


Capnography

A Comprehensive Educational Website
Designed, produced, and maintained by
Bhavani-Shankar Kodali MD
Associate Professor,
Department of Anesthesiology, Brigham and Women's Hospital,
Harvard Medical School, Boston, MA



Site launched August 2001 Edition 8, updated 2014

Physiology of capnography

Bhavani Shankar Kodali MD

Can Negative (a-ET)CO₂ differences occur?

YES

Where can they occur?

Healthy subjects during low frequency high tidal volume ventilation
Pregnant subjects
Infants and Children
After coming off cardiac bypass
During and after exercise.

What are the reasons for negative values?

Experimental errors

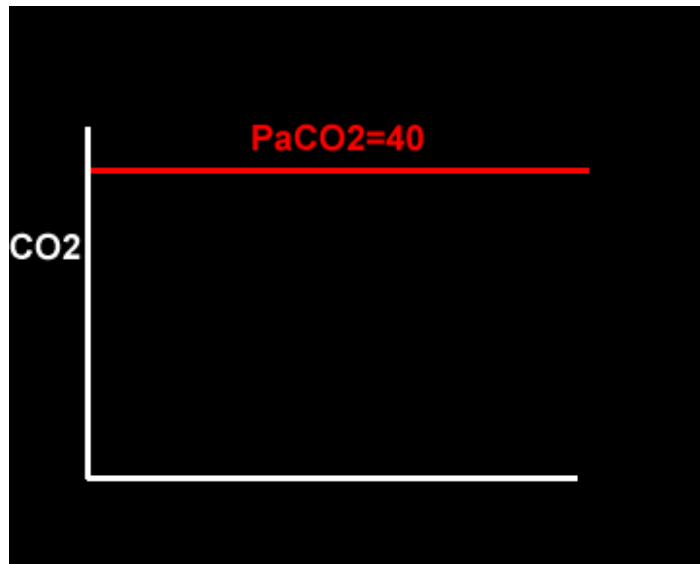
Rebreathing

Inadvertent addition of CO₂ to the inspired gases

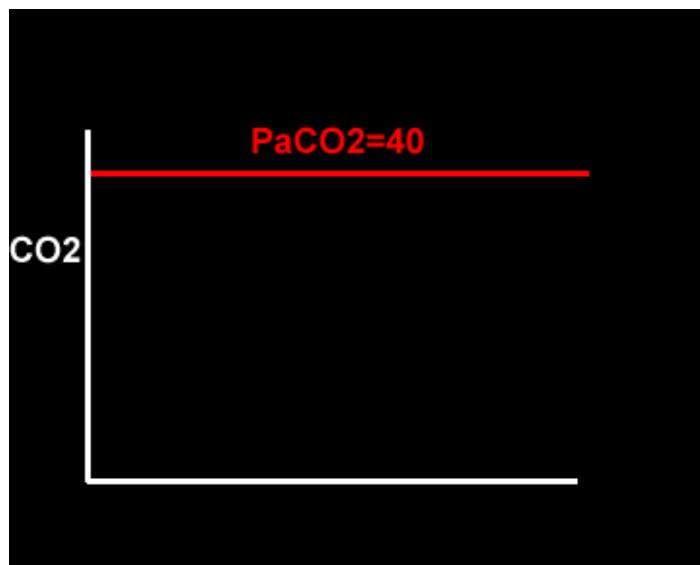
Physiological reasons

Four physiological reasons

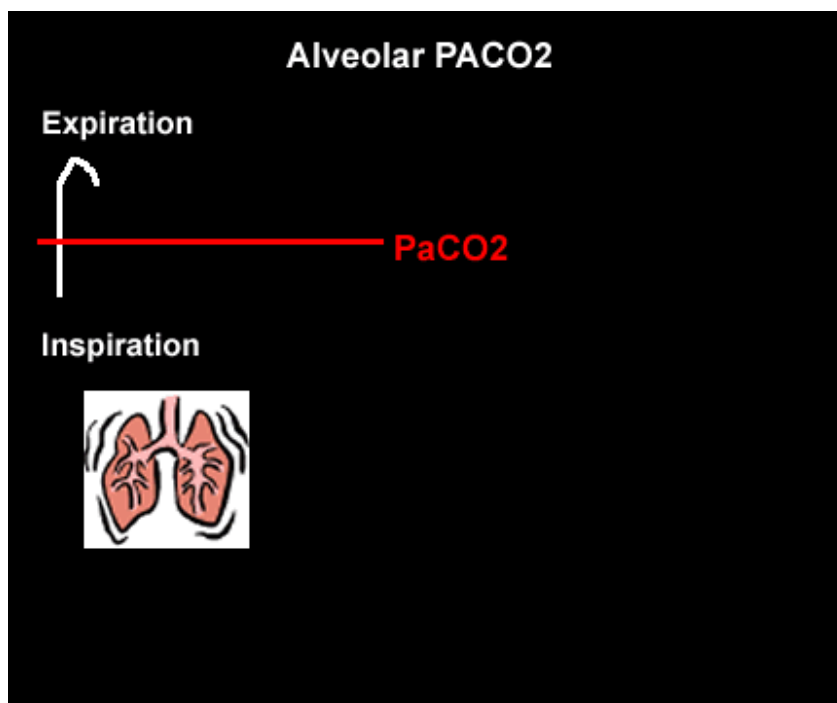
Smaller alveolar dead space and inherent upward slope of phase III



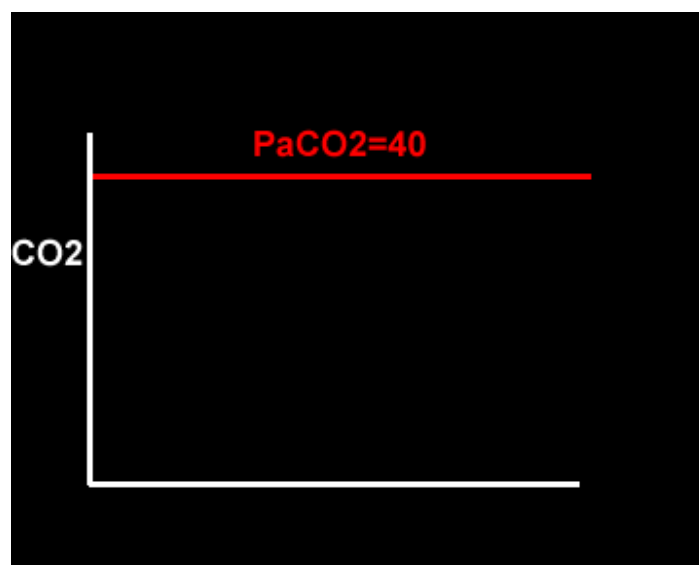
Increase in the slope of phase III



Exaggerated alveolar PCO₂ fluctuations during respiratory cycle due to increased CO₂ production and decreased FRC which make it more likely to **sample higher alveolar PCO₂** (sampled as PETCO₂) during expiration **greater than mean PaCO₂**



Occurrence of phase IV



Negative (a-ET)PCO₂ gradients:

Negative (a-ET)PCO₂ values were observed more than 30 yr ago by Nunn et al. during anesthesia although no explanation was offered.¹ Fletcher et al. observed negative or zero (a-ET)PCO₂ values in 12% of normal subjects during anesthesia and IPPV with large tidal volumes and low frequencies.² Negative values were also observed during anesthesia in 50% of pregnant subjects,³ in 8.1% of patients after post-cardiac bypass (PCB)⁴ and in 50% of infants.⁵

Piper reviewed several studies with negative arterial to end-tidal CO₂ differences in 1986 and

concluded that the reasons for the remarkably pronounced disagreement between the experimental data of different studies cannot be definitely identified and suggested that it is desirable that more observational and experimental data become available in future to review this subject.⁶ Since then several studies have reported negative differences as stated above. The following possible mechanism have been postulated to explain observed (a-ET)PCO₂ differences under various circumstances.

