



ORIGINAL ARTICLE

## Effects of the transcutaneous electrode temperature on the accuracy of transcutaneous carbon dioxide tension

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### Abstract

**Aim.** The harmful effect of hypocapnia on the neonatal brain emphasizes the importance of monitoring arterial carbon dioxide tension (PaCO<sub>2</sub>). Transcutaneous monitoring of carbon dioxide (tcPCO<sub>2</sub>) reduces the need for arterial blood sampling. Drawbacks are high electrode temperature causing risks of skin burning. The aim was to determine the accuracy and precision of tcPCO<sub>2</sub> at reduced electrode temperature. **Methods.** Forty newborns (GA 24.9–41.7) were included. Two tc-monitors were applied (TCM4, Radiometer, Copenhagen). Arterial blood gas sampling and monitoring of tcPCO<sub>2</sub>-level at different electrode temperatures was done simultaneously (39°C, 40°C, 41°C, 42°C, 44°C). Difference of PaCO<sub>2</sub> – tcPCO<sub>2</sub> was expressed as a percentage of the mean. **Results.** Mean PaCO<sub>2</sub> was 5.8 kPa [3.2; 7.9]. Bias (PaCO<sub>2</sub> – tcPCO<sub>2</sub>) increased from 5% at 44°C to 17% at 39°C, but did not differ significantly between 41°C and 40°C. The precision of the tcPCO<sub>2</sub> at each temperature ranged from +7–10%. After correction for the temperature-dependent overreading, we found increasing PaCO<sub>2</sub> – tcPCO<sub>2</sub> difference with increasing PaCO<sub>2</sub>, approx. 2% pr. kPa increase of CO<sub>2</sub>. Only mild transient erythema was observed. **Conclusion.** A lower electrode temperature in tcPCO<sub>2</sub>-monitoring increases systematic overreading of the tc-electrode. However, in very preterm babies, monitoring at 40°C or 41°C is possible provided a bias correction of 12–15% is applied.

**Key Words:** Infant: newborn, infant: premature, carbon dioxide, blood gas monitoring, transcutaneous, pulmonary ventilation

### Introduction

The harmful effects of hyperventilation and hypocapnia on the neonatal brain emphasize the importance of a stable arterial carbon dioxide tension (PaCO<sub>2</sub>) especially in mechanically ventilated newborns [1]. Transcutaneous monitoring (TCM) allows continuous monitoring of the transcutaneous carbon dioxide (tcPCO<sub>2</sub>) – a surrogate measure of the PaCO<sub>2</sub>. Hence, the need of arterial blood sampling is reduced.

The transcutaneous electrode conventionally is heated to 44°C to arterialize the capillary bed. The correlation between tcPCO<sub>2</sub> and PaCO<sub>2</sub> depends on the temperature in the capillary bed, the blood flow to the skin, and the local metabolic production of CO<sub>2</sub>. Drawbacks of the high electrode temperature are risk of skin burns [2,3]. The correlation between time and surface temperature in the causation of

cutaneous burns has been known for a long time [4]. Early studies in preterm infants showed a direct correlation between the duration of electrode application and persistence of erythema; while the application time is shortened [2]. Besides disturbance of the neonate, relocation of the electrode gives unstable values during stabilization.

In the early era of transcutaneous monitoring, several studies were performed, improving the monitors in use today [5,6]. This also includes studies with monitoring at different electrode temperatures [7–9]. Because of the fragile skin of preterm infants a lower electrode temperature would be appreciated. This could reduce the risk of skin burns and prolong the application time. However, a recent study in adults examined reduced electrode temperature using the TCM4, Radiometer Medical ApS, Denmark [10]. Due to bias, the authors concluded that the

electrode should be heated to at least 43°C to measure reliable tcPCO<sub>2</sub> [10].

The aim of this study was to determine the accuracy and precision of tcPCO<sub>2</sub> in preterm infants using TCM4, Radiometer Medical ApS, Denmark, with reduced electrode temperatures. Our assumption was that electrode temperatures of 39–41°C would be less damaging to the skin, since these temperatures are reached during systemic fever.

