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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 processes [3]. In addition, these techniques are being investigated from a clinical perspective and current evidence suggests that NIBS may be applicable to the treatment of multiple neurological and psychiatric disorders [4, 5].

Studies involving the use of NIBS begin by selecting a target brain area for stimulation. This process is typically informed by evidence from brain imaging, animal neurophysiology, or studies involving neurological patients. Subsequent steps include the selection of appropriate stimulation parameters and ensuring that the stimulation is delivered to the correct brain area. This latter point is particularly important as the stimulation effects are most pronounced in close proximity to the rTMS coil and tDCS electrodes [6]. Therefore, accurate, participant-specific localization of stimulation sites on the scalp is required for optimal stimulation [7].

A number of approaches can be used to identify the correct scalp position for stimulation. Single pulse TMS can be used to activate specific regions of the primary motor cortex resulting in motor evoked potentials (MEPs) within the corresponding peripheral muscle [8]. The scalp location

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 [1]. 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 V₁ whereby TMS can be used to induce moving phosphenes [2]. It has been shown that this technique is in good agreement with localization of V₁ using functional magnetic resonance imaging [3]. 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 participant-specific stimulation sites on the scalp is the 10–20-electrode system, which was originally designed for positioning EEG electrodes [4]. 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 theinion. 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 [5].

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 [6].

The combination of brain imaging and a neuronavigation system is the current gold standard in the field of NIBS [7] and may improve the results of NIBS-based therapeutic interventions [8–10]; 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 \$10,000. Techniques have been described that allow NIBS to be targeted using generic MRI datasets [11] or when structural but not functional MRI data are available for individual participants [12]. Furthermore, techniques for identifying optimal scalp locations for stimulation based on individual participant’s neuroanatomy are also available [13]. However, each of these approaches requires the use of a neuronavigation system. Here we describe a technique that allows the use of individual structural and functional MRI to guide NIBS in the absence of a neuronavigation system. The approach is based on vectors drawn on a mesh that is morphed to participant-specific MRI data. These mesh vectors are then transposed to the participant’s head by converting them to

head measurements anchored to anatomical landmarks. We report comparisons between measurements made using our technique and actual head measurements. We also give an example of how the technique can be used in combination with fMRI to localize a stimulation site for visual area V₁ in a single subject. Visual area V₁ was chosen for this example as it can be readily localized using fMRI and the corresponding scalp position cannot be identified based on a single anatomical landmark. Therefore, a number of measurements

