Introduction

$T_1$ maps can be computed from spoiled gradient recalled echo (SPGR) images acquired with different flip angles and/or repetition times (TRs). The function relating signal intensity to flip angle and TR is non-linear; however, a linear form proposed by Gupta in 1977 [1] is currently widely used [1-6]. Using this linearized model, $T_1$ has been estimated with a linear least squares (LLS) method, which has the advantage of being computationally efficient. However, our preliminary study found that the estimated $T_1$ using this LLS method was generally biased and over-estimated [7]. We propose a new weighted linear least squares (WLLS) approach that uses adjusted uncertainties in the fitting. The proposed WLLS method weights each data point with the uncertainty that corrects the noise contribution produced by the transformation of a nonlinear model to a linear one. Numerical and human brain data simulations are used to compare the accuracy of $T_1$ estimated using the LLS, WLLS, and nonlinear least squares (NLS) methods.

Theory

The measured SPGR signal intensity can be written as equation (1), where $\alpha_i$ is the flip angle, $M_0$ is the equilibrium longitudinal magnetization, and $E_i = \exp(-TR_i/T_1)$ [1, 2]. The NLS method estimates $T_i$ and $M_0$ from equation (1) by minimizing the objective function (2). Linear fitting can be used if all images in the dataset are collected with the same TR; equation (1) can be represented in linear form as equation (3). The LLS method estimates $T_i$ and $M_0$ from equation (3) by minimizing the objective function (4), where $y_i = x_i/\sin(\alpha_i)$, $x_i = x_i/\tan(\alpha_i)$, $b = E_i$, and $a = M_0(1-E_i)$. The proposed WLLS method also estimates $T_i$ and $M_0$ from equation (3) by minimizing another objective function (5), where $y_i$, $x_i$, $a$, and $b$ have the same definition as in the LLS method. The weighting function in equation (5) can be derived directly from equation (2) or by using error propagation analysis [8].

Simulations

We performed simulations of different experimental designs, and different expected values of $T_1$. Different signal to noise (SNR), expressed as $M_0/\sigma$, were simulated by adding (in quadrature) Gaussian noise with zero mean and variable standard deviation, $\sigma$, to the noise-free SPGR signals generated using equation (1). Results reported below are computed assuming $TR=10ms$ and $M_0=3000$, optimal flip angles were computed based on the true $T_1$ as suggested in [3]. Six SPGR images, consisting of three replicates of two flip angles without averaging, were used. The accuracy of the estimated $T_1$ was also tested on synthetic data derived from $T_1$ measurements in the human brain. The strategy used to create the synthetic human brain data was similar to that described in [9]. The resultant SPGR brain images have SNR = $M_0/\sigma$ ranging from 90-150 throughout most areas of the brain tissue.

Results

Fig. 1 shows the estimated $T_1$ using the LLS, WLLS, and NLS methods for SNR ranging from 30 to 300. WLLS and NLS produce estimates of $T_1$ with comparable accuracy at all SNR tested, while LLS overestimates $T_1$ progressively as SNR decreases. Fig. 2 shows the relative error of $T_1$ using the LLS, WLLS, and NLS methods for $T_1$ ranging from 600 to 2000 ms with a fixed SNR = 100. The relative error is defined as (true $T_1$ – estimated $T_1$)/true $T_1$. The bias of $T_1$ is corrected in WLLS with the relative error less than 5% regardless of the value of $T_1$. Fig. 3 shows maps of the relative error on the estimated $T_1$ in the synthetic human brain data using the LLS and WLLS methods. The results shown in Fig. 3 (b) and (c) were scaled in the range of ±20%, the gray background corresponds to zero. The relative error of LLS is consistently higher than that of WLLS in brain tissue, and is positive, indicating that $T_1$ is overestimated.

Discussion & Conclusion

In this work we show that the widely-used LLS method improperly weights the uncertainty, resulting in significant errors in $T_1$ estimation. For $T_1$ values ranging from 800 to 1600 ms, which cover most of the $T_1$ values in brain tissue, $T_1$ is overestimated by 10-20%. We propose a weighting approach for the linear model that uses properly weighted uncertainties to adjust the noise contribution produced by the linear transformation. The proposed weighted linear least squares method yields estimated $T_1$ with a precision and accuracy comparable to that obtained from nonlinear fitting while reducing the computation time significantly, enabling the generation of accurate $T_1$ maps “on the fly” at the scanner console.

References: