

# Correlation and agreement of regional cerebral oxygen saturation measured from sensor sites at frontal and temporal areas in adult patients undergoing cardiovascular anesthesia

Sirirat Tribuddharat<sup>1</sup>, Kriangsak Ngamsaengsirirup<sup>1</sup>, Phatcharakamon Mahothorn<sup>1</sup>, Thepakorn Sathitkarnmanee<sup>Corresp.</sup><sup>1</sup>

<sup>1</sup> Department of Anesthesiology, Faculty of Medicine, Khon Kaen University, Khon Kaen, Thailand

Corresponding Author: Thepakorn Sathitkarnmanee  
Email address: thepakorns@gmail.com

**Background.** The function and viability of the brain depend on adequate oxygen supply. A decrease in cerebral blood supply causing cerebral desaturation may lead to many neurological complications. Direct measurement of regional cerebral oxygen saturation (rScO<sub>2</sub>) assists in early detection and management. Near-infrared spectroscopy (NIRS) has been introduced for measuring rScO<sub>2</sub>. A pair of sensors are attached to the right and left forehead. However, there are some situations where the forehead of the patient is not accessible for sensor attachment—e.g., neurosurgery involving the frontal area; a bispectral index (BIS) sensor already attached; or a wound to the forehead—so, alternate sites for sensor attachment are required. The temporal area was proposed as an alternate site. The objective of this study was to assess the correlation and agreement of rScO<sub>2</sub> measured at the forehead vs. the temporal area. **Methods.** Adult patients undergoing cardiothoracic or vascular surgery were monitored for rScO<sub>2</sub> using 2 pairs of ForeSight sensors. The first pair (A1 and A2) were attached to the right and left forehead, while the second pair (B1 and B2) were attached to the right and left temporal area. The rScO<sub>2</sub> values measured from A1 vs. B1 and A2 vs. B2 were assessed for correlation and agreement using the Bland-Altman analysis. **Results.** Data from 19 patients with 14,364 sets of data were analyzed. The data from A1 vs. B1 and A2 vs. B2 showed moderate positive correlation ( $r = 0.627$ ;  $P < 0.0001$  and  $r = 0.548$ ;  $P < 0.0001$ ). The biases of A1 vs. B1 and A2 vs. B2 were -2.3% (95% CI -2.5 to -2.2;  $P < 0.0001$ ) and 0.7% (95% CI 0.6 to 0.8;  $P < 0.0001$ ). The lower and upper limits of agreement of A1 vs. B1 were -17.5% (95% CI -17.7 to -17.3) and 12.8% (95% CI 12.6 to 13.0). The lower and upper limits of agreement of A2 vs. B2 were -14.6% (95% CI -14.8 to -14.4) and 16.0% (95% CI 15.8 to 16.3). **Conclusions.** The rScO<sub>2</sub> values measured from sensors at the frontal and temporal areas

show a moderate correlation with sufficiently good agreement. The temporal area may be an alternative to the frontal area for cerebral oximetry monitoring.

1 **Correlation and agreement of regional cerebral oxygen saturation measured from sensor**  
2 **sites at frontal and temporal areas in adult patients undergoing cardiovascular anesthesia**

3

4 Sirirat Tribuddharat<sup>1</sup>, Kriangsak Ngamsaengsirirup<sup>1</sup>, Phatcharakamon Mahothorn<sup>1</sup>, Thepakorn  
5 Sathitkarnmanee<sup>1</sup>

6 <sup>1</sup>Department of Anesthesiology, Faculty of Medicine, Khon Kaen University, Khon Kaen,  
7 40002, Thailand

8

9 Corresponding Author:

10 Thepakorn Sathitkarnmanee

11 Department of Anesthesiology, Faculty of Medicine, Khon Kaen University. 123 Mitrapap Road,

12 Ampur Muang, Khon Kaen, 40002, Thailand Email address:

13 Email: [thepakorns@gmail.com](mailto:thepakorns@gmail.com)

14

15

16

17

18

19

20

21

## 22 Abstract

23 **Background.** The function and viability of the brain depend on adequate oxygen supply. A  
24 decrease in cerebral blood supply causing cerebral desaturation may lead to many neurological  
25 complications. Direct measurement of regional cerebral oxygen saturation (rScO<sub>2</sub>) assists in  
26 early detection and management. Near-infrared spectroscopy (NIRS) has been introduced for  
27 measuring rScO<sub>2</sub>. A pair of sensors are attached to the right and left forehead. However, there are  
28 some situations where the forehead of the patient is not accessible for sensor attachment—*e.g.*,  
29 neurosurgery involving the frontal area; a bispectral index (BIS) sensor already attached; or a  
30 wound to the forehead—so, alternate sites for sensor attachment are required. The temporal area  
31 was proposed as an alternate site. The objective of this study was to assess the correlation and  
32 agreement of rScO<sub>2</sub> measured at the forehead vs. the temporal area.

33 **Methods.** Adult patients undergoing cardiothoracic or vascular surgery were monitored for  
34 rScO<sub>2</sub> using 2 pairs of ForeSight sensors. The first pair (A1 and A2) were attached to the right  
35 and left forehead, while the second pair (B1 and B2) were attached to the right and left temporal  
36 area. The rScO<sub>2</sub> values measured from A1 vs. B1 and A2 vs. B2 were assessed for correlation  
37 and agreement using the Bland-Altman analysis.

