

Whole-Brain Functional and Diffusion Tensor MRI in Human Participants with Metallic Orthodontic Braces

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Conflicts of interest are listed at the end of this article.

See also the editorial by Dietrich in this issue.

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Background: MRI performed with echo-planar imaging (EPI) sequences is sensitive to susceptibility artifacts in the presence of metallic objects, which presents a substantial barrier for performing functional MRI and diffusion tensor imaging (DTI) in patients with metallic orthodontic material and other head implants.

Purpose: To evaluate the ability to reduce susceptibility artifacts in healthy human participants wearing metallic orthodontic braces for two alternative approaches: T2-prepared functional MRI and diffusion-prepared DTI with three-dimensional fast gradient-echo readout.

Materials and Methods: In this prospective study conducted from February to September 2018, T2-prepared functional MRI and diffusion-prepared DTI were performed in healthy human participants. Removable dental braces with bonding trays were used so that MRI could be performed with braces and without braces in the same participants. Results were evaluated in regions with strong (EPI dropout regions for functional MRI and the inferior fronto-occipital fasciculus for DTI) and minimal (motor cortex for functional MRI and the posterior limb of internal capsule for DTI) susceptibility artifacts. Signal-to-noise ratio (SNR), contrast-to-noise ratio for functional MRI, apparent diffusion coefficient and fractional anisotropy for DTI, and degree of distortion (quantified with the Jaccard index, which measures the similarity of geometric shapes) were compared in regions with strong or minimal susceptibility effects between the current standard EPI sequences and the proposed alternatives by using paired *t* test.

Results: Six participants were evaluated (mean age \pm standard deviation, 40 years \pm 6; three women). In brain regions with strong susceptibility effects from the metallic braces, T2-prepared functional MRI showed significantly higher SNR (37.8 ± 2.4 vs 15.5 ± 5.3 ; $P < .001$) and contrast-to-noise ratio (0.83 ± 0.16 vs 0.29 ± 0.10 ; $P < .001$), whereas diffusion-prepared DTI showed higher SNR (5.8 ± 1.5 vs 3.8 ± 0.7 ; $P = .03$) than did conventional EPI methods. Apparent diffusion coefficient and fractional anisotropy were consistent with the literature. Geometric distortion was substantially reduced throughout the brain with the proposed methods (significantly higher Jaccard index, 0.95 ± 0.12 vs 0.81 ± 0.61 ; $P < .001$).

Conclusion: T2-prepared functional MRI and diffusion-prepared diffusion tensor imaging can acquire functional and diffusion MRI, respectively, in healthy human participants wearing metallic dental braces with less susceptibility artifacts and geometric distortion than with conventional echo-planar imaging.

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With MRI, magnetic susceptibility effects can cause significant image artifacts such as signal reduction and image distortion, especially when using echo-planar imaging (EPI). Such susceptibility artifacts are commonly seen in the presence of metallic objects. Dental fillings and orthodontic braces containing various metals can cause large artifacts extending from the facial region

into the brain at EPI (1–3). Orthodontic treatments have become increasingly accessible and are commonly performed in the United States and around the world (see <https://www.aaoinfo.org>). Furthermore, susceptibility artifacts often affect patients undergoing MRI examinations (4,5), especially in regions close to surgical resection cavities, calcified structures, hemorrhages,

Abbreviations

BOLD = blood oxygenation level dependent, DTI = diffusion tensor imaging, EPI = echo-planar imaging, GRE = gradient echo, MPRAGE = magnetization-prepared rapid acquisition gradient-echo, SE = spin echo, SNR = signal-to-noise ratio, 3D = three-dimensional

Summary

T2-prepared blood oxygenation level–dependent functional MRI and diffusion-prepared diffusion tensor imaging with three-dimensional fast gradient-echo readout can significantly reduce susceptibility artifacts commonly seen at conventional echo-planar MRI, especially in the presence of metallic implants.

Key Results

- T2-prepared functional MRI and diffusion-prepared diffusion tensor imaging (DTI) had lower susceptibility artifacts compared with conventional echo-planar MRI and improved signal-to-noise ratio (SNR) in the regions affected by artifacts (functional MRI: 38 vs 16, $P < .001$; DTI: 5.8 vs 3.8, $P = .03$).
- T2-prepared functional MRI and diffusion-prepared DTI showed preserved SNR (functional MRI: 38 vs 37, $P = .38$; DTI: 5.8 vs 5.8, $P = .43$) in brain regions with strong or minimal susceptibility artifacts caused by the metallic braces, whereas the conventional echo-planar imaging methods showed significantly impaired SNR (functional MRI: 16 vs 45, $P = .02$; DTI: 3.8 vs 6.2, $P = .01$) in regions with strong susceptibility effects.
- Geometric distortion was reduced throughout the brain with diffusion-prepared DTI compared with conventional echo-planar imaging methods (measured by significantly increased Jaccard index in diffusion-prepared DTI: 0.95 vs 0.81, $P < .001$).

lesions that contain hemosiderin, and intracranial metallic implants (2,3).

