

# Design and Use of Tailored Hard-Pulse Trains for Uniformed Saturation of Myocardium at 3 Tesla

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**Complete and uniform saturation of myocardium is essential for quantitative myocardial perfusion imaging using the first pass of a contrast agent. At 3 T, inhomogeneities of both the static ( $B_0$ ) and radiofrequency ( $B_1$ ) magnetic fields have led to the use of adiabatic  $B_1$ -insensitive rotation type 4 (BIR-4) pulses, which in practice are constrained by radiofrequency (RF) heating. In this study, we propose the use of trains of weighted hard pulses that are optimized for the measured variation of  $B_0$  and  $B_1$  fields in the myocardium. These pulses are simple to design, and require substantially lower RF power when compared with BIR-4 pulses. In volunteers, at 3 T, we demonstrated that the proposed saturation pulse with three subpulses results in lower peak and lower average residual longitudinal magnetization over the heart, as compared with 8-msec BIR-4 pulses and conventional hard pulse trains ( $P < 0.05$ ). Magn Reson Med 60:997–1002, 2008. © 2008 Wiley-Liss, Inc.**

**Key words:** RF saturation pulses;  $B_1$  inhomogeneity; cardiac MRI; myocardial perfusion imaging

First-pass magnetic resonance (MR) myocardial perfusion imaging (MPI) is an established technique for the assessment of ischemic heart disease (1–3). Saturation recovery preparation is widely used to produce  $T_1$ -weighted images rapidly and with multiple slice coverage. Contrast-to-noise ratio (CNR) of myocardial wall enhancement during the first pass is critical to the performance of this technique. Recent work at 3 T has demonstrated improved CNR and showed that the diagnostic performance of 3-T MPI is superior to that of 1.5 T for the identification of both single- and multiple-vessel coronary disease (4). Despite the gain in CNR, there are still major technical challenges in 3-T cardiac imaging, such as static ( $B_0$ ) and radiofrequency ( $B_1$ ) magnetic field inhomogeneities (5–9).

Qualitative and quantitative MPI both rely on complete and uniform saturation of myocardium, and the performance of saturation pulses is sensitive to variations in the  $B_0$  and  $B_1$  fields. Recent studies have compared the effectiveness of different saturation pulses and have shown that the rectangular RF pulse train (10) and the adiabatic  $B_1$ -insensitive rotation type 4 (BIR-4) pulse (11) exhibit better saturation effectiveness than the conventional 90° rectangular hard pulse with 1.5-T and 3-T cardiac imaging (12–15). However, the conventional RF pulse train is more susceptible to  $B_1$  inhomogeneity than the BIR-4 pulse.

Conversely, the BIR-4 pulse has a higher specific absorption rate (SAR). These costs inherently limit their application at 3 T, where low RF power deposition and immunity to  $B_0$ - and  $B_1$ -field inhomogeneities are highly desirable.

In this work, we hypothesize that tailored pulse train designs, based on estimated  $B_0$  and  $B_1$  profiles that were measured a priori over the heart, can overcome both SAR and  $B_1$  inhomogeneity constraints. Compared with the conventional 90° pulse train, the tailored pulse train consists of hard pulses with unequal weighting. The tailored-pulse-train design is considered to improve the immunity to  $B_1$  variation (16), while maintaining low RF power. In this study, the performance of the proposed tailored pulse train is tested using simulations and in vivo experiments at 3 T.

## METHODS

### Experimental Methods

Experiments were performed on a commercial whole-body 3.0-T scanner (Signa Excite HD; GE Healthcare, Waukesha, WI) with gradients capable of 40-mT/m amplitude and 150-T/m/sec slew rate. A body coil was used for RF transmission and an 8-channel phased-array cardiac coil was used for signal reception. Parallel imaging was not used. In all studies, the transmit gain was calibrated using a standard pre-scan and the center frequency was adjusted over a three-dimensional (3D) region of interest containing the left ventricle (LV). Synchronization with the cardiac cycle was achieved with prospective triggering based on an electrocardiogram (ECG) signal. The institutional review board of the University of Southern California approved the imaging protocols. Each subject was screened for magnetic resonance imaging risk factors and provided informed consent in accordance with institutional policy.