38 **Results.** Data from 19 patients with 14,364 sets of data were analyzed. The data from A1 vs. B1  
39 and A2 vs. B2 showed moderate positive correlation ( $r = 0.627$ ;  $P < 0.0001$  and  $r = 0.548$ ;  $P$   
40  $< 0.0001$ ). The biases of A1 vs. B1 and A2 vs. B2 were -2.3% (95% CI -2.5 to -2.2;  $P < 0.0001$ )  
41 and 0.7% (95% CI 0.6 to 0.8;  $P < 0.0001$ ). The lower and upper limits of agreement of A1 vs. B1  
42 were -17.5% (95% CI -17.7 to -17.3) and 12.8% (95% CI 12.6 to 13.0). The lower and upper  
43 limits of agreement of A2 vs. B2 were -14.6% (95% CI -14.8 to -14.4) and 16.0% (95% CI 15.8  
44 to 16.3).

45 **Conclusions.** The rScO<sub>2</sub> values measured from sensors at the frontal and temporal areas show a  
46 moderate correlation with sufficiently good agreement. The temporal area may be an alternative  
47 to the frontal area for cerebral oximetry monitoring.

48

49

50

51

52

53

54

55

56

57

58

59

60

61

62

63

## 64 INTRODUCTION

65 The brain's normal function and viability depend on an adequate oxygen supply. The  
66 current methods to assess oxygen delivery to the brain include measuring blood pressure and  
67 arterial oxygenation via pulse oximetry or blood gas analysis which are surrogate parameters.(1)  
68 Near-infrared spectroscopy (NIRS), a non-invasive cerebral oximetry monitor, has been  
69 introduced to directly measure regional cerebral oxygen saturation (rScO<sub>2</sub>). (2-5) The light in the  
70 near-infrared spectrum can penetrate tissue containing bone and soft tissue/gray matter up to 2.5  
71 cm in depth. Sensors are placed at fixed distances from a light emitter, and algorithms subtract  
72 superficial from deep light absorption to provide an index of tissue oxygenation.(6)

73 Cardiac surgery may reduce cerebral perfusion due to a decrease in cardiac output and  
74 blood pressure leading to cerebral desaturation; thus, it is associated with many neurological  
75 complications.(7-10) It is the most common procedure benefitting from NIRS, which has  
76 resulted in lower rates of postoperative stroke, postoperative cognitive dysfunction (POCD), and  
77 postoperative delirium (POD) in adult cardiac surgery.(11) NIRS measurement does not rely on  
78 pulsatile flow as does pulse oximetry; rather, it averages the oxygenation of the artery, capillary,  
79 and venous flow of the underlying tissue. There is still no consensus on the criteria for cerebral  
80 desaturation, but the commonly used criterion is a greater than 20% reduction of rScO<sub>2</sub> from  
81 baseline or an absolute value of less than 50%, although a reduction greater than 10% was set as  
82 the threshold for early intervention.(12, 13) Since there is broad steady state variability among  
83 individuals and wide dynamic error in measurements, cerebral oximetry should be interpreted as  
84 a trend rather than an absolute rScO<sub>2</sub> value.(14, 15) A network meta-analysis revealed that by  
85 maintaining a rScO<sub>2</sub> > 80% of baseline, NIRS was associated with protection against  
86 POCD/POD in cardiac surgery (pooled odds ratio 0.34; 95% CI 0.14 to 0.85).(16)

87 In order to perform cerebral oximetry monitoring, a pair of adhesive sensors are attached  
88 to the right and left sides of the forehead of the patient to measure the rScO<sub>2</sub> of the frontal  
89 cortex.(5) The frontal cortex receives blood supply from two branches of the internal carotid  
90 artery: the anterior and the middle cerebral artery.(17) There are some situations where the  
91 forehead of the patient is not accessible for sensor attachment—*e.g.*, neurosurgery involving the  
92 frontal area; a bispectral index (BIS) sensor already attached; or a wound to the forehead—so,  
93 alternate sites for sensor attachment are required. The temporal area was proposed as an  
94 alternative site for sensor attachment for measuring temporal lobe rScO<sub>2</sub>. The temporal lobe  
95 forms the cerebral cortex along with the occipital lobe, the parietal lobe, and the frontal lobe. The  
96 temporal lobe receives its blood supply from the internal carotid system and the vertebrobasilar  
97 artery.(18) Furthermore, the structures covering frontal and temporal lobes—the skin,  
98 subcutaneous fat, thin layer of muscle, and skull—are comparable; thus, the rScO<sub>2</sub> measured  
99 from the forehead and temporal area should provide comparable information.

100 The objective of this study was to assess the correlation and agreement of rScO<sub>2</sub> as  
101 measured using sensors attached to the forehead vs. the temporal area. The proposed criterion for  
102 an acceptable agreement was a bias of less than 5%, with the lower and upper limits of  
103 agreement being within 15% of the bias.

#### 104 **METHODS**

105 The study was approved on July 30, 2021 by the Khon Kaen University Ethics  
106 Committee for Human Research (HE641311). The study was registered on October 9, 2021 at  
107 ClinicalTrials.com (NCT05087836). The study was performed as per the Declaration of Helsinki  
108 and the ICH GCP, and all participants gave written informed consent before being recruited.

