# Oscillations and effective connectivity in human corticothalamic networks

Mario Rosanova, MD, PhD
Department of clinical sciences «Luigi Sacco»
University of Milan, Milan, Italy

# -Characterize EEG oscillations triggered by TMS in healthy, awake subjects

-Measuring cortical effective connectivity in different models of unconsciousness

# -Characterize EEG oscillations triggered by TMS in healthy, awake subjects

-Measuring cortical effective connectivity in different models of unconsciousness

# -Characterize EEG oscillations triggered by TMS in healthy, awake subjects

-Measuring cortical effective connectivity in different models of unconsciousness

## Bedside assessment of consciousness by evaluating the patient's ability to interact with the external environment



### CRS-R

#### COMA RECOVERY SCALE-REVISED

©2004

Administration and Scoring Guidelines

Joseph T. Giacino, Ph.D. and Kathleen Kalmar, Ph.D.

Center for Head Injuries Edison, New Jersey



Johnson Rehabilitation Institution Affiliated with JFK Medical Center



Updated 11-1-05

#### JFK COMA RECOVERY SCALE - REVISED @2004

Record Form

This form should only be used in association with the "CRS-R ADMINISTRATION AND SCORING GUIDELINES" which provide instructions for standardized administration of the scale.

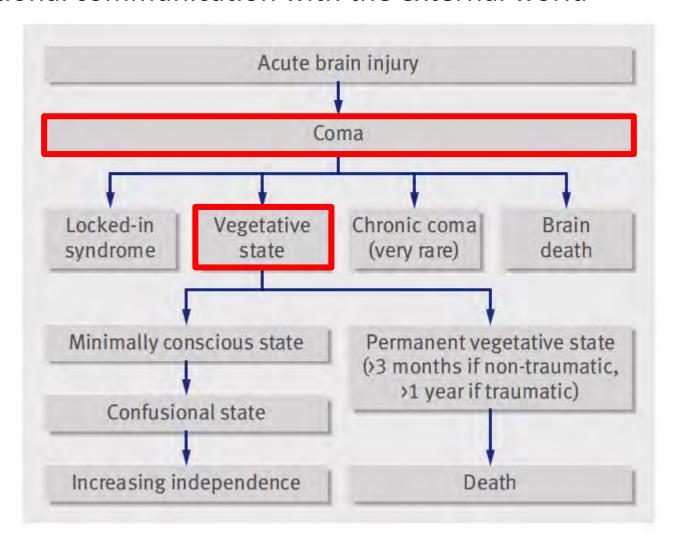
Patient:	Diagnosis:								Etiology:							
Date of Onset:		Date	e of	Adm	issi	on:										
Date																Г
Week	ADM	2	3	4	5	6	7	8	9	10	11	12	13	14	15	10
AUDITORY FUNCTION SCALE	AUM		_	7	_		ŕ		_	10	-	12	10	14	10	
4 - Consistent Movement to Command *																
3 - Reproducible Movement to Command *																Т
2 - Localization to Sound																
1 - Auditory Startle																Т
0 - None																
VISUAL FUNCTION SCALE																
5 - Object Recognition *																Т
4 - Object Localization: Reaching *																
3 - Visual Pursuit *																
2 - Fixation *																Т
1 - Visual Startle																Т
0 - None																
MOTOR FUNCTION SCALE																
6 - Functional Object Use <sup>†</sup>																Г
5 - Automatic Motor Response *																П
4 - Object Manipulation *																
3 - Localization to Noxious Stimulation *																
2 - Flexion Withdrawal																
1 - Abnormal Posturing																
0 - None/Flaccid																
OROMOTOR/VERBAL FUNCTION SCALE																
3 - Intelligible Verbalization *																Г
2 - Vocalization/Oral Movement																
1 - Oral Reflexive Movement																П
0 - None																
COMMUNICATION SCALE																
2 - Functional: Accurate †																
1 - Non-Functional: Intentional *																
0 - None																
AROUSAL SCALE																
3 - Attention																
2 - Eye Opening w/o Stimulation																L
1 - Eye Opening with Stimulation																
0 - Unarousable																
TOTAL SCORE																

Denotes emergence from MCS<sup>†</sup>
Denotes MCS \*

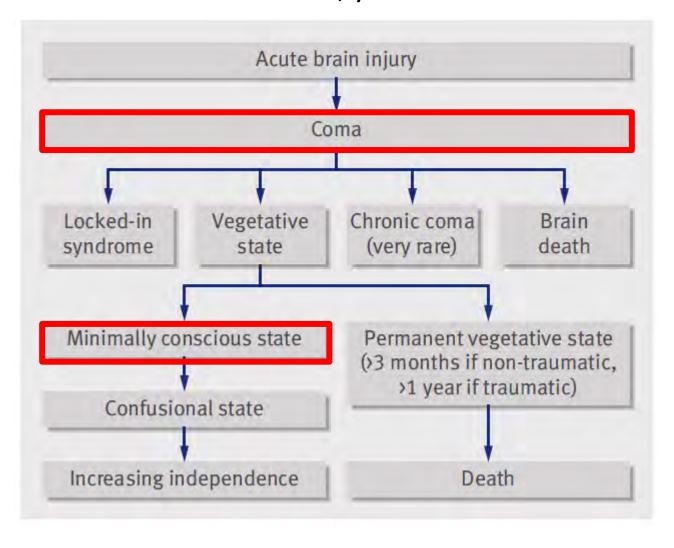




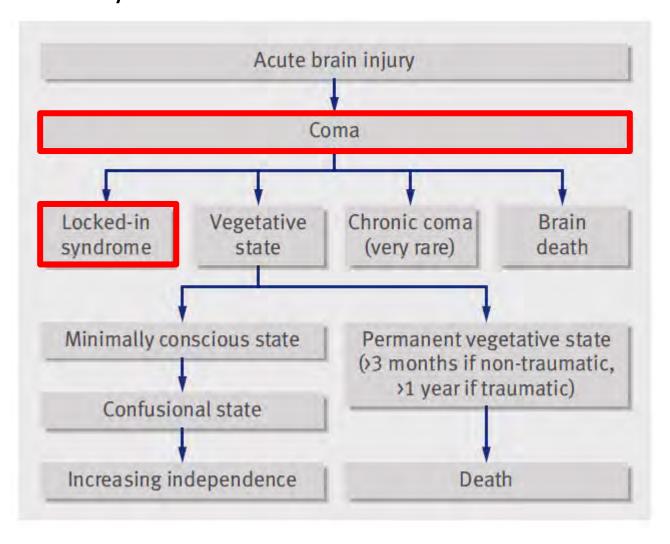
Vegetative State: the patient recovers sleep/wake cycles, yet there is no functional communication with the external world



