

# Detecting gamma frequency neural activity using simultaneous multiband EEG-fMRI

Makoto Uji<sup>a</sup>, Ross Wilson<sup>a</sup>, Susan T. Francis<sup>b</sup>, Karen J. Mullinger<sup>a,b\*</sup>, and Stephen D. Mayhew<sup>a\*</sup>

[s.d.mayhew@bham.ac.uk](mailto:s.d.mayhew@bham.ac.uk)

a. Centre for Human Brain Health (CHBH), University of Birmingham, Birmingham, UK.

b. SPMIC, School of Physics and Astronomy, University of Nottingham, Nottingham, UK.

\*equal leads in this work

**Introduction:** Synchronization of gamma frequency (>35Hz) EEG activity is linked to cognitive and sensory behaviour as well as being widely cited as the closest neuronal correlate of the BOLD fMRI signal<sup>[1]</sup>. However, the majority of gamma-BOLD studies were conducted in the visual<sup>[2,3]</sup> or auditory<sup>[4,5]</sup> modalities, therefore a deeper understanding necessitates extension to the motor domain. Simultaneous EEG-fMRI is an ideal method to investigate gamma-BOLD correlates non-invasively in humans, however, residual gradient artefacts typically obscure gamma frequency EEG activity when acquired with fMRI. Accelerated fMRI methods such as multiband (MB)<sup>[6,7]</sup> allow whole-brain coverage in a sparse fMRI scheme which incorporates MR gradient “quiet periods” thus potentially useful to overcome EEG gradient artefacts during fMRI acquisition.

**Aim: To assess: (1) the feasibility of simultaneous EEG-MBfMRI, both safety aspects related to the higher RF power of MB excitation and EPI image quality<sup>[8,9,10]</sup>; (2) the potential to investigate relationship between gamma activity and BOLD responses in motor cortex.**

## Methods

EEG data were acquired from 64 scalp channels at 5kHz (Brain Products). MB fMRI acquisition (Gyrottools) used a 3T Philips Achieva scanner.

### Safety testing – agar phantom

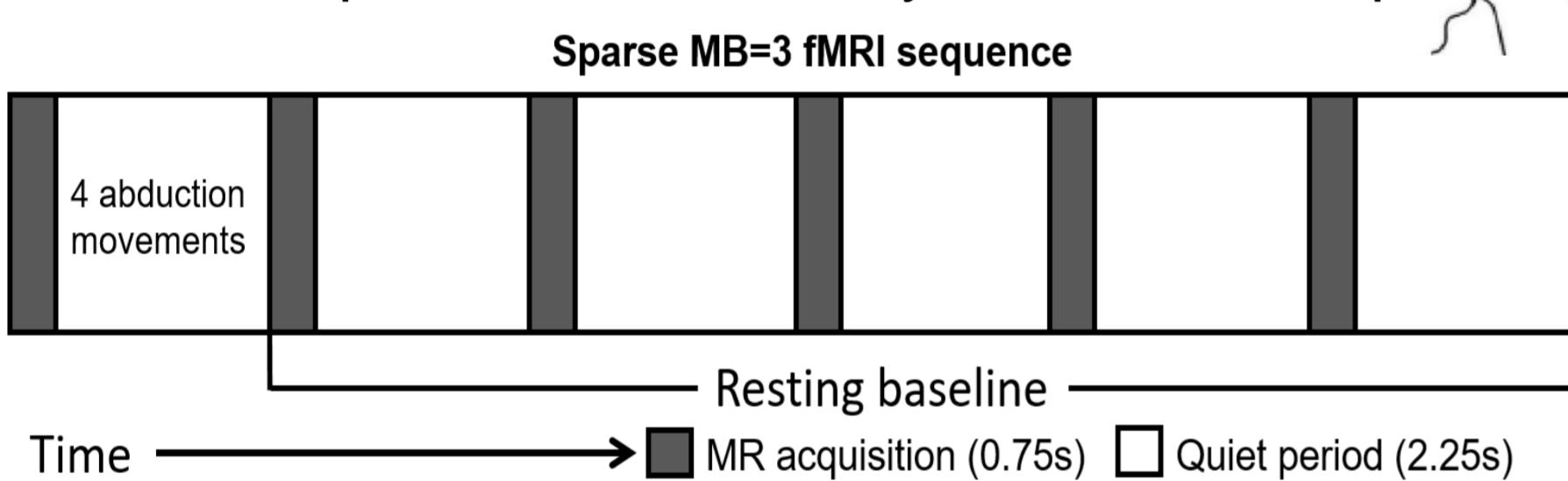
- The EEG cap was connected to a conductive agar head-shaped phantom
- Fibre-optic thermometers (Luxtron) monitored the temperature change during 20-minute scans with MB factor = 4 and SPIR fat suppression.
- Tested upper SAR limits using:
  - GE-EPI (TR/TE=1000/40ms, SENSE=2, slices=48, B1 RMS=1.09 $\mu$ T, SAR/head=22%)
  - PCASL-GE-EPI (TR/TE=3500/10ms, SENSE=2, slices=32, B1 RMS=1.58 $\mu$ T, SAR/head=46%)

