



# Comparison of ETCO<sub>2</sub> Value and Blood Gas PCO<sub>2</sub> Value of Patients Receiving Non-invasive Mechanical Ventilation Treatment in Emergency Department

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## Abstract

Capnography is the non-invasive measurement and graphic representation of the partial pressure of CO<sub>2</sub> in expiration. Although there are many studies in the literature comparing the partial pressure of carbon dioxide (pCO<sub>2</sub>) and end-tidal CO<sub>2</sub> (ETCO<sub>2</sub>) values in patients who underwent IMV (invasive mechanical ventilation), there are no studies showing their interchangeable applicability in patients who received NIMV (non-IMV). We aimed to evaluate whether the use of ETCO<sub>2</sub> in the treatment process can replace pCO<sub>2</sub> use in patients scheduled for NIMV treatment in the emergency department. Patients who applied to the emergency department with respiratory distress between March 2019 and January 2020, who were diagnosed with acute cardiogenic edema or acute chronic obstructive pulmonary disease (COPD) exacerbation, and who needed NIMV were included in the study. General characteristics of the patients and the pCO<sub>2</sub> and ETCO<sub>2</sub> values were measured in the blood gas 1 h after the NIMV application was started. 64.2% (99 patients) of the patients included in the study were male, and 35.8% (55 patients) were female. The mean age of the patients included in the study was 69.1 ± 12.2 years. The mean pCO<sub>2</sub> values were measured as 52.6 ± 13.2. The mean of ETCO<sub>2</sub> values measured simultaneously was 33.6 ± 10.1. There was a significant difference between the controlled pCO<sub>2</sub> values and ETCO<sub>2</sub> values at the first hour of NIMV treatment (Z: - 10.640, *p* < 0.001). The ETCO<sub>2</sub> level was found to be different in our patients who received NIMV treatment, which could not be used instead of the pCO<sub>2</sub> level.

**Keywords** Capnography · pCO<sub>2</sub> · ETCO<sub>2</sub> · Non-invasive mechanical ventilation

## Introduction

When the complaints at admission to the emergency department are evaluated, the rate of dyspnea and respiratory distress is 2–3% [1]. Mechanical ventilation (MV) is a frequently used method in treating patients with respiratory distress because it reduces gas exchange and respiratory work and has two types

of use: invasive mechanical ventilation (IMV) and non-invasive mechanical ventilation (NIMV) [2]. Blood gas analysis is an important indicator in the follow-up and treatment of patients, and especially the partial pressure (pCO<sub>2</sub>) value of the carbon dioxide (CO<sub>2</sub>) gas dissolved in the blood provides guidance for selecting the mode of the mechanical ventilator and adjusting its settings [3].

Capnography is the non-invasive measurement and graphic representation of the partial pressure of CO<sub>2</sub> in expiration. It is also called end-tidal CO<sub>2</sub> (ETCO<sub>2</sub>) because the measurement is taken from expiratory air. ETCO<sub>2</sub> is mostly used in the emergency department to verify the location of the endotracheal tube in the intubated patient, to monitor the spontaneous return during cardiopulmonary resuscitation (CPR) in cardiac arrest cases, and to monitor the ventilation status in procedural sedoanalgesia. Using ETCO<sub>2</sub> instead of pCO<sub>2</sub> in the monitoring of intensive care patients undergoing IMV is an increasingly common method [4, 5].

It was found that there is a 1–4 mmHg difference between the pCO<sub>2</sub> value and the ETCO<sub>2</sub> value in the use of

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capnography for the monitoring of intubated patients on a mechanical ventilator. However, it was found that the difference between pCO<sub>2</sub> and ETCO<sub>2</sub> increased in situations such as decreased lung perfusion, pulmonary embolism, dead space ventilation, and cardiac arrest [6]. Although there are many studies in the literature comparing pCO<sub>2</sub> and ETCO<sub>2</sub> values in patients who underwent IMV, there are no studies showing their interchangeable applicability in patients who received NIMV.

## Aims and Objectives

Herein, we aim to describe to evaluate whether the use of ETCO<sub>2</sub> in the treatment process can replace pCO<sub>2</sub> use in patients scheduled for NIMV treatment in the emergency department.

