

Aus der Kinderklinik und Kinderpoliklinik
im Dr. von Haunerschen Kinderspital
der Ludwig-Maximilians-Universität München

Direktor: Prof. Dr. Dr. Christoph Klein

**Verlaufsbeschreibung der regionalen zerebralen
Gewebssättigung bei reifen und gesunden Neugeborenen
unmittelbar nach Geburt**

Dissertation zum Erwerb des Doktorgrades der Medizin an der Medizinischen
Fakultät der Ludwig-Maximilians-Universität zu München

vorgelegt von:

Dr. med. univ. Elisabeth Beckenbach (geb. Kratky)

aus (Geburtsort): Graz, Österreich

Jahr: 2013

Mit Genehmigung der Medizinischen Fakultät der Universität München

Berichterstatter: PD Dr. Andreas Flemmer.....

Mitberichterstatter: Priv.Doz. Dr. med. Andreas Bender.....

Priv.-Doz. Dr. med. Steffen Berweck

Dekan: Prof. Dr. med. Dr. h.c. M. Reiser, FACR, FRCR.....

Tag der mündlichen Prüfung: 21.11.2013.....

Eidesstattliche Erklärung

Ich erkläre hiermit an Eides statt, dass ich die vorliegende Dissertation mit dem Thema „Verlaufsbeschreibung der regionalen zerebralen Gewebssättigung bei reifen und gesunden Neugeborenen unmittelbar nach Geburt“ selbständig verfasst, mich außer der angegebenen keiner weiteren Hilfsmittel bedient und

alle Erkenntnisse, die aus dem Schrifttum ganz oder annähernd übernommen sind, als solche kenntlich gemacht und nach ihrer Herkunft unter Bezeichnung der Fundstelle einzeln nachgewiesen habe.

München, am

Unterschrift:

(Elisabeth Beckenbach)

Inhaltsverzeichnis

INHALTSVERZEICHNIS.....	4
ABKÜRZUNGEN	5
LISTE DER PUBLIKATIONEN.....	6
1.1 PUBLIKATION I:.....	6
1.2 PUBLIKATION II.....	6
DEKLARATION KOAUTOREN	7
1.3 DEKLARATION KOAUTOREN PUBLIKATION I.....	7
1.4 DEKLARATION KOAUTOREN PUBLIKATION II	10
1.5 ZUSAMMENFASSUNG PUBLIKATION I.....	12
1.6 ZUSAMMENFASSUNG PUBLIKATION II.....	13
EINLEITUNG	14
PUBLIKATION I.....	16
PUBLIKATION II	25
DISKUSSION	30
DANKSAGUNG.....	33
LEBENSLAUF	34
LITERATURVERZEICHNIS.....	37

Abkürzungen

FTOE	Fractional tissue oxygen extraction
HR	Herzfrequenz (heart rate)
LM	Lebensminute
NG	Neugeborene(s)
NIRS	Nahinfrarotspektroskopie
rSO ₂ brain	regionale zerebrale Sättigung
SaO ₂	arterielle Sättigung (nicht-invasiv gemessen)

Liste der Publikationen

1.1 **Publikation I:**

Titel: ***Regional cerebral oxygen saturation in newborn infants in the first 15 min of life after vaginal delivery***

Autoren: Kratky E, Pichler G, Rehak T, Avian A, Pocivalnik M, Müller W, Urlesberger B.

Forschungsgruppe: Division of Neonatology, Department of Pediatrics, Medical University of Graz, Austria.

Erschienen: ***Physiological Measurement***, 2012 Jan;33(1):95-102. doi: 10.1088/0967-3334/33/1/95.

1.2 **Publikation II**

Titel: ***Regional Oxygen Saturation of the Brain during Birth Transition of Term Infants: Comparison between Elective Cesarean and Vaginal Deliveries***

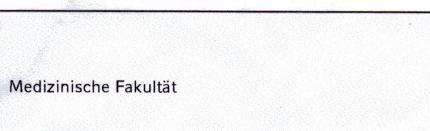
Autoren: Urlesberger B, Kratky E, Rehak T, Pocivalnik M, Avian A, Czihak J, Müller W, Pichler G.

Forschungsgruppe: Research Unit for Cerebral Development and Oximetry, Division of Neonatology, Department of Pediatrics, Medical University of Graz, Austria.

Erschienen: ***The Journal of Pediatrics***, September 2011, 159(3):404-8. doi: 10.1016/j.jpeds.2011.02.030., Epub 2011 Apr 9.

Deklaration Koautoren

1.3 Deklaration Koautoren Publikation I



Kumulative Dissertation

Bestätigung

gem. § 4a Abs. 3 und 5 Promotionsordnung für die Promotion zum Dr. med., Dr. med. dent und Dr. rer. biol. hum.
und
gem. § 7 Abs. 4 Promotionsordnung für die Promotion zum Dr. rer. nat. an der Medizinischen Fakultät

Elisabeth Beckenbach (geborene Kratky)

Doktorand

"Regional cerebral oxygen saturation in newborn infants in the first 15 min of life after vaginal delivery"

Titel der Publikation

Hiermit bestätige ich, dass keiner der zur Promotion eingereichten Fachartikel Gegenstand einer anderen (laufenden oder abgeschlossenen) Dissertation ist.

Elisabeth Beckenbach

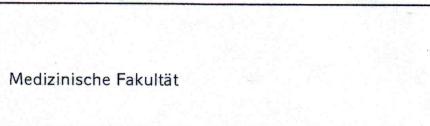
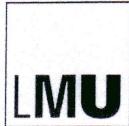
Unterschrift Doktorand

Folgende Ko-Autoren bestätigen mit ihrer Unterschrift

- ihren Arbeitsanteil (Inhalt und Umfang) an den eingereichten Veröffentlichungen,
- ihr Einverständnis zur Einreichung der Publikationen sowie,
- dass der jeweilige eingereichte Fachartikel nicht Gegenstand einer anderen (laufenden oder abgeschlossenen) Dissertation ist.

Name Ko-Autor	Arbeitsanteil (Inhalt und Umfang)	Unterschrift Ko-Autor
<u>1. Pichler Gerhard</u>	<u>Konzeption, Planung</u> <u>Durchführbarkeit</u> <u>Textbearbeitung, Einreichung</u> <u>Ethikautrag</u>	<u>Gerhard Pichler</u>
<u>2. Rehak Thomas</u>	<u>Statistik, Grafikgestaltung</u>	<u>Thomas Rehak</u>
<u>3. Arian Alexander</u>	<u>Messdatenerhebung</u> <u>Geräte einweisung</u>	<u>Arian Alexander</u>
<u>4. Pacivalnik Meagan</u>	<u>Konzeption, Textbearbeitung</u> <u>Korrekturen</u>	<u>Meagan Pacivalnik</u>
<u>5. Uhlendorfer Blumdt</u>		<u>Uhlendorfer Blumdt</u>

weitere Autoren bitte auf ein gesondertes Blatt



Kumulative Dissertation

Bestätigung

gem. § 4a Abs. 3 und 5 Promotionsordnung für die Promotion zum Dr. med., Dr. med. dent und Dr. rer. biol. hum.
und
gem. § 7 Abs. 4 Promotionsordnung für die Promotion zum Dr. rer. nat. an der Medizinischen Fakultät

Elisabeth Beckenbach (geborene Kratky)

Doktorand

"Regional cerebral oxygen saturation in newborn infants in the first 15 min of life after vaginal delivery"

Titel der Publikation

Hiermit bestätige ich, dass keiner der zur Promotion eingereichten Fachartikel Gegenstand einer anderen (laufenden oder abgeschlossenen) Dissertation ist.

Elisabeth Beckenbach

Unterschrift Doktorand

Folgende **Ko-Autoren** bestätigen mit ihrer Unterschrift

- ihren Arbeitsanteil (Inhalt und Umfang) an den eingereichten Veröffentlichungen,
- ihr Einverständnis zur Einreichung der Publikationen sowie,
- dass der jeweilige eingereichte Fachartikel nicht Gegenstand einer anderen (laufenden oder abgeschlossenen) Dissertation ist.

Name Ko-Autor

Arbeitsanteil (Inhalt und Umfang)

Unterschrift Ko-Autor

J. Müller Wilhelm

Konzeption, Supervision,
Interdisziplinäre Zusammenarbeit

Ulrich

2.

3.

4.

5.

weitere Autoren bitte auf ein gesondertes Blatt



LUDWIG-
MAXIMILIANS-
UNIVERSITÄT
MÜNCHEN

Medizinische Fakultät



Kumulative Dissertation

Bestätigung

gem. § 4a Abs. 3 und 5 Promotionsordnung für die Promotion zum Dr. med., Dr. med. dent und Dr. rer. biol. hum.
und
gem. § 7 Abs. 4 Promotionsordnung für die Promotion zum Dr. rer. nat. an der Medizinischen Fakultät

Elisabeth Beckenbach (geborene Kratky)

Doktorand

"Regional cerebral oxygen saturation in newborn infants in the first 15 min of life after vaginal delivery"

Titel der Publikation

Hiermit bestätige ich, dass keiner der zur Promotion eingereichten Fachartikel Gegenstand einer anderen (laufenden oder abgeschlossenen) Dissertation ist.

Elisabeth Beckenbach

Unterschrift Doktorand

Folgende **Ko-Autoren** bestätigen mit ihrer Unterschrift

- ihren Arbeitsanteil (Inhalt und Umfang) an den eingereichten Veröffentlichungen,
- ihr Einverständnis zur Einreichung der Publikationen sowie,
- dass der jeweilige eingereichte Fachartikel nicht Gegenstand einer anderen (laufenden oder abgeschlossenen) Dissertation ist.

Name Ko-Autor

Arbeitsanteil (Inhalt und Umfang)

Unterschrift Ko-Autor

1. Czihak Johanna

*Interdisziplinäre Zusammenarbeit
Teamarbeiten im Vereinsraum
Kolaboration mit Patienten*

Czihak X

2. _____

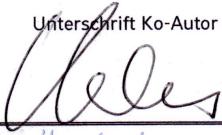
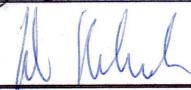
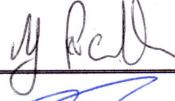
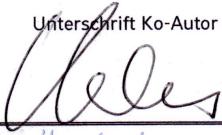
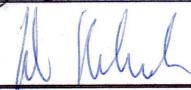
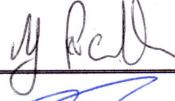
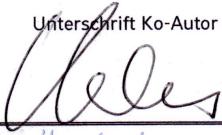
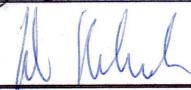
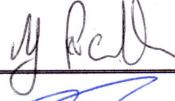
3. _____

