

Evaluation of the cardiopulmonary status using a noninvasive respiratory profile monitor in chronic obstructive lung disease patients during low-ventilation strategy

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Background Patients with chronic obstructive pulmonary disease (COPD) patients are susceptible to complications, especially volutrauma, during the period of mechanical ventilation; low ventilation is a safe strategy to avoid these complications. Noninvasive capnography is a suitable technique for monitoring and assessing the cardiac and the pulmonary status of these patients during the period of mechanical ventilation.

Objectives Assessment of the cardiac and the pulmonary status of two COPD patient groups receiving mechanical ventilation with a low tidal volume strategy using a noninvasive CO₂ respiratory profile monitor (volumetric capnography).

Patients and methods Forty patients were recruited in the respiratory ICU of Abbassia Chest Hospital; these patients were divided into two groups: 20 COPD patients with the predominant pathology of chronic bronchitis (CB) and 20 patients with the predominant pathology of emphysema disease, who presented with clinical and radiological evidence of chronic obstructive lung disease and were in need of mechanical ventilation. All the patients in the study were followed up three times per day until weaning; data were recorded on admission, after 24 h and before weaning using volumetric capnography.

Results There was significant correlation between EtCO₂ and arterial PCO₂ during the whole period of mechanical

ventilation in CB and emphysematous patients; the mean dead-space fraction was significantly higher in the emphysema group than in the CB group. There was a significant negative correlation between the mean values of V_d/V_t and the pulmonary capillary blood flow on admission and after 24 h in the emphysema group. The mean cardiac output, the mean stroke volume, and the pulmonary capillary blood flow increased significantly before extubation in the CB group, in contrast to the emphysema group in which there was an insignificant difference.

Conclusion Volumetric capnography could be helpful in assessing the severity of functional disturbances, and the use of more refined noninvasive parameters will be of value in managing and monitoring COPD patients during the whole period of mechanical ventilation. *Egypt J Broncho* 2014 8:44–50

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Keywords: capnography, chronic bronchitis, emphysema, mechanical ventilation

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Introduction

Invasive mechanical ventilatory support has two important considerations in respiratory failure caused by acute exacerbation of chronic obstructive pulmonary disease (COPD): they are minimizing regional overdistention and managing positive end-expiratory pressure (PEEP). Overdistention injury occurs when an excessive end-inspiratory alveolar 'stretch' physically damages alveolar structures and produces local and systemic inflammation (ventilator-induced lung injury). This stretch injury may be a consequence of excessive tidal volumes. This has led to recommendation to reduce tidal volumes (e.g. 5–7 ml/kg) to protect the lung in acute exacerbation of COPD [1]. Monitoring of the ventilated patient should focus on the assessment of patient response to and titration of mechanical ventilation, while avoiding complications [2]. The use of volumetric capnography in mechanically ventilated patients has the capability

to monitor not only lung mechanics but also cardiac and respiratory interaction noninvasively [3].

Patients and methods

Study design

Forty patients, who presented with clinical and radiological evidence of COPD with respiratory failure due to exacerbation and were in need of mechanical ventilation, were recruited in the respiratory ICU of Abbassia Chest Hospital in the period between 2011 and 2013. These patients were divided into two groups: 20 COPD patients with the predominant pathology of chronic bronchitis (CB) and 20 patients with the predominant pathology of emphysema disease; we used clinical criteria, clinical history with compatible physical findings, and evidence of hyperinflation on chest radiograph, in support of the diagnosis of COPD and for differentiation between

the two groups [4]. All patients were subjected to full history taking from a relative of the patient, clinical examination, chest radiography, computed tomography, ECG, echocardiography, and arterial blood gas measurement. Continuous infusion of midazolam and elective invasive mechanical ventilation and volumetric capnography monitoring were performed. All the patients in the study were followed up three times per day until weaning, and the data were recorded on admission, after 24 h and before weaning.

Monitoring

Variables measured included the cardiac output, the stroke volume, the pulmonary capillary blood flow (PCBF), EtCO₂, VCO₂, V_t/V_d, V_t alv, MV alv, Raw, PEEP, and compliance.

Data analysis

Analysis of data was performed using the Statistical Package for Social Sciences (SPSS version 15.0.1 for Windows; SPSS Inc., Chicago, Illinois, USA).

