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# Utilizing Induced Voxel Correlation in fMRI Analysis

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# Utilizing Induced Voxel Correlation in fMRI Analysis

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Department of Mathematics,  
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Department of Biophysics





## OUTLINE

**1. Reconstruction-Preprocessing**

**2. Induced Correlation**

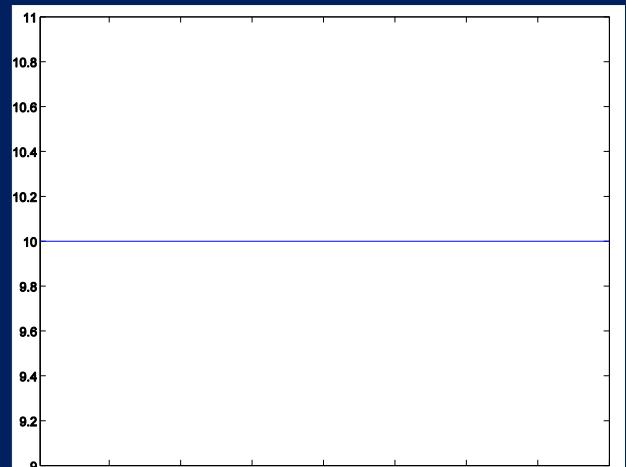
**3. Utilizing Induced Correlation**

**4. Results**

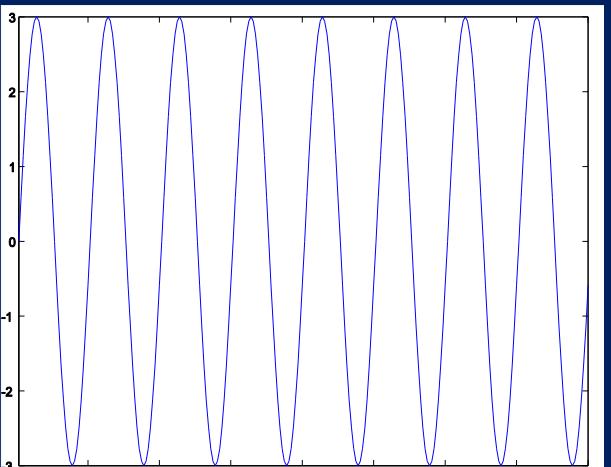
**5. Discussion**

# Reconstruction: 1D FT

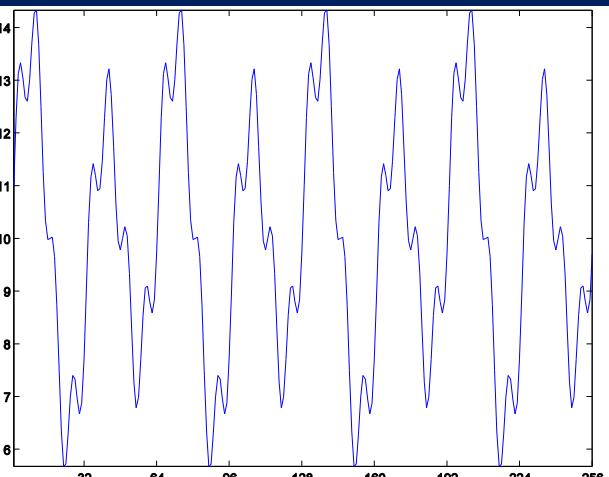
( $n=256, \Delta t=2$  s)



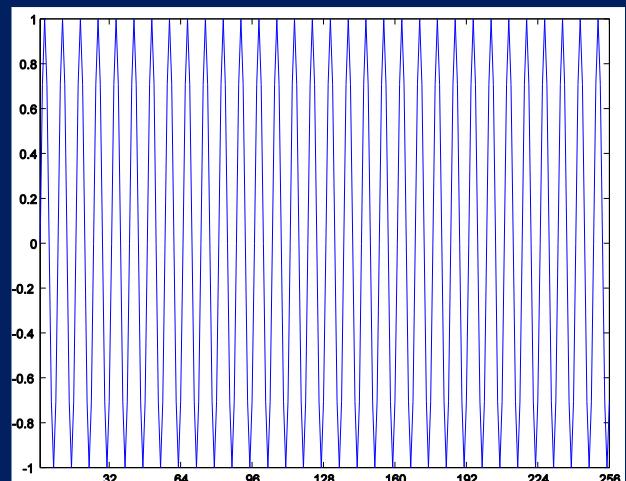
$$10\cos(2\pi 0/512t)$$



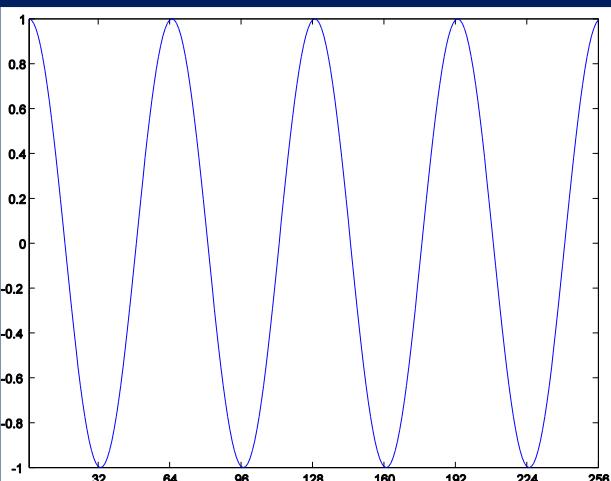
$$3\sin(2\pi 8/512t)$$



sum



$$\cos(2\pi 32/512t)$$

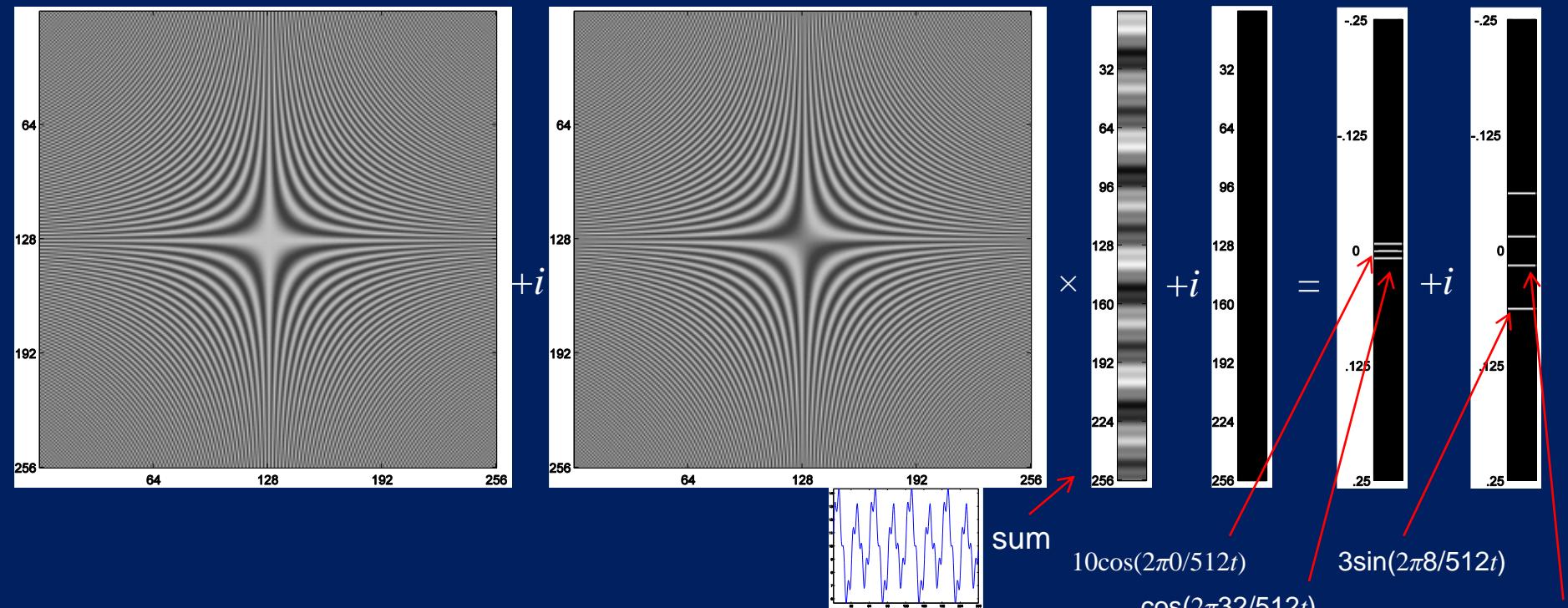


$$\sin(2\pi 4/512t)$$

# Reconstruction: 1D FT

( $n=256, \Delta t=2$  s)

$$(\overline{\Omega}_R + i \overline{\Omega}_I) \times (y_R + i y_I) = (f_R + i f_I)$$



There are lines at the frequency locations.

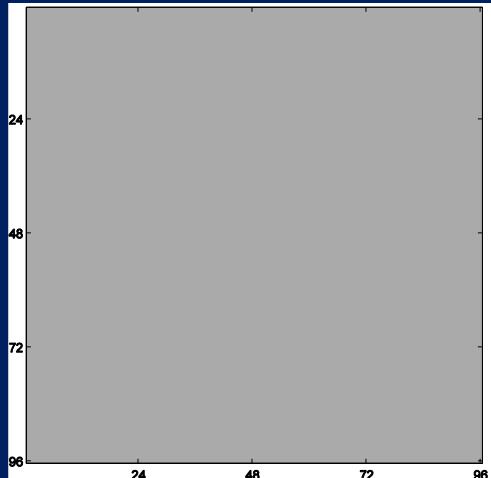
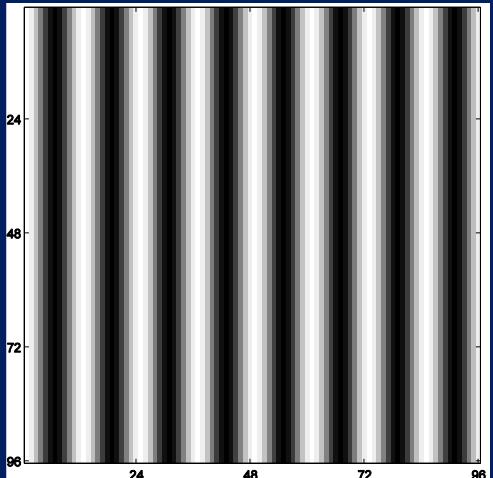
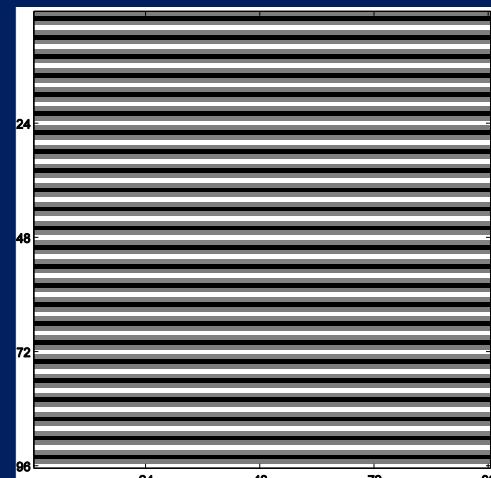
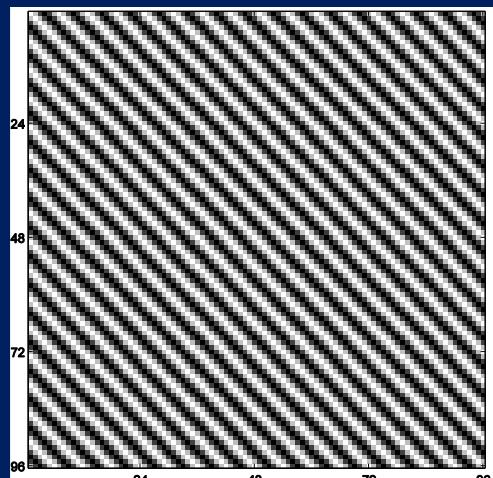
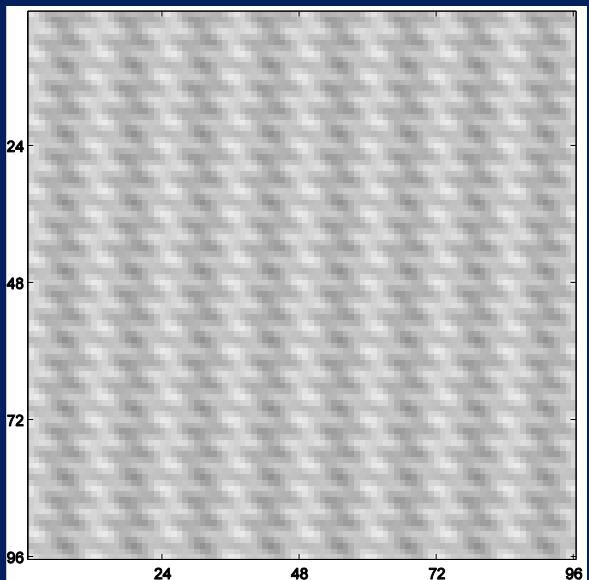
Real part (image) represents constituent cosine frequencies.

Imaginary part (image) represents constituent sine frequencies.

Intensity of the lines represents amplitude of that frequency.