## Methods

### Study Participants

This study was conducted as a prospective cross-sectional clinical study at a tertiary university hospital. Approval was obtained from the local ethics committee, and signed informed consent was obtained from all patients who participated in the study or from their first-degree relatives. Patients who applied to the emergency department with respiratory distress between March 2019 and January 2020, who were diagnosed with acute decompensated heart failure (ADHF) or acute chronic obstructive pulmonary disease (COPD) exacerbation, and who needed NIMV were included in the study. The need for NIMV was determined by clinical senior residents and specialist physicians. Being under 18 years of age, having pulmonary parenchymal diseases such as pneumonia, neuromuscular diseases, obstructive sleep apnea syndrome, obesity hypoventilation syndrome, patients who cannot tolerate NIMV treatment, and patients who did not agree to participate in the study were determined as the exclusion criteria. It was ensured that all patients who were treated were conscious, had sufficient cough and swallowing reflex, were clinically stable, and that there was no obstacle on the patient's face for the application of non-invasive ventilation.

Demographic information such as age, gender and history, vital signs (body temperature, pulse, blood pressure, respiratory rate), fingertip oxygen saturation (SpO<sub>2</sub>), main physical examination findings (rale, rhoncus, pretibial edema), and outcome conditions were noted in the patients who met the inclusion criteria. The pCO<sub>2</sub> and ETCO<sub>2</sub> values were measured in the blood gas 1 h after the NIMV application was started.

After the physician decided to apply NIMV in treatment, treatments of patients started with a non-return mask using the continuous positive airway pressure (CPAP) and bilevel

positive airway pressure (BiPAP) modes of the LTV 1200 (Vyaire Medical, Mettawa, Illinois) device. At the first hour of treatment, blood gas samples were taken to adjust the mode settings of the ventilator and to evaluate the treatment. Simultaneously, an A portable mainstream capnograph (Masimo EMMA Mainstream Capnograph; Masimo Corp., CA, USA) was placed between the connection of the mask and the hose, and ETCO<sub>2</sub> was measured. Venous blood gas samples were taken from the patients with 22G-tipped blood gas injectors. Testing of the samples taken were performed with the calibrated ABL800 FLEX © blood gas analyzer (ABL800) (Radiometer Medical ApS, Copenhagen, Denmark) device, and the pCO<sub>2</sub> value was recorded. The hospitalization and intubation needs of the patients were obtained from the records on the hospital automation system.

### Primary Endpoint

The primary endpoint of our study was the comparing pCO<sub>2</sub> and ETCO<sub>2</sub> values in patients who underwent NIMV.

### Secondary Endpoints

The secondary aim is to compare the results of pCO<sub>2</sub> and ETCO<sub>2</sub> the suitability of using interchangeably.

### Statistical Analysis

The data obtained were analyzed by using SPSS for Windows 22 package program. Descriptive numerical variables were presented as mean and standard deviation, and categorical variables were presented as number and percentage. Prior to the comparison of means, normal distribution was tested. The Mann-Whitney *U* test was used to compare the means in non-normally distributed groups, and the dependent groups' *t*-test was used to compare the means in normally distributed variables. Correlation analysis of variables was performed with the Pearson test for normally distributed data, and the Spearman test was used for non-normally distributed data. Bland-Altman analysis was used to determine the agreement between the two methods. Correlation coefficient *r* value of < 0.20 was considered very weak, 0.20–0.40 weak, 0.40–0.60 moderate, 0.60–0.80 high, > 0.80 very high correlation. *p* < 0.05 was considered statistically significant.

## Results

64.2% (99 patients) of the patients included in the study were male, and 35.8% (55 patients) were female. The mean age of the patients included in the study was 69.1 ± 12.2 years. 55.1% (85 patients) of the pre-diagnoses made by emergency department physicians were COPD, and 44.9% (69 patients)

were ADHF. Twenty-nine (19%) of the patients had no previously known diagnosis of COPD or ADHF and were evaluated as new diagnosis in the emergency department (newly diagnosed COPD, 4 patients; newly diagnosed ADHF, 25 patients). Physical examination findings were rale in 104 patients (67.6%), rhoncus in 92 patients (59.7%), and pretibial edema in 72 patients (46.7%).