### Image quality – 3 healthy adults

- Tested 5 GE-EPI sequences (Table.1).
- Image quality was assessed by comparing temporal SNR (tSNR) of grey matter voxels.

### EEG-fMRI motor study – 10 healthy adults (8M)

- 4 right-hand index finger abduction movements per trial with 16s interstimulus interval (Fig.1)
- All abductions performed in 1.5s period, each in response to an auditory cue, 1kHz beep.



**Figure 1: Experimental paradigm**

- 4 runs of 30 trials, 120 trials in total per subject
- Sparse GE-EPI: MB=3, SENSE=2, TR/TE=3000/40ms, 33 slices, FA=79°, volumes=192, voxels=3mm<sup>3</sup>, SAR/head < 7%
- EEG-MRI clocks synchronised
- EEG electrode positions digitised (Polhemus)

## Data analysis

### EEG

- Gradient and pulse artefacts were corrected, data downsampled (600Hz) and epoched -16–2s relative to auditory cue onset (BrainVision Analyzer2).
- Trials contaminated with movement artefacts were removed. Eye-blinks/movements were removed (ICA, EEGLAB) and data were average referenced.
- An LCMV beamformer<sup>[11]</sup> (noise regularization=1%) was employed with individual BEM head models (Fieldtrip<sup>[12]</sup>) to create T-stat images of changes in gamma (55–80Hz) power to finger abductions [active: 0–1.5s & passive: -9.0 to -7.5s windows].
- A broadband (1-120Hz) timecourse of neural activity was extracted from the peak T-stat location in the contralateral (left) motor cortex (cM1). Time-frequency spectrograms were calculated using multitaper wavelets<sup>[3]</sup>.
- The mean gamma power per trial (0-1.5s after auditory cue onset) was mean subtracted to form a regressor for fMRI analysis.