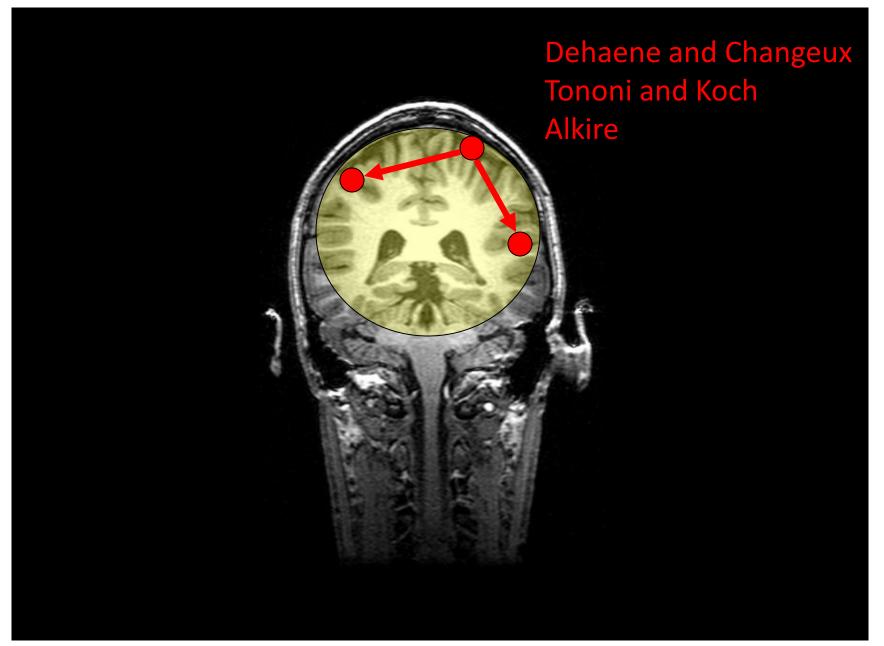
Minimally Conscious State: the patient shows inconsistent signs of interaction with the external world, yet no functional communication



Locked-In Syndrome: the patient is fully conscious, yet her/he cannot functionally communicate due to lesions of the motor pathways