4. _____

5. _____

weitere Autoren bitte auf ein gesondertes Blatt

1.4 Deklaration Koautoren Publikation II

	LUDWIG-MAXIMILIANS-UNIVERSITÄT MÜNCHEN	Medizinische Fakultät																			
Kumulative Dissertation																					
Bestätigung gem. § 4a Abs. 3 und 5 Promotionsordnung für die Promotion zum Dr. med., Dr. med. dent und Dr. rer. biol. hum. und gem. § 7 Abs. 4 Promotionsordnung für die Promotion zum Dr. rer. nat. an der Medizinischen Fakultät																					
<p>Elisabeth Beckenbach (geborene Kratky) Doktorand</p> <p>"Regional Oxygen Saturation of the Brain during Birth transition of Term Infants: Comparison between Elective and Vaginal Deliveries" Titel der Publikation</p> <p>Hiermit bestätige ich, dass keiner der zur Promotion eingereichten Fachartikel Gegenstand einer anderen (laufenden oder abgeschlossenen) Dissertation ist.</p> <p> Unterschrift Doktorand</p> <p>Folgende Ko-Autoren bestätigen mit ihrer Unterschrift</p> <table border="1"><thead><tr><th>Name Ko-Autor</th><th>Arbeitsanteil (Inhalt und Umfang)</th><th>Unterschrift Ko-Autor</th></tr></thead><tbody><tr><td>1. Ullesberger Berndt</td><td>Konzeption, Projektleiter Textverfassung</td><td></td></tr><tr><td>2. Rehak Thomas</td><td>Datenantrag, Konzeption</td><td></td></tr><tr><td>3. Pocivalnik Klimjan</td><td>Datenerhebung, Aggruppieren der Studienteilnehmer</td><td></td></tr><tr><td>4. Arvan Alexander</td><td>Statistik Technische Verarbeitung</td><td></td></tr><tr><td>5. Pichler Gerhard</td><td>Datenerhebung, Konzeption Textverfassung</td><td></td></tr></tbody></table> <p>weitere Autoren bitte auf ein gesondertes Blatt</p>				Name Ko-Autor	Arbeitsanteil (Inhalt und Umfang)	Unterschrift Ko-Autor	1. Ullesberger Berndt	Konzeption, Projektleiter Textverfassung		2. Rehak Thomas	Datenantrag, Konzeption		3. Pocivalnik Klimjan	Datenerhebung, Aggruppieren der Studienteilnehmer		4. Arvan Alexander	Statistik Technische Verarbeitung		5. Pichler Gerhard	Datenerhebung, Konzeption Textverfassung	
Name Ko-Autor	Arbeitsanteil (Inhalt und Umfang)	Unterschrift Ko-Autor																			
1. Ullesberger Berndt	Konzeption, Projektleiter Textverfassung																				
2. Rehak Thomas	Datenantrag, Konzeption																				
3. Pocivalnik Klimjan	Datenerhebung, Aggruppieren der Studienteilnehmer																				
4. Arvan Alexander	Statistik Technische Verarbeitung																				
5. Pichler Gerhard	Datenerhebung, Konzeption Textverfassung																				



Medizinische Fakultät



Kumulative Dissertation

Bestätigung

gem. § 4a Abs. 3 und 5 Promotionsordnung für die Promotion zum Dr. med., Dr. med. dent und Dr. rer. biol. hum.
und
gem. § 7 Abs. 4 Promotionsordnung für die Promotion zum Dr. rer. nat. an der Medizinischen Fakultät

Elisabeth Beckenbach (geborene Kratky)

Doktorand

"Regional Oxygen Saturation of the Brain during Birth transition of Term Infants: Comparison between

Elective and Vaginal Deliveries"

Titel der Publikation

Hiermit bestätige ich, dass keiner der zur Promotion eingereichten Fachartikel Gegenstand einer anderen (laufenden oder abgeschlossenen) Dissertation ist.

Unterschrift Doktorand

Folgende **Ko-Autoren** bestätigen mit ihrer Unterschrift

- ihren Arbeitsanteil (Inhalt und Umfang) an den eingereichten Veröffentlichungen,
- ihr Einverständnis zur Einreichung der Publikationen sowie,
- dass der jeweilige eingereichte Fachartikel nicht Gegenstand einer anderen (laufenden oder abgeschlossenen) Dissertation ist.

Name Ko-Autor

Arbeitsanteil (Inhalt und Umfang)

Unterschrift Ko-Autor

1.

Inhaltsliche Begleitung

2.

3.

4.

5.

weitere Autoren bitte auf ein gesondertes Blatt

1.5 Zusammenfassung Publikation I

Ziel dieser Studie war es die zerebrale Gewebssättigung (rSO_2 brain) beim reifen Neugeborenen (NG) unmittelbar nach der Geburt zu messen und diese mit nicht-invasiv präduktal gemessenen arteriellen Sättigungswerten zu vergleichen. In dieser prospektiven Studie wurden reife Neugeborene in den ersten 15 Lebensminuten nach vaginaler Spontangeburt und unauffälliger Anpassung inkludiert. Dabei wurde die rSO_2 brain mittels Nahinfrarotspektroskopie (NIRS) an der rechten Schläfe des Kindes gemessen. Zeitgleich wurde die Herzfrequenz (HR) und die arterielle Sättigung (SaO_2) am rechten Handgelenk (präduktal) mittels Pulsoxymetrie ermittelt. Diese drei Parameter (rSO_2 brain, HR, SaO_2) wurden von der 1. bis zur 15. Lebensminute kontinuierlich gemessen. Weiters wurde die regionale zerebrale Sauerstoffextraktion (Fractional tissue oxygen extraction, FTOE) mittels der Formel $(SaO_2 - rSO_2\text{brain})/SaO_2$ berechnet.

63 von 145 NG erfüllten die Einschlusskriterien.

Das Geschlechterverhältnis war 31 Mädchen (49,2%) und 32 Knaben (50,8%). rSO_2 brain zeigte von Minute 2 (39%) bis Minute 5 (69%) einen hoch signifikanten Anstieg. SaO_2 zeigte von Minute 2 (72%) bis Minute 14 (96%) einen signifikanten Anstieg. Die FTOE zeigte von Min 2 (0,47) bis Min 4 (0,3) einen signifikanten Abfall und einen neuerlichen Anstieg von Minute 8 bis 13.

Der Anstieg der Sauerstoffsättigung des Gehirns erfolgt nach Spontangeburt sehr rasch. Obwohl SaO_2 in den ersten 14 Minuten anstieg, zeigte die rSO_2 brain schon nach 5 Minuten keine weiteren Veränderungen mehr. Die zerebrale FTOE fiel in den ersten 4 Minuten ab, danach erreichte sie Normalwerte.

1.6 Zusammenfassung Publikation II

In dieser prospektiven Beobachtungsstudie wurde die Oxygenierung bei 63 NG nach Spontangeburt und 51 NG nach Kaiserschnittentbindung in den ersten 10 Lebensminuten miteinander verglichen. Auch hier wurde die rSO₂brain an der Stirn mittels NIRS gemessen. HR und SaO₂ wurde an der rechten Hand mittels Pulsoxymetrie ermittelt. Die FTOE wurde für jede Minute berechnet. Die Ergebnisse zeigten, dass zwischen der 4. und 8. Lebensminute die SaO₂ Werte bei den NG nach Kaiserschnitt signifikant niedriger waren im Vergleich zu NG nach Spontangeburt. Die Herzfrequenz der NG nach Kaiserschnitt zeigte sich während des gesamten Messzeitraumes von 10 Minuten signifikant niedriger. Bezuglich der rSO₂brain konnten keine signifikanten Unterschiede in den 2 Gruppen festgestellt werden. Obwohl sich SpO₂ und HR-Werte in der Kaiserschnitt-Gruppe deutlich niedriger waren, konnte bezüglich der zerebralen Sättigung kein Unterschied zwischen den Geburtsmodi nachgewiesen werden.

Einleitung

Die vorliegenden Publikationen sind aus einer Studie im Rahmen meiner Diplomarbeit an der Medizinischen Universität Graz (Österreich) entstanden. Seit mehreren Jahren beschäftigt sich die Forschungsgruppe der Neonatologie (Leiter Prof. Dr. Urlesberger) mit Veränderungen der Oxygenierung bei Neu und Frühgeborenen. Die Umstellungsprozesse nach der Geburt bieten diesbezüglich sehr interessante Aspekte. Als Mitglied dieser Forschungsgruppe durfte ich über einen längeren Zeitpunkt bei verschiedenen Studien mitwirken. Dabei entstand eine Studie, bei dem ich die Veränderungen der Oxygenierung an Neugeborenen unmittelbar nach der Geburt untersuchte. Als Projektleiterin hatte ich unterschiedliche Aufgabenbereiche, auf die ich im Folgenden näher eingehen möchte. Die Vorbereitungsphase beinhaltete das Einholen des Einverständnisses der Ethikkomission, Projektvorstellung im Kreißsaal-Team und Planung der Durchführbarkeit. In der Zeit von Oktober 2009 bis März 2010 rekrutierte ich das Patientenkollektiv und führte alle Messungen selbst durch. Gemeinsam mit den anderen Forschungsgruppenmitgliedern wurden die Messdaten analysiert und publiziert. In meiner Diplomarbeit¹ befasste ich mich ausführlich mit den medizinischen Grundlagen der Anpassung des Kreislaufs unter der Geburt, sowie den technischen Grundlagen der verwendeten Messgeräte und Messtechnologien. Weiters beschrieb ich die durchgeführte Studie und verglich sie mit den bisher publizierten Daten auf diesem Gebiet. In den letzten 2 Jahren konnten die Ergebnisse in zwei internationalen Zeitschriften publiziert werden. Diese sind Bestand meiner hier vorliegenden kumulativen Dissertation. Wie eingangs erwähnt, beschäftigt sich die Forschungsgruppe mit Veränderungen der zerebralen Gewebssättigung unmittelbar nach der Geburt.

„Der Übergang eines ungeborenen Kindes zu einem lebensfähigen Neugeborenen (NG) bedarf vieler physiologischer und teils komplexer Anpassungsleistungen. Es ist seit vielen Jahren bekannt, dass sich die Sauerstoffverhältnisse in dieser Übergangsphase drastisch verändern. Diese Veränderungen können mittels Pulsoxymetrie gemessen werden. Mit dieser Methode misst man die arterielle Sättigung (SaO_2), die den Prozentsatz an sauerstoffbeladenem Blutfarbstoff Hämoglobin anzeigt. Die engmaschige Überwachung dieser SaO_2 hat heutzutage in der intensiv-medizinischen Überwachung einen großen Stellenwert. Aus wissenschaftlichem Interesse haben sich einige Forschungsgruppen mit der Veränderung der SaO_2 von NG unmittelbar nach der Geburt beschäftigt^{2 3 4 5 6 7 8}. Diese erfährt bei gesunden NG innerhalb des Übergangs vom fetalen zum neonatalen Kreislauf einen rapiden Anstieg⁴⁻⁶. Bei nicht regelrechtem Ablauf dieser physiologischen Prozesse kann

es zu einer Mangelversorgung durch Sauerstoff (O_2) der Organe kommen. Das Gehirn ist das empfindlichste Organ des Menschen bezüglich O_2 -Mangel. Verschiedenste Ursachen vor, während und nach der Geburt können zu einem O_2 -Mangel führen und schwerwiegende Konsequenzen für die Entwicklung des Gehirns haben. Trotz aller Fortschritte der modernen Medizin im Bereich der intensivmedizinischen Betreuung von NG stellen hypoxisch-ischämische Gehirnveränderungen noch immer ein großes Problem dar.

Es ist noch weitgehend unerforscht, inwieweit sich die O_2 -Verhältnisse im Gehirn unter gesunden Bedingungen unmittelbar nach der Geburt verändern. Mehr Wissen darüber könnte in Zukunft eine abnormale O_2 -Versorgung des Gehirns erkennen lassen und für therapeutische Interventionen potentiell hilfreich sein.

Jöbsis⁹ beschrieb vor mehr als 30 Jahren eine Methode der Nahinfrarotspektroskopie (NIRS), welche Auskunft über die zerebrale Zirkulation gibt. Mit dieser Methode wurde es möglich, in verschiedenen Geweben im Körper Informationen über die regionale Gewebssättigung zu erhalten. In den letzten Jahrzehnten wurden einzelne Organsysteme wie die Niere, Gehirn, Leber und auch Muskelgewebe auf ihre O_2 -Sättigung näher untersucht. Es gibt auch Studien, die sich mit der zerebralen Gewebssättigung am ersten Lebenstag bei NG beschäftigten^{10 11 12}. Die Messgenauigkeit und die Aussagekraft dieser Methode stellen weiterhin ein großes Problem dar. Deshalb ist der klinisch-routinemäßige Einsatz bislang noch nicht erfolgt.

