EDITORIAL Comment

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Carbon dioxide production during cardiopulmonary bypass: continuous measure and clinical relevance

Carbon dioxide production during cardiopulmonary bypass derives from both the aerobic metabolism and the buffering of lactic acid produced by tissues under anaerobic conditions. Therefore, carbon dioxide removal monitoring is an important measure of the adequacy of perfusion and oxygen delivery. However, routine continuous monitoring of carbon dioxide removal is not widely applied.

Carbon dioxide (CO₂) removal monitoring during cardiopulmonary bypass (CPB) is considered a recommended guideline for practice by the American Society of Extracorporeal Technology and a standard of practice by the Australian New Zealand College of Perfusion.^{1,2} This monitoring is usually performed through capnometric analysis of gases from the exhaust port of the oxygenator. Despite its relative easiness, this practice is not routinely applied.

 $\rm CO_2$ tension (PCO_2) as measured by capnometry is representative of the complex interaction of $\rm CO_2$ production (VCO_2) by cells and its elimination by the natural or artificial lung. Therefore, this technique may provide important information, not only on the efficacy of $\rm CO_2$ removal, but even on the metabolic status of the peripheral organs and on the adequacy of their perfusion with respect to their oxygen (O₂) needs.³

The hollow-fibre/membrane oxygenators commonly used on CPB have totally different performance in terms of CO₂ clearance with respect to the natural lung. Actually, the ventilation/perfusion ratio of the natural lung is around 0.8 (higher at the top and lower at the bottom of the lung in the orthostatic position). An increase of this ratio, as happens in the case of low pulmonary blood flow, creates the conditions for a PvCO₂ – ePCO₂ gradient (Mixed venous carbon dioxide tension - Exhaled carbon dioxide tension). Conversely, artificial lungs have a much higher efficiency in clearing venous blood from the CO₂, with an optimal ventilation/perfusion ratio of around 0.4-0.5.⁴

This property of the artificial lung offers a great advantage for the measurement of VCO₂ during CPB. The ability of the oxygenator to clear off the CO₃, even at a low

ventilation/perfusion ratio, avoids the onset of a $PvCO_2 - ePCO_2$ gradient and the $ePCO_2$ measured at the exhaust port of the oxygenator can be reliably used to assess the VCO_2 .

Infra-red spectography is the most popular means currently used to monitor CO_2 and most centres that use oxygenator exhaust capnography routinely use sidestream devices because of their ease of use.

Initially, capnography was mainly used to measure metabolic CO_2 production and to estimate $PaCO_2$. Many studies show a good correlation between ePCO2 and uncorrected $PaCO_2$ during clinical cases,⁵⁻⁷ making it easier to maintain a specific target $PaCO_2$ during CPB. Although there is a good linear correlation between ePCO₂ and uncorrected $PaCO_2$, there is some deviation, especially during the rewarming phase. This deviation is the same for oxygenators of the same type and brand, but can differ between types and brands.

More recent research showed that VCO₂ is a good predictor of anaerobic metabolism⁸ and, as such, can help to reduce CPB-related morbidity. Indeed, CO₂-derived parameters are more rapid and sensitive than O2-derived parameters in detecting anaerobic metabolism.⁹ A combination of DO₂, VO₂ and VCO₂ parameters was significantly associated with the risk of postoperative renal insufficiency.¹⁰

The potential clinical relevance of VCO₂ monitoring goes far beyond the safety control of the maintenance of an adequate PaCO₂ of the patient throughout the CPB procedure. Certainly, the on-line measurement of ePCO₂ at the exhaust port of the oxygenator may prompt sweep gas adjustments to rapidly adjust the systemic PaCO₂ during the different phases of CPB. Increased values of ePCO₂ as an expression of increased VCO₂ can be found at the release of the cross-clamp, due to the reperfusion of the heart (anaerobic CO₂) or during the rewarming phases after deep hypothermia, as an expression of the increasing VO₂ (aerobic CO₂) and of the decreased solubility of CO₂. This last mechanism should be considered when assessing

 VCO_2 during CPB cooling and rewarming phases, where the changes in CO_2 solubility respectively decrease and increase the ePCO₂.

Nowadays, the routine use of low temperatures on CPB has been replaced by moderate hypothermia or normothermia in many institutions. However, CPB temperatures <28°C may still be used in congenital heart surgery and for specific interventions of high complexity. Within the setting of profound hypothermia, the changes in CO₂ solubility result in corresponding changes in pH, with a reflection on cerebral blood flow. To compensate for the low values of CO₂ at low temperatures, the pH- stat strategy considers the addition of exogenous CO₂ to the sweep gas.

Another exogenous source of CO_2 may come from the flooding of the surgical field to prevent the formation of large air bubbles inside the heart chambers. The finding of elevated values of $ePCO_2$ is rarely attributable to a failure of the oxygenator.