### Measurement of $B_0$ and $B_1$ Variation

$B_0$  and  $B_1$  maps were measured in 8 healthy subjects (1 female and 7 males, age  $29 \pm 4.7$  years, height  $177 \pm 5.8$  cm, weight  $70 \pm 7.1$  kg) with 6–8 parallel short-axis slices (6 subjects) and 1 short-axis slice (2 subjects). Cardiac  $B_0$  maps were obtained in a single breath-hold using cardiac-gated gradient-echo sequences with two echo times (TEs). Signal from fat was reduced using a fat-saturation prepulse. Imaging parameters were: field of view (FOV) = 30 cm; in-plane resolution = 2.6 mm, TE = 1.6 msec and 3.6 msec ( $\pm 250$ -Hz frequency range); repetition time (TR) = 12.8 ms; flip angle = 30°; and slice thickness = 5 mm.

Cardiac  $B_1$  maps were acquired in a single breath-hold using the cardiac gated saturated double-angle method (SDAM), as previously described (9,17,18). The transmit

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gain was calibrated for each individual subject. The  $B_1$  scale was computed as the measured flip angle divided by the prescribed flip angle. Imaging parameters were: FOV = 30 cm; in-plane resolution = 5 mm; TE = 2 msec; TR = 7.2 msec; prescribed flip angle = 60° and 120°; and slice thickness = 5 mm.

The LV myocardium was manually segmented in each image, and a composite 2D histogram was produced in  $B_0$ - $B_1$  space. A region of interest (ROI) was defined that contained  $B_0$  and  $B_1$  values representative of nearly all myocardial pixels from all subjects and all imaging slices. This ROI was used for subsequent tailored-pulse-train optimization.

### Optimization of Pulse Trains

Weighted hard-pulse trains ( $\alpha_1, \dots, \alpha_n$ ) of length  $n$  were designed to minimize the maximum residual longitudinal magnetization ( $M_z$ ) over the  $B_0$ - $B_1$  ROI. An exhaustive search based on numerical Bloch simulations was performed with the following cost function:

$$C = \max_{\text{ROI}} \left| \frac{M_z}{M_0} \right|. \quad [1]$$

Each  $\alpha_j$  ranged from 70° to 240° in steps of 1° for  $n \leq 3$ , and 5° for otherwise ( $4 \leq n \leq 6$ ). Minimum and maximum  $\alpha_j$  constraints were determined by 90° divided by the minimum and maximum  $B_1$  scale of the ROI. The peak  $B_1$  amplitude was fixed at 0.115G, a typical value of body-coil transmission in commercial scanners when imaging medium to large-sized humans. We ignored  $T_1$  relaxation between RF subpulses, and assumed the residual transverse magnetization was completely removed by the crusher gradients. Therefore, the subpulses in a set ( $\alpha_1, \dots, \alpha_n$ ) had no specific order, reducing the computation time.

### In Vivo Pulse Performance

In vivo tests of the proposed method were performed in 4 healthy volunteers (1 female and 3 males, age  $31 \pm 2.1$  years, height  $173 \pm 5.3$  cm, weight  $67 \pm 8.1$  kg) with three cardiac views (axial, short-axis, and four-chamber). The tailored pulse train with  $n = 3$  was compared with an 8-ms

BIR-4 pulse with tanh/tan (amplitude/frequency) modulation functions ( $\beta = 5$  and  $\tan(\kappa) = 80$ ) (11), and conventional pulse train using a saturation-no-recovery (SR) experiment, as previously described (12,13). Identical transmit gain and shim values were maintained during the comparison. A 2DFT fast gradient echo (FGRE) acquisition with a center-out  $k$ -space trajectory was used with a 10° flip angle. Proton-density (PD) weighted images with a 3° flip angle were also acquired in the same breath-hold for normalization. The pulse sequence was cardiac-gated and there was one heartbeat between SR and PD acquisition to allow full recovery of magnetization, resulting in a single breath-hold of 5 heartbeats (2 for SR images, 1 for recovery, and 2 for PD images). Each saturation pulse (BIR-4, pulse train, and tailored pulse train) was employed in the FGRE pulse sequence, and acquired in a separate breath-hold. Imaging parameters were: FOV = 30 cm; TE = 1.2 msec; TR = 3.4 msec, acquisition matrix =  $64 \times 64$ ; in-plane resolution = 4.7 mm; slice thickness = 5 mm; image acquisition time = 109 msec; and bandwidth =  $\pm 125$  kHz.