109 This was a prospective descriptive study. The sample size of 21 patients was based on the  
110 expected correlation coefficient of 0.6, an  $\alpha$ -value of 0.05, and a  $\beta$ -value of 0.2, with a 20% drop-  
111 out. The inclusion criteria were patients 1) of any sex; 2) age 18 or older; 3) undergoing elective  
112 cardiothoracic or vascular surgery at Srinagarind Hospital or Queen Sirikit Heart Center of the  
113 Northeast, Khon Kaen University, Khon Kaen, Thailand; and, 4) with an American Society of  
114 Anesthesiologists (ASA) classification II to III. The exclusion criteria were patients with 1) intra-  
115 cranial or carotid vascular disease; 2) previous surgery of face or brain; 3) abnormal anatomy or  
116 fibrosis of face; and/or 4) redo surgery.

117 All patients received standard cardiac anesthesia care. The intraoperative monitoring  
118 consisted of electrocardiogram, pulse oximetry, non-invasive blood pressure, temperature,  
119 capnography, anesthetic gas analyzer, and urine output. In addition, the rScO<sub>2</sub> was monitored  
120 using ForeSight sensors and a HemoSphere monitor (Edwards Lifesciences, Irvine, CA, USA).  
121 Each patient was monitored with 2 pairs of ForeSight sensors connected to the same  
122 HemoSphere monitor. The first pair (A1 and A2) were attached to the right and left forehead  
123 area, while the second pair (B1 and B2) were attached to the right and left temporal region of the  
124 patient (Figure 1). The proximal light source of the temporal sensor was located between the  
125 eyebrow's tail and the helix root outside the hairline, while the distal end of the deep detector  
126 was located upward below the hairline (Figure 2). The sensors were then secured by covering  
127 with Tegaderm films (3M, Minn, USA). The rScO<sub>2</sub> values from all sensors were recorded every  
128 20 seconds. After the operation, the data from all sensors were downloaded for analysis.

129 All patients received fentanyl 2-3  $\mu\text{g}\cdot\text{kg}^{-1}$  and midazolam 1-2 mg as premedication.  
130 Anesthesia was induced with propofol 2-3  $\text{mg}\cdot\text{kg}^{-1}$  or etomidate 0.3  $\text{mg}\cdot\text{kg}^{-1}$ . Endotracheal  
131 intubation was facilitated with cis-atracurium 0.2  $\text{mg}\cdot\text{kg}^{-1}$ . The ventilation was controlled using a

132 tidal volume of  $8 \text{ mL} \cdot \text{kg}^{-1}$  and a rate of 12-14 breaths  $\cdot \text{min}^{-1}$ , adjusted to maintain an end-tidal  
133  $\text{CO}_2$  close to 35 mmHg. The depth of anesthesia was maintained at 1 minimum alveolar  
134 concentration (MAC) using sevoflurane 1%-2% or desflurane 3%-6% with 50%  $\text{O}_2$  and air. The  
135 radial artery was cannulated and connected to a FloTrac transducer (Edwards Lifesciences,  
136 Irvine, CA, USA) linked to the HemoSphere monitor to measure invasive blood pressure, stroke  
137 volume variation (SVV), and cardiac index (CI). The internal jugular vein was cannulated and  
138 connected to another pressure transducer linked to the HemoSphere to measure the systemic  
139 vascular resistance index (SVRI). According to the institutional standard protocols, other specific  
140 anesthetic management regarding each surgical procedure was accomplished. Patient  
141 hemodynamics were optimized using the early goal-directed therapy (EGDT) protocol. The goals  
142 were: mean arterial pressure (MAP) 65-90 mmHg; urine output  $\geq 0.5 \text{ mL} \cdot \text{kg}^{-1} \cdot \text{h}^{-1}$ ;  $\text{SpO}_2 > 95\%$ ;  
143 and hematocrit  $\geq 30\%$ . For patients undergoing cardiac surgery with a cardiopulmonary bypass  
144 (CPB), the goals during CPB were MAP 50-75 mmHg, hematocrit 22%-25%,  $\text{PaO}_2$  150-200  
145 mmHg, and normocapnea. All patients received fluid and blood components to maintain a SVV  
146  $< 13\%$ , inotropic drugs to achieve a CI of  $2.2\text{-}4.0 \text{ L} \cdot \text{min}^{-1} \cdot \text{m}^{-2}$ , followed by vasoactive drugs to  
147 maintain a SVRI of  $1,500\text{-}2,500 \text{ dynes s}^{-1} \cdot \text{cm}^{-5} \cdot \text{m}^{-2}$ . Arterial blood gas and electrolytes were  
148 monitored hourly and corrected as required.

#### 149 **Statistical analysis**

150 Continuous data were presented as means  $\pm$  standard deviations (SD) or medians with an  
151 interquartile range, as appropriate. Categorical data were presented as numbers (%). Data from  
152 A1 vs. B1 and A2 vs. B2 were assessed for correlation coefficients. The agreements between  
153 data from A1 vs. B1 and A2 vs. B2 were evaluated using the Bland-Altman plot and analysis.  
154 The Bland-Altman analysis is a graphical method used to determine whether two methods of

155 measurement can be used interchangeably by plotting the calculated mean difference between  
156 the two methods of measurement (the ‘bias’) with 95% limits of agreement calculated from  
157  $\pm 1.96$  SD of the mean difference.(19) All statistical analyses were performed using MedCalc  
158 version 20.027.

## 159 RESULTS

160 Twenty-one patients were recruited between November 1, 2021 and March 10, 2022.  
161 There were two drop-outs (ID10 and ID20) due to sensor errors resulting in incomplete data.  
162 Therefore, the data (14,364 sets) from 19 patients were analyzed. A summary of the  
163 demographic and clinical data for the patients is presented in Table 1. Nearly half the participants  
164 were male, and the mean age of all participants was 57.3 years. The types of operation included  
165 cardiac, thoracic, video-assisted thoracoscopic, and endovascular stent surgery. The descriptive  
166 characteristics of data from A1, B1, A2, and B2 are presented in Table 2. The data from A1 vs.  
167 B1 and A2 vs. B2 show a moderate positive correlation (Table 3).