Apart from this, the measurement of the VCO₂ may offer important information on the adequacy of the perfusion in terms of DO₂ (pump flow x arterial O₂ content). In the pathophysiology of CO₂ production, under conditions of inadequate DO₂, there is an excess CO₂ production as a result of lactic acid buffering. Therefore, VCO₂ may be considered as an indirect marker of lactate increase.

The use of DO_2 and VCO_2 to guide the perfusion management are now included in the concept of the "Goal Directed Perfusion (GDP)".¹²⁻¹⁴ The GDP concept considers that the goal of perfusion is to maintain an adequate oxygen supply to all the organs, avoiding the patient entering into the anaerobic zone.

Despite the many possible applications of CO_2 -derived parameters during CPB, very few studies have been published in this area. The expanding concept of GDP will probably increase the interest of clinicians and researchers in these measurements. Further studies on the clinical relevance of CO_2 production monitoring are warranted.

The article from Valdir Filho¹⁵ has revived this subject. Shows an available technique to assess carbon dioxide production and removal and the clinically relevant applications of carbon dioxide-related variables as markers of the adequacy of perfusion during cardiopulmonary bypass.

REFERENCES

 Baker RA, Bronson SL, Dickinson TA, et al. Report from AmSECT's International Consortium for Evidence- Based Perfusion: American Society of Extracorporeal Technology Standards and Guidelines for Perfusion Practice: 2013. J Extra Corpor Technol 2013; 45: 156–166.

- The Australian and New Zealand College of Perfusion. Regulations and Guidelines for Perfusionists. Available at: http://esvc000803.wic050u.server-web.com/Documents/ ANZCP%20Regulations.pdf. Accessed December 17, 2015.
- Wasserman K, Whipp BJ, Casaburi R. Respiratory con- trol during exercise. In: NS Cherniack, JG Widdicombe (Eds), Handbook of Physiology, Section 3: The Respiratory System. American Physiological Society, Bethesda 1986, pp 595–619.
- Pybus DA, Lyon M, Hamilton J, Henderson M. Measuring the efficiency of an artificial lung: 1. Carbon dioxide transfer. Anaesth Intens Care 1991; 19: 421–443.
- 5. Alston RP, McNicol J. Oxygenator exhaust capnography: an in vitro evaluation. J Cardiothorac Anesth 1988; 2: 798–802.
- Zia M, Davies FW, Alston RP, Anaes FC. Oxygenator exhaust capnography: a method of estimating arterial carbon dioxide tension during cardiopulmonary bypass. J Cardiothorac Vasc Anesth 1992; 6: 42–45.
- Weightman WM, Sheminant MR. Oxygenator exhaust capnography as an index of arterial carbon dioxide ten- sion during cardiopulmonary bypass using a membrane oxygenator. Br J Anaesth 2000; 84: 536–537.
- Ranucci M, Isgrò G, Romitti F, Mele S, Biagioli B, Giomarelli P. Anaerobic metabolism during cardiopulmonary bypass: predictive value of carbon dioxide derived parameters. Ann Thorac Surg 2006; 81: 2189 –2195
- Mekontso-Dessap A, Castelain V, Anguel N, et al. Combination of venoarterial PCO2 difference with arteriovenous O2 content difference to detect anaerobic metabolism in patients. Intensive Care Med 2002; 28: 272–277.
- de Somer F, Mulholland JW, Bryan MR, Aloisio T, Van Nooten GJ, Ranucci M. O2 delivery and CO2 production during cardiopulmonary bypass as determinants of acute kidney injury: time for a goal-directed perfusion man- agement? Crit Care 2011; 15: R192.
- Dres M, Monnet X, Teboul J-L. Hemodynamic manage- ment of cardiovascular failure by using PCO2 venous- arterial difference. J Clin Monit Comput 2012; 26: 367–374.
- Dijoy L, Dean JS, Bistrick C, Sistino JJ. The history of goaldirected therapy and relevance to cardiopulmonary bypass. J Extra Corpor Technol 2015; 47: 90–94.
- Ranucci M, Aloisio T, Carboni G, et al. Acute kid- ney injury and hemodilution during cardiopulmonary bypass: a changing scenario. Ann Thorac Surg 2015; 100: 95–100.
- Ranucci M, Romitti F, Isgrò G, et al. Oxygen delivery during cardiopulmonary bypass and acute renal failure after coronary operations. Ann Thorac Surg 2005; 80: 2213–2220.
- 15. Assis-dos-Reis-Filho V, Lopes-de-Oliveira E, Scramim JF, Sanga MA, Arrais-dos-Santos M. Benefits of continuous monitoring of PCO2 obtained from a system applied to membrane oxygenator exhaustion of the cardiopulmonary bypass circuit. Rev Port Cir Cardiotorac Vasc. 2019; 26(3):205-208.