For image analysis, the SR images were normalized by the PD images (SR/PD) to remove the effect of receive coil sensitivity and other system imperfections, and then multiplied by  $\sin 3^\circ / \sin 10^\circ$  to compensate for the excitation angle difference. The resulting images reflect  $|M_z/M_0|$  and have a range from 0 (complete saturation) to 1 (no saturation). LV and right ventricle (RV) were manually defined based on the PD images. We computed the maximum and average  $|M_z/M_0|$  over the LV and LV+RV, and report mean  $\pm$  SD. All image analysis and numerical simulations were performed in MATLAB, version 7.0 (The Mathworks, Inc., Natick, MA).

## RESULTS

### Measurements of $B_0$ and $B_1$ Variation

Fig. 1a (top) illustrates the measured  $B_0$  and  $B_1$  maps from a mid-short-axis slice in one representative volunteer. A Gaussian mixture model (GMM) was initially applied to cluster a region by using the expectation maximization algorithm to estimate the order and parameters of the GMM (19). The 2D histogram of a given  $B_0$ - $B_1$  set and its clustered region (red line) are shown in Fig. 1a (bottom).

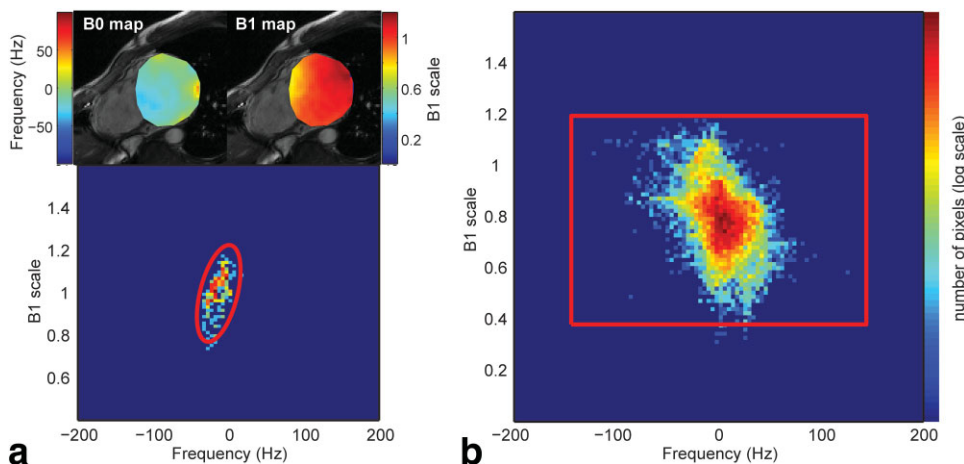


FIG. 1. Measured  $B_0$  and  $B_1$  inhomogeneity over the left ventricle at 3 T: (a)  $B_0$  and  $B_1$  maps from a single slice for 1 subject; and (b) the combined 2D histogram from all slices for all 8 subjects. The extent of the  $B_0$ - $B_1$  ROI (red rectangle) was based on the extent of the  $B_0$ - $B_1$  clusters (red oval) computed using Gaussian mixture models for each slice in each subject. The ROI contains 99.7% of all scattered pixels. [Color figure can be viewed in the online issue, which is available at [www.interscience.wiley.com](http://www.interscience.wiley.com).]

Table 1  
Prescribed Flip Angles, Max  $|M_z/M_0|$ , and Relative RF Power for Optimized Tailored Pulse Trains

n	Optimal weighting					Max $ M_z/M_0 $	Relative RF power <sup>a</sup>	
1	114°					0.737	1.3	
2	116°	231°				0.268	3.9	
3	96°	228°	141°			0.087	5.2	
4	120°	90°	180°	230°		0.043	6.9	
5	90°	110°	145°	205°	235°	0.016	8.7	
6	90°	170°	130°	105°	220°	240°	0.007	10.6

<sup>a</sup>Relative RF energy is defined as the pulse RF energy divided by the RF energy of a single 90° hard pulse (duration 0.5 ms). All pulses use a peak B1 amplitude of 0.115G.

