Respiration-dependent movement of the visceral pleura and lung surface with respect to the parietal pleura and chest wall can be easily visualized with transthoracic sonography given that the presence of air in the pleural space prevents sonographic visualization of visceral pleura movements. Mechanically ventilated patients with a pneumothorax require tube thoracostomy placement because of the high risk of tension pneumothorax. Small-bore catheters are now preferred in the majority of ventilated patients. Furthermore, if there are clinical signs of a tension pneumothorax, emergency needle decompression followed by tube thoracostomy is widely advocated. Patients with pneumothorax related to mechanical ventilation who have tension pneumothorax, a higher acute physiology and chronic health evaluation II score or PaO<sub>2</sub>/FiO<sub>2</sub> < 200 mmHg were found to have higher mortality.

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**Key words:** Barotrauma; Complication; Critical care; Mechanical ventilation; Pneumothorax

**Core tip:** Patients with pneumothorax related to mechanical ventilation (PRMV) have a high mortality rate. PRMV often occurs in the early stage of mechanical ventilation and it may recur on the other side of lung in a short period of time. Low compliance is associated with a high incidence of PRMV, with PRMV being more related to the underlying process than the ventilatory setting. PRMV patients with tension pneumothorax, higher acute physiology and chronic health evaluation score or  $PaO_2/FiO_2 < 200$  mmHg have a higher mortality.

Hsu CW, Sun SF. Iatrogenic pneumothorax related to mechanical ventilation. *World J Crit Care Med* 2014; 3(1): 8-14 Available from: URL: http://www.wjgnet.com/2220-3141/full/v3/i1/8.htm DOI: http://dx.doi.org/10.5492/wjccm.v3.i1.8

## INTRODUCTION

Pneumothorax, defined as the presence of air in the pleural space, is a serious complication of mechanical ventilation and is associated with increased morbid-



ity and mortality<sup>[1,2]</sup>. Pneumothorax can be categorized as primary, secondary, iatrogenic or traumatic according to etiology. Mechanical ventilation was found to be the common cause of iatrogenic pneumothorax in an intensive care unit (ICU)<sup>[3]</sup>. Pneumothorax is rare in intubated patients with normal lungs and most patients with pneumothorax related to mechanical ventilation (PRMV) have underlying lung diseases that range from primary obstructive lung disease to secondary pneumonia and acute respiratory distress syndrome (ARDS)<sup>[4-6]</sup>. In critical illness, pneumothoraces may be difficult to diagnose when they have different clinical presentations and their locations are atypical and complicated by other disease processes in unconscious patients<sup>[7,8]</sup>. Prompt recognition and treatment of pneumothorax is important to minimize morbidity and mortality<sup>[9]</sup> because if pneumothorax is not diagnosed quickly, once tension pneumothorax ensues, it usually has a malignant course leading to death if untreated<sup>[10]</sup>. Sudden fall in oxygen saturation followed by hypotension are often observed clinically in these patients<sup>[11,12]</sup>. Tension pneumothorax is more common in ventilated patients, occurring in 30%-97% percent of all pneumothoraces<sup>[13-17]</sup>.

Pneumothorax was found to be an independent predictor of mortality during mechanical ventilation<sup>[18]</sup> and was associated with a significant increase in the ICU length of stay, hospital stay and mortality in all mechanically ventilated patients<sup>[3]</sup>. The mortality rates are high, ranging from 46% to 77% if barotrauma is a complication of mechanical ventilation<sup>[2,3,14,15,19-24]</sup>.

## EPIDEMIOLOGY

Patients who received mechanical ventilation have an approximate incidence of barotrauma of 4%-15%<sup>[4,13,19,25]</sup>. It has been reported that a 14%-87% incidence of pneumothorax occurs depending on severity and duration of ARDS and mode of ventilator for management<sup>[1,21,26]</sup>. A prior study found that the incidence rate of pneumothorax decreased after the implementation of protective lung strategies in pediatric patients with severe ARDS<sup>[27]</sup>. The incidence of barotrauma has been reported to be as low as 0.5% in postoperative patients<sup>[23]</sup>.

## DISEASE ASSOCIATED WITH PNEUMOTHORAX

Development of pneumothorax is most closely correlated with underlying lung disease<sup>[1,21]</sup>. Pneumonia is an important predisposing factor in the development of pulmonary barotrauma in mechanically ventilated patients<sup>[28]</sup>. Necrotizing bacterial pneumonia can cause air leaks into the pleura that result in development of pneumothorax<sup>[29]</sup>.