# Reconstruction: 2D FT

(FOV=192 mm)

 $(n_x=n_y=96, \Delta x=\Delta y=2 \text{ mm})$  $10\cos(2\pi 0/96x)$  $1.5\cos(2\pi 8/96x)$  $\sin(2\pi 24/96y)$  $\cos(2\pi 4/96x+2\pi 4/96y)$ 

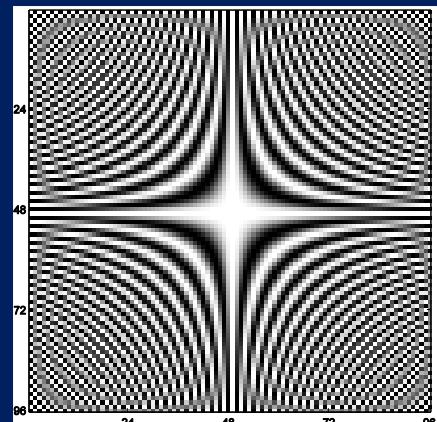
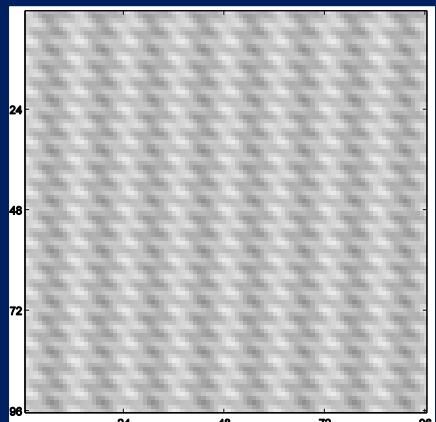
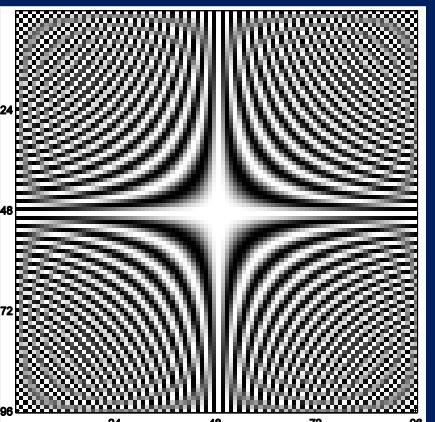
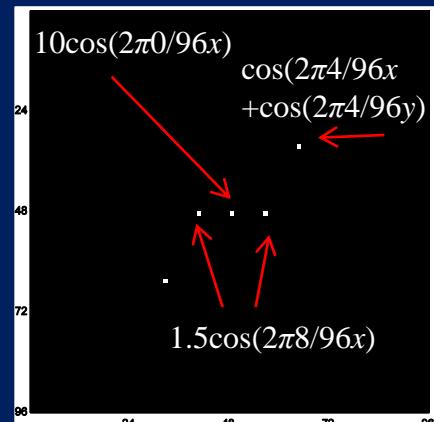
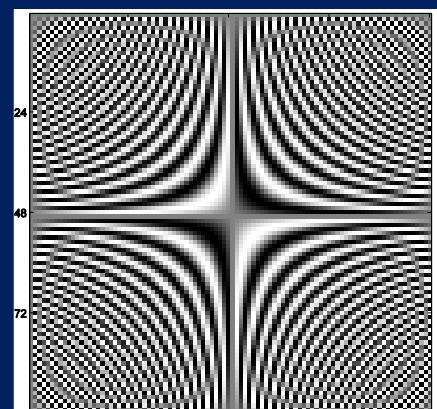
sum

# Reconstruction: 2D FT

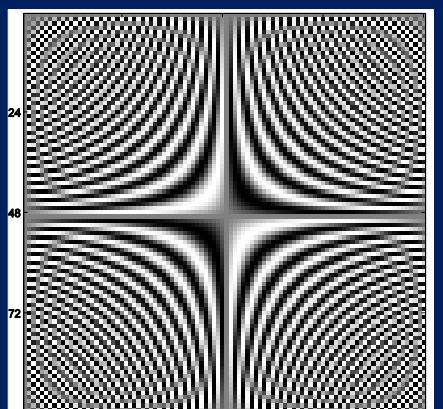
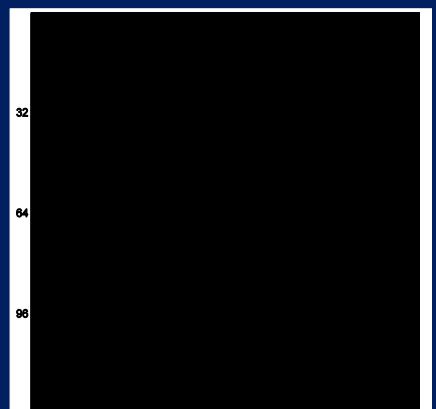
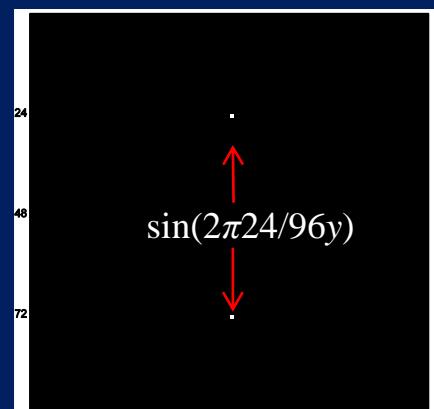
(FOV=192 mm)

 $(n_x=n_y=96, \Delta x=\Delta y=2 \text{ mm})$ 

$$(\overline{\Omega}_{yR} + i \overline{\Omega}_{yI}) \times (V_R + i V_I) \times (\overline{\Omega}_{xR} + i \overline{\Omega}_{xI})^T = (F_R + i F_I)$$

 $+i$  $\times$  $+i$  $=$  $+i$ 

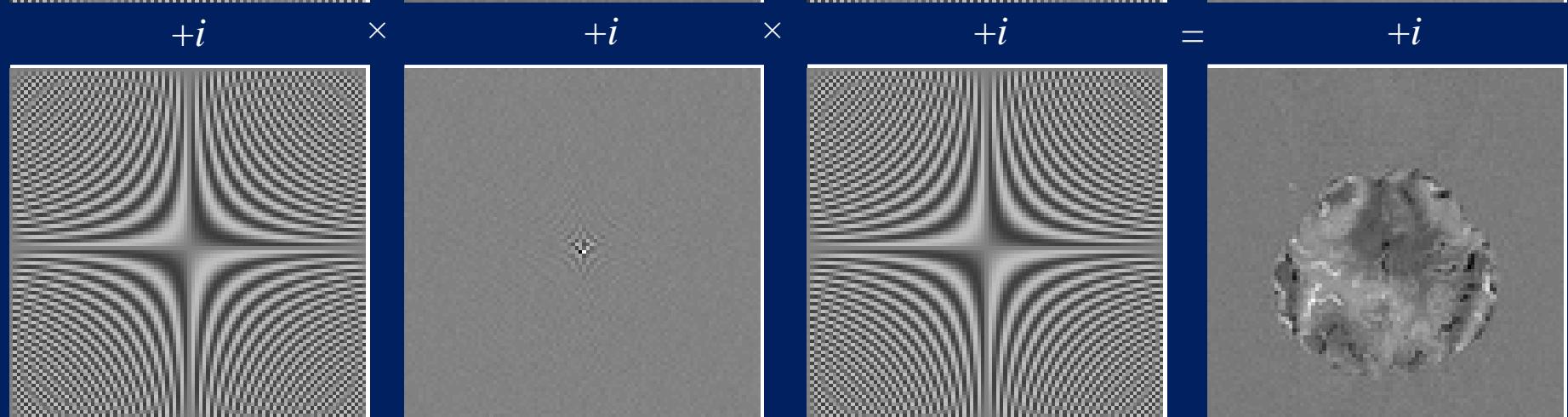
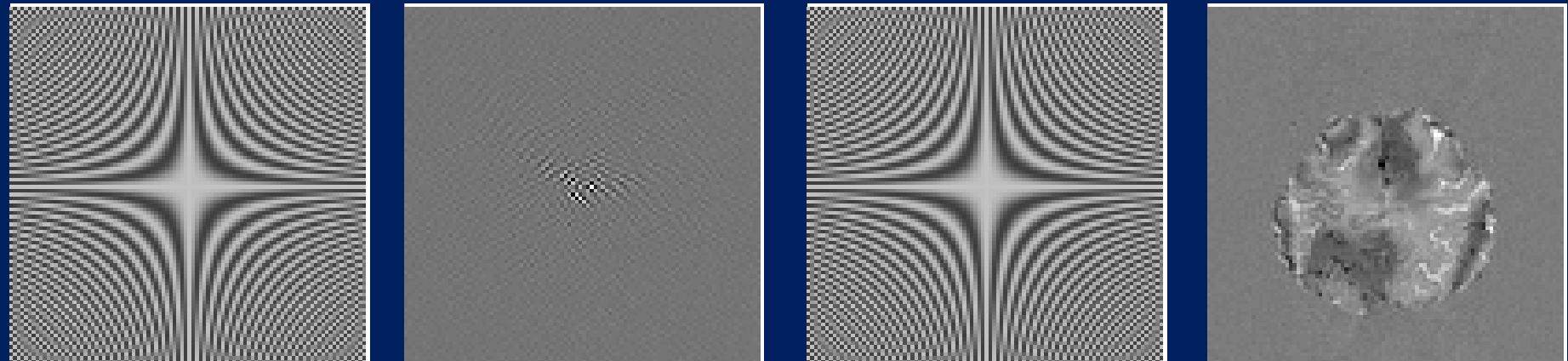
sum

 $\sin(2\pi 24/96y)$ 

# Reconstruction: 2D IFT

$$(\Omega_{yR} + i \Omega_{yI}) \times (F_R + i F_I) \times (\Omega_{xR} + i \Omega_{xI})^T = (V_R + i V_I)$$

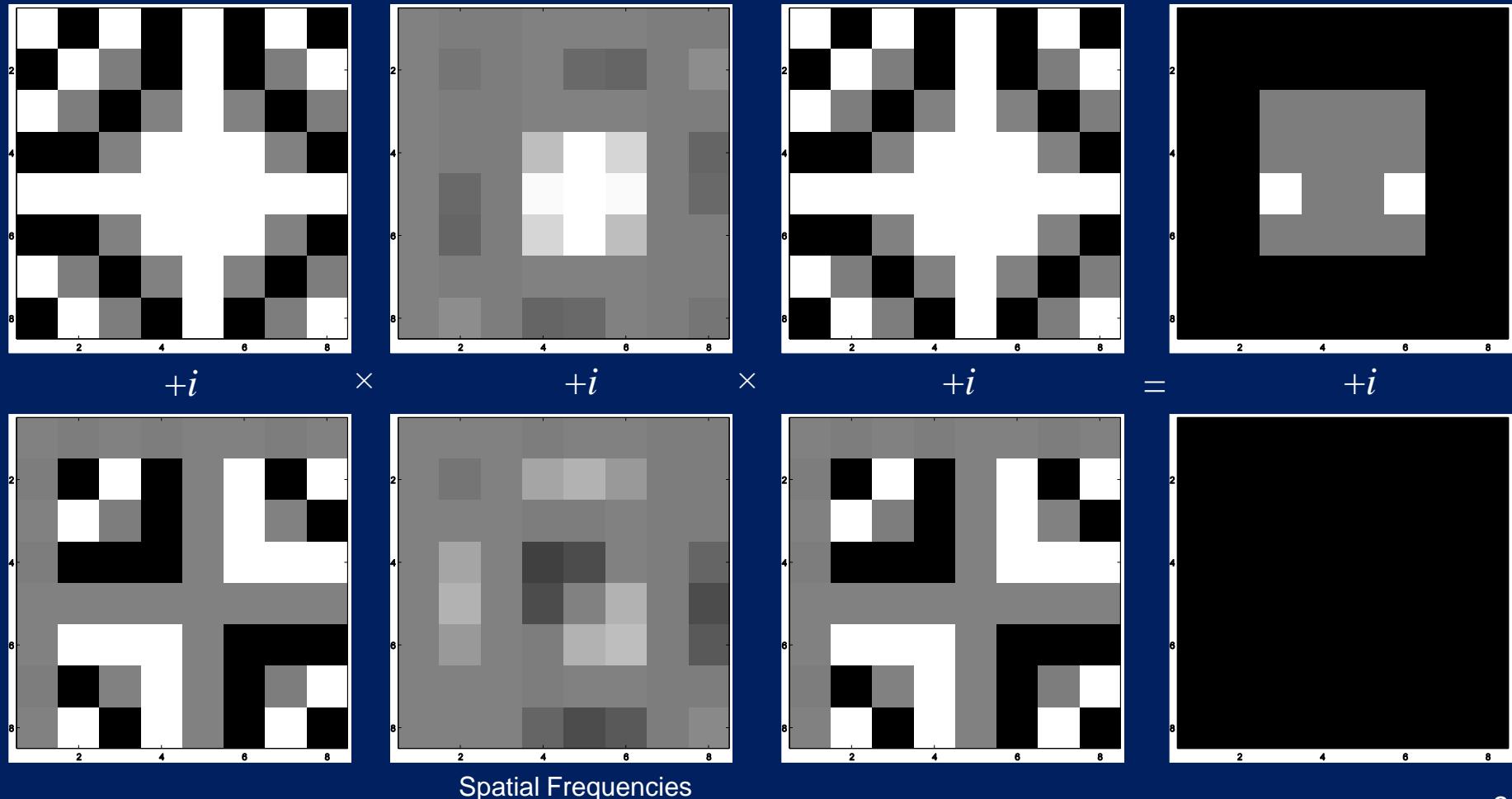
Measure  
(FOV=192 mm)  
( $n_x=n_y=96, \Delta x=\Delta y=2$  mm)