**Patients and methods**

The study was conducted in the NICU at the Copenhagen University Hospital, Rigshospitalet. The local ethics committee approved the study (J. nr. (KF) 01 283061). Parental consent was obtained in all cases.

This paper consists of two studies. In the first study, pCO<sub>2</sub> precision was examined at five different electrode temperatures (39, 40, 41, 42 and 44°C). Recruitment was from January to April 2006. Inclusion criteria were clinical indication of transcutaneous (TC) monitoring and presence of an umbilical or radial arterial catheter. A total of 20 newborns irrespective of gestational age (GA) were included (= GA mixed group).

In the second study, based on results from study one, comparisons of pCO<sub>2</sub> precision at selected electrode temperatures were done (40, 41 and 44°C). Recruitment was from November 2006 to June 2008. Inclusions criteria was GA below 34 weeks or birth weight below 1500 g, clinical indication of TC monitoring and presence of an umbilical or radial arterial catheter. A total of 20 preterm infants were included (= Preterm group). Inclusion and examination was within the neonatal period (first 28 days of life).

All newborns were monitored using two TC monitors simultaneously (TCM 4 with combined CO<sub>2</sub>/O<sub>2</sub> electrode E5280, Radiometer Medical ApS, Denmark). Because of higher values of pCO<sub>2</sub> when elevating skin temperature (anaerobic factor 4.5%/°C), the CO<sub>2</sub> production of epidermal cells ('metabolic' constant) adding to the pCO<sub>2</sub> measured transcutaneously, the monitor has a correction

algorithm integrated [11]. We used standard recommendations with the anaerobic factor (Severinghaus constant) turned on, and with a metabolic constant set at 0.5 kPa.

Using two TCM4 simultaneously in the first study, allowed one electrode to remain unchanged at 44°C while the temperature of the other electrode was increased every half hour (39, 40, 41, 42 and 44°C) (Figure 1).

Using two TCM4 simultaneously in the second study allowed a cross over design to investigate the impact of preheating of the skin on pCO<sub>2</sub> precision (Figure 1).

The TC electrodes were fixed to the skin of the trunk using a sticky fixation ring (E5260/E5280: Fixation ring 904 – 891; 30 mm; Radiometer Medical ApS, Denmark). The localization was not changed during the study. The electrode housing was 15 mm in diameter. After a few droplets of an electrolyte solution enhancing contact between electrode and skin, the electrode was fixed in the fixation ring (Electrolyte solution; Radiometer Medical ApS, Denmark).

The 'gold standard' to which the TC values were compared was the arterial pCO<sub>2</sub> (PaCO<sub>2</sub>).

All arterial blood samples were taken from the indwelling catheter. Blood was sampled in heparinized capillary tubes (100 microliter). Immediately after sampling the tube was turned slowly 20 times. Within 5 min the blood gases were analysed using Blood Gas Analyzer ABL 735 (Radiometer Medical ApS, Denmark). In the second study, the first blood sample was analysed in duplicate to estimate test-retest variation of arterial pCO<sub>2</sub>.

Arterial blood gas sampling and readings of the tcPCO<sub>2</sub>-level was done simultaneously. After changing the electrode temperature, a minimum of 30 min. was allowed for temperature equilibration and for stabilizing tcPCO<sub>2</sub> before blood sampling. Before measuring at a new electrode temperature, the sensor was calibrated. The monitor automatically performs this process using a built in calibration module and gas cylinder. Calibration of the sensor is recommended as a routine procedure in TC monitoring using TCM4. The sensor was re-membraned

Study 1

TCM4 monitoring	30 minutes	30 minutes	30 minutes	30 minutes	30 minutes
Electrode temperature	44°C	44°C	44°C	44°C	44°C
Electrode temperature	39°C	40°C	41°C	42°C	44°C

Study 2

TCM4 monitoring	30 minutes	30 minutes	30 minutes
Electrode temperature	40°C	41°C	44°C
Electrode temperature	44°C	44°C	40°C

Figure 1. The sequence of the transcutaneous electrode temperature changes in the two studies.

when needed (E5260/E5280: Membrane 904 – 892; Radiometer Medical ApS, Denmark).