Blood oxygenation level–dependent (BOLD) functional MRI and diffusion tensor imaging (DTI) are two widely used MRI techniques. Currently, EPI is the method of choice for most functional MRI and DTI studies. However, susceptibility artifacts commonly hamper its application. Several other sequences are less sensitive to susceptibility artifacts than EPI (for instance, the three-dimensional [3D] fast gradient-echo [GRE] sequence with short echo time that is commonly used in anatomic examinations such as magnetization-prepared rapid acquisition gradient-echo [MPRAGE]). However, the requirement of a long echo time to generate BOLD and diffusion contrasts during readout precludes the use of such approaches for functional MRI and DTI. Alternatively, however, BOLD and diffusion contrasts can be induced by using spin preparation modules, which separate contrast generation from the readout, thereby opening the possibility to use virtually any sequence for readout. A whole-brain T2-prepared BOLD functional MRI approach (6) was recently demonstrated, which uses a T2-prepared module (7) to induce BOLD contrast, immediately followed by a 3D fast GRE readout with short echo time (< 2 msec). T2-prepared BOLD functional MRI showed little susceptibility artifacts throughout the brain, and greater functional sensitivity than did GRE EPI BOLD functional MRI in regions near air-filled cavities in healthy participants (6) and around the lesions containing blood products in presurgical patients (8,9). Similarly, diffusion

contrast can be generated by using diffusion preparation modules (10,11). Among the various strategies, a double refocusing diffusion preparation module followed by a 3D fast GRE readout has been commonly adopted (11–13). Studies have shown that this technique can significantly reduce image distortion, and thus provide more reliable diffusion measures in regions with strong susceptibility artifacts at EPI (14).

Here, we apply T2-prepared BOLD functional MRI and diffusion-prepared DTI with 3D fast GRE readout in healthy participants wearing metallic orthodontic braces to evaluate the ability of these techniques to minimize susceptibility artifacts in the presence of metallic implants.

Materials and Methods

Written informed consent was obtained from six healthy participants in this institutional review board–approved and Health Insurance Portability and Accountability Act–compliant study. Inclusion criteria was age 18–100 years, both women and men. Standard exclusion criteria for MRI apply (15). Dental braces without magnetically activated components are considered safe for MRI (15).

Figure 1 illustrates the functional MRI and DTI sequences. The T2-prepared BOLD functional MRI sequence consists of a double refocusing T2-preparation module for generating the BOLD contrast, followed by a 3D fast GRE readout. The diffusion-prepared DTI sequence includes a diffusion-preparation module followed by 3D fast GRE. To minimize eddy current–related artifacts and to reduce T1 effects during readout, a stimulated-echo scheme (12,16) was adopted in diffusion-prepared DTI. Removable dental braces (Fig 2) with bonding trays were used so that MRI scans could be acquired with braces and without braces in the same participants. Five examinations were performed in each participant with an Ingenia 3.0-T MRI scanner (Philips Healthcare, Best, the Netherlands): MPRAGE, two-dimensional GRE EPI BOLD functional MRI, 3D T2-prepared BOLD functional MRI (repetition time, 2 sec; voxel size, $3.75 \times 3.75 \times 4$ mm³), two-dimensional spin-echo (SE) EPI DTI, and 3D diffusion-prepared DTI (voxel size, $2.5 \times 2.5 \times 2.5$ mm³). The examination order was chosen to be unique for each participant to avoid potential systematic biases. Optimized higher order shims using the toolbox described in Schär et al (17) and distortion correction (provided by the vendor) were applied in all EPI examinations. To assess BOLD functional MRI changes in the entire brain, a breath-hold task (4) was performed. A respiratory belt was placed around the participant's chest during the examinations. The point spread function was measured for each method (18).

Functional MRI data were analyzed by using statistical parametric mapping (SPM, version 12; Wellcome Trust Center for Neuroimaging, London, United Kingdom; <https://www.fil.ion.ucl.ac.uk/spm/>) and Matlab 2017a (MathWorks, Natick, Mass). Preprocessing steps include realignment, slice timing correction, coregistration, segmentation, and normalization. A general linear model was used to help detect functional