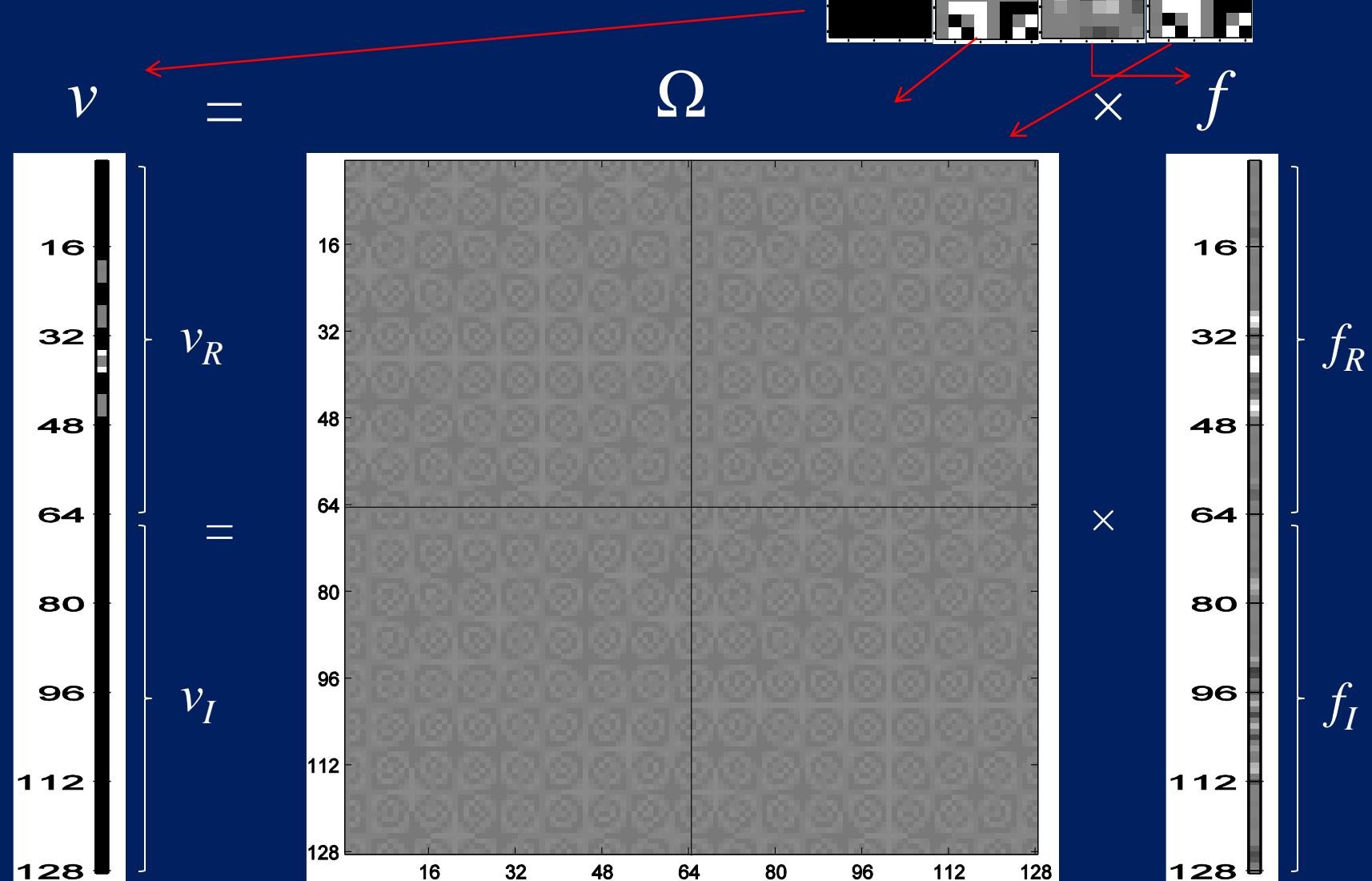
Spatial Frequencies

# Reconstruction: 2D IFT

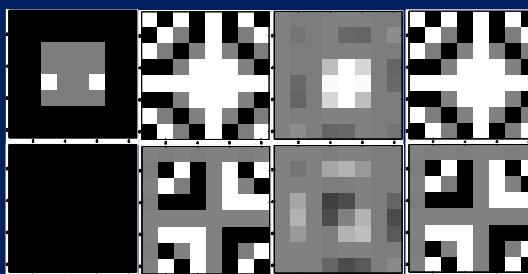
$$(\Omega_{yR} + i \Omega_{yI}) \times (F_R + i F_I) \times (\Omega_{xR} + i \Omega_{xI})^T = (V_R + i V_I)$$



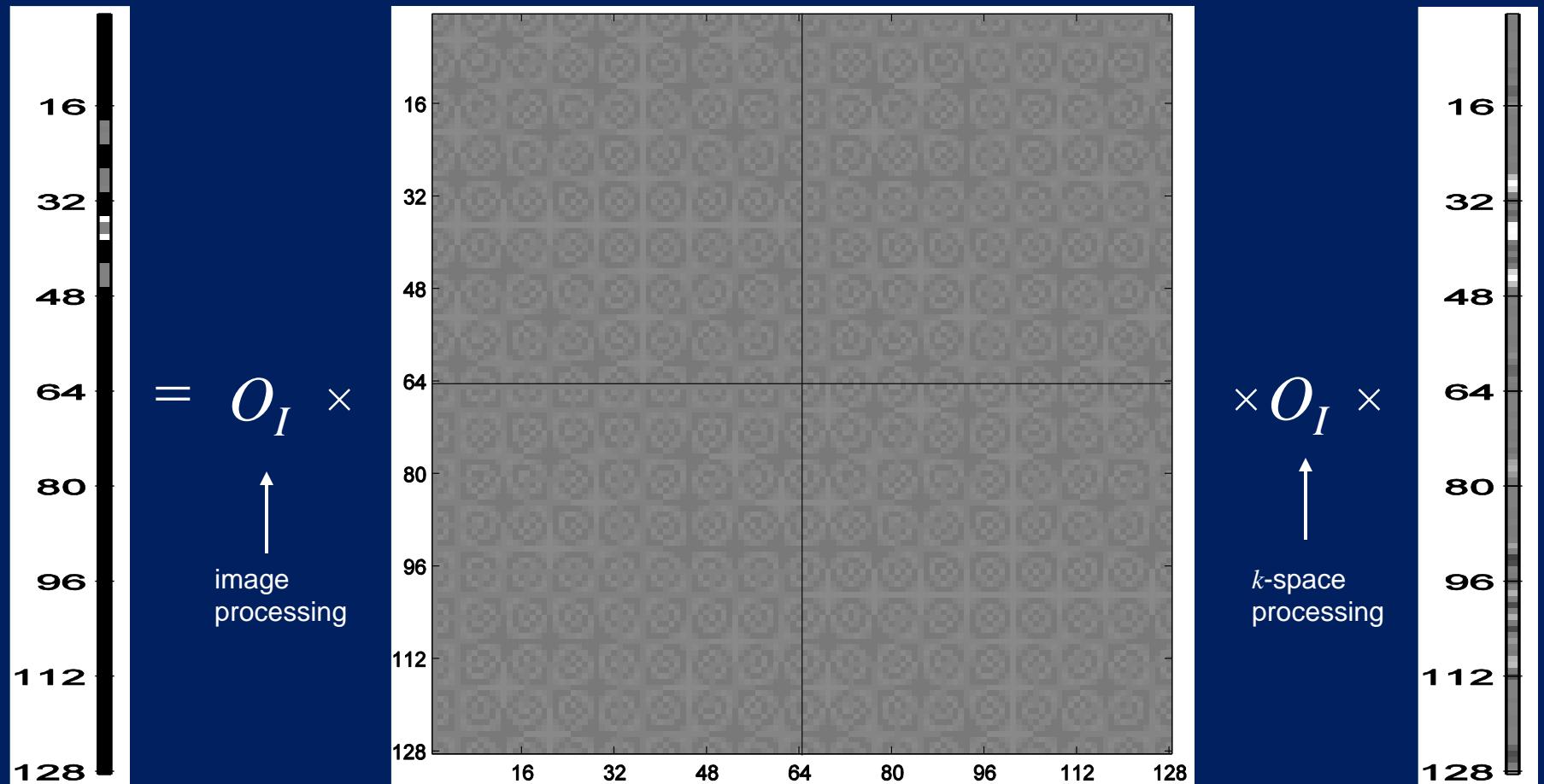
# Reconstruction: 2D IFT Isomorphism



# Reconstruction: Processing Image



$$v = O_I \times \Omega_a \leftarrow \text{adjusted} \times O_I \times f$$



# Reconstruction: Processing Image

$$\nu = O_I \times \Omega_a \times O_k \times f$$

These operators are:

$$f = P_C \mathcal{R} C \mathcal{F}$$

Permuted rows  
 $RI_{\dots}RI \rightarrow RR_{\dots}II$   
 Reverse rows  
 Censor artifacts

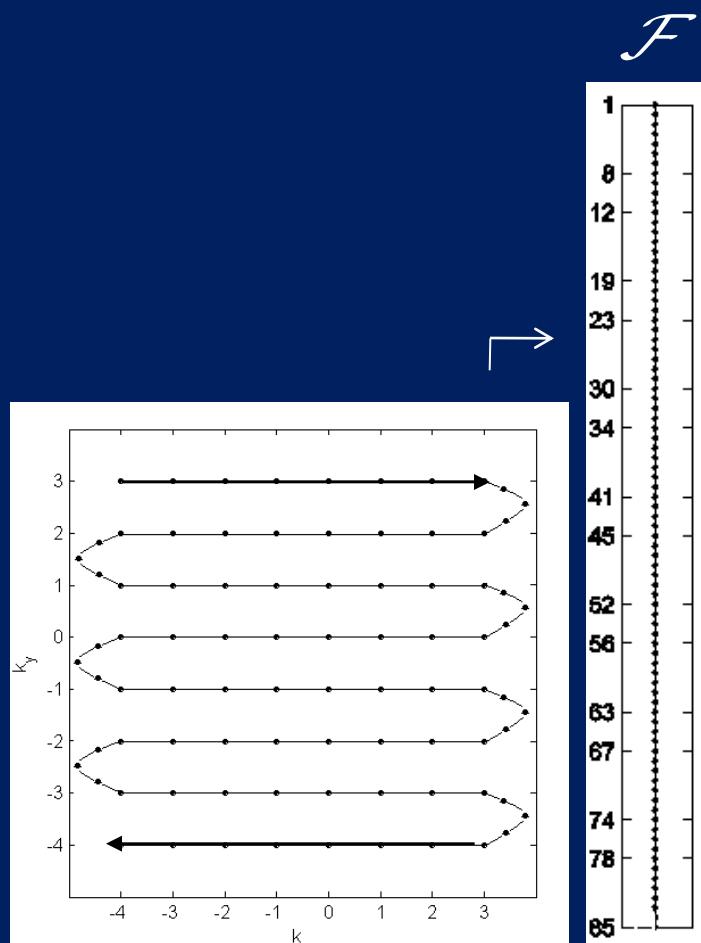
$$O_k = \mathcal{A} \mathcal{Z} \mathcal{H} \underbrace{P_R^{-1} \Omega_{row}^{-1} \Phi \Omega_{row} P_R}_{\text{Nyquist shift rows}} \text{IFT rows} \text{FT rows} \text{permute rows}$$

Apodize  
 Zero-fill  
 Homodyne  
 unpermute rows

$$\Omega_a = \Omega \text{ adjusted for } \Delta B \text{ and for } T_2^*.$$

$$O_I = I_2 \otimes S_m$$

Image smoothing



## Induced Correlation: Mean and Covariance

If  $E(f) = f_0$ , then for  $Of$ ,  $E(Of) = Of_0$ .

If  $\text{cov}(f) = \Gamma$ , then for  $Of$ ,  $\text{cov}(Of) = O\Gamma O^T$ .

This means that with  $v = \underbrace{O_I \Omega_a O_k f}_O$ .

$$E(v) = O_I \Omega_a O_k f_0$$

$$\text{cov}(v) = (O_I \Omega_a O_k) \Gamma (O_k^T \Omega_a^T O_I^T) = \Sigma_{2p \times 2p} \quad \leftarrow \text{Spatial Covariance}$$

$$\text{cor}(v) = R_\Sigma \quad \leftarrow \text{Spatial Correlation}$$

So even if  $\Gamma = \sigma_k^2 I$ , it is not necessarily true that  $\Sigma = \sigma_I^2 I$  !

