Should capnography be used as a guide for choosing a ventilation strategy in circulatory shock caused by severe hypothermia? Observational case-series study

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Abstract

Background: Severe accidental hypothermia can cause circulatory disturbances ranging from cardiac arrhythmias through circulatory shock to cardiac arrest. Severity of shock, pulmonary hypoperfusion and ventilation-perfusion mismatch are reflected by a discrepancy between measurements of CO₂ levels in end-tidal air (EtCO₂) and partial CO₂ pressure in arterial blood (PaCO₂). This disparity can pose a problem in the choice of an optimal ventilation strategy for accidental hypothermia victims, particularly in the prehospital period. We hypothesized that in severely hypothermic patients capnometry should not be used as a reliable guide to choose optimal ventilatory parameters.

Methods: We undertook a pilot, observational case-series study, in which we included all consecutive patients admitted to the Severe Hypothermia Treatment Centre in Cracow, Poland for VA-ECMO in stage III hypothermia and with signs of circulatory shock. We performed serial measurements of arterial blood gases and EtCO₂, core temperature, and calculated a PaCO₂/EtCO₂ quotient.

Results: The study population consisted of 13 consecutive patients (10 males, three females, median 60 years old). The core temperature measured in esophagus was 20.7–29.0 °C, median 25.7 °C. In extreme cases we have observed a Pa-EtCO₂ gradient of 35–36 mmHg. Median PaCO₂/EtCO₂ quotient was 2.15.

Discussion and Conclusion: Severe hypothermia seems to present an example of extremely large Pa-EtCO₂ gradient. EtCO₂ monitoring does not seem to be a reliable guide to ventilation parameters in severe hypothermia.

Keywords: Accidental hypothermia, Pulmonary ventilation, Capnography

Background

While end-tidal carbon dioxide (EtCO₂) monitoring is one of the objective standards set in the Intensive Care Society guidelines [1, 2] and is of particular use for verification of endotracheal tube placement [1], it does not seem to be a reliable guide to ventilation in profound shock states.

It was noted that abnormal EtCO₂ measurements on initial emergency department presentation correlate with bad prognosis both in adults and children [1]. Since cerebral blood vessels are sensitive to changes in partial pressure of CO₂ (PaCO₂), and hypocapnia induced by hyperventilation can lead to vasoconstriction and as a consequence worsening of secondary brain injury, it is advocated that ventilation parameters should be aimed at achieving “normocapnia”.

Pulmonary hypoperfusion and pulmonary ventilation – perfusion mismatch seem to play an important role among many factors determining extremely large Pa-EtCO₂
gradient observed in severe hypothermia victims. This discrepancy is further aggravated by a drop in blood temperature itself.

There is no published data on Pa-EtCO₂ gradient and reliability of EtCO₂ measurement in severe hypothermia. Based on our experience we hypothesize that in severely hypothermic patients capnometry should not be used as a reliable guide to choose optimal ventilatory parameters.

Methods
We carried out a retrospective observational case-series study. All patients admitted to the Severe Hypothermia Treatment Centre (SHTC-Cracow, Poland) with stage III hypothermia, that still had a circulation and features of shock, were enrolled [3]. All data analyzed was collected on admission. The measurement of the central temperature (Tc) was taken in the lower third esophagus, using single-use Smiths Medical 12Fr probes, coupled with a SpaceLab cardiomonitor. The value of EtCO₂ was estimated from the main stream with the use of a capnometer from the SpaceLab monitoring system.

Blood tests were assayed by routine automated laboratory techniques (Radiometer Copenhagen model ABL80). Blood gas analyses according to alpha-stat (blood gases measured at 37 °C) were performed in the central hospital laboratory, certified with a program by RIQAS (Randox Quality Assessment Scheme, UK). Simple plotting of PaCO₂ against EtCO₂ was performed. The study was approved by the Local Ethical Committee of the John Paul II Hospital in Cracow.

Results
The study population consisted of 13 patients (ten males, three females, median 60 age years). The core temperature measured in the oesophagus was 20.7–29 °C, median 25.7 °C. PaCO₂ values varied between 17 to 53.1 mmHg (median 25.5 mmHg), and EtCO₂ from 12 to 19 mmHg (median 17 mmHg).