We then chose a rectangular  $B_0$ - $B_1$  ROI to represent the expected  $B_0$ - $B_1$  variation based on the GMM clustered regions, and minimum and maximum  $B_0$ - $B_1$  values. Fig. 1b shows a 2D histogram of all 8 subjects and its corresponding rectangular ROI. The ROI overestimates the inhomogeneities of  $B_0$ - $B_1$  fields to consider all possible cases, and contains 99.7% of all scattered data points. The ROI spans resonance offsets of -144 Hz to +144 Hz, which is in good agreement with the range reported in the literature (5), and a  $B_1$  scaling range of 0.38–1.2, which contains the range obtained in a previous study (0.45–1.12) (9).

### Simulated Pulse Performance

The pulse train weighting was optimized to produce the minimum cost according to Eq. [1]. Table 1 contains the optimal flip angles for  $n = 1$ –6, and the corresponding max  $|M_z/M_0|$  and relative RF energies. The relative RF energy was calculated as the RF energy of a pulse train divided by the RF energy of a single 90° hard pulse (duration 0.5 msec). Fig. 2 plots the residual  $M_z/M_0$  and the RF energy as a function of  $n$  in simulation. The maximum values of  $|M_z/M_0|$  within the ROI for the tailored pulse

train were compared with those for the BIR-4 and pulse train. Because the rectangular ROI overestimates the actual behavior of  $B_0$ - $B_1$  inhomogeneity for each subject, we also computed the average  $|M_z/M_0|$  value and the standard deviation (SD) of  $M_z/M_0$  over the ROI. The tailored-pulse-train designs showed superior saturation performance compared with the conventional pulse train, with only a small increment in the relative RF energy.

When  $n = 3$ , the behavior of the residual  $M_z/M_0$  (max  $|M_z/M_0|$ , mean  $|M_z/M_0|$ , and SD  $M_z/M_0$ ) for the tailored pulse train indicates better saturation performance compared with the BIR-4 pulse, while maintaining lower relative RF energy. Fig. 3a–c shows pulse sequence diagrams for the three different saturation pulses considered in this study. The tailored pulse train ( $n = 3$ ) had a 7.7-msec pulse duration (Fig. 3c). The optimal pulse weighting was 96°, 228°, and 141° (0.548, 1.296, and 0.804 msec, respectively), and the order was chosen among all the six possible combinations by including  $T_1$  relaxation ( $T_1 = 1115$  msec) in the cost function. The spoiler gradients (1 ms) and crusher gradients (3 msec–2 msec–2 msec) were cycled to avoid stimulated echoes, as previously described (10,13,14). The tailored pulse train produced 4.2 times lower relative RF energy than the BIR-4 pulse, and 1.7 times higher relative RF energy than the pulse train.

Fig. 3d–f shows the saturation profiles as a function of off-resonance (horizontal axis) and  $B_1$  scale (vertical axis) using numerical Bloch simulation for the three saturation pulses. The ROI used for optimization is shown in red. The profile for the BIR-4 pulse (Fig. 3d) had excellent saturation for most of the  $B_1$  range, but its  $B_0$  bandwidth became less than  $\pm 110$  Hz as the  $B_1$  scale fell below 0.43. This can be problematic when both the  $B_0$  and  $B_1$  fields are severely distorted. The conventional pulse train (Fig. 3e) showed excellent  $B_0$  variation insensitivity due to its effective bandwidth of  $\pm 1000$  Hz, but the peak  $M_z/M_0$  became larger than 0.1 when the  $B_1$  scale was smaller than 0.7. This explains the weaker  $B_1$  immunity of the pulse train, as previously discussed. The tailored pulse train with the optimal weighting (Fig. 3f), however, improved the saturation performance, and produced excellent saturation effectiveness over the entire ROI.