168 According to the Bland-Altman analysis, the respective bias of A1 vs. B1 and A2 vs. B2  
169 was -2.3% (95% CI -2.5 to -2.2;  $P < 0.0001$ ) and 0.7% (95% CI 0.6 to 0.8;  $P < 0.0001$ ). The  
170 respective lower and upper limits of agreement for A1 vs. B1 and A2 vs. B2 are presented in  
171 Table 4. The Bland-Altman plots for A1 vs. B1 and A2 vs. B2 are presented in Figures 3 and 4.

172 The results of the Bland-Altman subgroup analysis for sex (male vs. female), age ( $< 60$  y  
173 vs.  $\geq 60$  y), and anesthetic time ( $< 300$  min vs.  $\geq 300$  min) revealed comparable biases—less than  
174  $\pm 5\%$ —and limits of agreements—within  $\pm 15\%$  of the biases (Table 5).

175

176

177 **DISCUSSION**

178           The results of the present study showed that the rScO<sub>2</sub> values measured from the  
179 temporal area slightly but significantly differed from the frontal area. NIRS sensors attached at  
180 different areas of the same subject yielded varying rScO<sub>2</sub> values. Kishi et al(20) investigated the  
181 effects of sensor location on rScO<sub>2</sub> as measured using a cerebral oximeter INVOS 4100 and  
182 applying the sensors to the right forehead (R), 1 cm lateral to R (R1), the left forehead (L), 1 cm  
183 lateral to L (L1), and the center of the forehead (C). They found that the rScO<sub>2</sub> values from R1  
184 ( $58 \pm 11\%$ ) and L1 ( $59 \pm 10\%$ ) were significantly lower than those from R ( $61 \pm 10\%$ ) and L ( $61$   
185  $\pm 11\%$ ), while the values from C ( $64 \pm 12\%$ ) were significantly higher than those at the other  
186 sites. Cho et al(21) revealed that the NIRS sensors located at the upper forehead, compared with  
187 the lower forehead, yielded lower rScO<sub>2</sub> values. The explanation is that the difference in  
188 structures over the brain and the depth of the brain surface affect the optical path length and light  
189 scatter. The absolute rScO<sub>2</sub> values, thus, may not reflect cerebral oxygenation status; instead, the  
190 change in rScO<sub>2</sub> values used as a trend monitor has more clinical relevance. Thus, the rScO<sub>2</sub>  
191 values derived from different scalp areas, with acceptable agreement, may be used for trend  
192 monitoring of change in rScO<sub>2</sub> values.

193           The current study revealed that the rScO<sub>2</sub> values—as measured by sensors on the frontal  
194 and temporal areas—demonstrated moderate correlation with sufficiently good agreement. The  
195 biases of data from A1 vs. B1 and A2 vs. B2 were marginal—less than 3%—with high  
196 precision—95% CIs range within 0.3%—indicating excellent agreement and confirming that the  
197 temporal area is an alternative site if the NIRS sensors cannot be attached to the forehead. The  
198 acceptable lower and upper limits of agreement were defined before the study to be within 15%  
199 of the bias because these limits should be narrower than the 20% reduction which is the criteria

200 of cerebral desaturation. The lower and upper limits of agreement of A1 vs. B1 and A2 vs. B2  
201 were approximately  $\pm 15\%$  of the bias, which is clinically acceptable since this margin is within  
202 the predefined criteria.

203 The data from 2 patients (ID10 and ID20) were not included in the final analysis due to  
204 sensor error. After confirming the correctness of the sensor attachment, the HemoSphere monitor  
205 continued to report incomplete data.

206 Since many factors may affect rScO<sub>2</sub> values, e.g., sex, age, and anesthetic time(20, 22), a  
207 subgroup analysis was performed and found that these factors did not affect the reading of rScO<sub>2</sub>  
208 values in the current study. The biases and limits of agreements of these subgroups were  
209 comparable to the primary outcomes. The results of the current study differ from those of Kishi  
210 et al(20) who found a significant negative correlation between age and rScO<sub>2</sub> values. This may  
211 be due to differences in the age range of the subjects (7-89 y in the study of Kishi et al vs. 21-78  
212 y in the current study).

213 The explanation for the agreement of rScO<sub>2</sub> values reading from the frontal and temporal  
214 areas is that both the frontal and temporal lobes are parts of the cerebral cortex with common  
215 blood supplies and covering structures, thus the rScO<sub>2</sub> values measured from these areas should  
216 be comparable. Even though the differences were statistically significant, they are so trivial that  
217 there is no clinical relevance. Since the baseline rScO<sub>2</sub> values in cardiac surgery have high  
218 variability,(22) the trend in values is more clinically relevant.(4)

219 Many factors affect rScO<sub>2</sub> value including arterial carbon dioxide, cardiac output, arterial  
220 blood pressure, hemoglobin concentration, neural excitation, and anesthesia depth.(4) These  
221 factors should be optimized and taken into account to correct cerebral desaturation. Skin color

222 does not affect the rScO<sub>2</sub> reading.(4) Scalp hair follicles (SHF) have strong impact on NIRS  
223 measurement by decreasing the detected light intensity signal by 15% - 80% resulting in a  
224 miscalculation of rScO<sub>2</sub> by 11.7% to 292.2% linearly at SHF density varied from 1% to 11% in  
225 Asian human.(23) The authors tried to attach the NIRS sensors to the shaved occiput and the  
226 temporal area including the hairline and found no rScO<sub>2</sub> reading on the HemoSphere monitor.  
227 Thus, the NIRS sensor should not be attached to the scalp area where there are hair follicles.