Chronic obstructive pulmonary disease and asthma are also common underlying diseases associated with pneumothorax. High airway pressures are required to overcome severe bronchial obstruction. However, because of a variability of obstruction in the different airways, there is a mal-distribution of mechanical tidal volume, which promotes gas trapping and non-uniform alveolar distension. That is why these patients are at risk of pneumothorax<sup>[30]</sup>.

PRMV is most closely correlated with ARDS<sup>[31]</sup>, a heterogeneous disease in which the lung is physiologically small and with low compliance<sup>[32]</sup>. The dependent lung regions tend to be collapsed and the lung regions subject to high pressure overinflation and alveolar rupture are nondependent regions when positive end-expiratory pressure (PEEP) inflates and recruits some of collapsed regions during mechanical ventilation<sup>[33,34]</sup>. Subpleural and intrapulmonary air cysts occur in ARDS patients and the rupture of these air cysts may be a cause of pneumothorax<sup>[14]</sup>. Whether the pneumothorax in ARDS arises from overinflation of normal lung regions or from cyst rupture has not yet been conclusively established<sup>[21]</sup>.

Other diseases such as lung cancer, tuberculosis, bronchiectasis, cystic fibrosis, idiopathic pulmonary fibrosis, sarcoidosis, histiocytosis X, Marfan's syndrome, Ehlers-Danlos syndrome, lymphangioleiomyomatosis, rheumatoid arthritis and other connective tissue disease may predispose to the development of a pneumothorax<sup>[35]</sup>.

## PATHOPHYSIOLOGY AND RISK FACTORS FOR PRMV

An early investigation reported that peak airway pressure over 50 cm H<sub>2</sub>O is associated with increased risk of alveolar rupture during mechanical ventilation<sup>[4]</sup>. There have also been correlations made between high peak airway pressure and the development of pneumothorax<sup>[15,36]</sup>. High PEEP had been reported to be associated with pneumothorax<sup>[1]</sup> but several studies have found no such relationship<sup>[15,17,23,28,37]</sup>. Increased pressure is not enough by itself to produce alveolar rupture, with some studies demonstrating that pneumothorax is related to high tidal volume<sup>[37]</sup>. In animal studies, there is evidence to conclude that lung overdistension rather than high airway pressure is the primary cause of alveolar and interstitial injury<sup>[38,39]</sup>.

A clinical study showed that when plateau pressure was maintained lower than 35 cm H<sub>2</sub>O, pneumothorax was unavoidable<sup>[40]</sup>. Although one study showed that the incidence of pneumothorax decreased after implementation of protective lung strategies<sup>[27]</sup>, several recent studies comparing low tidal volume with conventional ventilation in ARDS failed to demonstrate any reduction in barotrauma when low tidal volumes were used<sup>[41-43]</sup>. Previous literature showed the incidence of barotrauma did not relate to the ventilatory settings<sup>[22,31,41,44]</sup> but a low compliance was associated with a high incidence of barotrauma, which suggested that barotrauma has been more related to the underlying process than the ventilatory setting<sup>[40]</sup>.

## CLINICAL MANIFESTATIONS OF PRMV

Pneumothorax is secondary to ruptured alveoli and dis-



#### Hsu CW et al. Ventilator-related pneumothorax

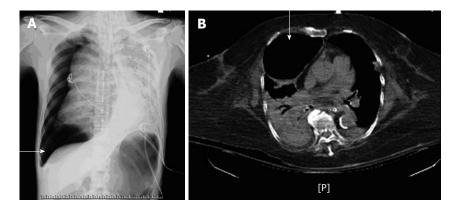


Figure 1 A chest radiograph demonstrates (A) and computed tomography (B) right pneumothorax (arrows). Air may lie in the costophrenic angle extending more inferiorly than usual (A, arrow).

section of air along the vascular sheaths passing to the mediastinum, subcutaneous tissue and retroperitoneum. Therefore, it may manifest as pulmonary interstitial emphysema, pneumomediastinum, pneumoperitoneum or subcutaneous emphysema<sup>[32]</sup>. Patients may present with tachycardia, chest pain, tachypnea, agitation, hypotension, cyanosis or consciousness change. Tachycardia is the most common finding.

It has been observed that the median time from mechanical ventilation to the development of pneumothorax was 4 d<sup>[3]</sup>. Anzueto *et al*<sup>[45]</sup> reported that 80% of their patients developed pneumothorax within the first 3 d of mechanical ventilation. Gattinoni *et al*<sup>[21]</sup> found that late ARDS (> 2 wk of mechanical ventilation) patients had an increased incidence of pneumothoraces when compared to early ARDS (less than 1 wk of mechanical ventilation) patients<sup>[21]</sup>. However, Gammon *et al*<sup>[31]</sup> reported that pneumothoraces occurred most commonly during the early phase of ARDS, with a declining risk over time.