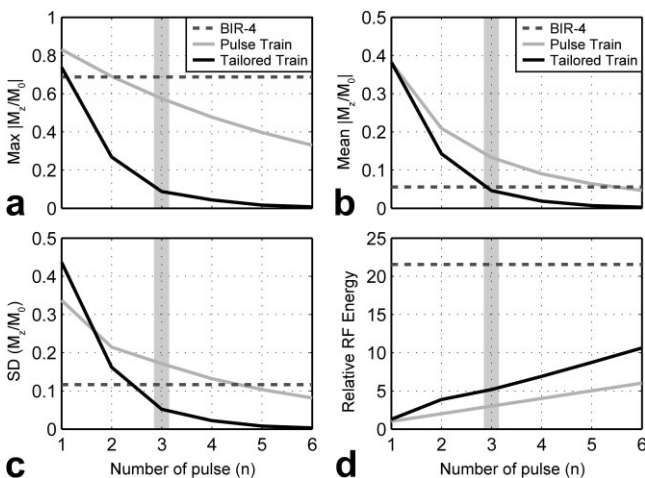


FIG. 2. Behavior of residual  $M_z$  and the corresponding RF energy as a function of  $n$ . (a) Max  $|M_z/M_0|$ , (b) mean  $|M_z/M_0|$ , (c) SD of  $M_z/M_0$ , and (d) relative RF energy were simulated for the BIR-4 pulse (dotted line), conventional pulse train (gray line), and tailored pulse train (black line). The tailored pulse train when  $n \geq 3$  shows superior saturation performance than the BIR-4 and pulse train while maintaining lower relative RF energy than the BIR-4 pulse.

### In Vivo Pulse Performance

Fig. 4 shows representative normalized SR images with the different saturation pulses in three cardiac views. Cardiac cine images (left column) are displayed to provide an anatomical reference. Table 2 summarizes the saturation

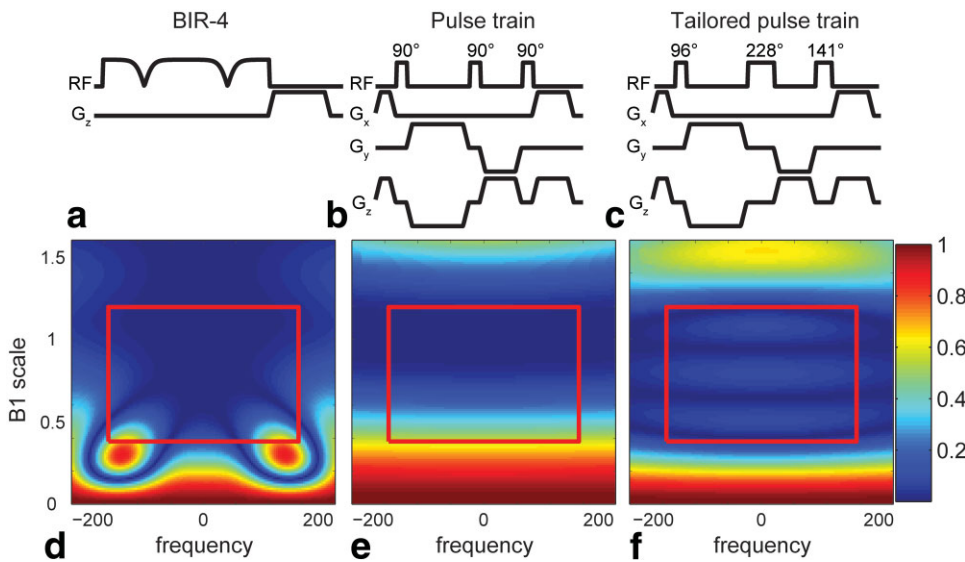


FIG. 3. Saturation pulse sequence diagrams for (a) the BIR-4, (b) conventional pulse train, and (c) tailored pulse train ( $n = 3$ ), and corresponding simulated  $|M_z/M_0|$  profiles (d–f). The  $B_0$ – $B_1$  ROI is indicated by the red box. The simulation profile for the tailored pulse train (f) shows reduced peak  $|M_z/M_0|$  within the ROI. [Color figure can be viewed in the online issue, which is available at [www.interscience.wiley.com](http://www.interscience.wiley.com).]

