Department of Neurological Surgery

# Clinical Applications of Navigated TMS in Acute Neurological Injury

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## Induction



# **Transcranial Induction**

#### epartment of Neurological Surgery

## Where it all began...



#### A novel approach for documenting naming errors induced by navigated transcranial magnetic stimulation

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Anomia Paraphasias

#### ABSTRACT

Transcranial magnetic stimulation (TMS) is widely used both in basic research and in clinical practice. TMS has been utilized in studies of functional organization of speech in healthy volunteers. Navigated TMS (nTMS) allows preoperative mapping of the motor cortex for surgical planning. Recording behavioral responses to nTMS in the speech-related cortical network in a manner that allows off-line review of performance might increase utility of nTMS both for scientific and clinical purposes, e.g., for a careful preoperative planning.

Four subjects participated in the study. The subjects named pictures of objects presented every 2.5 s on a computer screen. One-second trains of 5 pulses were applied by nTMS 300 ms after the presentation of pictures. The nTMS and stimulus presentation screens were cloned. A commercial digital camera was utilized to record the subject's performance and the screen clones. Delays between presentation, audio and video signals were eliminated by carefully tested combination of displays and camera. An experienced neuropsychologist studied the videos and classified the errors evoked by nTMS during the object naming. Complete anomias, semantic, phonological and performance errors were observed during nTMS of left

fronto-parieto-temporal cortical regions. Several errors were detected only in the video classification. nTMS combined with synchronized video recording provides an accurate monitoring tool of behavioral TMS experiments. This experimental setup can be particularly useful for high-quality cognitive paradigms and for clinical purposes.

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## nTMS Language Mapping

#### RESEARCH—HUMAN—CLINICAL STUDIES

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‡Department of Neurosurgery, Chaitié Universitätsmedizin, Berlin, Germany; SDepartment of Neurosurgery, Kinikum rechts der Isar, Technische Universität München, Germany; @Neostim Ltd., Hekniki, Finland and Neuroscience Unit, Institute of Biomedicine/Physiology, University of Heksinki, Finland; ||BioMag Laboratory, HUS Medical Imaging, Hospital Oktrict of Heksinki and Uusimaa, Helsinki, Finland; #Department of Intraopentice Neurophysiology, St. Luke's-Roosevet Hospital, New York, New York; \*"Laboratory for Human and Experimental Neurophysiology, School of Medidine, University of Split, Split, Croats A Comparison of Language Mapping by Preoperative Navigated Transcranial Magnetic Stimulation and Direct Cortical Stimulation During Awake Surgery

**BACKGROUND:** Navigated transcranial magnetic stimulation (nTMS) is increasingly used in presurgical brain mapping. Preoperative nTMS results correlate well with direct cortical stimulation (DCS) data in the identification of the primary motor cortex. Repetitive nTMS can also be used for mapping of speech-sensitive cortical areas.

**OBJECTIVE:** The current cohort study compares the safety and effectiveness of preoperative nTMS with DCS mapping during awake surgery for the identification of language areas in patients with left-sided cerebral lesions.

**METHODS:** Twenty patients with tumors in or close to left-sided language eloquent regions were examined by repetitive nTMS before surgery. During awake surgery, language-eloquent cortex was identified by DCS. nTMS results were compared for accuracy and reliability with regard to DCS by projecting both results into the cortical parcellation system.

**RESULTS:** Presurgical nTMS maps showed an overall sensitivity of 90.2%, specificity of 23.8%, positive predictive value of 35.6%, and negative predictive value of 83.9% compared with DCS. For the anatomic Broca's area, the corresponding values were a sensitivity of 100%, specificity of 13.0%, positive predictive value of 56.5%, and negative predictive value of 100%, respectively.

**CONCLUSION:** Good overall correlation between repetitive nTMS and DCS was observed, particularly with regard to negatively mapped regions. Noninvasive inhibition mapping with nTMS is evolving as a valuable tool for preoperative mapping of language areas. Yet its low specificity in posterior language areas in the current study necessitates further research to refine the methodology.

KEY WORDS: Awake surgery, Language, Preoperative mapping, Transcranial magnetic stimulation, Tumor





## and any anguage Mapping

**RESEARCH—HUMAN—CLINICAL STUDIES** 



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## Language mapping with navigated repetitive TMS: Proof of technique and validation



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# Motor Mapping with TMS

Reference	Number of patients	Mean distance nTMS to DCS	Mean distance fMRI/MEG to DCS
Krings, 1997	2	<10 mm	-
Picht, 2009	10	3 mm	-
Kantelhardt , 2010	1	<5 mm	-
Picht, 2011	23	8 mm	14 mm (fMRI)
Forster, 2011	10	13 mm	21 mm (fMRI)
Krieg, 2012	14	4 mm	10 mm (fMRI)
Paiva, 2012	6	4 mm	-
Tarapore, 2012	8	2 mm	5 mm (MEG)
Coburger, 2012	30	-	23% fMRI not feasible
Krieg, 2013	24	5 mm	9 mm (fMRI)
Vitikainen, 2013	13	11 mm	-
Mangraviti, 2013	8	8 mm	13 mm (fMRI)
Frey, 2014	82	6 mm	-