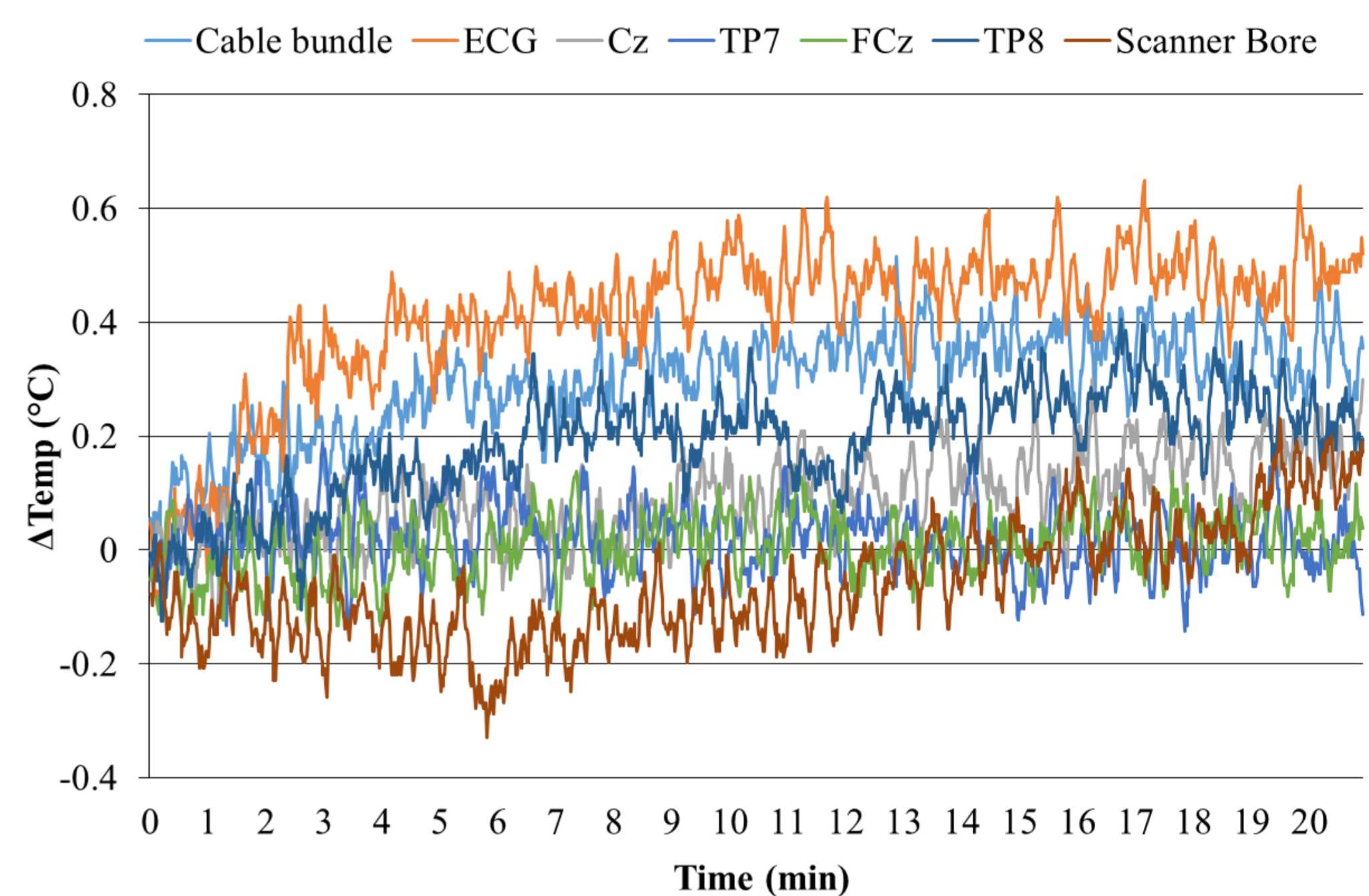
### fMRI

- Data were motion corrected, smoothed (5mm) and normalised to the MNI template (FSL).
- First-level GLM analysis employed 2 regressors: 1) boxcar abduction movement, 2) parametric modulation of single-trial gamma activity, convolved with the double-gamma HRF.
- Data were grouped over runs and subjects using second-level fixed and third-level mixed effects.

## Heating effects

- The greatest temperature change seen for the GE-EPI sequence was 0.6°C in the ECG electrode (Fig 2)
- The PCASL-GE-EPI resulted in the greatest heating effect (ECG ~0.9°C)

**Figure 2: Temperature changes at EEG electrodes, cable bundle and a control location on the scanner bore during a 20-minute GE-EPI scan. Temperature changes were calculated relative to a 5-minute baseline recording made before the scan.**