Das Ziel meiner Arbeiten war es, im Rahmen einer prospektiven klinischen Studie Informationen über die zerebrale Gewebssättigung in den ersten 15 Lebensminuten bei gesunden NG zu erhalten und eine Korrelation zu den arteriell gemessenen Werten darzustellen. Diese Studie wurde mittels einer Kooperation zwischen der Abteilung für Neonatologie (Universitätsklinik für Kinder- und Jugendheilkunde Graz) und dem Kreißsaal der Universitätsklinik für Gynäkologie und Geburtshilfe Graz möglich.“¹

Regional cerebral oxygen saturation in newborn infants in the first 15 min of life after vaginal delivery

Elisabeth Kratky¹, Gerhard Pichler^{1,3}, Thomas Rehak¹,
Alexander Avian², Mirjam Pocivalnik¹, Wilhelm Müller¹
and Berndt Urlesberger¹

¹ Division of Neonatology, Department of Pediatrics, Medical University of Graz, Austria

² Institute for Medical Informatics, Statistics and Documentation, Medical University of Graz, Austria

E-mail: pichler.gerhard@klinikum-graz.at

Received 30 August 2011, accepted for publication 15 November 2011

Published 15 December 2011

Online at stacks.iop.org/PM/33/95

Abstract

The objective of this study was to evaluate regional oxygen saturation of the brain during immediate transition after birth, and to correlate it with pre-ductal arterial oxygen saturation in newborn infants. The prospective observational study including newborn infants in the first 15 min after spontaneous vaginal delivery and uncomplicated transitional period was undertaken. Regional cerebral oxygen saturation (rSO_2 brain) was measured using near-infrared spectroscopy. Arterial oxygen saturation (SpO_2) and heart rate (HR) were measured on the right wrist by pulse oximetry. rSO_2 brain, SpO_2 and HR measurements were started immediately after birth and were performed in the first 15 min of life. Cerebral fractional tissue oxygen extraction (FTOE) was calculated for each minute. Of 145 newborn infants, 16 were included and the gender allocation was 31 females (49.2%) and 32 males (50.8%). rSO_2 brain increased rapidly from 39% (2 min) to 69% (5 min), SpO_2 increased from 72% (2 min) to 96% (14 min) and FTOE showed a significant decrease from minute 2 (0.47) until minute 4 (0.30) and an increase between 8 to 13 min. rSO_2 brain increased rapidly after vaginal delivery. Although SpO_2 increased within the first 14 min after delivery, rSO_2 brain showed no further significant changes after 5 min. FTOE decreased in the first 4 min and reached standard values subsequently.

Keywords: near-infrared spectroscopy, neonate, cerebral, oxygenation, transitional period

³ Author to whom any correspondence should be addressed.

Introduction

Arterial oxygen saturation (SpO_2) undergoes tremendous changes during transition from the foetus to newborn. Those changes can be seen clinically in the colour of the newborn's skin turning from cyanotic to pink but also can be measured objectively and non-invasively by pulse oximetry. In 1971, Oski and Delivoria-Papadopoulos (1971) already reported low oxygen saturation levels of about 60% in the newborn immediately after birth. Since then many reports have described the behaviour of oxygen saturation in this first phase of life. The oxygen saturation needs more than 5 min to attain levels >80% and almost 10 min to reach 90% (Kamlin *et al* 2006, Rabi *et al* 2006, Dawson *et al* 2007, House *et al* 1987). However, the SpO_2 measured on the right hand/wrist gives only indirect information on oxygen supply of the brain. As the brain is the most vulnerable organ system regarding oxygen deficiency, the adequate supply of oxygen to the brain is vital to provide accurate functioning.

After the introduction of near-infrared spectroscopy (NIRS) (Jöbsis 1977), the method allowed only the use of relative parameter changes in clinical use for years (Litscher and Schwarz 1997). A new variant of NIRS, called spatially resolved spectroscopy, introduced a new parameter, the regional tissue oxygen saturation ($r\text{SO}_2$) (Suzuki *et al* 1999), which is an absolute parameter for calculating the ratio of oxygenated haemoglobin to total haemoglobin. With this technology, it is possible to measure $r\text{SO}_2$ in different organ systems (brain, kidney, liver, muscle and others) or body regions (pre- and post-ductal peripheral tissue).

Knowing that SpO_2 changes significantly in the newborn infant after birth, interest has grown to monitor the changes of regional cerebral oxygen saturation ($r\text{SO}_2\text{brain}$). Some reports described the $r\text{SO}_2\text{brain}$ in pre-term infants on the first day of life (Sorensen and Greisen 2009) and during the first weeks of life (Verhagen *et al* 2007, Naulaers *et al* 2002, 2003). Only a few studies measured cerebral oxygenation during the transitional period in healthy term infants (Isobe *et al* 2002, Fauchère *et al* 2010, Urlesberger *et al* 2010, 2011). The majority of studies investigated infants born after caesarean delivery. Therefore, recently we conducted a study, comparing infants after vaginal deliveries and elective caesarean sections (Urlesberger *et al* 2011). The publication focused on behaviour of SpO_2 and $r\text{SO}_2\text{brain}$ between minutes 3 and 10 after birth. Until now there was no publication of the detailed behaviour of $r\text{SO}_2\text{brain}$ after spontaneous vaginal delivery. Therefore, this study describes the behaviour of SpO_2 and $r\text{SO}_2\text{brain}$ from minutes 2 to 15 in these healthy newborn infants after spontaneous vaginal delivery for the first time, and thus presenting physiological data of a perfectly normal transitional process in detail.

Methods

In this prospective observational study, we included newborn infants over 37 weeks of gestational age after uncomplicated pregnancy. Only infants after spontaneous vaginal delivery and uncomplicated transitional period were included. Infants after vacuum/ forceps assisted delivery or caesarean section were excluded. All infants with any suspected malformations, need of respiratory support or supplemental inspired oxygen during transition were excluded.

Instruments

The Invos Cerebral/Somatic Oximeter Monitor (Covidien, Mansfield, MA, USA) was used with the neonatal transducer. The transducer contains a light-emitting diode and two sensors at different distances. The Invos calculates the $r\text{SO}_2$, which is expressed as the percentage of oxygenated haemoglobin (oxygenated haemoglobin/total haemoglobin). The transducer was

positioned on the left fronto-parietal forehead of the infant. The sample rate of rSO₂ was 8 s (0.13 Hz).

SpO₂ and the heart rate (HR) were measured with the Intelli-Vue MMS X2 Monitor (Philips, Eindhoven, the Netherlands). The transcutaneous transducer was placed on the right wrist (pre-ductal) of the infant. Values of the SpO₂ and rSO₂ were stored every second.

Procedure

As soon as the whole body of the infant was delivered, a stopwatch was started to record the age in minutes. Immediately after birth, the skin on the left forehead was cleaned with a towel to reduce the amount of amniotic fluid and vernix. Both could have a negative influence on the signal quality detection. Straight after the NIRS transducer was put on the left fronto-parietal region of the infant, the transducer was fixed with a red headband (Medijet nCPAP Bonnet, Size Medium). Subsequently, the transcutaneous pulse oximetry transducer was positioned on the right wrist of the newborn as quickly as possible. In the mean time, the cord was clamped and the infants were placed on the naked skin of the mother's chest to have close skin-to-skin contact. The infants were positioned in prone position and covered with a warm towel. All values were stored continuously in both devices. A neonatologist observed the transition of the newborn infants and recorded Apgar (Apgar 1953) scores at 1, 5 and 10 minutes. The Apgar score is a simple and repeatable method to quickly assess the clinical situation of newborn infants immediately after birth. Five items (skin colour, HR, reflex irritability, muscle tone and respiration) are scored with a range from 0 to 2, and summed up (resulting in ten points for a healthy normal newborn after transition). After 15 min, both transducers and the headband were taken off. Our procedure ensured an uninterrupted bonding phase (relationship) between child and mother, i.e. there was no need to take the child away from the mother as a result of our measurements. All measurements were done by the same investigator (EK).

Quality criteria

To detect and eliminate artefacts, the following quality criteria for rSO₂brain were used. For physiological reasons, the SpO₂ should not be equal or below the rSO₂brain. Such data were considered not to be physiological and, therefore, the arterial as well as the regional saturation values were excluded.

Ethics

The Regional Committee on Biomedical Research Ethics approved the project. Written informed consent was obtained from the mothers/parents before birth of their infant.

Statistics

Mean values of arterial oxygen saturations and rSO₂brain were calculated for each minute after birth. Data are presented as mean and 95% confidence interval (CI). In addition, the 'fractional tissue oxygen extraction' (FTOE) was calculated (SpO₂-rSO₂brain)/SpO₂) (Naulaers *et al* 2007). A linear mixed model with a fixed effect for time and a first-order autoregressive covariance structure was used for the calculation of significant differences to endpoint of measurements (15 min value). A *p*-value of <0.05 (*) was considered statistically significant and a *p*-value <0.01 (**) as highly significant. Statistical analysis was performed using SPSS 17.0.3 (SPSS Inc., Chicago, IL, USA, 2010).

Table 1. Demographic data and patient's characteristics.

Gestational age (weeks)	40 (1.3)
Birthweight (g)	3370 (437)
Headcircumference (cm)	34.9 (1.4)
APGAR- 1 min	9 (0.5)
APGAR- 2 min	10 (0)
pH umbilical artery	7.28 (0.08)

Mean (SD).

Table 2. Mean (CI) of oxygen saturation parameter, HR and FTOE for minute 2 to minute 15.

Minute	rSO ₂ brain (%)	SpO ₂ (%)	HR (1/min)	FTOE
2	39 (33–46)*	72 (69–76)*	146 (134–158)	0.47*
3	49 (43–56)*	78 (75–81)*	157 (145–169)*	0.38*
4	61 (55–67)*	85 (82–88)*	160 (148–172)*	0.30*
5	69 (63–75)*	89 (86–92)*	160 (148–172)*	0.24
6	73 (67–79)	92 (89–95)*	157 (146–169)*	0.21
7	75 (69–82)	94 (91–97)*	155 (144–167)*	0.19
8	77 (71–83)	95 (92–98)*	153 (142–165)	0.18*
9	78 (72–84)	95 (92–98)*	152 (141–163)	0.18*
10	77 (72–83)	95 (92–98)	152 (141–163)	0.19*
11	77 (72–82)	95 (92–98)*	151 (141–161)	0.19*
12	77 (74–82)	95 (93–98)*	149 (139–159)	0.19*
13	76 (72–81)	96 (93–98)*	148 (139–157)	0.20*
14	76 (72–80)	96 (94–98)*	148 (140–156)	0.21
15	75 (72–78)	97 (95–98)	149 (143–154)	0.22

*Significant differences ($p < 0.05$) to endpoint of measurements (15 min value).

Results

Between October 2009 and August 2010, 134 newborn infants fulfilled inclusion criteria primarily (before delivery). Sixty-one (46%) infants were excluded because the mode of delivery was finally not spontaneous vaginal but vacuum/forceps assisted ($n = 37$) or caesarean sections ($n = 24$). Moreover, four (3%) infants were excluded because they needed respiratory support after delivery and six (4%) were not assessed due to organizational reasons (i.e. two deliveries at the same time). The gender allocation of the remaining 63 (47%) was 32 males and 31 females (table 1).

By 2 min of age, saturation signals could be measured for at least 85% of infants. Therefore, the data presentation begins at 2 min. From minute 4 to minute 15, saturation signals were available for at least 96.8% of the infants. Six (0.7%) out of 812 rSO₂brain values failed quality criteria and subsequently were removed (the corresponding SpO₂ was removed too, six (0.7%) out of 804).