- (1) Descriptive statistics:
 - (a) Parametric data were expressed as range and mean ± SD.
 - (b) Nonparametric data were expressed as frequency and percentage.
- (2) Analytical statistics:
 - (a) Paired *t*-test.
 - (b) Independent sample *t*-test.
 - (c) Analysis of variance test.

Results

Forty participants were chosen randomly. Twenty of them had CB: this group consisted of 85% men, with a mean age of 63.75 years; and 20 participants had emphysema: all of them in this group were men with a mean age of 59 years. CB patients in this study

were successfully weaned from mechanical ventilation within 3–9 days, whereas emphysematous patients took longer (3–15 days). Tracheostomy operation was performed in 12.5% patients in the emphysematous group at a median time of 15 days from the beginning of mechanical ventilation; the main demographic data are illustrated in Table 1.

On admission, PO₂ in the CB group was significantly less than in the emphysema group with *P* value of 0.013 as shown in Fig. 1. Significantly better values of PH and PCO₂ were observed in the CB group than in the emphysema group after 24 h of mechanical ventilation, with *P* values of 0.014 and 0.009, respectively, as shown in Figs. 2 and 3.

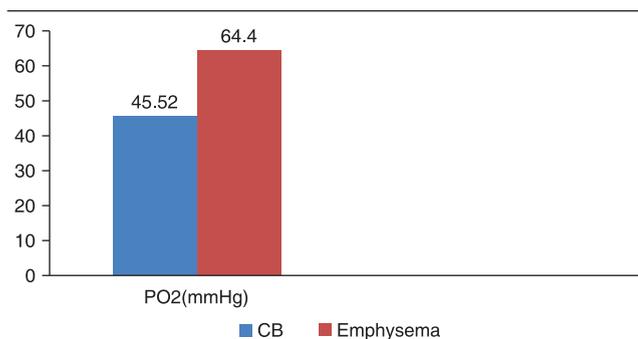
In the CB group, there was significant improvement in the hemodynamic parameters during the whole period of mechanical ventilation with regard to blood pressure, pulse, and temperature. In the emphysema group, there were no significant differences in the hemodynamic parameters except for pulse, which showed significant improvement until trial of weaning. On comparison, there was no significant difference between the two groups.

Table 1 Demographic data of the two groups (n = 20)

	CB group	Emphysema group	<i>P</i>
Sex [N (%)]			
Male	17 (85)	20 (100)	0.22
Female	3 (15)	0	
Age (years)			
Range	53–86	45–79	0.13
Mean ± SD	63.75 ± 10.06	59 ± 9.5	
Tracheostomy [N (%)]	0	5 (25)	0.055
Duration of MV (days)			
Range	3–9	3–15	0.0001
Mean ± SD	4 ± 1.5	8.1 ± 3.7	

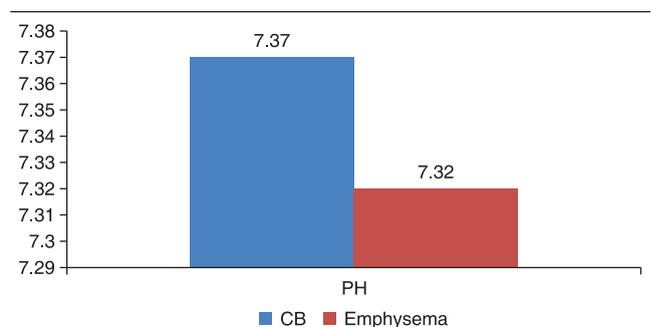
CB, chronic bronchitis; MV, mechanical ventilation.

Fig. 1



Comparison between chronic bronchitis (CB) and emphysema groups regarding PO₂: a significant decrease was observed in PO₂ in the CB group compared with the emphysema group (*t* = 2.57, *P* = 0.013).