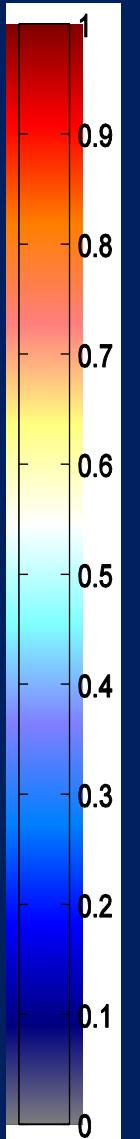
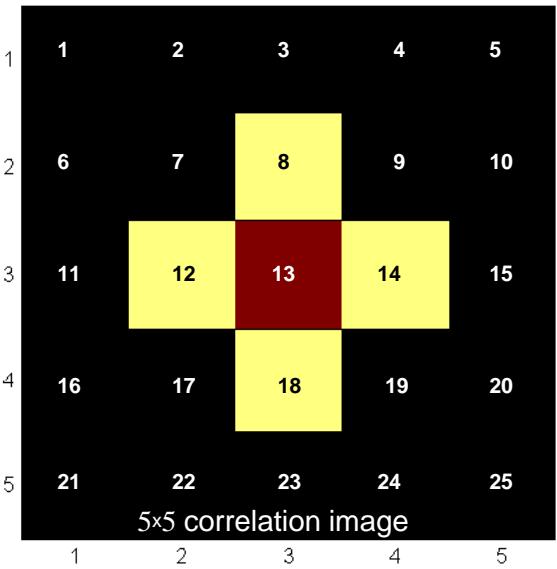
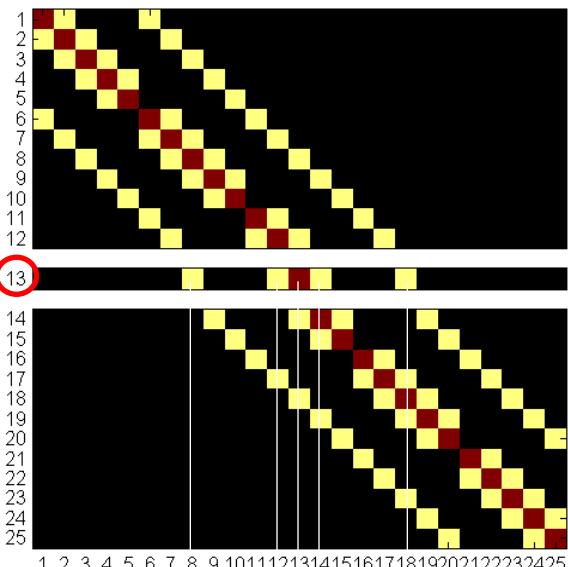
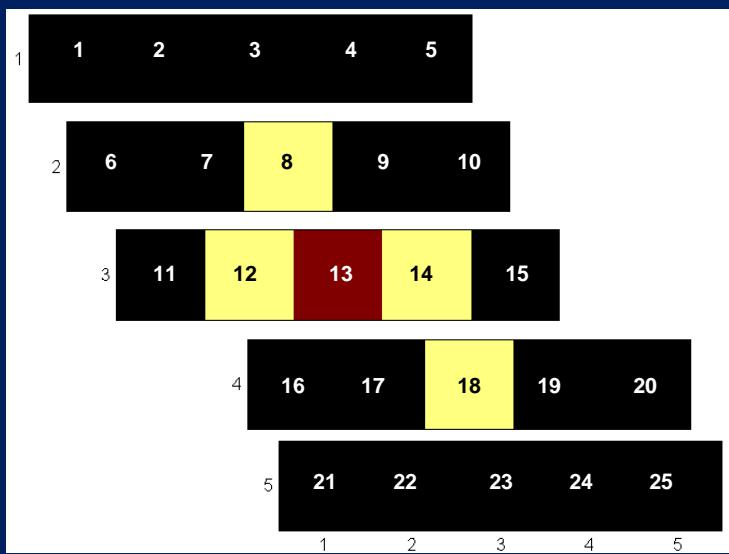
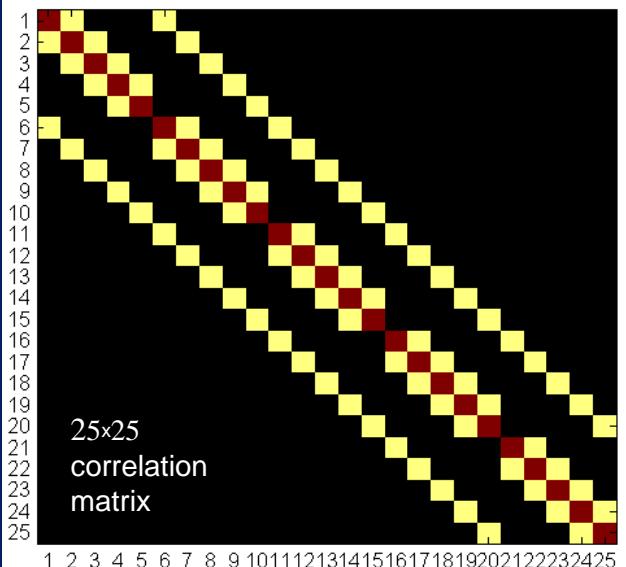
This has  $H_0$  fMRI noise and fcMRI connectivity implications!

# Induced Correlation: Matrix to Image

1	2	3	4	5
6	7	8	9	10
11	12	13	14	15
16	17	18	19	20
21	22	23	24	25

5x5 image

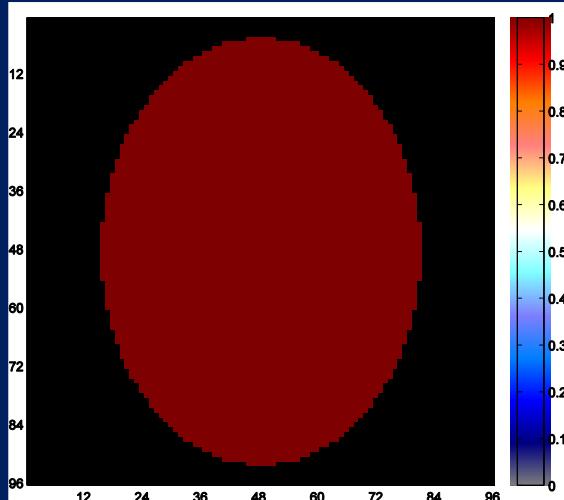
cor =



$$f(t) = \int \int \rho(x, y) e^{-t/T_2^*(x, y)} e^{-i\gamma\Delta B(x, y)t} e^{-i2\pi(k_x x + k_y y)} dx dy$$

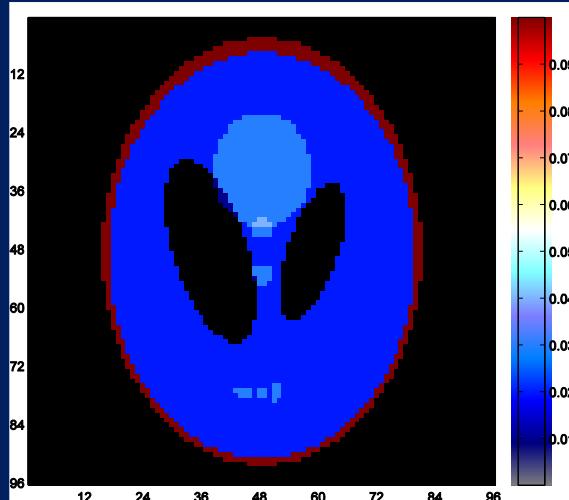
# Induced Correlation: Simulation Parameters

$\rho(x, y)$

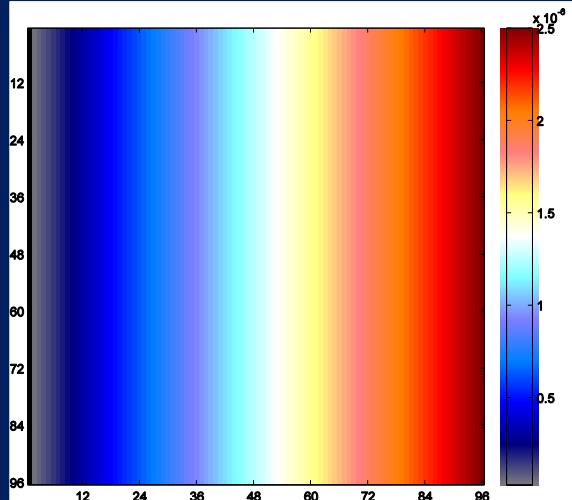


$\Delta t$

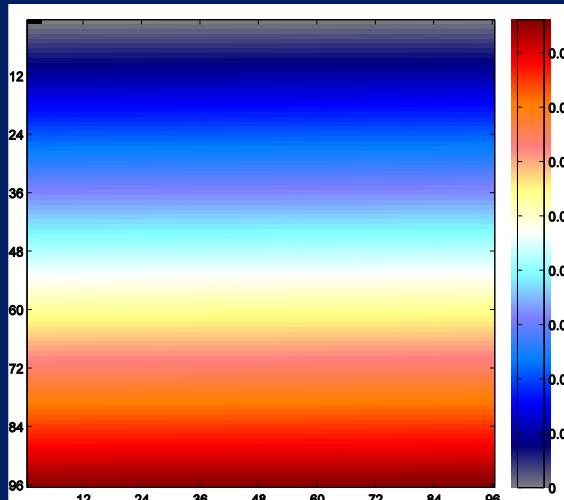
$T_2^*(x, y)$



$\Delta B_\theta(x, y)$

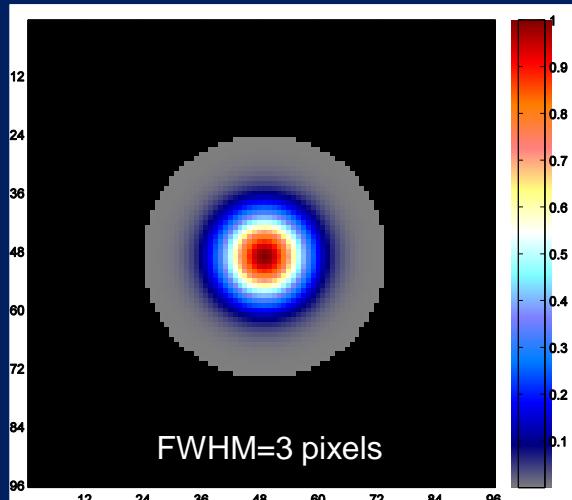


$\mathcal{A} \neq I$



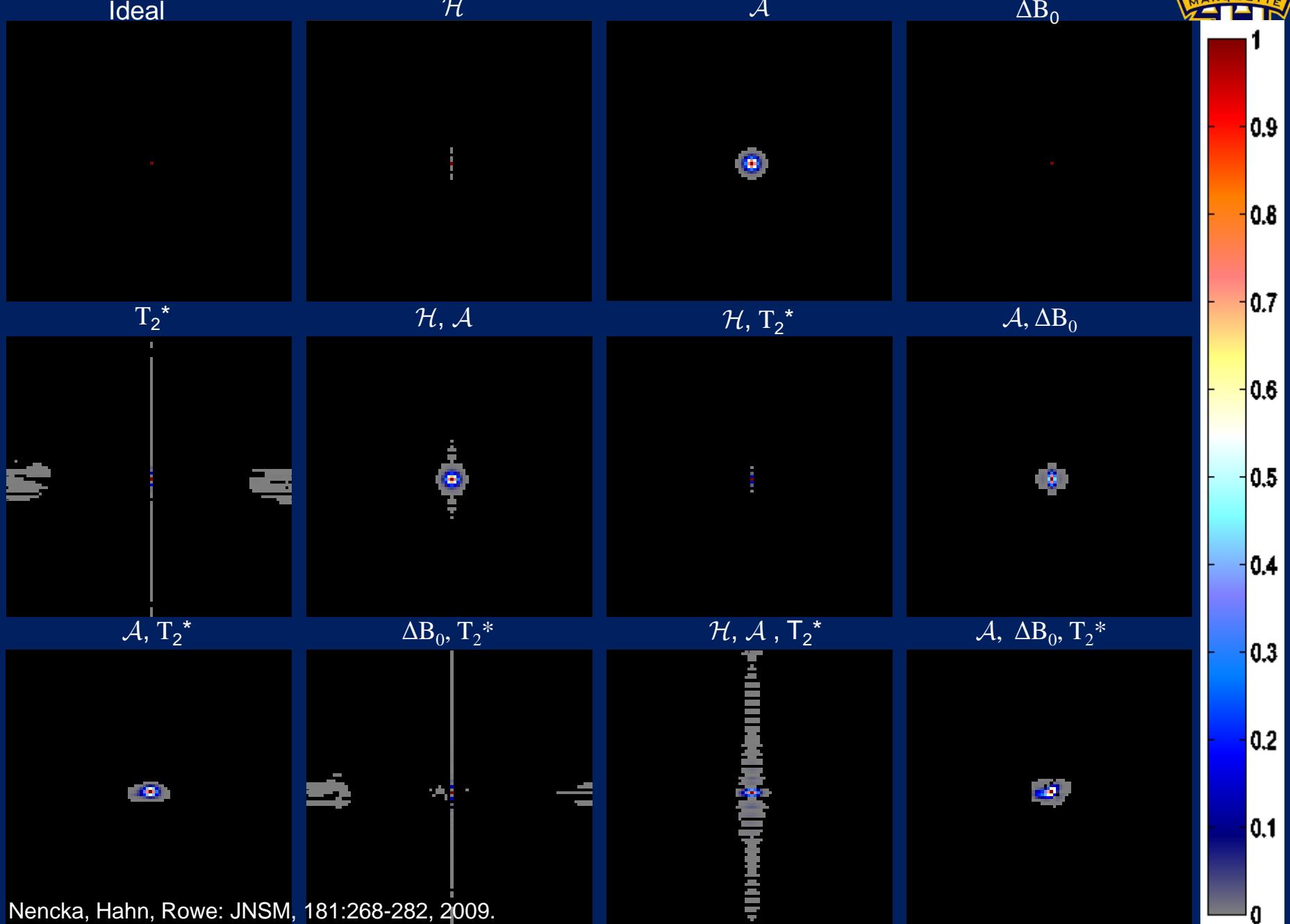
$\mathcal{H} \neq I$   
16 over scan lines