Approximately 7.4%-10% of patients with first episode pneumothorax will develop second episode pneumothorax on the other lung during mechanical ventilation<sup>[3,31]</sup>. Clinically, we should take notice of these patients because the pneumothorax might attack the lung on the other side.

Tension pneumothorax is common in mechanically ventilated patients<sup>[13-17]</sup>. Mechanical ventilation will increase gas flow through pleura defects, allowing more air to pass per unit time, and resulting in a more rapid intrapleural pressure rise with earlier mechanical compressive effects and rapid progress to cardiorespiratory collapse and death<sup>[46]</sup>.

## DIAGONOSIS

The diagnosis of pneumothorax in critical illness is established from patients' history, physical examination and radiological investigation. The factors related to the underlying lung disease are important in the history. Examination of the respiratory and cardiovascular systems may help establish the diagnosis of pneumothorax. However, examination findings may vary according to the size of pneumothorax and presence of limited cardiorespiratory reserve<sup>[47]</sup>, with patients with a small pneumothorax (one involving < 15% of the hemithorax) possibly having a normal physical examination. Careful inspection and repeated auscultation of the chest is therefore crucial. Contralateral tracheal deviation, hyperresonant percussion over the chest and decreased breathing sounds might be noted. Reduction in tidal volume during pressure controlled ventilation and increased airway pressure with volume controlled ventilation might be found from ventilators. A pulsus paradoxus on the arterial trace and increased central venous pressure from central venous catheterization may be observed<sup>[8,9,46,47]</sup>.

It should be noted that many of the above findings are nonspecific and have not been a reliable indicator of pneumothorax given that dyspnea severity can be out of proportion to the size of the pneumothorax. The radiographic data thus remains the gold standard for the diagnosis of pneumothorax<sup>[8]</sup>. The chest radiograph may show radiolucent hemithorax with apparent lung edge and absent lung marking. Mediastinal emphysema was the initial manifestation of extra-alveolar air<sup>[1]</sup>. Although an erect posteroanterior chest radiograph may help to estimate the pneumothorax size<sup>[48]</sup>, it is not practical in critical illness. However, the supine anteroposterior chest radiographs are available in the ICU, although the appearances of a pneumothorax on a supine radiograph may be different from the classic appearances on an erect radiograph. Air may lie in the costophrenic angle extending more inferiorly than usual (Figure 1A), which has been called deep sulcus sign<sup>[49]</sup>. Concurrent lung disease may lead to different distributions of free air in the pleural space than in relatively normal lungs<sup>[50]</sup> or it may lead to loculated gas collections if there is associated pleural disease<sup>[51]</sup>. Some radiographic findings that may be confused with pneumothorax are skin folds, visceral gas within the gas and emphysema bullae. A chest computed tomography (CT) examination, in contrast, can differentiate these diagnoses and is the gold standard test for both the diagnosis and sizing of pneumothorax (Figure 1B)<sup>[52]</sup>. CT is an excellent tool to differentiate bullous lung disease and prevent unnecessary drainage attempts that may result in the creation of a parenchymal-pleural fistula<sup>[53]</sup>. Chest radiographs may not distinguish between emphysematous bullae and pneumothorax, thus possibly resulting in potentially catastrophic insertion of intercostal chest drain into emphysematous bullae in mechanically ventilated patients. Unfortunately, it is not always practical or safe



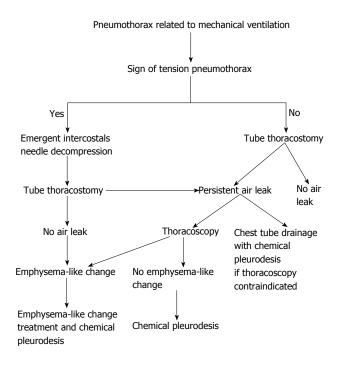


Figure 2 An algorithmic approach to treatment of pneumothorax related to mechanical ventilation.

to transport critically ill patients for a CT scan to exclude pneumothorax, particularly when the patient is hemodynamically unstable.