228 POCD is a permanent drop in cognitive function after surgery that interferes with quality  
229 of life of the patients. Cerebral desaturation was identified to be associated with POCD.(24) The  
230 significant risk factors for POCD are increasing age and maximum percentage drop in rScO<sub>2</sub> >  
231 11%.(25) A network meta-analysis showed that maintaining rScO<sub>2</sub> > 80% of the baseline value  
232 was protective against POCD/POD in cardiac surgery patients.(16) Thus, rScO<sub>2</sub> monitoring for  
233 cerebral oxygenation during cardiac surgery is essential. The standard site for NIRS sensor  
234 attachment is the forehead, but an alternative site is needed in case the forehead is not available.

235 The temporal area was chosen as an alternative to the forehead because this is the only  
236 scalp area that has no hair follicles with similar depth from scalp to brain tissue. Furthermore,  
237 these two areas reflect the frontal and temporal lobes, which form the cerebral cortex with a  
238 common blood supply.(17, 18) The near-infrared spectrum of NIRS can penetrate the tissue up to  
239 2.5 cm(6); thus, the sensors attached to these areas can accurately measure the oxygen saturation  
240 of the underlying brain tissue resulting in comparable rScO<sub>2</sub> values.

241 The Bland-Altman analysis was selected to compare the rScO<sub>2</sub> values measured from  
242 sensors attached to the frontal and temporal areas because this method quantifies agreement  
243 between two quantitative measurements by assessing the bias between the mean differences and  
244 constructing limits of agreement.(19) The method does not judge whether those biases and limits

245 are acceptable or not. Acceptable limits must be defined before the study based on clinical  
246 essentiality. The biases and limits of agreement of this study remain within the proposed criteria,  
247 indicating that the temporal area may be used as a site for attaching NIRS sensors if the frontal  
248 area is not feasible.

#### 249 **Limitations**

250 A limited number of patients undergoing elective cardiothoracic or vascular surgery at a  
251 single tertiary center were included, so the results may not be generalizable to other contexts. In  
252 addition, this study used the ForeSight Elite sensors and HemoSphere monitoring that again may  
253 not be generalizable to other types of monitors. Therefore, a further multi-center study with a  
254 larger sample size with more types of monitors is recommended to validate our findings.

#### 255 **CONCLUSIONS**

256 The rScO<sub>2</sub> values measured from the sensors attached to the frontal and temporal areas  
257 showed moderate correlation. The biases and limits of agreement remained within the predefined  
258 criteria indicating a good agreement. The temporal area may thus be used as an alternative site to  
259 the frontal area for cerebral oximetry monitoring. The identification of cerebral desaturation  
260 should be based on the trend—in changes to rScO<sub>2</sub> values from baseline—rather than the  
261 absolute rScO<sub>2</sub> values.

262

#### 263 **ACKNOWLEDGMENTS**

264 The authors thank Mr. Bryan Roderick Hamman under the aegis of the Publication  
265 Clinic, Khon Kaen University, Thailand, for assistance with the English-language presentation of  
266 the manuscript.

267 **REFERENCES**

- 268 1. Pollard V, Prough DS, DeMelo AE, Deyo DJ, Uchida T, Stoddart HF. Validation in  
269 volunteers of a near-infrared spectroscope for monitoring brain oxygenation in vivo. *Anesth*  
270 *Analg* 1996;82(2):269-77.
- 271 2. Wahr JA, Tremper KK, Samra S, Delpy DT. Near-infrared spectroscopy: theory and  
272 applications. *J Cardiothorac Vasc Anesth* 1996;10(3):406-18.
- 273 3. Owen-Reece H, Smith M, Elwell CE, Goldstone JC. Near infrared spectroscopy. *Br J*  
274 *Anaesth* 1999;82(3):418-26.
- 275 4. Vegh T. Cerebral Oximetry in General Anaesthesia. *Turk J Anaesthesiol Reanim*  
276 2016;44(5):247-9.
- 277 5. Hogue CW, Levine A, Hudson A, Lewis C. Clinical Applications of Near-infrared  
278 Spectroscopy Monitoring in Cardiovascular Surgery. *Anesthesiology* 2021;134(5):784-91.
- 279 6. Stepan J, Hogue CW, Jr. Cerebral and tissue oximetry. *Best Pract Res Clin Anaesthesiol*  
280 2014;28(4):429-39.
- 281 7. Hogue CW, Jr., Palin CA, Arrowsmith JE. Cardiopulmonary bypass management and  
282 neurologic outcomes: an evidence-based appraisal of current practices. *Anesth Analg*  
283 2006;103(1):21-37.
- 284 8. Gottesman RF, McKhann GM, Hogue CW. Neurological complications of cardiac  
285 surgery. *Semin Neurol* 2008;28(5):703-15.
- 286 9. Selnes OA, Gottesman RF, Grega MA, Baumgartner WA, Zeger SL, McKhann GM.  
287 Cognitive and neurologic outcomes after coronary-artery bypass surgery. *N Engl J Med*  
288 2012;366(3):250-7.

