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# An Automatic and Improved MEG Method for Diagnosing and Monitoring of Neuronal and Psychiatric Disorders with a Novel Approach for Removing Correlated Noises

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## BACKGROUND

Traumatic brain injury (TBI) is a leading cause of sustained impairment in military and

civilian populations. However, mild (and some moderate) TBI can be difficult to diagnose, with only a ~10% positive-finding rate when using conventional neuroimaging methods of CT and MRI. The diffusion tensor imaging (DTI), which is still under development, has a positive-finding rate of ~20-30% in mild TBI patients.

Furthermore pre-surgical functional brain mapping of the brain regions mediating sensation, movement, and language can facilitate surgical planning and reduce function loss.

# **TECHNOLOGY DESCRIPTION**

UCSD researchers have developed a sensitive and automated diagnostic method using magnetoencephalography (MEG) to detect loci of neuronal injury and abnormal neuronal networks, not visible with CT and MRI. The invention is a fast MEG source imaging technique based on a Vector-based Spatial-Temporal Analysis for L1-minimum-norm solution, (namely Fast-VESTAL) to obtain the source amplitude images of resting-state MEG signals for different frequency bands. The Fast-VESTAL technique consists of two steps: 1) obtain L1-minimum-norm MEG source images for the dominant spatial modes of sensor-waveform covariance matrix, and 2) obtain accurate source time-courses with millisecond temporal resolution, using an inverse operator constructed from the spatial source images of Step 1.

Localizing normal and abnormal resting-state human brain signals: TBI patients generate abnormal low-frequency magnetic activity (ALFMA, peaked at 1–4 Hz) that can be measured and localized by magnetoencephalography (MEG). By applying the fast -VESTAL to selectively process the ALFMA frequency-domain, one only need to analyze the complicated MEG source signal for a few frequency bins instead of thousands of time samples in a given time window. MEG slow-wave source imaging is substantially more sensitive to mild TBI than CT, MRI, and DTI. The positive detection rate of mild TBI was 87% using MEG slow-wave source imaging for ALFMA (Huang et al., 2012). A whole brain normative database was also established for all different frequency bands (alpha band 8-12Hz, beta band 15-30 Hz, gamma band 30-100 Hz, theta band 4-7 Hz, and delta band 1-4 Hz). Using Fast-VESTAL, the first set of source images have been obtained for all the neuronal generators in the above frequency bands (Huang et al., 2014), since Hans Berger discovered the human alpha wave in 1924.

Localizing normal and abnormal brain functions in evoked brain responses for pre-surgical functional brain mapping: The application of Fast-VESTAL to human MEG median-nerve responses further demonstrated its power in reconstructing source time-courses that were highly consistent with known electrophysiology of the human somatosensory system. Fast-VESTAL provided the first set of comprehensive MEG source-amplitude images that covered the entire brain in standard atlas coordinates for different frequency bands of resting-state signals and the Fast-VESTAL solution with a low computational cost. In addition, a new objective pre-whitening method, OPWM, when used in conjunction with the Fast-VESTAL provides a novel approach to remove correlated noises. In addition to the somatosensory function, Fast-VESTAL has also been used to examine the receptive and expressive language areas (i.e., Wernicke and Broca's areas) as well as motor areas in many pre-surgical patients.

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## **OTHER INFORMATION**

**KEYWORDS** MEG, Magnetoencephalography, Traumatic brain injury, Slow-wave, Blast, Motor Vehicle Accident, Sport Injury, Alzheimer's, Post-traumatic Stress Disorder, PTSD, Neuronal, Psychiatric Disorders, VESTA

### **CATEGORIZED AS**

#### Medical

- Diagnostics
- Disease: Central Nervous
- System
- Imaging

**RELATED CASES** 2014-024-0, 2012-354-0, 2010-340-0

## **ADVANTAGES**

1) Extremely low computational costs;

2) Ability to localize and resolve a large number (up the limit determined by the number of MEG sensors) of focal and distributed neuronal

sources with any degrees of correlations;

3) Ability to obtain accurate source time-courses, and hence the accurate source-based functional connectivity at poor SNR conditions, even at

SNRs in the negative dB ranges;

- 4) Extremely low in the signal leakage of the Fast-VESTAL solution to other areas where no sources exist;
- 5) Facilitating imaging registration and group analysis by providing voxel-based whole brain imaging of MEG signal;

6) Improved sensitivity of detecting injuries and abnormalities in mild TBI and PTSD.

## **APPLICATIONS**

Diagnosis and monitoring of patients with:

• mild and moderate TBI due to blast and non-blast causes; and

• other neuronal and psychiatric disorders, especially the early stages of the disorders, including but not limited to Alzheimer's, dementia,

multiple sclerosis, autism, schizophrenia, post-traumatic stress disorder (PTSD).

## **RELATED MATERIALS**

Huang MX, Huang CW, Robb A, Angeles A, Nichols SL, Baker DG, Song T, Harrington DL, Theilmann RJ, Srinivasan R, Heister D, Diwakar M, Canive JM, Edgar JC, Chen YH, Ji Z, Shen M, El-Gabalawy F, Levy M, McLay R, Webb-Murphy J, Liu TT, Drake A, Lee RR. MEG source imaging method using fast L1 minimum-norm and its applications to signals with brain noise and human resting-state source amplitude images. Neuroimage. 2014 Jan 1;84:585-604. doi: 10.1016/j.neuroimage.2013.09.022. Epub 2013 Sep 19. - 09/19/2013
Huang MX, Nichols S, Robb A, Angeles A, Drake A, Holland M, Asmussen S, D'Andrea J, Chun W, Levy M, Cui L, Song T, Baker DG, Hammer P, McLay R, Theilmann RJ, Coimbra R, Diwakar M, Boyd C, Neff J, Liu TT, Webb-Murphy J, Farinpour R, Cheung C, Harrington DL, Heister D, Lee RR. An automatic MEG low-frequency source imaging approach for detecting injuries in mild and moderate TBI patients with blast and non-blast causes. Neuroimage. 2012 Jul 16;61(4):1067-82. doi: 10.1016/j.neuroimage.2012.04.029. Epub 2012 Apr 20. - 04/20/2012

## **PATENT STATUS**

Country	Туре	Number	Dated	Case
United States Of America	Issued Patent	10,433,742	10/19/2019	2014-024
United States Of America	Issued Patent	9,883,812	02/06/2018	2010-340

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