Large tidal volume and low frequency ventilation result in (i) better ventilation of dependent well-perfused alveoli which improves V/Q matching (small area of alveolar dead space as above in figure I). (ii) Gas emptying from slow alveoli to reach the mouth, whereas it would have remained in the airways with small frequent breaths. Under these circumstances the low V/Q areas (alveoli with higher PCO₂) make a more substantial contribution to the gas exchange. The net effect of these factors is to enable the terminal part of phase III to exceed mean PaCO₂, resulting in negative (a-ET)PCO₂.²

Alveolar PCO₂ varies cyclically, being lowest at end-inspiration and highest at end-expiration. However, because of mixing in the heart and syringe, PaCO₂ sampled at the radial artery is the spatial and temporal mean of alveolar PCO₂ (Riley's physiological integrator) and therefore it is quite possible for PETCO₂ to exceed the sampled PaCO₂. The increased cardiac output and increased CO₂ production, reduced FRC and low compliance associated with pregnancy may result in greater cyclical variations in alveolar PCO₂ during a respiratory cycle and also in more alveoli with long time constants. During expiration, PACO₂ increases towards PVC₂ (partial pressure of mixed venous CO₂) more rapidly in pregnant subjects because a larger amount of CO₂ is evolved into a lung which becomes smaller as expiration continues. Further, pregnant subjects resemble the obese in some features namely reduced FRC and low total compliance and hence may exhibit a biphasic slope reminiscent of phase IV of the nitrogen closing volume test. The PCO₂ of most alveolar gas is less than PaCO₂ but, in the terminal part of the expirate, PCO₂ rises rapidly and may exceed PaCO₂. The combined effect of these two mechanisms increases the slope of phase III (Figure 4) and the likelihood of sampling a PETCO₂ greater than PaCO₂.^{3,7-12} The presence of a wide range of V/Q mismatching and reduced FRC may result in negative (a-ET)PCO₂ values in patients after cardiopulmonary bypass.^{4,7} Increased CO₂ production and reduced FRC may be responsible for the negative (a-ET)PCO₂ values observed in infants.⁵

1. Nunn JF, Hill DW. Respiratory dead space and arterial to end-tidal CO₂ tension difference in anesthetized man. *J Appl Physiol* 1960;15:383-9.
2. Fletcher R, Jonson B. Dead-space and the single breath test for carbon dioxide during anaesthesia and artificial ventilation. *Br J Anaesth* 1984;56:109-19.
3. Shankar KB, Moseley H, Kumar Y, Vemula V. Arterial to end-tidal carbon dioxide tension difference during cesarean section anaesthesia. *Anaesthesia* 1986;41:698-702.
4. Russell GB, Graybeal JM, Strout JC. Stability of arterial to end-tidal carbon dioxide gradients during postoperative cardiorespiratory support. *Can J Anaesth* 1990;37:560-6.
5. Rich GF, Sconzo JM. Continuous end-tidal CO₂ difference sampling within the proximal

endotracheal tube estimates arterial CO₂ tension in infants. *Can J Anaesth* 1991;38:201-3.

6. Piper Johannes. Blood-gas equilibrium of carbon dioxide in lungs: a continuing controversy. *J. Appl Physiol* 1986;60:1-8.

7. Shankar KB, Moseley H, Kumar Y. Negative arterial to end-tidal gradients. *Can J Anaesth* 1991;38:260-1.

8. Shankar KB, Moseley H, Kumar Y, Vemula V, Krishan A. Arterial to end-tidal carbon dioxide tension difference during anaesthesia for tubal ligation. *Anaesthesia* 1987;42:482-6.

9. Jones NL, Robertson DG, Kane JW. Differences between end-tidal and arterial PCO₂ in exercise. *J Appl Physiol* 1979;47:954-60.

10. Fletcher R. Arterial to end-tidal CO₂ tension differences. *Anaesthesia* 1987;42:210-1.

11. Bhavani-Shankar K, Moseley H, Kumar AY, Delph Y. Capnometry and anaesthesia. *Can J Anaesth* 1992;39:6:617-32.

12. Bhavani-Shankar K, Kumar AY, Moseley HSL, Hallsworth RA. Terminology and the current limitations of time capnography: A brief review. *J Clin Monit* 1995;11:175-82.

[Back](#)

[Main](#)