The mean pCO<sub>2</sub> values were measured as 52.6 ± 13.2. The mean of ETCO<sub>2</sub> values measured simultaneously was 33.6 ± 10.1. There was a significant difference between the controlled pCO<sub>2</sub> values and ETCO<sub>2</sub> values at the first hour of NIMV treatment ( $Z: -10.640, p < 0.001$ ). However, when the correlation of pCO<sub>2</sub> and ETCO<sub>2</sub> values was examined, a moderate positive correlation was found ( $p < 0.001$ , Spearman  $r: 0.569$ ) (Table 1).

BiPAP mode was used in 107 (69.4%) patients, and CPAP mode was used in 47 (30.6%) patients in NIMV treatment. The mean systolic and diastolic blood pressure, respiratory

rate, and O<sub>2</sub> saturation levels were found to be significantly higher in CPAP mode usage group compared to the BiPAP mode usage group ( $p < 0.05$  for all markers). The mean pCO<sub>2</sub> and ETCO<sub>2</sub> values were found to be significantly higher in BiPAP mode usage group compared to the CPAP mode usage group ( $p < 0.001$ ) (Table 2). Scatter plot of ETCO<sub>2</sub> vs. pCO<sub>2</sub> levels is shown in Fig. 1. Bland-Altman plot illustrating the agreement between pCO<sub>2</sub> and ETCO<sub>2</sub> is shown in Fig. 2. It was observed that 59.1% (91 patients) of the patients included in the study were hospitalized according to their outcomes. 35.7% (55 patients) of the patients who were hospitalized were admitted to the intensive care unit. 25.4% (14 patients) of the patients who were taken into the intensive care unit resulted in endotracheal intubation.

## Discussion

ETCO<sub>2</sub> monitoring with capnometer is a method routinely used in intensive care units to evaluate the respiratory state and rate in patients receiving IMV treatment, and it is advantageous as it does not require intervention. While the use of invasive ETCO<sub>2</sub> has become widespread by clinicians who monitor patients in intensive care units, it has been thought that CO<sub>2</sub> in physiological dead spaces may reflect the CO<sub>2</sub> pressure in the blood. In the literature, the relationship between ETCO<sub>2</sub> and pCO<sub>2</sub> was first shown with 31 patients in 2000 [7]. In the literature, there are many studies evaluating the usability of ETCO<sub>2</sub> instead of pCO<sub>2</sub> in blood gas of patients undergoing IMV treatment [8, 9]. In 2017, in a study by Nassar et al., correlation coefficients between pCO<sub>2</sub> and ETCO<sub>2</sub> values were found to be between 0.6 and 0.9 [10].

A recent study investigating simultaneous ETCO<sub>2</sub> and pCO<sub>2</sub> measurements in blood gas in SIMV (synchronized intermittent mandatory ventilation), CPAP, and T-tube modes of IMV therapy demonstrated a moderate correlation between measured pCO<sub>2</sub> and ETCO<sub>2</sub> values.<sup>9</sup> In 2013, Singh et al. conducted a study in the neonatal intensive care unit and found a moderate correlation between the ETCO<sub>2</sub> level and pCO<sub>2</sub> level in patients treated with IMV [11]. In a study by Fujimoto et al. conducted in 2019 with 30 patients with respiratory failure and 18 patients with hypercapnia, a significant difference was found between ETCO<sub>2</sub> and pCO<sub>2</sub> [12].

