

Monitoraggio della profondità dell'anestesia

Zaccaria Ricci

Azienda Ospedaliero-Universitaria Meyer, Firenze



European Society of
Regional Anaesthesia
& Pain Therapy
ESRA ITALIA



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CONGRESSO
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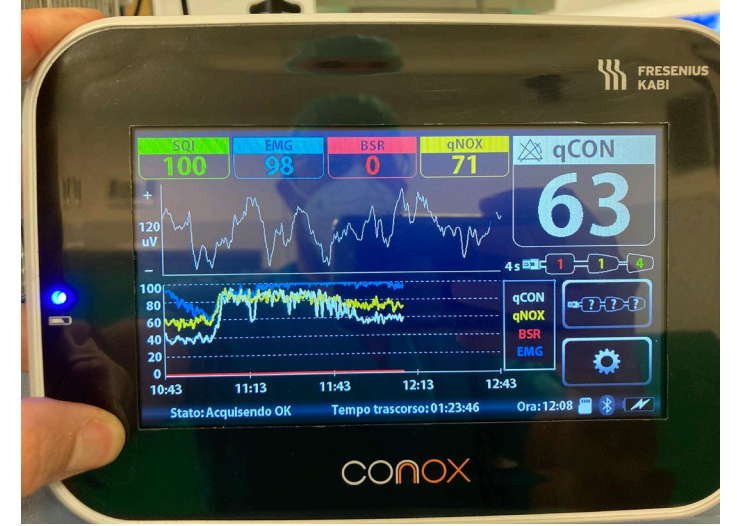
*7-9
Novembre
2024*

Presidenti del Congresso:
Vanni Agnoletti
Domenico Pietro Santonastaso
Andrea Tognù











PROCESSED EEG

Proprietary (secret) algorithm gets the necessary EEG information and digests them in order to provide a single resuming number

A Narrative Review Illustrating the Clinical Utility of Electroencephalogram-Guided Anesthesia Care in Children

Choon Looi Bong, FRCA,* Gustavo A. Balanza, MD,† Charis Ern-Hui Khoo, FANZCA,*
Josephine Swee-Kim Tan, MMed (Anaes),* Tenzin Desel, BA,† and Patrick Lee Purdon, PhD†

1) Avoiding oversedation

- Children undergoing general anesthesia with (and without) NMBA
- Children with Atypical Neurodevelopment or Neuropsychiatric Disorders
- Children with Altered Levels of Consciousness Before Induction of Anesthesia
- Neonates and Infants

2) Special circumstances

- TIVA
- Hemodynamic instability
- CPB
- Brain states (i.e., ketamine vs GABAergic vs alpha2)
- [Children with locoregional anesthesia]



Neuromonitoring in paediatric anaesthesia

Andrew Davidson^{a,b} and Justin Skowno^{c,d}

- There are few studies, which demonstrate better outcomes in paediatric anaesthesia with EEG-derived depth monitors in children and there is no evidence that any particular EEG-derived depth monitor is superior in children.
- The EEG during anaesthesia is fundamentally different in infants and children.

Proprietary algorithms are made on adults' neurophysiology. Go back to basics

Normal Adult Brain Waves

Awake with
mental activity



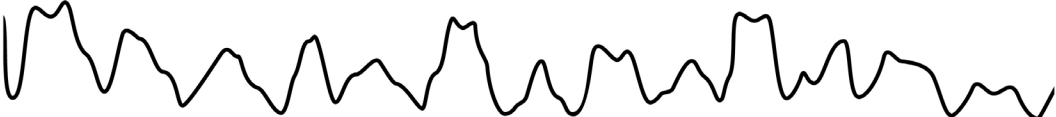
Beta
14-30 Hz

Awake and
resting



Alpha
8-13 Hz

Sleeping

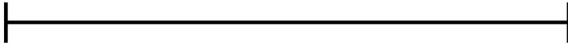


Theta
4-7 Hz

Deep sleep



Delta
<3.5 Hz



1 sec

Anesthesiology 2015; 123:937–60

Clinical Electroencephalography for Anesthesiologists

Part I: Background and Basic Signatures

Patrick L. Purdon, Ph.D., Aaron Sampson, B.S., Kara J. Pavone, B.S., Emery N. Brown, M.D., Ph.D.

GABA agonist drugs

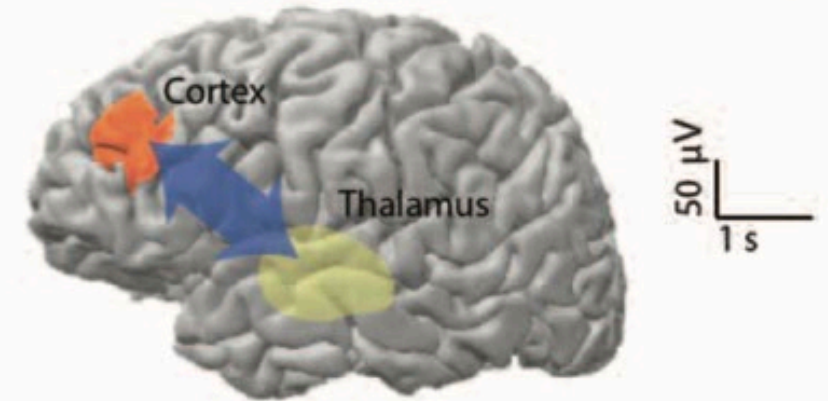
During consciousness, there is broadband communication between the thalamus and the frontal cortex with beta and gamma activity in the electroencephalogram.

Modeling studies suggest that during propofol-induced unconsciousness the spatially coherent alpha oscillations are highly structured rhythms in thalamocortical circuits

We postulate that the highly organized coherent alpha oscillations most likely prevent normal communications between the thalamus and cortex, whereas the incoherent slow oscillations represent an impediment to normal intracortical communication

B Thalamocortical Circuits Underlying Propofol-Induced Alpha Oscillations

Conscious

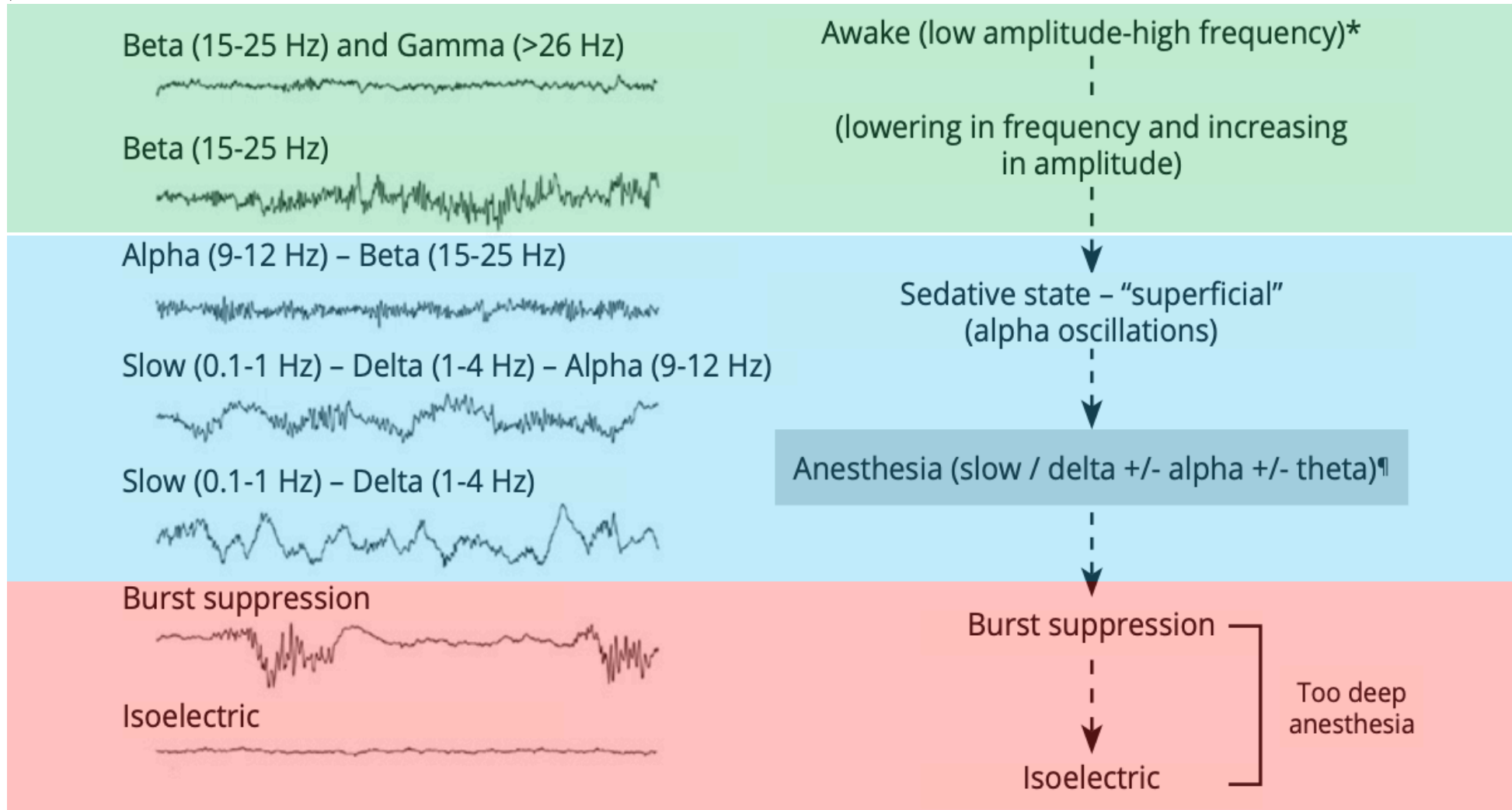


Anesthesiology 2015; 123:937–60

Clinical Electroencephalography for Anesthesiologists

Part I: Background and Basic Signatures

Patrick L. Purdon, Ph.D., Aaron Sampson, B.S., Kara J. Pavone, B.S., Emery N. Brown, M.D., Ph.D.



AVOIDING OVERSEDATION: CHILDREN UNDERGOING GA

Received: 25 March 2023 | Revised: 5 June 2023 | Accepted: 6 June 2023

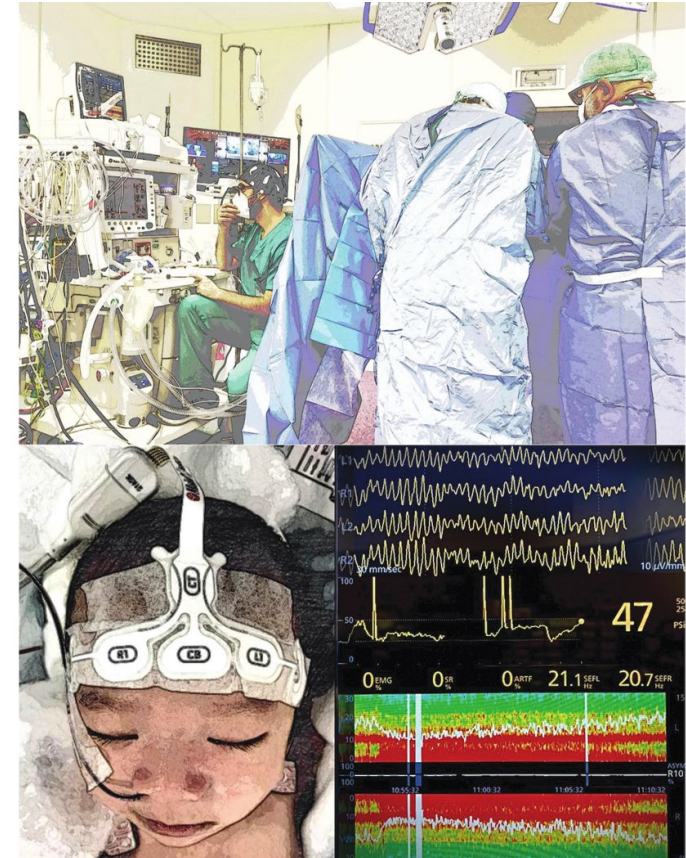
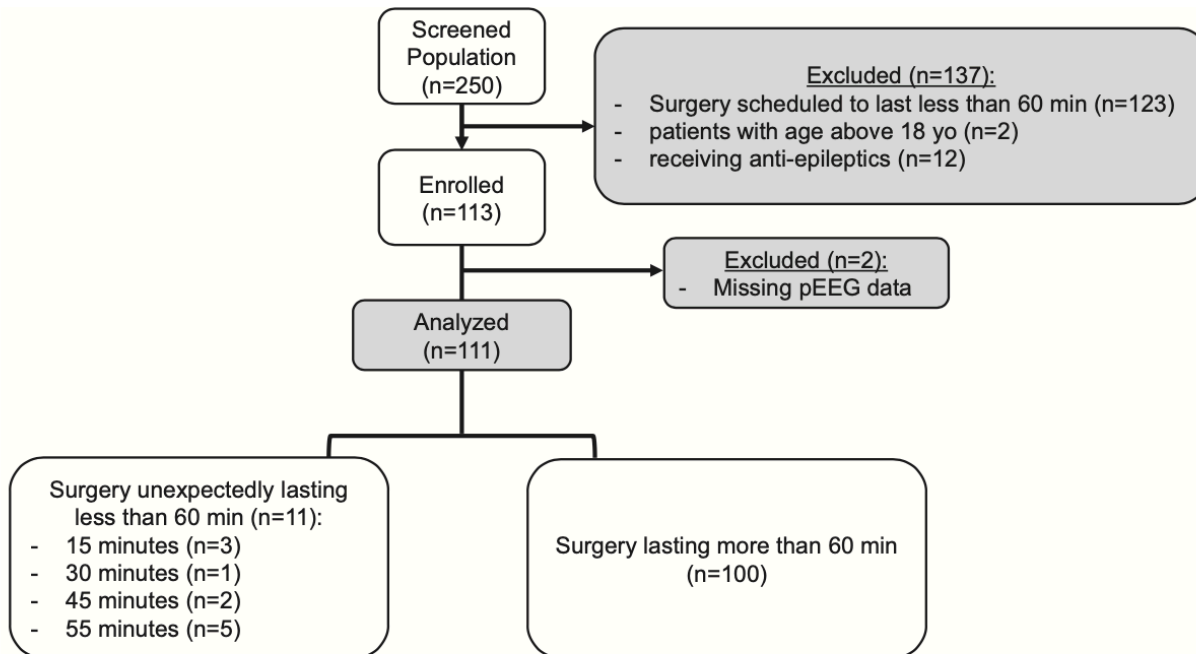
DOI: 10.1111/pan.14711