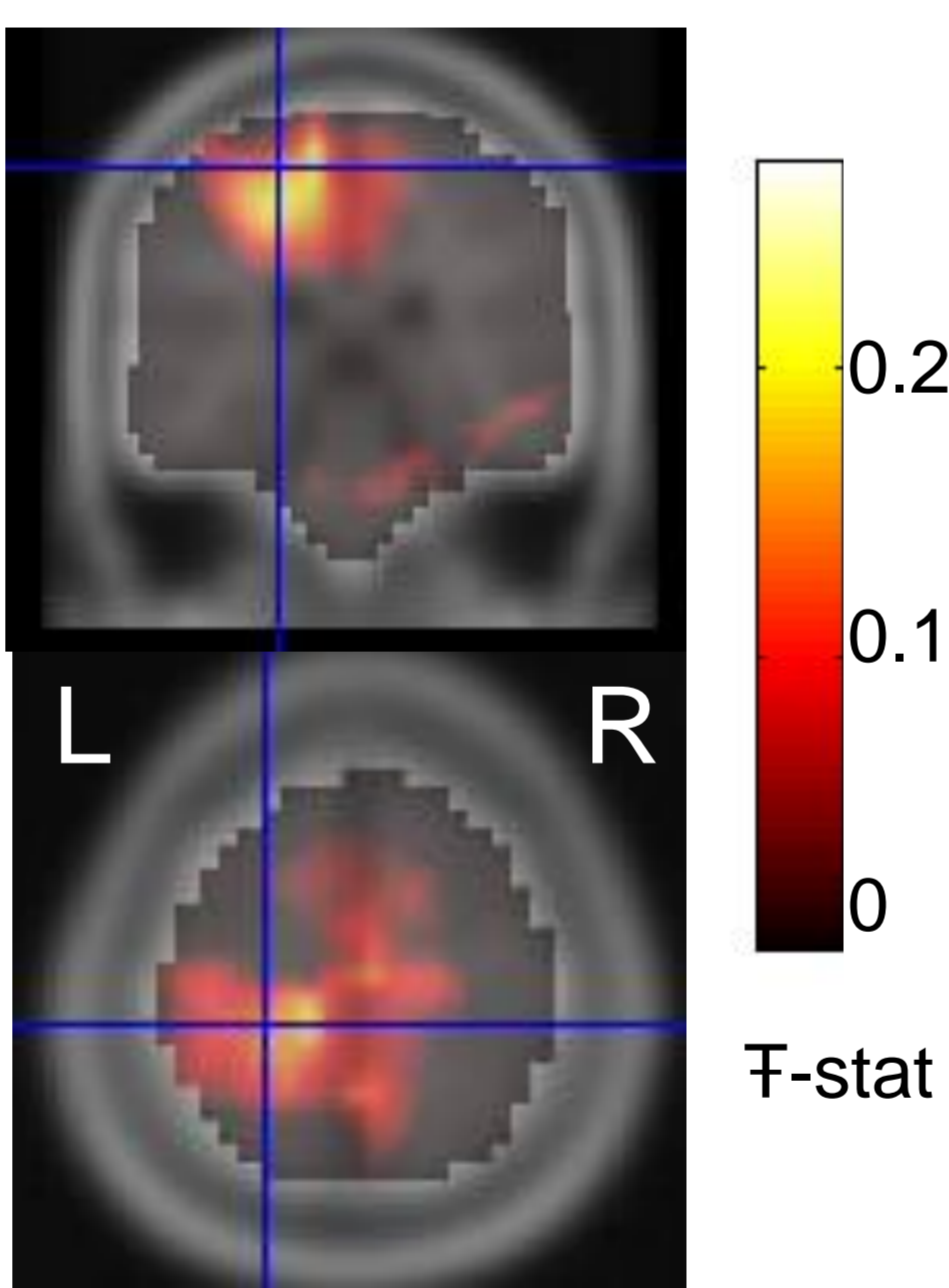
## Image quality

- Little variability in tSNR between sequences
- MB=3, sparse acquisition chosen for further experiments (red box)

**Table 1: Mean tSNR calculated during five MR sequences: MB: 1-3; acquisition type = equidistant or sparse. All other parameters were constant: TR/TE=3060/40ms, SENSE=2, slices=36, FA=79°, voxels=3mm<sup>3</sup>, volumes=41**

Multiband Factor	Slice acquisition spacing	tSNR $\pm$ SD
1	Equidistant	74 $\pm$ 40
2	Equidistant	72 $\pm$ 39
2	Sparse	67 $\pm$ 37
3	Equidistant	68 $\pm$ 37
3	Sparse	74 $\pm$ 38