- 289 10. Krause M, Morabito JE, Mackensen GB, Perry TE, Bartels K. Current Neurologic  
290 Assessment and Neuroprotective Strategies in Cardiac Anesthesia: A Survey to the Membership  
291 of the Society of Cardiovascular Anesthesiologists. *Anesth Analg* 2020;131(2):518-26.
- 292 11. Zheng F, Sheinberg R, Yee MS, Ono M, Zheng Y, Hogue CW. Cerebral near-infrared  
293 spectroscopy monitoring and neurologic outcomes in adult cardiac surgery patients: a systematic  
294 review. *Anesth Analg* 2013;116(3):663-76.
- 295 12. Murkin JM, Arango M. Near-infrared spectroscopy as an index of brain and tissue  
296 oxygenation. *Br J Anaesth* 2009;103 Suppl 1:i3-13.
- 297 13. Deschamps A, Hall R, Grocott H, Mazer CD, Choi PT, Turgeon AF, et al. Cerebral  
298 Oximetry Monitoring to Maintain Normal Cerebral Oxygen Saturation during High-risk Cardiac  
299 Surgery: A Randomized Controlled Feasibility Trial. *Anesthesiology* 2016;124(4):826-36.
- 300 14. Henson LC, Calalang C, Temp JA, Ward DS. Accuracy of a cerebral oximeter in healthy  
301 volunteers under conditions of isocapnic hypoxia. *Anesthesiology* 1998;88(1):58-65.
- 302 15. Bickler PE, Feiner JR, Rollins MD. Factors affecting the performance of 5 cerebral  
303 oximeters during hypoxia in healthy volunteers. *Anesth Analg* 2013;117(4):813-23.
- 304 16. Ortega-Loubon C, Herrera-Gomez F, Bernuy-Guevara C, Jorge-Monjas P, Ochoa-  
305 Sangrador C, Bustamante-Munguira J, et al. Near-Infrared Spectroscopy Monitoring in Cardiac  
306 and Noncardiac Surgery: Pairwise and Network Meta-Analyses. *J Clin Med* 2019;8(12):2208.
- 307 17. El-Baba RM, Schury MP. Neuroanatomy, Frontal Cortex. *StatPearls*. Treasure Island  
308 (FL)2022.
- 309 18. Patel A, Biso G, Fowler JB. Neuroanatomy, Temporal Lobe. *StatPearls*. Treasure Island  
310 (FL)2022.

- 311 19. Giavarina D. Understanding Bland Altman analysis. *Biochem Med (Zagreb)*  
312 2015;25(2):141-51.
- 313 20. Kishi K, Kawaguchi M, Yoshitani K, Nagahata T, Furuya H. Influence of patient  
314 variables and sensor location on regional cerebral oxygen saturation measured by INVOS 4100  
315 near-infrared spectrophotometers. *J Neurosurg Anesthesiol* 2003;15(4):302-6.
- 316 21. Cho AR, Kwon JY, Kim C, Hong JM, Kang C. Effect of sensor location on regional  
317 cerebral oxygen saturation measured by INVOS 5100 in on-pump cardiac surgery. *J Anesth*  
318 2017;31(2):178-84.
- 319 22. Chan MJ, Chung T, Glassford NJ, Bellomo R. Near-Infrared Spectroscopy in Adult  
320 Cardiac Surgery Patients: A Systematic Review and Meta-Analysis. *J Cardiothorac Vasc Anesth*  
321 2017;31(4):1155-65.
- 322 23. Fang X, Pan B, Liu W, Wang Z, Li T. Effect of Scalp Hair Follicles on NIRS  
323 Quantification by Monte Carlo Simulation and Visible Chinese Human Dataset. *IEEE Photonics*  
324 *Journal* 2018;10(5):3901110.
- 325 24. Casati A, Fanelli G, Pietropaoli P, Proietti R, Tufano R, Danelli G, et al. Continuous  
326 monitoring of cerebral oxygen saturation in elderly patients undergoing major abdominal surgery  
327 minimizes brain exposure to potential hypoxia. *Anesth Analg* 2005;101(3):740-7.
- 328 25. Lin R, Zhang F, Xue Q, Yu B. Accuracy of regional cerebral oxygen saturation in  
329 predicting postoperative cognitive dysfunction after total hip arthroplasty: regional cerebral  
330 oxygen saturation predicts POCD. *J Arthroplasty* 2013;28(3):494-7.

331

332

**Table 1** (on next page)

Demographic and clinical data of the patients (n = 19)

1 **Table 1 Demographic and clinical data of the patients (n = 19)**

	Value
<b>Male</b>	10 (52.6)
<b>Age (y)</b>	57.3 ± 16.6
<b>Body weight (kg)</b>	62.0 ± 10.8
<b>Height (cm)</b>	165.2 ± 8.0
<b>ASA classification</b>	
<b>II</b>	9 (47.4)
<b>III</b>	10 (52.6)
<b>Underlying diseases</b>	
<b>Hypertension</b>	7 (36.8)
<b>Diabetes mellitus</b>	2 (10.5)
<b>Myocardial ischemia</b>	3 (15.8)
<b>Dyslipidemia</b>	2 (10.5)
<b>Chronic kidney disease</b>	3 (15.8)
<b>Type of operations</b>	
<b>CABG</b>	3 (15.8)
<b>OPCAB</b>	1 (5.3)
<b>Valvular</b>	3 (15.8)
<b>Valvular and aortic root</b>	2 (10.5)
<b>Thoracic</b>	3 (15.8)
<b>VATs</b>	4 (21.0)
<b>Endovascular stent</b>	2 (10.5)
<b>Cone operation</b>	1 (5.3)
<b>Mean arterial pressure (mmHg)</b>	87.0 ± 16.6
<b>Heart rate (bpm)</b>	71.3 ± 13.1
<b>SpO<sub>2</sub> (%)</b>	99.0 ± 2.9
<b>End-tidal CO<sub>2</sub></b>	33.1 ± 4.4
<b>Temperature (°C)</b>	35.8 ± 0.5
<b>Hemoglobin (g/dL)</b>	12.3 ± 2.0
<b>P/F ratio (mmHg)</b>	369.3 ± 121.9
<b>Blood sugar (mg/dL)</b>	126.8 ± 41.4
<b>Anesthesia time (min)</b>	314.3 ± 110.9