All three measured parameters (rSO₂brain, SpO₂ and HR) and the calculated FTOE are presented in table 2. Mean rSO₂brain increased rapidly from 39% at 2 min to 69% at 5 min; subsequently, no significant changes occurred. Mean SpO₂ increased from 72% at 2 min up to 96% at 14 min. Only at 10 min there was no significant difference, but it was very close to significance ($p = 0.056$). The time course of SpO₂ and rSO₂brain are presented in figure 1 and of FTOE in figure 2. FTOE values show a decline from 0.47 at 2 min to 0.3 at 4 min. Subsequently, a slight increase from 8 min to 13 min was observed.

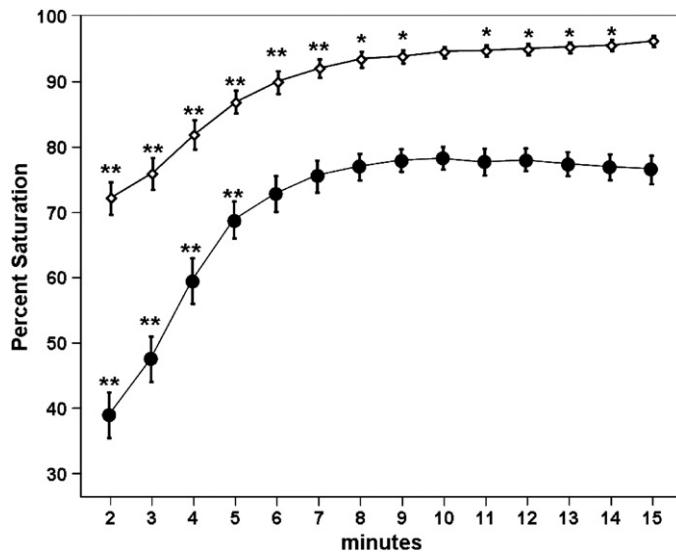


Figure 1. Arterial oxygen saturation (SpO_2) (open diamonds) and regional oxygen saturation of the brain (rSO_2brain) (closed circles) from minute 2—to minute 15 (mean, 95%CI). * $p < 0.05$, ** $p < 0.001$: difference to value minute 15.

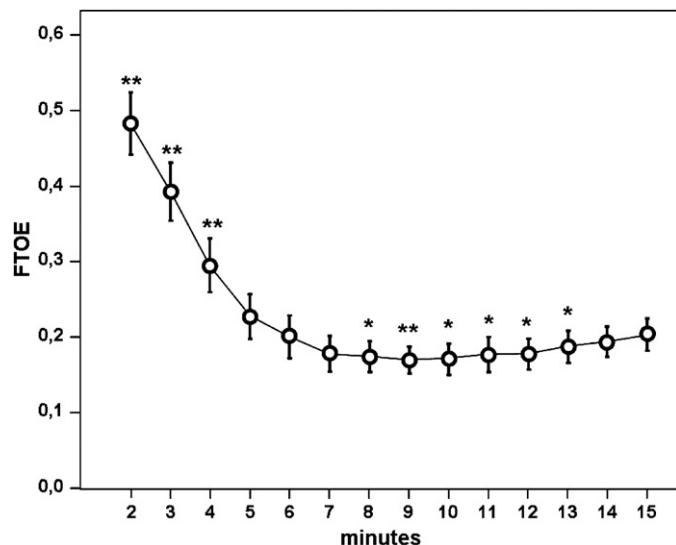


Figure 2. Time course of fractional oxygen tissue extraction FTOE (mean, 95%CI). * $p < 0.05$, ** $p < 0.001$: difference to value minute 15.

Discussion

We analysed changes in regional oxygen saturation in the brain and simultaneous changes in pre-ductal SpO_2 during transitional period in healthy newborn infants after spontaneous vaginal delivery. Our group performed these measurements for the first time in a significant number of newborns. The lack of data till now is certainly due to the difficulty to measure

in a delicate situation immediately after vaginal delivery. Nevertheless, we tried to ensure an uninterrupted bonding for mother and child. Comparing spontaneous vaginal delivery and caesarean section we have already analysed data from minute 3 to minute 10¹⁶. In this study we present data from minute 2 to minute 15 in more detail.

rSO₂ depends on the local balance of oxygen delivery and oxygen consumption, and the regional arterial/venous volume ratio of veins (70–80%), capillaries (5%) and arteries (15–25%) (Watzman *et al* 2000). The main component of tissue oxygen saturation is venous blood, thus representing oxygen saturation after oxygen consumption by the tissue. Therefore, the values for rSO₂ tend to be close to venous oxygen saturation, but do not equate venous saturation (Watzman *et al* 2000).

In our study rSO₂brain showed a significant increase within the first 5 min after birth, thereafter there was no longer any significant change. In contrast to that, SpO₂ values showed a significant increase during the whole observation period (but minute 10). So far all publications on SpO₂ values described a period up to minute 10. Our results show a constant further increase beyond minute 10 until minute 14. These results clearly show a different behaviour of arterial versus rSO₂brain, the latter showing no further increase after 5 min. We speculate, that there may be a difference in oxygen delivery to the brain compared to the body. Recently, we described significantly lower values of SpO₂ and HR in caesarean-delivered infants, but there were no differences in rSO₂ brain values between infants after vaginal and caesarean delivery (Urlesberger *et al* 2011).

Fauchère *et al* (2010) published data on rSO₂brain measured in 17 healthy term infants after caesarean section using a different NIRS device (NIRO 300, Hamamatsu, Japan). They also observed a plateau of rSO₂ after a steep increase until 7 min. Their absolute values for rSO₂ were lower compared to this study using a different measurement device. Isobe *et al* (2002) also investigated healthy term infants after vaginal delivery ($n = 20$) and elective caesarean section ($n = 7$). They did not measure rSO₂brain directly but calculated a value for rSO₂ of the brain. They also observed a plateau of cerebral oxygen saturation 8.5 min after birth (Isobe *et al* 2002). The observational periods differed between those studies: two lasted for 10 min (Fauchère *et al* 2010, Urlesberger *et al* 2010), and one for 15 min (Isobe *et al* 2002).

By calculating the FTOE, information on oxygen consumption is gained (Naulaers *et al* 2003). An increase in fractional oxygen extraction indicates an increase of oxygen consumption because of increased cerebral metabolic rate, or a decreased oxygen delivery, or both (van den Berg *et al* 2010, Kissack *et al* 2005, 2004). In this study, FTOE values are very high in the first few minutes of life. This is probably due to the low oxygen supply in arterial blood at that time (SpO₂ 60% at 1 min) so that the brain extracts almost half (FTOE of 0.47) of the available oxygen in the blood. After a few minutes, the percentage of oxygenated haemoglobin in the blood increases so that the brain can extract smaller amount of oxygen to be adequately provided and FTOE values reach standard values of 20–30%. This significant fall in FTOE lasted only until minute 4. Thereafter, an increase of FTOE was observed from 8 to 13 min. The increase was small, and this may not be relevant clinically. We can only speculate about the reasons for this behaviour. Cerebral oxygen delivery is a product of blood flow and oxygen content. On one hand, a decrease in HR with a consecutive decrease in cardiac output may have reduced cerebral oxygen delivery again. There was a trend of decreasing HR at the same time, but there were no significant changes. Nevertheless, it has been shown in a newborn dog model that even significant changes in cardiac output were compensated and did not result in changes in cerebral blood flow (CBF) (Camp *et al* 1982). As SpO₂ did show a significant increase during this period, oxygen content of arterial blood should not have been decreased. Another possible explanation could be first sensory orientation and/or starting motor movements in

this phase of transition. This may lead to higher oxygen demand and therefore to a higher oxygen extraction in the brain.

Our results show that oxygen supply of the brain is provided very quickly although the increase of SpO_2 takes much longer. This might indicate some sort of preference of oxygen supply to the brain compared to other organ systems. We recently showed (Urlesberger *et al* 2010) that regional oxygen saturation increase in muscle tissue takes 7 min, whereas the brain showed no significant changes 5 min after birth.

The slight increase after 8 min was an unexpected observance. At that moment there were no corresponding changes in HR or SpO_2 .

At the moment we do not know, whether or not rSO_2 of the brain may differ significantly between different regions of the brain. Recently, Lemmers and Van Bel (2009) have shown, that especially during situations with unstable arterial saturation, differences can exist between the left and right hemisphere. Therefore, we used the same fronto-parietal region in all infants.

As we have no detailed information on cardio-circulatory changes, behaviour of CBF and blood gas changes (PaO_2 and PaCO_2), we cannot define the impact of these parameters on changes in oxygen saturation parameters. Myocardial function is crucial during haemodynamic transition to extrauterine life. Beside left ventricular output and blood pressure, CBF is furthermore regulated by cerebral autoregulation. Cerebral autoregulation is influenced by changes in PaO_2 and PaCO_2 . PaCO_2 seems to have even more influence on CBF than mean arterial blood pressure (Noori *et al* 2009). There is a negative correlation between PaCO_2 and CBF (Sorensen and Greisen 2006). We saw a steep increase of SpO_2 , which certainly has influence on cerebral oxygen extraction (Kissack *et al* 2005), but whether there was a concomitant change in CBF is unknown. In the situation of normal transition, one would expect that the increase in SpO_2 (reassuring improving gas exchange via the lungs) would be accompanied by a decrease in pCO_2 (resulting in an increase of CBF).

One has to emphasize, that at the moment comparison of absolute values of rSO_2 has to be done with caution. The absolute value of rSO_2 lacks the precision (Sorensen and Greisen 2006) to be used as a quantitative variable, but it can be used as a trend monitor with helpful clinical information (van Bel *et al* 2008). Comparison of values of rSO_2 attained with different technical devices may be problematic (Thavasothy *et al* 2002, Gagnon *et al* 2002). However, there are some comparative studies done between the two most used devices: the INVOS system and the NIRO 300 (Hamamatsu, Japan) (Thavasothy *et al* 2002, Gagnon *et al* 2002). The standard deviation of the difference between the two monitors was 7.5%, but they showed a very good correlation of the trend values (Pryds *et al* 1988).

In conclusion, this paper is able to show a rapid increase of cerebral oxygen saturation in newborn infants during the first few minutes after vaginal delivery. Further studies are needed to gain more normative data from healthy term infants. These studies will show the potential value of rSO_2 measurements of the brain during neonatal transition (and define their value in evaluation of the use of supplemental oxygen at the resuscitation table in future).

