## Practical Optimum Experimental Designs for Fast T1 Relaxometry with SPGR Sequences

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Introduction: Knowledge of the longitudinal relaxation time  $T_1$  is necessary in many quantitative MRI applications.  $T_1$  mapping using variable flip angle SPGR acquisitions is an attractive choice due to its speed and accuracy [1]. Critical to the efficiency of the method is the choice of flip angles. A pair of optimized angles, *ideal angles*, may be easily computed for a single  $T_1$  value [1]. However, more angles may be necessary for optimized performance over a range of  $T_1$  values observed in both healthy and pathological tissues and different tissue types. A weighted genetic algorithm (wGA) was used to optimize estimation accuracy for a single  $T_1$  value and yielded a 10 flip angle design with improved performance over a wider  $T_1$  range [2]. In this work, we describe a method for automatic selection of  $T_1$  mapping flip angles, which explicitly optimizes the performance of  $T_1$  mapping for a wide range of  $T_1$  values and demonstrates performance similar to the 10 flip angle wGA design with as few as 3 excitation flip angles, which may allow development of more flexible and efficient  $T_1$  mapping protocols.

**Theory:** The SPGR signal equation is given by  $S = M_0(1-E_1)\sin\alpha/(1-E_1\cos\alpha)$ , where  $M_0$  is equilibrium longitudinal magnetization,  $\alpha$  - excitation flip angle,  $T_R$ repetition time, and  $E_1 = e^{-T_R/T_1}$ . The lowest limit of the variance of  $T_1$  estimate  $\sigma^2(\xi, T_1)$  is equal to  $[\mathbf{M}^{\cdot 1}]_{2,2}$ , where **M** is the Fisher information matrix with

elements  $[\mathbf{M}]_{kl} = \sum_{i=1}^{n} \left( w_i \frac{\partial S(\xi_i, \mathbf{p})}{\partial p_k} \frac{\partial S(\xi_i, \mathbf{p})}{\partial p_l} \right)$  [3]. Here,  $\mathbf{p} = [M_0 T_1]$ , k, l = 1..2, and  $\xi_i$  is a vector of control variables for the  $i^{th}$  measurement ( $\xi_i = \alpha_i$ ). The weight

 $w_i$  indicates the relative accuracy of the  $i^{th}$  measurement that is acquired, and is equal to the number of time averages for a particular measurement. We chose to optimize  $T_1$  mapping efficiency given by  $\Gamma(T_1) = T_1 / (\sigma(\xi, T_1) \sqrt{t})$ , where t is total acquisition time. It is desirable to get maximized and uniform efficiency in the

expected  $T_1$  range given by  $T_1^{(j)}$ , j=1...m. We propose several optimization criteria to fulfill this goal - minimum efficiency  $\left(C_1 = \min\left[\Gamma\left(\xi, T_1^{(j)}\right)\right]\right)$ , algebraic mean

$$\left(C_2 = \sum_{j=1}^m \Gamma\left(\xi, T_1^{(j)}\right)\right), \text{ and geometric mean} \left(C_3 = \left[\sum_{j=1}^m \Gamma^{-1}\left(\xi, T_1^{(j)}\right)\right]^{-1}\right) \text{ in the range.}$$

Methods and Results: In our simulations, the criteria were maximized using simulated annealing optimization to avoid local minima [4]. The flip angles were sought in the range  $[1^{\circ}, 60^{\circ}]$ . The optimization range was [0.1, 5] s. We compared the new methods against the wGA algorithm,  $(FA=(2,3,4,5,7,9,11,14,17,22), T_R=5 \text{ ms})$ , the heuristic algorithm in [5], and the variance minimization approach of [6]. All three criteria improved mean efficiency gain compared to the wGA algorithm using just 3 flip angles (Fig. 1). Criterion  $C_2$  yielded a performance curve similar to the wGA design.  $C_2$  and

 $C_3$  are best at maximizing efficiency, while  $C_1$  and  $C_3$  provide more uniform response in the range (Fig. 1). It is clear

from Table 1 that increasing the number of flip angles from 3 to 10 results in an average gain of only 5-7% in maximum values of the optimization criteria. Furthermore, flip angles of the 10 point design are clustered around 3 values (for Criteria 1 and 2). Hence, we suggest that a 3 flip angle design may be most efficient both in terms of  $T_1$  mapping efficiency and minimum number of required measurements. Several practical 3 flip angle designs for several repetition times are given in Table 2. Figure 2 shows comparison of our method with method of Koay et al. [6].

