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Methodological Issues in EEG-correlated Functional MRI Experiments

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Abstract. The acquisition of functional MRI data in conjunction with the recording of EEG inside the scanner is a new technique that offers the possibility of combining the excellent time resolution of EEG and the spatial resolution of MRI. Three practical considerations are of immediate concern for the acquisition of EEG-correlated functional MR images. These are patient safety, EEG quality and image quality. We review the interactions that arise from the introduction of the EEG recording equipment inside the MR scanner and their consequence for patient safety, EEG quality and image quality. Preventive and corrective measures are then described.

1. Introduction

EEG-correlated functional MRI (EEG/fMRI) is a general term that refers to fMRI experiments with the recording of EEG inside the MR scanner. EEG-triggered fMRI is a specific type of EEG/fMRI experiments in which fMRI acquisitions are triggered after spontaneous EEG events.

EEG/fMRI experiments offer the potential to localize the generators of scalp EEG events using a direct measurement of physiological activity, namely fMRI, as opposed to relying solely on modelling techniques. Several studies have applied EEG-triggered fMRI to epilepsy patients and have shown the potential of this technique to identify focal BOLD-signal increases associated with interictal epileptiform discharges [1,2,3]. Until now the wider application has been limited mainly by problems due to safety concerns and compatibility problems of EEG and MRI. We review the important practical issues of patient safety, EEG quality, and MR image artifacts during EEG/fMRI.

2. Patient Safety Issues

To the medical physicist, introducing wires directly connected to the head into the complex and intense electromagnetic environment of a MR scanner should trigger >alarm signals=. The

potential health hazards in EEG/fMRI, or more generally EEG/MRI, experiments can be classified into two categories: first, the risks associated with the interactions between the patient, the EEG leads and the MRI scanner-s electromagnetic fields; second, the risks associated with the handling of patients and the EEG recording equipment in the MR environment. We will concentrate mainly on the former; see [4] for a more general discussion of safety issues in MRI. While it is possible that EEG recording equipment will become fully integrated into MR scanners in the future, at present EEG/MRI remains an uncommon procedure and therefore it is crucial to ensure that all EEG equipment introduced in the vicinity of the MR scanner should be tested for MR-compatibility [4,5,6] and a specific EEG/MRI patient and equipment handling protocol be devised.

There are three types of electromagnetic fields used in a MR imager that can lead to potential safety hazards: the main static magnetic field, the time-varying magnetic gradient fields, which have different orientations and operate at ~ 1 kHz; and the radio-frequency (RF) pulsed field generated by the headcoil which is homogeneous in the central region of the headcoil and operates at 42.6 MHz/T. MR headcoils are designed to maximise the RF energy which is transmitted to the body via the magnetic field and minimise the electric contribution [8,9]. Nonetheless, the electric component can be a potential health hazard when wires are placed in the vicinity of the RF transmitting coil [10,11].

1.1 Conducting Loops and Other Mechanisms

Low-impedance conduction through the patient represents a potential hazard as currents may be induced in loops placed in the time-varying gradients and RF fields, and due to body movement in the static field. In EEG/MRI, conducting loops will inevitably be present in order to measure electrical potentials on the head. Although these normally have a high impedance (due to the EEG amplifiers) low-impedance loops may form as follows: exposed lead in contact with the patient; two leads in direct electrical contact; a single lead bending on itself and current being able to flow through the insulation at RF frequencies; and failure of the EEG pre-amplifier circuit. Most of the above represent single-fault conditions. However, pre-amplification circuits which have a low-pass filter at the sfront end= could also provide relatively low-impedance in the RF bandwidth. Similarly, a conducting loop could be formed due to the capacitance between parallel leads, with the induced current flowing through the patient. Currents in the leads may cause excessive heating of current-limiting resistors, and must therefore be investigated. Heating due to eddy currents induced within the electrodes and other conducting media in contact with the patient must also be assessed [12,13,14]. Our results for standard electrodes and gel revealed negligible heating, even when using sequences with high RF output [7].

1.2 Specific Health Risks and Relevant Official Safety Guidelines

Whereas currents are invariably induced in the body in the course of normal MR imaging, resulting in a certain amount of heating (which is limited to specific levels under current safety regulations), conducting loops provide a concentration of current in metallic components and therefore a high current density in the adjacent tissue. Referred to as 'contact currents', these represent a well documented health risk [15] and can result in ulcers (~0Hz) [16], electric shock or stimulation (<100 kHz) and heating (>100 kHz) [17,18].