Takahashi et al, Neurosurg. Focus, 2013



## **Facial Recognition**

## Non-invasive Mapping of Face Processing by Navigated Transcranial Magnetic Stimulation

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**REVIEW ARTICLE - NEUROSURGICAL TECHNIQUES** 

### Protocol for motor and language mapping by navigated TMS in patients and healthy volunteers; workshop report

nTMS Mapping Consensus

Sandro M. Krieg<sup>1</sup> · Pantelis Lioumis<sup>2</sup> · Jyrki P. Mäkelä<sup>2</sup> · Juha Wilenius<sup>3</sup> · Jari Karhu<sup>4</sup> · Henri Hannula<sup>4,5</sup> · Petri Savolainen<sup>4</sup> · Carolin Weiss Lucas<sup>6</sup> · Kathleen Seidel<sup>7</sup> · Aki Laakso<sup>8</sup> · Mominul Islam<sup>9</sup> · Selja Vaalto<sup>3</sup> · Henri Lehtinen<sup>10</sup> · Anne-Mari Vitikainen<sup>2</sup> · Phiroz E. Tarapore<sup>11</sup> · Thomas Picht<sup>12</sup>

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#### Abstract

*Introduction* Navigated transcranial magnetic stimulation (nTMS) is increasingly used for preoperative mapping of motor function, and clinical evidence for its benefit for brain tumor patients is accumulating. In respect to language mapping with repetitive nTMS, literature

reports have yielded variable results, and it is currently not routinely performed for presurgical language localization. The aim of this project is to define a common protocol for nTMS motor and language mapping to standardize its neurosurgical application and increase its clinical value.

The navigated TMS workshop group; Helsinki Meeting 2016

All authors contributed equally to this work

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## **New Frontiers...**





### nTMS – An Adjunct Neuro Exam

- Quantitative (MEPs)
- Repeatable
- Reproducible
- Highly sensitive and specific
- Does not require patient participation

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### **Acute Traumatic Brain Injury**





#### **Post-operative Day 1**

- RMT APB- Left 45% / Right 47%
- Latency 21-24ms





#### **Post-operative Day 4**

### Intermittently elevated ICP (20-26)



- Decrease in amplitude
- Monophasic morphology
- Ionger (and asymmetric) latency



### **Post-operative Day 4 (Late)**

		1.1.15.		
			-	
1.abp right				υμν
	-4µV 3min5s247.52ms		-2μV -25.00ms	0 0ms
2.EDC				υμν
				0 0ms
			-	
			-	85uV
3.adm right				
				28.0ms
			-	



#### **Post-operative Day 5**

### Persistently elevated ICP (25-35)





#### **Lessons** Learned

- nTMS can be used in an acute inpatient setting, in a critically ill patient
- MEPs may change with the neurophysiology
  - Amplitude
  - Morphology
  - Latency
    - SO...
- It's possible!





## Acute Cervical Spinal Cord Injury



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### **Post-operative Day 1**

#### Nexstim Navigation Stimulation Digitization Overlay Settings Graph Settings Registration Stimulation target Perform registration. Add Stimulation target ... Slice scan (mm) 3D peeling Depth (mm) Axial depth (mm) Sagittal Corona Axial 250-250-50-200-200-200 200-40-150-150-30-150-150-100-100-100-50-50-50-50-10-<u>\_</u> : 0-0-1 125 125 100 21.9 100 Navigation control-Display control Head tracker 0 Stimulation coil Crosshair (F1) Digitizing pen (F2) Pan Registration performed Zoom (F3) Tracking unit state Rotate (F4) Enable Tracking unit Original (F5) 1.1.17. Stimulus: 107 s 546 ms, 40% (visible) 1.1.18. Stimulus: 122 s 221 ms, 60% (visible) 1.1.19. Stimulus: 124 s 912 ms, 60% (visible) 1,1,20, Stimulus: 127 s 145 ms, 60% (visible) 1.1.21. Stimulus: 135 s 100 ms, 60% (visible) 1.1.22. Stimulus: 141 s 284 ms, 60% (visible) 1.1.23. Stimulus: 153 s 101 ms, 60% (visible) 1.1.24. Stimulus: 158 s 482 ms, 60% (visible) 1.2. Stimulus Sequence 1.3. Stimulus Sequence 1.4. Stimulus Sequence 1.5. Stimulus Sequence Description: Bilateral ... Coil name: Focal Coil Stimulator name: TMS II - 1.5. Stimuli 1.5.2. Stimulus: 39 s 260 ms, 100% (visible) 1.5.3. Stimulus: 42 s 879 ms, 100% (visible) >

Quit

Close

session

New

session

Open

session



### **Post-operative Days 1-2**



#### Day 2 (delirious)





Crosshair (V/m

Aim: Total (mm)

Depth (mm)

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### **Post-operative Day 3**



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### **Post-operative Day 3**







#### **Lessons** Learned

- MEPs correlate closely with motor examination in acute cervical spinal cord injury
- MEPs can be extremely valuable for following the exam in a patient who is unable to participate
- Therapeutic intervention may be guided successfully by following MEP.

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## **Aneurysmal SAH**





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## **Post-Bleed Day 4**



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## **Post-Bleed Day 7**



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### **A Silver Lining?**



APB

ED

#### ADM



#### Lessons Learned

- MEPs can be used to follow cerebral ischemia in real-time
- The localization of function is specific enough that it can distinguish different vascular territories
- Hypoperfused brain tissue ("area at risk") demonstrates functional changes before it displays



#### Conclusions

- nTMS can be safely utilized in the acute inpatient setting
- Exact reproducibility between studies is critical (software-based point selection)
- nTMS provides neurophysiological data that would otherwise be unobtainable
- Trends in MEPs over time seem to have clinical significance
- nTMS can help guide clinical management in a variety of neurological injury states



#### **Future Directions**

- More patients
- More disease states
- Effects of anesthetics?
- Medium and longterm follow-up studies



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