

# Monitoring and optimization of the microcirculation during CPB

# John M. Murkin

Department of Anesthesia & Perioperative Medicine, London Health Sciences Centre, University Hospital, London, ON, Canada *Correspondence to:* John M. Murkin, MD, FRCPC, FACI (Hon). London Health Sciences, University Hospital, Main Building – Room C3-112, London, ON N6A 5A5, Canada. Email: John.Murkin@lhsc.on.ca.

**Abstract:** In this review the impact of cardiopulmonary bypass (CPB) on the microcirculation is discussed. The impact of conventional non-pulsatile perfusion is contrasted with the microcirculatory impact of pulsatile CPB which is associated with better preservation of microcirculatory function. Evidence that alterations in microcirculatory function that persist following cardiac surgery are associated with adverse clinical outcomes is reviewed. Recent studies using novel techniques employing near infrared reflectance spectroscopy (NIRS) and sublingual microscopy to explore alterations in microcirculatory function during CPB are reviewed and the implications of these observations for studies investigating minimally invasive extracorporeal circulation (MiECC) are discussed.

**Keywords:** Cardiopulmonary bypass (CPB); microcirculation; near-infrared reflectance spectroscopy (NIRS), tissue oximetry; minimally invasive extracorporeal circulation (MiECC); non-pulsatile bypass

Submitted Dec 06, 2018. Accepted for publication Feb 27, 2019. doi: 10.21037/jtd.2019.02.100 View this article at: http://dx.doi.org/10.21037/jtd.2019.02.100

Cardiac surgery can be associated with organ dysfunction and the degree of organ dysfunction is associated with length of ICU stay and mortality (1). In a majority of cardiac surgical procedures cardiopulmonary bypass (CPB) is used to facilitate surgery. It has been shown that conventional CPB (cCPB) can be associated with significant decreases in both cerebral blood flow (CBF) and cerebral metabolic rate for oxygen (CMRO2), during laminar flow in comparison with use of a pulsed waveform (2). Pulsatile CPB when compared with cCPB has also been associated with decreased incidences of end organ dysfunction (3). Although there is evidence that high risk patients, particularly those with renal failure, have improved outcomes with pulsatile CPB (4), currently non-pulsatile CPB remains standard of care in a majority of cardiac centers. Until recently it has been difficult to examine tissue microcirculatory perfusion during cardiac surgery and CPB.

CPB is known to induce an inflammatory reaction and release of inflammatory mediators that may exert effects on the microcirculation similar to severe sepsis. De Backer *et al.* observed alterations in microcirculatory function during and after cardiac surgery in 15 patients using Side Stream Dark Field Imaging (SSDFI) that were present after induction of anesthesia and persisted for 24 hours post-operatively (5). Very few studies have been published describing the microcirculation during and after cardiac surgery using non-invasive monitoring. While the mechanism for decreased organ perfusion with cCPB remains speculative, it is reported that non-pulsatile perfusion during cardiac surgery results in decreased microcirculatory oxygenation in the skin and gut (6).

Near-infrared reflectance spectroscopy (NIRS) uses analysis of absorptance spectra in the near-infrared range (680 to 1,100 nm) to determine regional tissue hemoglobin oxygen saturation (7). Near-infrared light readily penetrates biological tissues and is absorbed mainly by hemoglobin. According to Beer's Law, the near-infrared spectroscopy signal is limited to vessels with a diameter less than 1 mm (arterioles, capillaries, and venules). Commercially available tissue NIRS (tNIRS) devices detect relative percentages of oxygenated and de-oxygenated blood in tissue to determine tissue oxygen saturation. tNIRS has been used to assess oxygen balance in the microcirculation of skeletal muscles of shock patients (8). Since cCPB invokes a significant inflammatory response impairing microcirculatory perfusion (9), tNIRS is increasingly being utilized to assess S1490