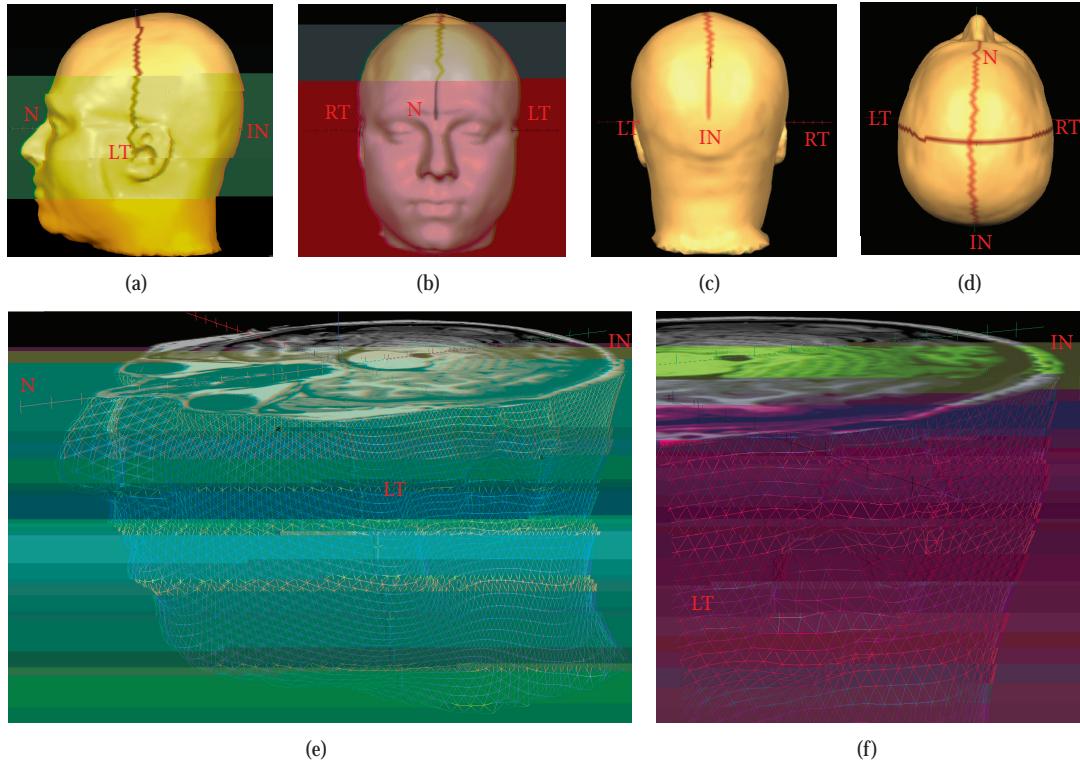
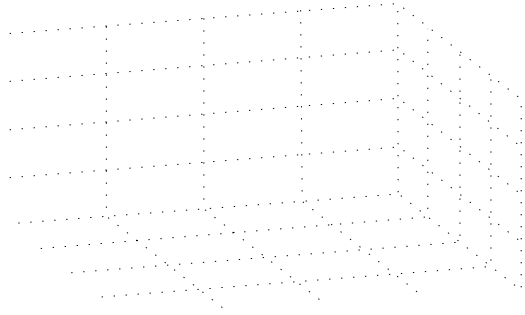
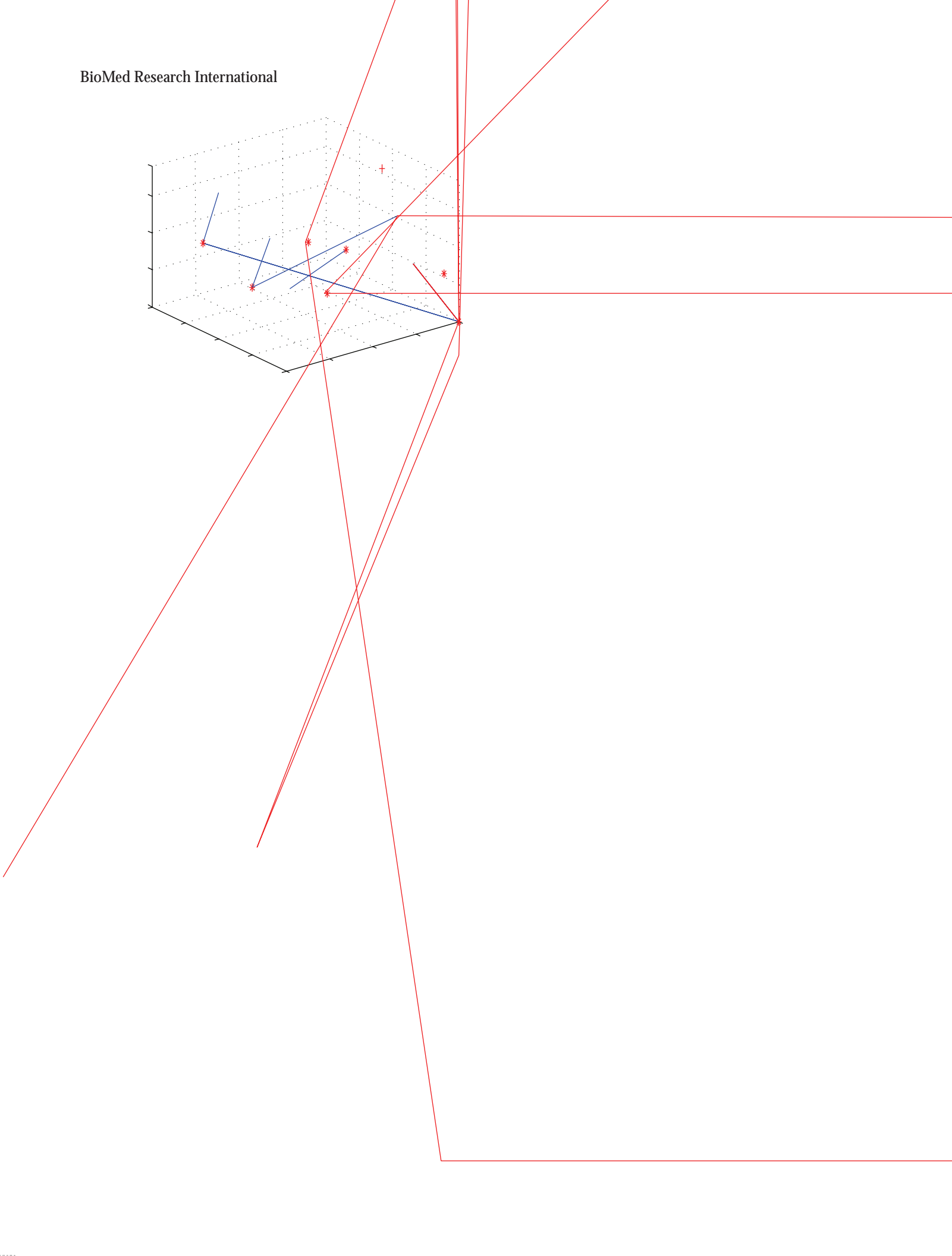