BW=250 kHz  
 $\Delta t = 4 \mu s$   
 EES=0.96 ms  
 TE=50.0 ms

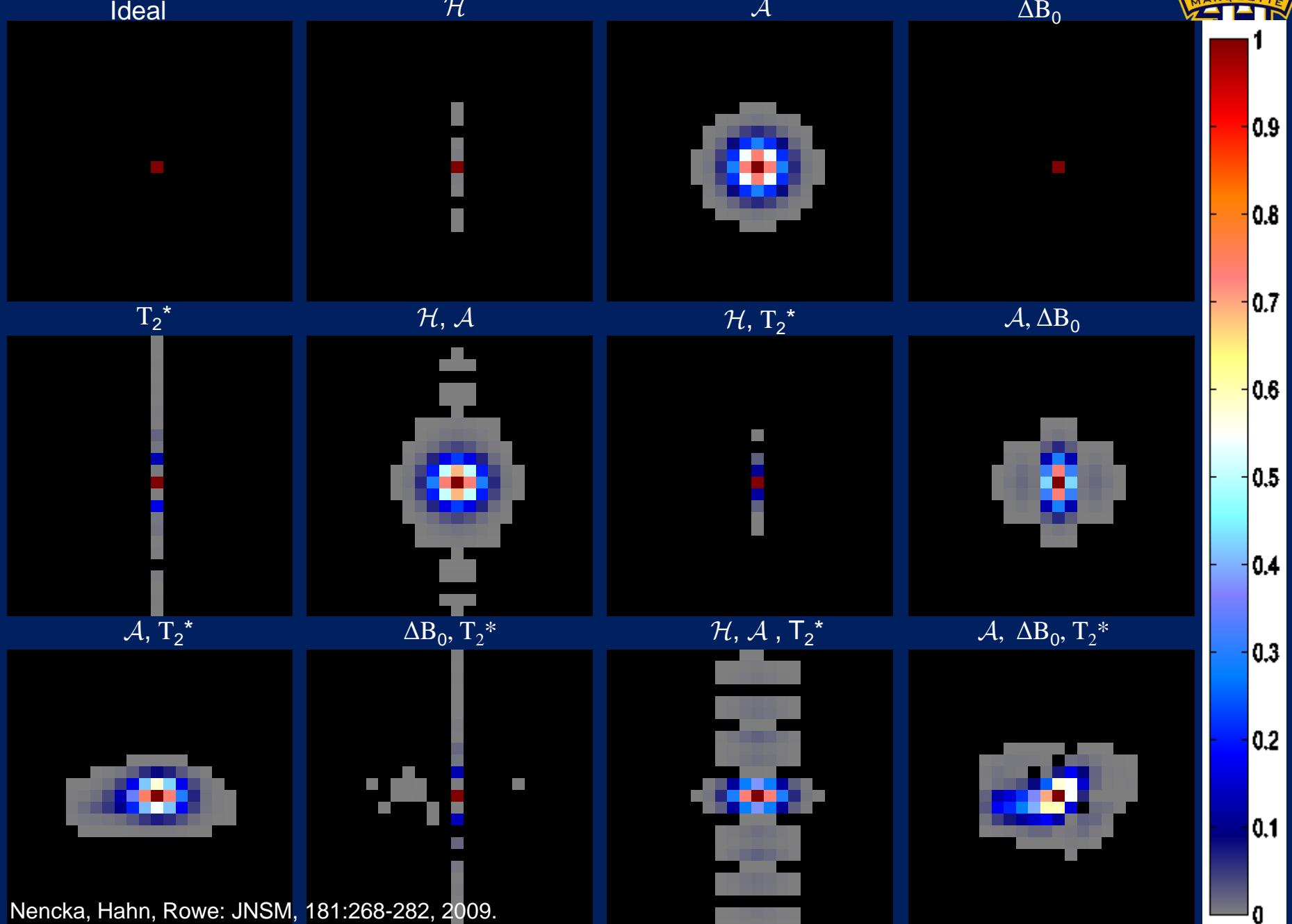


FWHM=3 pixels

Rowe

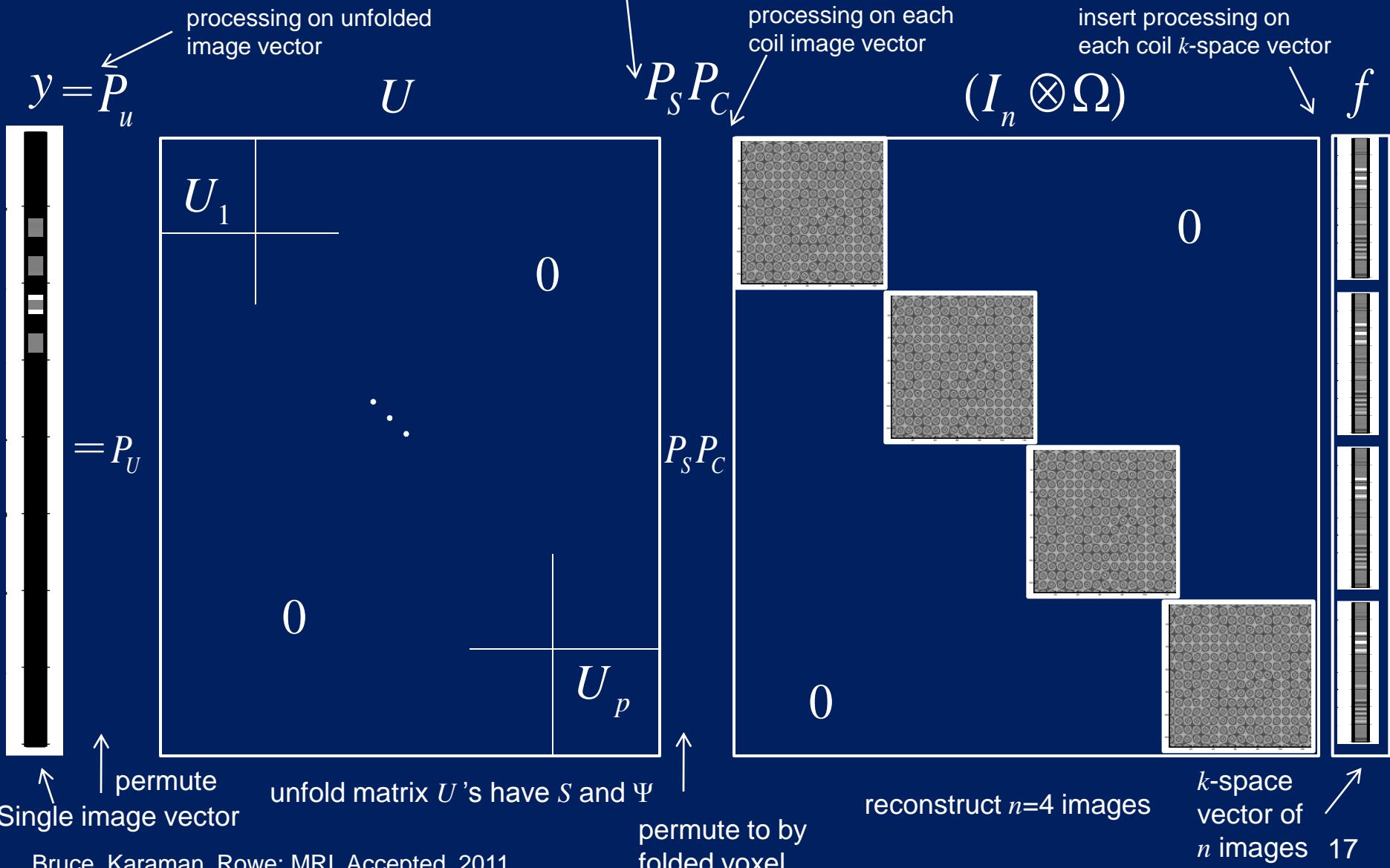
Induced Correlation: Magnitude<sup>2</sup>

Rowe

Induced Correlation: Magnitude<sup>2</sup> Zoomed

here is  $a$  for each voxel

# Induced Correlation: SENSE Multi Coil Combine



## Induced Correlation: SENSE Multi Coil Combine

$$y = \underbrace{\begin{bmatrix} O_I & P_u & U & P_S P_C & (I_n \otimes \Omega_a O_k) \end{bmatrix}}_O f$$

where

$f = (f_1, \dots, f_n)'$  are coil  $k$ -space

$O_k$  is  $k$ -space preprocessing

$\Omega_a$  is adj. inverse Fourier matrix     $\Omega_a = \Omega$     adjusted for  $\Delta B$   
and for  $T_2^*$

$P_u$ ,  $P_S$ ,  $P_C$ , permutation matrices

$U$  SENSE unfolding matrix

$O_I$  is image space preprocessing

$$f = P_C \mathcal{R} C \mathcal{F}$$

$$O_k = \mathcal{A} \mathcal{Z} \mathcal{H} \underbrace{P_R^{-1} \Omega_{row}^{-1} \Phi \Omega_{row} P_R}_{\text{adjusted for } \Delta B \text{ and for } T_2^*}$$

$$O_I = I_2 \otimes S_m$$

k-space vector  
censor uturns  
row reverse  
permute

Image smoothing

## Induced Correlation: SENSE Multi Coil Combine Statistical Expectation and Covariance.

If  $E(f) = f_0$ , then for  $Mf$ ,  $E(Mf) = Mf_0$ .

If  $\text{cov}(f) = \Gamma$ , then for  $Mf$ ,  $\text{cov}(Mf) = M\Gamma M'$ .

This means that with  $y = Of$ ,

$$E(y) = Of_0 \quad \text{and} \quad \text{cov}(y) = O\Gamma O' = \Sigma_{2p \times 2p}$$

$$\Rightarrow \text{cor}(v) = D_\Sigma^{-1/2} \Sigma D_\Sigma^{-1/2}$$

So even if  $\Gamma = \sigma^2 I$ , it is not necessarily true that  $\Sigma = \sigma^2 I$  !

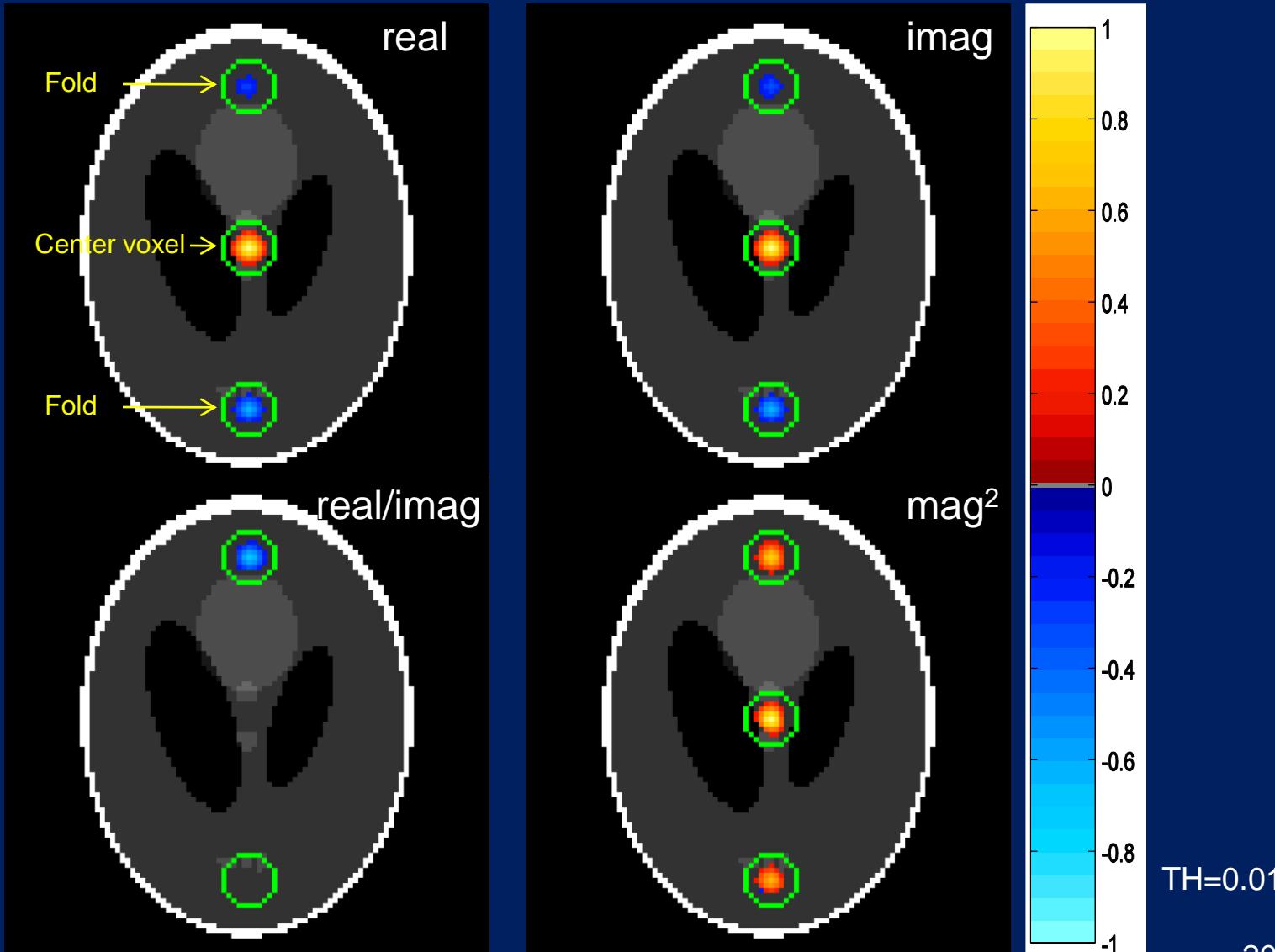
This has  $H_0$  fMRI noise and fcMRI connectivity implications!

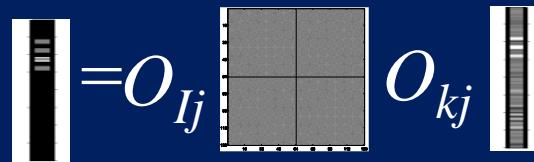
Correlations induced about the center voxel.

