



Original Article

The association between end-tidal carbon dioxide and arterial partial pressure of carbon dioxide after cardiopulmonary bypass pumping in cyanotic children

Behrang Nooralishahi^{1*}, Rozhin Faroughi², Hooman Naghashian², Ashkan Taghizadeh³, Mohammadjavad Mehrabian¹, Mehdi Dehghani Firoozabadi¹

¹Tehran Heart Center, Tehran University of Medical Sciences, Tehran, Iran

²School of Medicine, Tehran University of Medical Sciences, Tehran, Iran

³Children's Medical Center Hospital, Tehran University of Medical Sciences, Tehran, Iran

Article info

Article History:

Received: 1 May 2021

Accepted: 30 October 2021

published: 21 November 2021

Keywords:

End-Tidal Carbon Dioxide
Carbon Dioxide Pressure
Cyanosis
Children

Abstract

Introduction: Evidence suggests the high capability of non-invasive assessment of the End-tidal carbon dioxide (ETCO₂) in predicting changes in arterial carbon dioxide pressure (PCO₂) following major surgeries in children. We aimed to compare ETCO₂ values measured by capnography with mainstream device and ETCO₂ values assessed by arterial blood gas analysis before and after cardiopulmonary bypass pumping in cyanotic children.

Methods: This cross-sectional study was performed on 32 children aged less than 12 years with ASA II suffering cyanotic heart diseases and undergoing elective cardiopulmonary bypass pumping. Arterial blood sample was prepared through arterial line before and after pumping and arterial blood gas (ABG) was analyzed. Simultaneously, the value of ETCO₂ was measured by capnography with mainstream device.

Results: A significant direct relationship was found between the changes in ETCO₂ and arterial PCO₂ ($r=0.529$, $P=0.029$) postoperatively. According to significant linear association between postoperative change in ETCO₂ and arterial PCO₂, we revealed a new linear formula between the two indices: $\Delta\text{PCO}_2=0.89 \times \text{ETCO}_2-0.54$. The association between arterial PCO₂ and ETCO₂ remained significant adjusted for gender, age, and body weight.

Conclusion: the value of ETCO₂ can reliability estimate postoperative changes in arterial PCO₂ in cyanotic children undergoing cardiopulmonary bypass pumping.

Introduction

The partial pressure of arterial carbon dioxide (PaCo₂) is one of the effective and determining factors of blood pH, so its changes can cause many problems for critically ill patients. Because the likelihood of these changes during anesthesia is very high and it is not possible to continuously monitor PaCo₂ directly, so during anesthesia, End-tidal CO₂ (EtCO₂) monitoring is used to estimate PaCo₂, which is one of the standard monitoring methods during anesthesia and is often used as a non-invasive method for patients under anesthesia as well as in recovery and intensive care units.¹ Capnography as a tool for measuring EtCO₂ has become more common in recent years. Continuous measurement of EtCO₂ is one of the methods used in the operating room for evaluation of hemodynamic state during general anesthesia and also in intubated patients. But it can even be a non-invasive, fast and reliable method for predicting PaCo₂ in non-intubated patients.^{2,3} This measurement makes it possible to estimate arterial PaCo₂ without the need for arterial blood sampling. If there is a constant relationship

between PaCo₂ and EtCO₂, this method is reliable and there is no need for frequent arterial blood extraction. Using this method can provide necessary information about the patient's respiratory status quantitatively with high reliability. Therefore, in any type of monitor, the relationship between measured EtCO₂ and arterial blood PaCO₂ is an important and significant point. The difference between EtCO₂ and PaCO₂ represents the pulmonary dead space. Acute increase in dead space, such as in pulmonary embolism, widens this gap. This criterion also indicates the presence of shunting. In some studies, the association between EtCO₂ and PaCO₂ has been established, but the accuracy of this method is still debated.⁴⁻⁷ In a study comparing the relationship between EtCO₂ and PaCO₂ in the prone and supine positions, it was found that under normal circumstances, EtCO₂ is a useful tool for monitoring PaCO₂, but its accuracy may decrease when using different surgical techniques and in different positions.⁸

Overall, measuring and monitoring EtCO₂ is an important aspect of critical patient care. While EtCO₂



*Corresponding Author: Behrang Nooralishahi, Email: b-nooralishahi@sina.tums.ac.ir