References

- Apgar V 1953 A proposal for a new method of evaluation of the newborn infant *Curr. Res. Anesth. Analg.* **32** 260–7
Camp D, Kotgal U and Kleinman L 1982 Preservation of cerebral autoregulation in the unanesthetized hypoxicemic newborn dog *Brain Res.* **241** 207–13
Dawson J A, Davis P G, O'Donnell C P F, Kamlin C O F and Morley C J 2007 Pulse oximetry for monitoring infants in the delivery room: a review *Arch. Dis. Child. Fetal Neonatal Ed.* **92** F4–7
Fauchère J C, Schulz G, Haensse D, Keller E, Ersch J, Bucher H U and Wolf M 2010 Near-infrared spectroscopy measurements of cerebral oxygenation in newborns during immediate postnatal adaptation *J. Pediatr.* **156** 372–6
Gagnon R E *et al* 2002 Comparison of two spatially resolved NIRS oxygenation indices *J. Clin. Monit. Comput.* **17** 385–91

- House J T, Schultetus R R and Gravenstein N 1987 Continuous neonatal evaluation in the delivery room by pulse oximetry *J. Clin. Monit.* **3** 96–100
- Isobe K, Kusaka T, Fujikawa Y, Okubo K, Nagano K, Yasuda S, Kondo M, Itoh S, Hirao K and Onishi S 2002 Measurement of cerebral oxygenation in neonates after vaginal delivery and cesarean section using full-spectrum near infrared spectroscopy *Comp. Biochem. Physiol. A: Mol. Integr. Physiol.* **132** 133–8
- Jöbsis F F 1977 Noninvasive infrared monitoring of cerebral and myocardial oxygen sufficiency and circulatory parameters *Science* **198** 1264–7
- Kamlin C O, O'Donell C P, Davis P G and Morley C J 2006 Oxygen saturation in healthy infants immediately after birth *J. Pediatr.* **148** 585–9
- Kissack C M, Garr R, Wardle S P and Weindling A M 2004 Cerebral fractional oxygen extraction in very low birth weight infants is high when there is low left ventricular output and hypotension *Pediatr. Res.* **55** 400–5
- Kissack C M, Garr R, Wardle S P and Weindling A M 2005 Cerebral fractional oxygen extraction is inversely correlated with oxygen delivery in the sick, newborn, preterm infant *J. Cereb. Blood Flow Metab.* **25** 545–53
- Lemmers P and Van Bel F 2009 Left-to-right differences of regional cerebral oxygen saturation and oxygen extraction in preterm infants during the first days of life *Pediatr. Res.* **65** 226–30
- Litscher G and Schwarz G 1997 *Trancranial Cerebral Oximetry* (Vienna: Papst Science Publishers)
- Naulaers G, Meyns B, Miserez M, Leunens V, Van Huffel S, Casaer P, Weindling M and Devlieger H 2007 Use of tissue oxygenation index and fractional tissue oxygen extraction as non-invasive parameters for cerebral oxygenation. A validation study in piglets *Neonatology* **92** 120–6
- Naulaers G, Morren G, Van Huffel S, Casaer P and Devlieger H 2002 Cerebral tissue oxygenation index in very premature infants *Arch. Dis. Child. Fetal Neonatal Ed.* **87** F189–92
- Naulaers G, Morren G, Van Huffel S, Casaer P and Devlieger H 2003 Measurement of tissue oxygenation index during the first three days in premature born infants *Adv. Exp. Med. Biol.* **510** 379–83
- Noori S, Stavroudis T A and Seri I 2009 Systemic and cerebral hemodynamics during the transitional period after premature birth *Clin. Perinatol.* **36** 723–36
- Oski F A and Delivoria-Papadopoulos M 1971 The shift to the left *Pediatrics* **48** 853–6
- Pryds O, Greisen G and Trojaborg W 1988 Visual evoked potentials in preterm infants during the first hours of life *Electroencephalogr. Clin. Neurophysiol.* **71** 257–65
- Rabi Y, Yee W, Chen S Y and Singhal N 2006 Oxygen saturation trends immediately after birth *J. Pediatr.* **148** 590–4
- Sorensen L C and Greisen G 2006 Precision of measurement of cerebral tissue oxygenation index using near-infrared spectroscopy in preterm neonates *J. Biomed. Opt.* **11** 054005
- Sorensen L C and Greisen G 2009 The brains of very preterm newborns in clinically stable condition may be hyperoxygenated *Pediatrics* **124** e958–63
- Suzuki S, Takasaki S, Ozaki T and Kobayashi Y 1999 A tissue oxygenation monitor using NIR spatially resolved spectroscopy *Proc. SPIE* **3597** 582–92
- Thavasothy M, Broadhead M, Elwell C, Peters M and Smith M 2002 A comparison of cerebral oxygenation as measured by the NIRO 300 and the INVOS 5100 Near-Infrared Spectrophotometers *Anaesthesia* **57** 999–1006
- Urlesberger B, Grossauer K, Pocivalnik M, Avian A, Müller W and Pichler G 2010 Regional oxygen saturation of the brain and peripheral tissue during birth transition of term infants *J. Pediatr.* **157** 740–4
- Urlesberger B, Kratky E, Rehak T, Pocivalnik M, Avian A, Czihak J, Müller W and Pichler G 2011 Regional oxygen saturation of the brain during birth transition of term infants: comparison between elective cesarean and vaginal deliveries *J. Pediatr.* **159** 404–8
- van Bel F, Lemmers P and Naulaers G 2008 Monitoring neonatal regional cerebral oxygen saturation in clinical practice: value and pitfalls *Neonatology* **94** 237–44
- van den Berg E, Lemmers P M A, Toet M C, Klaessens J H G and van Bel F 2010 Effect of the 'InSurE' procedure on cerebral oxygenation and electrical brain activity of the preterm infant *Arch. Dis. Child. Fetal Neonatal Ed.* **95** F53–8
- Verhagen E A, ter Horst H J, Keating P and Bos A F 2007 The course of cerebral oxygen saturation and oxygen extraction during the first two weeks of life in pre term infants *Acta Paediatr.* **96** (Suppl. 456) 140
- Watzman H M *et al* 2000 Arterial and venous contributions to near-infrared cerebral oximetry *Anesthesiology* **93** 947–53

Regional Oxygen Saturation of the Brain during Birth Transition of Term Infants: Comparison between Elective Cesarean and Vaginal Deliveries

Berndt Urlesberger, MD, Elisabeth Kratky, MD, Thomas Rehak, MD, Mirjam Pocivalnik, MD, Alexander Avian, MSc, Johanna Czihak, MD, Wilhelm Müller, MD, and Gerhard Pichler, MD

Objective To evaluate differences in regional oxygen saturation of the brains of term infants of vaginal or cesarean deliveries.

Study design Vaginal delivery ($n = 63$) and elective cesarean delivery infants were prospectively evaluated for the first 10 minutes after delivery. Peripheral arterial oxygen saturation (SpO_2) and heart rate were measured on the right hand using pulsoximetry with near infrared spectroscopy. Regional oxygen saturation of the brain ($r\text{SO}_2\text{brain}$) was measured. Fractional tissue oxygen extraction was calculated for each minute.

Results From 4 to 8 minutes, SpO_2 values for cesarean delivery infants were significantly lower than for vaginally delivered infants. Heart rate of the cesarean delivery infants was significantly lower throughout the whole observation period. There was no difference between groups in $r\text{SO}_2\text{brain}$. Fractional tissue oxygen extraction only differed at minute 10.

Conclusions Although SpO_2 and heart rate were significantly lower in cesarean-delivered infants, there were no differences in $r\text{SO}_2\text{brain}$ with respect to mode of delivery. (*J Pediatr* 2011;159:404-8).

The transition from fetus to newborn is a complex physiological process. In recent years, interest has grown in the use of pulsoximetry to monitor arterial oxygen saturation during this transitional period.¹⁻³ All newborn infants have oxygen desaturation at birth. A newborn infant undergoing normal postnatal transition needs more than 5 minutes to attain an arterial oxygen saturation >80% and almost 10 minutes to reach 90%. Several studies have shown that there are significant differences in the time course of arterial oxygen saturation (SpO_2) according to the mode of delivery. Newborn infants after elective caesarean delivery have lower SpO_2 values during transition compared with infants after vaginal delivery.¹⁻³

There is an ongoing discussion about the use of supplemental oxygen during neonatal resuscitation because it is not known which oxygen concentration is appropriate for preterm and term infants during resuscitation.^{3,4} Because the brain is the most vulnerable organ system of the infant, a more direct way to assess its oxygenation in a simple noninvasive way would be useful. A new approach to cerebral oxygenation is to measure regional oxygen saturation using near infrared spectroscopy (NIRS).

Spatially resolved spectroscopy, a new NIRS method, was recently introduced for evaluation of regional oxygen saturation ($r\text{SO}_2$).⁵ With this technology, it is possible to measure regional tissue oxygen saturation in different organ systems (brain, kidney, liver, muscle, and others) or body regions (preductal and postductal peripheral tissues). There are some reports of regional oxygen saturation of the brain ($r\text{SO}_2\text{brain}$) on the first day of life,⁶⁻¹⁰ during the first weeks, and during birth transition.¹¹⁻¹³

A question remains as to whether $r\text{SO}_2$ of the brain shows significant differences during transition according to mode of delivery in analogy to SpO_2 behavior. Therefore, we measured $r\text{SO}_2$ brain in newborn infants after vaginal delivery and elective cesarean delivery to evaluate whether there was a significant difference.

Methods

In this prospective, observational study, we included newborn infants >37 weeks gestational age delivered after an uncomplicated pregnancy. Two groups were formed according to the mode of delivery, either vaginal delivery or delivery by elective cesarean delivery. All infants with malformations were excluded. Only infants after uncomplicated vaginal delivery were included. Infants from vacuum-assisted or forceps deliveries were excluded. All infants in need of any respiratory support or supplemental inspired oxygen during transition were excluded. Only infants with uncomplicated transitional periods were included.

CBF	Cerebral blood flow
HR	Heart rate
NIRS	Near infrared spectroscopy
$r\text{SO}_2$	Regional oxygen saturation
$r\text{SO}_2$ brain	Regional oxygen saturation of the brain
SpO_2	Peripheral arterial oxygen saturation

From the Research Unit for Cerebral Development and Oximetry, Division of Neonatology, Department of Pediatrics (B.U., E.K., T.R., M.P., W.M., G.P.); the Institute for Medical Informatics, Statistics, and Information (A.A.); and the Department of Obstetrics and Gynecology (J.C.), Medical University of Graz, Austria

The authors declare no conflicts of interest.

0022-3476/\$ - see front matter. Copyright © 2011 Mosby Inc.
All rights reserved. 10.1016/j.jpeds.2011.02.030

For NIRS measurements, the Invos Cerebral/Somatic Oximeter Monitor (Covidien, Mansfield, Massachusetts) was used with the neonatal transducer. The transducer contains a light-emitting diode and two sensors at different distances. The Invos calculates the rSO₂, which is expressed as the percentage of oxygenated hemoglobin (oxygenated hemoglobin/total hemoglobin). The transducer was positioned on the left frontoparietal region of the forehead of each infant, regardless of mode of delivery. The sensor on the forehead was secured with a bandage. Arterial oxygen saturation and heart rate (HR) were measured with the IntelliVue MP30 Monitor (Philips, Eindhoven, The Netherlands) in the cesarean delivery group and the IntelliVue MMS X2 Monitor (Philips) in vaginal delivery group. The transducer was placed on the right hand/wrist. In the cesarean delivery group, blood pressures were measured once 5 minutes after birth with the IntelliVue MP30 Monitor (Philips). Rectal and peripheral temperatures were measured with the IntelliVue MP30 Monitor (Philips).

All variables were stored continuously in a multichannel system, "alpha-trace digital MM" (B.E.S.T. Medical Systems, Vienna, Austria), for subsequent analysis. Values of the arterial and regional oxygen saturation were stored every second, and the sample rate of rSO₂ was 8 seconds (0.13 Hz). The Regional Committee on Biomedical Research Ethics approved the project. Written informed consent was obtained before the birth of the infant.

Procedure

A stopwatch was started when the cord was clamped.

Vaginal Delivery Group

Immediately after birth, the NIRS transducer was applied to the left forehead (rSO₂brain), and, concurrently, transcutaneous pulsoximetry was started on a preductal level (right hand) (SpO₂). After that, the child was given to the mother, and for the rest of the 10-minute transition period, the infant was positioned in prone position on the mother's chest. A neonatologist observed the transition of the newborn infant and recorded APGAR scores at 1, 5, and 10 minutes. Measurement of blood pressure and rectal temperature were omitted because of the delicate measurement situation immediately after delivery and to ensure an uninterrupted bonding of mother and child.