## Induced Correlation: SENSE Multi Coil Combine

$N_X=96$   
 $N_Y=96$   
 $n = 4$   
 $A = 3$   
FWHM=3

Functional connectivity implications





## Induced Correlation: Extend to Time Series

Reconstruction of  $n$  images described as:

$$\boldsymbol{\nu} = \begin{matrix} I \\ R \\ K \\ f \end{matrix}$$

$2np \times 1$

↑      ↑      ↗  
diagonal

$$\begin{aligned} K &= BlkDiag(O_{Kt}) \\ R &= BlkDiag(\Omega_{at}) \\ I &= BlkDiag(O_{It}) \end{aligned}$$

$$\left( \begin{matrix} \boldsymbol{\nu}_1 \\ \vdots \\ \boldsymbol{\nu}_n \end{matrix} \right) = \left( \begin{matrix} O_{I1}\Omega_{a1}O_{k1} & & & 0 \\ & \ddots & & \\ 0 & & O_{In}\Omega_{an}O_{kn} & \left( \begin{matrix} f_1 \\ \vdots \\ f_n \end{matrix} \right) \end{matrix} \right) \left( \begin{matrix} & & & \\ & & & \\ & & & \\ \uparrow & & & \uparrow \\ \text{image-space} & & & \text{k-space} \\ \text{vector of} & & & \text{vector} \\ n \text{ images} & & & n \text{ images} \end{matrix} \right)$$

↑      ↑      ↗  
Reconstruction  
 $k$ -space operation  
matrix for  $n$  images

image 1  
 $k$ -space  
vector

image  $n$   
 $k$ -space  
vector

# Induced Correlation: Extend to Time Series

 $v =$ 
 $IRK$ 
 $\times f$ 

$$\begin{matrix}
 v & = & IRK & \times & f \\
 \left[ \begin{array}{c} 16 \\ 32 \\ 48 \\ 64 \\ 80 \\ 96 \\ 112 \\ 128 \end{array} \right] & \left\{ \begin{array}{l} V_1 \\ \text{image 1} \\ \text{vector} \end{array} \right\} & O_{I1} & & \left[ \begin{array}{c} 16 \\ 32 \\ 48 \\ 64 \\ 80 \\ 96 \\ 112 \\ 128 \end{array} \right] & \left\{ \begin{array}{l} f_1 \\ \text{image 1} \\ k\text{-space} \\ \text{vector} \end{array} \right\} \\
 & \vdots & & & & \vdots \\
 \left[ \begin{array}{c} 16 \\ 32 \\ 48 \\ 64 \\ 80 \\ 96 \\ 112 \\ 128 \end{array} \right] & \left\{ \begin{array}{l} V_n \\ \text{image } n \\ \text{vector} \end{array} \right\} & 0 & \dots & \left[ \begin{array}{c} 16 \\ 32 \\ 48 \\ 64 \\ 80 \\ 96 \\ 112 \\ 128 \end{array} \right] & \left\{ \begin{array}{l} f_n \\ \text{image } n \\ k\text{-space} \\ \text{vector} \end{array} \right\}
 \end{matrix}$$

$O_{I1}$        $\Omega_{a1}$        $O_{k1}$        $0$        $\dots$        $O_{In}$        $\Omega_{an}$        $O_{kn}$

# Induced Correlation: Extend to Time Series

(dyn  $\Delta B_0$ ,  $\Delta x$ ,  $\Delta t$ , freq filt)

$$y = T \cdot P \cdot IRK \cdot f$$

$$\begin{pmatrix} y_1 \\ \vdots \\ y_p \\ 2np \times 1 \end{pmatrix} = \begin{pmatrix} O_{T1} & & 0 \\ & \ddots & \\ 0 & & O_{Tp} \end{pmatrix}_{2np \times 2np} P \begin{pmatrix} O_{I1}\Omega_{a1}O_{k1} & & 0 \\ & \ddots & \\ 0 & & O_{In}\Omega_{an}O_{kn} \end{pmatrix}_{2np \times 2np} \begin{pmatrix} f_1 \\ \vdots \\ f_n \\ 2np \times 1 \end{pmatrix}$$

voxel 1 temporal processing  
voxel  $p$  series filter      permute from measurements by image to by voxel

$$y = \begin{pmatrix} y_1 \\ \vdots \\ y_n \end{pmatrix} \quad \xleftarrow{\text{voxel 1 temporally processed series}}$$

voxel  $j$  temporally processed series

$$y_j = \begin{pmatrix} y_{Rj} \\ y_{Ij} \end{pmatrix}_{n \times 1} \quad j = 1, \dots, p$$

$n$  reals

## Induced Correlation: Mean and Covariance

If  $E(f) = f_0$ , then for  $E(Of) = Of_0$ .

If  $\text{cov}(f) = \Gamma$ , then for  $\text{cov}(Of) = O\Gamma O^T$ .

This means that with  $y = \underbrace{TPIRKf}_O$ .

$$E(y) = TPIRKf_0$$

$$\text{cov}(y) = (TPIRK)\Gamma(K^T R^T I^T P^T T^T) = \Sigma_{2np \times 2np}$$

$$\text{cor}(y) = R_\Sigma \quad \leftarrow \text{Spatio-Temporal Correlation}$$

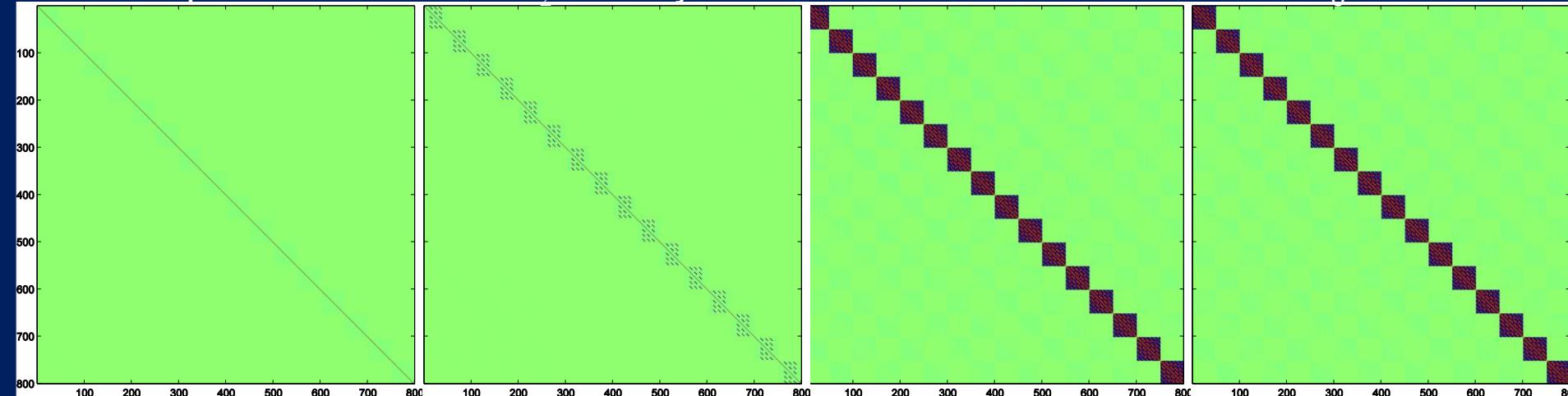
Spatio-Temporal  
Covariance  
HUGE

So even if  $\Gamma = \sigma_k^2 I$ , it is not necessarily true that  $\Sigma = \sigma_I^2 I$  !

This has  $H_0$  fMRI noise and fcMRI connectivity implications!

# Induced Correlation: Example $5 \times 5$ image 8 TRs 2 slices

No Operations

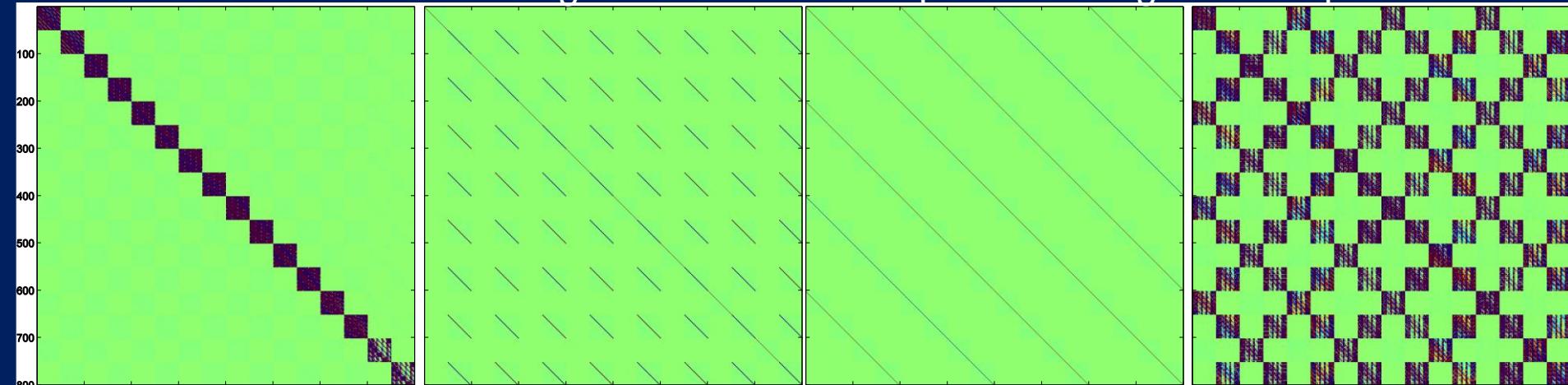
 $T_2^*$  DecayApodization,  $\mathcal{A}$  $\Delta B_0$  Error

Motion Correction

Timing Correction

Temporal Filtering

All Operations



$$\Sigma = OIO^T \rightarrow R_\Sigma$$

800x800                    800x800

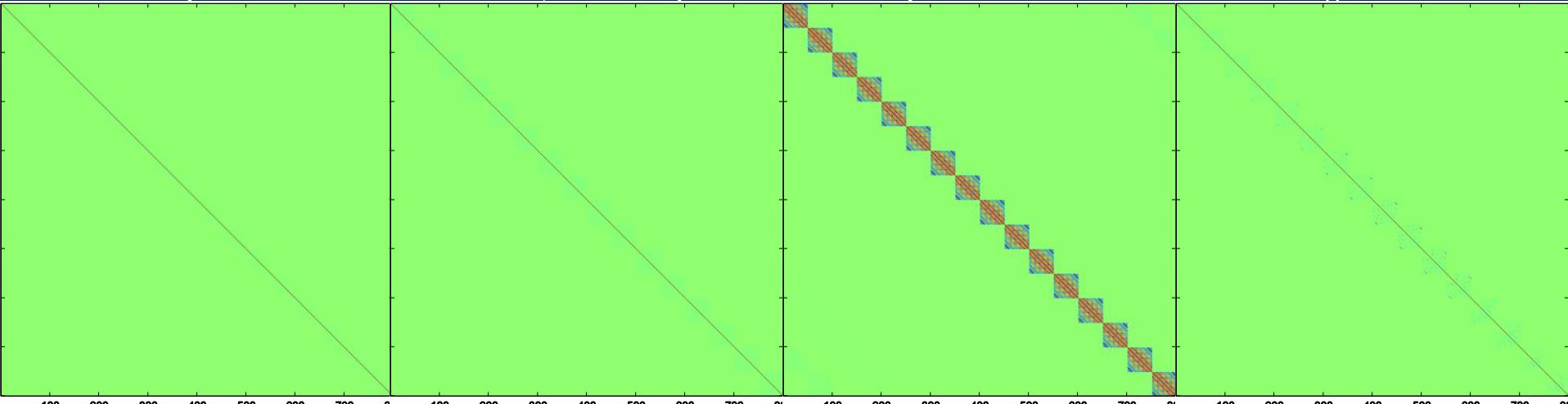
# Induced Correlation: Example 5x5 image 8 TRs 2 slices