To date, there are no specific official safety guidelines for EEG/MRI. However, relevant general guidelines for physiological monitoring (mainly derived from experience with ECG monitoring inside MR scanners) have been published [4,19,20]. For ECG/MRI, the main risk is that of RF burns at the points of contact between the ECG leads and the patient [19,20]. Therefore, it is recommended to minimize loops, avoid wires from crossing each other and limit currents by using higher-resistance leads. As the amplitude of EEG signal is approximately two orders of magnitude lower than ECG, such leads are likely to give significant degradation of the EEG signal, and therefore guidelines specific to EEG/MRI are required.

Safety guidelines and investigative work which address the same basic physical interactions as described above can be used to establish safety limits: 1) recommendations on maximum contact currents by the National Radiological Protection Board, UK [15]; 2) safety guidelines on the use of electrical equipment from the International Electrotechnical Committee "Medical electrical equipment: specification for general safety requirements" (*IEC601:1*) [21]; 3) published research on RF-burn hazard in animals [22]. In our own study of EEG/MRI safety issues, we have chosen the lowest value across all standards and guidelines [7] (all are RMS): 0.01 mA in the normal operating mode (NOM) and 0.05 mA for the single-fault condition (SFC) for DC; 0.1 mA (NOM) and 0.5 mA (SFC) at low frequency (~1 kHz); and 10 mA (NOM and SFC) for RF.

Heating of objects in sustained contact with the body must not exceed a temperature of 41 °C [22].

1.3 Estimates of Induced Voltages

Table I is a summary of the induced voltages and value of current-limiting resistor required for the switching gradients, RF pulses and loop movement in the static field taken separately. These are obtained from the application of Faraday's induction law [23] and for a GE Signa 1.5 T Echospeed (GE Medical Systems, Milwaukee, USA). The details of the derivation of the formulas and calculation of the voltage and resistor values can be found in [7]. In these calculations, we used the concept of a worst case low impedance loop with an area of 400 cm². For our the relevant parameter values are: $s_{max} = 120 \text{ T m}^{-1} \text{ s}^{-1}$; the length of the headcoil is 0.4 m. The loading function of the body weight H(w) was obtained from our scanner=s manufacturer. Our measurements have shown that K = 1. In order to assess a worst case, for RF we take the example of a fast spin-echo sequence with magnetization transfer contrast, with a SAR_{avg} value of 0.06 W/kg, which is near the allowed maximum.

TABLE I. Formulas for induced EMF's, maximum voltage values for 1.5T Signa EchoSpeed and minimum current-limiting resistor value that ensures safety with regards to current and heating.

Mechanism	Induced EMF	max(V _{RMS} , Signa EchoSpeed 1.5T) (Volt)	min(R _{safe}) (©)
Gradient switching	$ V_{G_Z, \max} = \sqrt{3} s_{\max} A z$	1.66	11.7
RF pulses	$V_{rms} = \omega_0 A K H(w) \sqrt{SAR_{wg}}$	53.0	11.9н10 ³
Body movement m	$ V_{mov} = B_0 \frac{dA}{dt}$	0.2	0.17

A: loop area;

 s_{max} : maximum gradient slew rate;

z: distance from gradient coil isocentre;

 \acute{E}_0 : Larmor frequency;

 B_0 : main field magnitude;

K: geometric gain factor for loop position in the head coil and RF B₁-field inhomogeneity;

H(w): empirical function of the subject=s body weight;

SAR_{avg}: average whole-body Specific Absorption Rate.

The values for the resistor that ensures patient safety must take into account the heating of the resistor that may arise when the maximum allowed current flows through it. Heat dissipation in the current-limiting resistors must be limited such that the temperature increase is less than $20~^{\circ}\text{C}$ and therefore:

$$R > \frac{R_{th}}{20} V_{rms}^2$$

where R_{th} is the thermal resistance (EC/Watt) [7]. In table I, we have used a specific type of resistor (2-Watt carbon-composition resistor, type 101-4, Vitrohm GmbH, Germany), for which R_{th} is 85. Therefore, the minimum value of resistance that ensures that both the maximum allowable current at any given frequency and the resistor temperature increase are not exceeded is 11.9k©. This method has been used in all our EEG/fMRI experiments (approximately 60) and no adverse effects have been observed or reported by the subjects. Furthermore, no visually identifiable difference in background noise was apparent in the EEG, demonstrating that this value of resistance can be used without compromising EEG quality.