© 2021 The Author(s). This is an open access article distributed under the terms of the Creative Commons Attribution License (<http://creativecommons.org/licenses/by/4.0>), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

monitoring was initially used by physicians to verify the location of endotracheal tubes and mechanical ventilation of patients in the emergency department, today, it is increasingly used for purposes such as monitoring the quality of cardiac resuscitation and assessing the causes of bronchospasm.⁹ In addition, studies have been performed to measure EtCO₂ to predict relative PaCO₂ or blood bicarbonate levels.¹⁰ Based on the gas sampling method, infrared monitoring of PaCO₂ is divided into two mainstream and sidestream methods. Sidestream capnography depends on the amount of gas removed from where the endotracheal tube is attached to the anesthesia machine. If the gas extraction rate is higher than the exhaust gas, the sample is mixed with fresh gas and the measurement accuracy will not be correct.¹¹ In the Mainstream type, because the capnometer is placed directly on the endotracheal tube, the reaction time is faster and the probability of error is low, but there is a possibility of the endotracheal tube closing due to the weight of the sensor.¹² Sidestream capnographs are highly dependent on the amount of gas that separates from the main breath system and reaches the monitor.¹³ In the present study, mainstream devices were used. In the present study, we aimed to compare EtCO₂ values measured by capnography with mainstream device and PaCO₂ values assessed by arterial blood gas analysis before and after cardiopulmonary bypass pump in cyanotic children. In fact, we attempted to determine the accuracy of measuring EtCO₂ by direct capnography and the effect of pumping on its measurement in cardiac cyanotic children.

Materials and Methods

This cross-sectional study was performed on 32 children aged less than 12 years with ASA II suffering cardiac cyanotic disorders and undergoing elective surgical correction at Pediatric Medical Center in Tehran between January and March 2019. All subjects were operated on by a single team and also had the same anesthesia procedure. In this regard, reluctance of the child's parents to cooperate in the project, recent colds, airway abnormalities, difficult intubation, high arterial pulmonary pressure, and emergency surgery were considered as the exclusion criteria. The study protocol was ethically approved by the ethical committee at Tehran University of Medical Sciences. The basis for patients' choice is heart surgery, which requires that an arterial blood sample be taken from the patient during the operation, and from this point of view, there is no moral problem in taking an arterial blood sample.

After standard monitoring and induction of anesthesia which were the same in all patients, after initial premedication with fentanyl (2 µg/kg), induction of anesthesia was scheduled with fentanyl (5 µg/kg) plus midazolam (0.05 mg/kg) and cisatracurium (0.15 mg/kg) and achieving sufficient depth of anesthesia, tracheal intubation was performed through the nose

and patients were connected to a ventilator. Immediately after intubation, the arterial line was inserted using 20 or 22 gauge arterial catheters from the radial artery of the patient's right hand. Maintenance of anesthesia was performed with isoflurane (1.5%) and oxygen (100%). The dose of fentanyl was repeated at intervals of one hour. Arterial blood sample was prepared through arterial line before and after cardiopulmonary bypass pumping (CPB) and arterial blood gas (ABG) was analyzed by the GEM Premier 3000 machine. Simultaneously, the value of EtCO₂ was measured by capnography with mainstream device. Such measurements were done both before and after CPB. Hemodynamic parameters including systolic and diastolic blood pressure, heart rate and arterial oxygen saturation were also assessed before and after cardiopulmonary bypass.

For statistical analysis, results were presented as mean ± standard deviation (SD) for quantitative variables and were summarized by frequency (percentage) for categorical variables. The values of study parameters before and after the procedure were compared using the Paired t test or Wilcoxon test. The agreement between the values of EtCO₂ assessed by capnography and PaCO₂ measured by arterial blood gas analysis was also examined by the Kappa agreement test. For the statistical analysis, the statistical software SPSS version 23.0 for windows (IBM, Armonk, New York) was used.

Results

Overall, 32 children (14 male and 18 female) were included into the study. The age of participants ranged from 6 days to 12 years that 31.2% were neonates, 43.8% were infants, and 25% aged higher than 2 years. The mean weight of the patients was also 7.98 ± 6.22kg. Due to non-parametric distribution of weight in the study subjects, the median of this index was determined as 8.0 kg (first quartile = 6.25, third quartile = 9.75).