No Operations

$T_2^*$  Decay

Apodization,  $\mathcal{A}$

$\Delta B_0$  Error

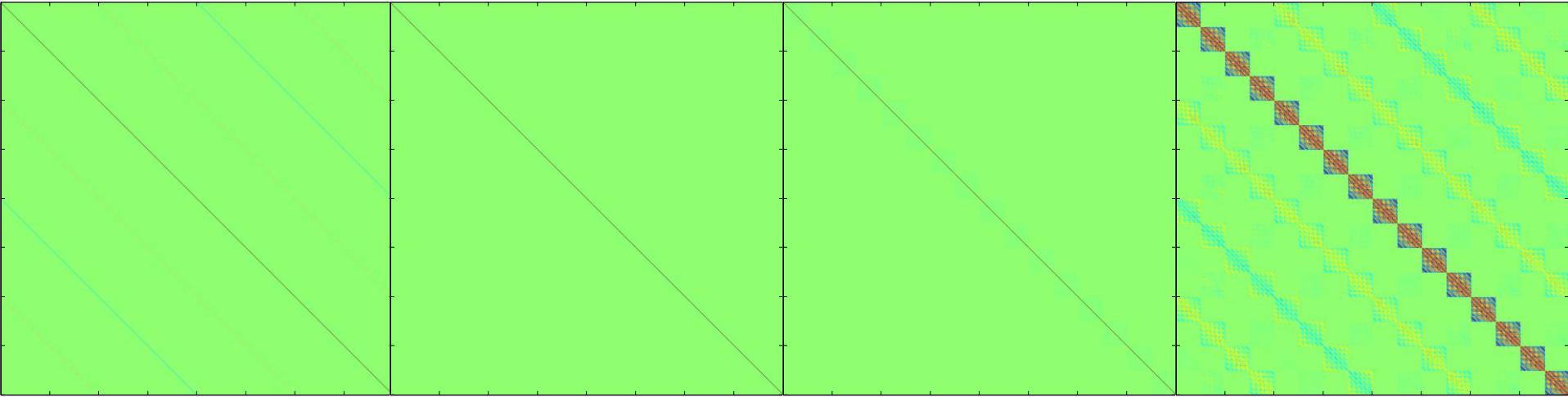


Temporal Filtering

Timing Correction

Motion Correction

All Operations

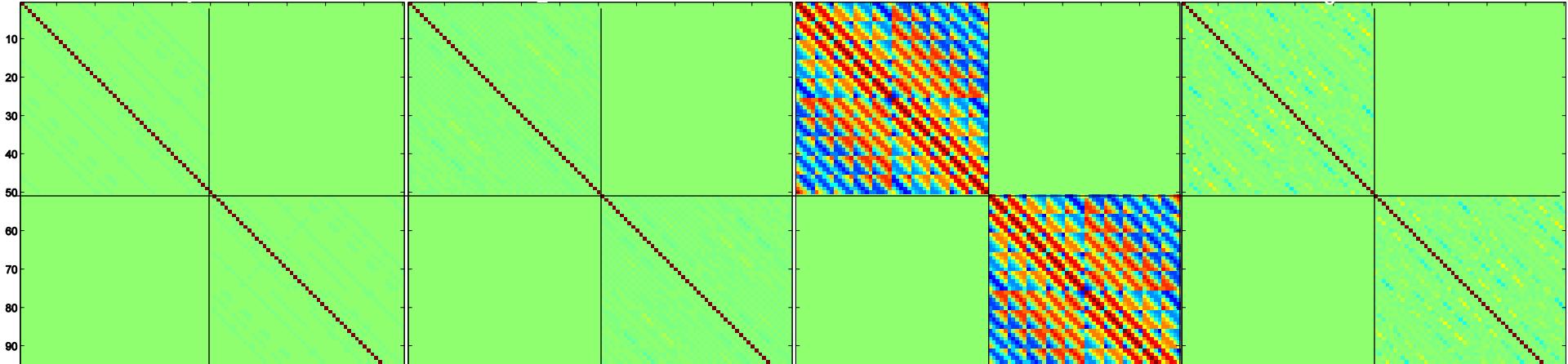


$$\Sigma = OIO^T \rightarrow R_S$$

100x100                    100x100

# Induced Correlation: Example 5x5 image 8 TRs 2 slices

No Operations

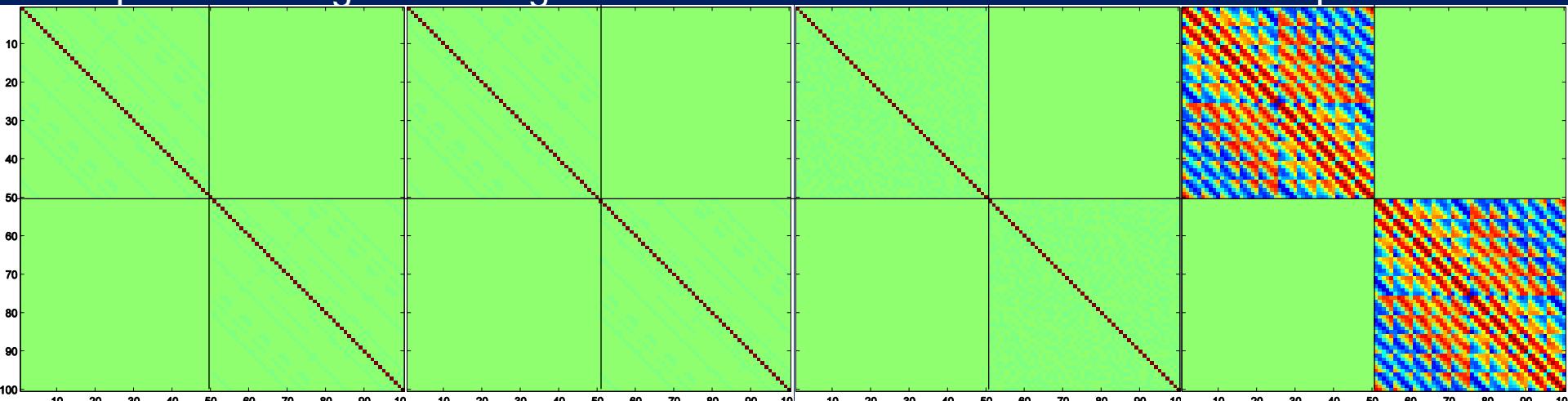
 $T_2^*$  DecayApodization,  $\mathcal{A}$  $\Delta B_0$  Error

Temporal Filtering

Timing Correction

Motion Correction

All Operations



$$\Sigma = OIO^T \rightarrow R_T$$

16x16                            16x16

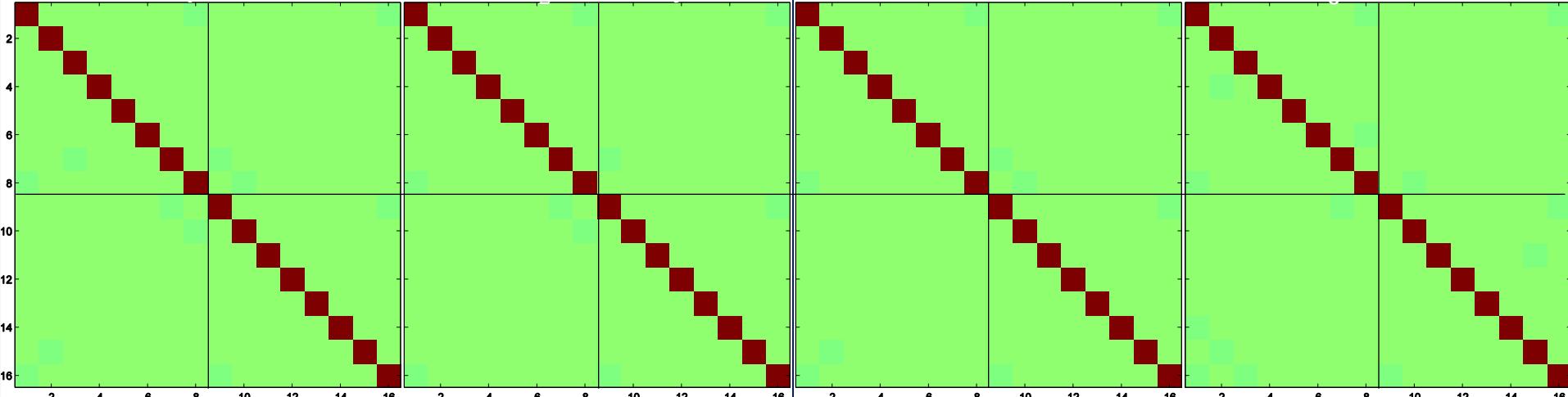
# Induced Correlation: Example 5x5 image 8 TRs 2 slices

No Operations

$T_2^*$  Decay

Apodization,  $\mathcal{A}$

$\Delta B_0$  Error

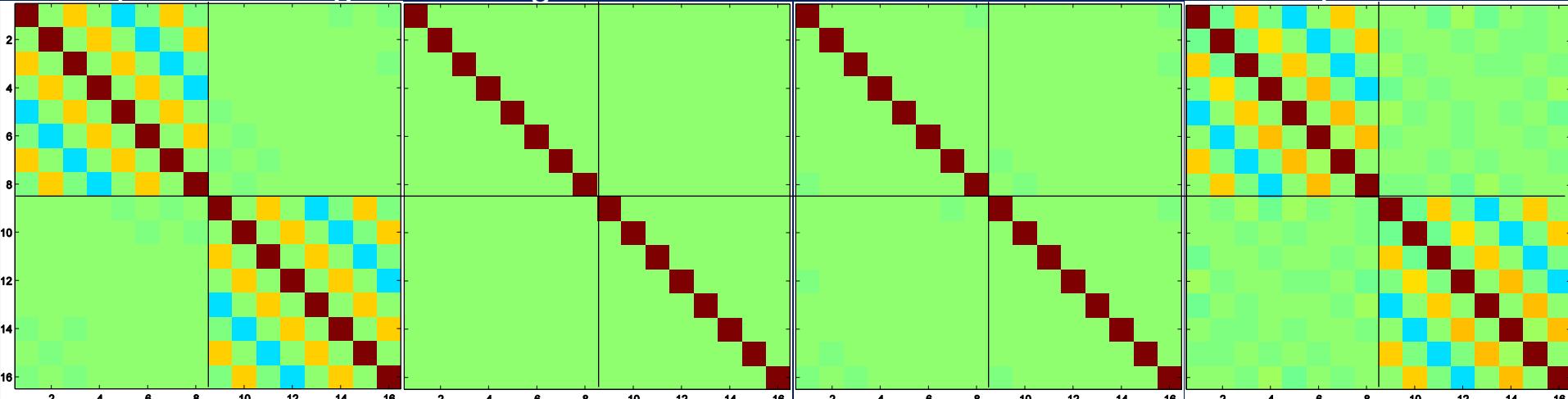


Temporal Filtering

Timing Correction

Motion Correction

All Operations



## Utilizing Induced Correlation:

Since for all voxels  
is

$$\Sigma_{2np \times 2np} = \begin{pmatrix} \Sigma_1 & & \Sigma_{jk} \\ & \ddots & \\ \Sigma'_{jk} & & \Sigma_p \end{pmatrix}$$

Then for voxel  $j$ , the R-I cov

$$\Sigma_j_{2n \times 2n} = \begin{pmatrix} \Sigma_{jR} & \Sigma_{jRI} \\ \Sigma'_{jRI} & \Sigma_{jI} \end{pmatrix}$$

and the magnitude<sup>2</sup> covariance is

$$\begin{aligned} \sigma_j &= \text{tr}(\Sigma_j) + \mu'_j \mu_j \\ \Lambda_{jj} &= 2\text{tr}(\Sigma'_j \Sigma_j) + 4\mu'_j \Sigma_j \mu_j , \\ \Lambda_{jk} &= 2\text{tr}(\Sigma'_{jk} \Sigma_{jk}) + 4\mu'_j \Sigma_{jk} \mu_k \end{aligned}$$

## Utilizing Induced Correlation:

Complex-Valued

$$C_j = \begin{pmatrix} \cos \theta_{j1} & & 0 \\ & \ddots & \\ 0 & & \cos \theta_{jn} \end{pmatrix} \quad S_j = \begin{pmatrix} \sin \theta_{j1} & & 0 \\ & \ddots & \\ 0 & & \sin \theta_{jn} \end{pmatrix}$$

$$\begin{pmatrix} y_{jR} \\ y_{jI} \end{pmatrix} = \begin{pmatrix} C_j X \beta_j \\ S_j X \beta_j \end{pmatrix} + \begin{pmatrix} \eta_{jR} \\ \eta_{jI} \end{pmatrix},$$

↑  
Compute activation individually for each voxel.

Magnitude-Only (assuming high SNR)

$$m_j = X \beta_j + \varepsilon_j,$$

↑  
Compute activation individually for each voxel.

Can form larger spatio-temporal model.