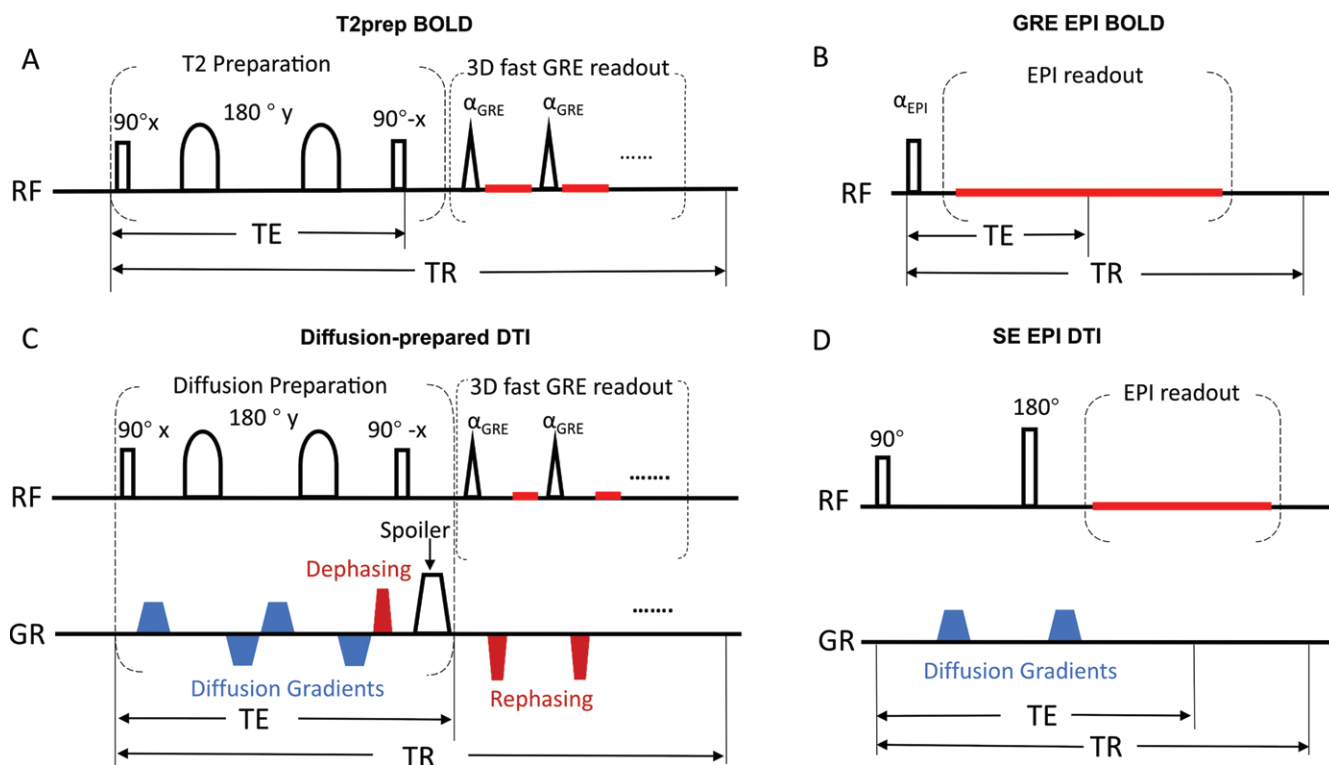


Figure 1: Image shows pulse sequence diagrams of, A, three-dimensional (3D) T2-prepared (T2prep) blood oxygenation level-dependent (BOLD) functional MRI, B, conventional two-dimensional multislice gradient-echo (GR) echo-planar imaging (EPI) BOLD functional MRI, C, 3D diffusion-prepared diffusion tensor imaging (DTI), and, D, conventional 2D multislice spin-echo (SE) EPI DTI. One entire image volume was acquired in a single repetition time (TR) period in all sequences to avoid well-known phase errors in multishot approaches. Details of these pulse sequences are described in Appendix E1 (online). RF = radiofrequency, TE = echo time.



Figure 2: Photograph shows custom-made dental braces that can be easily mounted on and removed from participants' teeth. Brackets made of stainless steel are bonded to a pair of indirect bonding trays made of hard plastic. Beta titanium archwire is seated and secured in bracket slots.

activation (adjusted $P < .05$, cluster size ≥ 3). Motion parameters recorded from the respiratory belt were regressed out. Functional MRI results were compared in two manually delineated regions of interest in each participant: one with strong susceptibility artifacts (dropout) and one covering bilateral motor cortex with minimal susceptibility artifacts at EPI.

DTI data were processed by using MRI Studio (<https://www.mristudio.org>). Signal-to-noise ratio (SNR), apparent diffusion coefficient, and fractional anisotropy were compared in two manually drawn regions of interest: bilateral inferior fronto-occipital fasciculus with strong susceptibility artifacts and bilateral posterior limb of internal capsule with minimal susceptibility artifacts at EPI. Geometric distortion on diffusion tensor images was visualized by using Slicer of FSL (version 6.0; <https://fsl.fmrib.ox.ac.uk/>) and quantified by using the Jaccard index that ranges from zero to one, indicating no overlap to complete agreement, respectively, between the geometric shapes of the diffusion tensor image and reference structural images.

Statistical Analysis

The sample size was determined from power analysis by using the approach described by Cohen et al (19) based on the average effect size (approximately 1.3) reported in previous studies (8) with power of 0.8 and significance set at $\alpha = .05$ (type I error, two tailed).

Paired t tests were performed to compare the results from respective functional MRI and DTI methods. More details on MRI pulse sequences, data analysis, and dental braces are provided in Appendix E1 (online).