RESEARCH REPORT

Pediatric Anesthesia WILEY

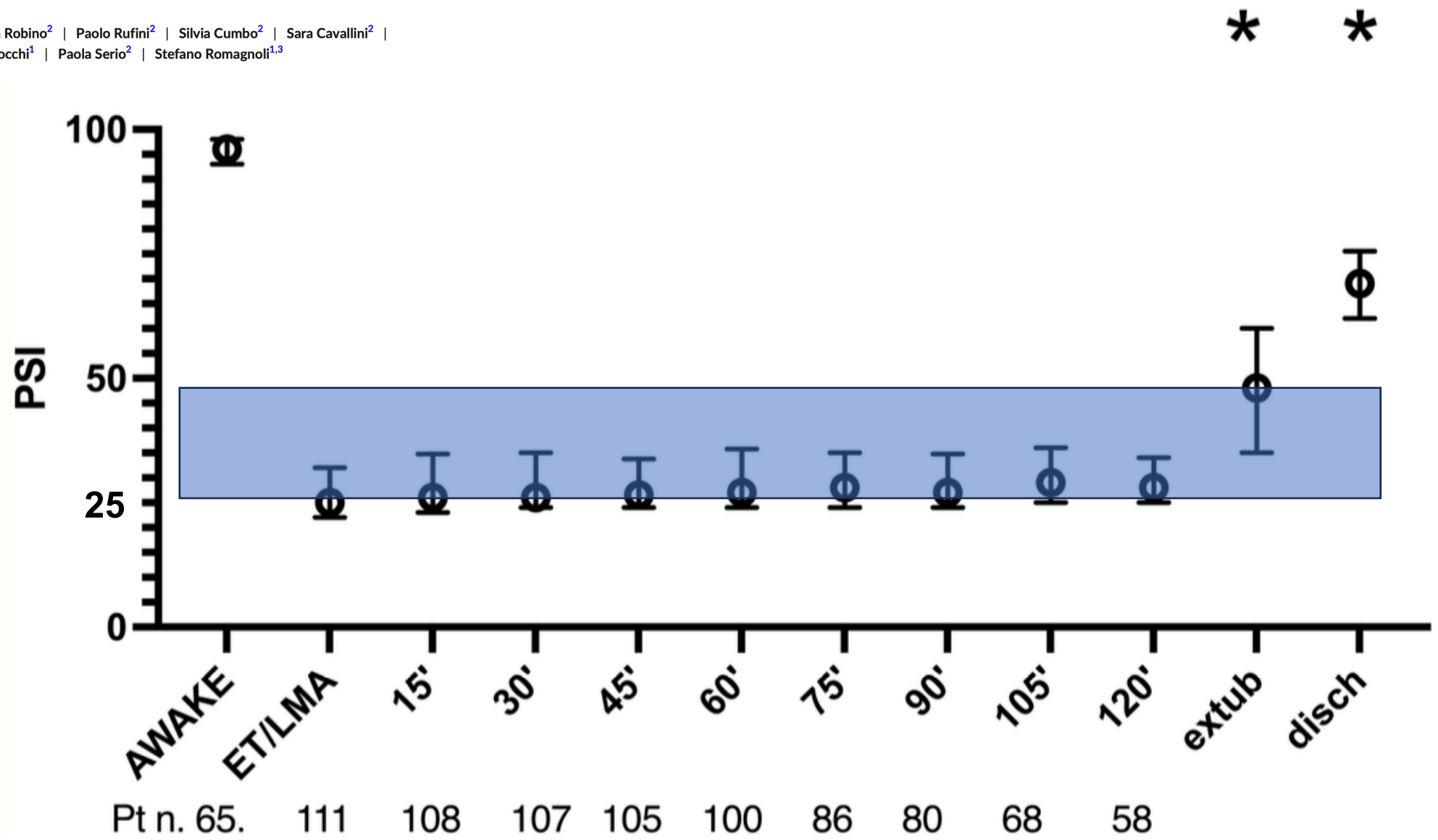
Monitoring anesthesia depth with patient state index during pediatric surgery

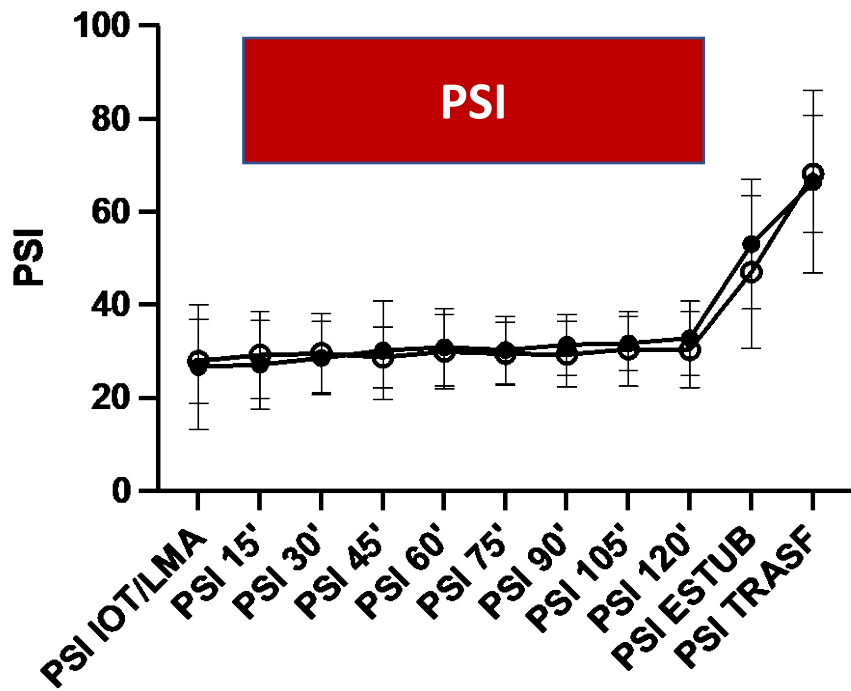
Zaccaria Ricci^{1,2} | Chiara Robino² | Paolo Rufini² | Silvia Cumbo² | Sara Cavallini² | Lorenzo Gobbi¹ | Agata Brocchi¹ | Paola Serio² | Stefano Romagnoli^{1,3}



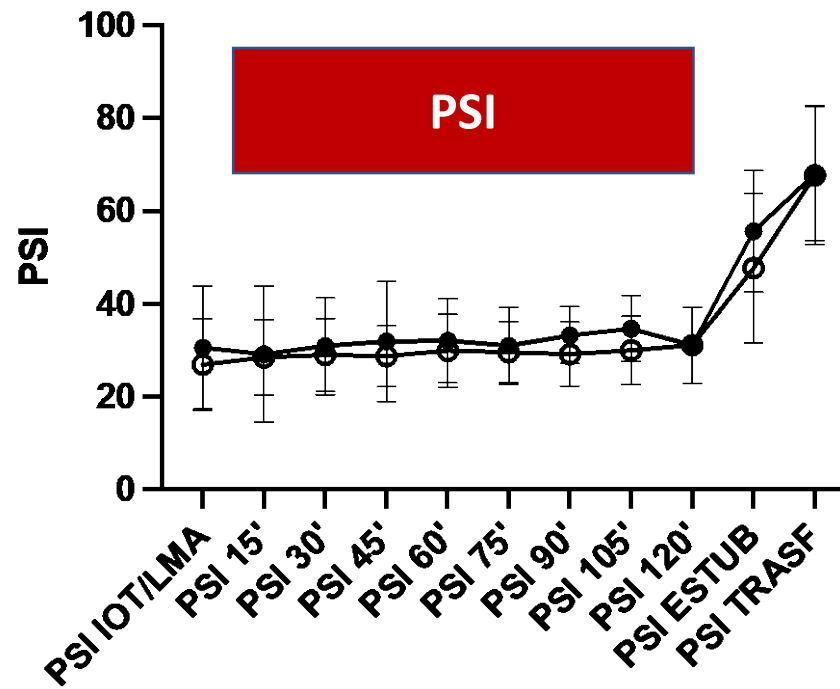
Monitoring anesthesia depth with patient state index during pediatric surgery

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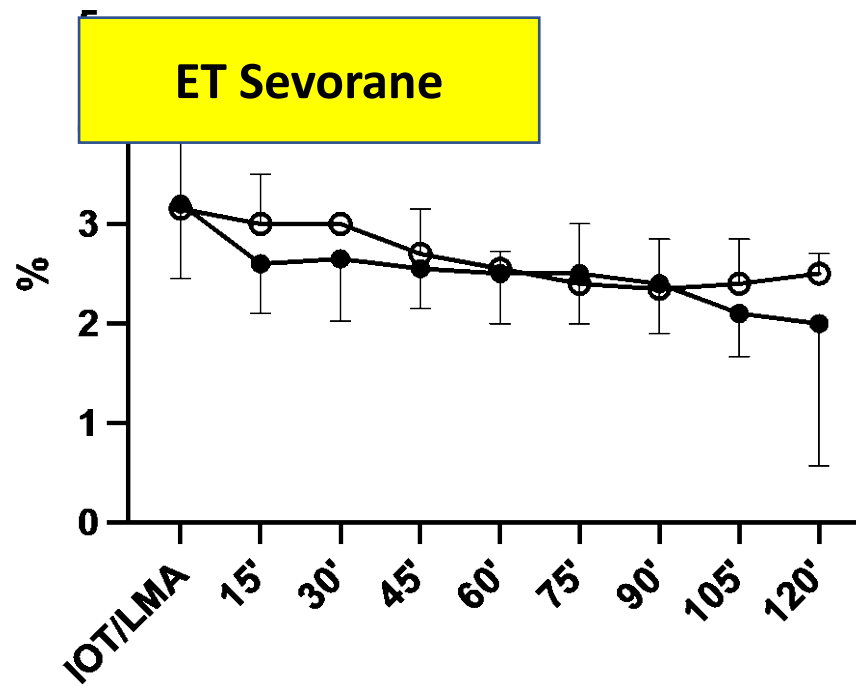




● AG
○ ALR



○ INHAL
● TIVA



○ perid
● gener

VLS?
OPEN?
ROBOT?



Prevalence of Isoelectric Electroencephalography Events in Infants and Young Children Undergoing General Anesthesia

Ian Yuan, MD,* William P. Landis, BS,* Alexis A. Topjian, MD,* Nicholas S. Abend, MD,† Shih-Shan Lang, MD,‡ Jimmy W. Huh, MD,* Matthew P. Kirschen, MD, PhD,*† Janel L. Mensinger, PhD,§ Bingqing Zhang, MPH,* and Charles D. Kurth, MD*

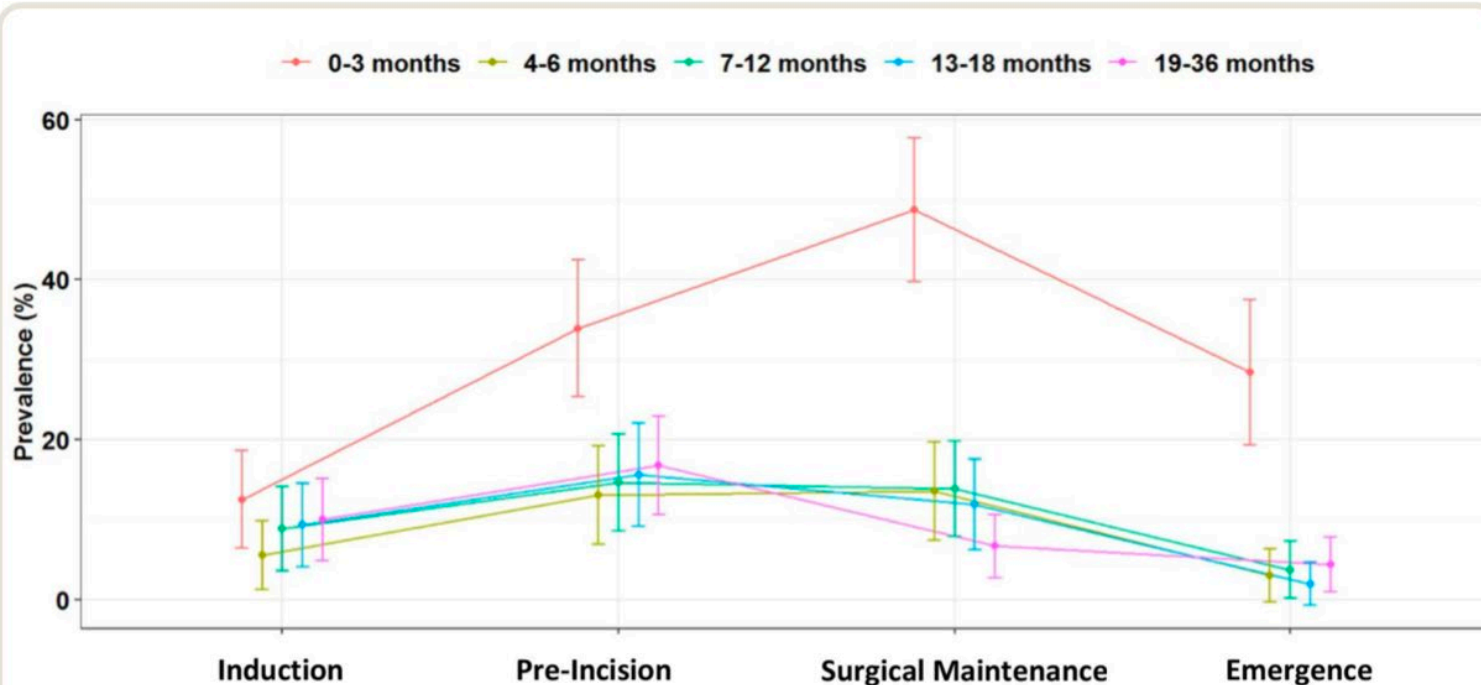


Fig. 3. Prevalence of isoelectric electroencephalography stratified by age groups. Median (dot) and 95% CI (vertical lines) displayed.