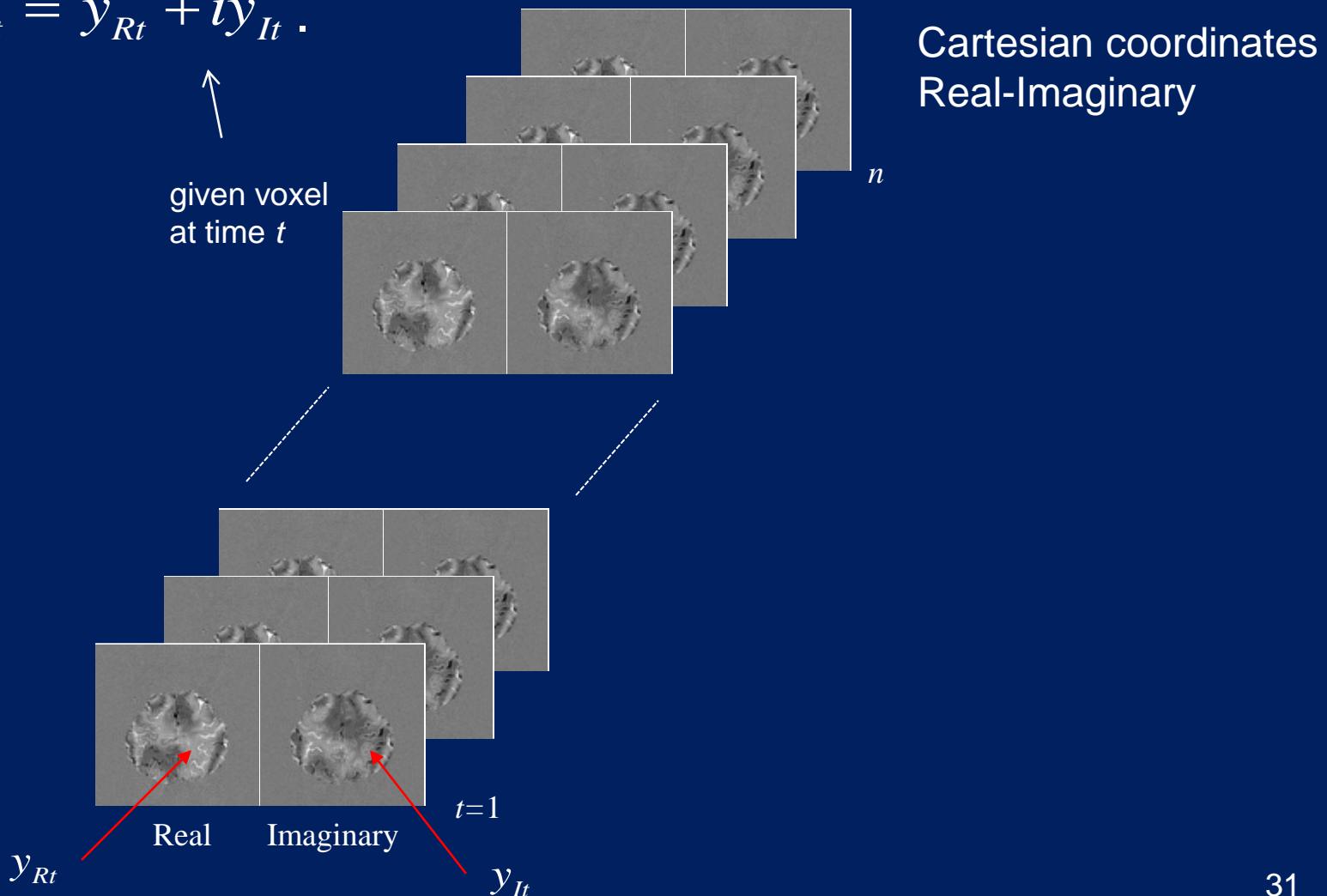
$$\eta_j \sim N(0, \Sigma_j)$$

Incorporate  
Induced  
Covariance

$$\varepsilon_j \sim N(0, \Lambda_j)$$

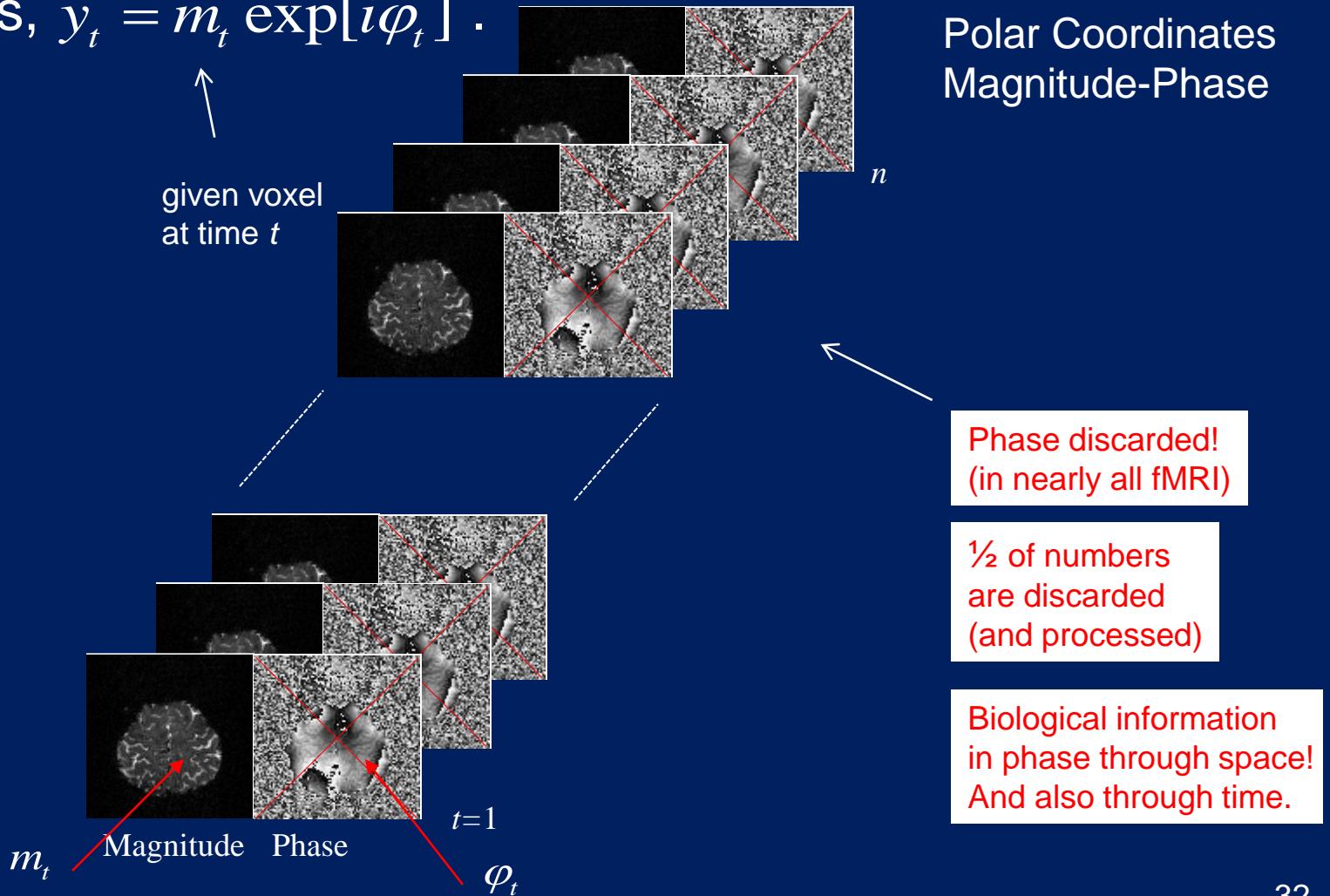
# Utilizing Induced Correlation: Complex fMRI

The fMRI data is truly complex-valued images and voxel time series,  $y_t = y_{Rt} + iy_{It}$ .



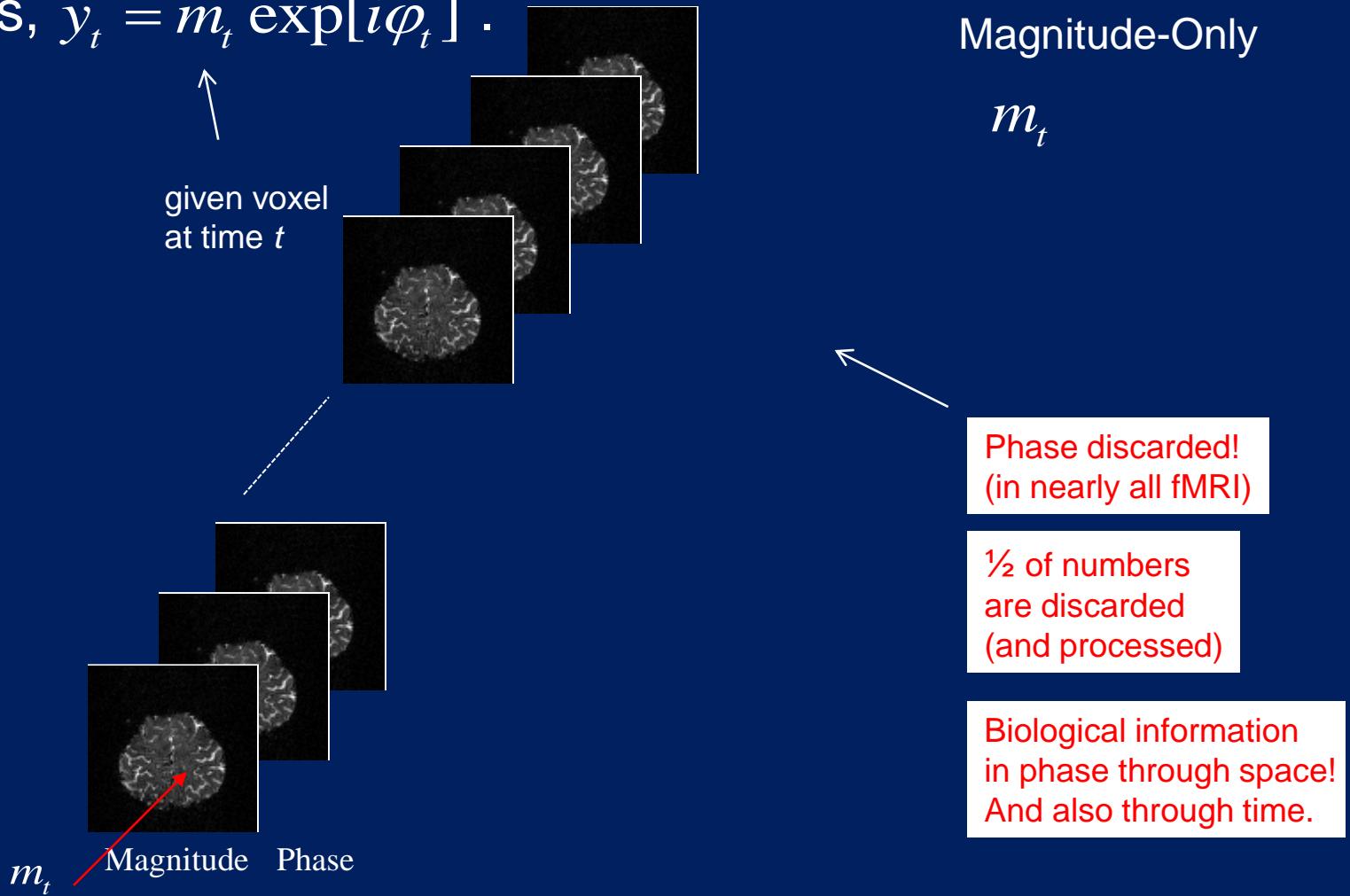
# Utilizing Induced Correlation: Magnitude-Only fMRI

Complex-valued images to magnitude and phase images and time series,  $y_t = m_t \exp[i\varphi_t]$ .



# Utilizing Induced Correlation: Magnitude-Only fMRI

Complex-valued images to magnitude and phase images and time series,  $y_t = m_t \exp[i\varphi_t]$ .



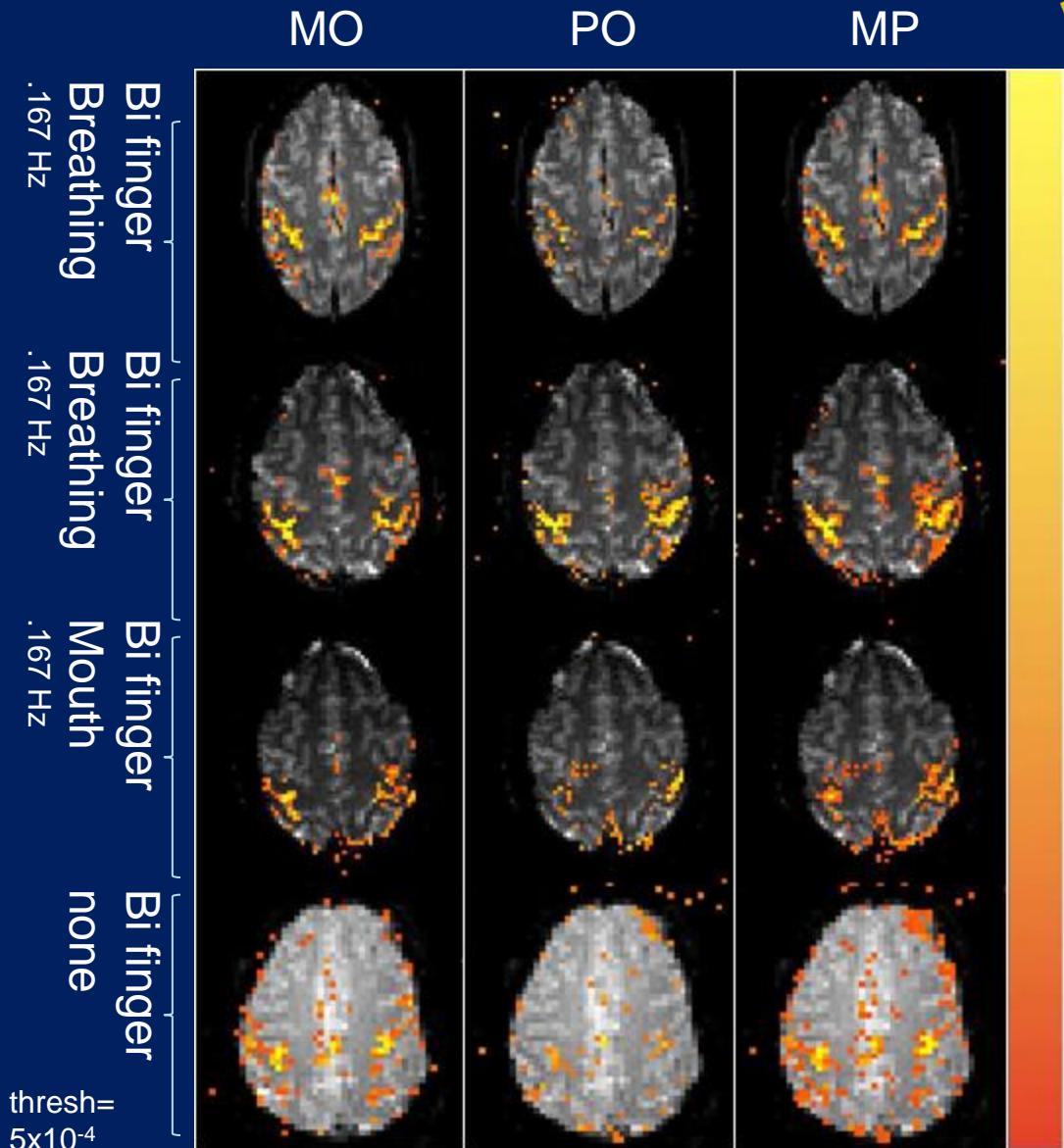
## Results: Independent

20s off+16x(8 s on 8 s off), 276 TRs  
 12 axial slices, 96 x 96, FOV = 24 cm  
 TH = 2.5 mm, TR = 1 s, TE = 34.6 ms  
 FA = 45°, BW = 125 kHz, ES = .708 ms

20s off+16x(8 s on 8 s off), 276 TRs  
 10 axial slices, 96 x 96, FOV = 24 cm  
 TH = 2.5 mm, TR = 1 s, TE = 42.8 ms  
 FA = 45°, BW = 125 kHz, ES = .768 ms

20s off+16x(8 s on 8 s off), 276 TRs  
 10 axial slices, 96 x 96, FOV = 24 cm,  
 TH = 2.5 mm, TR = 1 s, TE = 42.8 ms  
 FA = 45°, BW = 125 kHz, ES = .768 ms

20s off+10x(8 s on 8 s off), 180 TRs  
 9 axial slices, 64 x 64, FOV = 24 cm  
 TH = 3.8 mm, TR = 1 s, TE = 26.0 ms  
 FA = 45°, BW = 125 kHz, ES = .680 ms



Rowe: NIMG, 25:1310-1324, 2005.  
 Rowe: MRM, to appear, 2009.

Hahn, Nencka, Rowe: NIMG, 742-752, 2009.  
 Hahn, Nencka, Rowe: HBM, Online, 2011.

$$\Delta B_t = \frac{\arg \left( I_t \sum_{j=1}^n \left( \frac{I_j^*}{|I_j|} \right) \right)}{\gamma TE}$$

## Discussion:

When DATA ANALYSTS preprocess RESEARCHERS data,

THEY change the mean and covariance structure.

Many preprocessing operations have been shown

to modify or induce a correlation.

WE need to utilize this correlation in OUR analysis model!



# Thank You

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