#### INTRACORTICAL COMMUNICATION

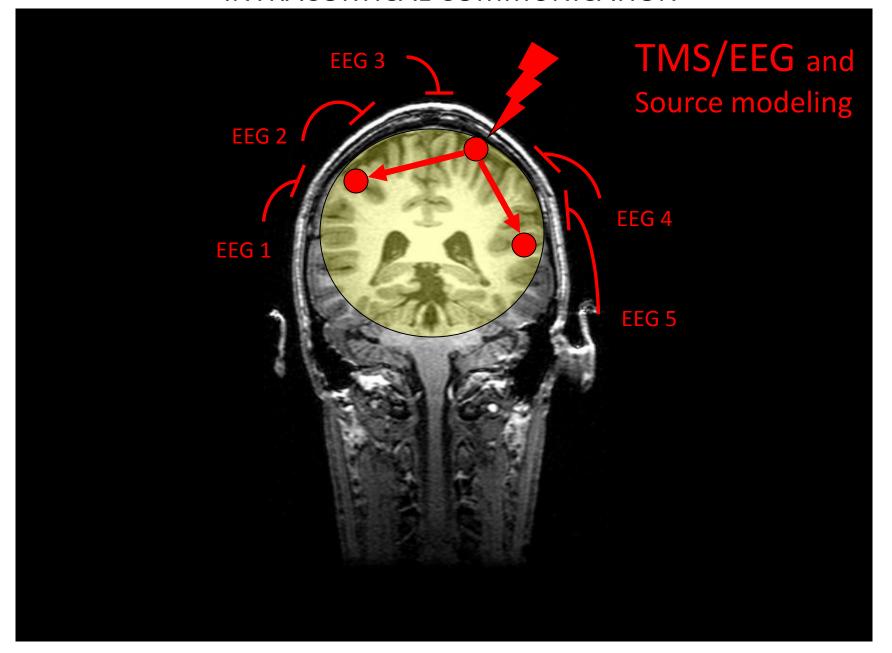


#### INTRACORTICAL COMMUNICATION



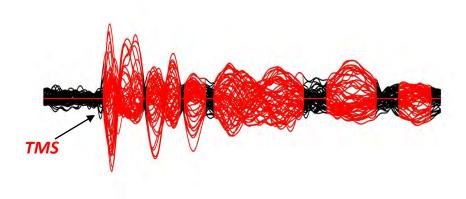
Dynamical Causal Modelling

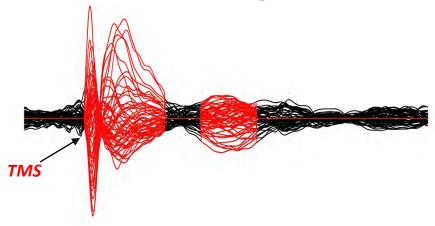
#### INTRACORTICAL COMMUNICATION



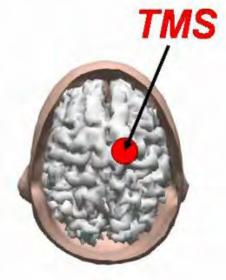


## Sleep

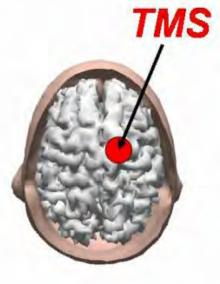




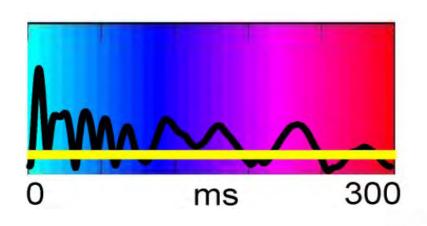
0 ms

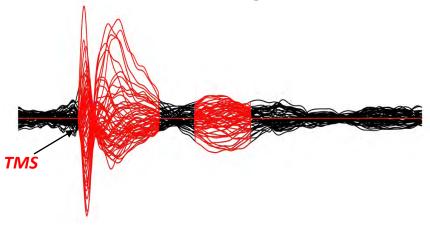


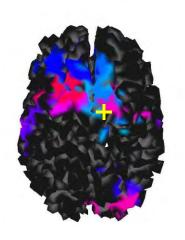
0 ms

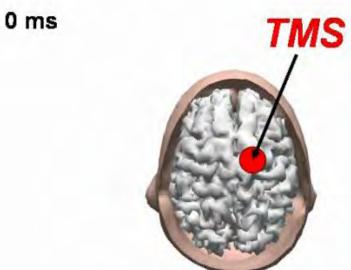


## Sleep

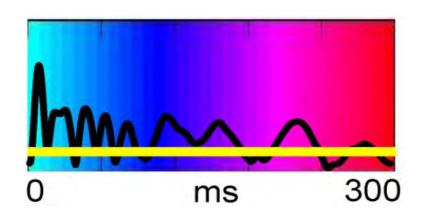


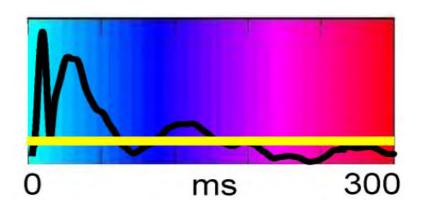


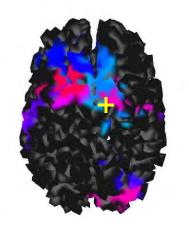


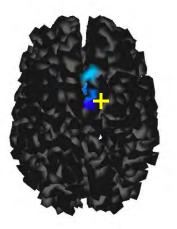


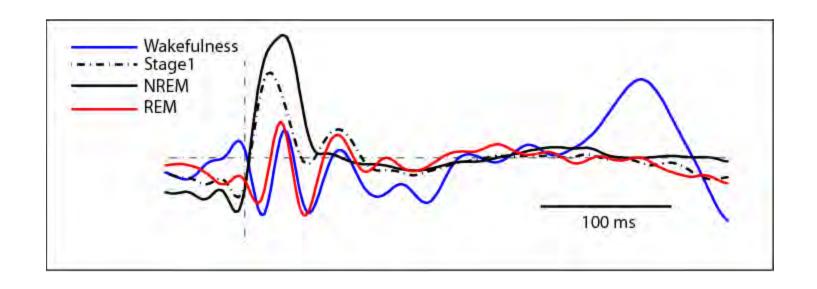