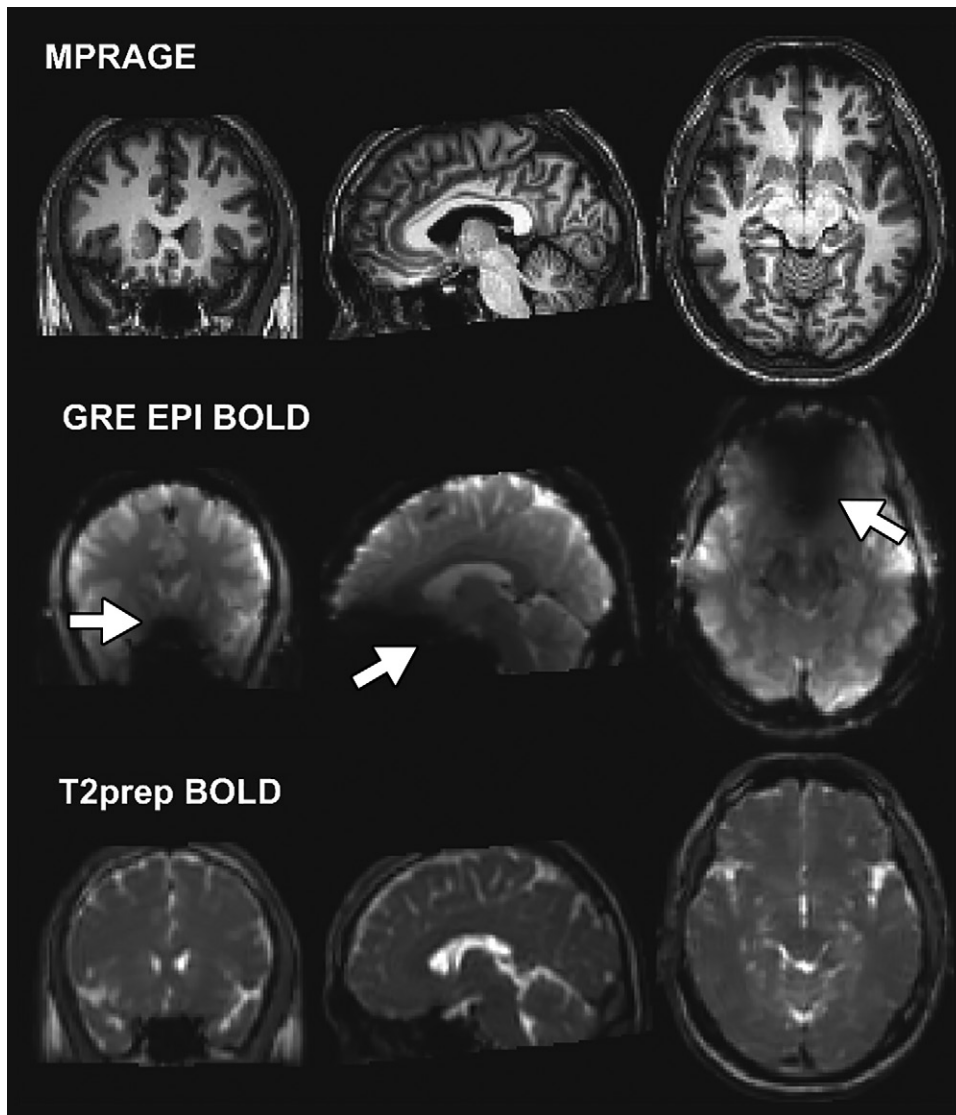


Figure 3: Representative anatomic (magnetization-prepared rapid acquisition gradient-echo [MPRAGE]), gradient-echo (GRE) echo-planar imaging (EPI) blood oxygenation level–dependent (BOLD), and T2-prepared (T2prep) BOLD images acquired in participant wearing titanium dental brace at 3.0 T. Image plane from left to right: coronal, sagittal, and axial. Signal dropouts can be seen on EPI-based images in regions close to brace (arrows). T2-prepared BOLD images were not affected by strong susceptibility effect.

Results

Study Participants

Six healthy participants (mean age \pm standard deviation, 40 years \pm 6; three women) were recruited for this study. Each participant underwent a 60-minute MRI session.

Functional MRI Results

Figure 3 demonstrates representative functional MRIs from one participant wearing braces. Large signal voids are seen on GRE EPI BOLD functional MRI scans, mainly in the orbitofrontal

and ventromedial prefrontal cortex (subsequently defined as the dropout region). Little artifact was visible in the entire brain on MPRAGE and T2-prepared BOLD functional MRI scans. Typical activation maps during breath hold are shown in Figure 4. While activation was detected in most regions with minimal susceptibility artifacts using both methods, fewer activations were detected in the dropout region with GRE EPI BOLD than with T2-prepared BOLD functional MRI.

Quantitative Comparison: T2-prepared BOLD Functional MRI versus GRE EPI BOLD Functional MRI

The group-averaged quantitative results from all participants ($n = 6$) are summarized in Table 1. In the dropout region, temporal SNR, percentage signal change, and contrast-to-noise ratio were all significantly higher with T2-prepared BOLD functional MRI than with GRE EPI. In the motor cortex with minimal susceptibility artifacts, GRE EPI BOLD showed slightly higher (not significant) temporal SNR ($P = .05$), percentage signal change ($P = .56$), and contrast-to-noise ratio ($P = .21$) than did T2-prepared BOLD functional MRI. When the examinations were repeated

without braces, temporal SNR, percentage signal change, and contrast-to-noise ratio in both the dropout and motor regions were slightly higher (not significant; dropout region: $P = .06$, $P = .33$, $P = .06$; motor region: $P = .10$, $P = .91$, $P = .18$) with GRE EPI BOLD functional MRI than with T2-prepared BOLD functional MRI.

Quantitative Comparison: Functional MRI with Braces versus without Braces

Temporal SNR, percentage signal change, and contrast-to-noise ratio in the dropout region were not significantly different with braces and without braces in T2-prepared BOLD

Figure 4: Activation maps overlaid on original functional MRI scans from A, gradient-echo echo-planar imaging (EPI) and B, T2-prepared (T2prep) blood oxygenation level–dependent functional MRI approaches, respectively, in axial imaging plane from healthy participant wearing titanium dental braces during breath-hold task. Activated voxels are highlighted with their corresponding t scores from general linear model analysis with identical statistical threshold. Range of t scores is indicated by scale bar.