Cesarean Delivery Group

All infants were dried and wrapped with warmed towels. Immediately after arrival at the resuscitation table, the NIRS transducer was applied to the left forehead (rSO₂brain), and transcutaneous pulsoximetry was started on a preductal level (right hand) (SpO₂). The infant was positioned supine and was breathing room air. A neonatologist observed the transition of the newborn infant and recorded APGAR scores at 1, 5, and 10 minutes. A clinical assessment at 10 minutes was done before the newborn infant was given to the parents. Fractional tissue oxygen extraction was calculated for each minute [(SpO₂-rSO₂)/SpO₂] ^{xiv}¹⁴.

Quality Criteria

To detect and eliminate artifacts, the following quality criteria for rSO₂ measurements were used. For physiological reasons, the SpO₂ should not be equal or below the rSO₂. Such data were considered to be not physiological, and therefore the arterial as well as the regional saturation values were excluded. Furthermore, if ≥5 rSO₂ data in any category failed quality criteria (≤50% data completeness), the patient was excluded from the study.

Statistics

Mean values of arterial oxygen saturations and regional oxygen saturations were calculated for each minute after birth. Data are presented as mean and 95% CI or mean ± SD. A linear mixed model with a fixed effect for time and a first-order autoregressive covariance structure was used for calculation of significant differences between both groups. To compare patient characteristics, *t* tests were used. A *P* value of <.05 was considered significant. Statistical analysis was performed using SPSS 17.0.3 (SPSS Inc, Chicago, Illinois, 2009).

Results

Between October 2009 and August 2010, 185 infants fulfilled inclusion criteria and thus were included in the study (all before delivery). In the cesarean delivery group, 51 newborn infants fulfilled inclusion criteria initially. Two infants (4%) were excluded because of the need of respiratory support after delivery. Five infants (10%) were excluded because SpO₂ values were not able to be measured within 10 minutes. The sex allocation of remaining 44 infants in the cesarean delivery group was 23 male and 21 female (**Table**).

In the vaginal delivery group, 134 infants fulfilled inclusion criteria initially (before delivery). Four (3%) infants were excluded because of the need of respiratory support after delivery, six infants (4%) were not assessed because of organizational reasons (two deliveries at the same time), and 61 (46%) infants were excluded because they failed vaginal delivery (24 infants, cesarean delivery; 35 infants, vacuum-assisted delivery; and two infants, forceps delivery). The sex allocation of remaining 63 infants in vaginal delivery group was 32 male and 31 female (**Table**).

Table. Demographic data and patients characteristics

	Cesarean delivery	Vaginal delivery
Gestational age, wk	38.5 (1.1)	40 (1.3)
Body weight, g	3197 (448)	3369 (437)
Head circumference, cm	34.6 (1.1)	34.9 (1.4)
APGAR: 1 min	9 (0.2)	9 (0.5)
APGAR: 5 min	10 (0.15)	10 (0)
pH Umbilical artery	7.29 (0.03)	7.28 (0.08)
Mean arterial blood pressure, mm Hg	49.3 (8.6)	
Temperature rectal, Celsius	36.7 (0.04)	

Values are mean (SD).

By 3 minutes, both saturation signals (rSO_2 and SpO_2) could be measured for at least 90% of infants. Therefore, the data presentation begins at 3 minutes. In the cesarean delivery group, 10 (1.8%) of 559 rSO_2 values failed quality criteria and subsequently were removed (the corresponding SpO_2 values also were removed, 10 (1.8%) of 556). In the vaginal delivery group, six (0.7%) of 812 rSO_2 values failed quality criteria and subsequently were removed (the corresponding SpO_2 value also was removed, six (0.7%) of 804).

From 4 to 8 minutes, SpO_2 values of the cesarean delivery group were significantly lower. HR of the cesarean delivery group was significantly lower throughout the whole observation period (Figure 1). There was not any difference between the groups in rSO_2 brain. Fractional tissue oxygen extraction only differed at minute 10 (Figure 2).

Discussion

We analyzed changes of the regional tissue oxygen saturation of the brain and simultaneous changes of the preductal arterial oxygen saturation during transition in a large group of healthy infants after vaginal delivery in comparison with cesarean delivery infants. Regional tissue oxygen saturation depends on the local balance of oxygen delivery and oxygen consumption and the regional arterial/venous volume ratio,

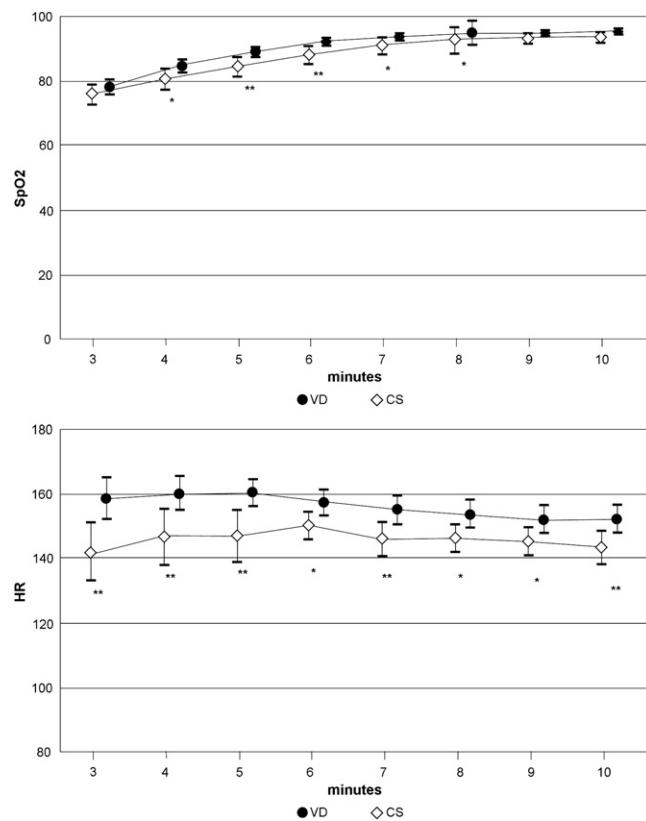


Figure 1. Course of arterial oxygen saturation and heart rate (mean, 95% CI; * $P < .05$, ** $P < .001$, comparison between groups).

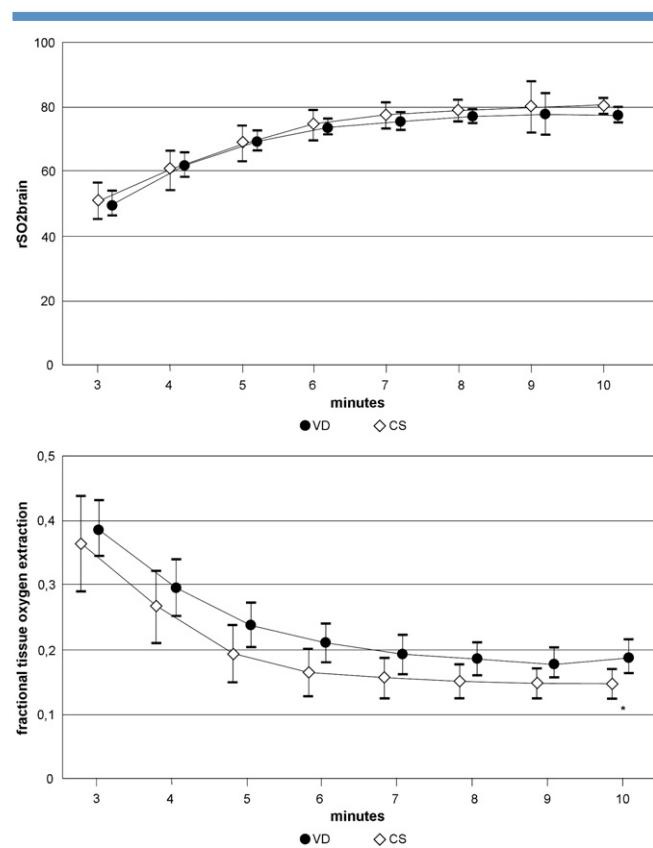


Figure 2. Course of regional cerebral oxygen saturation and fractional tissue oxygen extraction (mean, 95% CI; * $P < .05$, comparison between groups).

reflecting oxygen saturation in veins (70% to 80%), capillaries (5%) and arteries (15% to 25%).¹⁵ The main component of tissue oxygen saturation is venous blood, thus representing oxygen saturation after oxygen consumption by the tissue. Therefore, the values for regional oxygen saturation tend to be close to venous oxygen saturation but do not equate venous saturation.

Rabi et al² demonstrated that term infants born by cesarean delivery had modestly lower oxygen saturations and required longer to reach a stable oxygen saturation $\geq 85\%$ in the immediate newborn period compared with infants born vaginally. The mean difference at 5 minutes was 3%. Very similar results were reported by Kamlin et al.¹ Recently, Dawson et al¹⁶ showed in a large group of patients that there were significantly lower SpO_2 values in infants delivered by cesarean birth from 1 until 5 minutes after birth. Thereafter, the SpO_2 values did not differ. Furthermore, Dawson et al¹⁷ did show that HR was significantly lower in newborn infants after cesarean delivery (from 2 to 10 minutes). Therefore, the course of the SpO_2 and HR values of the present study are in accordance with the literature.

Although there were significantly lower SpO_2 and HR values in the cesarean delivery group, there was not any significant difference in rSO_2 brain between both groups. Years ago, before spatially resolved spectroscopy was introduced

to measure rSO₂, Isobe et al¹¹ calculated a value for regional cerebral oxygen saturation in a small number of infants (6 infants after cesarean delivery, 20 infants after vaginal delivery) during transition. They, too, did not find a significant difference in cerebral oxygen saturation within the first 10 minutes of life. It is difficult to differentiate whether oxygen delivery and/or oxygen consumption of the brain were different between the two groups. Cerebral oxygen delivery is a product of blood flow and oxygen content. Because cardiac output in neonates is mainly influenced by HR, one would expect a reduction in oxygen delivery with reduced HR (cesarean delivery group). Nevertheless, in a newborn dog model, even significant changes in cardiac output were compensated and did not result in changes in cerebral blood flow (CBF).¹⁸ Because oxygen content of the arterial blood also influences oxygen delivery, significantly lower SpO₂ in the cesarean delivery group would be expected to result in reduced cerebral oxygen delivery. Nevertheless, in a newborn lamb model, changes of arterial oxygen content were accompanied by reciprocal changes in CBF to maintain constant cerebral oxygen delivery.¹⁹ The authors of both studies explained their results by cerebral autoregulatory response mechanisms.

The fractional tissue oxygen extraction may give information on differences in oxygen extraction and/or consumption.^{1,20,21} Cerebral oxygen extraction is inversely correlated to oxygen delivery in preterm infants.^{20,21} In these studies, there was no correlation of changes in mean arterial blood pressure and cerebral fractional oxygen extraction. In our study, there was a trend to lower values in cesarean delivery group, but only at 10 minutes was there a significant difference. As extraction is increased with decreased oxygen content, one would have expected that lower oxygen delivery in the cesarean delivery group (due to significantly lower HR and SpO₂) would result in increased instead decreased extraction values to explain missing differences in rSO₂brain. Because it is the other way around, behavior of rSO₂brain cannot be explained with differences in oxygen extraction. Therefore, observations of our study may be due cerebral autoregulatory response.

Vaginally delivered infants had significantly higher catecholamine levels at birth compared with infants born by cesarean delivery.²² In hemodynamically stable near-term sheep, dopamine infusion was associated with cerebral vasoconstriction.²³ Therefore, besides autoregulation, one further explanation for our observation may be that rSO₂brain after vaginal delivery was reduced as the result of catecholamine-induced vasoconstriction.