Table 1 summarizes the value of study parameters before and after procedure. Of all study parameters assessed postoperatively, the increase in systolic and diastolic blood pressures, the increase in arterial oxygen saturation despite the decrease in FiO₂ were significant after procedure when compared to before. The results related to association of the postoperative change in ETCO₂ and changing other study parameters (Table 2) showed only a significant direct relationship between ETCO₂ and arterial PCO₂ ($r = 0.529$, $p = 0.029$). According to significant linear association between postoperative change in ETCO₂ and arterial PCO₂, we revealed the following linear formula between the two indices: $\Delta\text{PCO}_2 = 0.89 \times \text{ETCO}_2 - 0.54$. In this regard, for each unit of increase in ETCO₂, the amount of change in arterial PCO₂ will be equal to 0.35 units (Figure 1). Using the multivariable linear regression model with the presence of baseline variables including gender, age and body weight (Table 3), the association between arterial PCO₂ and ETCO₂ remained significant

Table 1. The value of study parameters before and after procedure

Parameter	Before procedure	After procedure	P value
Systolic blood pressure, mm Hg	76.71 ± 15.73	84.50 ± 12.00	0.031
Diastolic blood pressure, mm Hg	44.29 ± 10.89	50.35 ± 11.75	0.050
PEEP	3.00 ± 0.01	3.31 ± 0.70	0.096
Arterial pH	7.34 ± 0.08	7.36 ± 0.05	0.336
Arterial HCO ₃ , meq/L	22.87 ± 3.93	23.90 ± 3.92	0.190
Arterial PO ₂	60.19 ± 27.37	128.50 ± 89.32	0.008
Arterial PCO ₂	42.29 ± 7.28	41.65 ± 6.69	0.720
ETCO ₂	32.12 ± 5.02	32.00 ± 3.82	0.912
FiO ₂	76.75 ± 6.93	68.44 ± 10.18	0.019
Heart rate, /minute	127.88 ± 24.31	131.12 ± 20.26	0.354

ETCO₂: End-Tidal Carbon Dioxide; FiO₂: Fraction of Inspired Oxygen; HCO₃: Bicarbonate; PCO₂: Partial Pressure of Carbon Dioxide; PEEP: Positive End Expiratory Pressure;

Table 2. The association between the change in ETCO₂ parameter and change in other study indices

Parameter	R coefficient	P value
Systolic blood pressure, mm Hg	0.315	0.217
Diastolic blood pressure, mm Hg	0.146	0.577
PEEP	-0.410	0.114
Arterial pH	-0.239	0.373
Arterial HCO ₃ , meq/L	0.221	0.411
Arterial PO ₂	0.015	0.955
Arterial PCO ₂	0.529	0.029
FiO ₂	0.447	0.083
Heart rate, /minute	0.017	0.950

ETCO₂: End-Tidal Carbon Dioxide; FiO₂: Fraction of Inspired Oxygen; HCO₃: Bicarbonate; PCO₂: Partial Pressure of Carbon Dioxide; PEEP: Positive End Expiratory Pressure;

Table 3. The association between ETCO₂ and PaCO₂ values adjusted for baseline parameters

Item	Unstandardized Coefficients		T score	P value
	Beta	Standard Error		
Change in ETCO ₂	0.911	1.447	2.040	0.044
Male gender	2.641	3.527	0.749	0.468
Age	0.001	0.004	-0.081	0.937
Weight	0.033	1.728	0.046	0.964

ETCO₂: End-Tidal Carbon Dioxide

adjusted for baseline parameters.

Discussion

The change in both arterial PCO₂ and ETCO₂ parameters were not significant after cardiopulmonary bypass pumping in cyanotic children remained statistically insignificant and only hemodynamic indices including systolic and diastolic blood pressures as well as arterial oxygen saturation significantly changed that would be related to anesthesia and also procedural-related changes. However, we revealed a direct strong association between

the changes in arterial PCO₂ and ETCO₂ indices after surgery indicating high value of assessing the changes in ETCO₂ in predicting postoperative changes in arterial PCO₂. The assessment of arterial PCO₂ is very important to predicting critical postsurgical outcome especially hemodynamic instability that can be closely linked to patients' poor outcome. In this regard and aided by the found linear association between PCO₂ and ETCO₂ parameters, we will be able to easily and noninvasively predict abnormal arterial PCO₂ occurs after major surgeries.

Some other similar studies attempted to assess the link between arterial PCO₂ and ETCO₂ in both children and adults especially following major surgeries, however some believe that ETCO₂ might underestimate the value of PCO₂ in children under anesthesia. In a recent study by Arvind Chandrakantan et al,¹⁴ ETCO₂ underestimated venous PCO₂ values in neonates and infants under general anesthesia, but they introduced transcutaneous CO₂ (tcPCO₂) as a valuable parameters to closely approximate venous blood gas values. Wahba et al¹⁵ also showed that the changes in ETCO₂ did not accurately indicate the direction and extent of the change in arterial PCO₂. However, similar to our results, some studies emphasized the value of ETCO₂ in predicting the change in hemodynamic changes following major surgeries particularly cardiac surgeries. As similarly shown by Donati et al,¹⁶ ETCO₂ recordings were reliable, and can help in maintaining normocarbia during the short but unstable period associated with rewarming following cardiac surgery. Shibutani et al¹⁷ also revealed that decreases in ETCO₂ quantitatively reflected the decreases in CO₂ elimination. Burrows¹⁸ also indicated that PETCO₂ correlated closely with the arterial PCO₂ in the normal and acyanotic groups. Finally, Bissonnette et al¹⁹ showed that single breath ETCO₂ measurements from the proximal end of the endotracheal tube accurately estimate the arterial PCO₂ in infants and children.