## Sleep

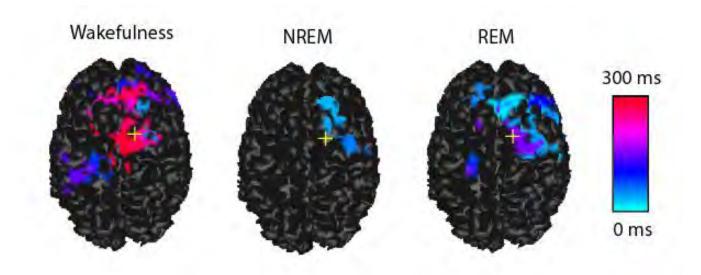






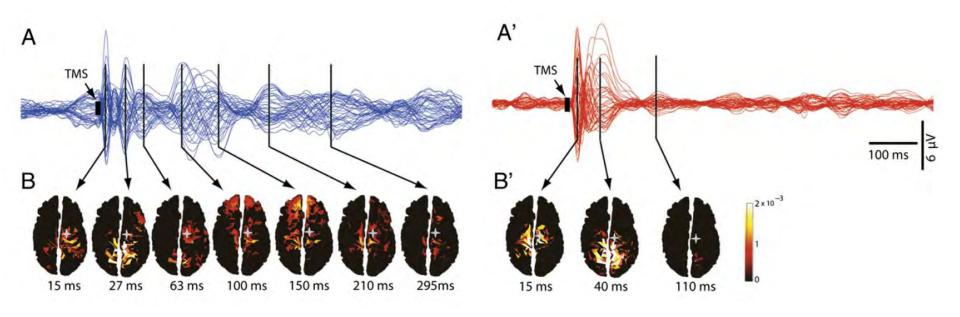






Massimini et al., Cogn. Neurosci, 2010

# Anaesthesia (Midazolam)



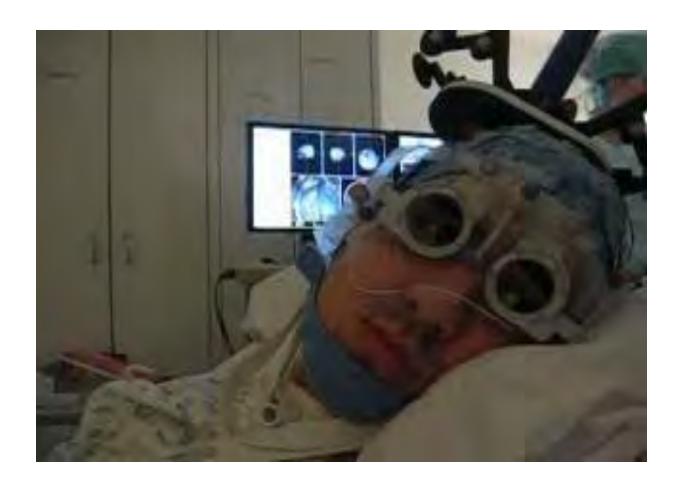


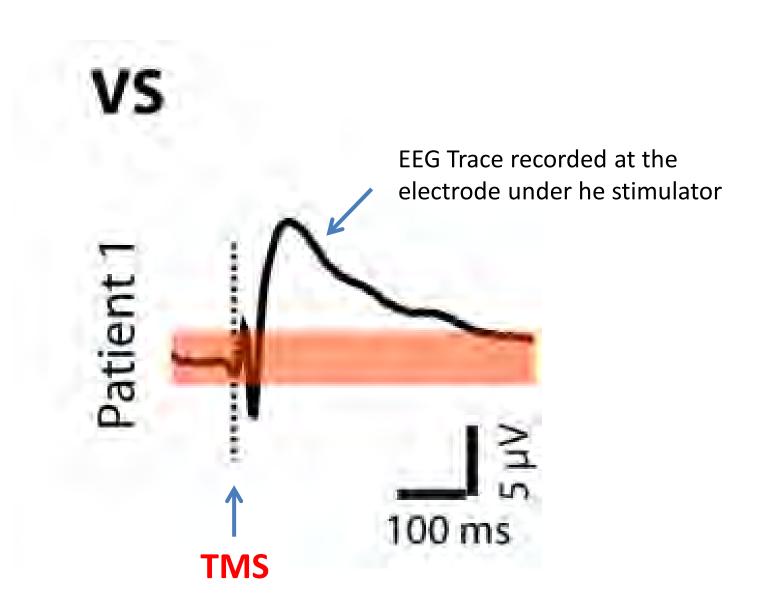
Transcranial Magnetic Stimulation combined with EEG and source modelling in non-communicating brain-injured patients:

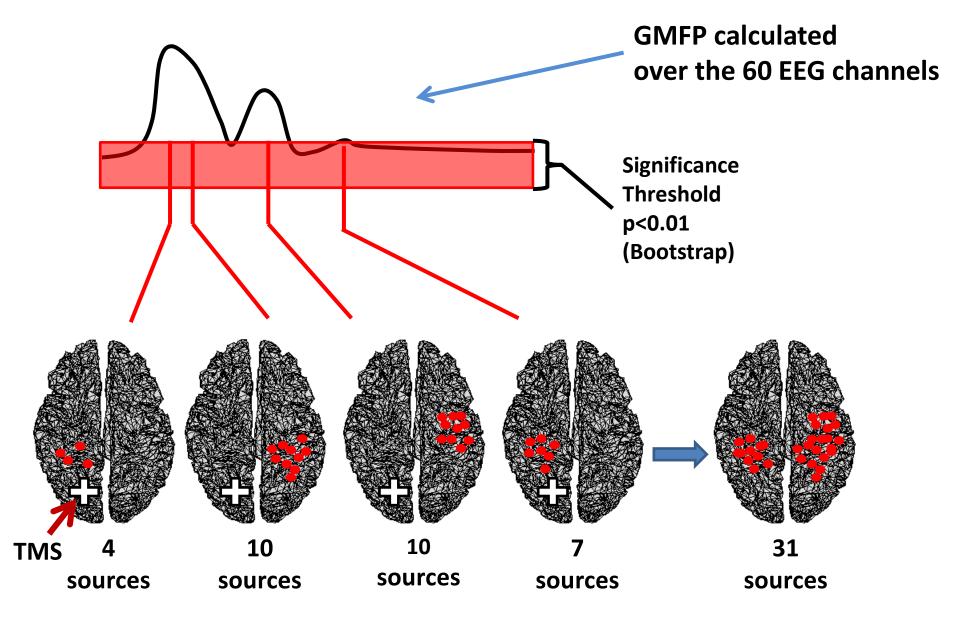
Vegetative State (**VS**) and Minimally Conscious State (**MCS**) Locked-In Syndrome (**LIS**)

#### **Vegetative State**

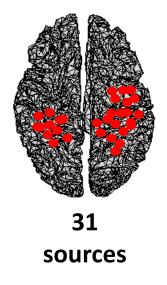
(vigilance without awareness)

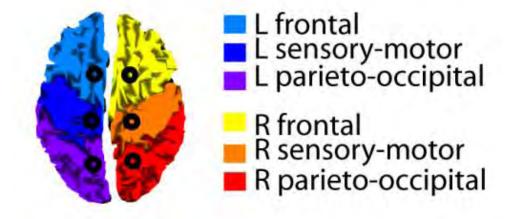


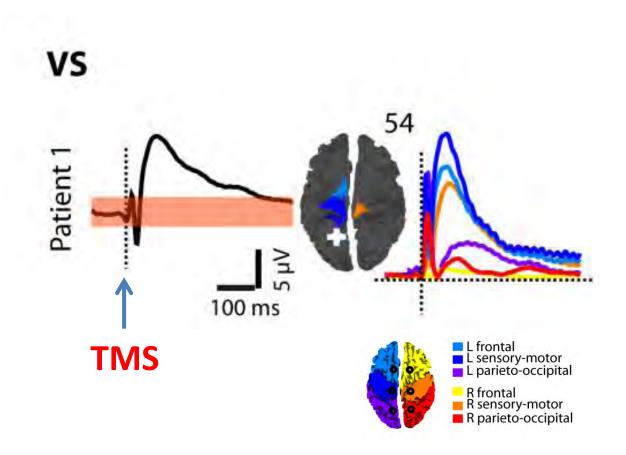


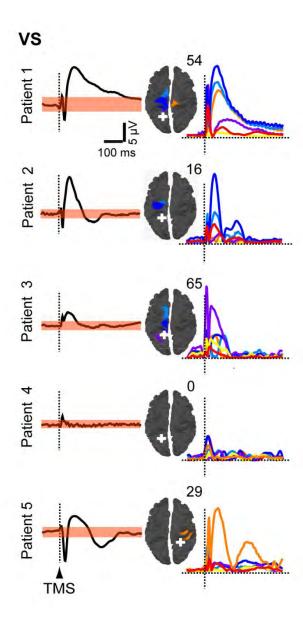


Source Modelling was computed by means of Minimum Norm Estimate (MNE)