Fig. 2

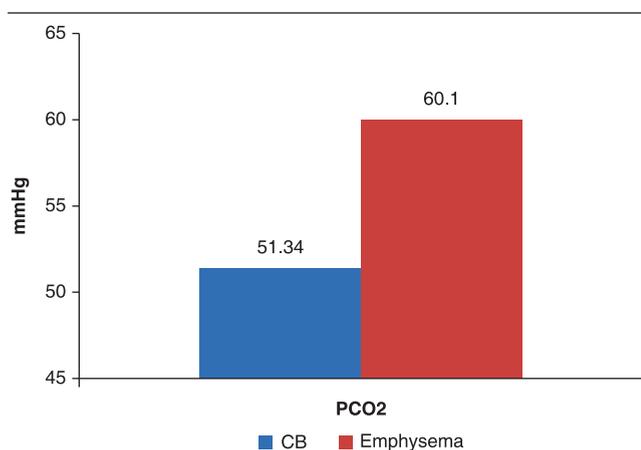


Comparison between the two groups regarding PH after 24 h: a significant decrease in PH was observed in the emphysema group compared with the chronic bronchitis (CB) group (*t* = 2.86, *P* = 0.014).

Regarding the cardiac parameters, pulmonary blood flow, cardiac output, and stroke volume, there was no significant difference between the two groups during the whole period of mechanical ventilation. At baseline in CB, the mean stroke volume and the cardiac index were 49.5 ml/beat and 2.43 l/min/m², respectively, and increased to 69.1 ml/beat ($P < 0.01$) and 3.1 l/min/m² ($P < 0.01$), respectively, before extubation; also, the mean PCBF level and the cardiac output were 3.6 and 5.2 l/min, respectively, at baseline compared with 4.2 l/min ($P < 0.05$) and 6.1 l/min ($P = 0.05$), respectively, before extubation. In emphysematous patients, only the stroke volume at baseline of 47.2 ml/beat increased significantly to 61.3 ml/beat ($P < 0.05$) before extubation as shown in Table 2.

On admission, the mean values of the dead-space fraction, the peak expiratory flow (PEF), and the PEEP measured among emphysema patients group were significantly higher than the values among patients in the CB group, with P values of 0.000, 0.004, and 0.004, respectively, as shown in Fig. 4a, whereas the mean values of EtCO₂ and V_t in the CB group were significantly higher than the values in the emphysema

Fig. 3



Comparison between the two groups regarding PCO₂ after 24 h: a significant increase in PCO₂ was observed in the emphysema group compared with the chronic bronchitis (CB) group ($t = 2.79$, $P = 0.009$).

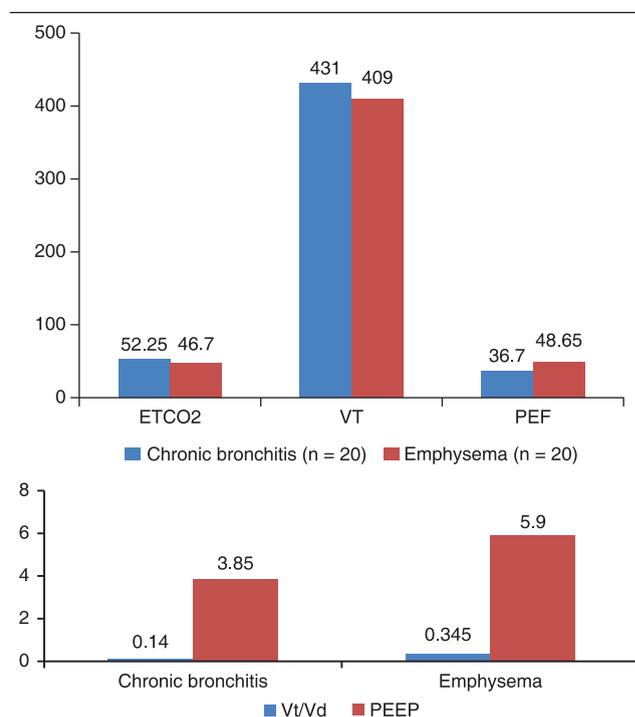
group, with P values of 0.025 and 0.044, respectively, as shown in Fig. 4b.

After 24 h, the mean dead-space fraction, the PEF and the PEEP were significantly increased in the emphysema group compared with the CB group, with P values of 0.000, 0.021, and 0.002, respectively, as shown in Fig. 5.

Before extubation, the mean dead-space fraction, the PEF, and the PEEP were significantly higher in the emphysema group than in the CB group, with P values of 0.048, 0.015, and 0.002, respectively (Fig. 6).

With regard to the two groups studied, the end-tidal carbon dioxide was significantly correlated to the arterial partial pressure of carbon dioxide on admission, after 24 h and before extubation.