Transthoracic sonography is a diagnostic tool for pneumothorax<sup>[54-56]</sup>. There are several advantages of ultrasonography over standard chest radiography and CT scanning, including the lower radiation, portability, realtime imaging and the ability to easily perform dynamic and repeat evaluation. The main use of ultrasonography for assessment of pneumothorax lies in its capacity to rule out a pneumothorax. Respiration-dependent movement of the visceral pleura and lung surface with respect to the parietal pleura and chest wall can be easily visualized with real-time transthoracic sonography. This characteristic is known as lung sliding or gliding sign<sup>[56-58]</sup>. At the boundary between the pleura and the ventilated lung, intensive band-like reverberation echoes (comettail artifacts) are evoked during breathing movement<sup>[54,57]</sup>. M-mode images of patients without pneumothorax show breath-dependent movements as a single, thin comet-tail artifact. The presence of air in the pleural space prevents sonographic visualization of visceral pleura movements and the gliding sign and comet-tail artifacts disappear<sup>[57,58]</sup>. M-mode images of patients with pneumothorax show the immobility of these artifacts known as frozen echoes under breathing movement.

One study showed transthoracic ultrasound was more sensitive than chest radiography in the detection of pneumothorax<sup>[59]</sup>. Another study comparing ultrasonography to CT scan and chest radiographs for diagnosis of occult pneumothorax revealed that the use of ultrasonography detected 92% of occult pneumothoraces diagnosed with CT scan<sup>[60]</sup>. In trained hands, an ultrasound examination may obviate the need for empiric tube thoracostomy for suspected tension pneumothorax.

## TREATMENT AND PREVENTION

Most mechanically ventilated patients with a pneumothorax require tube thoracostomy placement because of the high risk of tension pneumothorax<sup>[61]</sup>. The strategy of managing pneumothorax is shown in Figure 2. It is not advisable to wait for a radiograph if there are clinical signs of a tension pneumothorax. Emergency needle decompression followed by tube thoracostomy is widely advocated<sup>[46]</sup>. In critically ill patients with minimal pulmonary reserve, even a small pneumothorax can have adverse cardiopulmonary effects<sup>[62]</sup>. Positive pressure ventilation can exacerbate air leaks and prevent pleural healing, potentially causing a rapid increase in the size and severity of existing pneumothorax.

The traditional treatment for pneumothorax in mechanically ventilated patients has been chest tube thoracostomy<sup>[61]</sup>, an image-guided small catheter whose size ranges from 7 to 10 Fr, to become an effective therapeutic option for pneumothorax<sup>[63]</sup>. A retrospective review of 62 ventilated patients who underwent small-bore chest tube drainage as the primary management of pneumothorax found a 68.6% success rate, defined as no residual air seen in the follow-up chest radiograph, and with no major complications<sup>[64]</sup>. These results compare favorably with previous data showing a success rate of 55% with the same definition for success as in the previous study with a large bore tube<sup>[20]</sup>.

In patients with a persistent air leak or failure of the lung to expand, early thoracic surgical consultation is generally requested within 3-5 d<sup>[65]</sup>. Although surgical intervention is considered very effective and safe with a low recurrence rate, the studies reaching this conclusion do not include critically ill patients<sup>[66,67]</sup>.

Muscle relaxants might be effective to decrease the incidence of pneumothorax for patients with ARDS. A recent study by Papazian *et al*<sup>[68]</sup> reported a significant reduction in pneumothoraces in patients with severe ARDS who received 48 h of paralysis.

#### OUTCOMES

Mechanically ventilated patients with pneumothorax had a significantly higher mortality rate than those without pneumothorax. de Lassence *et al*<sup>[3]</sup> reported that iatrogenic pneumothorax was associated with a greater than twofold increase in the risk of death. In terms of iatrogenic pneumothorax, the mortality rate of patients with ventilator-related pneumothorax was significantly higher than that of patients with procedure-related pneumothorax<sup>[20]</sup>. Ventilator-related pneumothorax patients with tension pneumothorax had a higher risk of death<sup>[20]</sup>.

## CONCLUSION

Pneumothorax is a medical emergency and a disease with a high mortality rate; it requires a careful awareness, prompt recognition and intervention to reduce morbidity and mortality. Most patients with PRMV have underlying pulmonary diseases, the most common of which are pneumonia, ARDS and obstructive lung disease. Pneumothorax presentation and radiographic findings may be subtle or atypical. Whereas a CT scan is the gold standard for diagnosis, the use of ultrasonography has benefits in both diagnosis and management of pneumothorax. Most of the ventilated patients with a pneumothorax require immediate treatment with tube thoracostomy because of the high risk of progression to a tension pneumothorax. Small-bore catheters are now preferred in the majority of ventilated patients.

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> P- Reviewers: Kastelik JA, Kawai H L- Editor: Roemmele A E- Editor: Liu SQ







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