effectiveness for the different saturation pulses. Both the BIR-4 and tailored pulse showed comparable saturation performance over the LV, whereas the conventional pulse train became less effective in septal myocardium. The peak and average residual  $|M_z/M_0|$  over the LV for the tailored pulse train were smaller than those for the BIR-4 and conventional pulse train, but only the peak value was significantly different ( $P < 0.05$ ). The residual  $|M_z/M_0|$  within the LV+RV behaved differently, and the BIR-4 pulse demonstrated inferior performance compared with the tailored pulse train. The peak and average residual  $|M_z/M_0|$  over the LV+RV for the tailored pulse train were significantly smaller than those for the BIR-4 and conventional pulse train ( $P < 0.05$ ). Note that the measured values were influenced by low SNR and  $T_1$  recovery during the acquisition, and therefore could not represent the absolute saturation effectiveness.

## DISCUSSION

The proposed modification of RF pulse trains was implemented to increase immunity to  $B_0$  and  $B_1$  variation. We demonstrated the train of tailored hard RF pulses to have superior saturation effectiveness in simulation and in vivo, with the cost of 1.7-times-higher RF power than the conventional  $90^\circ$  pulse train. Compared with the BIR-4 pulse, the tailored pulse showed superior saturation over the heart while requiring 4.2 times less RF power. This alleviates the SAR constraints and should enable more frequent use of saturation pulses or more extensive coverage of the myocardium in 3-T MPI (e.g., more slices per heartbeat).

We confined the total number of subpulses to three in this validation study, because three subpulses were the minimum number that was predicted to produce satura-

FIG. 4. Representative normalized SR images with the (a) BIR-4, (b) conventional pulse train, and (c) tailored pulse train in three cardiac views. The proposed saturation pulse demonstrates better saturation performance over the LV and RV than the BIR-4 and pulse train.

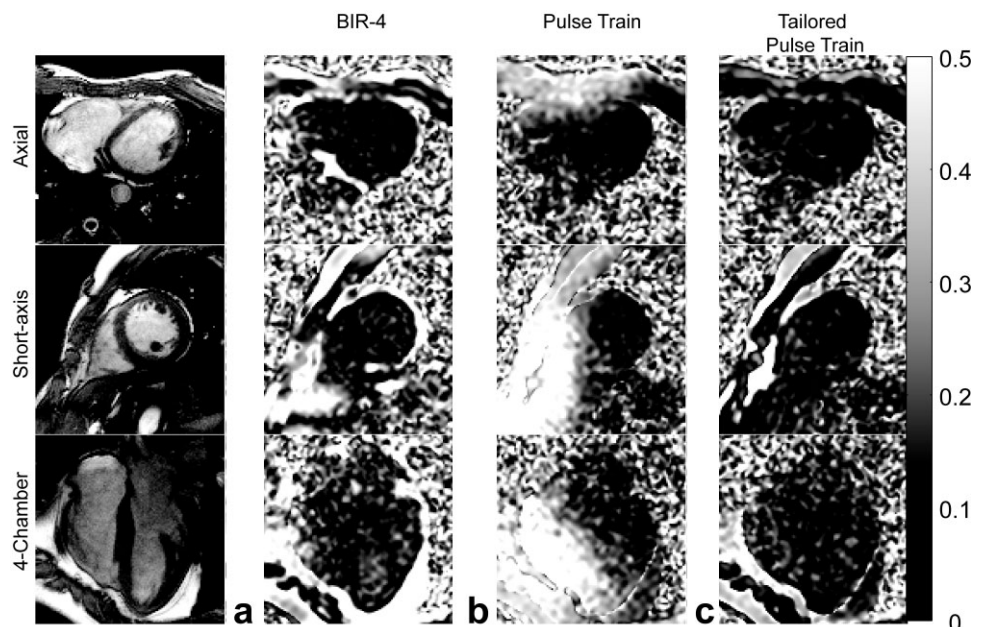


Table 2  
In Vivo Performance of Myocardial Saturation Pulses at 3 Tesla

	RF Pulse duration	Relative RF power	LV		LV and RV	
			Max $ M_z/M_0 $	Mean $ M_z/M_0 $	Max $ M_z/M_0 $	Mean $ M_z/M_0 $
BIR-4	8.0 msec	21.6	$0.28 \pm 0.066$	$0.11 \pm 0.022$	$0.57 \pm 0.259$	$0.14 \pm 0.046$
Conventional pulse train	6.5 msec	3	$0.32 \pm 0.133$	$0.12 \pm 0.055$	$0.81 \pm 0.335$	$0.21 \pm 0.093$
Tailored pulse train	7.7 msec	5.2	$0.22 \pm 0.062$	$0.09 \pm 0.017$	$0.26 \pm 0.086$	$0.097 \pm 0.019$