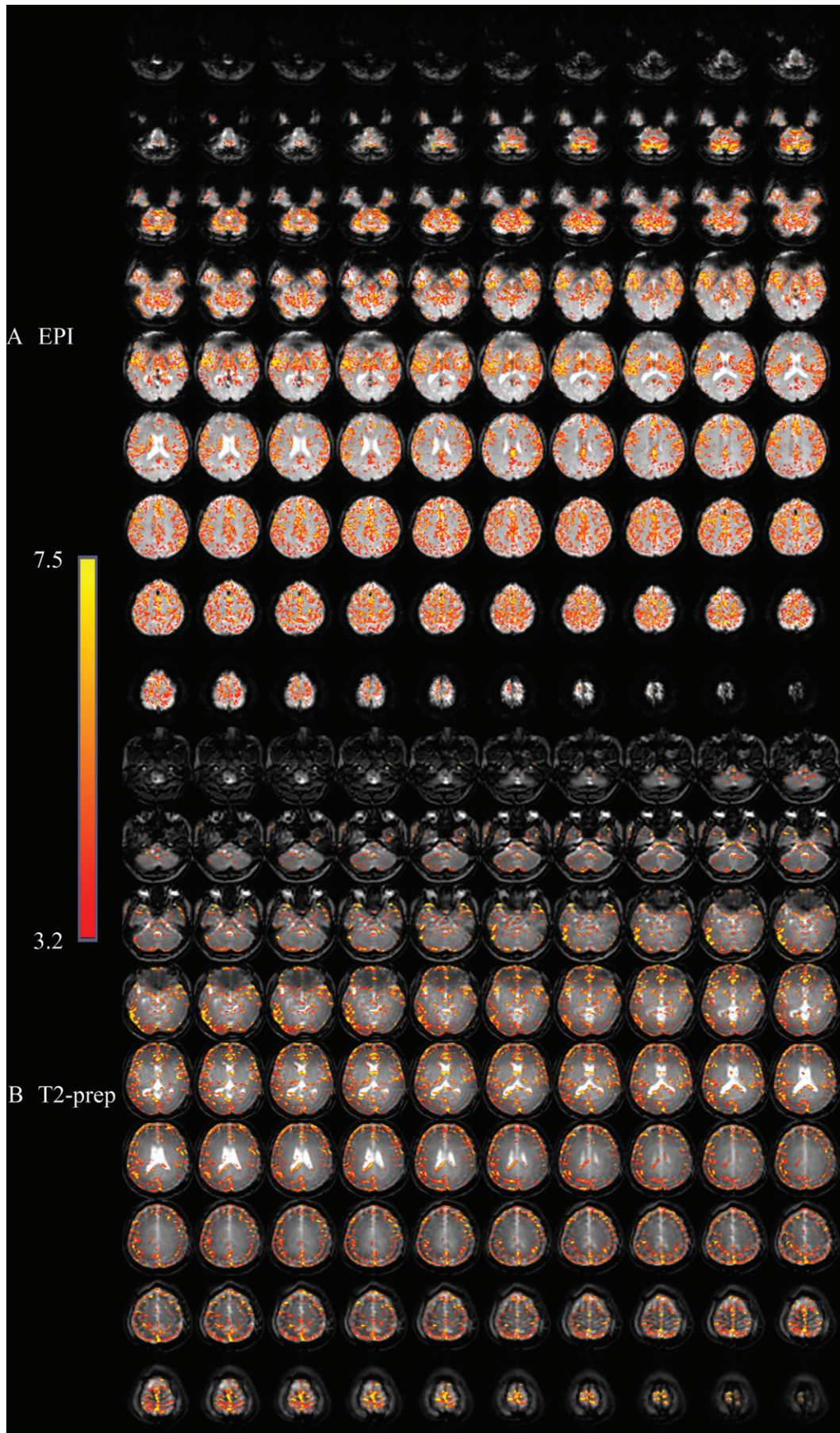


Table 1: Group-averaged Quantitative Results from All Participants (n = 6) for Comparison of the GRE EPI BOLD and T2-prepared BOLD Functional MRI Approaches

Parameter	Temporal Signal-to-Noise Ratio	Percentage Signal Change	Contrast-to-Noise Ratio
With braces			
Dropout region*			
T2-prepared	37.8 ± 2.4	2.3 ± 0.7	0.83 ± 0.16
EPI	15.5 ± 5.3	1.8 ± 0.23	0.29 ± 0.10
P value†	<.001	<.001	<.001
Motor cortex			
T2-prepared	41.6 ± 5.9	2.4 ± 0.7	1.1 ± 0.5
EPI	48.9 ± 7.6	2.6 ± 0.4	1.4 ± 0.4
P value	.05	.56	.21
Without braces			
Dropout region			
T2-prepared	37.0 ± 2.5	2.93 ± 0.54	1.1 ± 0.1
EPI	45.3 ± 6.3	2.83 ± 0.43	1.3 ± 0.2
P value	0.06	0.33	0.06
Motor cortex			
T2-prepared	44.4 ± 2.2	2.8 ± 0.6	1.3 ± 0.6
EPI	52.8 ± 5.8	2.8 ± 0.4	1.5 ± 0.1
P value	0.10	0.91	0.18

Note.—Unless otherwise specified, data are means ± standard deviation. Percentage signal change is relative signal change between breath hold and normal breathing. More details are described in Methods and Appendix E1 (online). BOLD = blood oxygenation level dependent, EPI = echo-planar imaging, GRE = gradient echo.

* Dropout region refers to the area showing large signal wipeout in echo-planar images in participants wearing metallic dental braces, which mainly includes the orbitofrontal and ventromedial prefrontal cortex.

† P < .05.

functional MRI ($P = .38$, $P = .18$, $P = .14$), but were significantly improved with GRE EPI BOLD without braces (Table E1 [online]). The results in the motor cortex in each method were not significantly different with braces and without braces (T2-prepared BOLD functional MRI: $P = .22$, $P = .47$, $P = .17$; GRE EPI BOLD: $P = .20$, $P = .27$, $P = .19$).

DTI Results

Figure 5 demonstrates typical raw diffusion-weighted images, apparent diffusion coefficient, and color-coded fractional anisotropy maps from one participant wearing braces. On diffusion-weighted images and apparent diffusion coefficient maps, no obvious artifact was visible in diffusion-prepared DTI in the entire brain, whereas susceptibility artifacts were substantial in SE EPI DTI. Fractional anisotropy maps color-coded by V1 orientation from SE EPI DTI showed spurious results in the inferior frontal lobe near the braces, affecting visualization of the inferior fronto-occipital fasciculus. Compared with MPRAGE, geometric distortion (Fig 6) was minimal in diffusion-prepared DTI, but was substantial in SE EPI DTI (significantly lower Jaccard index in each slice; $P < .001$). The degree of distortion (Jaccard index) in SE EPI DTI varied with the location of slice. However, diffusion-prepared diffusion tensor images appeared smoother than did SE EPI diffusion tensor images.

