Cerebral near-infrared spectroscopy insensitively detects low cerebral venous oxygen saturations after stage 1 palliation

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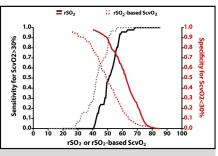
ABSTRACT

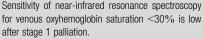
Background: Measurement of cerebral venous oxyhemoglobin saturation $(ScvO_2)$ is considered a gold standard in assessing the adequacy of tissue oxygen delivery (DO_2) after the stage 1 palliation (S1P), with $SvO_2 < 30\%$ often representing severely compromised DO_2 . Regional oxygenation index (rSO_2) based on near-infrared resonance spectroscopy (NIRS) frequently is used to screen for compromised DO_2 , although its sensitivity to detect severe abnormalities in SvO_2 is uncertain.

Methods: $ScvO_2$ was measured by co-oximetry from the internal jugular vein as clinically indicated in 73 neonates after S1P. These values were compared with cerebral rSO₂ (FORE-SIGHT; CASMED) via mixed effects model linear regression, Bland-Altman analysis, and sensitivity analysis. Because NIRS devices measure a composite of arterial and venous blood, we calculated an rSO₂-based ScvO₂ designed to remove arterial contamination from the rSO₂ signal: rSO₂-based ScvO₂ = (rSO₂ – arterial oxygen saturation × 0.3)/0.7.

Results: Among 520 time-matched pairs of ScvO_2 and cerebral rSO₂, the slope of the relationship between rSO₂ and ScvO_2 (after we adjusted for effects of hemoglobin) was 0.37 ± 0.04 with only modest correlation (r² = 0.39), and mean bias of +8.26. When ScvO_2 was <30%, cerebral rSO₂ was <30 in less than 1%, <40 less than 1%, and <50 in 45.7% of data points; specificity of rSO₂ in the same range is >99%. Correction of rSO₂ for arterial contamination significantly decreased mean bias (+3.03) and improved the sensitivity of rSO₂ to detect $\text{ScvO}_2 <30$ to 6.5% for rSO₂ <30, 29% for rSO₂ <40, and 77.4% for rSO₂ <50.

Conclusions: Cerebral rSO₂ in isolation should not be used to detect low ScvO₂, because its sensitivity is low, although correction of rSO₂ for arterial contamination may improve sensitivity. Cerebral rSO₂ of 50 or greater should not be considered reassuring, though values below 30 are specific for low ScvO₂. (J Thorac Cardiovasc Surg 2017; \blacksquare :1-7)





Central Message

Cerebral near-infrared resonance spectroscopy (regional oxygenation saturation [rSO₂]) insensitively detects cerebral venous oxyhemoglobin saturation less than 30% in the range of clinical interest (ie, rSO₂ 30-50), a problem improved by correcting for arterial contamination (rSO₂-based cerebral venous oxyhemoglobin saturation).

Perspective

Although cerebral near-infrared resonance spectroscopy (regional oxygenation saturation) is commonly used to detect low cardiac output, the sensitivity of cerebral near-infrared resonance spectroscopy to detect venous oxyhemoglobin saturation less than 30% is low in the range of clinical interest after stage 1 palliation. This problem is ameliorated by correction of the signal for arterial contamination by backcalculating venous oxyhemoglobin saturation from regional oxygenation saturation and arterial oxygen saturation.

See Editorial Commentary page XXX.

Optimization of systemic oxygen delivery (DO_2) during the early postoperative period after stage 1 palliation (S1P) requires accurate assessment of venous oxyhemoglobin saturation (SvO₂). Although true mixed venous blood sampling is anatomically precluded in single-ventricle anatomy, sampling from the superior vena cava (ie, cerebral venous

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Abbreviations and Acronyms	
DO_2	= systemic oxygen delivery
	= near-infrared resonance spectroscopy
r^2	= correlation coefficient
rSO ₂	= regional oxygenation saturation
S1P	= stage 1 palliation
SaO_2	= arterial oxygen saturation
ScvO ₂	= cerebral venous oxyhemoglobin saturation
SvO_2	= venous oxyhemoglobin saturation

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oxyhemoglobin saturation, ScvO₂) frequently is used as a representation of the adequacy of DO₂ after S1P.¹ When oxygen consumption exceeds DO₂, tissue hypoxia develops, which increases oxygen dissociation from hemoglobin and causes venous oxyhemoglobin desaturation. In the single-ventricle population, it has been suggested that ScvO₂ <30% represents the anaerobic threshold² and that maintaining ScvO₂ >50% in the early postoperative period is associated with improved survival.^{1,3,4} Thus, accurate assessment of ScvO₂ after S1P is vital to the postoperative management of these patients.

