## Fast Regularized Reconstruction Tools for QSM and DSI

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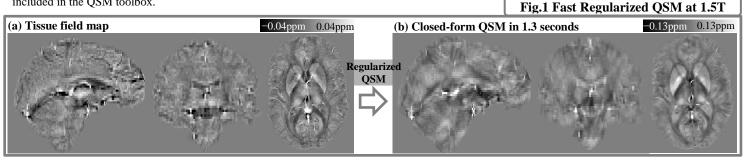
**Software Download:** http://web.mit.edu/berkin/www/software.html

**ℓ**<sub>2</sub>-regularized Reconstruction: admits closed-form solutions that can be computed efficiently. Herein, Matlab tools that achieve dramatic computational speed-up relative to iterative algorithms while retaining good reconstruction performance are presented. Two representative applications of these developed tools, Quantitative Susceptibility Mapping (QSM) and Diffusion Spectrum Imaging (DSI), are demonstrated.

**I. OSM** yields a map of the tissue magnetic susceptibility,  $\chi$ , that lends itself to applications such as estimation of tissue iron concentration and venous oxygenation. The mapping requires the solution of an inverse problem of the form  $\mathbf{F}^H \mathbf{D} \mathbf{F} \chi = \boldsymbol{\phi}$ , where  $\mathbf{F}$  is the Fourier transform,  $\mathbf{D}$  is a diagonal matrix with entries  $1/3 - k_z^2/k^2$ ,  $\chi$  is the unknown susceptibility distribution and  $\boldsymbol{\phi}$  is the measured tissue phase. Since the kernel  $\mathbf{D}$  undersamples the frequency content of  $\chi$  along a cone in k-space, the inversion is facilitated by regularization that imposes prior knowledge [1].

Fast Regularized OSM: involves minimization of  $||\mathbf{F}^H \mathbf{D} \mathbf{F} \chi - \boldsymbol{\phi}||_2^2 + \lambda \cdot ||\mathbf{G} \chi||_2^2$ , where  $\mathbf{G} = [\mathbf{G}_x; \mathbf{G}_y; \mathbf{G}_z]$  is the gradient operator in three dimensions and  $\lambda$  is a regularization parameter. The minimizer  $(\mathbf{F}^H \mathbf{D}^2 \mathbf{F} + \lambda \cdot \mathbf{G}^H \mathbf{G})^{-1} \mathbf{F}^H \mathbf{D} \mathbf{F} \boldsymbol{\phi}$  can be computed efficiently given that the matrix inversion is rapidly performed. The gradient along the x-axis can be expressed as  $\mathbf{G}_x = \mathbf{F}^H \mathbf{E}_x \mathbf{F}$ , where  $\mathbf{E}_x$  is a diagonal matrix with entries  $\mathbf{E}_x(i,i) = 1 - e^{(-2\pi\sqrt{-1}k_x(i,i)/N_x)}$ , which is the k-space representation of the difference operator  $\delta_x - \delta_{x-1}$ . Here,  $k_x$  is the k-space index and  $N_x$  is the matrix size along x, and  $\mathbf{G}_y$  and  $\mathbf{G}_z$  are similarly defined. With this formulation, a closed-form solution  $\widetilde{\chi} = \mathbf{F}^H \mathbf{D}[\mathbf{D}^2 + \lambda \cdot (\mathbf{E}_x^2 + \mathbf{E}_y^2 + \mathbf{E}_z^2)]^{-1} \mathbf{F} \boldsymbol{\phi}$  is obtained. The total cost is two FFTs and multiplication of diagonal matrices.

**QSM Methods and Results:** 3D SPGR data were acquired on a healthy subject at 1.5T with resolution  $0.94 \times 0.94 \times 2.5 \text{mm}^3$  and TR/TE = 58 ms/40 ms. Background phase was removed using dipole fitting [3].  $\lambda = 1.5 \cdot 10^{-2}$  was chosen based on the L-curve heuristic. Fig.1 shows the tissue phase  $\phi$  and the  $\chi$  map reconstructed with the proposed closed-form method. For comparison, iterative conjugate gradient (CG) algorithm was employed to minimize the same objective function (results not shown). After 100 CG iterations, the root-mean-square error (RMSE) between the closed-form and iterative reconstructions was 0.3%. The processing time was 1.3 seconds for the closed-form solution and 29 minutes for the CG algorithm. Both methods are included in the OSM toolbox.



<u>II. DSI</u> involves acquisition of the full q-space samples and yields a complete description of the diffusion probability density function (pdf) for target voxels at the expense of long imaging times (~1 hour). Significant benefit in Compressed Sensing (CS) reconstruction of DSI data from undersampled q-space was demonstrated when a dictionary trained for sparse representation was utilized [4] rather than Wavelet and Total Variation (TV) [5]. However, computation times of these CS methods are on the order of *days* for full-brain processing.

Fast Dictionary-Based DSI: In place of typical  $\ell_1$ -regularized algorithms, two fast and simple  $\ell_2$ -based methods are proposed. These  $\ell_2$ -based methods rely on prior information that is built in to the dictionary and thereby forgoing the need to perform  $\ell_1$  optimization during the reconstruction.

<u>i)Tikhonov regularization:</u> Given a training set of example pdfs, K-SVD algorithm [6] is used to find a dictionary **D** that achieves sparse representation of the training pdfs. For each voxel, the following is solved:  $min_x \|\mathbf{F}_{\Omega}\mathbf{D}x - \mathbf{q}\|_2^2 + \alpha \cdot \|\mathbf{x}\|_2^2$ , where  $\mathbf{F}_{\Omega}$  is the undersampled Fourier transform,  $\mathbf{q}$  denote the undersampled q-space data,  $\mathbf{x}$  are the dictionary transform coefficients, and  $\alpha$  is a regularization parameter.

<u>ii) Principal Component Analysis (PCA)</u>: After subtracting the average pdf  $p_{mean}$  from the training pdf dataset, PCA is applied to produce a matrix of principal pdfs  $\mathbf{Q}$ . A reduced-dimensionality representation is obtained by generating the matrix  $\mathbf{Q}_T$  from the first T columns of  $\mathbf{Q}$ . The target pdf is estimated from undersampled q-space by solving:  $min_{pca} \|\mathbf{F}_{\Omega}\mathbf{Q}_T pca - (\mathbf{q} - \mathbf{F}_{\Omega}p_{mean})\|_2^2$ , where pca are the PCA coefficients.

**DSI Methods and Results:** As the minimizers in the proposed methods can be expressed in closed form, the computational cost is a single matrix-vector multiplication per voxel. Dictionary training is based on data from a subject *different* from the test subject. Optimal regularization parameters were determined using this training dataset. 515 direction, 2.3 mm isotropic DSI data with  $b_{max} = 8000 \text{ s/mm}^2$  were acquired using the 3T Connectom system. Fig.2 depicts reconstruction times and errors in the pdf space at two undersampling factors, R=3 and 9. The proposed methods demonstrate *1000-fold* speed-up relative to the previous dictionary-based algorithm in [4] with comparable reconstruction quality, while substantially reducing the reconstruction error relative to the Wavelet+TV method in [5]. Both of the proposed methods are included in the DSI toolbox.