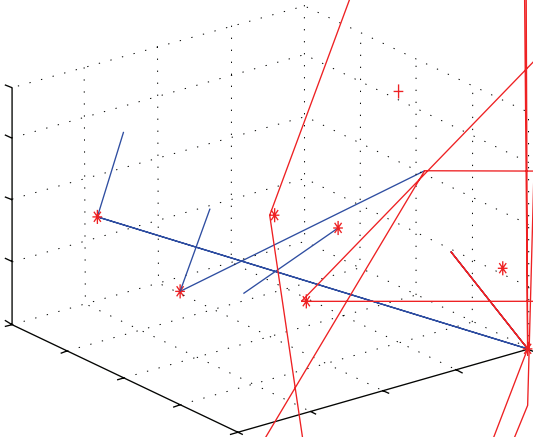
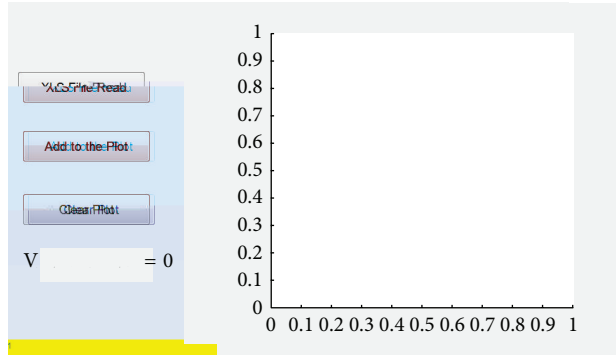