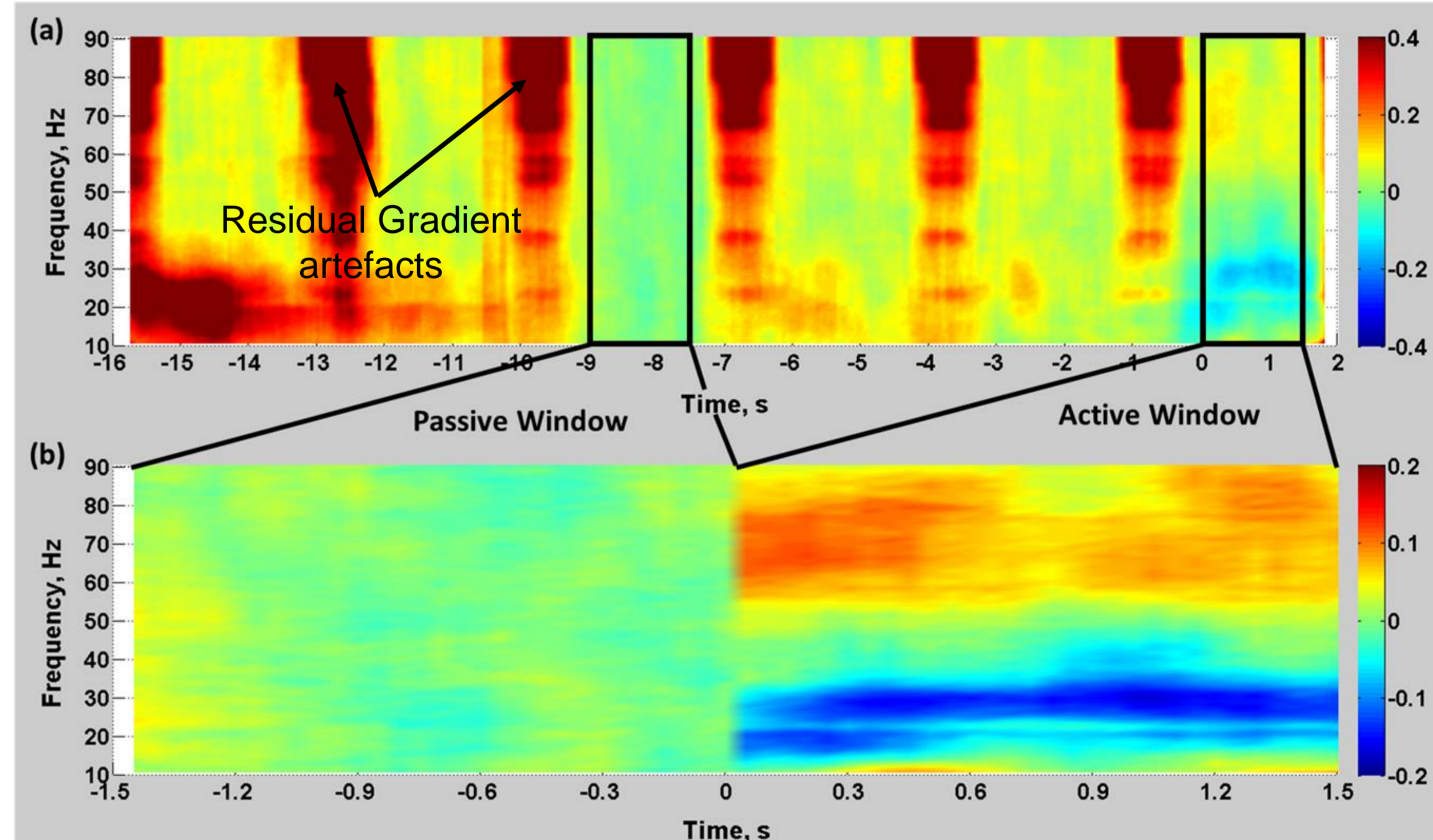
## EEG-fMRI motor study



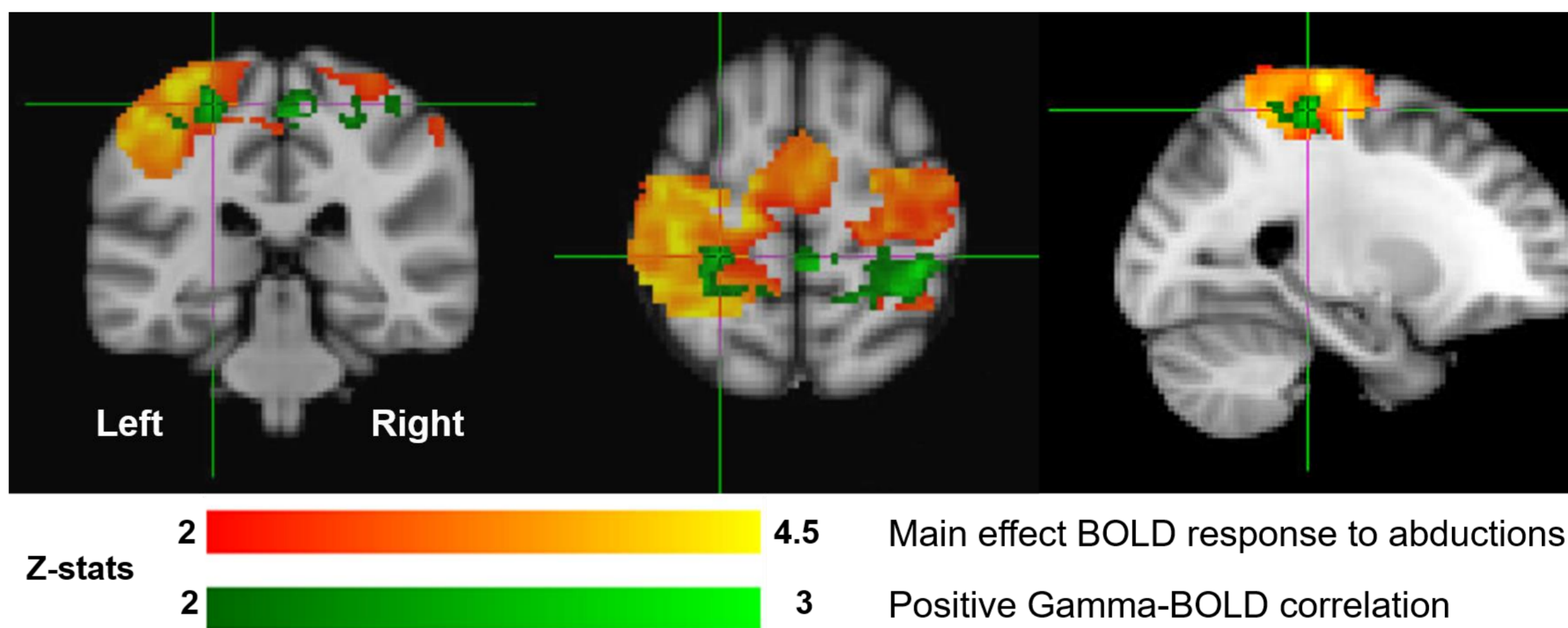
**Figure 3: Group mean beamformer T-stat maps of increase in gamma-power during abductions compared to baseline. Cross hair show mean peak location over subjects.**

- Increase in cM1 (left) gamma power during right-hand finger abductions (Fig 3&4).
- Main effect BOLD activation to the abductions and positive gamma-BOLD correlation in cM1 (Fig 5).
- Correlation was focal to the central sulcus and motor hand-knob, supporting a tight coupling between BOLD fMRI and gamma EEG responses<sup>[3,13]</sup>.

## Results



**Figure 4: Group mean time-frequency spectrograms of cM1 EEG responses: a) 18s whole-trial duration: shows residual gradient artefacts during EPI acquisition; b) Gamma power increases and beta (15-30Hz) power decreases are seen during the active window (0-1.5s) compared with passive window (-9 to -7.5s).**



**Figure 5: Group average fMRI mixed effects results. Main effects and single-trial gamma correlations are cluster corrected with p<0.05, masked to motor cortex.**

## Summary

- Safety:** MB EEG-fMRI acquisition was safe for GE-EPI with MB factor = 4. However, in future work safety testing of the specific MB sequence implementation is still needed.
