



Research Article

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Pulmonary Barotrauma in COVID-19 Patients Under Non-invasive Ventilation Support: Case Series

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Abstract

Purpose: Pulmonary barotrauma (PBT) is a complication typically seen in intubated patients but is a rare event in those using non-invasive ventilation (NIV) support. In this case report, we identified five patients with COVID-19 who developed PBT on NIV support in a hospital of southern Brazil from March to April 2021.

Results: All the patients were non-smokers with no pre-existing lung disease. They required oxygen therapy which included escalation from nasal cannula to NIV. PBT was diagnosed later in the course of the disease (17.6 ± 4.3 days) and after a mean time of 6 days on NIV support.

Conclusions: We found high PBT rate in COVID-19 patients. The patients developed PBT without being subjected to high levels of EPAP. Therefore, clinicians should be aware of the high risk of PBT among COVID-19 patients, since barotrauma correlates with longer ICU stay and hospitalization.

Keywords: Pulmonary barotrauma, COVID-19, Non-invasive ventilation

Introduction

The new coronavirus pandemic has been worrying the world since 2019. In order to improve clinical outcomes, early recognition of clinical deterioration and appropriate respiratory support are crucial in coronavirus disease-19 (COVID-19) patients. Most patients who become seriously ill with COVID-19 do so primarily due to an acute respiratory illness [1] and about 14% will require non-invasive ventilation (NIV) support [2].

Despite novel therapies and ventilation strategies, morbimortality remains high for COVID-19 patients, often due to

complications associated with invasive mechanical ventilation (IMV) support, such as pulmonary barotrauma (PBT) [3]. On the other hand, barotrauma is a rare complication of NIV support and tends to be only seen in patients who are chronically using this therapy [4,5]. In the related viral epidemic of SARS in 2002, approximately 10% of patients on NIV were described to develop PBT, with diffuse alveolar damage as the suggested mechanism [6]. However, there is little data on the incidence of barotrauma in critically ill patients with COVID-19, especially in those using only NIV.

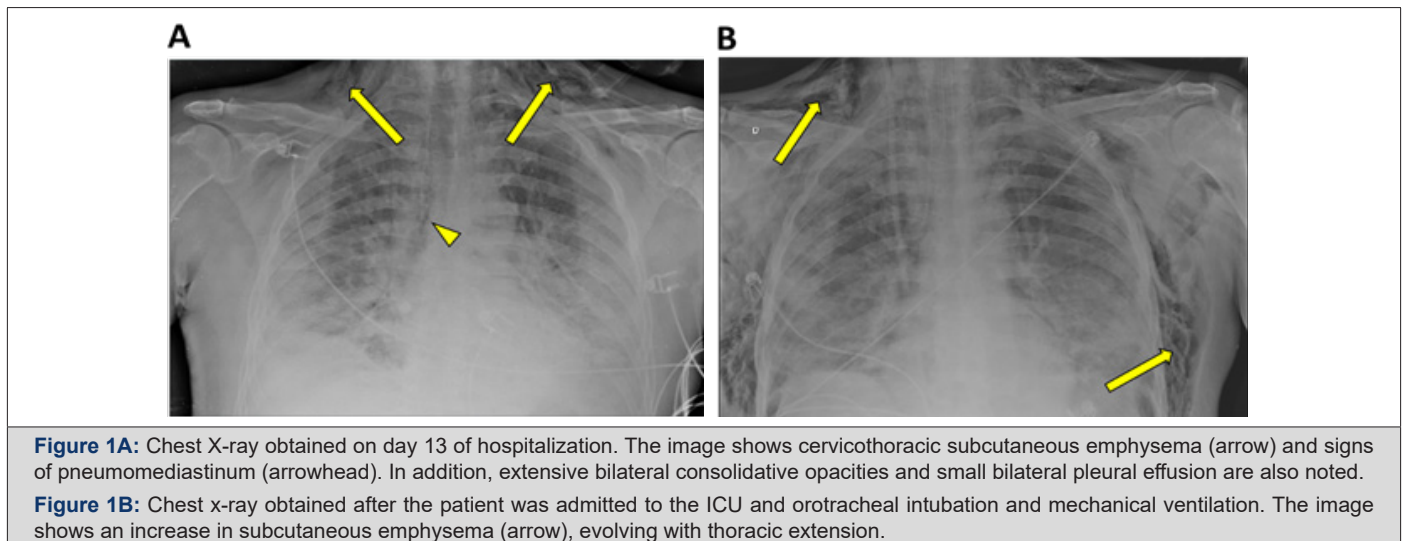


Herein, we describe five cases of PBT in patients with COVID-19 using NIV along with a brief literature review on the possible pathophysiologic mechanisms. The cases were selected from a cohort of patients who were hospitalized between March 10 and April 1 of 2021 in a semi-intensive care unit at a tertiary hospital in southern Brazil.