ANESTHESIOLOGY

Isoelectric Electroencephalography in Infants and Toddlers during Anesthesia for Surgery: An International Observational Study

Ian Yuan, M.D., Ting Xu, M.D., Justin Skowno, M.B.Ch.B., Ph.D., Bingqing Zhang, M.P.H., Andrew Davidson, M.B.B.S., M.D., Ph.D., Britta S. von Ungern-Sternberg, M.D., Ph.D., David Sommerfield, M.D., Jianmin Zhang, M.D., Xingrong Song, M.D., Ph.D., Mazhong Zhang, M.D., Ph.D., Ping Zhao, M.D., Ph.D., Huacheng Liu, M.D., Ph.D., Yifei Jiang, M.D., Ph.D., Yunxia Zuo, M.D., Ph.D., Jurgen C. de Graaff, M.D., Ph.D., Laszlo Vutskits, M.D., Ph.D., Vanessa A. Olbrecht, M.D., M.B.A., Peter Szmuk, M.D., Charles D. Kurth, M.D., for the BRAIN Collaborative Investigators*

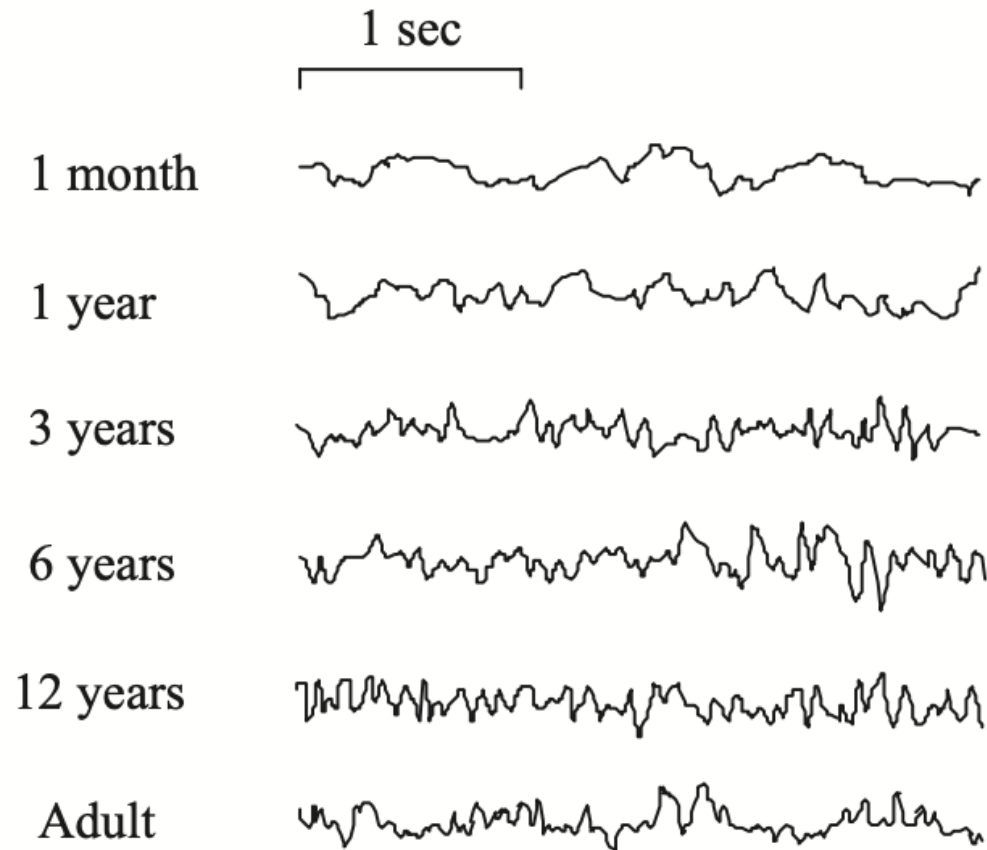
ANESTHESIOLOGY 2022; 137:187-200

REVIEW ARTICLE

The EEG signal: a window on the cortical brain activity

Isabelle Constant & Nada Sabourdin

Department of Anesthesiology, Armand Trousseau Hospital, AP-HP, UPMC, Paris, France



Developmental stages in detail:

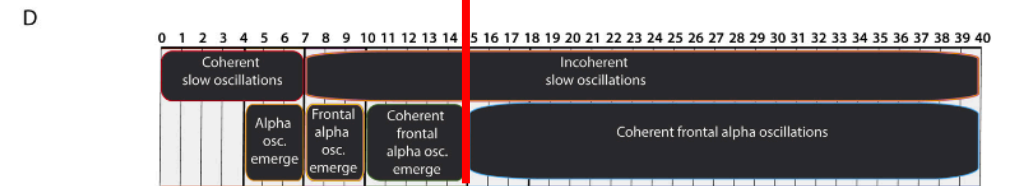
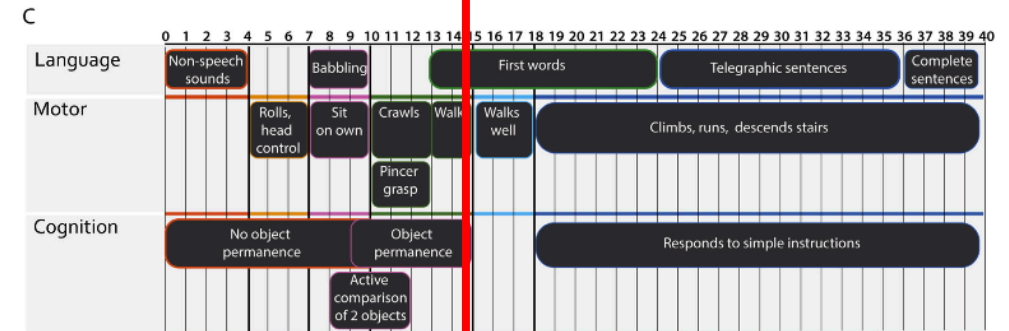
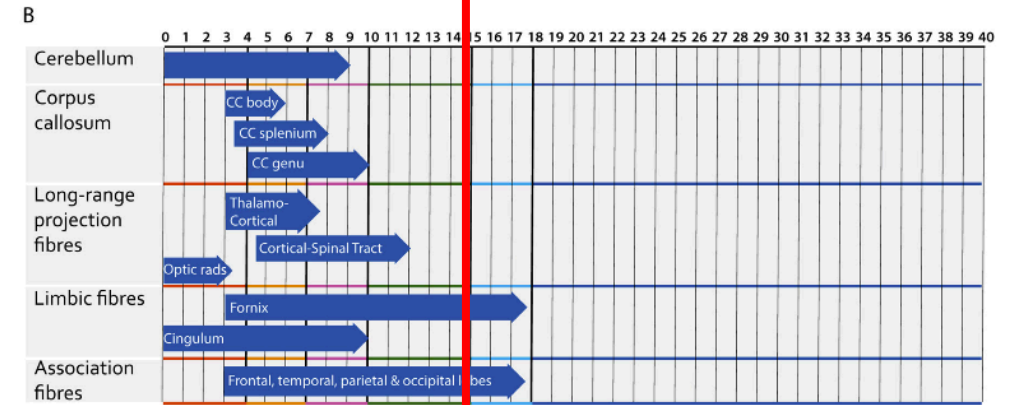
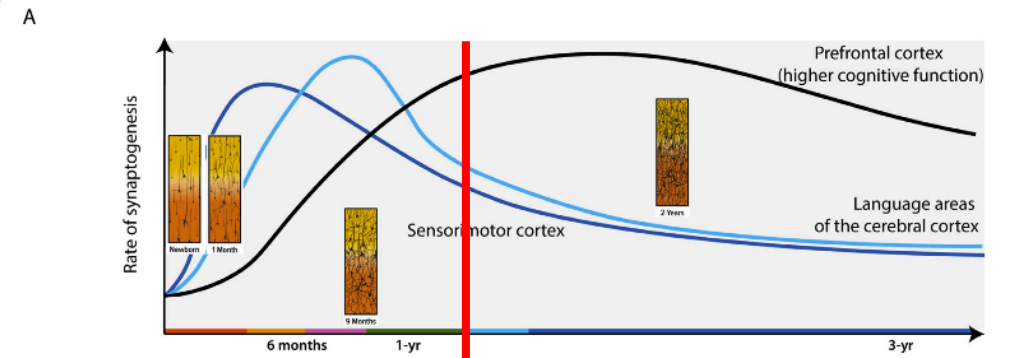
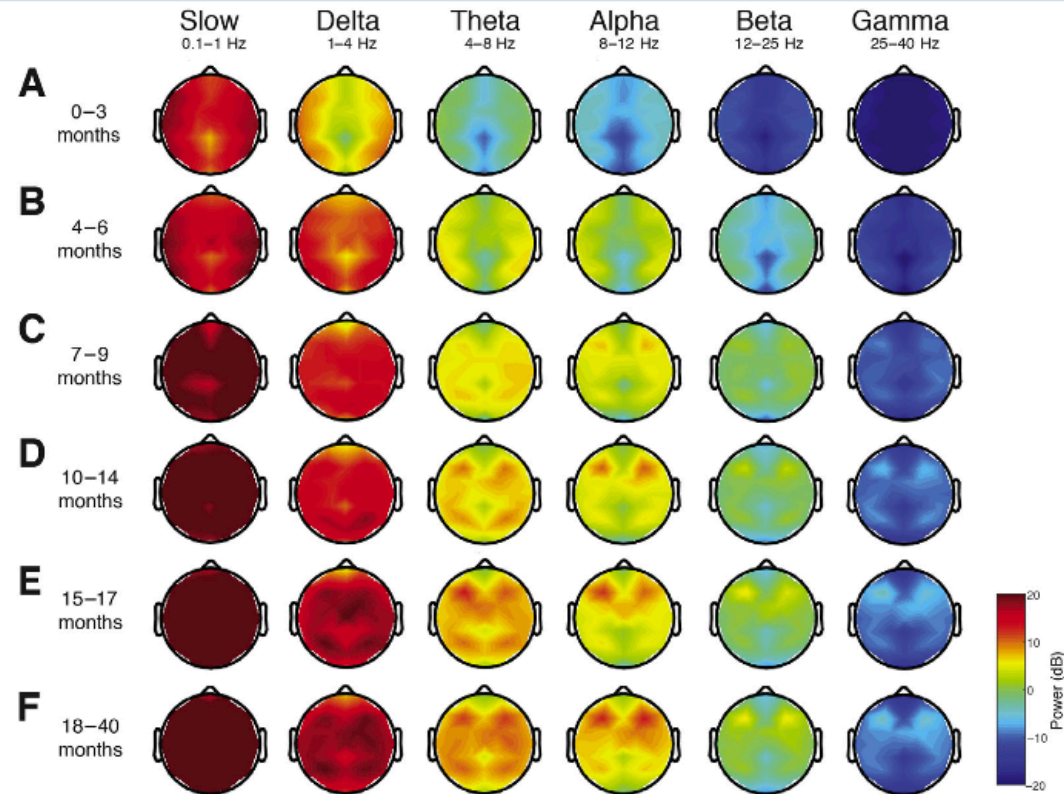
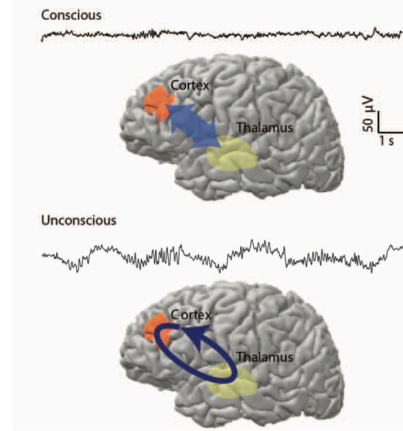
- ✓ From birth to 2 months: Sleep spindles appear.
- ✓ From 3 to 5 months: Parieto-occipital sinusoidal activity appears, announcing alpha rhythm, with increasing frequency: from 4–6 Hz initially to 8–9 Hz by the age of 3.
- ✓ In the normal child aged 3 years and older: The alpha parieto-occipital rhythm, initially discreet and slow (8 Hz), with high voltage and asymmetry. Theta rhythm is abundant, diffuse, and mixed with alpha in the posterior leads and predominates. Slow waves appear, grouped in bursts, and become rhythmic at 2–3 Hz.
- ✓ From 3 to 10 years: Alpha rhythm becomes more important and abundant

Electroencephalographic markers of brain development during sevoflurane anaesthesia in children up to 3 years old

L. Cornelissen^{1,2,*}, S. E. Kim^{3,5}, J. M. Lee^{1,3}, E. N. Brown^{2,3,4}, P. L. Purdon^{2,4} and C. B. Berde^{1,2}

¹Department of Anesthesiology, Critical Care and Pain Medicine, Boston Children's Hospital, Boston, MA, USA, ²Department of Anaesthesia, Harvard Medical School, Boston, MA, USA, ³Department of Brain and Cognitive Sciences, Massachusetts Institute of Technology, Cambridge, MA, USA and ⁴Department of Anesthesia, Critical Care and Pain Medicine, Massachusetts General Hospital, Harvard Medical School, Boston, MA, USA