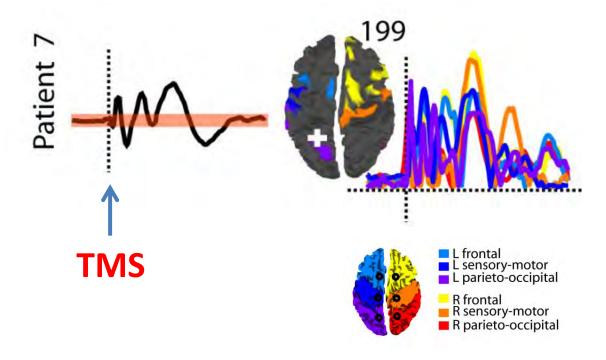


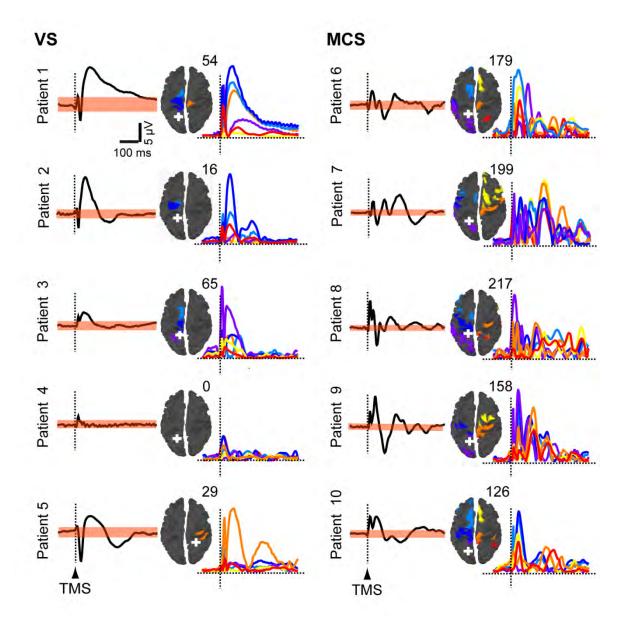




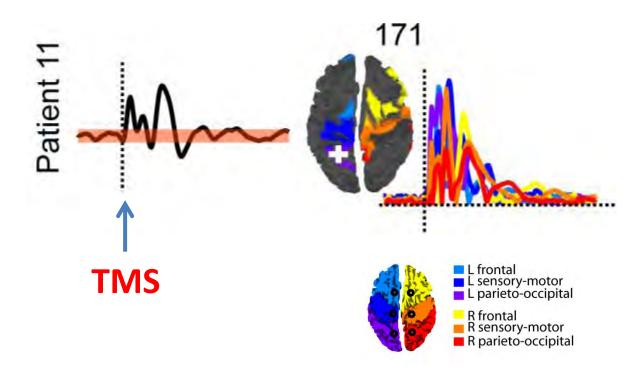


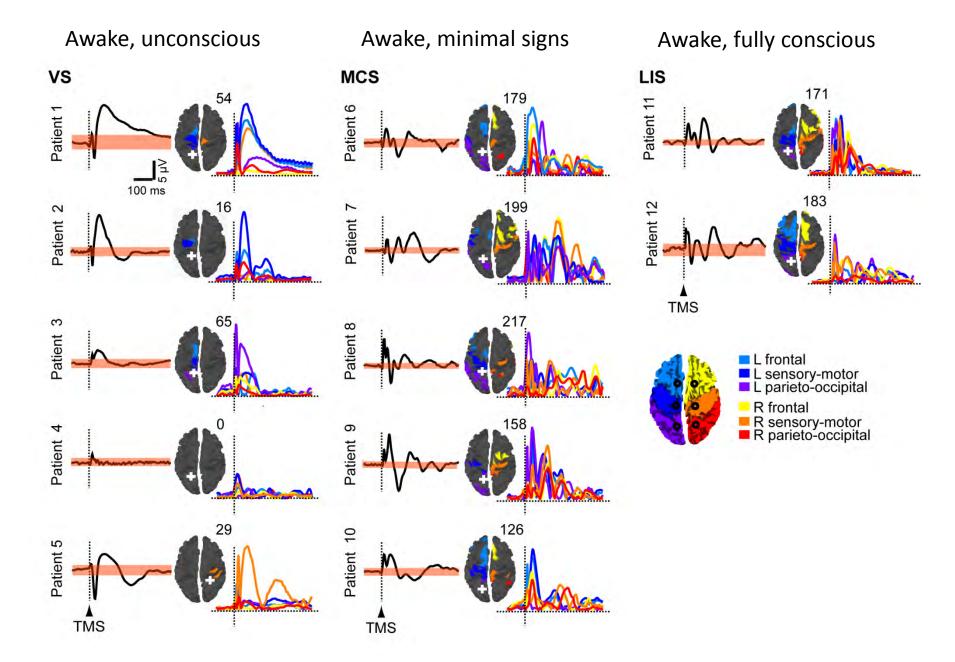
## **MCS**

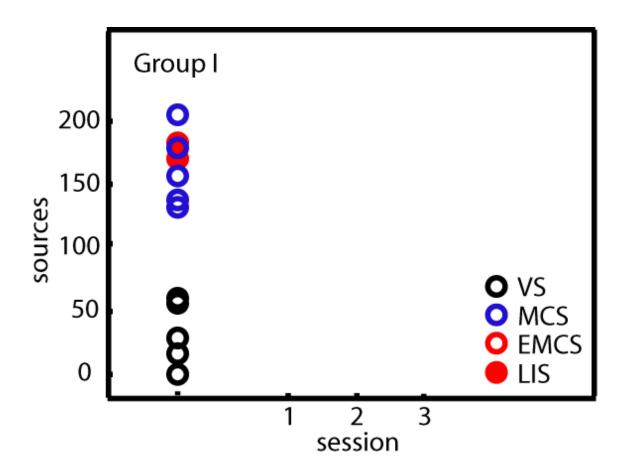




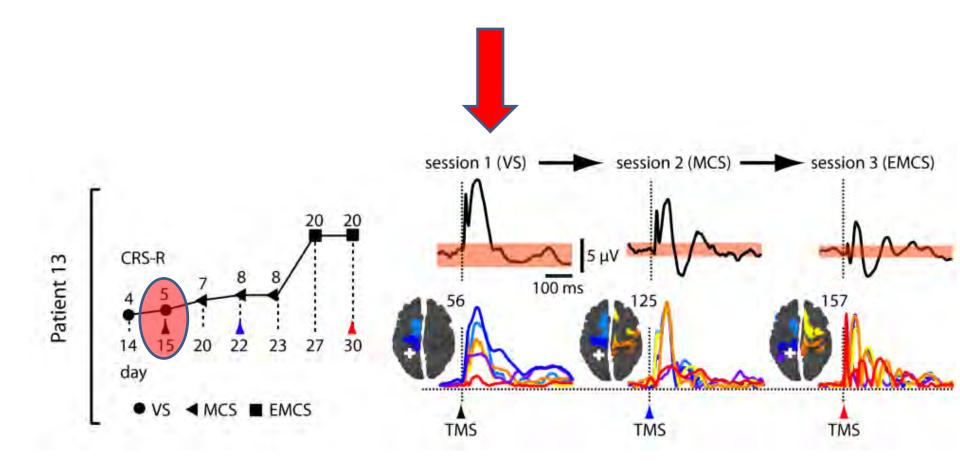
## LIS

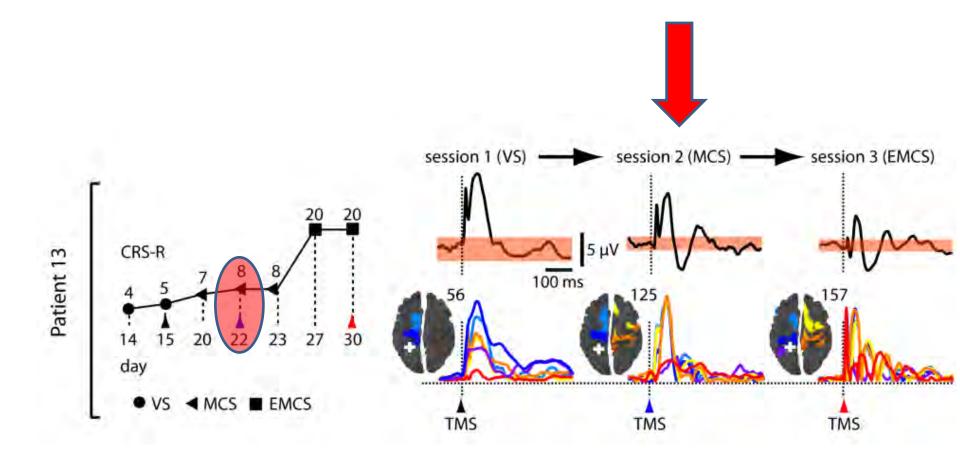