Fig. 4



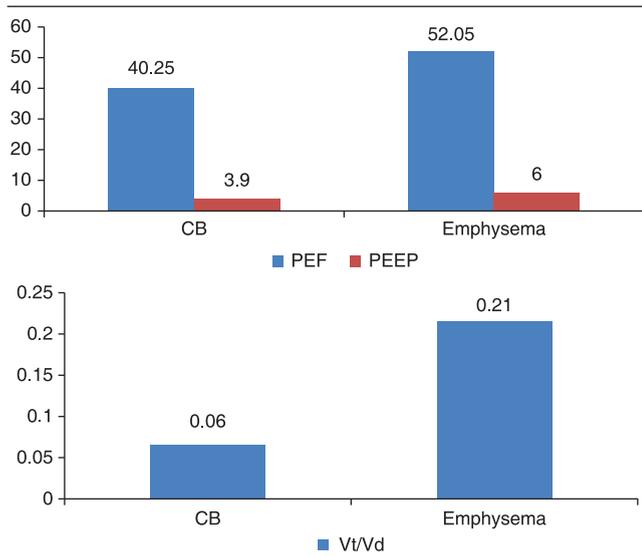
The difference between chronic bronchitis (CB) and emphysema groups with regard to their (a) EtCO₂, peak expiratory flow (PEF), and tidal volume (VT) and (b) V_t/V_d and positive end-expiratory pressure (PEEP) on admission.

Table 2 Cardiac parameters of the chronic bronchitis and the emphysema groups by capnography

	Variables	Baseline (mean ± SD)	After 24 h (mean ± SD)	Before extubation (mean ± SD)	P
Chronic bronchitis	COP (l/min)	5.25 ± 1.4	5.3 ± 1.3	6.1 ± 1.3	0.05
	SV (ml/beat)	49.5 ± 14.8	54.6 ± 15.2	69.1 ± 16.5	<0.01
	CI (l/min/m ²)	2.43 ± 0.5	2.5 ± 0.5	3.1 ± 0.7	<0.01
	PCBF (l/min)	3.6 ± 0.7	3.7 ± 0.8	4.2 ± 0.9	<0.05
Emphysema	COP (l/min)	5.3 ± 1.5	5.7 ± 1.2	5.4 ± 1.5	>0.05
	SV (ml/beat)	47.2 ± 17.8	59.3 ± 16.5	61.3 ± 16.9	<0.05
	CI (l/min/m ²)	2.8 ± 0.7	2.8 ± 0.7	2.8 ± 0.8	>0.05
	PCBF (l/min)	3.7 ± 1.03	3.7 ± 0.9	3.5 ± 1.03	>0.05

CI, cardiac index; COP, cardiac output; PCBF, pulmonary capillary blood flow; SV, stroke volume.

Fig. 5



The difference between chronic bronchitis (CB) and emphysema groups with regard to their (a) peak expiratory flow (PEF) and positive end-expiratory pressure (PEEP) and (b) V_t/V_d after 24 h.

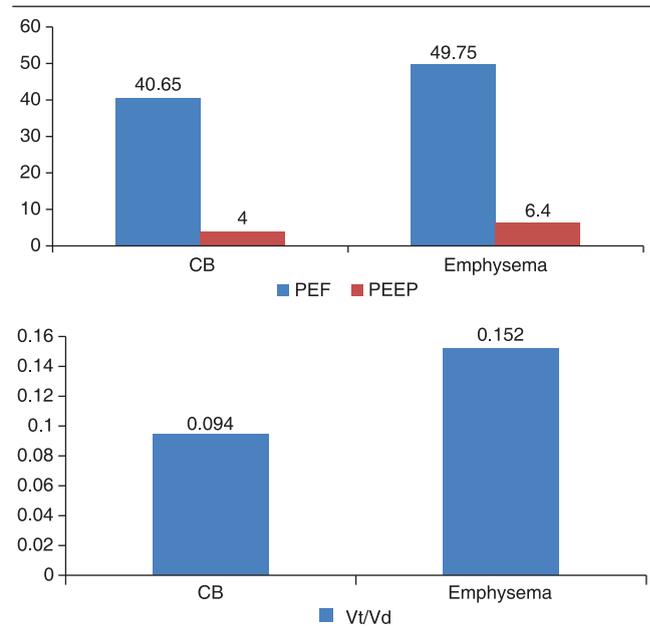
In the CB group, there was a highly significant negative correlation between the dead-space fraction and the cardiac output during all periods of mechanical ventilation. There was also a significant negative correlation between the two variables in the emphysema group on admission and after 24 h.

