

Do Low-density Cerebral Oximetry Measures Accurately Detect Variability of Cerebral Perfusion During Cardiac Surgery?

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Introduction

- Cerebral oxygenation and perfusion are important clinical parameters, since hypoxia is the primary cause of neurological injuries [1].
- These parameters may help guide intraoperative monitoring during procedures associated with neurological complications [2,3].
- While stroke and cognitive dysfunction are complications of many surgical procedures, the incidence following cardiac surgery remains highest [4,5]. Consequently, prompt identification of cerebral hypoxia before irreversible injury occurs is paramount.
- Devices currently approved by the Food and Drug Administration (FDA) provide non-invasive monitoring of cerebral oxygen saturation based on low-density configurations of transmitters and sensors [6,7,8].
- Intraoperative cerebral monitoring using cerebral oximetry, coupled with intervention to keep brain oxygen saturation above a fixed threshold, has not significantly reduced the incidence of stroke following cardiac surgery [9].
- Given the heterogeneity of cardiac surgical patients, who commonly present with
 one or more pre-existing risks (e.g., hypertension, diabetes, cerebral / coronary /
 peripheral atherosclerotic disease), it is uncertain whether oximetry based on
 sampling small areas over the frontal lobe is representative of regional cerebral
 perfusion.
- In this study we have explored whether low-density optical sensor arrays, derived from small subsets of four larger arrays, are able to provide representative measures of the true state of regional cerebral perfusion in patients undergoing cardiac surgery.

Methods

- · Six heart surgery patients were recruited for this study (Table 1).
- Optodes were attached to a headgear and arranged in arrays covering 4 sites (Fig 1).
- Optical recordings and a record of surgical supportive events were taken during surgery using a Near-Infrared Spectroscopic (NIRS) imaging system (NIRx DYNOTcompact) and examined retrospectively to identify clinically significant events.
- An anesthesia monitor (Dräger-Narkomed 6000) was used during surgery to simultaneously measure physiological parameters such as mean systemic arterial and pulmonary arterial pressures.
- A global estimate of light-source intensity variability and superficial hemodynamic fluctuations was computed.
- Least-squares linear regression was used to subtract the contribution of the global factor from each raw data time series [10]. The normalized difference method [11] was used to recover time series of volumetric images from the pre-processed data (see Fig. 2).
- Ànalysis of hemoglobin (Hb) levels was done using a modified Beer-Lambert law (MBL) [12] algorithm to compute time series of oxy, deoxy and total Hb (Hb_{oxy}-Hb_{otas}) for all channels.
- Superficial-signal correction was carried out for each channel as depicted in Fig 3.
 Channels labeled S1D4 and S4D1 are a reciprocal pair, with the roles of S and D interchanged.
- The signals measured contain cerebral and overlying tissue contributions. In contrast, channels S1D3 and S4D2 consist largely of signals from overlying tissue.
 This would appear to allow for selective removal of the superficial-tissue component of the data recorded by the S1D4 and S4D1 channels [13].
- Correlation coefficients are computed for all pairs of channels having the same S-D separation distance (see Fig. 4).
- The superficial-signal corrected time series of Hb_{oxy}, Hb_{deoxy}, & Hb_{btal}, for all channels with S-D separation ≥ 3cm, were compared in regard to the magnitudes of the detected Hb concentration changes. A large, relatively abrupt change in Hb level is taken as an indication that a clinically significant event is occurring (see Fig. 5(a)). The overall sensitivity—i.e., percentage of S-D channels that report the clinically significant event—was calculated, as a function of the trigger level (see Fig. 5(b)).

Results

Subject	Gender	Age	Surgery	Measurement Duration	Hemodynamic Event
1	F	65	CABG – Off-pump converted to on-pump beating heart	5h 18m	VF arrest / crash on CPB
2	F	73	CABG – Off-pump	5h 38m	VF / Cardioversion
3	М	67	AVR – On-pump	3h 00m	Cardiac Cannulation
4	F	58	CABG – On-pump	4h 32m	Initiation of CPB
5	F	58	CABG – On-pump	3h 27m	Low CPB flow rate
6	F	27	AVR – On-pump	4h 44m	Retrograde Autologous Priming of CPB

Table 1. Clinical information for the study participants. Abbreviations: AVR = aortic valve replacement, CABG = coronary artery bypass graft, CPB = cardiopulmonary bypass, VF = ventricular fibrillation.

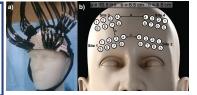


Figure 1. a) Photograph of headgear with the four-site optode array; b) Diagram of the optode arrangements in each site, and their inter-site distances.