The present study has some shortcomings. At the moment, we do not know whether or not rSO₂ of the brain may differ significantly between different regions of the brain. Recently, Lemmers et al²⁴ showed that especially during situations with unstable arterial saturation, differences can exist between the left and right hemispheres. Therefore, we used the same frontoparietal region (left side) in all infants. We calculated the power of the present study. Estimating a mean difference of

10% of rSO₂brain as significant, the probability of type II error is 0.02.

Because we have no detailed information on cardiocirculatory changes, behavior of CBF, and blood gas changes (PaO₂ and PaCO₂), we cannot define the impact of these variables on changes in oxygen saturation. Myocardial function is crucial during hemodynamic transition to extrauterine life. Besides left ventricular output and blood pressure, CBF is furthermore regulated by cerebral autoregulation. Cerebral autoregulation is influenced by changes in blood gas (PaO₂ and PaCO₂). PaCO₂ appears to have even more influence on CBF than mean arterial blood pressure.²⁵ There is a positive correlation between cerebral rSO₂ and PaCO₂ and a negative correlation between fractional tissue oxygen extraction and PaCO₂.²⁶ We saw a steep increase of SpO₂, which certainly has influence on cerebral oxygen extraction,²⁰ but, whether there was a concomitant change in CBF is unknown. In the situation of normal transition, one would expect that the increase in SpO₂ (reassuring improving gas exchange via the lungs) would be accompanied by a decrease in PaCO₂ (resulting in a decrease of rSO₂).

We emphasize that comparisons of absolute values of rSO₂ must be done with caution. The absolute value of regional tissue oxygen saturation lacks the precision²⁷ to be used as a quantitative variable, but it can be used as a trend monitor with helpful clinical information.²⁸ Comparison of values of rSO₂ attained with different technical devices may be problematic.^{29,30} However, there are some comparative studies between the two most used devices, the INVOS system and the NIRO 300 (Hamamatsu, Japan).^{29,30} The standard deviation of the difference between the two monitors was 7.5%, but they showed a very good correlation of the trend values.³⁰ Furthermore, because the calculation of cerebral fractional tissue oxygen extraction values uses cerebral rSO₂ but preductal SpO₂ values, there must be some caution with interpretation.

In conclusion, we show that rSO₂brain remained unchanged in situations with significant changes in HR and SpO₂. Therefore, oxygen delivery to the brain cannot be estimated reliably with SpO₂. Further studies are needed in term and preterm infants to gain normative data from a range of infants. These studies show the potential value of rSO₂ measurements of the brain during neonatal transition and demonstrate their value in the evaluation of supplemental oxygen for resuscitation. ■

We thank the parents for allowing us to study their infants, as well as the midwives, nurses, and physicians involved in the treatment of the neonates. We also thank Professor Andrea Berghold (Institute for Medical Informatics, Statistics and Information, Medical University Graz) for statistical counseling. We also thank Evelyne Ziehenberger for her help in realization of the study.

Submitted for publication Nov 15, 2010; last revision received Jan 27, 2011; accepted Feb 24, 2011.

Reprint requests: Dr Berndt Urlesberger, MD, Abteilung Neonatologie Univ. Klinik für Kinder- und Jugendheilkunde, Auenbruggerplatz 30, 8036 Graz, Austria. E-mail: berndt.urlesberger@medunigraz.at

References

1. Kamlin CO, O'Donnell CP, Davis PG, Morley CJ. Oxygen saturation in healthy infants immediately after birth. *J Pediatr* 2006;148:585-9.
2. Rabi Y, Yee W, Chen SY, Singhal N. Oxygen saturation trends immediately after birth. *J Pediatr* 2006;148:590-4.
3. Dawson JA, Davis PG, O'Donnell CPF, Kamlin COF, Morley CJ. Pulse oximetry for monitoring infants in the delivery room: a review. *Arch Dis Child Fetal Neonatal Ed* 2007;92:F4-7.
4. Sorensen LC, Greisen G. The brains of very preterm newborns in clinically stable condition may be hyperoxygenated. *Pediatrics* 2009;124:958-63.
5. Suzuki S, Takasaki S, Ozaki T, Kobayashi Y. A tissue oxygenation monitor using NIR spatially resolved spectroscopy. *Proc SPIE* 1999;3597:582-92.
6. Verhagen EA, ter Horst HJ, Keating P, Bos AF. The course of cerebral oxygen saturation and oxygen extraction during the first two weeks of life in preterm infants. *Acta Paediatrica* 2007;96(Suppl 456):140.
7. Grossauer K, Pichler G, Schmöller G, Zotter H, Mueller W, Urlesberger B. Comparison of peripheral and cerebral tissue oxygenation index in neonates. *Arch Dis Child Fetal Neonatal Ed* 2009;94:F156.
8. Naulaers G, Morren G, Van Huffel S, Casaer P, Devlieger H. Cerebral tissue oxygenation index in very premature infants. *Arch Dis Child Fetal Neonatal Ed* 2002;87:F189-92.
9. Naulaers G, Morren G, Van Huffel S, Casaer P, Devlieger H. Measurement of tissue oxygenation index during the first three days in premature born infants. *Adv Exp Med Biol*. 2003;510:379-83.
10. van den Berg E, Lemmers PMA, Toet MC, Klaessens JHG, van Bel F. Effect of the "InSurE" procedure on cerebral oxygenation and electrical brain activity of the preterm infant. *Arch Dis Child Fetal Neonatal Ed* 2010;95:F53-8.
11. Isobe K, Kusaka T, Fujikawa Y, Okubo K, Nagano N, Yasuda S, et al. Measurement of cerebral oxygenation in neonates after vaginal delivery and cesarean section using full-spectrum near infrared spectroscopy. *Comp Biochem Physiol. Part A Mol Integr Physiol* 2002;132:133-8.
12. Fauchere JC, Schulz G, Haensse D, Keller E, Ersch J, Bucher HU, et al. Near-infrared spectroscopy measurements of cerebral oxygenation in newborns during immediate postnatal adaptation. *J Pediatr* 2010;156:372-6.
13. Urlesberger B, Grossauer K, Pocivalnik M, Avian A, Müller W, Pichler G. Regional oxygen saturation of the brain and peripheral tissue during birth transition of term infants. *J Pediatr* 2010;157:740-4.
14. Naulaers G, Meyns B, Miserez M, Leunens V, Van Huffel S, Casaer P, et al. Use of tissue oxygenation index and fractional tissue oxygen extraction as non-invasive parameters for cerebral oxygenation: a validation study in piglets. *Neonatology* 2007;92:120-6.
15. Watzman HM, Kurth CD, Montenegro LM, Rome J, Steven JM, Nicolson SC. Arterial and venous contributions to near-infrared cerebral oximetry. *Anesthesiology* 2000;93:947-53.
16. Dawson J, Kamlin CO, Vento M, Wong C, Cole TJ, Donath SM, et al. Defining the reference range for oxygen saturation for infants after birth. *Pediatrics* 2010;125:e1340-7.
17. Dawson JA, Kamlin CO, Wong C, te Pas AB, Vento M, Cole TJ, et al. Changes in heart rate in the first minutes after birth. *Arch Dis Child Fetal Neonatal Ed* 2010;95:F177-81.
18. Camp D, Kotgal U, Kleinman L. Preservation of cerebral autoregulation in the unanesthetized hypoxic newborn dog. *Brain Res* 1982;241:207-13.
19. Jones MD, Jr., Trystman RJ, Simmons MA, Molteni RA. Effects of changes in arterial O₂ content on cerebral blood flow in the lamb. *Am J Physiol* 1981;240:H209-15.
20. Kissack CM, Garr R, Wardle SP, Weindling AM. Cerebral fractional oxygen extraction is inversely correlated with oxygen delivery in the sick, newborn, preterm infant. *J Cereb Blood Flow Metab* 2005;25:545-53.
21. Kissack CM, Garr R, Wardle SP, Weindling AM. Cerebral fractional oxygen extraction in very low birth weight infants is high when there is low left ventricular output and hypocarbia but is unaffected by hypotension. *Pediatr Res* 2004;55:400-5.
22. Agata Y, Hiraishi S, Misawa H, Han JH, Oguchi K, Horiguchi Y, et al. Hemodynamic adaptations at birth and neonates delivered vaginally and by Cesarean section. *Biol Neonate* 1995;68:404-11.
23. Gleason CH, Robinson R, Harris A, Maycock D, Trystman R. Cerebrovascular effects of intravenous dopamine infusions in fetal sheep. *J Appl Physiol* 2002;92:717-24.
24. Lemmers P, Van Bel F. Left-to-right differences of regional cerebral oxygen saturation and oxygen extraction in preterm infants during the first days of life. *Pediatr Res* 2009;65:226-30.
25. Noori S, Stavroudis TA, Seri I. Systemic and cerebral hemodynamics during the transitional period after premature life. *Clin Perinatol* 2009;36:723-36.
26. Vanderhaegen J, Naulaers G, Vanhole C, de Smet D, van Huffel S, Vanhaesbrouck S, et al. The effect of changes in tPCO₂ on the fractional tissue oxygen extraction—as measured by near-infrared spectroscopy—in neonates during the first days of life. *J Eur Paediatr Neurol* 2009;13:128-34.
27. Sorensen LC, Greisen G. Precision of measurement of cerebral tissue oxygenation index using near-infrared spectroscopy in preterm neonates. *J Biomed Optics* 2006;11:054005.
28. van Bel F, Lemmers P, Naulaers G. Monitoring neonatal regional cerebral oxygen saturation in clinical practice: value and pitfalls. *Neonatology* 2008;94:237-44.
29. Thavasothy M, Broadhead M, Elwell C, Peters M, Smith M. A comparison of cerebral oxygenation as measured by the NIRO 300 and the INVOS 5100 near infrared spectrophotometers. *Anaesthesia* 2002;57:999-1006.
30. Gagnon RE, Macnab A, Gagnon F, Blackstock D, LeBlanc J. Comparison of two spatially resolved NIRS oxygenation indices. *J Clin Monit Comput* 2002;17:385-91.

Diskussion

Beide vorgelegten Publikationen beinhalten eine ausführliche Diskussion, sodass dieser Teil lediglich eine kleine Zusammenfassung beinhaltet und kürzlich erschienene Publikationen auf diesem Gebiet erwähnt.

In der Literatur findet man nur wenige direkt vergleichbare Studien. Beide vorgelegten Arbeiten beschäftigen sich mit den physiologischen Veränderungen der zerebralen Oxygenierung unmittelbar nach der Geburt. In Publikation I handelt es sich um gesunde Neugeborene (n=63) nach Spontangeburt. „Der Grund für die wenig vorhandenen Studien bei spontan geborenen Kindern ist am ehesten aus ethischen und organisatorischen Gründen gegeben. Um diese Messungen zu erhalten, muss die Intimsphäre einer Geburt zu einem gewissen Grad gestört werden. Es bedarf vieler organisatorischer, persönlicher und technischer Faktoren, um eine solche Studie durchzuführen zu können.“¹

In Publikation II bestand das Patientenkollektiv zusätzlich auch aus gesunden NG nach Kaiserschnitt. Diese werden ohnehin primär zur Erstversorgung vom Kinderarzt betreut, und damit räumlich von der Mutter getrennt. Somit entstanden keine zusätzlichen Unterbrechungen bzw. Störfaktoren der Mutter-Kind-Interaktion während der ersten 15 Lebensminuten. Die Väter durften nach einigen Minuten ihr NG sehen und nach der Messphase zu sich nehmen.

Zusammengefasst konnte Publikation I einen signifikanten und raschen Anstieg der rSO₂brain in Lebensminute (LM) 1-5 zeigen. Die SaO₂ hingegen zeigte einen langsameren signifikanten Anstieg von LM 2-15.

