#### **FMRI Statistical Brain Activation From k-Space Data**

Daniel B. Rowe, Ph.D. dbrowe@mcw.edu

Department of Biophysics Division of Biostatistics Graduate School of Biomedical Sciences



#### **MR** Physics

**Bloch Equation** 

$$\frac{d\mathbf{M}}{dt} = \gamma \mathbf{M} \times \mathbf{B} + \frac{1}{T_1} (\mathbf{M}_0 - \mathbf{M}_z) - \frac{1}{T_2} (\mathbf{M}_x + \mathbf{M}_y)$$

Solving the Bloch equation yields transverse magnetization

$$\begin{split} M_{xy}(x,y,z,t) = &M_{xy0}(x,y,z)e^{-t/T_2}e^{-i\gamma B_0 t}e^{-i\gamma \int [G_x(t)x + G_y(t)y + G_z(t)z]dt} \\ \text{By Faraday's law of induction, the signal at time } t \text{ is} \\ s(t) = &\int_x \int_y M(x,y)e^{-i2\pi k_x(t)x}e^{-i2\pi k_y(t)y} \ dx \ dy \end{split}$$

Take the IFT of the spatial frequencies to get our magnetzation image.

$$\hat{M}(x,y) = \sum_{k_x} \sum_{k_y} \hat{s}(k_x,k_y) e^{i2\pi k_x x} e^{i2\pi k_y y}$$

The magnetzation is related to our PSD and is complex valued.

Rowe, MCW

#### **Pulse Sequence**

How do we get signal that is spatial frequencies?

We apply  $G_x \& G_y$  magnetic field gradients to encode then we measure the complex-valued DFT of the object.



(a) Gradient Echo-EPI Pulse Sequence

(b) Corresponding *k*-Space Trajectory

Kumar, Welti and Ernst: NMR Fourier Zeugmatography, J. Magn. Reson. 1975

Haacke et al.: Magnetic Resonance Imaging: Physical Principles and Sequence Design, 1999.

#### **Complex Image Reconstruction I**

### $S(k_x, k_y) = S_R(k_x, k_y) + iS_I(k_x, k_y)$ , the complex-valued DFT of object



(a) real:  $96 \times 96$ 

(b) imaginary:  $96 \times 96$ 

FOV=192 mm, mat=96 $\times$ 96, vox=2 mm<sup>3</sup>

#### **Complex Image Reconstruction I**

## complex-valued 2D IFT $(\Omega_{yR} + i\Omega_{yI}) * (S_R + iS_I) * (\Omega_{xR} + i\Omega_{xI})^T = (M_R + iM_I)$



FOV=192 mm, mat=96imes96, vox=2 mm $^3$ 

#### **Complex Image Reconstruction I**

Due to the imperfect Fourier encoding, the IFT reconstructed object is complex-valued,  $M(x, y) = M_R(x, y) + iM_I(x, y)$ .



(a) Real image,  $y_R$ 

(b) Imaginary image,  $y_I$ 

FOV=192 mm, mat=96 $\times$ 96, vox=2 mm<sup>3</sup>

This occurs over time in fMRI and results in complex-valued images and voxel time course observations,  $y_t = y_{Rt} + iy_{It}$ .



Most fMRI studies transform from real-imaginary rectangular coordinates to magnitude-phase polar coordinates,  $M(x, y) = R(x, y)e^{i\Phi(x, y)}$ .



(a) Magnitude,  $r_t = \sqrt{y_{Rt}^2 + y_{It}^2}$ 

(b) Phase,  $\phi_t = \operatorname{atan}_4(y_{It}/y_{Rt})$ 

# Collect a sequence of these reconstructed images over time. Form voxel time courses, $y_t = r_t e^{i\phi_t}$ .



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Time series are complex-valued or bivariate with phase coupled means.



The  $y_R$  and  $y_I$  time courses have related vector length info! This is a time series from actual human experimental data!

Time series are complex-valued or bivariate with phase coupled means.