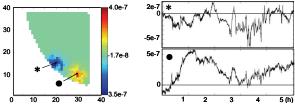


Figure 2. Left: 2D horizontal section through the volumetric image of Hboxy, recovered from Site-1 data collected near the 2.5-hr mark, for one of the study participants. Right: time series of the recovered Hboxy concentrations (time-varying perturbations with respect to a baseline mean value) in two selected pixels, labeled by the ** and ** symbols.

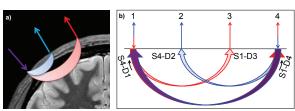
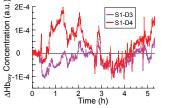


Figure 3. a) Cartoon illustrating the principle of the small-array cerebral oxygenation probe: light received by the detector located closer to the source (blue) predominantly traverses a superficial volume of tissue, while the detector located farther from the source (red) receives light that passes through both extra-cerebral and cortical tissue. b) A 180° rotation of the probe consisting of \$1, D3 and D4 (red arrows) transforms it into the probe consisting of \$4, D2 and D1 (blue arrows).



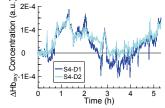
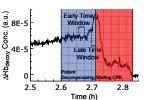


Figure 4. Hb_{ow} time series recorded from a set of channels having the geometry depicted in Fig. 3(b). The curves for channels S1-D4 and S4-D1 (3 cm S-D separation, reciprocal pair) are nearly superimposable (r = 0.995), while the time series for S1-D3 and S4-D2 (2 cm) are substantially different.



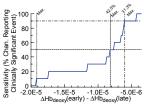


Figure 5. a) ΔHb_{decoxy} time series during clinically significant event, for one selected S-D channel. b) Percentage of S-D channels (fixed separation) that report clinically significant event, as a function of the triocer level.

Conclusion

- While subjects were under general anesthesia, their intraoperative cerebral perfusion remained highly heterogeneous.
- Minor changes in source-detector pair location result in notably differing signal recordings.
- FDA-approved non-invasive cerebral oximetry devices, based on low-density arrays, are unlikely to yield accurate representation of complex heterogeneous cerebral perfusion.
- In contrast, a tomographic imaging method with a rich array of optodes would retain the possibility to capture time-varying heterogeneous spatial maps of cerebral perfusion.

References

- [1] Zwienenberg-Lee M, Kim KD, Muizelaar JP. Injury to the Central Nervous System. In Souba WW, Fink MP, Jurkovich GJ, Kaiser LR, Pearce WH, Pemberton JH, Soper NJ (ed.): ACS Surgery: Principles and Practice, 7th Edition. New York, WebMD, 2008
- [2] Casati A, Spreeafico M, Putzu M, et al. New technology for noninvasive brain monitoring: continuous cerebral oximetry. Minerva Anestesiol 2006; 72:605-25
- [3] Edmonds Jr HL, Ganzel BL, Austin III EH. Cerebral Oximetry for Cardiac and Vascular Surgery Seminars in Cardiothoracic and Vascular Anesthesia 2004; 8:147-166
- [4] Moller JT, Cluitmans P, Rasmussen LS, et al: Long-term postoperative cognitive dysfunction in the elderly ISPOCD1 study (see comments). International Study of Post-Operative Cognitive Dysfunction (ISPOCD) Investigators. Lancet 1998; 351:857
- [5] Savino JS, Cheung AT. Cardiac Anesthesia. In Cohn LH (ed.): Cardiac Surgery in the Adult, 3rd Edition. New York, McGraw-Hill, 2008:281-314
- [6] CASMED: www.casmed.com/foresight.html
- [7] Somanetics: www.somanetics.com/invos-system
- [8] Nonin: www.nonin.com/Regional Oximetry
- [9] Murkin JM, Adams SJ, Novic RJ, et al. Monitoring brain oxygen saturation during coronary bypass surgery: a randomized prospective study. Anesth Analg 2007;104:51-8
- [10] White BR, Snyder AZ, Cohen AL, Petersen SE, Raichle ME, Schlaggar BL, Culver JP. Restingstate functional connectivity in the human brain revealed with diffuse optical tomography, NeuroImage 2009; 47:148–156
- [11] Pei Y, Graber HL, and Barbour RL. Influence of systematic errors in reference states on image quality and on stability of derived information for DC optical imaging. Applied Optics 2001; 40:5755-5769
- [12] Schmitz CH, Klemer DP, Hardin RE, Katz MS, Pei Y, Graber HL, Levin MB, Levina RD, Franco NA, Solomon WB, and Barbour RL. Design and implementation of dynamic near-infrared optical tomographic imaging instrumentation for simultaneous dual-breast measurements. Applied
- [13] Fischer GW. Recent advances in application of cerebral oximetry in adult cardiovascular surgery. Semin Cardiothorac Vasc Anesth. 2008; 12:60-9

Acknowledgements

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