In the emphysema group, there was a highly significant negative correlation between dead-space fractions and PCBF on admission and it also had a significant negative correlation after 24 h (Tables 3–6).

Discussion

This study enrolled 40 COPD patients and differentiated them into two groups, according to the history, clinical findings, and radiological findings of each group: there were 20 COPD patients, with the predominant pathology of CB, and 20 patients with the predominant pathology of emphysema diseases. This selective methodology was partially in agreement with the methodology used by Farah and Makhoul [5], who enrolled COPD patients and dealt with them as one group without differentiated them into the CB and the emphysematous groups, but their study depended on history and clinical findings to diagnose the disease. Frazier *et al.* [6] measured the mean cardiac output by volumetric capnography, which was 5.3 l/min at baseline during mechanical ventilation and increased significantly to 6.5 l/min during the weaning trial by continues positive airway pressure (CPAP) trial ($P = 0.036$). These were in agreement with the results in the present study as there was a significant increase in the mean cardiac output measured by volumetric

Fig. 6



The difference between chronic bronchitis (CB) and emphysema groups with regard to their (a) peak expiratory flow (PEF) and positive end-expiratory pressure (PEEP) and (b) V_t/V_d before extubation.

capnography of the CB group at baseline (5.25 ± 1.4 l/min), which reached 6.1 ± 1.3 l/min before extubation, whereas in the emphysematous group, there was no significant difference between the two stages. In our study, the cardiac output of the two groups was recorded using the partial CO_2 rebreathing technique, which yielded a mean cardiac output at baseline of 5.25 ± 1.4 and 5.3 ± 1.5 l/min in the CB and the emphysematous groups, respectively. This coincides with the study of Jérôme *et al.* [7] which enrolled 20 consecutive mechanically ventilated patients who had acute respiratory distress syndrome (ARDS). They measured the cardiac output after a 2-h period of hemodynamic stability. The mean cardiac output value was 5.8 ± 1.7 l/min with the partial carbon dioxide rebreathing technique. Their study compared this method with thermodilution in which the mean cardiac output was 6.7 ± 1.9 l/min and there was significant correlation between the two methods. The mean stroke volume at baseline was 49.5 ± 14.8 and 47.2 ± 17.8 ml/beat, respectively, and increased significantly to 69.1 ± 16.5 and 61.3 ± 16.9 ml/beat in the CB and the emphysema groups, respectively, during weaning. These results were in agreement with Frazier *et al.* [6] who studied the hemodynamic function during baseline mechanical ventilation and during a trial of CPAP by the carbon dioxide rebreathing technique: the mean stroke volume was 52 ± 36 ml/beat at baseline during mechanical ventilation and increased significantly to 78 ± 38 ml/beat ($P < 0.001$) during the CPAP trial. The pulmonary blood flow was measured noninvasively by the partial

Table 3 Correlation between the end-tidal carbon dioxide and the arterial partial pressure of carbon dioxide

EtCO ₂ /PaCO ₂	On admission	After 24 h	Before extubation
Chronic bronchitis			
<i>r</i>	0.7	0.9	0.8
<i>P</i>	0.00	0.00	0.00
Emphysema			
<i>r</i>	0.23	0.3	0.7
<i>P</i>	0.3	0.1	0.00

Table 4 Correlation between external positive end-expiratory pressure and airway resistance

External PEEP/Raw	On admission	After 24 h	Before extubation
Chronic bronchitis			
<i>r</i>	-0.09	-0.125	-0.513
<i>P</i>	0.66	0.58	0.021
Emphysema			
<i>r</i>	0.12	0.14	0.33
<i>P</i>	0.61	0.53	0.146

There was a significant negative correlation between the external PEEP and airway resistance before extubation among the chronic bronchitis group. PEEP, positive end-expiratory pressure.

Table 5 Correlation between the dead-space fraction ratio V_d/V_a and the cardiac output

V_d/V_a /COP	On admission	After 24 h	Before extubation
Chronic bronchitis			
<i>r</i>	-0.55	0.74	-0.803
<i>P</i>	0.008	<0.0001	<0.0001
Emphysema			
<i>r</i>	0.59	-0.48	-0.12
<i>P</i>	0.006	0.029	0.59

COP, cardiac output.

