## Research Article

## MRI Guided Brain Stimulation without the Use of a Neuronavigation System

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A key issue in the field of noninvasive brain stimulation (NIBS) is the accurate localization of scalp positions that correspond to targeted cortical areas. The current gold standard is to combine structural and functional brain imaging with a commercially available "neuronavigation" system. However, neuronavigation systems are not commonplace outside of specialized research environments. Here we describe a technique that allows for the use of participant-specific functional and structural MRI data to guide NIBS without a neuronavigation system. Surface mesh representations of the head were generated using Brain Voyager and vectors linking key anatomical landmarks were drawn on the mesh. Our technique was then used to calculate the precise distances on the scalp corresponding to these vectors. These calculations were verified using actual measurements of the head and the technique was used to identify a scalp position corresponding to a brain area localized using functional MRI.

## 1. Introduction

Noninvasive brain stimulation (NIBS) techniques such as repetitive transcranial magnetic stimulation (rTMS) and transcranial direct current stimulation (tDCS) allow for the temporary modulation of neural activity within the human brain. rTMS involves the induction of weak electrical currents within targeted regions of the cortex via brief, time-varying magnetic fields produced with a hand-held coil [1]. tDCS employs head-mounted electrodes, which allow for a weak direct current to interact with the underlying cortex [2]. NIBS can be used to investigate the role of individual brain areas in specific cognitive, behavioral, or perceptual processl

 that evokes the strongest MEP can then be used as the location for rTMS or tDCS. A comparable technique also exists for the visual cortex whereby single pulse TMS of the occipital pole can be used to evoke the percept of a phosphene [8]. The scalp location that induces the most robust phosphene or a phosphene in a specific visual field location can be used for visual cortex stimulation. A similar technique can be used for motion sensitive, extra-striate visual area V5 whereby TMS can be used to induce moving phosphenes [9]. It has been shown that this technique is in good agreement with localization of V5 using functional magnetic resonance imaging [10]. However, it is not possible to use this approach outside of the motor and visual cortices because most brain regions do not produce acute neurophysiological or perceptual effects in response to single pulse TMS.

An alternative technique for identifying participantspecific stimulation sites on the scalp is the 10–20-electrode system, which was originally designed for positioning EEG electrodes [11]. This approach defines a grid of positions on the scalp that are separated by 10% or 20% of the distance between anatomical landmarks such as the nasion and the inion. This approach has been used successfully in a large number of brain stimulation studies; however, the mapping of particular 10–20 system locations to specific brain areas can vary across participants [12].

Another alternative is to use structural and functional brain imaging techniques to localize specific brain areas in individuals with millimetre resolution. A number of frameless stereotactic navigation systems exist for real-time coregistration of a participant to their own MRI images. Tools such as a "pointer" or a TMS coil can also be registered within the volume. These systems typically involve ultrasound devices or infrared cameras and a number of reference targets mounted on the head and NIBS apparatus. When used in combination with structural and functional MRI images these "neuronavigation" systems allow for precise identification of the scalp position corresponding to a particular brain area [13].

The combination of brain imaging and a neuronavigation system is the current gold standard in the field of NIBS [14] and may improve the results of NIBS-based therapeutic interventions [15–20]; however, there are some disadvantages. These include difficulty in using these systems for studies of posterior brain areas that can fall outside of the neuronavigation system's field of view and, most importantly, the high cost of these systems, which can exceed \$50,000. Techniques have been described that allow NIBS to be targeted using generic MRI datasets [21



FIGURE 1: A 3D mesh morphed to the structural MRI data of a representative participant. Panels (a)–(d) show the anatomical landmarks that were used as anchor points for scalp distance calculations marked on a T1-volume surface mesh created using Brain Voyager. N: nasion, RT and LT: right and left tragi, respectively, and IN: inion. The lines connecting the anatomical landmarks are "patches of interest" (POIs) drawn in Brain Voyager that link adjacent triangles in the mesh. Panels (e) and (f) show close-up views of the mesh without the surface coloring. The mesh has been cut axially at the level of the inion. The smooth surface of the head is represented using triangular elements and each of these elements is defined by its tricorners.

subroutines within Brain Voyager. A general linear analysis was conducted and the results were visualized as t-maps on the anatomical image. Area V5 was identified as a region in the appropriate anatomical location that responded significantly more strongly to dynamic than static grating stimuli (FDR corrected q < 0.01). The precise location of V5 was defined as the location of the peak voxel within the V5 region.

2.3. Comparison of Measurements Made on the Surface Mesh and the Head. Four anatomical landmarks were identified on each surface mesh: the nasion, the left and right tragi, and the inion (Figures 1(a)–1(d)). The shortest paths between the nasion and inion and the left and right tragi that passed through the center point of the head (Cz) were then marked on the surface of the 3D mesh and exported as "patches of interest" (POIs) within Brain Voyager. After this, the *x*, *y*, and *z* coordinates oe the mesh nodes that formed the POI were exported from Brain Voyager in XLS format and read into the MATLAB analysis environment for distance calculations. The actual distances between the two tragi and the nasion and inion were also measured for each participant using a tape measure. An investigator masked to the results of the MATLAB analysis made these measurements. 2.4. MATLAB Operations. A Graphical User Interface was created in MATLAB to import the coordinate matrix of the POI exported from the Brain Voyager environment. Since the aim was to develop a widely applicable tool, the software does not require the use of Brian Voyager. Rather, the software is capable of reading a coordinate matrix from a text/MS-Excel file as this format is an export option in most image postprocessing software packages. The file must have three columns (x, y, z), which conform to the following format:

$$Nodal_{vector} = (Node(n), Node(n), Node(n)).$$
 (1)

Here, n is the index of the nodal coordinate in the vector matrix. Subscripts x, y, and z indicate the Cartesian tricoordinates of the vector's nodal points.

The MATLAB code opens the text-based input file and searches for the first line of the nodal coordinate series. Next, it reads consecutive coordinates until the pattern is broken; that is, no further coordinates are listed. The nodes of the POI/vector can then be viewed immediately in 3D (Figure 2).

There are two main issues to be addressed when calculating scalp distances from POIs measured on a surface mesh. (1) The majority of packages that provide surface meshes (Brain Voyager included) export POIs across the mesh in a proprietary format that makes it difficult to identify adjacent







(a)

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