

## Characterizing the Inherent and Noise-Induced Errors in Actual Flip Angle Imaging

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**Target Audience:** Researchers interested in measuring and characterizing the flip angle term in an image.

**Purpose:**  $T_1$  mapping requires knowing the flip angle,  $\theta$ , or equivalently  $\cos \theta$ ; it can be determined using the TR-interleaved spoiled gradient-recalled echo AFI (actual flip angle imaging) sequence.<sup>1</sup> AFI makes an approximation and uses non-linear processing, two issues that can affect both the accuracy and precision of the calculation. Our objective is to theoretically determine these errors for  $\cos \theta$  (*i.e.*, the estimate of  $\cos \theta$ ) as a function of  $T_1$  and the various AFI acquisition parameters in order to provide an overall error.

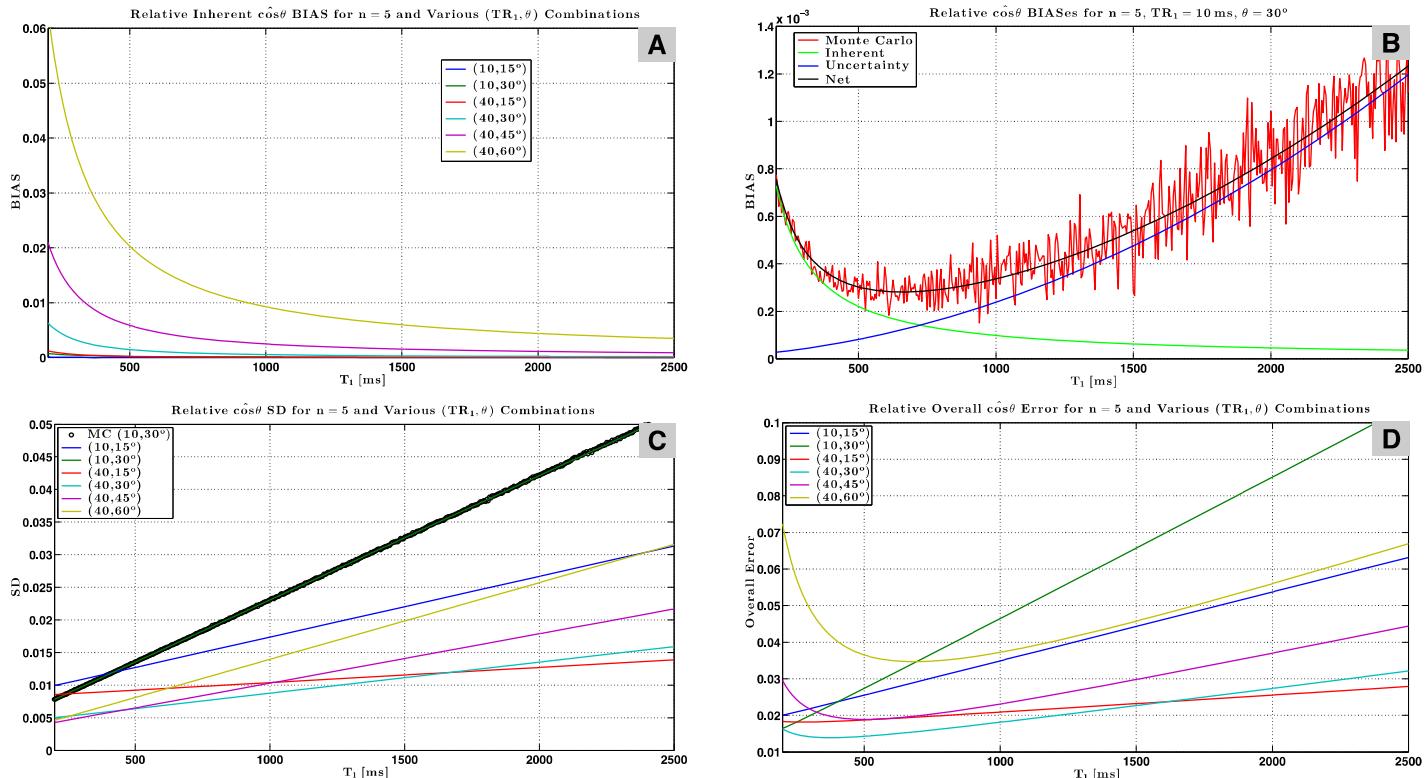
**Methods:** AFI signals  $S_1$  and  $S_2$  are processed to give  $\cos \theta = (Rn - 1)/(n - R)$ , where  $R = S_2/S_1$  and  $n = \text{TR}_2/\text{TR}_1$ . The aforementioned AFI approximation leads to an *inherent* bias, whereas noise produces an *uncertainty* bias; the *net* bias is the sum. Noise also leads to an increase in the variance ( $\sigma^2$ ); the relative bias (a measure of accuracy) of random variable  $z$  is  $\Delta z/z$ , and its relative standard deviation (a measure of precision) is  $\sigma_z/z$ . These can be determined analytically using uncertainty analysis.<sup>2,3</sup> All work was done using Matlab (8.1 R2013a; MathWorks, Natick, MA). The theoretical results were verified numerically (*i.e.*, via Monte Carlo simulation) by generating  $10^5$  instances at each  $T_1$  value, adding normally distributed noise, then calculating the mean and standard deviation (SD) accordingly. The maximum theoretical SNR ( $= M_0/\sigma_S$ ) was set to 1000. The overall error (to roughly 95% confidence) is approximated by the absolute value of the net bias plus two standard deviations.