Figure 1: A 3D mesh morphed to the structural MRI data of a representative participant. Panels (a)–(d) show the anatomical landmarks that were used as anchor points for scalp distance calculations marked on a T₁-volume surface mesh created using Brain Voyager. N: nasion, RT and LT: right and left tragi, 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.

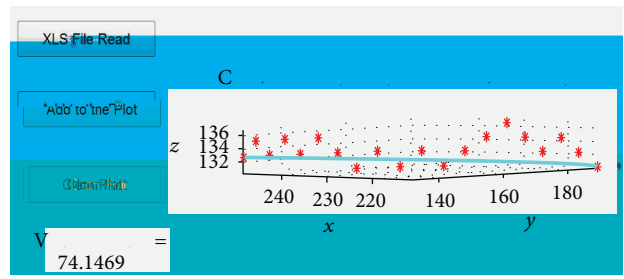
subroutines within Brain Voyager. A general linear analysis was conducted and the results were visualized as t-maps on the anatomical image. Area V was identified as a



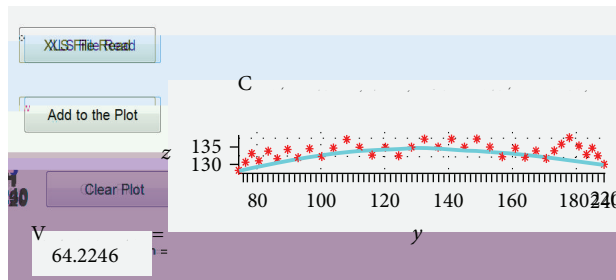




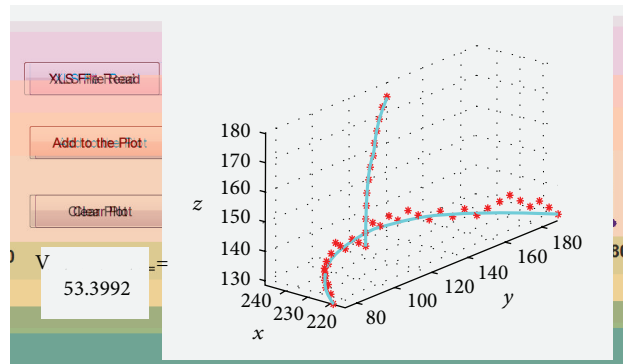
(a)



(b)



(c)



(d)

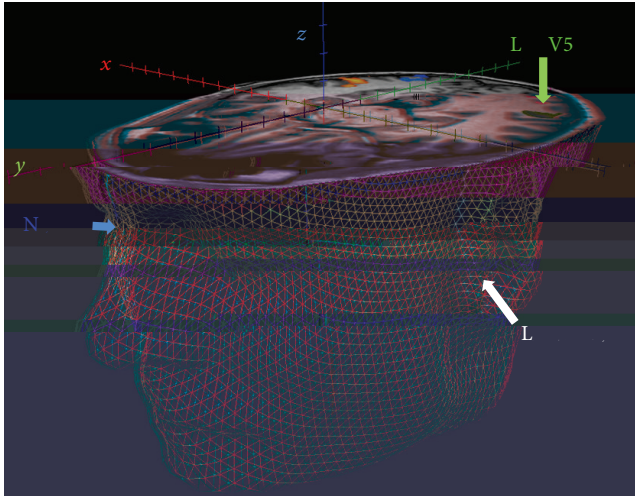


Figure 1: Localization of a scalp position above V5 in the left hemisphere. The axial cut through the Brain Voyager mesh was positioned to reveal the most active voxel in left V5. The lines drawn on the mesh show the POIs that were used to identify the scalp location corresponding to left V5. Blue: nasion to transverse plane, white: tragus to transverse plane, red: intersection of nasion vector and transverse plane to intersection of tragus vector and transverse plane, and green: extension of the vector to the scalp position above area V5. Orange regions indicate areas of functional activation in response to the V5 localization scans. See the main text for a detailed description of this procedure.

not supported. This issue is also relevant to the use of neuronavigation systems. Selection of the optimal stimulation site is a complex process as the electrical current generated by NIBS techniques interacts with the head and brain anatomy in ways that are unique to each participant [15–17]. A number of techniques for identifying optimal NIBS sites based on MRI data have been developed. These could be combined with our approach for transposing stimulation sites to the head to further improve the targeting of NIBS when neuronavigation systems are not available.

Conflicts of Interest

The authors declare that there is no conflict of interests regarding the publication of this paper.

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