In extreme cases we have observed a Pa-EtCO₂ gradient of 35–36 mmHg. Median PaCO₂/EtCO₂ quotient was 2.15 (blood gases measured at 37 °C). Figure 1 summarizes the parameters of patients in stage III hypothermia who still had a circulation.

Discussion
General guidelines for ventilatory support do not cover special population of severe hypothermia patients (Swiss Stage III and IV, Table 1) [4, 5]. Some experts recommend that the respiratory rate of mechanical ventilation should be lower [6], others prefer the ventilation rate to be normal [7]. The Wilderness Medical Society guidelines state that in intubated patients, without the possibility of EtCO₂ control, it is recommended to decrease the respiratory rate by half in relation to the value in normothermia. At the same time, in patients in which capnometry is available, it is recommended to maintain EtCO₂ in normal range [8]. In the latest review of the current knowledge about hypothermia, there is an emphasis on the maintenance of normocapnia in order to prevent arrhythmia related to hyper- or hypoventilation [9]. Maintenance of normoventillation and normocapnia in patients in hypothermia is not an easy task. In mild, therapeutic hypothermia, such as in the ICU, normocapnia is achieved and maintained in only about 55% [10]. Unfortunately, even the EtCO₂ does not solve the problem. It has been ascertained that in mild, therapeutic hypothermia (36 – 32 °C), the gradient between PaCO₂ and EtCO₂ may increase 2.5-fold and be as high as 18.7 mmHg [11].

During the prehospital period, the only practical way to assess PaCO₂ is by indirect measurement of end-tidal CO₂ (EtCO₂). In normothermia, the Pa- EtCO₂ gradient is usually 4–6 mmHg, so the EtCO₂ values may be easily

![Fig. 1 Pa-EtCO₂ gradient parameters of patients in stage III hypothermia](image-url)
treated as a baseline for establishing parameters of normoventilation.

However, in our opinion, in significantly decreased core temperatures this is an unreliable guide to ventilation because of the profound metabolic, circulatory and respiratory disturbances, especially within the ventilation - perfusion mismatch, which accompanies severe hypothermia. This has been confirmed by the results obtained in our patients.

The impact of hypothermia on the partial pressure of CO$_2$ in arterial blood and acid-base balance has been known for years, especially in cardiac surgery. Two methods of interpreting blood gas results have been proposed. One is the pH-stat strategy in which ventilation is adjusted to maintain PaCO$_2$ at 40 mmHg at the patient's current body temperature. Such a correction is difficult to calculate, and is hardly ever used in the adult population [12]. Most centres, including ours, do not use this method, and prefer to use the alpha-stat strategy instead. In this approach, ventilation is adjusted to maintain PaCO$_2$ at 40 mmHg at 37 °C, meaning that PaCO$_2$ will be $<$40 mmHg in hypothermia. The alpha-stat strategy is recommended nowadays for patients with hypothermia [5, 9, 13].

The difference between EtCO$_2$ and PaCO$_2$ (blood gases measured at 37 °C) are generally consistent with the observations conducted by Sitzwohl et al., although this study only looked at patients with mild intra-operative hypothermia (32–36 °C) [11]. The small number of observations obtained so far does not allow us to carry out statistical analysis. Nevertheless, there is a definite trend of increasing Pa-EtCO$_2$ gradient above 1 as the core temperature falls, although the correlation is not linear, as presented on the Fig. 1. In our opinion, such a large gradient is a result of both increased CO$_2$ solubility as temperature decreases and the increase in ventilation-perfusion disorders, including low cardiac output. Interestingly, Sitzwohl et al. found that the mode of ventilation did not have a significant effect on the Pa-EtCO$_2$ gradient. Unfortunately, no data concerning the actual parameters of ventilation in the pre-hospital phase has been recorded in our patients, though we hope to obtain this data in a future study.

At present, it is unclear whether the recommendation to mildly hypoventilate patients with hypothermia has clinical significance [8]. It is worth noting that based on local guidelines, SHTC coordinators routinely recommend the use of normoventilation during transport as part of a lung-protective ventilation strategy ($V_t$ = 6–7 ml/kg of ideal body weight, PEEP 5 mmHg and ventilatory rate = 10/min), and it should be regarded as the optimal ventilation strategy in adults. We also advocate for avoidance of manual bag-valve ventilation due to its tendency to hyperventilate.