Some groups have used venous oximetric catheters to permit continuous monitoring.5 These catheters are somewhat larger in caliber than standard catheters, may be more challenging to insert, and exhibit significant drift in calibration during use, such that their use is not widespread. Thus, a mainstay of patient monitoring is the use of surface oximetry with near-infrared spectroscopy (NIRS) to measure tissue O₂ saturation. Cerebral oximetry is a noninvasive optical technology that relies on the relative transparency of biological tissues (eg, scalp and skull) to near-infrared light to measure brain oxygen saturation in an uncertain mixture of arterioles, capillaries, and venules. Commercial cerebral oximeters use continuous-wave, spatially resolved spectroscopy to measure the ratio of oxyhemoglobin to total hemoglobin, providing a cerebral hemoglobin oxygen saturation. The unit is percent (%) and is referred variably to as regional oxygen saturation (rSO_2) , tissue oxygen saturation, or tissue oxygenation index, depending on the manufacturer. The ratio of venous to arterial blood in the sampled volume is approximately 70% to 30%,⁶ although it can vary significantly (as much as 85%/15%).⁷ Although hemoglobin is distributed unevenly within arteries, arterioles, capillaries, venules, and veins, the venous to arterial ratio is estimated and fixed during the calibration and validation of the monitor (eg, 70%-75% venous/25%-30% arterial⁸⁻¹⁰). Because NIRS is used commonly as a noninvasive screening tool for inadequate DO₂ after S1P, we examined the sensitivity and specificity of NIRS to detect low ScvO₂ in the postoperative setting.

METHODS

Patients

The following retrospective study was approved by the Institutional Review Board at Boston Children's Hospital. We retrospectively studied 84 consecutive patients undergoing single-ventricle palliation for hypoplastic left heart syndrome at our institution between January 2013 and May 2016. Cerebral rSO₂ (applied to the forehead without a preference or laterality) was measured as a part of routine clinical care during this period (FORE-SIGHT tissue oximeter; CASMED, Branford, Conn). Patients who did not have cerebral NIRS data recorded during the postoperative period (n = 3) were excluded from the study. During this period, placement of an internal jugular vein catheter with the tip in the proximal superior vena cava (at or above the junction with the right atrium) was also part of routine clinical practice. Blood sampled from the superior vena cava was analyzed routinely by co-oximetry in the hospital's core laboratory (Radiometer ABL 800; Radiometer America, Brea, Calif). Presence and position of the tip of the internal jugular catheter on the first chest radiograph after admission to the intensive care unit was assessed by 2 independent investigators (E.R. and J.K.). Data were excluded for patients in whom the tip of the catheter was deep within the right atrium (n = 8), leaving a total of 73 patients. Data also were excluded during the times that patients were on extracorporeal membrane oxygenation support. A total of 15,336 hours of patient monitoring were assessed.

Data Extraction and Processing

 $ScvO_2$ and hemoglobin were extracted automatically (Boston Children's Hospital 360; MicroStrategy, Inc, Tysons Corner, Va) from the hospital database in time- and patient-stamped fashion. Similarly, rSO₂ was extracted from 1 of 2 sources. In a subset of patients (n = 20, 26.3%), the NIRS device was tethered to the bedside monitor and the data continuously (ie, every 5 seconds) stored in a database and subsequently exported (T3 Data Collection and Analytics Software System; Etiometry, Inc, Boston, Mass) and down-sampled to every 5-minute median values (MatLab, R2015b, 8.6; MathWorks Inc, Natick, Mass). In the remainder, NIRS data were entered manually by the bedside nurse at least twice per shift, and these data were normalized to elapsed time after removal of the aortic crossclamp (the final was chosen if multiple) via a customized MatLab script.

Data Analyses

Time-matched rSO₂ measurements were compared with ScvO₂ measurements with the use of several techniques. First, the partial correlation coefficient and slope of the relationship were measured with mixed-effects model linear regression to account for repeated measures and to control for hemoglobin with ScvO₂ as the outcome. Second, the mean bias and standard deviation (limits of agreement) between the 2 measurements were compared by Bland-Altman analysis corrected for replicate measurements.¹¹ Third, the sensitivity and specificity of NIRS to detect ScvO₂ critical thresholds were assessed with receiver operating characteristic analysis by defining the presence of deficient oxygen delivery (ie, disease present) as ScvO₂ <30%, <40%, or <50%. Sensitivity and specificity sensitive to detect ScvO₂ <30%.

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