Patient 1

A 68-year-old male with a past medical history of hypertension presented to the hospital with 11-day-onset respiratory symptoms and progressively worsening dyspnea. He had a positive RT-PCR for SARS-CoV-2 at the third day of symptoms. At the emergency department, the patient received amoxicillin-clavulanate and completed a 7-day course of treatment. Dexamethasone was

started on day 1, and intermittent bilevel positive pressure airway (BPAP) support was started on day 4. Due to lack of respiratory improvement, he underwent a CT scan, which revealed signs of organizing pneumonia (COP), and prednisone therapy at 1 mg/kg/day was started. Throughout the following days, the patient progressively needed more aggressive ventilatory parameters to maintain targeted oxygen saturation. On day 13 of hospitalization, daily medical examination found crepitation under cervical skin which was not present on the previous day. A bedside chest X-ray found cervical subcutaneous emphysema and signs of pneumomediastinum [Figure 1]. After 2 days, he experienced a significant clinical deterioration, requiring IMV and was transferred to the ICU.



Patient 2

A 69-year-old male with an unremarkable prior medical history and a positive RT-PCR for SARS-CoV-2 was admitted to the hospital on the 7th day of respiratory symptoms. Prior to admission, he had used azithromycin, hydroxychloroquine, ivermectin and colchicine. Oxygen therapy with nasal cannula (NC) was started and dexamethasone was prescribed. A CT scan upon admission showed ground-glass opacities in both lungs and at least 40% of parenchymal impairment. Three days later, the patient was transferred to the ICU due to respiratory deterioration, and intermittent BPAP was started. On day 4, the patient was requiring longer periods under NIV support. A chest X-ray was performed [Figure 2] and identified cervical subcutaneous emphysema and a discrete pneumothorax area on the right lung. During ICU admission, the patient required IMV. In the follow-up, he was diagnosed with ventilator-associated pneumonia, evolving with lung abscess and septic pulmonary embolism, and was treated with multiple broad-spectrum antimicrobial regimens.

Patient 3

A male patient, 47 years old with no previous medical history, was admitted to the hospital with a 10-day history of high fever, dry cough and worsening respiratory symptoms. He also tested positive for SARS-COV-2. At the emergency room, he presented with low oxygen saturation and high respiratory frequency rate. On day 2 of hospitalization, the patient required escalation from NC to intermittent BPAP due to worsening of respiratory condition and was encouraged to self-prone. On day 4, due to neck subcutaneous crepitation, a chest X-ray was performed and found subcutaneous emphysema and signs of pneumomediastinum [Figure 3], without significant clinical repercussion, which were conservatively managed. After 2 days, he required IMV and was transferred to the ICU. Due to clinical deterioration, a CT angiography was performed and found pulmonary thromboembolism (PTE) signs, and anticoagulation was promptly started. On ICU, the patient evolved favorably and after 10 days was discharged to a non-intensive care unit.

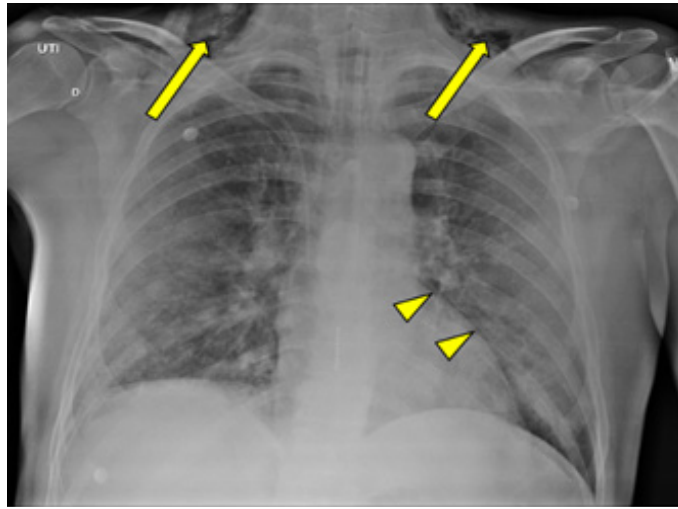


Figure 2: Chest X-ray performed on day 4 of hospitalization, after admission on ICU. The image showed bilateral infiltrative opacities, cervical subcutaneous emphysema (arrows) and signs of pneumomediastinum (arrowheads).