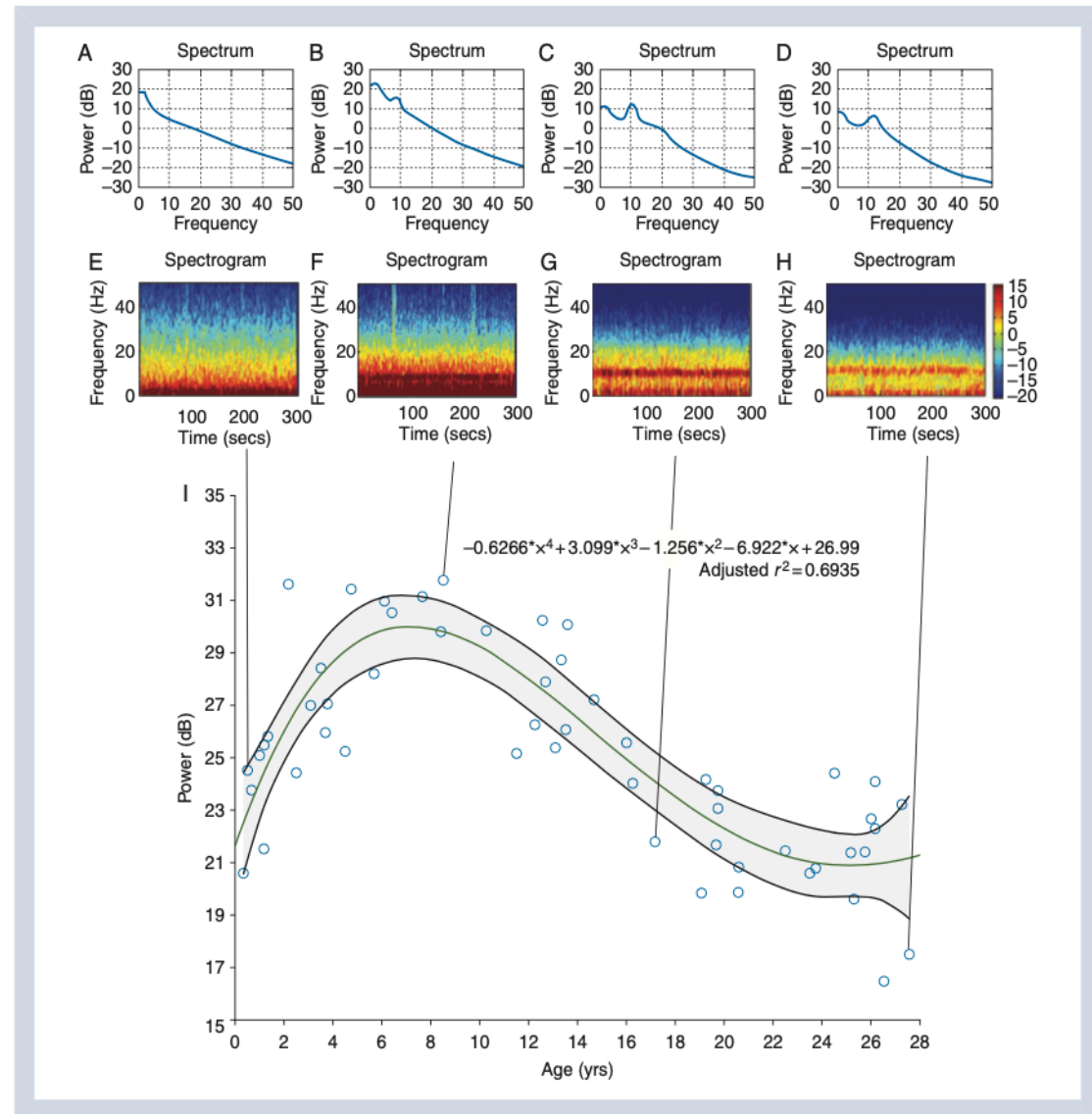
B Thalamocortical Circuits Underlying Propofol-Induced Alpha Oscillations



Age-dependency of sevoflurane-induced electroencephalogram dynamics in children

O. Akeju^{1,3,*}, K. J. Pavone^{1,†}, J. A. Thum^{3,5,†}, P. G. Firth^{1,2}, M. B. Westover^{2,3}, M. Puglia^{1,3}, E. S. Shank^{1,3}, E. N. Brown^{1,3,4,5,6}, and P. L. Purdon^{1,3,4,*}

- EEG recorded during routine care of patients between 0 and 28 yr of age (n=54), using **power spectrum** and **coherence** methods.
- The power spectrum quantifies the energy in the EEG at each frequency, while the coherence measures the frequency-dependent correlation or synchronization between EEG signals at different scalp locations.
- 5 age groups: <1 yr old (n=4), 1–6 yr old (n=12), >6–14 yr old (n=14), >14–21 yr old (n=11), >21–28 yr old (n=13).
- **EEG power significantly increased from infancy through ~6 yr, subsequently declining to a plateau at approximately 21 yr. Alpha (8–13 Hz) coherence, a prominent EEG feature associated with sevoflurane-induced unconsciousness in adults, is absent in patients <1 yr.**

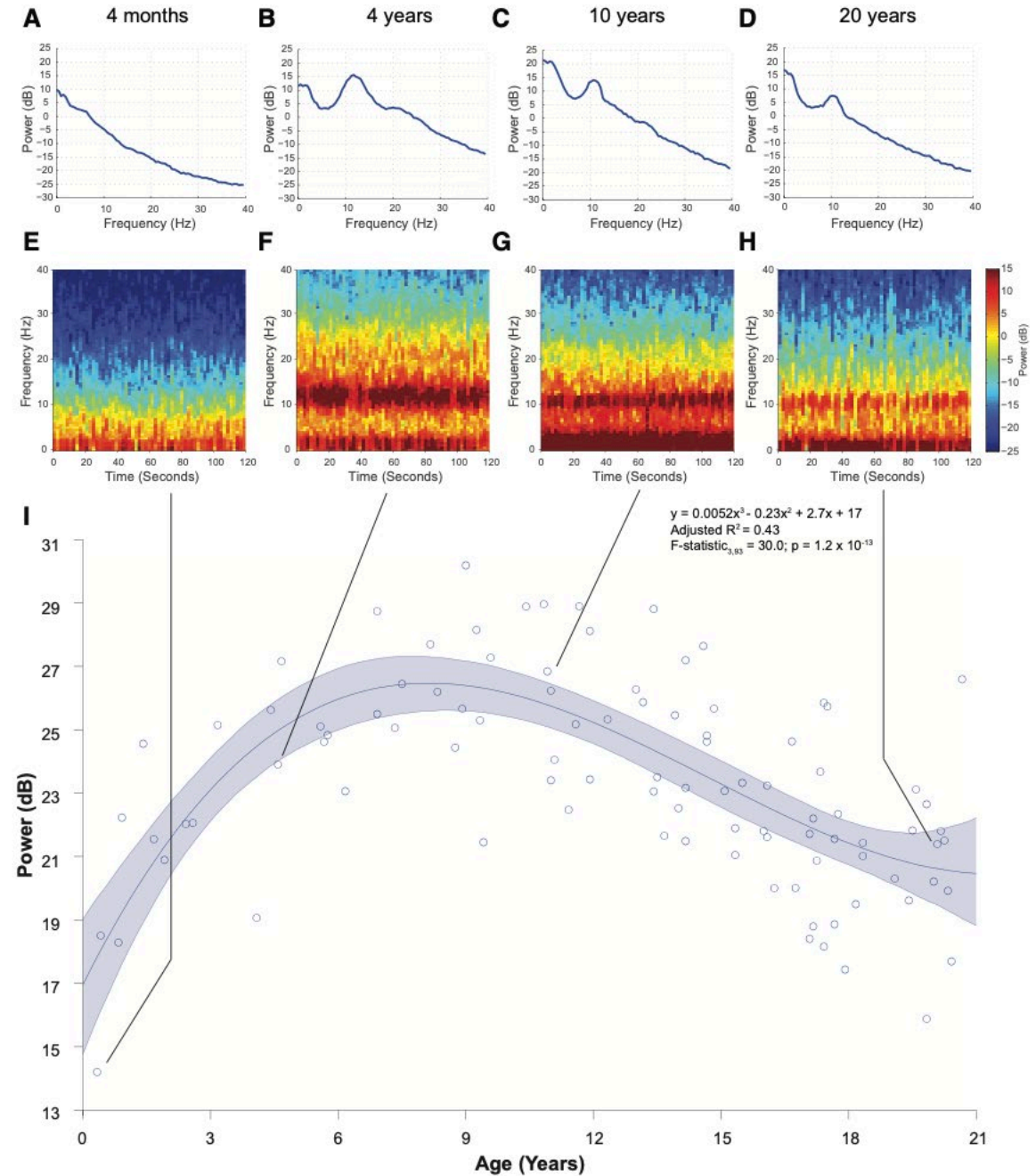


A Prospective Study of Age-dependent Changes in Propofol-induced Electroencephalogram Oscillations in Children

Johanna M. Lee, A.B., Oluwaseun Akeju, M.D., M.M.Sc., Kristina Terzakis, Kara J. Pavone, B.S., Hao Deng, M.B.B.S., M.P.H., Timothy T. Houle, Ph.D., Paul G. Firth, M.B., Ch.B., Erik S. Shank, M.D., Emery N. Brown, M.D., Ph.D., Patrick L. Purdon, Ph.D.

Anesthesiology 2017

- Total electroencephalogram power (0.1 to 40 Hz) peaked at approximately 8 yr old and subsequently declined with increasing age.
- For patients greater than 1 yr old, the propofol-induced electroencephalogram structure was qualitatively similar regardless of age, featuring slow and coherent alpha oscillations.
- For patients under 1 yr of age, frontal alpha oscillations were not coherent.



Age-dependency of sevoflurane-induced electroencephalogram dynamics in children

O. Akeju^{1,3,*}, K. J. Pavone^{1,†}, J. A. Thum^{3,5,†}, P. G. Firth^{1,2}, M. B. Westover^{2,3}, M. Puglia^{1,3}, E. S. Shank^{1,3}, E. N. Brown^{1,3,4,5,6}, and P. L. Purdon^{1,3,4,*}

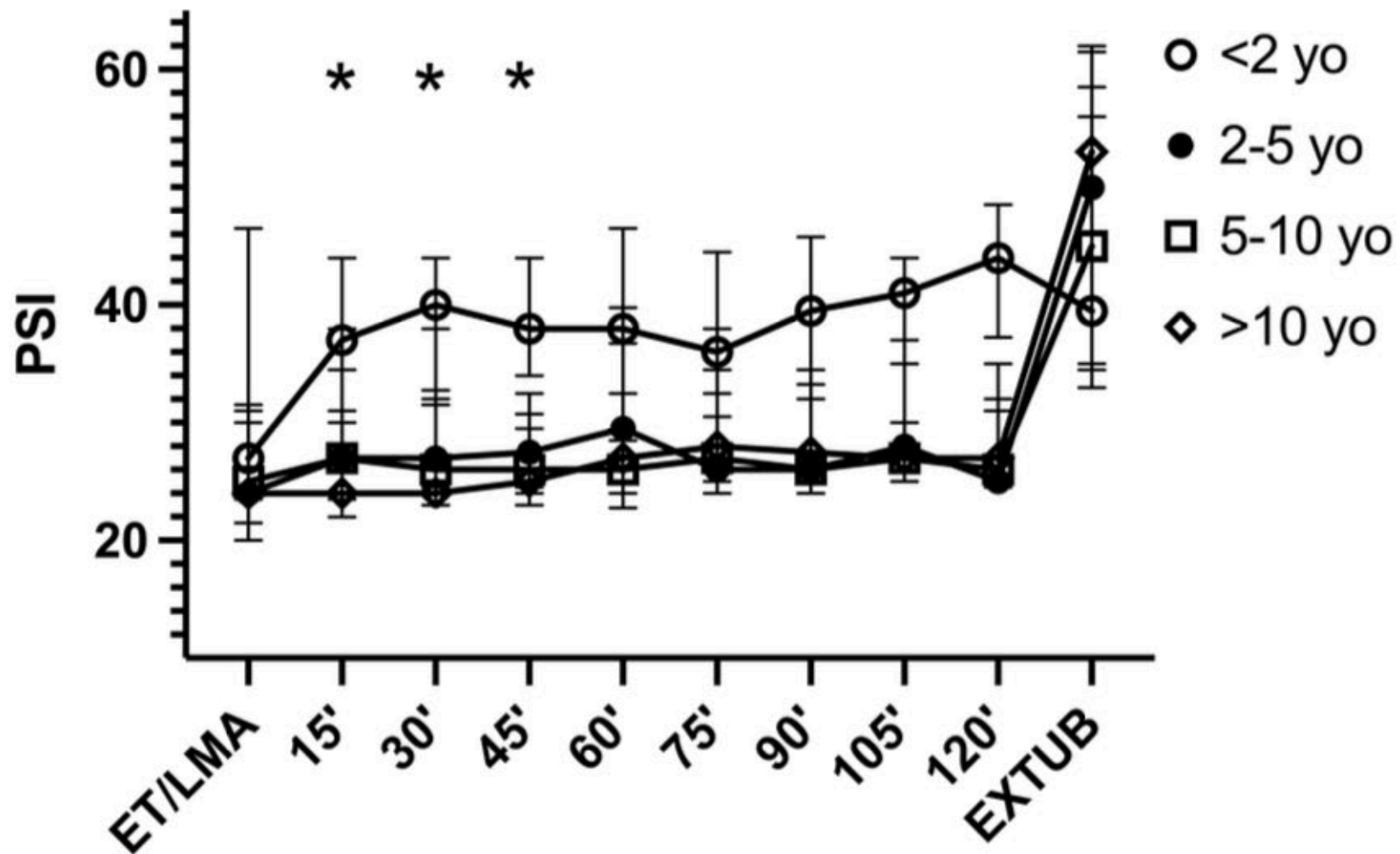
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- ✓ These higher-frequency bands is elevated in children greater than 1 yr of age at surgical concentrations of anaesthesia when compared with adults.
- ✓ This important difference could cause index-based depth of anaesthesia monitors to compute a falsely elevated index value in children. In such instances, these falsely elevated index values, could lead to increased anaesthetic drug dosing beyond what is required.

Monitoring anesthesia depth with patient state index during pediatric surgery

Zaccaria Ricci^{1,2} | Chiara Robino² | Paolo Rufini² | Silvia Cumbo² | Sara Cavallini² | Lorenzo Gobbi¹ | Agata Brocchi¹ | Paola Serio² | Stefano Romagnoli^{1,3}



CLINICAL INVESTIGATION

Electroencephalographic delta and alpha oscillations and phase-amplitude coupling in paediatric patients undergoing propofol-based general anaesthesia

Luai Zakaria^{1,2,3,4}, Adela Desowska^{2,3}, Charles...

¹Department of Anesthesiology, Perioperative and Pain Medicine, Harvard Medical School, Boston, MA, USA and ³Massachusetts General Hospital, Boston, MA, USA

The absence of phase-amplitude coupling cannot exist in children under the age of 10. A clinical state of unconsciousness is still observed in these patients. Together with our findings, this emphasises the complex nature of how anaesthetics produce a state of unconsciousness.