Real-Imaginary or Magnitude-Phase time courses have all info! Recent work indicates that phase time courses may exhibit TRPCs Menon, 2002; Hoogenrad et al., 1998; Borduka et al., 1999; Chow et al., 2006;

#### **Complex Time Series Activation**

Block-designed experiment: Off-On-Off-...-On-Off task



Complex Magnitude w/ Constant Phase (CP) Activation<sup>1,2</sup>
Complex Magnitude &/or Phase (CM) Activation<sup>3</sup>
Real Magnitude-Only (MO/UP) Activation (Discard Phase)<sup>4,5</sup>
Real Phase-Only (PO) Activation (Discard Magnitude)<sup>6</sup>

<sup>1</sup>Rowe and Logan: NeuroImage, 23:1078-1092, 2004.
<sup>3</sup>Rowe: NeuroImage, 25:1310-1324, 2005b.
<sup>5</sup>Rowe and Logan: NeuroImage, 24:603-606, 2005.

<sup>2</sup>Rowe: NeuroImage 25:1124-1132, 2005a.
<sup>4</sup>Bandettini et al.: Magn Reson Med, 30:161-173, 1993.
<sup>6</sup>Rowe, Meller, Hoffmann: J Neurosci Meth, 161:331-341, 2007.

#### **Complex Time Series Activation**

$$\begin{pmatrix} y_{Rt} \\ y_{It} \end{pmatrix} = \begin{pmatrix} \rho_t \cos \theta_t \\ \rho_t \sin \theta_t \end{pmatrix} + \begin{pmatrix} \eta_{Rt} \\ \eta_{It} \end{pmatrix}, \quad \begin{pmatrix} \eta_{Rt} \\ \eta_{It} \end{pmatrix} \sim N(0, \Sigma) .$$

 $y_{Rt}$  and  $y_{It}$  are the real and imaginary observations at time t  $\eta_{Rt}$  and  $\eta_{It}$  are the real and imaginary noise terms at time t

$$> Magnitude activation in complex data 
 $\rho_t = x'_t \beta = \beta_0 + \beta_1 x_{1t} + \dots + \beta_{q_1} x_{q_1t}$ 
I: Magnitude-only<sup>4,5</sup>  $\theta_t = \theta_{t'}$ 
II: Constant Phase<sup>1,2</sup>  $\theta_t = \theta$ 
III: Linear Phase<sup>3</sup>  $\theta_t = \gamma_0 + \gamma_1 u_{1t} + \dots + \gamma_{q_2} u_{q_2t}$$$

> Phase-only Activation in complex data,  $\rho_t = \rho_{t'}$  $\theta_t = \gamma_0 + \gamma_1 u_{1t} + \dots + \gamma_{q_2} u_{q_2t}$ 

<sup>1</sup>Rowe and Logan: NeuroImage, 23:1078-1092, 2004.
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#### **Complex Image Reconstruction II**

## complex-valued 2D F FT $(\bar{\Omega}_{yR} + i\bar{\Omega}_{yI}) * (M_R + iM_I) * (\bar{\Omega}_{xR} + i\bar{\Omega}_{xI})^T = (S_R + iS_I)$



#### **Complex Image Reconstruction II**

# $\begin{array}{rll} \text{complex-valued 2D I FT} \\ (\Omega_{yR}+i\Omega_{yI}) & \ast & (S_R+iS_I) & \ast & (\Omega_{xR}+i\Omega_{xI})^T = & (M_R+iM_I) \end{array}$



Stack rows of reals then imaginaries for s.  $\Omega$  from  $\Omega_{xR}$ ,  $\Omega_{xI}$ ,  $\Omega_{yR}$ ,  $\Omega_{yI}$ .

#### **Complex Image Reconstruction II**



Unstack m to real then imaginary rows to form image.

Rowe, Nencka, Hoffmann: JNMeth, 2007.





Now convert the m that has Real/Imaginaries stacked for images to y that has the Real/Imaginaries stacked for voxels

$$y = P * \qquad m$$
$$y = P * \qquad (I_n \otimes \Omega)s$$
$$\begin{pmatrix} y_1 \\ \vdots \\ y_p \end{pmatrix} = P * \begin{pmatrix} \Omega & 0 \\ & \ddots \\ 0 & \Omega \end{pmatrix} \begin{pmatrix} s_1 \\ \vdots \\ s_n \end{pmatrix}$$

## Amazingly can convert from m to y via a permutation matrix P. y = P



But of course the y's are obs in each voxel for complex activation



Estimate  $\beta$ 's, $\theta$ 's, and  $\sigma^2$ 's from my complex models Compute activation from my complex activation models.