Statistical analysis was done using SPSS version 12.0 (Chicago). To estimate the repeatability of the blood gas analysis we calculated the standard deviation of the differences between the pairs of blood gas measurements. Univariate ANOVA was used to test the influence of electrode temperature on accuracy ( $\Delta$ tcPCO<sub>2</sub> – PaCO<sub>2</sub>). 95% limits of agreement were calculated as described by Bland and Altman [12]. As the difference between transcutaneous and arterial pCO<sub>2</sub> increased with increasing pCO<sub>2</sub>, the difference between transcutaneous and arterial values was expressed as a percentage of the mean ( $100 \times (\text{diff. tcPCO}_2 - \text{PaCO}_2 / \text{mean tcPCO}_2 - \text{PaCO}_2)$ ). The Kruskal-Wallis test was used for comparisons.

## Results

The median gestational age in the GA mixed group was 31.1 weeks [24.9; 41.7] and median birth weight 1822 g [735; 3800 g] (Table I).

Diagnoses were manifold: Prematurity ( $n = 11$ ), asphyxia ( $n = 1$ ), meconium aspiration syndrome ( $n = 1$ ), necrotizing enterocolitis ( $n = 1$ ), congenital heart disease ( $n = 3$ ), oesophageal atresia ( $n = 1$ ), extracorporeal membrane oxygenation ( $n = 1$ ) and no diagnosis specified ( $n = 2$ ).

In the preterm group, the median gestational age was 28.9 weeks [25.6; 32.0] and median birth weight 1006 g [590 g; 1900 g] (Table I). All infants ( $n = 20$ ) were preterm, however other diagnoses included pneumothorax ( $n = 4$ ), rhesus immunization ( $n = 1$ ), persistent fetal circulation ( $n = 1$ ) and lung bleeding ( $n = 1$ ).

No infants were excluded due to instability, since this was not a part of the protocol. However, if there were any events that potentially could influence results, it was remarked. In the GA mixed group, two infants had remarks: ‘restless’ ( $n = 1$ ) and ‘apnoea, bradycardia and desaturation of prematurity’ ( $n = 1$ ), while in the preterm group four infants had remarks: ‘restless’ ( $n = 2$ ), ‘increasing oxygen demand’ ( $n = 2$ ), ‘apnoea, bradycardia and desaturation of prematurity’ ( $n = 4$ ).

Arterial pCO<sub>2</sub> ( $n = 220$ ) was normally distributed (range 3.2–7.9; mean 5.8 kPa  $\pm$  1.0 kPa) and

was not associated with electrode temperature ( $p = 0.9$ ). The repeatability of arterial pCO<sub>2</sub> measurements ( $n = 19$ ) was  $0.07 \pm 0.15$  kPa.

By defining a maximum of 1 kPa bias between tcPCO<sub>2</sub> and PaCO<sub>2</sub>, we examined how many of the paired samples met these qualifications. When using an electrode temperature of 39°C 45% had a bias below 1 kPa (9/20 measurements), at 40°C 73% (44/60 measurements), at 41°C 68% (27/40 measurements), at 42°C 90% (18/20 measurements) and at 44°C 94% had a bias below 1 kPa (75/80 measurements).

The mean PaCO<sub>2</sub> – tcPCO<sub>2</sub> difference (bias) increased from 5% at 44°C to 17% at 39°C, but did not differ significantly between 41°C and 40°C (median 14.8% vs. 11.8%) (Figures 2 & 3). The precision of the tcPCO<sub>2</sub> at each temperature ranged from 7–10%. After correction for the temperature-dependent overreading, we found increasing PaCO<sub>2</sub> – tcPCO<sub>2</sub> difference with increasing PaCO<sub>2</sub>, an approx. 2% pr. kPa increase of pCO<sub>2</sub>.

All outliers were above the 95% confidence interval. These outliers occurred primarily at low electrode temperature. Sequence of electrode temperatures, including preheating of the skin to 44°C, had no influence on precision. We did not find any significant electrode drift after each session by control measurement using the calibration gas. No skin lesions apart from mild transient erythema were observed.