Table 6 Correlation between the dead-space fraction V_d/V_a and the pulmonary capillary blood flow

V_d/V_a /PCBF	On admission	After 24 h	Before extubation
Chronic bronchitis			
<i>r</i>	0.040	0.05	-0.09
<i>P</i>	0.86	0.8	0.69
Emphysema			
<i>r</i>	-0.6	-0.5	-0.1
<i>P</i>	0.005	0.02	0.6

PCBF, pulmonary capillary blood flow.

CO₂ rebreathing technique: the mean pulmonary blood flow was 3.6 ± 0.7 l/min in CB at baseline and it significantly increased before extubation to reach 4.2 ± 0.9 l/min, in contrast to the emphysema group, in which there was an insignificant difference, with a mean PBF of 3.7 ± 1.03 l/min at baseline and 3.5 ± 1.03 l/min before extubation. These results are in agreement with the study of Jérôme *et al.* [7], in which the mean value of PCBF was 4.6 ± 1.3 l/min by the carbon dioxide rebreathing technique. In this study, before extubation, the mean dead-space fraction measured by volumetric capnography in the emphysema group was 0.15 ± 0.1 , which was significantly higher

than that of the CB group (0.09 ± 0.06); this was in agreement with González-Castroa *et al.* [8] who enrolled 76 patients for mechanical ventilation, out of whom 14 patients were diagnosed to have COPD with exacerbation, nine patients were diagnosed to have pneumonia, and the remaining had other causes: the mean value of V_d/V_t in the 59 extubated patients was 0.48 ± 0.09 , whereas in the 17 patients with failed extubation, the mean value of V_d/V_t was 0.65 ± 0.08 . Our result demonstrated that the mean dead-space fraction measured by volumetric capnography on admission and before extubation reduced from 0.345 to 0.15, respectively, in the emphysema group, which was significantly higher than that of CB group, which was 0.14 and reduced to 0.09, respectively. These results were in agreement with Kallet *et al.* [9], who measured the ratio of the physiologic dead-space to tidal volume V_d/V_t with volumetric capnography before therapy with human recombinant activated protein C, and found a reduction in V_d/V_t from 0.55 to 0.27. In the present study, before extubation, there were two modes of weaning used: pressure support and T piece. There was no significant difference between the two modes between the two groups. The mean values of dynamic compliance were 41.05 and 43.00 ml/cmH₂O, the static compliance were 25.8 and 26.90 ml/cmH₂O, airway resistance was 24.30 and 26.75 cmH₂O/l/s, and the mean dead-space fraction values were 0.15 and 0.09 in the emphysema group and the CB group, respectively. These results were comparable to those of El Ghamrawy *et al.* [10] who used BIPAP compared with pressure support for the weaning of 32 COPD patients with acute respiratory failure. The respiratory system static, dynamic compliance, and the resistance were calculated by equations, but the ratio of dead-space to tidal volume was calculated automatically from capnography and displayed on the ventilator screen. They did not use the non-rebreathing CO₂ monitor as in our study. The mean level of respiratory dynamic compliance with BIPAP was 21.8 ml/cmH₂O, which was significantly lower than its level with PS (25.0 ml/cmH₂O); the static compliance was 38.9 ± 11.3 ml/cmH₂O with BIPAP, which was insignificantly higher than its level with PS (39.3 ± 12.1 ml/cmH₂O); these values were less than our values, and this may be explained by the fact that they used the method of calculation. The mean level of resistance with bi-level positive air pressure (BIPAP) was 28.3 cmH₂O/l/s, which was significantly higher than the corresponding level with PS (22.8 cmH₂O); these values coincided with our result. The mean dead-space ventilation was 0.57 and 0.54 with BIPAP and PS, respectively, with no significant difference, but these results were slightly higher than our result. In the present study, the *P*(a-et) CO₂ gradient decreased from 25.15 ± 13.6 on admission to 8.15 ± 6.2 before extubation in the emphysema