Quantitative Comparison: Diffusion-prepared DTI versus SE EPI DTI

Table 2 summarizes the group-averaged quantitative DTI results from all participants ($n = 6$). Apparent diffusion coefficient ($P = .88$) and fractional anisotropy ($P = .33$) were not significantly different between diffusion-prepared DTI and SE EPI DTI, and SNR was slightly higher in SE EPI DTI (not significant; $P = .05$) in the posterior limb of internal capsule with minimal susceptibility artifacts. In the inferior fronto-occipital fasciculus close to the braces, SNR was significantly diminished in SE EPI DTI, leading to suspicious apparent diffusion coefficient and fractional anisotropy, whereas diffusion-prepared DTI showed greater SNR and reasonable apparent diffusion coefficient and fractional anisotropy consistent with the literature (20). When the examinations were repeated without braces, apparent diffusion coefficient, fractional anisotropy, and SNR in both pos-

terior limb of internal capsule ($P = .52$, $P = .11$, $P = .45$) and inferior fronto-occipital fasciculus ($P = .77$, $P = .19$, $P = .27$) became not significantly different between SE EPI DTI and diffusion-prepared DTI.

Quantitative Comparison: DTI with Braces versus DTI without Braces

Apparent diffusion coefficient, fractional anisotropy, and SNR in inferior fronto-occipital fasciculus were not significantly different with braces and without braces in diffusion-prepared DTI ($P = .37$, $P = .35$, $P = .43$), but were significantly improved in SE EPI DTI without braces (Table E2 [online]). The results in posterior limb of internal capsule in each method were not significantly different with braces and without braces (diffusion prepared DTI: $P = .17$, $P = .33$, $P = .28$; SE EPI DTI: $P = .21$, $P = .23$, $P = .31$).

Point Spread Function

The full width at half maximum, or FWHM, of point spread function was not significantly different between 3D T2-prepared BOLD functional MRI (readout × phase-encoding - y × phase-encoding - $z = 7.3$ mm × 7.4 mm × 7.7 mm) and two-dimensional GRE EPI BOLD functional MRI (readout × phase-encoding - $y = 7.3$ mm × 7.6 mm). The FWHM for 3D diffusion-prepared DTI with stimulated echo (5.7 mm × 6.5 mm × 6.7 mm) and two-dimensional SE EPI DTI (5.5 mm ×

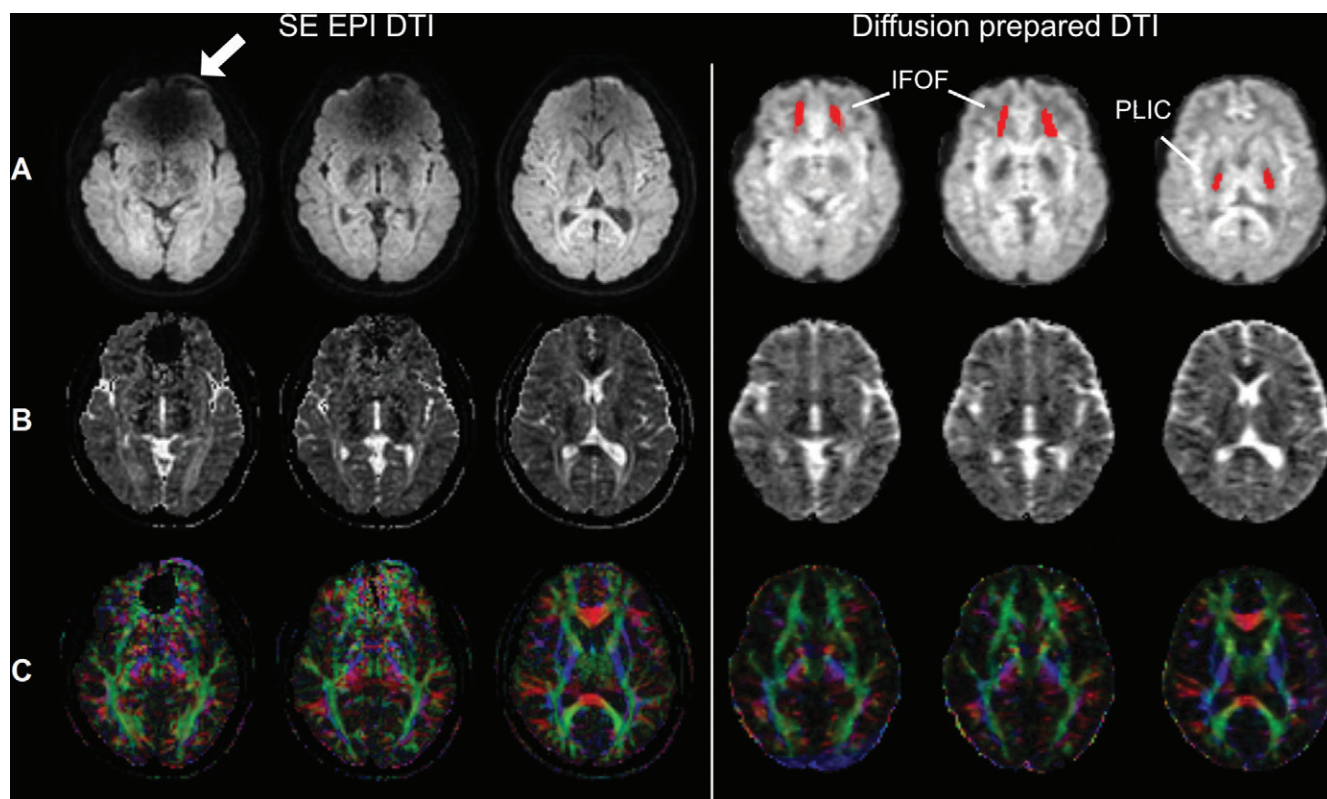


Figure 5: Spin-echo (SE) echo-planar imaging (EPI) and diffusion-prepared diffusion tensor imaging (DTI) axial images acquired at 3.0 T in participant wearing metallic dental braces. A, Raw diffusion-weighted images, B, calculated apparent diffusion coefficient maps, and, C, fractional anisotropy map color coded by V1 (principal eigenvector) orientation (standard red, green, and blue convention). Susceptibility artifacts were observed on SE echo-planar image in regions close to braces (arrow). No obvious artifacts were seen on diffusion-prepared diffusion tensor image. Regions of interest of inferior fronto-occipital fasciculus (IFOF) and posterior limb of internal capsule (PLIC) used in subsequent quantitative analysis are highlighted on diffusion-prepared diffusion tensor images with red.