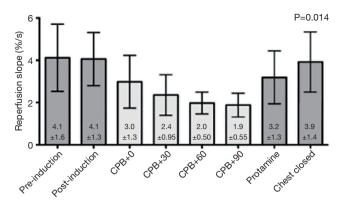


Figure 1 Reperfusion slope at various measurement points pre, during and after cardiopulmonary bypass (CPB) showing progressive decrease in microcirculatory vasoreactivity with duration of non-pulsatile CPB. Error bars and  $\pm$  values represent 95% CI [reproduced with permission from Smith RS, Murkin JM. (11)].

tissue microcirculation during CPB,

The first studies using tNIRS and vascular occlusion test (VOT) in conjunction with SSDFI demonstrated similar directional changes with both techniques when comparing pulsatile and non-pulsatile perfusion during CPB (10). In a randomized cohort study of 20 high risk patients, O'Neill *et al.* have recently reported that compared with cCPB, roller pump generated pulsatility is associated with improved microcirculatory blood flow and tissue oxygen saturation (10).

Recent studies using tNIRS have demonstrated a progressive reduction in microcirculatory vasoreactivity during cCPB (11). Using a pneumatic cuff to evoke a quantifiable ischemic challenge, the vascular occlusion test (VOT) uses tNIRS to measure the vasoreactivity of peripheral tissue during and after CPB and cCPB was shown in a small study to induce a progressive reduction in microcirculatory responsivity with longer duration cCPB that returned to baseline after separation from cCPB as shown in *Figure 1* (11). This study did raise the concern that the progressive decrease in microvascular vasoreactivity seen during cCPB, if not restored to baseline after separation from CPB, could be a harbinger of end organ dysfunction. Further studies have amplified this concern (12).

In a study of 254 patients undergoing cardiac and thoracic surgery in whom tNIRS and VOT were employed sequentially before and after surgery, although conventional hemodynamic values, such as cardiac output and blood pressure, did not differ between the groups, on multivariable regression and linear analyses low VOT recovery slope on postoperative day 1 was associated with increases of composite complications and hospital length of stay (12).

A more recent study again using tNIRS and VOT during cCPB and pulsatile CPB has shown better preservation of microcirculatory reactivity with pulsatile perfusion (10). As such it is likely that cCPB induces progressive reduction in microcirculatory perfusion—not only as shown in peripheral muscle, but similar phenomena likely also occur in deep tissue and organs and may well contribute to end organ dysfunction associated with cCPB (13). Use of pulsatile CPB has been associated with decreased rates of renal failure (14).

Taken together these studies would indicate that cCPB induces progressive decrease in microcirculatory vasoreactivity and that those patients in whom abnormal VOT—indicative of microcirculatory dysfunction—persists into the postoperative period, have a significantly greater incidence of end-organ dysfunction.

Use of minimally invasive extracorporeal circulation (MiECC) for CPB has been associated with a variety of improved clinical outcomes in comparison to cCPB (15). It is emphasized that MiECC represents not only a low volume circuit technology but rather that it engenders an approach that is focused on minimizing all procedural aspects of cardiac surgery from amount of fluids administered to duration of postoperative ventilation. Various endpoints including a survival benefit and reduced transfusion and higher hemoglobin levels, as well as better preservation of renal function and lowered levels of inflammatory biomarkers have all been associated with MiECC and may well reflect improved microcirculatory perfusion (15).

As such, it is tNIRS and VOT by enabling continuous and non-invasive assessment of tissue microcirculatory perfusion and oxygenation during CPB—when coupled with clinical outcomes—that will further help evaluate and improve circulatory support techniques. The role of MiECC and its impact on microcirculatory flow will be investigated and should provide further evidence of the fundamental role preservation of the microcirculation plays in helping to further improve patient outcomes.

#### **Acknowledgments**

None.

## Footnote

*Conflicts of Interest:* The author has no conflicts of interest to declare.

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**Cite this article as:** Murkin JM. Monitoring and optimization of the microcirculation during CPB. J Thorac Dis 2019;11(Suppl 10):S1489-S1491. doi: 10.21037/jtd.2019.02.100

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