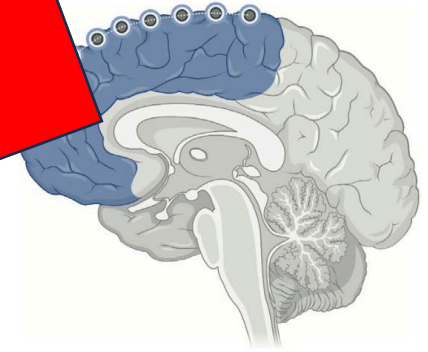
Although a single unifying mechanism is unlikely, the underlying mechanisms could involve decreased integration of neuronal signals, decreased functional connectivity, a generalised neuronal disruption of rhythmic oscillations, or all three

Phase amplitude coupling

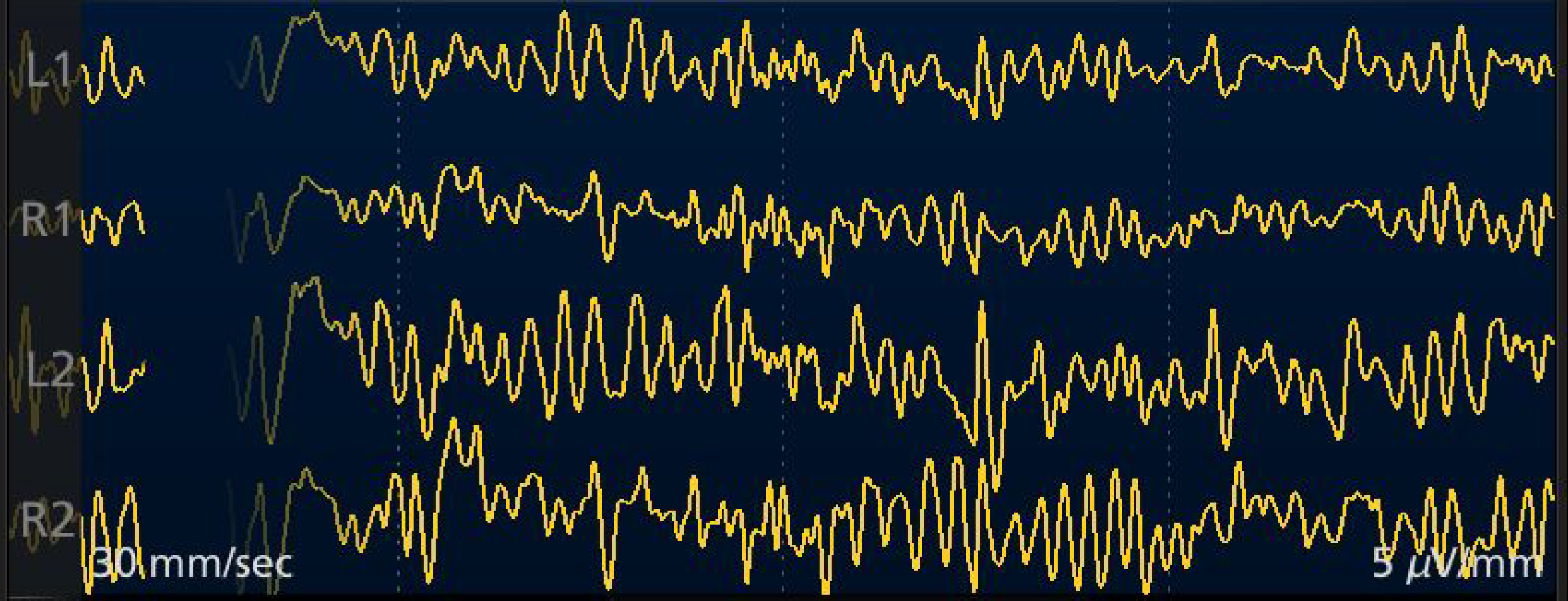
Higher frequency

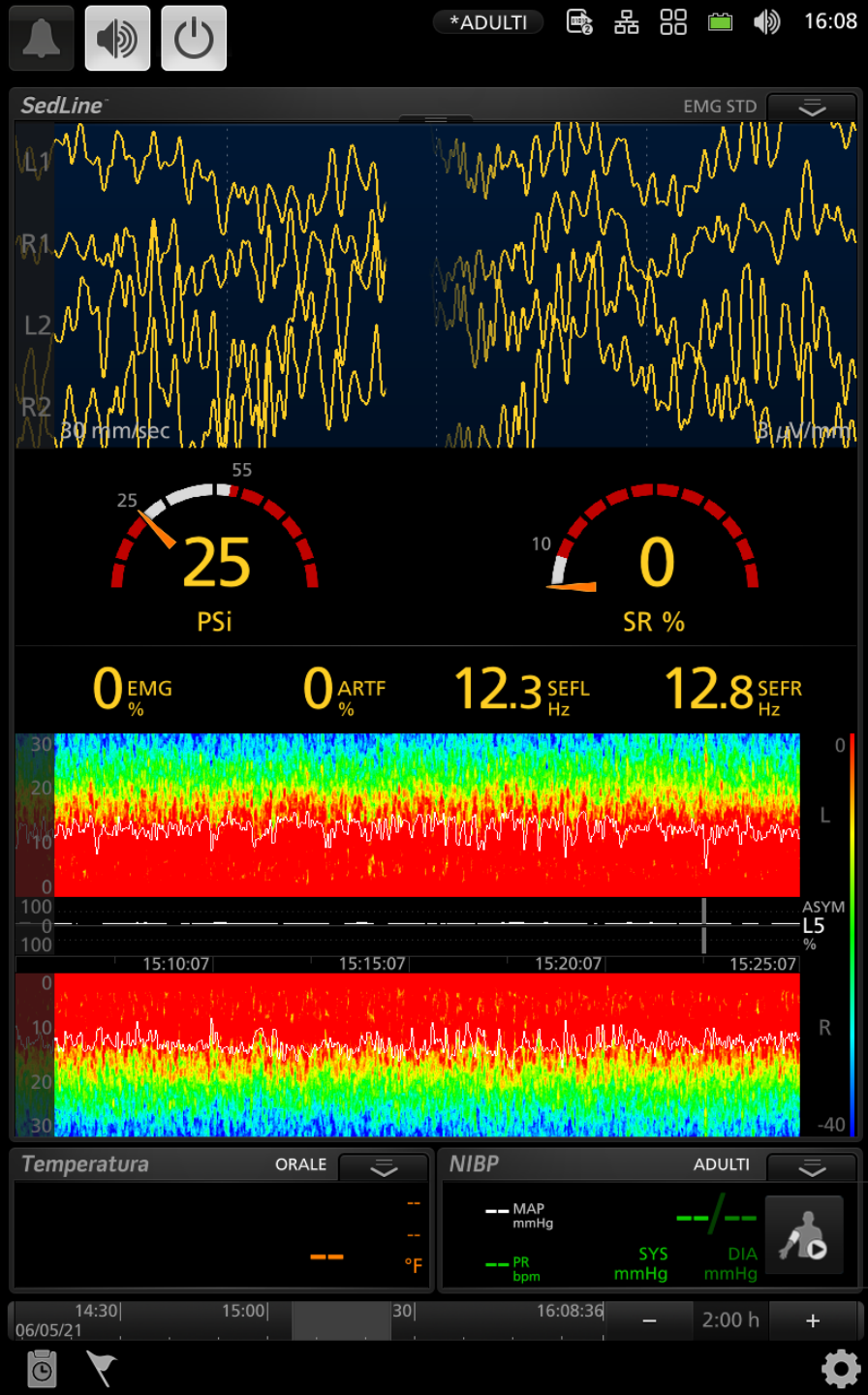
Down-state

Frontal EEG



FRAGILE

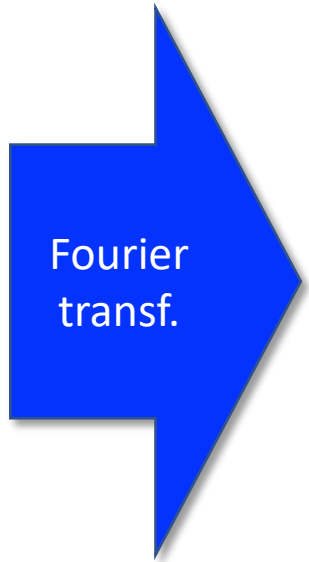




These anesthesia-induced oscillations are readily visible in the electroencephalogram

What about spectrogram?

Unprocessed EEG Waveform



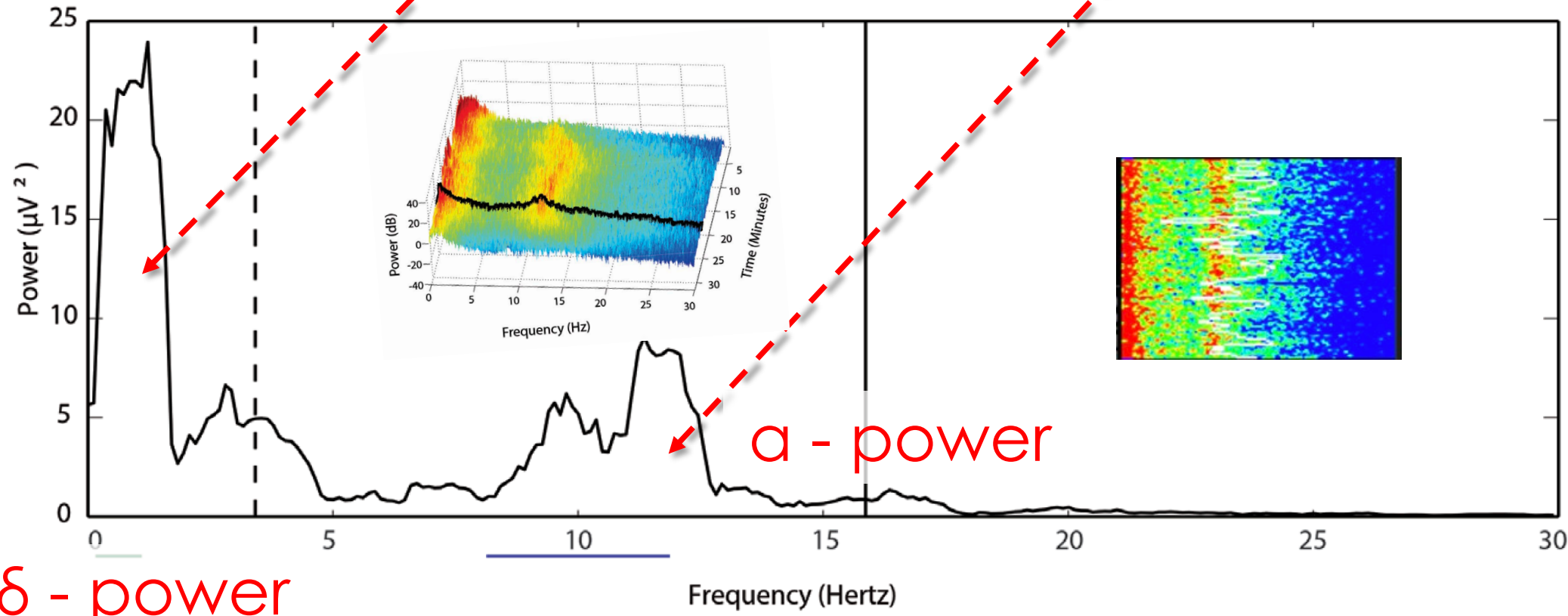
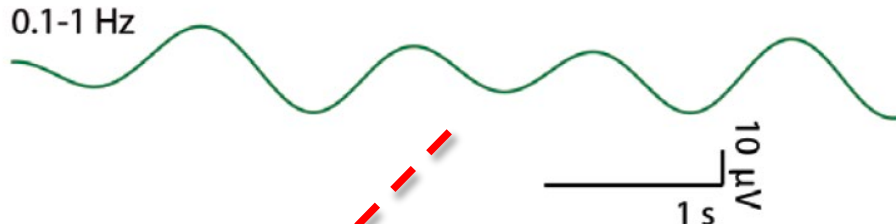
8-12 Hz

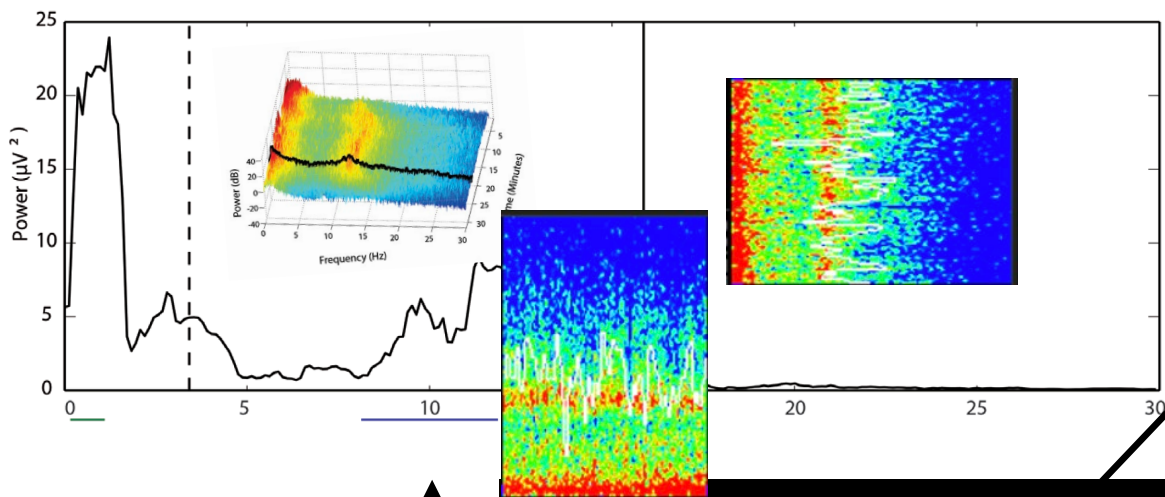


0.1-1 Hz



Spectral analysis → spectrum





Alpha

Delta

Beta-gamma

Burst-suppression

Spectrum (3D)
 →
 Density Spectral Array (2D)