In Publikation II wurden SaO₂, rSO₂brain, HF und FTOE Werte in den ersten 15 LM bei NG bezüglich ihres Geburtsmodus (Kaiserschnitt versus Spontanpartus) miteinander verglichen. Dabei konnte gezeigt werden, dass NG nach Kaiserschnitt (n=43) zwischen LM 4-8 signifikant niedrigere SaO₂ Werte sowie eine niedrigere HF während des gesamten Messzeitraumes hatten. Bezüglich der rSO₂brain konnte jedoch kein Unterschied zwischen den Gruppen beobachtet werden.

Eine erst kürzlich veröffentlichte Arbeit von Almaazmi et al.¹³ verglich ebenfalls die zerebrale Oxygenierung bei NG in den ersten 10 LM nach Kaiserschnitt (n=22), Spontanpartus (n=20) und Vakuumextraktion (n=2). In dieser Publikation konnte ebenfalls kein Unterschied bezüglich der zerebralen Oxygenierung zwischen den 3 Gruppen festgestellt

werden. Die mediane zerebrale Gewebssättigung betrug in LM 2 36-42% und stieg auf 62-77% in LM 8 an.

Fuchs et al.¹⁴ beschrieb erstmals die zerebrale Oxygenierung bei 51 extrem Frühgeborenen (medianes Gestationsalter 27,8 Schwangerschaftswoche). In diesem Kollektiv konnte ein Anstieg von rSO₂brain-Werte von 37% am Ende der LM 1 auf 61-84% in LM 7 beobachtet werden. Damit konnten die Autoren erstmals zeigen, dass auch bei extrem Frühgeborenen die Messung der zerebrale Oxygenierung in den ersten Lebensminuten möglich ist.

„Faucheré et al.¹² untersuchten mit Hilfe einer anderen NIRS-Technik 20 NG nach Kaiserschnitt und konnten auch ein rSO₂brain Plateau nach 7 Minuten zeigen. Isobe et al.¹¹ beschrieben in ihrer Studie die Messung von 7 NG nach Spontangeburt. In dieser Studie wurde aber wiederum eine andere Messart verwendet, sodass diese nur indirekt miteinander vergleichbar sind. Sie beschreiben ebenfalls einen Anstieg der zerebralen Gewebssättigung mit nachfolgendem Plateau 8,5 Minuten nach der Geburt.“¹

„Die Interpretation und die Vergleichbarkeit verschiedener Geräte bei der Messung der regionalen Sättigung muss mit Vorsicht geschehen¹⁵. Die bisher entwickelten Technologien weisen eine schlechte Präzision auf¹⁶. Van Bel et al.¹⁷ verfassten eine Arbeit über den Wert und die Mängel des Monitorings von der regionalen zerebralen Geweboxygenierung bei NG. Aufgrund der großen intra- und interpersonellen Variabilität kann diese Technik nicht als robuste quantitative Messung der zerebralen Oxygenierung herangezogen werden. Verwendet man diese Technik jedoch bloß als Trendmonitoring des individuellen Patienten, könnte dies hilfreiche klinische Informationen bringen¹⁷. Auch der Vergleich verschiedener Studien, die rSO₂brain Werte erheben, ist aufgrund der verschiedenen verwendeten Technologien sehr schwierig¹⁸.“¹ Die unterschiedlichen Technologien basieren auf verschiedenen Berechnungen bezüglich arteriellen und venösen Sättigungsanteilen^{19,20} sowie unterschiedlich großen Sensoren²¹.

Eine neue Technologie mit Laserlicht (FORE-SIGHT) verspricht eine bessere Übereinstimmung der zerebral gemessenen Werte mit invasiv gemessenen.^{22,20}

Zusammengefasst kann man sagen, dass mittels NIRS eine niedrige zerebrale oder andere Gewebssättigung nicht-invasiv detektiert werden kann. Dies kann hilfreiche klinische Hinweise auf eine niedrige kardiale Auswurfleistung geben bzw. eine niedrige Perfusion des Gewebes anzeigen²³.

Gerade in der Erstversorgung kranker Neu- und Frühgeborener gibt es weltweit noch keine einheitlichen Richtlinien, welche genauen Sauerstoffsättigungen erreicht werden sollen und

die damit verbundene Menge an supplementärem Sauerstoff. Die Kurz- und Langzeitfolgen von Hyperoxygenierung in der Erstversorgung sind noch nicht ausreichend erforscht.

Inwieweit die zerebrale Geweboxygenierung ihren Stellenwert in der diagnostischen Routine finden wird, ist bisher unklar. Dafür sind noch viele Studien in den nächsten Jahren notwendig. Viele Arbeitsgruppen beschäftigen sich aktuell mit der Erstellung der Normwerte für die verschiedenen Altersgruppen und Krankheitsbilder in der Neonatologie.

Danksagung

Zuerst möchte ich meinem Doktorvater PD Dr. Andreas Flemmer dafür danken meine kumulative Doktorarbeit zu betreuen. Ich freue mich auf weitere Forschungsprojekte mit ihm in der nahen Zukunft.

Mein größter Dank gilt meinem Forschungsgruppenleiter Univ. Prof. Dr. Berndt Urlesberger. Die wertvollen Erfahrungen, die ich im Forschungsteam sowie im Kreißsaal machen durfte, waren für meinen weiteren persönlichen und beruflichen Werdegang sehr wichtig und hilfreich.

Ein ganz großer Dank gilt all den Familien und Neugeborenen, die uns ihr Vertrauen geschenkt haben und sich bereit erklärt haben an den Studien teilzunehmen.

Ganz besonders bedanken möchte ich mich bei meinem Ehemann und meiner gesamten Familie, die mich stets unterstützen und mich auf meinem Lebensweg begleiten.

Literaturverzeichnis

-
- ¹ Kratky E, Zerebrale Oxygenierung bei reifen Neugeborenen in den ersten fünfzehn Lebensminuten nach Spontangeburt. Diplomarbeit 2011, Medizinische Universität Graz;
- ² House JT, Schultetus RR, Gravenstein N. Continuous neonatal evaluation in the delivery room by pulse oximetry. *J Clin Monit* 1987;3:96-100.
- ³ Toth B, Becker A, Seelbach-Gobel B. Oxygen saturation in healthy newborn infants immediately after birth measured by pulse oximetry. *Arch Gynecol Obstet*. 2002 Apr;266(2):105-7.
- ⁴ Kamlin CO, O'Donell CP, Davis PG, Morley CJ. Oxygen saturation in healthy infants immediately after birth. *J Pediatr* (2006) vol. 148 (5) pp. 585-9.
- ⁵ Rabi Y, Yee W, Chen SY, Singhal N. Oxygen saturation trends immediately after birth. *J Pediatr* (2006) vol. 148 (5) pp. 590-4.
- ⁶ Finer N, Leone T. Oxygen saturation monitoring for the preterm infant: the evidence basis for current practice. *Pediatr Res* (2009) vol. 65 (4) pp. 375-80.
- ⁷ Dawson JA, Davis PG, O'Donell CP, Kamlin CO. Pulse oximetry for monitoring infants in the delivery room: a review. *Arch Dis Child Fetal Neonatal Ed* (2007) vol. 92 (1) pp. F4-7.
- ⁸ Mariani G, Dik PB, Ezquer A, Aguirre A, Esteban ML, Perez C, Fernandez Jonusas S, Fustiñana C. Pre-ductal and post-ductal O₂ saturation in healthy term neonates after birth. *J Pediatr*. 2007 Apr;150(4):418-21.
- ⁹ Jöbsis FF. Noninvasive, infrared monitoring of cerebral and myocardial oxygen sufficiency and circulatory parameters. *Science* 1977 Dec 23;198(4323):1264-7.
- ¹⁰ Sorensen LC, Greisen G. The brains of very preterm newborns in clinically stable condition may be hyperoxygenated. *Pediatrics*. 2009 Nov;124(5):e958-63. Epub 2009 Oct 19.
- ¹¹ Isobe K, Kusaka T, Fujikawa Y, Okubo K, Nagano K, Yasuda S, Kondo M, Itoh S, Hirao K, Onishi S. Measurement of cerebral oxygenation in neonates after vaginal delivery and cesarean section using full-spectrum near infrared spectroscopy. *Comp Biochem Physiol, Part A Mol Integr Physiol* (2002) vol. 132 (1) pp. 133-8.
- ¹² Fauchère JC, Schulz G, Haensse D, Keller E, Ersch J, Bucher HU, Wolf M. Near-infrared spectroscopy measurements of cerebral oxygenation in newborns during immediate postnatal adaptation. *J Pediatr* (2010) vol. 156 (3) pp. 372-6.

-
- ¹³ Almaazmi M, Schmid MB, Havers S, Reister F, Lindner W, Mayer B, Hummeler HD, Fuchs H. Cerebral Near-Infrared Spectroscopy during Transition of Healthy Term Newborns. *Neonatology.* 2013 Feb;20;103(4):246-251.
- ¹⁴ Fuchs H, Lindner W, Buschko A, Almazam M, Hummeler HD, Schmid MB. Brain oxygenation monitoring during neonatal resuscitation of very low birth weight infants. *J Perinatol.* 2012 May;32(5):356-62. doi: 10.1038/jp.2011.110. Epub 2011 Aug 18.
- ¹⁵ Ferrari M, Mottola L, Quaresima V. Principles, techniques, and limitations of near infrared spectroscopy. *Can J Appl Physiol* 2004 Aug;29(4):463-87.
- ¹⁶ Sorensen LC, Greisen G. Precision of measurement of cerebral tissue oxygenation index using near-infrared spectroscopy in preterm neonates. *J Biomed Opt* (2006) vol. 11 (5) pp. 054005.
- ¹⁷ van Bel F, Lemmers P, Naulaers G. Monitoring neonatal regional cerebral oxygen saturation in clinical practice: value and pitfalls. *Neonatology* (2008) vol. 94 (4) pp. 237-44.
- ¹⁸ Gagnon RE, Macnab AJ, Gagnon FA, Blackstock D, JeBlanc JG. Comparison of two spatially resolved NIRS oxygenation indices. *J Clin Monit Comput* (2002) vol. 17 (7-8) pp. 385-91.
- ¹⁹ Benni PB, Chen B, Dykes FD, Wagoner SF, Heard M, Tanner AJ, Young TL, Rais-Bahrami K, Rivera O, Short BL. Validation of the CAS neonatal NIRS system by monitoring vv-ECMO patients: preliminary results. *Adv Exp Med Biol.* 2005;566:195-201.
- ²⁰ Rais-Bahrami K, Rivera O, Short BL. Validation of a noninvasive neonatal optical cerebral oximeter in veno-venous ECMO patients with a cephalad catheter. *J Perinatol.* 2006 Oct;26(10):628-35. Epub 2006 Aug 10.
- ²¹ Morris N, Pichler G, Pocivalnik M, Brandner A, Müller W, Urlesberger B. Cerebral regional oxygen saturation (crSO₂): are different sensors comparable? *Paediatr Anaesth.* 2012 Jun 12. doi: 10.1111/j.1460-9592.2012.03895.x.
- ²² MacLeod DB, Ikeda K, Keifer J, Moretti E, Ames W. Validation of the CAS adult cerebral oximeter during hypoxia in healthy volunteers. *ANESTH ANALG* 2006; 102; S-162
- ²³ Sorensen LC, Maroun LL, Borch K, Lou HC, Greisen G. Neonatal cerebral oxygenation is not linked to foetal vasculitis and predicts intraventricular haemorrhage in preterm infants. *Acta Paediatr.* 2008 Nov;97(11):1529-34. doi: 10.1111/j.1651-2227.2008.00970.x. Epub 2008 Jul 31.