tion performance superior to both the conventional pulse train and BIR-4. Tailored pulse trains with more than three subpulses may improve the saturation performance (max  $|M_z/M_0|$  and mean  $|M_z/M_0|$ ) with a small increment in RF power and pulse duration, as shown in Table 1. It should be noted that BIR-4 pulses can also be optimized based on  $B_0$ - $B_1$  maps—for example, when choosing amplitude and frequency modulation functions. Fig. 5 shows the mean  $|M_z/M_0|$  as a function of  $\beta$  and  $\tan(\kappa)$  with a fixed RF pulse duration (8 msec). This simulation suggests that an optimized BIR-4 pulse can reduce average residual  $M_z$  by 4.3% while increasing RF power by 8.3%.

Paramagnetic contrast agents during the first pass, and stents, sternal wire, and other metal implants can potentially create additional  $B_0$  inhomogeneity due to increased susceptibility effects (20), widening the  $B_0$ - $B_1$  ROI. The tailored pulse train consists of RF pulses with different

bandwidths, and has an effective  $B_0$  bandwidth of  $\pm 390$  Hz. Therefore, the additional off-resonance caused by the contrast agents is not expected to degrade the saturation performance of the tailored pulse. However, the saturation performance of the BIR-4 pulse, due to its limited bandwidth, can be impacted. The effective bandwidth in the tailored pulse train can be increased by either lowering the maximum  $\alpha_i$  constraint in the cost function or increasing the peak  $B_1$  amplitude in the pulse design.

First-pass contrast agents also cause a reduction in  $T_1$  relaxation time, and longitudinal recovery during saturation may affect pulse performance. Fig. 6 shows the results of a numerical simulation of the residual  $|M_z/M_0|$  within the ROI with different  $T_1$  relaxation times (50–2000 msec). The BIR-4 pulse showed the least  $T_1$  dependency with  $<2\%$  variation for all cases (max  $|M_z/M_0|$ , mean  $|M_z/M_0|$ , and SD  $M_z/M_0$ ), but the tailored pulse train also maintained comparable saturation performance in the short  $T_1$  regime. Based on the robustness to off-resonance and  $T_1$  dependency, we would expect the proposed saturation pulse to maintain its performance under the conditions of dynamic contrast-enhanced MRI.

The  $B_1$  scale range in the ROI (0.38–1.2) overestimates the actual  $B_1$  behavior of each subject because the individual  $B_1$  distribution never spans the entire  $B_1$  scale range. This can be partially attributed to errors in the transmit gain calibration and, specifically, the fact that transmit gain is typically calibrated based on a signal from an entire slice or prescribed volume. Similar to localized shimming and localized center frequency adjustment, the localized calibration of the transmit gain specifically to the heart would be desirable to produce a more accurate and tighter expected  $B_1$  scale range.

The proposed pulse designs can also be applied to other areas of the body such as the abdomen. Tailored pulse trains should be optimized with respect to the  $B_0$ - $B_1$  ROI of each application, which in the abdomen could include fat.

In conclusion, we have described and experimentally validated the use of tailored pulse trains that optimally saturate myocardium at 3 T, based on measurements of typical  $B_0$  and  $B_1$  field variation. Weighted hard pulse trains are simple to design, require substantially lower RF power compared with BIR-4 pulses, and show higher  $B_1$  insensitivity compared with conventional hard pulse trains. Furthermore, the proposed saturation pulse demonstrated lower peak and average residual  $M_z/M_0$  over the heart at 3 T, compared with a standard 8-msec BIR-4 pulse and a conventional hard pulse train ( $P < 0.05$ ). Tailored pulse trains may therefore have an important beneficial role in quantitative first-pass myocardial perfusion imaging at 3 T.