EtCO$_2$ monitoring in hypothermic victims should be used not only as proof of correct positioning of the endotracheal tube, but also, as a sign of preserved pulmonary flow, while circulatory instability is reflected by a fall of EtCO$_2$. Critically low values suggest cardiac arrest. Such observation may be particularly valuable in the event of cardiac arrest with pulseless electrical activity (PEA), whose confirmation can be very difficult in the pre-hospital phase, where no ultrasound and invasive blood pressure measurement are available.

Conclusions
A very high Pa-EtCO$_2$ gradient is found in patients with severe hypothermia.

In patients with severe hypothermia, the EtCO$_2$ values should not be used as the main criterion for the selection of ventilatory parameters.

The optimal ventilatory technique in patients with hypothermia should be mechanical lung protective ventilation.

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Availability of data and materials
Please contact author for data requests.

Authors’ contributions
TD designed the study, supervised data collection, took part in manuscript preparation, contributed substantially to the revision of the manuscript; takes responsibility for the paper as a whole. SK supervised data collection, took part in manuscript preparation, contributed substantially to the revision of the manuscript. AJ, PP, MZ contributed substantially to the revision of the manuscript. TS, RG, JP, JKK, RD provided advice on study design, contributed to the revision of the manuscript. All authors read and approved the final manuscript.

Competing interests
The authors declare that they have no competing interests.

Consent for publication
Written informed consent was obtained from the patients for publication of their individual details in this manuscript. The consent form is held by the authors and is available for review by the Editor-in-Chief.

Table 1 Swiss Stage of Hypothermia

| Hypothermia stage | Clinical findings                  | Core temperature (if available) |
|-------------------|-----------------------------------|---------------------------------|
| I (mild)          | Conscious; shivering              | 35–32 °C                        |
| II (moderate)     | Impaired consciousness; may or may not be shivering | <32–28 °C                      |
| III (severe)      | Unconscious; vital signs present   | <28 °C                          |
| IV                | Vital signs absent                | Variable                        |
Ethics approval and consent to participate
The prospective observational case-series study was approved by the Local Ethical Committee of the John Paul II Hospital in Cracow.

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References
1. The Intensive Care Society Guidelines Capnography in the Critically Ill Available at http://www.ics.ac.uk/ICS/guidelines-and-standards.aspx. Accessed 21 Jan 2017.
2. Donald MJ, Paterson B. End tidal carbon dioxide monitoring in prehospital and retrieval medicine: a review. Emerg Med J. 2006;23:728–30.
3. Darocha T, Kosirski S, Jarosz A, et al. Severe Accidental Hypothermia Center. Eur J Emerg Med. 2015;22:288–91.
4. Durer B, Brugger H, Syme D. The medical on-site treatment of hypothermia: ICA-MEDCOM recommendation. High Alt Med Biol. 2003;4:99–103.
5. Brown DJA. Hypothermia. In: Tintinalli JE, editor. Emergency Medicine. 8th ed. New York: McGraw Hill; 1997.
6. Socialstyrelsen. Hypothermia – cold induced injuries. Stockholm: National Board of Health and Welfare; 1997.
7. Lloyd EL. Accidental hypothermia. Resuscitation. 1996;32:111–2.
8. Zafren K, Gusbrecht GG, Danzl DF, et al. Wilderness-Medical Society practice guidelines for the out-of-hospital evaluation and treatment of accidental hypothermia: 2014 update. Wilderness Environ Med. 2014;25(4 Suppl):S66–85.
9. Paal P, Gordon L, Strapazzon G, et al. Accidental hypothermia—an update. Scand J Trauma Resusc Emerg Med. 2016;24:111. doi:10.1186/s13049-016-0303-7.
10. Falkenbach P, Kämäräinen A, Mäkelä A, et al. The arterial to end-tidal carbon dioxide gradient increases with uncorrected but not with temperature-corrected PaCO2 determination during mild to moderate hypothermia. Anesth Analg. 1998;86:1131–6.
11. Abdul Aziz KA, Meduoye A. Is pH-stat or alpha-stat the best technique to follow in patients undergoing deep hypothermic circulatory arrest? Interact Cardiovasc Thorac Surg. 2010;10:271–82.
12. Kempainen RR, Brunette DD. The evaluation and management of accidental hypothermia. Respir Care. 2004;49:192–205.