Higher Field Strengths

The equations given above can be used to obtain theoretical estimates of the induced voltages for a range of main and gradient field slew rates. However, factors such as RF coil transmission homogeneity and the gradient switching waveform should be considered when trying to implement EEG/fMRI in a different scanner. In certain cases it may be possible to adapt the above equations to the specific parameters of the new scanner, for example in the case of a different gradient switching waveform, whereas in other cases new experimental work may be required. For RF-induced voltages, the general form of the equation in Table I should apply to any scanner, however the exact form of the factors K and H(w) may vary.

2. EEG Quality

Currently, degradation of EEG quality is a limiting factor for many applications of EEG/fMRI. Here we review the basic principles of the generation of the artifacts, review a study of their magnitude and spatial distribution and discuss means of limiting them.

Artifacts in the EEG can be generated as a result of two mechanisms: a change in the magnetic flux through a loop will induce an electromotive force in the loop and movement of blood normal to a magnetic field leads to induced potentials [24]. As a general rule, it is advised to twist electrode leads together from the subject-s head to the amplifier inputs (to minimise inter-electrode lead loops) and to fix the leads securely to a stationary surface (to minimise lead movement) in order to minimize artifacts due to bulk movement of the subject and/or the EEG leads.

2.1 Image Acquisition Artifact

The changing magnetic fields applied during image acquisition may induce emfs in the electrode leads and the subject's head. These emfs can be of large amplitude and are mainly generated by the switching gradients. Their impact depends on the EEG/fMRI experiment paradigm used. For EEG-triggered fMRI, providing the images are acquired in a brief period, this artifact does not present a significant problem [3]. For simultaneous EEG/fMRI, this artifact can be more significant.

No method for removing the image acquisition artifact from EEG recordings has been reported yet. Although low-pass filtering before the amplifier front-end can reduce effectively the contribution due to RF interference and amplifier non-linearity for certain imaging sequences [25], overlap of the EEG and gradient interference spectra suggests that linear filtering alone will not remove this artifact without significant distortion of the EEG waveform. Laudon et al. have reported a method for minimizing this artifact in ECG recordings [25]. The amount of artifact reduction reported is unlikely to improve the usefulness of multi-channel EEG recordings sufficiently. Kreger and Giordano have described an adaptive filtering technique for removing this artifact from ECG signals that

may provide a useful approach for multi-channel EEG [26].

2.2 Pulse Artifact

Movement of the subject's head and electrode leads may alter the area normal to the static field of a loop between electrode leads, giving an induced emf in the leads. Other movements such as swallowing, talking or head turning cause gross artifacts that are amplified inside the scanner (Fig. 1). These may mimic epileptiform activity, although recording from a movement detector can assist in differentiating these [27]. In contrast, small body movements due to pulse and the blood flow effect may produce a pulse artifact which is a more significant problem as it may be widespread on the scalp. This problem has been reported by different centres, using different EEG recording equipment [27,28,29,30].

The amplitude of the pulse artifact for bipolar derivations in 6 subjects was measured in a 1.5 T scanner. We found that the pulse artifact was largest in channels with frontal-polar or fronto-central electrodes and showed large intra- and inter-individual variations in amplitude [31].

Hence, with the exception of large amplitude (>200 $\frac{1}{4}$ V) frontal EEG events or lower amplitude events in the posterior/temporal regions, reliable EEG event detection is difficult in many subjects unless a method for removing the pulse artifact is used. In addition, the amplitude of pulse artifact is directly related to B_0 and hence pulse artifact for scanners with higher B_0 will present a greater problem.

We have developed an on-line pulse artifact removal system that subtracts an on-going average of the artifact waveform for each EEG channel (see figure 1) [31]. This method was evaluated in 6 subjects using spectral analysis and by expert EEG readers, showing a dramatic improvement in EEG quality [31]. We have now used this system in approximately 60 EEG-triggered fMRI studies and adequate quality EEG was obtained in all cases (see figure 2).

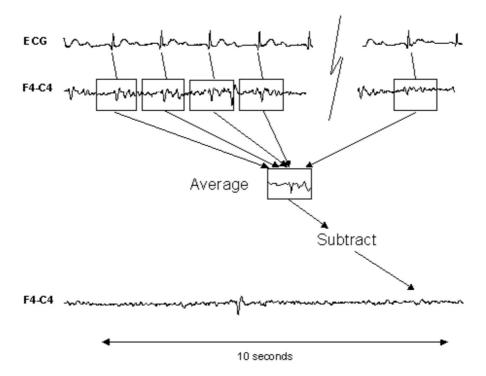


Figure 1. Schematic of the on-line pulse artifact subtraction scheme. A running average of the artifact, calculated within a running window centered on the QRS complexes detected in the ECG signal, is subtracted from the current EEG signal. See [31] for more details.