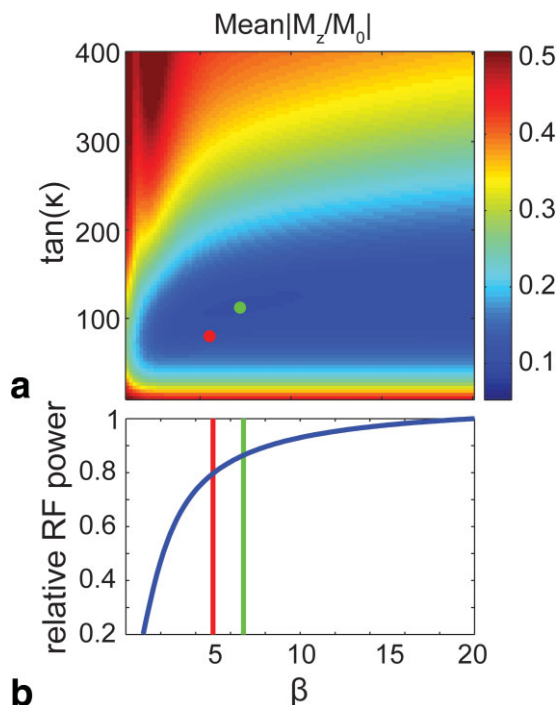


FIG. 5. Optimization of BIR-4 amplitude and frequency modulation function parameters  $\beta$  and  $\tan(\kappa)$ . Simulations show (a) mean  $|M_z/M_0|$  over the ROI as a function of  $\beta$  and  $\tan(\kappa)$ , and (b) relative RF power, which is a function of  $\beta$  only. The red dot (relative RF power = 0.80;  $\beta = 5$ ;  $\tanh(\kappa) = 80$ ) indicates the pulse used in this study, and the green dot (relative RF power = 0.86;  $\beta = 6.8$ ;  $\tanh(\kappa) = 112$ ) indicates the pulse with the lowest residual magnetization. [Color figure can be viewed in the online issue, which is available at [www.interscience.wiley.com](http://www.interscience.wiley.com).]

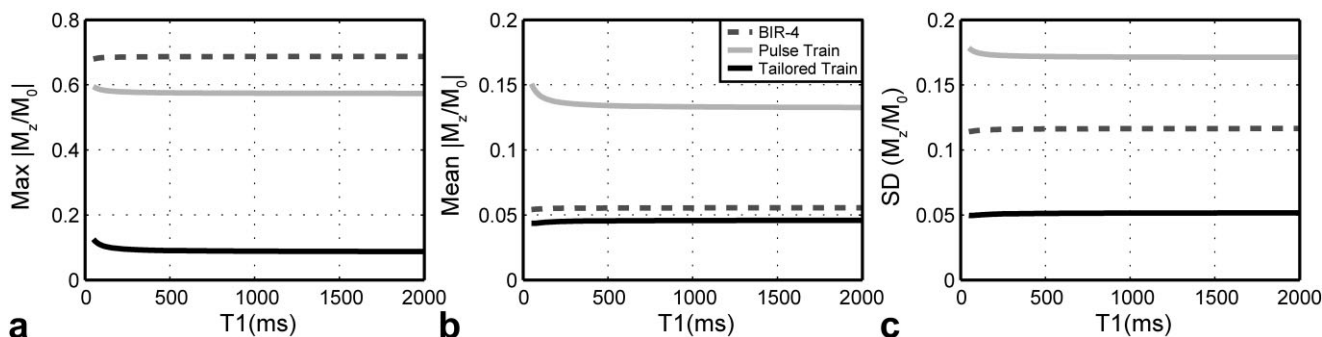


FIG. 6. The saturation effectiveness as a function of  $T_1$ -relaxation times. **(a)**  $\text{Max } |M_z/M_0|$ , **(b)**  $\text{mean } |M_z/M_0|$ , and **(c)**  $\text{SD } M_z/M_0$  were simulated for the BIR-4 pulse (dotted line), pulse train (gray line), and tailored pulse train (black line).

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