 $\begin{array}{ll} \rho_t = \beta_0 + \beta_1 x_{1t} + \dots + \beta_{q_1} x_{q_1t} \\ \text{I: Constant Phase} & A_j = I_n \cos \theta_j \text{ and } B_j = I_n \sin \theta_j \\ \text{II: Unrestricted Phase} & A_{jt} = \cos \theta_{jt} \text{ and } B_{jt} = \sin \theta_{jt} \\ \text{III: Linear Phase} & \theta_{jt} = \gamma_{0j} + \gamma_{1j}t + \dots + \gamma_{q_2j}u_{q_2t} \\ i \text{ is versel} \quad i = 1 \dots n \end{array}$ 

j is voxel, j = 1, ..., pt is time, t = 1, ..., n

#### **Activations in Human Experimental Data**

Imaging Parameters: 1.5T GE Signa 5 axial slices of 128x128 96 acq.-2.0833mm<sup>2</sup> 128 recon.-1.5625mm<sup>2</sup> FOV =20cm TR=1000ms TE=47ms FA=90°

Task: RH male Bilateral sequential finger tapping light triggered Block design 16 off  $+ 8 \times (160n+160ff)$ ;

Rowe and Logan: NeuroImage, 23:1078-1092, 2004.

#### z Maps: 5% Bonferroni Threshold



#### (a) UP/MO

#### (b) CP

#### (c) PO



Can also do directly from k-space data

$$\begin{pmatrix} s_1 \\ \vdots \\ s_n \end{pmatrix} = P^{-1} (I_n \otimes \Omega^{-1}) \begin{pmatrix} X\beta_1 \cos \theta_1 \\ X\beta_1 \sin \theta_1 \\ \vdots \\ X\beta_p \cos \theta_p \\ X\beta_p \sin \theta_p \end{pmatrix} + \begin{pmatrix} \epsilon_1 \\ \vdots \\ \epsilon_n \end{pmatrix}$$
$$\begin{pmatrix} s_1 \\ \vdots \\ s_n \end{pmatrix} = \underbrace{P^{-1} \left[ I_n \otimes \Omega^{-1} (I_2 \otimes X) \right]}_{\text{Known}} \begin{pmatrix} \beta_1 \cos \theta_1 \\ \beta_1 \sin \theta_1 \\ \vdots \\ \beta_p \cos \theta_p \\ \beta_p \sin \theta_p \end{pmatrix} + \begin{pmatrix} \epsilon_1 \\ \vdots \\ \epsilon_n \end{pmatrix}$$

Estimate  $\beta$ 's, $\theta$ 's (same as before), and  $\gamma^2$ 's for  $\epsilon$ 's Compute activation from previous complex activation models. Can model actual measurements s with original error structure  $\epsilon$ .

#### **Statistical Implications**

Showed that  $m = \Omega s$ .

If  $E(s) = s_0$  and  $var(s) = \Gamma$ then  $E(m) = \Omega s_0$  and  $var(m) = \Omega \Gamma \Omega^T$ 

And vise versa from multivariate statistics.

There are pre-processing adjustments performed on the data in k-space to correct the signal.

This is done pre-reconstruction and pre-magnitude-only time series.

This is done to your data and you may not know it. -Activation and connectivity implications

#### **Statistical Implications**

These adjustments can be written as linear operations, A.

 $\begin{array}{ll} \mbox{If} & {\rm E}(s) = s_0 & \mbox{ and } {\rm var}(s) = \Gamma \\ \mbox{and} & s_A = As \end{array} \end{array}$ 

then  $\mathsf{E}(s_A) = As_0$  and  $\mathsf{var}(As) = A\Gamma A^T$ and  $\mathsf{E}(m_A) = \Omega As_0$  and  $\mathsf{var}(m) = \Omega A\Gamma A^T \Omega^T$ 

These adjustments modify the signal and noise.

These adjustments induce correlations between voxels (connectivity).

We should account for these adjustments (simpler model in k-space).

#### Discussion

Described complex k-space measurement.

Described complex image reconstruction.

Described complex fMRI activation.

Described relationship between voxel and k-space measurements.

Showed activation from k-space measurements.

Postulated implications of adjustments to k-space measurements.

Further research is needed...

#### Thank You.