

# Optimal Experiment Design

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## 1 Signal Equation

The signal equation is

$$S_{ij}(M_0, T_1, T_2) = M_0 f_i(T_1, T_2) \frac{1 - E_j(T_1)}{1 - E_j(T_1) f_T(T_1, T_2)}, \quad (1)$$

where  $M_0$  is the initial longitudinal magnetization and  $T_1$  and  $T_2$  are relaxation parameters. The EPG function is

$$f_i(T_1, T_2) = \text{EPG}(\theta_1^i, T_1, T_2, T_s), \quad (2)$$

where  $\theta_1^i$  indicates the first  $i$  refocusing pulses and  $T_s$  is the echo spacing. The recovery function is

$$E_j(T_1) = e^{-\frac{(\text{TR}_j - (T+1) \times T_s)}{T_1}}, \quad (3)$$

where  $\text{TR}_j$  is the  $j^{\text{th}}$  repetition time.

The Fisher Information Matrix (FIM) consists of the following six terms:

$$\mathbf{I} = \begin{bmatrix} \frac{2}{\sigma^2} \sum_{ij} \left( \frac{\partial S_{ij}}{\partial T_1} \frac{\partial S_{ij}}{\partial T_1} \right) & \frac{2}{\sigma^2} \sum_{ij} \left( \frac{\partial S_{ij}}{\partial T_1} \frac{\partial S_{ij}}{\partial T_2} \right) & \frac{2}{\sigma^2} \sum_{ij} \left( \frac{\partial S_{ij}}{\partial T_1} \frac{\partial S_{ij}}{\partial M_0} \right) \\ \cdot & \frac{2}{\sigma^2} \sum_{ij} \left( \frac{\partial S_{ij}}{\partial T_2} \frac{\partial S_{ij}}{\partial T_2} \right) & \frac{2}{\sigma^2} \sum_{ij} \left( \frac{\partial S_{ij}}{\partial T_2} \frac{\partial S_{ij}}{\partial M_0} \right) \\ \cdot & \cdot & \frac{2}{\sigma^2} \sum_{ij} \left( \frac{\partial S_{ij}}{\partial M_0} \frac{\partial S_{ij}}{\partial M_0} \right) \end{bmatrix}. \quad (4)$$

For ease of notation, a function  $f(x, y, z)$  will be referred to as  $f(x)$  when it is understood that the other parameters are constants in the expression. Also, the partial derivative of a function  $f(x, y, z)$  w.r.t. parameter  $x$  will be denoted by  $f'(x)$ . We have

$$S'_{ij}(M_0) = f_i \frac{1 - E_j}{1 - E_j f_T}, \quad (5)$$

$$S'_{ij}(T_1) = M_0 (1 - E_j(T_1)) \frac{(f'_i(T_1) (1 - E_j(T_1)) - f_i(T_1) E'_j(T_1))}{(1 - f_T E_j)^2} \times \frac{(1 - f_T(T_1) E_j(T_1)) + f_i(T_1) (f'_T(T_1) E_j(T_1) + f_T(T_1) E'_j(T_1))}{(1 - f_T E_j)^2}, \quad (6)$$

$$S'_{ij}(T_2) = M_0 (1 - E_j) \frac{(f'_i(T_2) (1 - f_T(T_2) E_j) + f_i(T_2) f'_T(T_2) E_j)}{(1 - f_T E_j)^2}. \quad (7)$$

The inverse of the FIM is a lower bound on the covariance matrix of an unbiased estimator of the parameters  $\boldsymbol{\theta} = (T_1, T_2, M_r)$ . Thus it reasons to minimize some metric of sub-matrix of  $\mathbf{I}^{-1}$ . Group the FIM into

$$\mathbf{I} = \begin{bmatrix} \mathbf{I}_{11} & \mathbf{I}_{12} \\ \mathbf{I}_{21} & \mathbf{I}_{22} \end{bmatrix} \quad (8)$$

where  $\mathbf{I}_{11}$  represents the  $2 \times 2$  sub-matrix. Then the Schur complement of  $\mathbf{I}$  w.r.t.  $\mathbf{I}_{22}$  is

## 2 EPG Function

The EPG function is a recursive function of the signal magnetization and refocusing flip angles. For  $T$  refocusing pulses, the EPG function  $\mathbf{f} : \mathbb{R}_+^2 \rightarrow \mathbb{R}_+^T$  is

$$\mathbf{f}(T_1, T_2, \theta_1^T) = [f_1 \quad \cdots \quad f_T]^T. \quad (9)$$

Let the magnetization state after the  $i^{\text{th}}$  refocusing pulse be  $\mathbf{s}_i \in \mathbb{C}^{3(T+1)}$ . The first entry of  $\mathbf{s}_i$  represents the coherent transverse magnetization,  $M_{xy}$ , and the third entry represents the coherent longitudinal magnetization,  $M_z$ . The observed signal after the  $i^{\text{th}}$  pulse is then

$$f_i(T_1, T_2, \theta_1^i) = [1 \quad 0 \quad 0 \quad \cdots \quad 0] \mathbf{s}_i(T_1, T_2, \theta_1^i) \quad (10)$$

The initial magnetization state (after the 90 degree excitation pulse) is given by

$$\mathbf{s}_0 = \text{vec} \left( \begin{bmatrix} 1 & 0 & 0 & \cdots & 0 \\ 1 & 0 & 0 & \cdots & 0 \\ 0 & 0 & 0 & \cdots & 0 \end{bmatrix} \right). \quad (11)$$

One progression of the EPG states is given by

$$\mathbf{s}_i = \mathbf{E}(\mathbf{G}\mathbf{R}_i\mathbf{G}\mathbf{E}\mathbf{s}_{i-1} + \mathbf{E}_0) + \mathbf{E}_0, \quad (12)$$

where  $\mathbf{E}$ ,  $\mathbf{E}_0$ ,  $\mathbf{G}$ , and  $\mathbf{R}_i$  are defined in the conventional way for EPG. Thus,

$$\begin{aligned} \nabla \mathbf{s}_i(T_1) &= \nabla \mathbf{E}(T_1) (\mathbf{G}\mathbf{R}_i\mathbf{G}\mathbf{E}(T_1)\mathbf{s}_{i-1}(T_1) + \mathbf{E}_0(T_1)) \\ &\quad + \mathbf{E}(T_1) (\mathbf{G}\mathbf{R}_i\mathbf{G} (\nabla \mathbf{E}(T_1)\mathbf{s}_{i-1}(T_1) + \mathbf{E}(T_1)\nabla \mathbf{s}_{i-1}(T_1)) + \nabla \mathbf{E}_0(T_1)) + \nabla \mathbf{E}_0(T_1), \end{aligned} \quad (13)$$

$$\begin{aligned} \nabla \mathbf{s}_i(T_2) &= \nabla \mathbf{E}(T_2)\mathbf{G}\mathbf{R}_i\mathbf{G} (\mathbf{E}(T_2)\mathbf{s}_{i-1}(T_2) + \mathbf{E}_0) \\ &\quad + \mathbf{E}(T_2) (\mathbf{G}\mathbf{R}_i\mathbf{G} (\nabla \mathbf{E}(T_2)\mathbf{s}_{i-1}(T_2) + \mathbf{E}(T_2)\nabla \mathbf{s}_{i-1}(T_2))). \end{aligned} \quad (14)$$