Discussion: We have presented a method for automated selection of flip angles for optimum performance of SPGR-based  $T_1$  mapping over a wide range of  $T_1$  values. We found that excellent efficiency could be gained with only 3 flip angles. The method outperformed both the 10 flip angle wGA and the 3 flip angle designs of [6] (Figs. 1,2).

Our results are different from those of Koay et al., where flip angles were clustered around only 2 values; our studies showed clustering around 3 values. It is clear from Fig. 2 that narrowing the optimization range improves the performance of Koay's method. Hence, we suggest that Koay's' method may be useful for narrower  $T_1$  ranges. Another difference is that the optimization of Koay's targeted the variance of  $T_1$  estimates, i.e., absolute error. Conversely, we targeted the optimization of relative errors in a  $T_1$  range which focuses on the precision. It is interesting to note that the  $C_2$  -design (2,8,19) was close to the empirical design in [5] obtained combining similar angles from the set of ideal flip angles for 2 representative T<sub>1</sub> values from the range. In general, such a heuristic search may not always be well posed, if ideal flip angles are not clustered. For example, the set of the ideal flip angles is not clustered for TR=20 ms (3,7,17,38). Using an automatic search as proposed here enables an optimized design with flip angles (3,14,38).

The criteria  $C_2$  and  $C_3$  introduced here demonstrated excellent performance for optimum designs. Surprisingly, they are not used as widely as  $C_1$  [7,8]. Particular choice of a design criterion should be guided by application requirements.  $C_2$  and  $C_3$  are best at maximizing efficiency, while  $C_1$  and  $C_3$  provide more uniform response. This optimization framework may be useful for optimization of any quantitative MRI technique.

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References: [1] Deoni SCL, et al. MRM 2003, 49, 515. [2] Deoni S, et al. MRM 2004, 51:194. [3] Atkinson Donev. Optimum Experimental Designs, Oxford, 1992. [4] Press WH et al. Numerical Recipes in C, Cambridge University Press, 1992. [5] Cheng HLM et al. 2006, 55:566. [6] Koay C, et al. ISMRM 2007, 1794. [7] Cercignani M et al. MRM 2006, 56:803 [8] Pineda AR, MRM 2005, 54:625.

	C <sub>1</sub>	C <sub>2</sub>	C <sub>3</sub>
3 FA	FA=(1,7,10)	FA=(2,8,19)	FA=(2,8,12)
	v=7.29	v=5.34	v=1.92
10 FA	FA=(1,6,11)	FA=(2,8,18)	FA=(1,2,8,9,16)
	w=(5,4,1), v=7.74	w=(4,3,3), v=5.41	w=(3,2,4,1,1), v=2.0

**Table 1.** Comparison of several designs for  $T_{\rm R} = 5$  ms (FA – flip angles, **Table 2.** Practical optimized  $T_{\rm 1}$  mapping w-number of repetitions, v-final value of optimization flip angles).



Figure 1.  $T_1$ NR efficiency curves from maximizing several optimization criteria. Mean efficiency gains from proposed designs (3 flip angles) relative to wGA curve (10 flip angles) are given by 1.14 (C1), 1.19 (C2), 1.22 (C3) ( $T_R = 5 \text{ ms}$ )



Figure 2. Comparison of  $T_1$ NR efficiency curves for new design (2,8,19) and method of Koay [4] (KD) (2,10,10) for two  $T_1$  ranges.

TR	C <sub>1</sub>	C <sub>2</sub>	C <sub>3</sub>
10 ms	(2,8,13)	(2,10,28)	(2,10,19)
15 ms	(2,10,16)	(3,13,33)	(3,13,22)
20 ms	(2,12,19)	(3,14,38)	(3,15,26)

experiment designs.