↑
 Hz

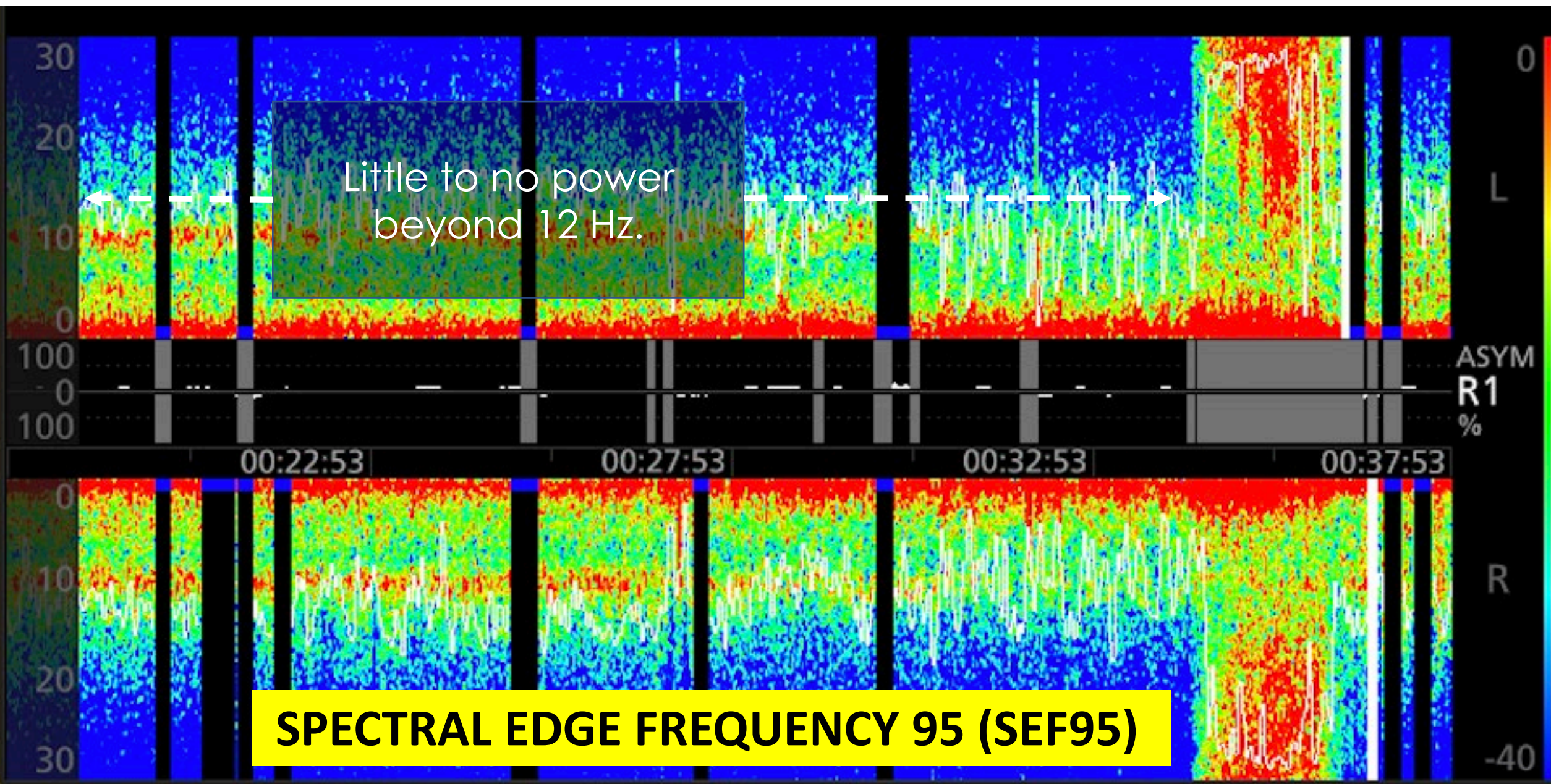


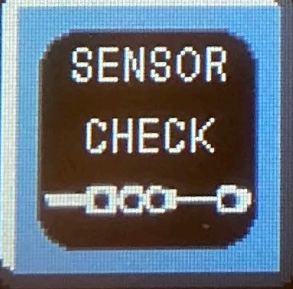
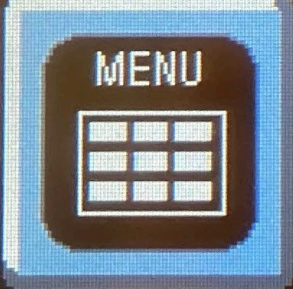
Left

Power (dB)
 $(10 \log_{10} (\text{amplitude})^2)$

Right

TIME →





SEF 12 MF 03

18 Mar 2024 17:16:11

IO9J

BIS 35

H

30 Hz

13 Hz

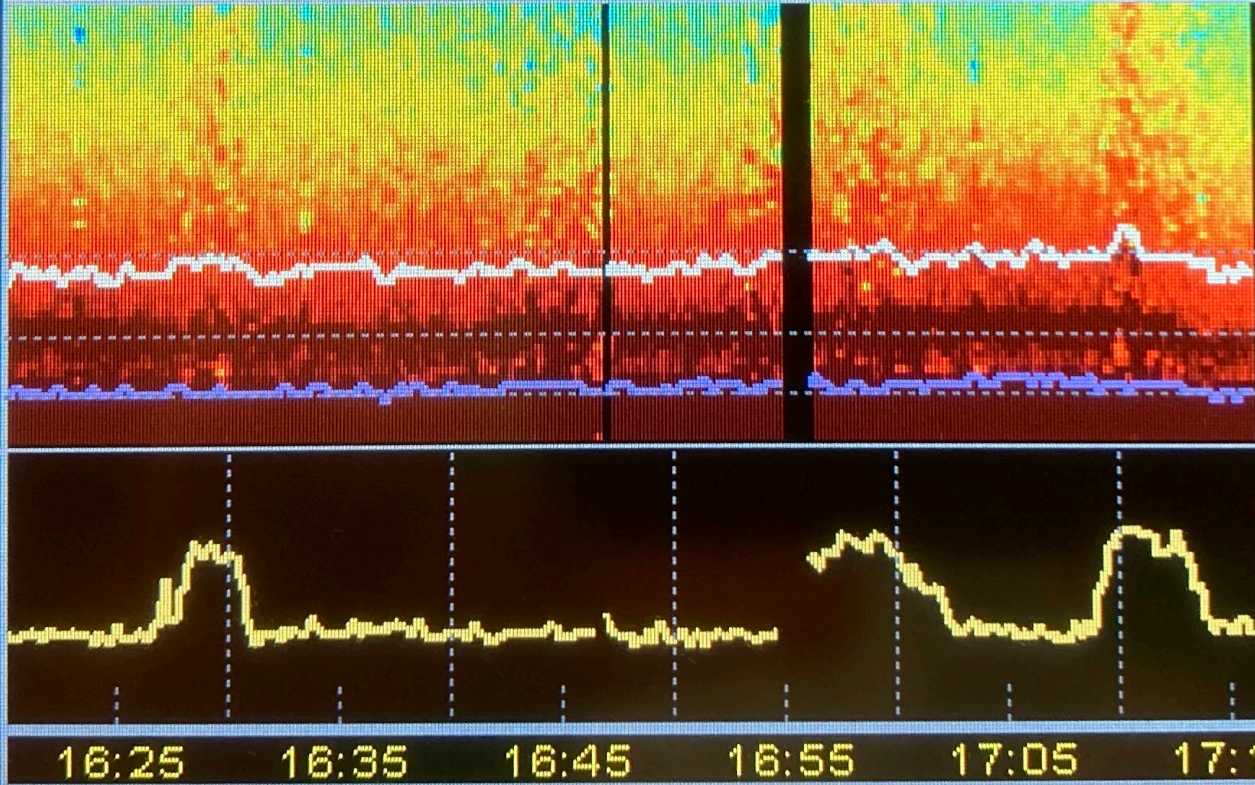
8 Hz

4 Hz

100

50

0



Beta

Alpha

Theta

Delta



16:25

16:35

16:45

16:55

17:05

17:15

EXPERTS' OPINION

Electroencephalographic Density Spectral Array monitoring in pediatric anesthesia: clinical background and practical applications

Iris J. de HEER, Frank WEBER *

Sevoflurane maintenance

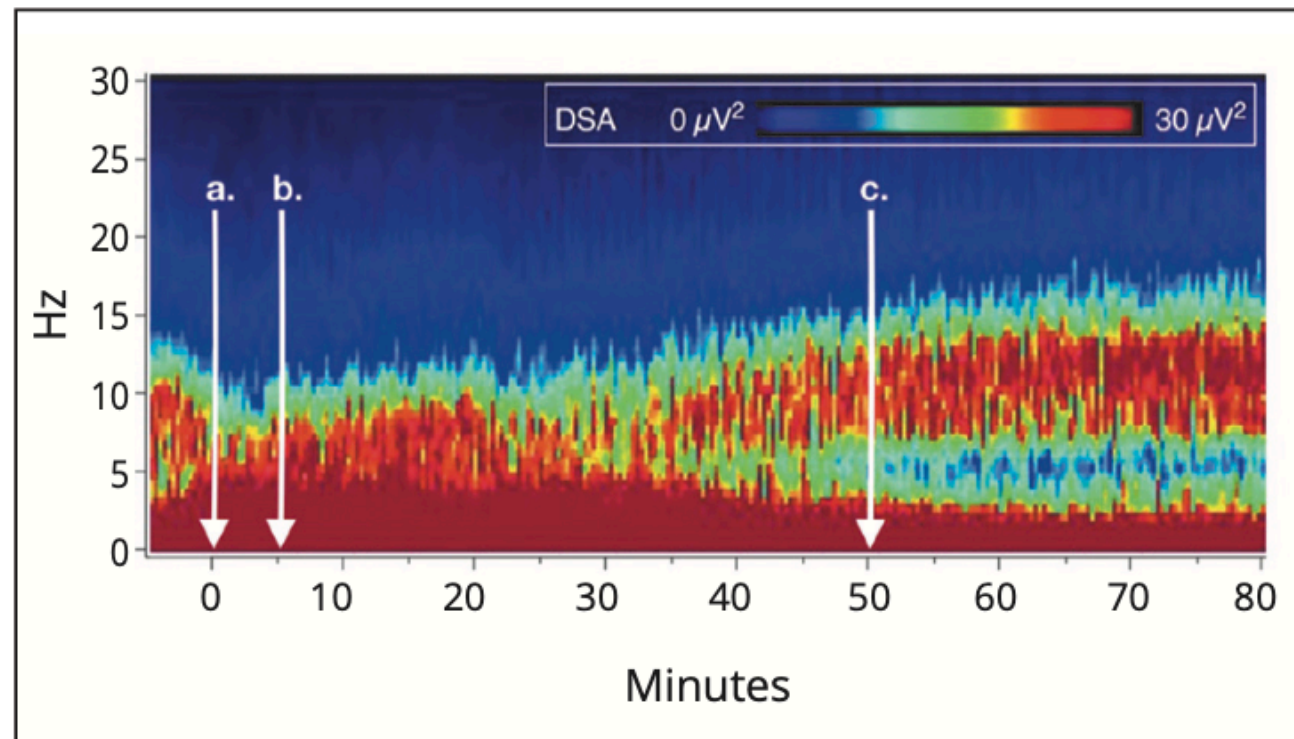
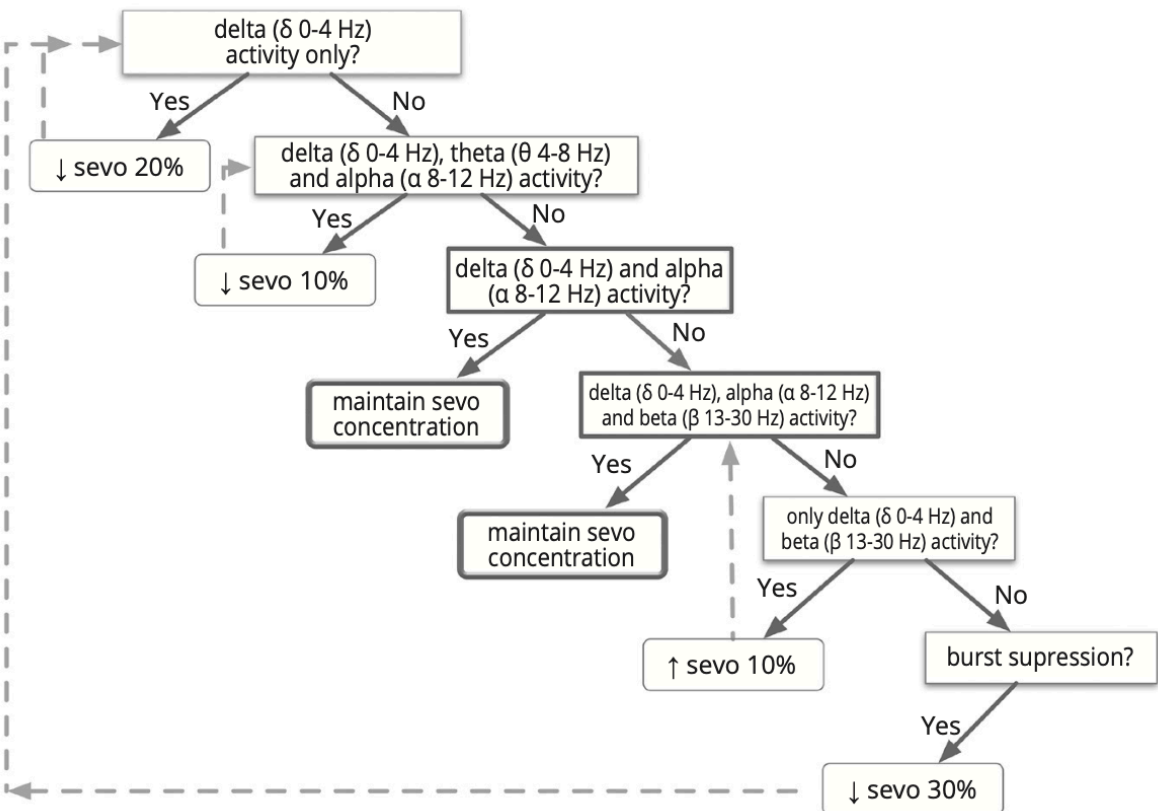


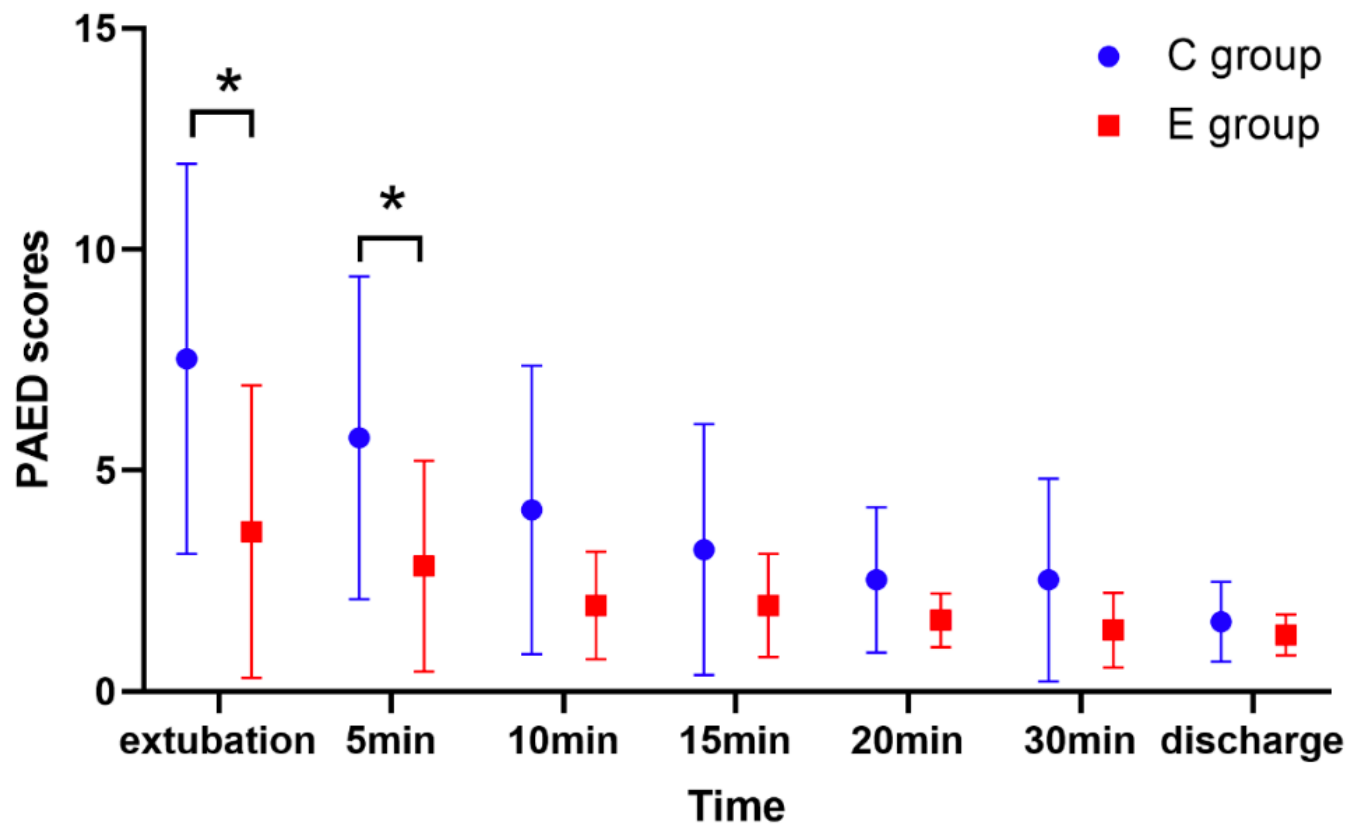
Figure 2.—Example of a DSA pattern during sevoflurane anesthesia in 9 months old infant.

a. IV induction with propofol followed by the onset of delta, theta and alpha oscillations; b. start sevoflurane maintenance, initial a Etsevo concentration of 3.0% was used followed by an increase in power of the theta oscillations and lower frequency alpha oscillations; c. EtSevo concentration of 2.0% was maintained followed by the presence of delta, alpha and beta oscillations.

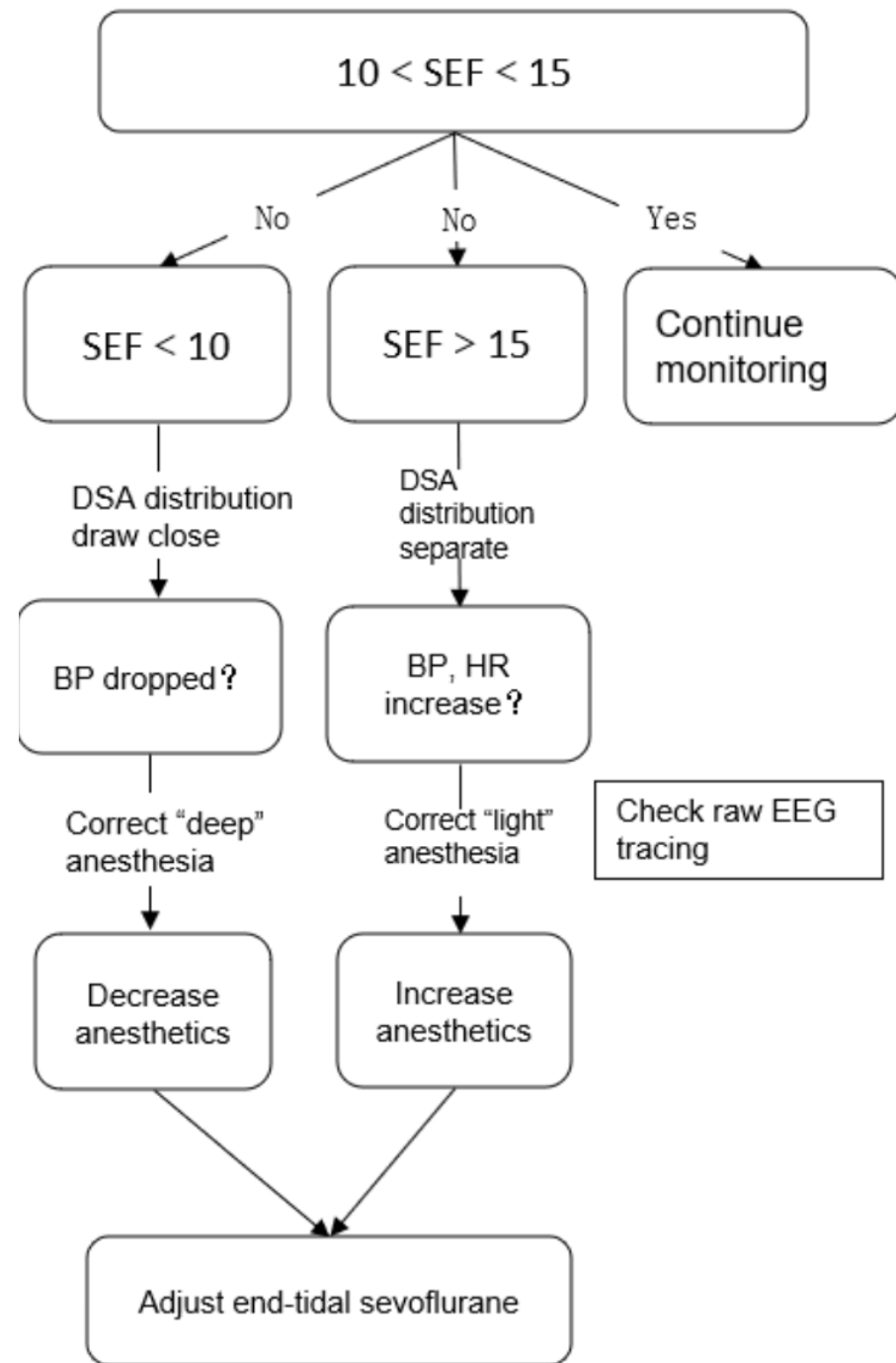
Article

EEG-Parameter-Guided Anesthesia for Prevention of Emergence Delirium in Children

Yaqian Han ^{1,2,†}, Mengrong Miao ^{1,2,†}, Pule Li ³, Yitian Yang ^{1,2}, Hui Zhang ^{1,2}, Beibei Zhang ^{1,2}, Mingyang Sun ^{2,*} and Jiaqiang Zhang ^{1,*}

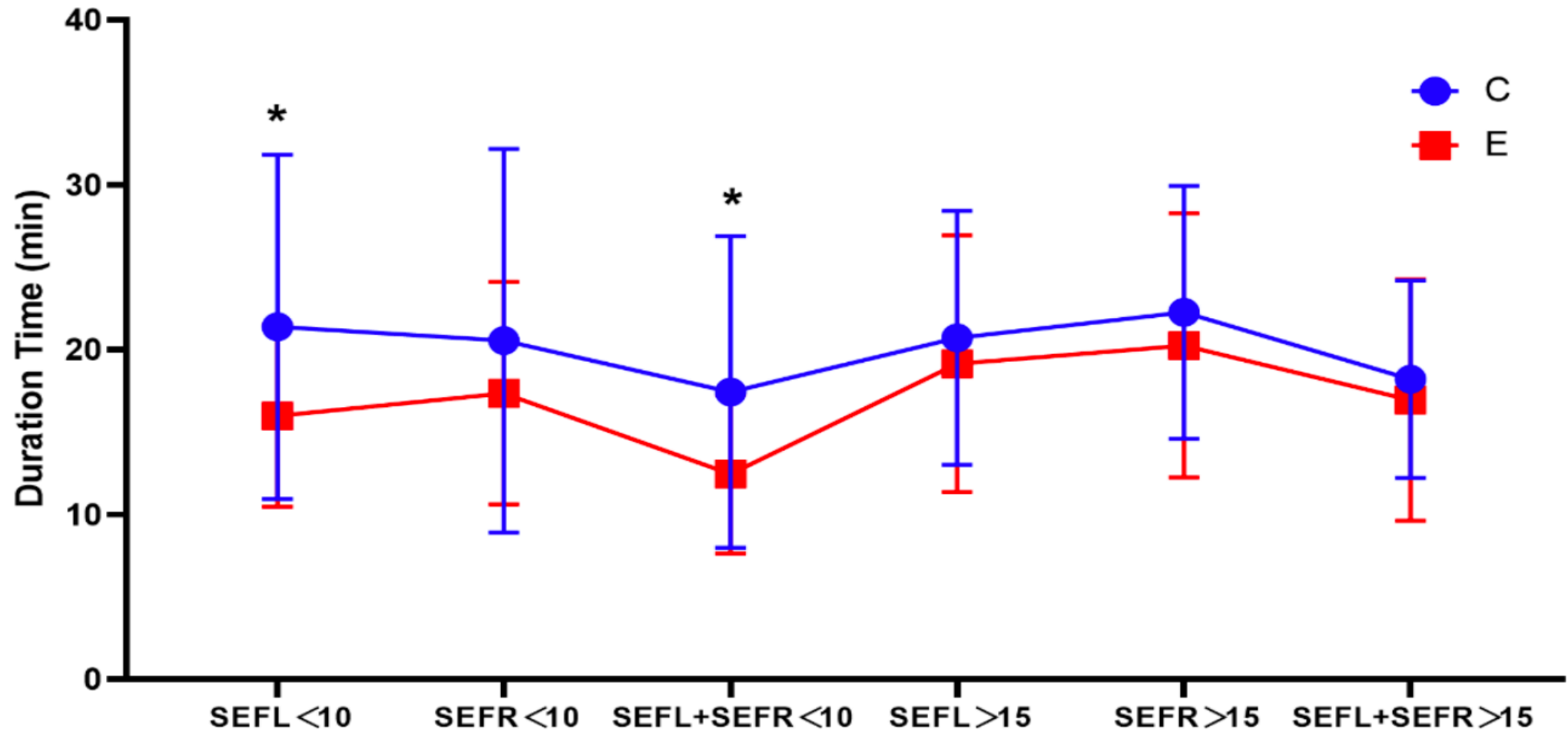


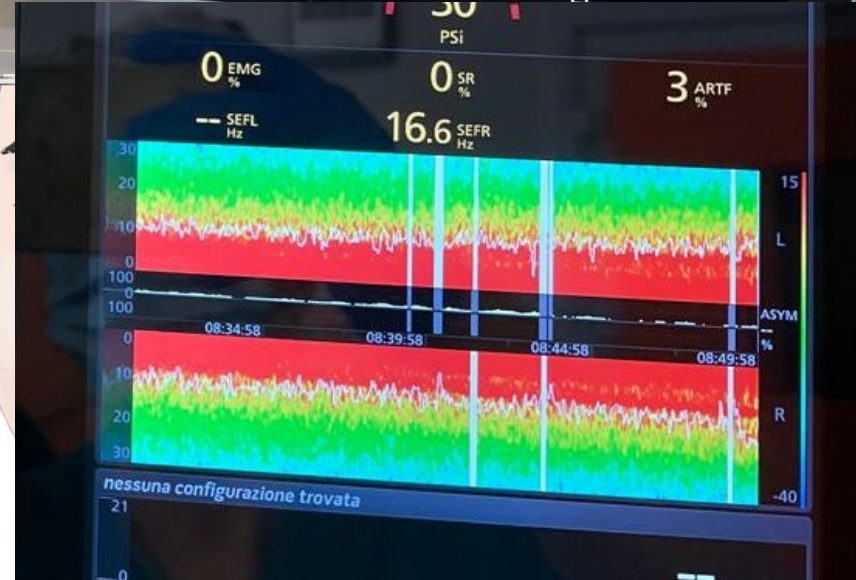
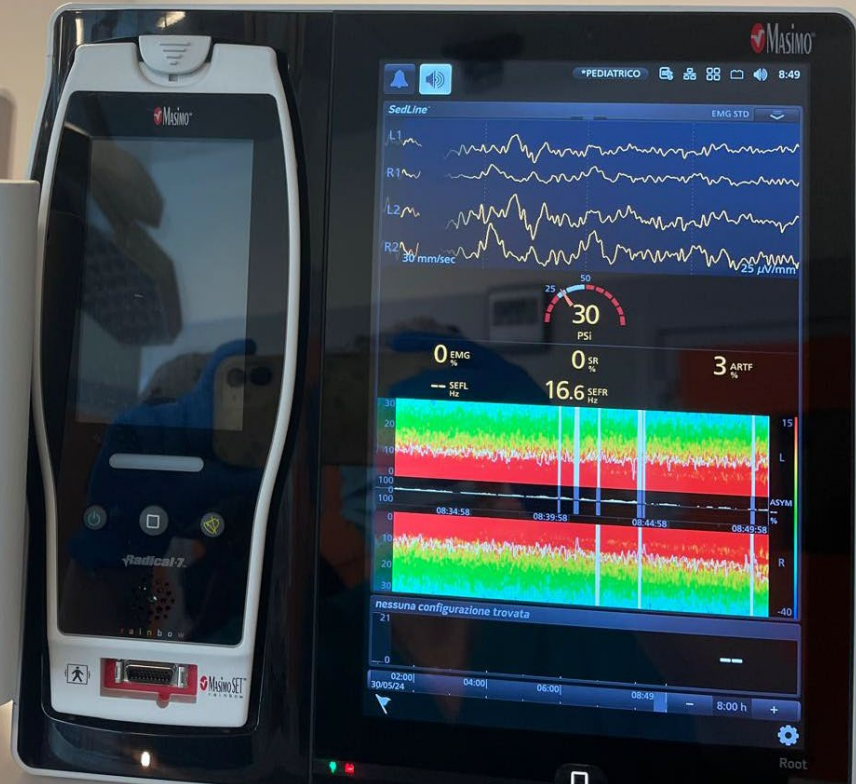
Data are expressed as means \pm SDs, medians (interquartile range). ASA, American Society of Anesthesiologists.