2

3 Data are presented as mean ± SD or number (%)

**Table 2** (on next page)

Descriptive data of A1, B1, A2, B2 (19 patients)

1 **Table 2** Descriptive data of A1, B1, A2, B2 (19 patients)

	A1	B1	A2	B2
<b>N (sample sets)</b>	14,364	14,364	14,364	14,364
<b>Mean (%)</b>	71.4	73.7	72.1	71.3
<b>Median (%)</b>	71.0	74.0	71.0	71.0
<b>Standard deviation (%)</b>	7.0	9.9	7.2	8.9
<b>Minimum (%)</b>	48	48	50	47
<b>Maximum (%)</b>	89	98	94	95

2

3

**Table 3** (on next page)

Correlation between A1 vs. B1 and A2 vs.-B2 (19 patients)

1 **Table 3** Correlation between A1 vs. B1 and A2 vs.-B2 (19 patients)

	A1-B1	A2-B2
<b>Sample sets</b>	14,364	14,364
<b>Correlation coefficient r</b>	0.627	0.548
<b>Significant level</b>	$P < 0.0001$	$P < 0.0001$
<b>95% Confidence interval for r</b>	0.617 to 0.637	0.536 to 0.559

2

3

**Table 4**(on next page)

Bland-Altman statistics of A1 vs. B1 and A2 vs. B2 (19 patients)

1 **Table 4** Bland-Altman statistics of A1 vs. B1 and A2 vs. B2 (19 patients)

	A1-B1 (n = 14,364)	A2-B2 (n = 14,364)
<b>Bias (%)</b>	-2.3 (95% CI -2.5 to -2.2)	0.7 (95% CI 0.6 to 0.8)
<b><i>P</i> value</b>	< 0.0001	< 0.0001
<b>Lower limit of agreement (%)</b>	-17.5 (95% CI -17.7 to -17.3)	-14.6 (95% CI -14.8 to -14.4)
<b>Upper limit of agreement (%)</b>	12.8 (95% CI 12.6 to 13.0)	16.0 (95% CI 15.8 to 16.3)

2

3

**Table 5** (on next page)

Bland-Altman statistics of A1 vs. B1 and A2 vs. B2: subgroup analysis

1 **Table 5** Bland-Altman statistics of A1 vs. B1 and A2 vs. B2: subgroup analysis

		A1-B1	A2-B2
<b>Sex:</b> <b>Male</b> (n = 5,686)	Bias (%)	-1.8 (95% CI -2.0 to -1.7)	4.7 (95% CI 4.5 to 4.9)
	<i>P</i> value	< 0.0001	< 0.0001
	Lower limit of agreement (%)	-15.0 (95% CI -15.3 to -14.7)	-7.9 (95% CI -8.2 to -7.6)
	Upper limit of agreement (%)	11.3 (95% CI 11.0 to 11.6)	17.3 (95% CI 17.0 to 17.6)
<b>Sex:</b> <b>Female</b> (n = 8,678)	Bias (%)	-2.7 (95% CI -2.8 to -2.5)	-1.9 (95% CI -2.0 to -1.7)
	<i>P</i> value	< 0.0001	< 0.0001
	Lower limit of agreement (%)	-17.5 (95% CI -17.8 to -17.2)	-16.9 (95% CI -17.1 to -16.6)
	Upper limit of agreement (%)	12.3 (95% CI 12.0 to 12.6)	13.1 (95% CI 12.8 to 13.4)
<b>Age:</b> <b>&lt; 60 y</b> (n = 7,509)	Bias (%)	-0.2 (95% CI -0.4 to -0.0)	4.9 (95% CI 4.8 to -5.1)
	<i>P</i> value	0.0338	< 0.0001
	Lower limit of agreement (%)	-15.1 (95% CI -15.4 to -14.8)	-8.1 (95% CI -8.4 to -7.9)
	Upper limit of agreement (%)	14.1 (95% CI 13.8 to 14.4)	18.0 (95% CI 17.7 to 18.2)
<b>Age:</b> <b>≥ 60 y</b> (n = 6,855)	Bias (%)	-4.7 (95% CI -4.8 to -4.5)	-3.9 (95% CI -4.0 to -3.7)
	<i>P</i> value	< 0.0001	< 0.0001
	Lower limit of agreement (%)	-18.3 (95% CI -18.6 to -18.0)	-16.4 (95% CI -16.6 to -16.1)
	Upper limit of agreement (%)	8.9 (95% CI 8.7 to 9.2)	8.6 (95% CI 8.4 to 8.9)
<b>Anesthetic time:</b> <b>&lt; 300 min</b> (n = 3,648)	Bias (%)	-1.2 (95% CI -1.4 to -1.0)	1.8 (95% CI 1.5 to 2.0)
	<i>P</i> value	< 0.0001	< 0.0001
	Lower limit of agreement (%)	-14.7 (95% CI -15.1 to -14.3)	-11.4 (95% CI -11.8 to -11.0)
	Upper limit of agreement (%)	12.4 (95% CI 12.0 to 12.7)	14.9 (95% CI 14.5 to 15.3)
<b>Anesthetic time:</b>	Bias (%)	-2.7 (95% CI -2.9 to -2.6)	0.4 (95% CI 0.2 to 0.5)