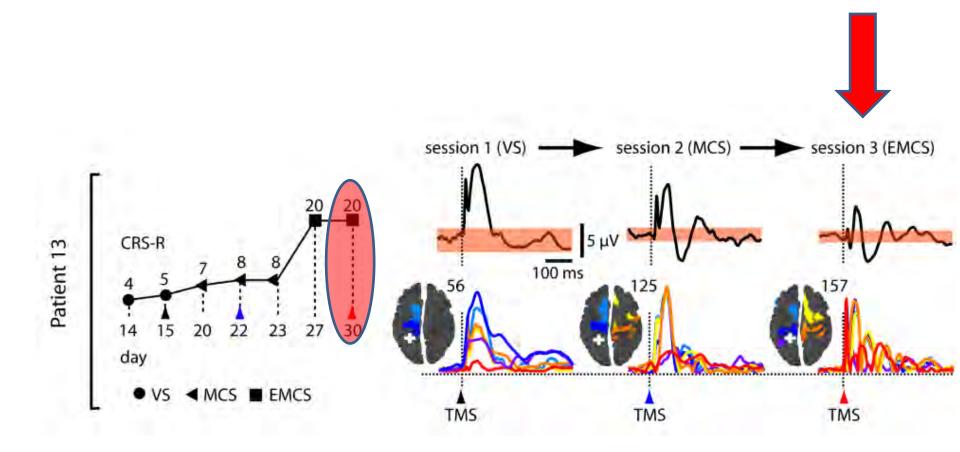




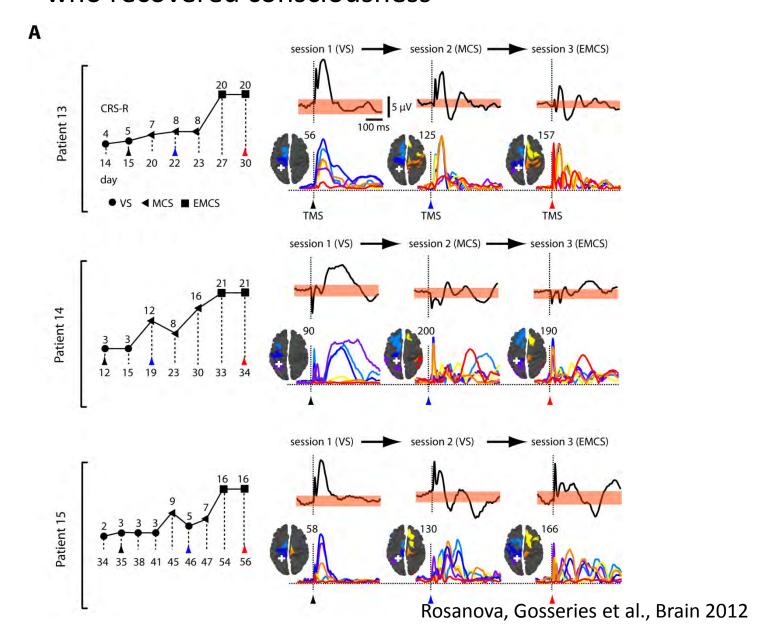
#### Longitudinal measurements in acute patients



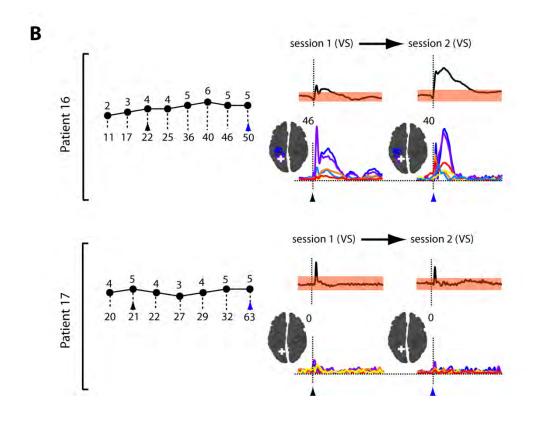




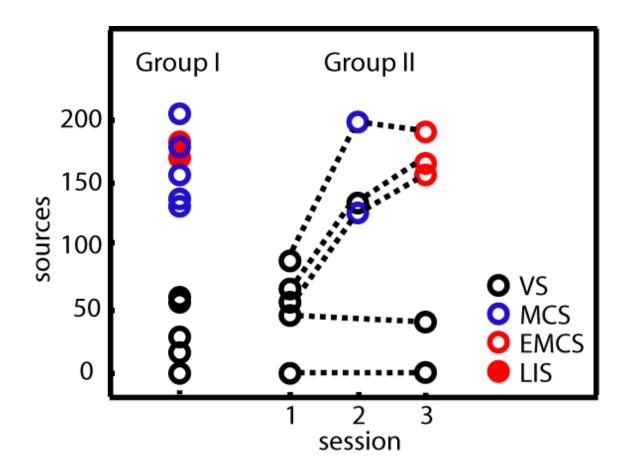
# Longitudinal measurements in the three patients who recovered consciousness