As another important finding in the present study was

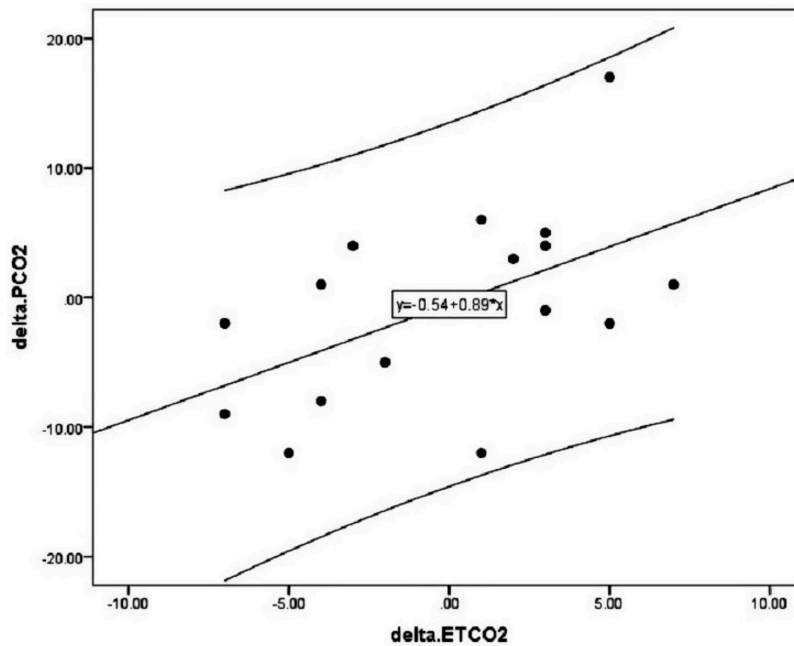


Figure 1. The linear association between ETCO₂ and arterial PCO₂

that the value of ETCO₂ measured was affected by gender, age or body weight of children. However it was shown that the value of this parameter might be affected by both parameters of patients' age as well as the site of measuring ETCO₂. As the first point, it was shown that ETCO₂ monitoring may not be reliable in older ages especially in adults, but it is contrarily valuable in children.¹⁹ Moreover, ETCO₂ measurements obtained from the proximal end of the endotracheal tube could not accurately predict arterial PCO₂ values in infants and children, however proximal measurements of ETCO₂ can accurately estimate arterial PCO₂.²⁰ Thus, measuring ETCO₂ obtained from proximal zone of tube can accurately estimate postoperative PCO₂ among children undergoing cardiac procedures.

Conclusion

Despite significant differences between ETCO₂ and arterial CO₂ in anesthetized cyanotic children before CPB, these measures become more relevant post CPB and after surgical correction; probably due to enhanced pulmonary blood circulation.

Acknowledgements

There are no acknowledgements to be mentioned in this study.

Competing Interest

There is no competing interest to be mentioned in this study.

Ethical Approval

This study was approved by the ethical committee of Tehran University of Medical sciences with approval I.D. IR.TUMS.CHMC. REC.1398.043