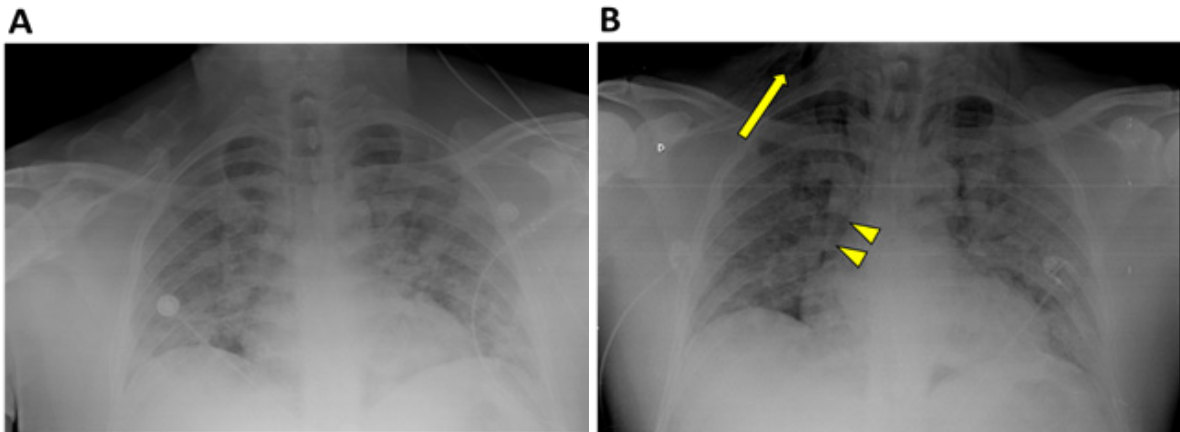


Figure 3A: Chest X-ray performed on admission, which found only extensive bilateral pulmonary opacities.

Figure 3B: Chest X-ray performed on day 4 of hospitalization. The findings include bilateral infiltrative opacities, cervical subcutaneous emphysema (arrow) and signs of pneumomediastinum (arrowheads), the last two which were not present on the previous X-ray.

Patient 4

A 49-year-old male patient with hypertension and in use of losartan was admitted to the hospital with a 12-day history of flu-like symptoms. RT-PCR was positive for COVID-19. He was treated previously to admission with ivermectin, hydroxycloloquin and azithromycin. The patient required oxygen therapy which included escalation from NC to intermittent BPAP due to worsening hypoxemia and was recommended to self-prone. CT scan performed short after admission found approximately 75% of pulmonary parenchyma impairment. BPAP was then changed to continuous positive airway pressure (CPAP) of 10cm H₂O, still

in intermittent use. On day 8 of hospitalization, while on CPAP of 12 cmH₂O, the patient presented with respiratory deterioration. At bedside examination, he presented with cervical crepitation and was diagnosed with subcutaneous emphysema. Due to rapid respiratory failure, he immediately required intubation and was transferred to the ICU. A control bedside X-ray performed shortly after intubation observed not only subcutaneous emphysema, but also a pneumothorax on the right lung, requiring surgical chest tube placement [Figure 4]. Despite initial clinical improvement, with adequate pulmonary reexpansion, the patient evolved unfavorably and later succumbed to his illness.

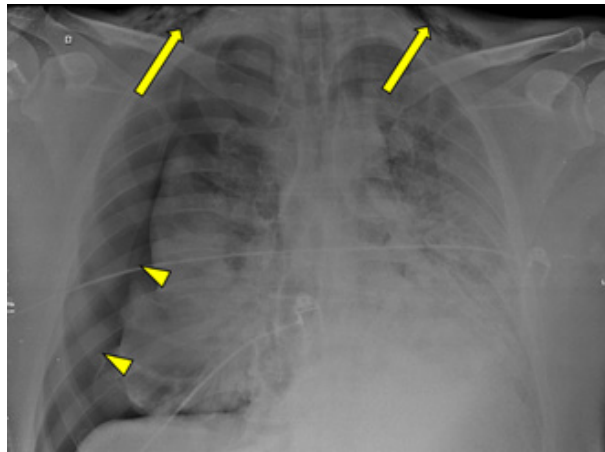


Figure 4: On day 8, a control chest X-ray performed short after intubation found cervical subcutaneous emphysema (arrows) and a large pneumothorax on the right with subtotal lung collapse (arrowheads), requiring surgical chest tube placement.