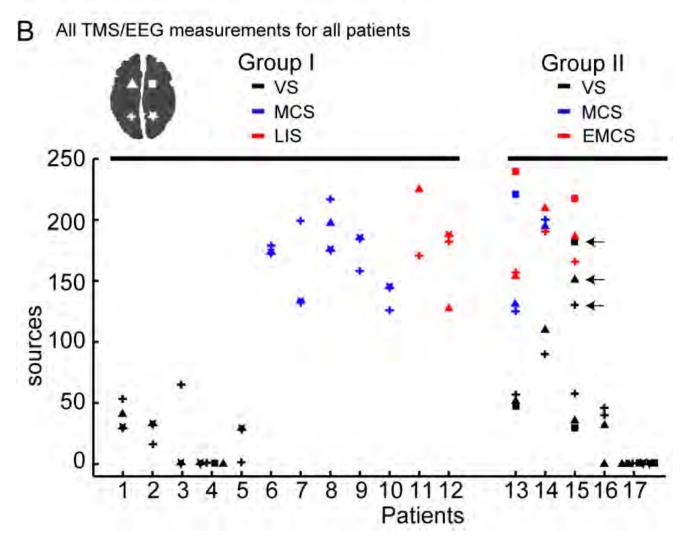
# Longitudinal measurements in the two patients who did not recover consciousness



Effective connectivity values for all patients in sessions where left parietal cortex was stimulated

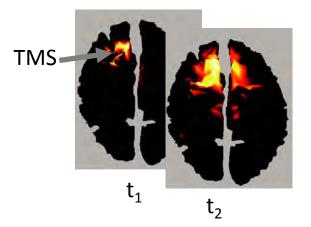


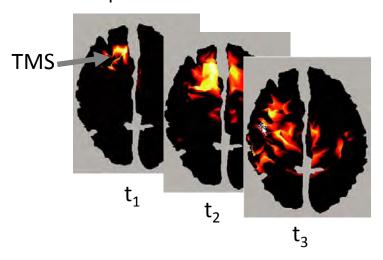
Effective connectivity values for all patients and in all sessions (also right parietal and frontal areas)