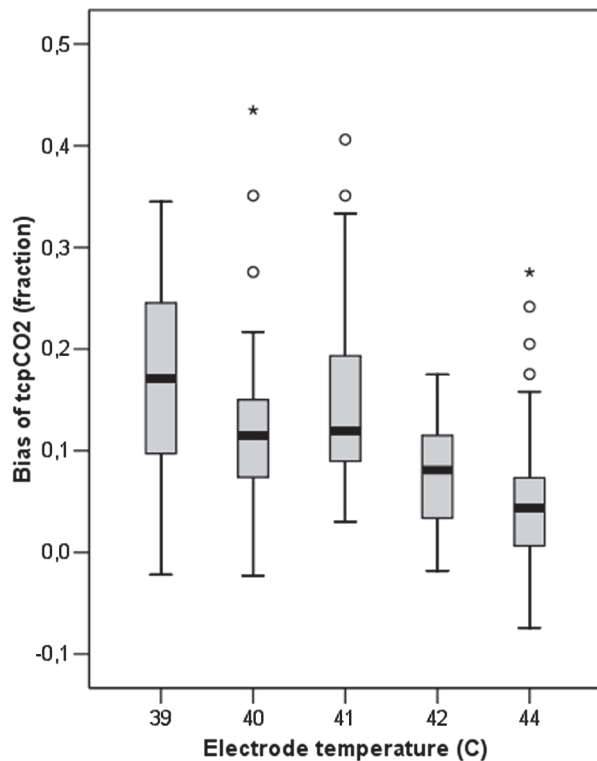


Figure 2. The difference between transcutaneous pCO<sub>2</sub> and arterial pCO<sub>2</sub> increases at lower electrode temperatures. The median difference is +12% at 40–41°C. Outliers also occur at high electrode temperature.

Table I. Group characteristics.

	GA mixed group (N = 20)	Preterm group (N = 20)
Gestational age (weeks)	31.1 [24.9; 41.7]	28.9 [25.6; 32.0]
BW (grams)	1822 [735; 3800]	1006 [590; 1900]
Nasal CPAP/Mechanical ventilation	50%/35%	55%/45%
No oxygen demand	55%	30%
Oxygen supplement (% O <sub>2</sub> )	21% [21%; 55%]	28% [21%; 70%]
Dopamine	0%	10%

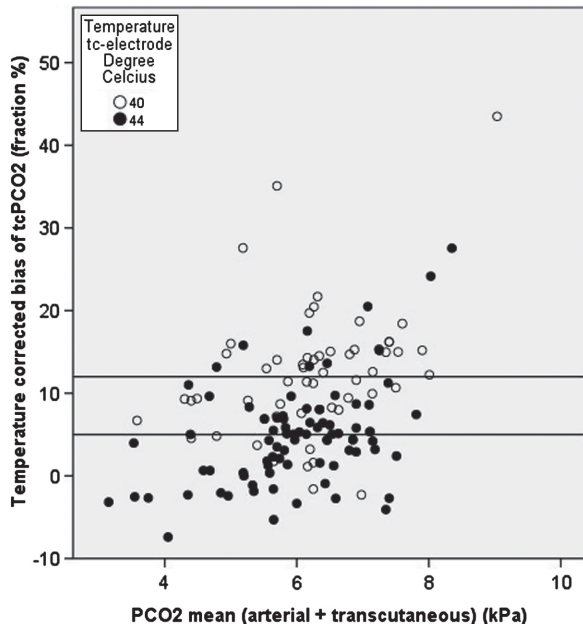


Figure 3. This figure shows the mean pCO<sub>2</sub> (kPa) versus the temperature corrected bias of TcPCO<sub>2</sub> (fraction, %). The horizontal lines show the mean bias of TcPCO<sub>2</sub> at 40°C (12%) and 44°C (5%). Correction with 12–15% when using 40°C will approximate measurements to the arterial pCO<sub>2</sub>.