Patient 5

A 62-year-old female patient, with previous medical history of hypertension and in use of losartan, was admitted to the hospital with worsening dyspnea and hypoxemia. She was on the 7th day of symptoms and had a positive RT-PCR for COVID-19. She was treated with 5 days of azithromycin and dexamethasone 6 mg was also prescribed. She initially required oxygen therapy with NC and later needed escalation to CPAP of 10cm H₂O due to worsening dyspnea. CT scan with 50% of pulmonary parenchyma impairment. On day 7 of hospitalization, she changed ventilatory support from CPAP to BPAP due to CO₂ retention and began to self-prone. A new CT scan was performed to investigate the respiratory worsening and signs of COP were observed. Dexamethasone was then suspended, and prednisone at 1 mg/kg/day was prescribed. She continued requiring increasing ventilator parameters and was under BPAP support with inspiratory positive airway pressure (IPAP) of 20cm

H₂O and expiratory positive airway pressure (EPAP) of 12cm H₂O. After 3 days on continuous NIV, the patient developed respiratory failure and daily routine examination found cervical crepitation. Due to established respiratory insufficiency, intubation was required, but she rapidly evolved with cardiopulmonary arrest. Adequate cardiopulmonary resuscitation was performed by the medical team, without success.

Discussion

In our study, 56 patients were admitted to the semi-intensive care unit with severe respiratory disease due to COVID-19 infection. Out of these, 21 were women and 35 were male patients. Virtually every patient used NIV at some point during hospitalization. All the case-patients were non-smokers with no pre-existing lung disease, both considered risk factors for barotrauma [7]. The ventilator parameters and demographic characteristics of these patients are summarized in [Table 1].

Table 1: Demographic and clinical characteristics of PBT patients.

	Patient 1	Patient 2	Patient 3	Patient 4	Patient 5	Mean (SD)
Age (Years)	68	69	47	49	62	59 (10.4)
Gender	Male	Male	Male	Male	Female	-
Smoker	No	No	No	No	No	-
Corticosteroids	Yes	Yes	Yes	Yes	Yes	-
Days of illness to PBT	24	14	13	19	18	17.6 (4.3)
Days of NIV to PBT	9	4	3	5	9	6 (2.8)
Tidal volume to predicted body weight (ml/kg)	8.8	8.6	4.9	8.7	10.7	8.3 (1.9)
CPAP or IPAP previous to PBT (cmH ₂ O)	16	10	10	12	20	13.6 (4.3)
EPAP previous to PBT (cmH ₂ O)	11	8	8	-	12	9.7 (2.0)

All the values are presented as mean ± standard deviation (SD). PBT = Pulmonary barotrauma; NIV = Non-invasive ventilation; CPAP = Continuous positive air pressure; IPAP = Inspiratory positive air pressure; EPAP = Expiratory positive air pressure.

The patients developed PBT later in the course of the disease (17.6 ± 4.3 days) and after an average time of 6 days using NIV support. The most recent guidelines recommend a target tidal volume (V_t) < 9 ml/kg when using NIV, since high volumes (> 9.2 or 9.5 mL/kg) under NIV are associated with increased mortality [8]. In our case, the mean V_t was 8.3 ± 1.9 ml/kg.

Barotrauma is seen in 5-12% of patients with acute respiratory distress syndrome (ARDS) receiving IMV but is a rare event in patients receiving NIV [9]. A recent study found that the rates of barotrauma in COVID-19 patients receiving IMV are significantly higher than the observed in other ARDS causes (15% vs. 5%) [10]. In our cohort, the incidence of PBT in patients under NIV support was approximately 9%, probably demonstrating that even in non-mechanically ventilated patients this rate is still high.

One possible explanation for the association between high incidence of PBT and COVID-19 is that the virus-induced ARDS could be a unique disease entity with a novel pathophysiology [11,12]. Kahn et al. suggested that the COVID-19 probably causes a heightened inflammatory response, even more intense than other infectious diseases due to our lack of prior immunity, manifested as diffuse alveolar damage [13]. Gattinoni et al. [11] have recently described that patients with COVID-19 ARDS present an atypical form of the syndrome. In this study, the most important observation was a dissociation between the patient's relatively well-preserved lung mechanics (high compliance) and the severity of hypoxemia [11]. The standard of ARDS management is the use of lung-protective strategies that combine low tidal volume and relatively high PEEP [14]. However, high levels of PEEP may lead to lung overdistension associated with an increase of alveolar dead space [15].

The occurrence of subcutaneous emphysema and pneumomediastinum is generally related to diffuse alveolar damage causing rupture and air leakage. Patients with a longer duration of illness have more diffuse alveolar damage, with a greater extent of pulmonary impairment, which, added to alveolar overdistension by using high PEEP, could lead to an increased risk of developing barotrauma.

Conclusion

In patients who are being treated with NIV and who present with clinical signs of respiratory failure, intubation should be prioritized to avoid excessive intrathoracic negative pressures and self-inflicted lung injury [11,16]. However, regardless of the underlying mechanism, our case report indicates that clinicians should be aware of the high rate of barotrauma seen among patients with COVID-19 using NIV, since it correlates with poor outcomes

[17]. Moreover, it also encourages clinicians to have a low threshold to suspect PBT in COVID-19 patients.

Disclosure statement

The authors have no conflicts of interest to declare.

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