<b>≥ 300 min</b> (n = 10,716)			
<i>P</i> value	< 0.0001		< 0.0001
Lower limit of agreement (%)	-17.3 (95% CI -17.6 to -17.1)	-14.5 (95% CI -14.8 to -14.2)	
Upper limit of agreement (%)	11.9 (95% CI 11.6 to 12.2)	15.4 (95% CI 15.2 to 15.7)	

2

3

# Figure 1

Locations of two pairs of ForeSight sensors

(A) A1 and A2 are the locations for the first pair of sensors on the right and left forehead areas. (B) B1 and B2 are the locations for the second pair of sensors on the right and left temporal areas.



## Figure 2

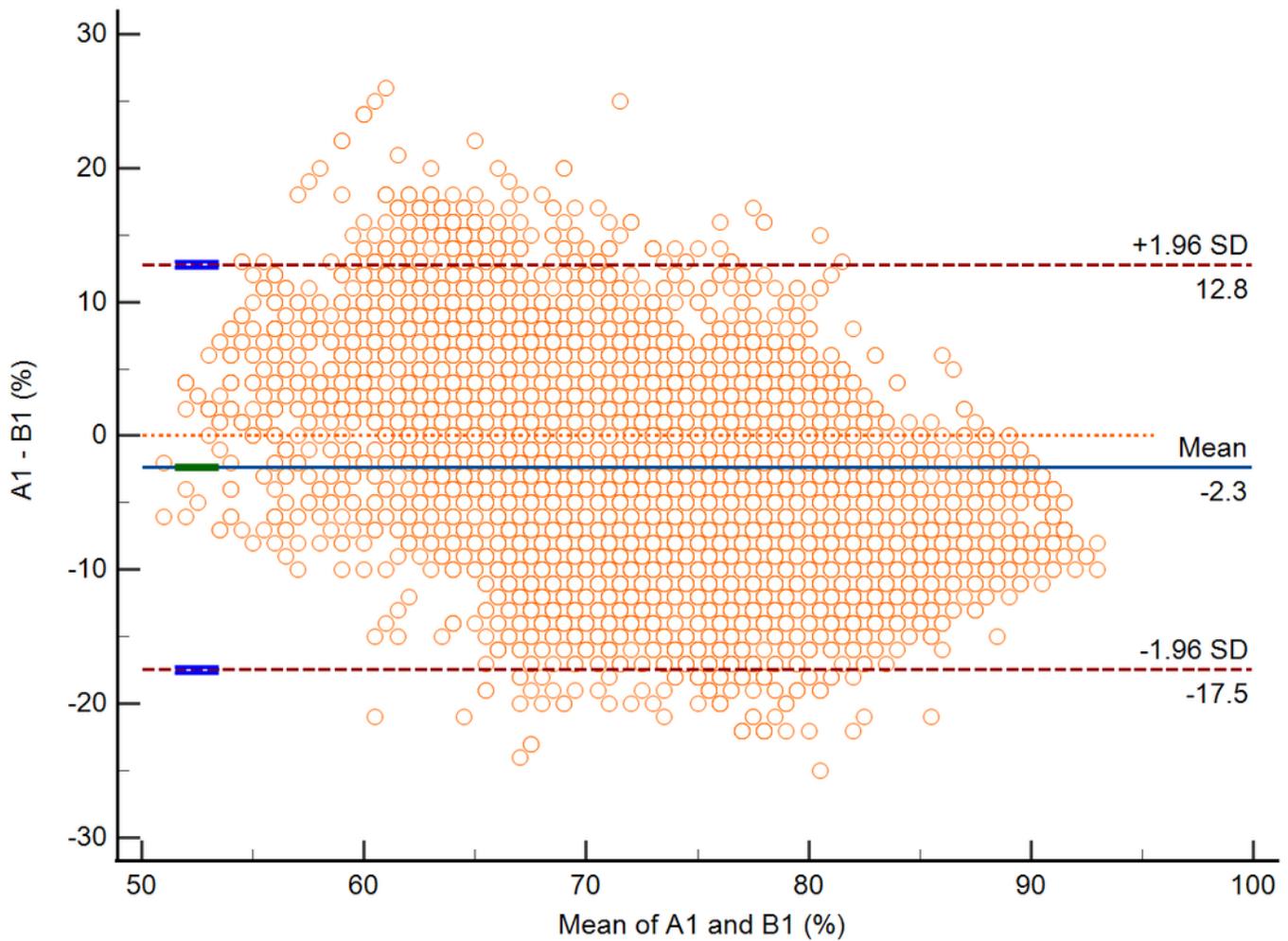
The location of the temporal sensor

The proximal light source was located between the eyebrow's tail and the helix root outside the hairline, while the distal end of the deep detector was located upward below the hairline.



## Figure 3

The Bland-Altman plot of data from A1 vs. B1



## Figure 4

The Bland-Altman plot of data from A2 vs. B2