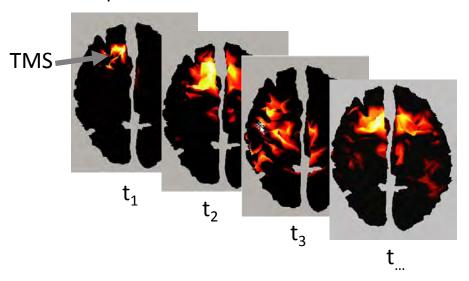


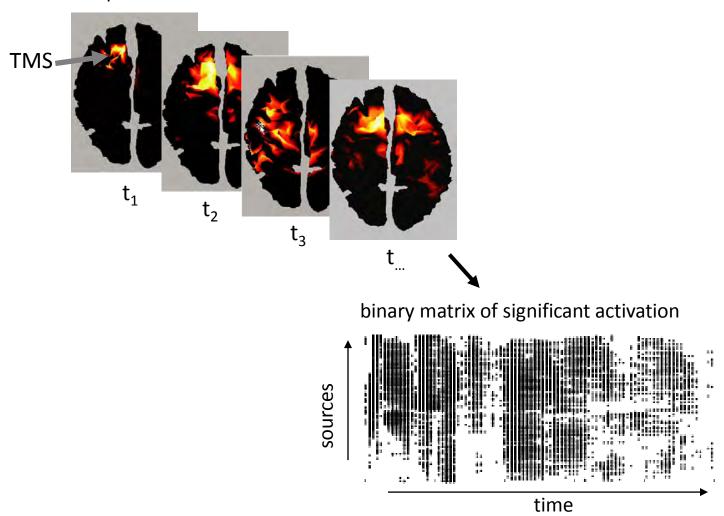


 $t_1$ 

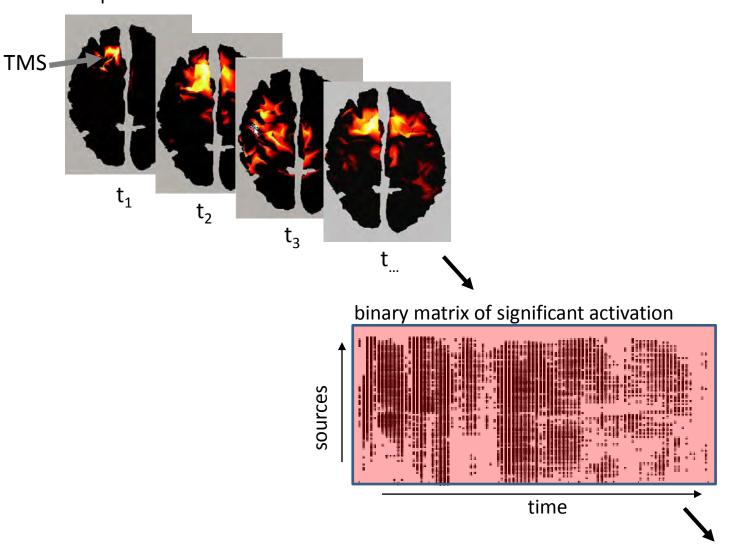






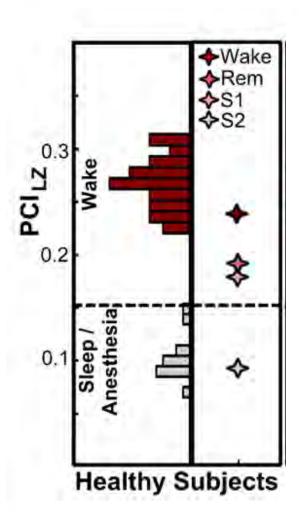


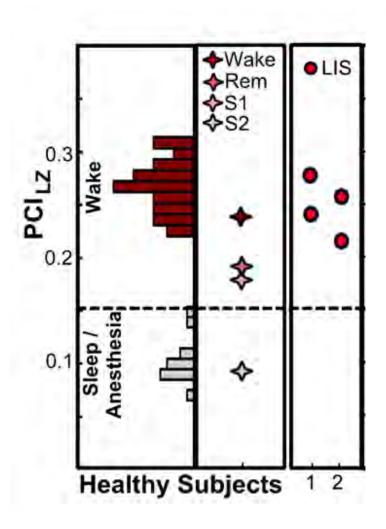
non-parametric statistics at the source level

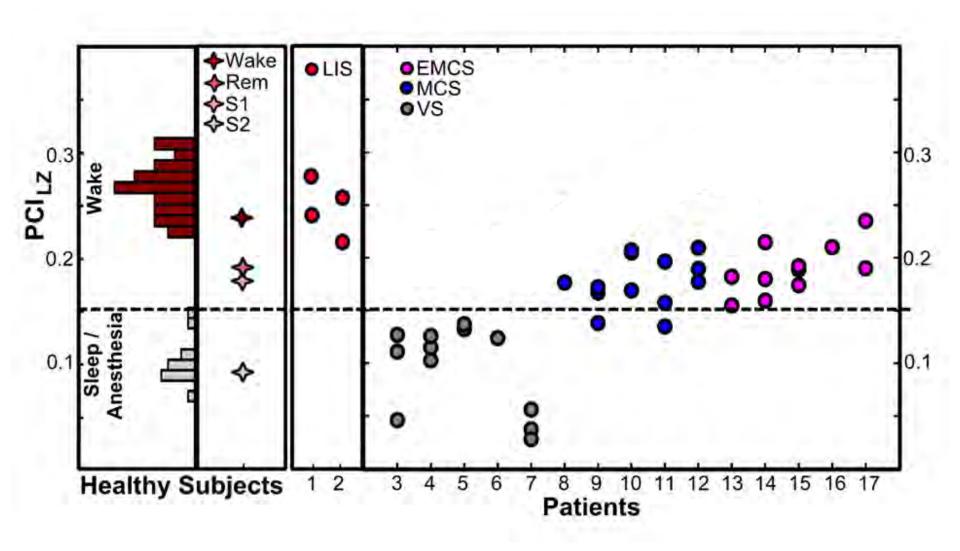


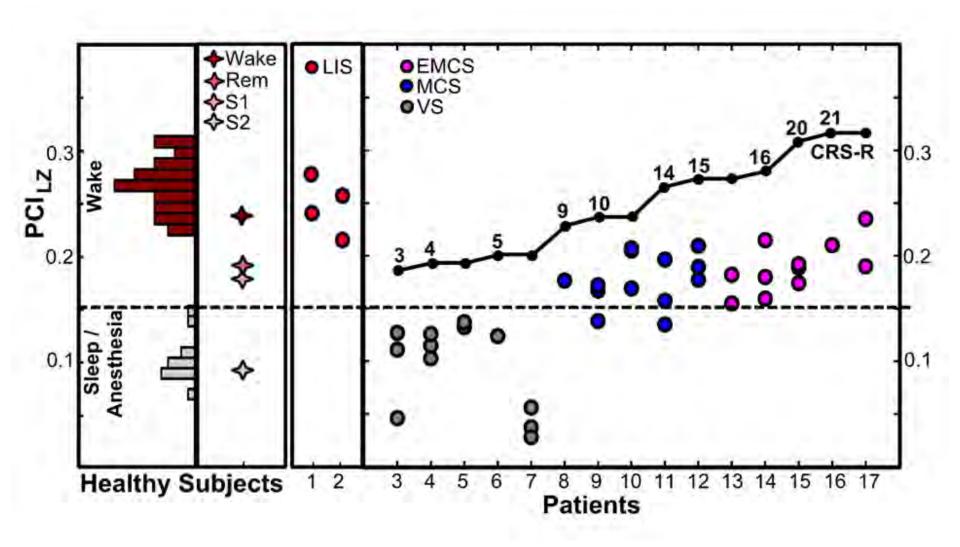
**BITS** 

(Lempel-Ziv-Welch Complexity)

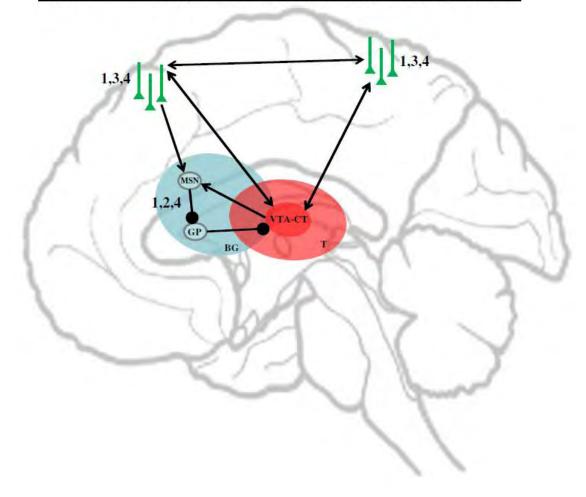








#### Possible mechanisms of action of central thalamic DBS in the injured brain



Shah and Schiff, Eur J Neurosci 2010

#### Thanks!

MILAN: MADISON: LIEGE:

Silvia Casarotto Fabio Ferrarelli Melanie Boly

Andrea Pigorini Michael Murphy Olivia Grosserie

Adenauer Casali Michael Peterson Marie-Aurelie Bruno

Matteo Fecchio Brady Riedner Pierre Boveroux

Simone Sarasso Giulio Tononi Steven Laureys

Marcello Massimini

#### Fundings:

EU-grant-224328-PredictAD STSM – COST 2009 Pur 2009 (Univ Milan) Prin 2008 (Italian Gov)