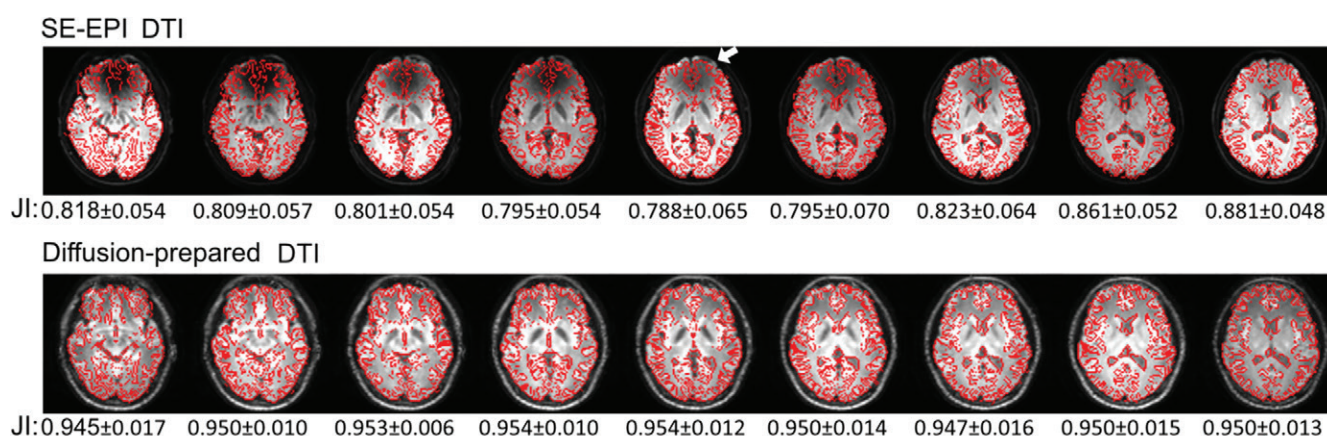


Figure 6: Geometric distortion in axial spin-echo (SE) echo-planar imaging (EPI) (top row) and diffusion-prepared (bottom row) diffusion tensor images when compared with anatomic magnetization-prepared rapid acquisition gradient-echo (MPRAGE) images in same participant wearing metallic dental braces. Edges of brain structures obtained from coregistered MPRAGE images are shown in red contour lines on mean diffusion-weighted images from corresponding diffusion tensor imaging (DTI) approaches. Mismatch between red contour lines and edge of structures shown in diffusion tensor images illustrates geometric distortion artifacts (eg, in frontal area indicated by arrow). Mean Jaccard index (JI) is calculated for each slice and is listed under each image. SE EPI DTI showed significantly lower JI in each corresponding slice ($P < .01$).

5.2 mm) was not significantly different on the readout direction, but it was approximately 25% wider on the first phase-encoding direction in diffusion-prepared DTI. The FWHM for diffusion-prepared DTI without stimulated echo was 5.7 mm \times 5.3 mm \times 5.5 mm, which was narrower on the phase-encoding directions than was diffusion-prepared DTI with stimulated echo.

Higher Spatial Resolution

We used relatively low spatial resolution here to match routine clinical examinations at 3.0 T and to compare the methods. Figure E1 (online) shows the feasibility for the proposed methods to acquire whole-brain images with a voxel size of 1.5 \times 1.5 \times 1.5 mm³ at a field strength of 3.0 T.

Table 2: Group-averaged Quantitative Results from All Participants (n = 6) for Comparison of the Spin-Echo EPI and Diffusion-prepared DTI Approaches

Parameter	Signal-to-Noise Ratio	ADC (10 ⁻³ mm ² /sec)	Fractional Anisotropy
With braces			
Inferior fronto-occipital fasciculus			
Diffusion-prepared	5.8 ± 1.5	0.75 ± 0.08	0.45 ± 0.07
Spin-echo EPI	3.8 ± 0.7	0.16 ± 0.16	0.10 ± 0.12
<i>P</i> value*	.03	<.001	<.001
Posterior limb of the internal capsule			
Diffusion-prepared	5.7 ± 1.1	0.73 ± 0.04	0.61 ± 0.03
Spin-echo EPI	7.0 ± 1.3	0.72 ± 0.03	0.59 ± 0.05
<i>P</i> value	.05	.88	.33
Without braces			
Inferior fronto-occipital fasciculus			
Diffusion-prepared	5.8 ± 1.1	0.79 ± 0.23	0.49 ± 0.02
Spin-echo EPI	6.2 ± 0.5	0.80 ± 0.18	0.45 ± 0.01
<i>P</i> value	.27	.77	.19
Posterior limb of the internal capsule			
Diffusion-prepared	6.2 ± 1.6	0.68 ± 0.27	0.60 ± 0.01
Spin-echo EPI	7.0 ± 1.5	0.69 ± 0.20	0.58 ± 0.01
<i>P</i> value	.45	.52	.11

Note.— Unless otherwise specified, data are means ± standard deviation. More details are described in Methods and Appendix E1 (online). ADC = apparent diffusion coefficient, EPI = echo-planar imaging, DTI = diffusion tensor imaging.

* *P* < .05.