Article
EEG-Parameter-Guided Anesthesia for Prevention of Emergence Delirium in Children

Yaqian Han ^{1,2,†}, Mengrong Miao ^{1,2,†}, Pule Li ³, Yitian Yang ^{1,2}, Hui Zhang ^{1,2}, Beibei Zhang ^{1,2}, Mingyang Sun ^{2,*} and Jiaqiang Zhang ^{1,*}



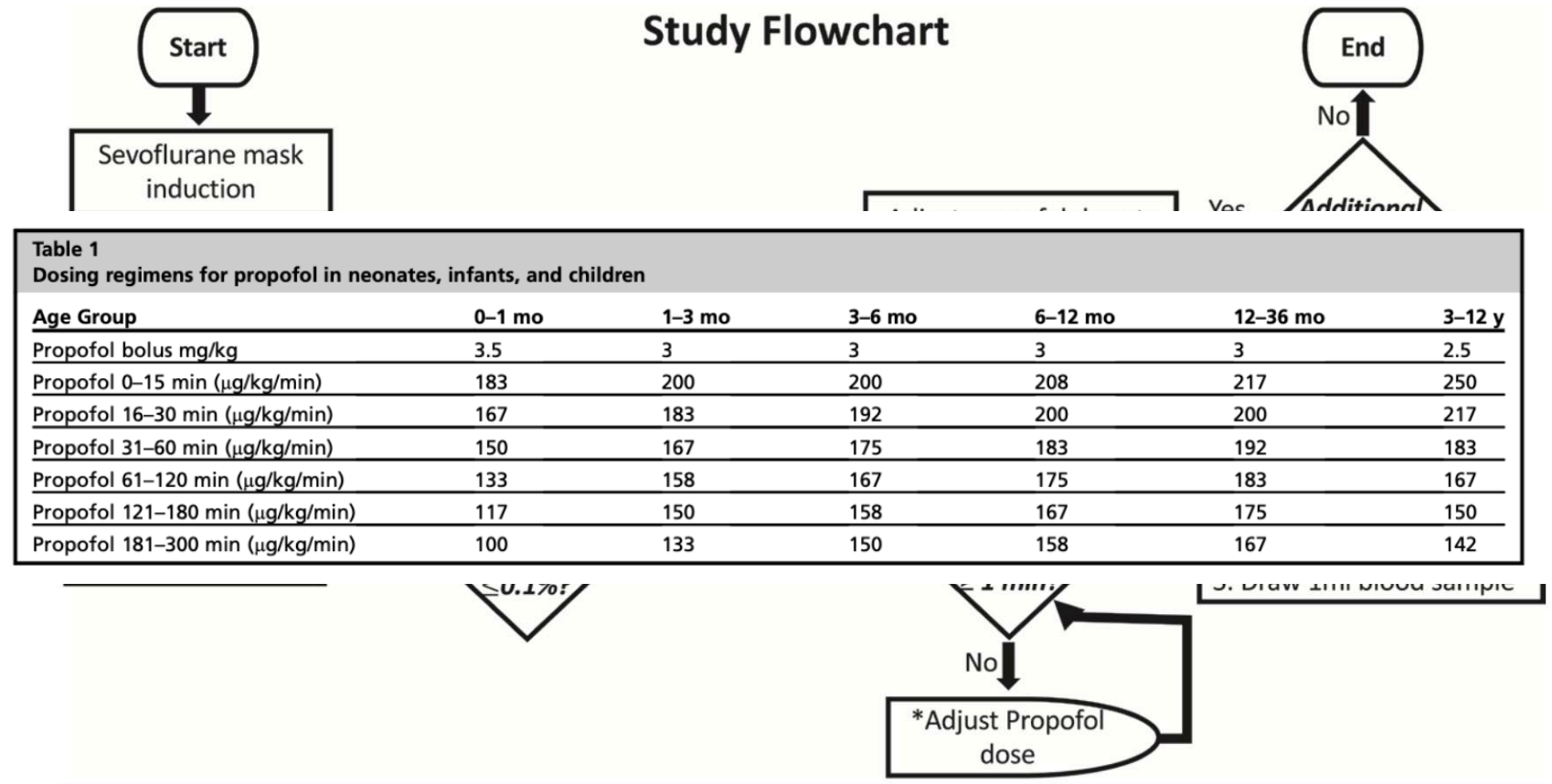


ANESTHESIOLOGY

Electroencephalographic Indices for Clinical Endpoints during Propofol Anesthesia in Infants: An Early-phase Propofol Biomarker-finding Study

Ian Yuan, M.D., M.Eng.,
Annery G. Garcia-Marcinkiewicz, M.D., M.S.C.E.,
Bingqing Zhang, M.P.H., Allison M. Ulrich, M.D.,
Georgia Georgostathi, B.S.E., B.A.,
Richard M. Missett, D.O., Shih-Shan Lang, M.D.,
James L. Bruton, M.S., C. Dean Kurth, M.D.

ANESTHESIOLOGY 2024; 141:353–64

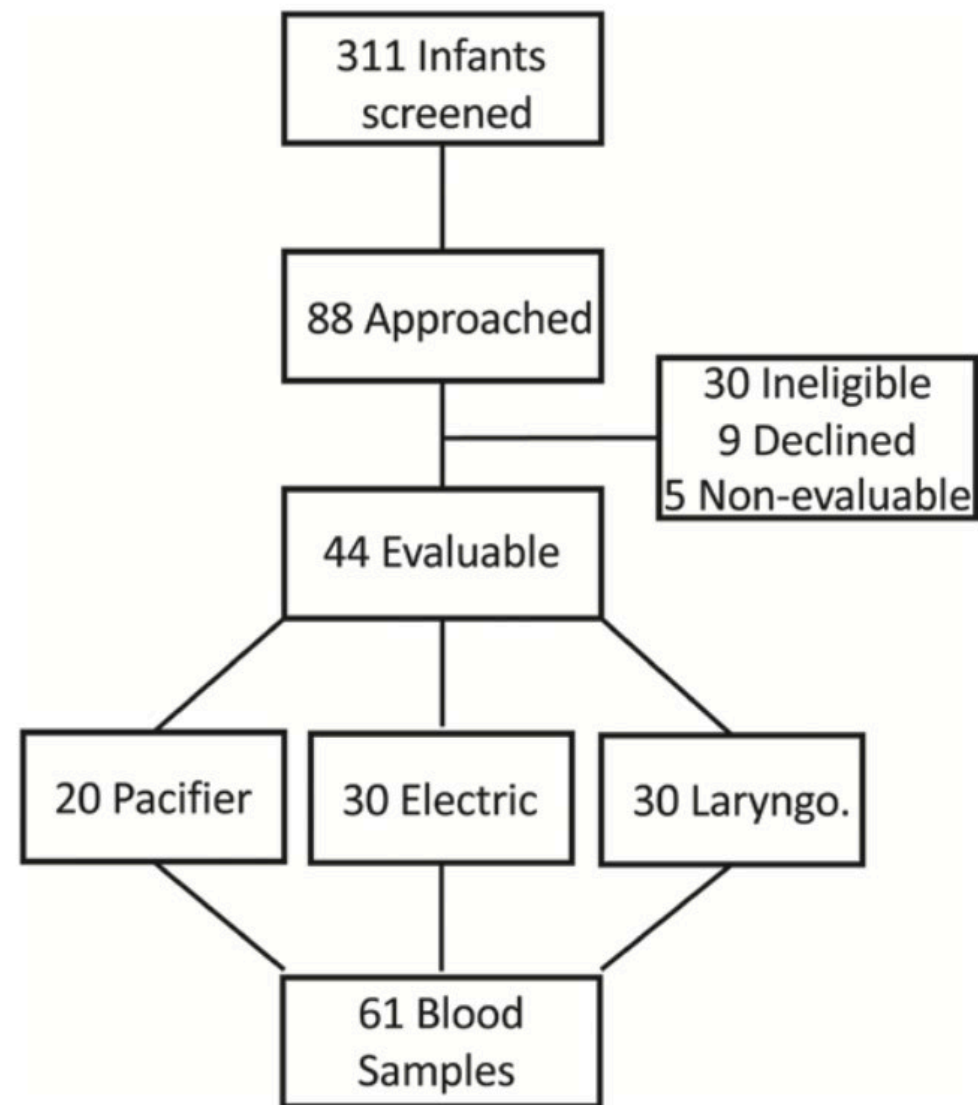


- The SEF95 for the first infant was targeted at 20, 14, and 10 Hz for oral pacifier, electrical stimulation, and laryngoscopy, respectively.
- If the first infant had a positive response the next infant will have SEF95 target decreased by 2 Hz for the same stimulus.
- The SEF95 target was achieved by adjusting propofol dose in each infant using a dosing table as a starting guide and confirmed by steady-state SEF95.

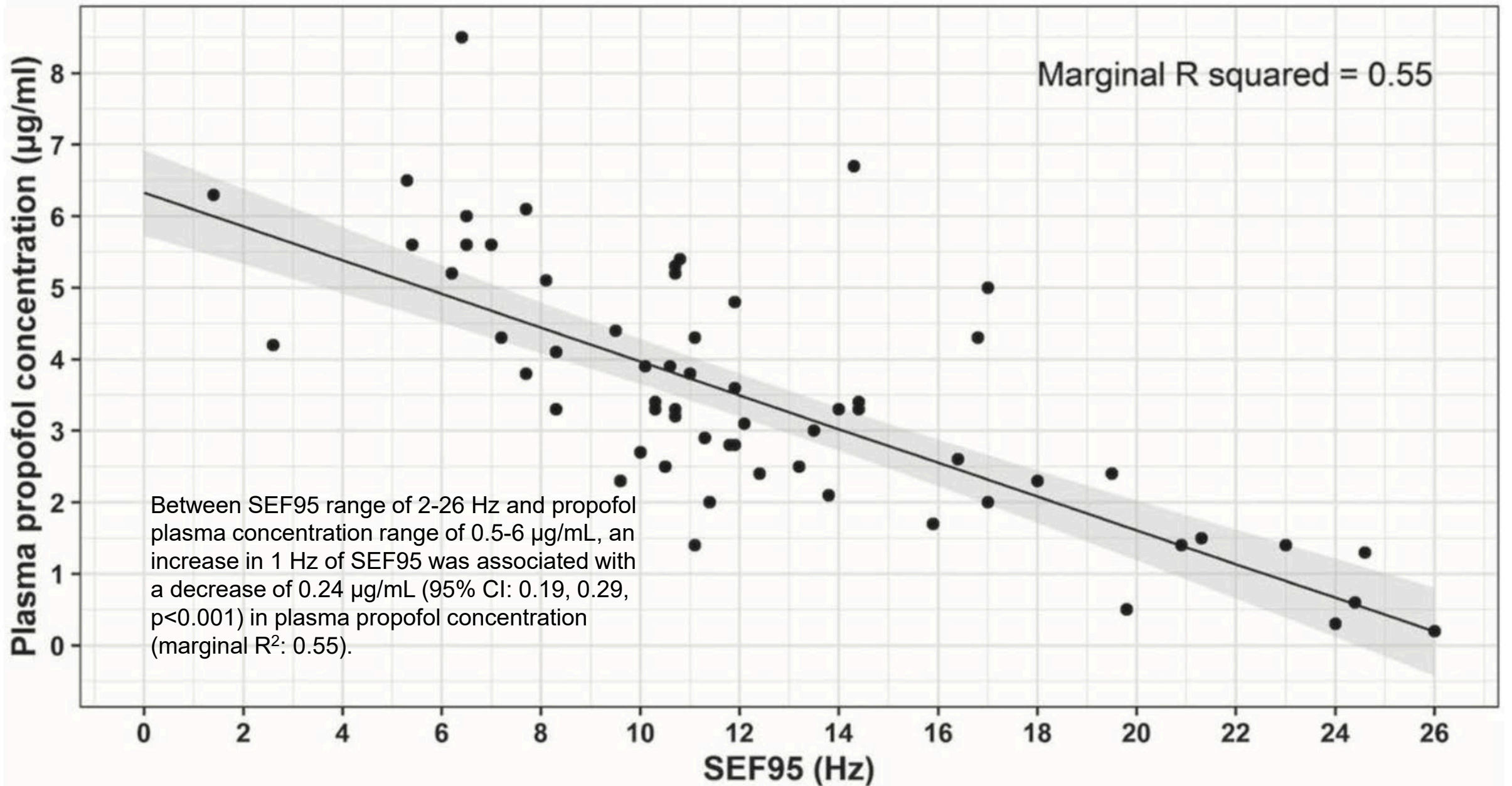
ANESTHESIOLOGY

Electroencephalographic Indices for Clinical Endpoints during Propofol Anesthesia in Infants: An Early-phase Propofol Biomarker-finding Study

Inclusion criteria: ASA 1 or 2, muscle relaxant not planned for laryngoscopy/intubation, and anticipated surgery duration < 2h 40min. This time limit was required by the IRB, citing an FDA warning of anesthetic neurotoxicity in children under 3 years for longer than 3h of anesthesia exposure.



Exclusion criteria: infants undergoing emergency surgery, seizure disorder treated with anticonvulsant, congenital or acquired brain malformation, deformities of forehead, known or potential difficult airway, and allergy to propofol



QUESTIONS

- No fentanyl?
- No roc?
- 3->12 months?
- Burst suppression?
- Sevoflurane?
- Locoregional?
- Electrical stimuli?
- LMA?

«The original dose-finding studies to determine median alveolar concentration used volatile anesthetic as the sole agent, as this was necessary to provide foundational knowledge.

Similarly, our study provides foundational knowledge of propofol for key clinical anesthesia endpoints, paving the way for future studies to determine responses to a combination of propofol with remifentanyl or other drugs often during clinical care»

CONCLUSIONS

1. Neuromonitoring should become standard of care in pediatric anesthesia, at least in patients >2 yo
2. Developing ages still have many unanswered questions
3. pEEG guided anesthesia should target the lowest anesthetic dosages for the optimal anesthesia depth (threshold?)
4. Avoidance of BS is associated with lower hemodynamic instability and potentially less postoperative neurologic impairment
5. Raw traces, spectrograms and SEF are useful for objective pEEG interpretation in different ages and with different technology
6. **A lot of research is due in this field and innovation is warranted (monitoring of power, relative power, nociception, delirium)**