## Discussion

The aim of this study was to investigate how reduction of the electrode temperature influences accuracy and precision in tcPCO<sub>2</sub> measurements in newborn infants. Lower electrode temperature reduces the risk of skin burning and allows longer periods of continued monitoring [13].

The bias of tcPCO<sub>2</sub> using the normal electrode temperature of 44°C was approximately 5%. We had set the metabolic constant at 0.53 kPa (4 mmHg), which appears to be too low for our population.

Lower electrode temperatures were associated with higher bias, but at 41°C the bias was still typically less than 1 kPa. Precision was unchanged. We therefore conclude that it is reasonable to monitor tcPCO<sub>2</sub> with a temperature of 41°C, particularly if the bias of 12–15% is accounted for. However, changing the electrode temperature requires an increased clinical awareness. With measurements close to the normal PaCO<sub>2</sub> range an error of 1 kPa will have no great consequences, however readings in the subnormal range of PaCO<sub>2</sub> can have a significant impact on cerebral perfusion. On our ward, we recommend an arterial blood gas if there is any doubt of the actual level of PaCO<sub>2</sub> and in particular if the tcPCO<sub>2</sub> is out of the normal range. At 41°C, the tcPCO<sub>2</sub> cannot be relied upon, because the capillary bed will not be sufficiently arterialized. The best procedure would be to deactivate the pO<sub>2</sub> readout of the TCM4. In most situations, oxygen treatment of newborns can be sufficiently monitored by pulseoximetry.

Our results agree reasonably with previous studies. A study in neonates found a good correlation between tcPCO<sub>2</sub> and arterial CO<sub>2</sub> at 42°C and even 37°C [7]. In adults, a good correlation was found between transcutaneous and capillary pCO<sub>2</sub> at 37, 39, 41, 43 and 45°C, provided the skin was preheated to 45°C [14]. The correlation coefficient, however, ignores bias. Fanconi and colleagues reduced the electrode temperature to 42°C and had reliable results, but only after topical metabolic inhibition [13]. Furthermore, after 12 h continuous use some neonates had skin burns [13]. In adults, the TOSCA with an electrode temperature of 42°C had a bias of +0.02 kPa, only [15]. The TOSCA monitor initially preheats the skin to 45°C for rapid arterialization before measurements and measurements were performed at the earlobe, using an ear clips. We did not find any effect of using high temperature before low temperature, or vice versa.

Surprisingly, in our study the bias was pCO<sub>2</sub>-dependent. The transcutaneous electrode uses an electrochemical principle: tcPCO<sub>2</sub> is measured by determining the pH of an electrolyte layer separated from the skin by a highly permeable membrane. Diffusing CO<sub>2</sub> changes the pH of the electrolyte solution and the pH changes are proportional to the logarithm of pCO<sub>2</sub>-changes. Since tissue heating increases tissue pCO<sub>2</sub> by 4.5%/°C and some CO<sub>2</sub> is produced locally, the tcPCO<sub>2</sub> is higher than the arterial pCO<sub>2</sub> and corrections for this are included in the monitor. For optimal use, the electrolyte and membrane must be changed every week, and the sensor must also be calibrated whenever a new patient is monitored. This was done meticulously for our two experimental monitors and furthermore control measurements using the calibration gas were done after measurements in all infants. If out of the acceptable range, recalibration and/or change of membrane was done. Therefore we cannot explain the pCO<sub>2</sub> dependency of the bias. Combined with the upwards skew of the distribution of transcutaneous-arterial pCO<sub>2</sub> difference, it meant that occasionally tcPCO<sub>2</sub> overestimated significantly, and this occurred more often at high arterial tcPCO<sub>2</sub>. In practical terms this means that a very high tcPCO<sub>2</sub> must be used with care. Safe practice would be taking a blood sample before significant clinical action, e.g. intubation for mechanical ventilation.

In conclusion, a lower transcutaneous electrode temperature increases the systematic overreading of tcPCO<sub>2</sub>. Monitoring at 40 or 41°C, however, appears reasonable since the typical bias is less than 1 kPa. In some cases with high arterial pCO<sub>2</sub> the overreading was clinically significant and care should be taken that this does not have undue clinical consequences.

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