In 2017, Schwarz et al. evaluated 3 measurements of pCO<sub>2</sub> values at the beginning, 30th minute and 60th minute with the Sidestream ETCO<sub>2</sub> measurements taken by cannula from the endotracheal tube of patients under IMV treatment in the Bland-Altman analysis. The mean pCO<sub>2</sub> values of all patients were found to be 42.4 ± 8.6 mmHg, and the mean of ETCO<sub>2</sub> values were 36 ± 7.5 mmHg. A significant correlation was found between the pCO<sub>2</sub> value and ETCO<sub>2</sub> value. In addition, it was emphasized that ETCO<sub>2</sub> was so inconsistent that it could not be used instead of pCO<sub>2</sub> [13]. In our study, the mean

**Table 1** Baseline characteristics of studied patients

Variables	Values (n = 154)
Age (years)	
Mean ± standard deviation	69.1 ± 12.2
Gender, n (%)	
Male	99 (64.2)
Female	55 (35.8)
Vital signs	
SBP, mm Hg	147.1 ± 37.3
DBP, mm Hg	85.3 ± 9.4
Heart rate, beats/min	102.3 ± 22.3
Body temperature	36.3 ± 0.2
Pre-diagnoses, n (%)	
Chronic obstructive pulmonary disease	85 (55.1)
Acute Decompensated Heart Failure	69 (44.9)
Clinical findings, n (%)	
Ralles	104 (67.6)
Ronchus	92 (59.7)
Pretibial edema	72 (46.7)
Previous history, n (%)	
Hypertension	25 (16.2)
Diabetes mellitus	48 (31.2)
Others	11 (7.1)
1st hour analysis	
pCO <sub>2</sub>	52.6 ± 13.2
ETCO <sub>2</sub>	33.6 ± 10.1
Outcome	
ICU admission	55 (35.7)

SBP systolic blood pressure, DBP diastolic blood pressure, PCO<sub>2</sub> partial pressure of carbon dioxide, ETCO<sub>2</sub> end-tidal carbon dioxide, ICU intensive care unit

**Table 2** Baseline characteristics of treatment options

Variables	CPAP ( <i>n</i> = 47 )	BIPAP ( <i>n</i> = 107)	<i>p</i>
Age (year)	71.66 ± 12.45	67.84 ± 11.84	0.061
Sex (male/female)	30/17	69/38	0.539
Mean systolic blood pressure	163.74 ± 41.75	140.18 ± 32.59	0.002
Mean diastolic blood pressure	97.45 ± 27.81	82.84 ± 16.68	0.002
Mean respiratory rate	30.23 ± 7.29	26.95 ± 7.61	0.014
Mean O <sub>2</sub> saturation (%)	88.43 ± 7.41	84.68 ± 10.12	< 0.001
Mean pulse/min	106.3 ± 21.71	100.11 ± 22.59	0.180
ETCO <sub>2</sub> levels	31.02 ± 9.28	34.81 ± 10.17	< 0.001
PaCO <sub>2</sub> levels	45.91 ± 10.52	55.54 ± 13.44	0.031

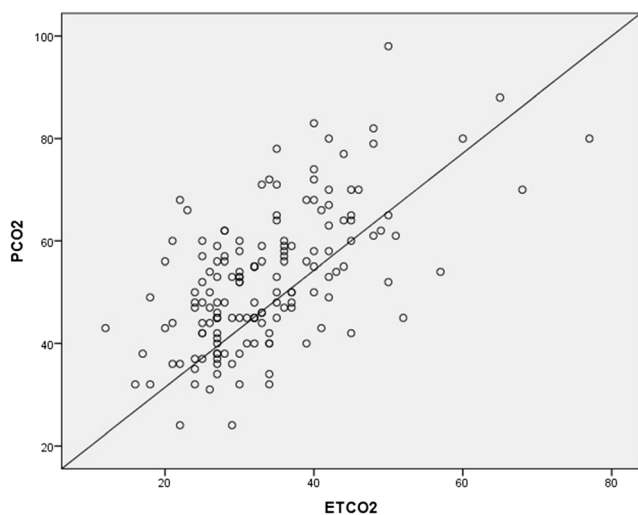
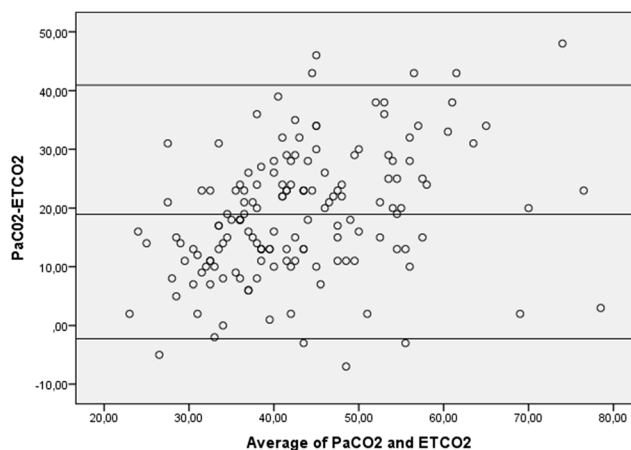
pCO<sub>2</sub> value was measured as 52.6 ± 13.2 mmHg in the blood gas sample taken at the first hour of NIMV treatment. The mean of ETCO<sub>2</sub> values measured simultaneously was 33.6 ± 10.1 mmHg. The reason why the pCO<sub>2</sub> value is higher than the literature may be that the study was conducted in the emergency department and the patients included in our study were evaluated at the first treatment without providing respiratory and metabolic stabilization.