Funding

There are no funding or financial interests in this study

References

- Selby ST, Abramo T, Hobart-Porter N. An update on end-tidal CO₂ monitoring. *Pediatr Emerg Care*. 2018;34(12):888-892. doi:10.1097/pec.0000000000001682
- Øberg EB, Jørgensen BG, Berthelsen PG. End-tidal CO₂ in the diagnosis of fluid responsiveness - a systematic review. *Dan Med J*. 2019;66(9):A5560.
- Donald MJ, Paterson B. End tidal carbon dioxide monitoring in prehospital and retrieval medicine: a review. *Emerg Med J*. 2006;23(9):728-730. doi:10.1136/emj.2006.037184
- Richardson M, Moulton K, Rabb D, Kindopp S, Pische T, Yan C, et al. **Capnography for Monitoring End-Tidal CO₂ in Hospital and Pre-hospital Settings: A Health Technology Assessment**. Ottawa, ON: Canadian Agency for Drugs and Technologies in Health; 2016.
- Defilippis V, D'Antini D, Cinnella G, Dambrosio M, Schiraldi F, Procacci V. End-tidal arterial CO₂ partial pressure gradient in patients with severe hypercapnia undergoing noninvasive ventilation. *Open Access Emerg Med*. 2013;5:1-7. doi:10.2147/oaem.s43070
- Belpomme V, Ricard-Hibon A, Devoir C, Dileseigres S, Devaud ML, Chollet C, et al. Correlation of arterial PCO₂ and PETCO₂ in prehospital controlled ventilation. *Am J Emerg Med*. 2005;23(7):852-859. doi:10.1016/j.ajem.2005.04.011
- Casati A, Gallioli G, Scandroglio M, Passaretta R, Borghi B, Torri G. Accuracy of end-tidal carbon dioxide monitoring using the NBP-75 microstream capnometer. A study in intubated ventilated and spontaneously breathing nonintubated patients. *Eur J Anaesthesiol*. 2000;17(10):622-626. doi:10.1046/j.1365-2346.2000.00731.x
- Magi E, Multari G, Recine C, Barberio O, Becattini G, Tellini A. [Difference between arterial and end-tidal carbon dioxide tension during surgery of lumbar herniated disk in general anesthesia]. *Minerva Anesthesiol*. 1994;60(7-8):381-386. [Italian].
- Sulemanji DS, Marchese A, Wysocki M, Kacmarek RM. Adaptive support ventilation with and without end-tidal CO₂ closed loop control versus conventional ventilation. *Intensive Care Med*. 2013;39(4):703-710. doi:10.1007/s00134-012-2742-6

10. Hochwald O, Borenstein-Levin L, Dinur G, Jubran H, Ben-David S, Kugelman A. Continuous noninvasive carbon dioxide monitoring in neonates: from theory to standard of care. **Pediatrics**. 2019;144(1). doi:10.1542/peds.2018-3640
11. Williams E, Dassios T, Greenough A. Assessment of sidestream end-tidal capnography in ventilated infants on the neonatal unit. **Pediatr Pulmonol**. 2020;55(6):1468-1473. doi:10.1002/ppul.24738
12. Kugelman A, Zeiger-Aginsky D, Bader D, Shoris I, Riskin A. A novel method of distal end-tidal CO₂ capnography in intubated infants: comparison with arterial CO₂ and with proximal mainstream end-tidal CO₂. **Pediatrics**. 2008;122(6):e1219-1224. doi:10.1542/peds.2008-1300
13. Bhat YR, Abhishek N. Mainstream end-tidal carbon dioxide monitoring in ventilated neonates. **Singapore Med J**. 2008;49(3):199-203.
14. Chandrakantan A, Jasiewicz R, Reinsel RA, Khmara K, Mintzer J, DeCristofaro JD, et al. Transcutaneous CO₂ versus end-tidal CO₂ in neonates and infants undergoing surgery: a prospective study. **Med Devices (Auckl)**. 2019;12:165-172. doi:10.2147/meder.s198707
15. Wahba RW, Tessler MJ. Misleading end-tidal CO₂ tensions. **Can J Anaesth**. 1996;43(8):862-866. doi:10.1007/bf03013040
16. Donati F, Maille JG, Blain R, Boulanger M, Sahab P. End-tidal carbon dioxide tension and temperature changes after coronary artery bypass surgery. **Can Anaesth Soc J**. 1985;32(3 Pt 1):272-277. doi:10.1007/bf03015142
17. Shibutani K, Muraoka M, Shirasaki S, Kubal K, Sanchala VT, Gupte P. Do changes in end-tidal PCO₂ quantitatively reflect changes in cardiac output? **Anesth Analg**. 1994;79(5):829-833. doi:10.1213/00000539-199411000-00002
18. Burrows FA. Physiologic dead space, venous admixture, and the arterial to end-tidal carbon dioxide difference in infants and children undergoing cardiac surgery. **Anesthesiology**. 1989;70(2):219-225. doi:10.1097/00000542-198902000-00007
19. Bissonnette B, Lerman J. Single breath end-tidal CO₂ estimates of arterial PCO₂ in infants and children. **Can J Anaesth**. 1989;36(2):110-112. doi:10.1007/bf03011429
20. Badgwell JM, McLeod ME, Lerman J, Creighton RE. End-tidal PCO₂ measurements sampled at the distal and proximal ends of the endotracheal tube in infants and children. **Anesth Analg**. 1987;66(10):959-964.