group, and the $P(a\text{-et})\text{CO}_2$ gradient decreased from 10.78 ± 10.84 to 5.35 ± 3.884 in the CB group. Defilippis *et al.* [11] studied the $P(a\text{-et})\text{CO}_2$ gradient in 20 patients with severe dyspnea and hypercapnia undergoing noninvasive ventilation. The $P(a\text{-et})\text{CO}_2$ gradient was measured at subsequent times: T0 (admission) was 60.7 ± 17.7 mmHg; it decreased progressively, reaching 8.4 ± 8 mmHg at T6 h and 4.7 ± 6.7 mmHg at T12 h, which was lower than the baseline value. A positive correlation was found between EtCO_2 and PaCO_2 values ($r = 0.89, P = 0.001$). In the present study, with regard to the CB group, EtCO_2 was significantly correlated to PaCO_2 on admission ($r = 0.7, P = 0.00$), after 24 h ($r = 0.9, P = 0.00$), and before extubation ($r = 0.8, P = 0.00$); however, in the emphysema group, the EtCO_2 was significantly correlated to the PaCO_2 only before extubation ($r = 0.7, P = 0.00$). These results were in agreement with Yosefy *et al.* [12], who enrolled 73 patients in their study. The medical diagnosis included 55–75.3% patients with pulmonary edema, 14–19.2% patients with exacerbation of COPD, and 4–5.5% patients with exacerbation of bronchial asthma. Significant correlation was found between EtCO_2 and arterial PCO_2 ($r = 0.792$). Razi *et al.* [13] confirmed a good correlation between the mean of EtCO_2 and PaCO_2 in each of the modes of SIMV, CPAP, and T-tube between EtCO_2 tensions with PaCO_2 measurements in mechanically ventilated patients with the following correlation coefficients and P values: SIMV ($r = 0.893, P < 0.0001$), CPAP ($r = 0.841, P < 0.0001$), and T-tube ($r = 0.923, P < 0.0001$), respectively. In the present study, in the emphysematous group, there was a highly significant negative correlation between the mean values of dead-space fractions V_t/V_d (0.3 ± 0.1) and PCBF on admission (3.7 ± 1.03) and after 24 h. These results were in agreement with the result of Tusman *et al.* [14] who tested whether the V_d/V_t can detect states of low PBF in a noninvasive manner. Fifteen patients who had undergone cardiopulmonary bypass (CPB) were studied. During constant ventilation, volumetric capnography was performed and the V_d/V_t was recorded. Before CPB, V_d/V_t was 0.36 ± 0.05 . During weaning from CPB, V_d/V_t decreased with increasing PBF. At CPBs of 80, 60, 40, and 20%, V_d/V_t values were $0.64 \pm 0.06, 0.55 \pm 0.06, 0.47 \pm 0.05,$ and 0.40 ± 0.04 , respectively ($P < 0.001$). After CPB, V_d/V_t values were similar to that at baseline (0.37 ± 0.04). Traditional approaches to mechanical ventilation use tidal volumes of 10 to 15 ml/kg of body weight, which result in volutrauma [15]. A prospective cohort of 361 ICUs from 20 countries with a total of 5183 mechanically ventilated patients with limited tidal volumes 6–8 ml/kg were included in the study. This study found that barotrauma was present in 2.9% of the patients with COPD, 6.3% of the patients with

asthma, 10.0% of the patients with chronic interstitial lung disease, 6.5% of the patients with acute respiratory distress syndrome, and 4.2% of the patients with pneumonia [16]. In the present study, barotrauma was not recorded in 40 patients in whom the low-ventilation strategy was used.

Conclusion

Low ventilation is a safe strategy to avoid complications such as volutrauma and barotrauma. Noninvasive capnography is a simple and safe bedside method for noninvasive estimation of pulmonary and cardiac parameters for monitoring COPD patients during the period of mechanical ventilation. There was significant correlation between EtCO_2 and arterial PCO_2 throughout the period of mechanical ventilation in CB and emphysematous patients; hence, monitoring PetCO_2 provided a good noninvasive assessment of hypercapnic episodes during weaning from mechanical ventilation. The $P(a\text{-et})\text{CO}_2$ gradient decreased before extubation in emphysematous and CB patients, and this indicates a change in perfusion to help with weaning. The mean dead-space fraction was significantly higher in the emphysema group than in the CB group throughout the period of mechanical ventilation. Also, there was a significant negative correlation between the mean values of dead-space fractions V_t/V_d and the PCBF in the emphysematous patients. Hence, early and repeated measurements of the pulmonary and the cardiac function by this monitor could provide clinicians with valuable information for prognosis and disease monitoring.

Acknowledgements

Conflicts of interest

There are no conflicts of interest.

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