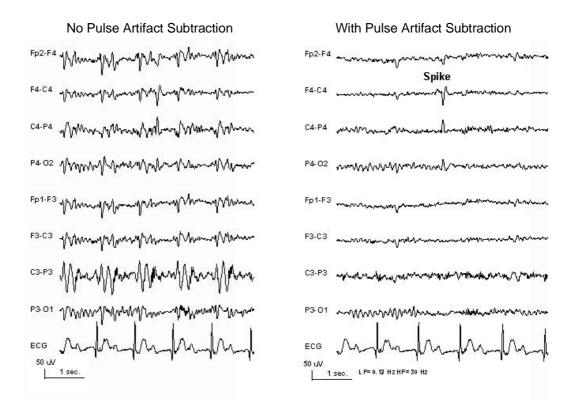


Figure 2. Illustration of the pulse artifact and the result of subtraction. The left-hand side shows simulated EEG traces obtained by adding two segments of EEG: one recorded inside the MRI scanner on a normal subject containing the pulse artifact and one segment containing a spike recorded on a patient. The traces on the right show the result of pulse artifact subtraction.

3. Image Quality

EEG recording inside the MR scanner can degrade MR image quality. The effects of EEG-recording equipment on the MR image quality have been addressed only by a few studies [28,29,32]. Two main mechanisms have been identified whereby EEG recording can compromise MR image quality: first, local signal drop-outs and geometric distortions from EEG electrode assemblies due to their magnetic susceptibility and the possible presence of eddy currents; second, increased noise due to electromagnetic radiation emitted by the EEG recording equipment.

3.1 Local Image Artifacts

In our system, the EEG electrode assembly consists of a standard gold EEG electrode, electrode adhesive, conducting gel and wire, with the addition of a current-limiting resistor soldered between the electrode and wire. Each of those components can cause artifacts due to magnetic susceptibility effects. To quantify these artifacts, we placed different components of EEG electrode assemblies on a spherical phantom and measured the maximum extent of the artifact from the surface of the object. We found that all components cause local artifacts and that there were large differences in the extent of the drop out. In some cases the artifact extended beyond 7.5mm inside the phantom, which is the average depth of the cortex from the scalp [32].

Based on the results of our experiments electrode assemblies comprising of gold electrodes,

cermet film resistors, heat shrink insulation, and copper wire would seem to give acceptable artifact and ease of use. In volunteer experiments, these electrode assemblies led to artifacts clearly smaller than the minimum scalp-cortex distance (Fig. 3).

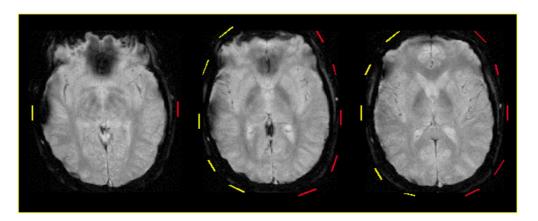


Figure 3. High resolution EPI of a subject with two types of EEG electrode assemblies applied on the scalp. On the right side of the head, non-optimized electrode assemblies (silver/silver chloride electrodes and carbon composition resistor; position shown in yellow) caused artifacts intruding into the cortex. Optimized electrode assemblies (gold electrodes, cermet film resistors; position shown in red) used on equivalent positions on the left side of the head did not compromise the cortical signal.

3.2 Image Noise Due to Electromagnetic Radiation from the EEG Recording Electronics

With regard to electromagnetic noise radiated from EEG equipment we observed significant coherent noise in the images. This is presumably due to broadband RF signals generated by fast switching signals in the EEG digitising circuit, containing a component in the frequency range detected by the receiver chain of the MRI scanner. This noise is likely to vary between specific devices, the location of the device with respect to the headcoil and the characteristics of the RF receive chain of the imager (frequency, bandwidth, etc). By encasing the EEG equipment inside an appropriate shielding box and fitting RF filters to all electrical connections to the equipment, the noise disappeared completely [29,32].

4. Conclusions

We have seen that the introduction of EEG recording equipment into the complex environment of the MR scanner gives rise to a number of interactions between the two systems and the patient. In particular, we have shown that large currents can be induced in conductors in contact with the patient and that these represent a potential health hazard. To this effect we have described a simple modification to the EEG recording system that should ensure patient safety. These interactions can degrade the quality of the EEG and image data by introducing artifacts. Means of avoiding or removing

these artifacts have been described. We have now completed more than 65 EEG/fMRI studies successfully using the safety and corrective measures recommended above.

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