In our study, similar to the literature, a significant difference was found between the ETCO<sub>2</sub> measurement and pCO<sub>2</sub> level (*Z*: -10.640, *p* < 0.001). For this reason, we think that it is not appropriate to use the ETCO<sub>2</sub> value numerically instead of pCO<sub>2</sub> level. Similar to the literature, the results of our study showed a moderate positive correlation between the pCO<sub>2</sub> value and ETCO<sub>2</sub> value (*p* < 0.001, *r*: 0.569). In the literature we reviewed, all the publications in which the ETCO<sub>2</sub> level was measured consisted of patients who were intubated or received IMV treatment in intensive care units. We think that the correlation coefficient between the ETCO<sub>2</sub> measurement and pCO<sub>2</sub> level was lower due to the fact that our patient group consisted of patients receiving NIMV treatment in the emergency department. The reason for this may be that the

non-functioning dead space area is more in NIMV applied patients, that there is an uncontrolled air leak outside the system, or that the device sensor used for mainstream measurement is fogged.

ETCO<sub>2</sub> is mostly recommended to be used in patients receiving IMV treatment under anesthesia and in the intensive care unit. Despite its use under these conditions, it has been found in studies in the literature that the use of ETCO<sub>2</sub> instead of pCO<sub>2</sub> value is limited [14]. In our study, in addition to the literature, we think that the ETCO<sub>2</sub> level can be used to evaluate the treatment monitoring in patients receiving NIMV treatment.

This study has certain limitations. The first of these was that the first hour observation time was quite short. Another limitation of our study was that although our study was designed prospectively, it was conducted with a relatively small number of patients. Furthermore, the single-center design of our study increased bias. Another limitation was the fact that the NIMV decision was not made by the same physicians may have influenced the results. In addition, the lack of evaluation of the time from the onset of symptoms to hospital admission may have affected the results.

**Fig. 1.** Scatter plot of ETCO<sub>2</sub> versus pCO<sub>2</sub>.**Fig. 2.** Bland–Altman plot illustrating the agreement between ETCO<sub>2</sub> and pCO<sub>2</sub>

## Conclusion

The ETCO<sub>2</sub> level was found to be different in our patients who received NIMV treatment, which could not be used instead of the pCO<sub>2</sub> level. However, the positive moderate correlation between the ETCO<sub>2</sub> level and pCO<sub>2</sub> level and the monitoring of NIMV treatment with the ETCO<sub>2</sub> level may have benefits such as predicting an unexpected increase in pCO<sub>2</sub> levels or making the decision to intubate patients faster.

**Author Contribution** HU, FS, and OFK conceptualized the idea. HU, FS, and CB reviewed the literature and collected the data with the help of FS and CB formulated the tables. HU, FS, CB, and OFK wrote the main draft of the manuscript. HU and FS supervised the project.

**Data Availability** Database is available upon request.

**Code Availability** Not applicable

## Declarations

**Ethics approval** This study was approved by the ethics committee of the Antalya Training and research hospital. All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional and/or national research committee and with the 1964 Helsinki Declaration and its later amendments or comparable ethical standards.

**Consent to Participate** Informed consent is applicable for this study.

**Consent for Publication** Not applicable

**Conflict of Interest** The authors declare that they have no conflict of interest.

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