**Results:** The relative inherent bias of  $\cos \theta$  is  $(Rn - 1)/[\cos \theta (n - R)] - 1$ ; this error decreases as  $T_1$  increases (Figure A). The relative uncertainty bias is given by  $(n^2 - 1)(1 + Rn)/[(n - R)^2(Rn - 1)\text{SNR}_{S_1}^2]$ , which increases with  $T_1$  (Figure B), in contradistinction to the inherent bias. The relative SD of  $\cos \theta$  is given by:  $(n - R)(Rn - 1)/[\text{SNR}_{S_1}(n^2 - 1)(1 + R^2)^{1/2}]$  (Figure C). Relative overall error estimates are provided in Figure D.

**Discussion:** The inherent bias of  $\cos \theta$  decreases as  $\theta$  or  $\text{TR}_1$  decrease, or as  $T_1$  increases since the AFI approximation is  $\text{TR}_{1,2} \ll T_1$ . The net bias is the superposition of the inherent and uncertainty biases, so different  $(n, \text{TR}_1, \theta)$  yield varying proportions. The SD of  $\cos \theta$  is essentially the SD of  $S_1$  since  $n$  and  $R$  are constant with respect to  $T_1$  for a given  $(\text{TR}_1, \text{TR}_2, \theta)$ . The overall error of  $\cos \theta$  shows a significant dependence on  $(\text{TR}_1, \theta)$ , and generally increases with  $T_1$  (contrary to the inherent bias).

**Conclusion:** The analysis herein allows one to tailor the AFI parameters to characterize inherent *vs.* noise-induced errors and minimize overall error. It provides a theoretical and best-case scenario of the error associated with the estimated  $\cos \theta$  maps.

**References:** <sup>1</sup>Yarnykh, Magn Reson Med 2007;57:192-200. <sup>2</sup>Ku, J Res Nat Bur Stand (Eng & Instr) 1966;70C:263-273. <sup>3</sup>Meyer, "Data Analysis for Scientists and Engineers", Wiley Series, New York: John Wiley & Sons, Inc., 1975.



**Figure:** Analytical relative  $\cos \theta$  inherent bias (A), Monte Carlo/inherent/uncertainty/net biases (B), standard deviation (C), and overall error (D) versus  $T_1$  for  $n = 5$  and various combinations of  $(\text{TR}_1, \theta)$ . The y-axis has been capped to 5% in (C) and 10% in (D).