- Data quality:** MB-fMRI provided “quiet periods” which allowed reliable measurement of EEG gamma and beta responses. The addition of the EEG to MB-fMRI did not significantly degrade MR image quality.
- Neurovascular coupling:** Our results support a tight, positive coupling between BOLD fMRI and gamma EEG responses in motor cortex, similar to previous reports in visual cortex<sup>[14]</sup>
- Our work shows the potential of combining EEG-MB fMRI for advanced study of human brain function**

[1] Fries Annual Review of Neuroscience 32:2009. [2] Logothetis Nature 453:2008. [3] Scheeringa et al. Neuron 69:2011. [4] Mukamel et al. Science 309:2005. [5] Mulert et al. Neuroimage 49:2010. [6] Feinberg et al. PLoS One 5:2010 [7] Moeller et al. Magn Reson Med 63:2010 [8] Auerbach et al. Magn Reson Med 69:2013 [9] Mullinger et al. Int J Psychophysiol 67:2008 [10] Chen et al. Neuroimage 104:2015 [11] Van Veen et al. IEEE Transactions on Biomedical Engineering 44:1997 [12] Oostenveld et al. Comput Intell Neurosci 2011:2011 [13] Logothetis J Neurosci 23:2003 [14] Logothetis et al. Nature 412:2001