Discussion

Susceptibility artifacts caused by metallic objects at echo-planar imaging (EPI) have been a major limitation for functional MRI and diffusion tensor imaging (DTI) in participants wearing metallic orthodontic material and other head implants. This study was designed to evaluate the ability of two alternative approaches: T2-prepared blood oxygenation level-dependent (BOLD) functional MRI and diffusion-prepared DTI to reduce such susceptibility artifacts in healthy human participants wearing metallic orthodontic braces. T2-prepared BOLD functional MRI and diffusion-prepared DTI showed preserved signal-to-noise ratio (SNR) in the entire brain, whereas conventional EPI approaches showed significantly reduced SNR in regions with strong susceptibility effects (functional MRI: 37.8 ± 2.4 vs 15.5 ± 5.3, *P* < .001; DTI: 5.8 ± 1.5 vs 3.8 ± 0.7, *P* = .03). Because our main goal was to compare the two types of methods in the presence of metallic dental braces, a breath-hold task was chosen as a simple and robust whole-brain functional MRI stimulation. The functional MRI approaches have also been evaluated in other tasks, including language mapping in presurgical patients (6,8).

SE echo-planar images have less signal dropout than do GRE echo-planar images as signal dropouts caused by through-plane susceptibility gradients are refocused. However, dropouts can still occur at SE EPI due to in-plane susceptibility gradients (21) and B1 inhomogeneity, as shown in our SE EPI DTI data with orthodontic braces. Even when complete dropout is avoided, in-plane susceptibility gradients can cause nonlinear

distortion that is difficult to correct for (often needing additional DTI examinations with opposite phase-encoding direction [22]), signal reduction (thus compromising SNR), and erroneous diffusion measures (23). We showed substantially reduced distortion throughout the brain in T2-prepared BOLD functional MRI and diffusion-prepared DTI compared with EPI.

For functional MRI, the T2*-weighted GRE BOLD contrast detected with GRE EPI is more sensitive to the BOLD effect than the T2-weighted SE BOLD contrast measured in T2-prepared BOLD when susceptibility artifacts are minimal. However, when susceptibility artifacts became prominent, the BOLD sensitivity dropped substantially with GRE EPI, but was largely preserved in the entire brain in T2-prepared BOLD functional MRI. Our data showed that compared with GRE EPI, contrast-to-noise ratio in T2-

prepared BOLD functional MRI was 15%–25% lower in the motor cortex (minimal susceptibility artifacts) but was greater than 150% higher in regions affected by the braces. SE EPI BOLD is another commonly used T2-weighted SE BOLD approach. A quantitative comparison between SE EPI BOLD and T2-prepared BOLD functional MRI was performed previously (6). However, unlike GRE EPI BOLD, SE EPI BOLD is rarely used clinically due to its high-power deposition and other limitations (6). We are currently developing a T2*-prepared BOLD functional MRI method.

The stimulated-echo scheme (12,16) used in diffusion-prepared DTI halves signal intensity compared with the same sequence without stimulated echo. Therefore, compared with SE EPI DTI, SNR in diffusion-prepared DTI was 10%–20% lower in regions with minimal susceptibility artifacts, but was 40%–50% higher in regions affected by the braces. Another drawback is that diffusion-prepared diffusion tensor images appear smoother than do SE EPI diffusion tensor images; this is a result of the widened point spread function on phase-encoding direction mainly caused by a longer echo train due to the dephasing and rephrasing gradients needed to generate the stimulated echo. This is supported by our point spread function measurement with and without the stimulated-echo scheme. However, the stimulated-echo scheme is important to suppress eddy-current artifacts and T1-relaxation effects that can cause erroneous diffusion measures (12,16), which we think outweigh the relatively minor loss in SNR and spatial resolution.

There are a wide variety of materials that constitute orthodontic braces (24,25) and other head implants. Therefore, our results

should be extrapolated with caution to implants made with different materials. When the susceptibility gradients exceed the readout bandwidth of 3D fast GRE, significant dropouts will still be present. The readout bandwidth with 3D fast GRE is typically much higher (>500 Hz/pixel) than with EPI (<100 Hz/pixel, phase-encoding direction). For materials with difference in susceptibility (with respect to water) less than 30 ppm (1) (eg, most alloys of precious metals in electrodes for deep brain stimulation or electroencephalography), 3D fast GRE can provide whole-brain largely artifact-free images (1). For materials with difference in susceptibility (with respect to water) less than 1500 ppm (1) (eg, titanium [about 200 ppm], gold [about -25 ppm], nickel [about 125 ppm]), 3D fast GRE typically shows no susceptibility artifact in regions greater than 20 mm from the implants (1–3). The distance between teeth and brain is usually greater than 50 mm in adults. In children or adolescents with orthodontic material, the distance is shorter, but should still be well above 20 mm. However, in regions immediately surrounding such materials, multispectral imaging (26) may be the only existing MRI technique to reduce such near-metal artifacts.

There were limitations to our study. The contrasts induced in the preparation modules decrease during the 3D fast GRE echo train. To minimize this effect, a centric phase-encoding profile was used in 3D fast GRE, in which the center of k -space, which determines the gross signal intensity in MRI scans, is acquired immediately after the preparation modules when the target contrast is strongest. The contrasts for higher spatial frequencies, such as the boundaries between cerebrospinal fluid and tissue, are smaller. Parallel imaging techniques can substantially shorten the 3D fast GRE echo train and thus reduce this effect. We showed that high-spatial-resolution whole-brain images can be acquired with the proposed methods by using advanced parallel imaging techniques. Finally, the results were evaluated in a small number of healthy participants without pathologic findings, which posed another limitation and warrants further investigation.

In summary, our results showed that T2-prepared blood oxygenation level–dependent functional MRI and diffusion-prepared diffusion tensor imaging can be useful alternatives for human brain mapping when echo-planar imaging fails to provide robust signals in the presence of metallic implants such as dental braces.

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