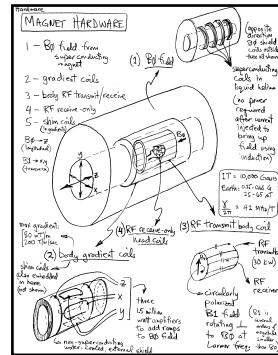


# Foundations of Neuroimaging

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13 Nov 2022

(124 pages)



<b>Hardware</b>	<b>2</b>
<b>Spin and Precession</b>	<b>3</b>
<b>Bloch equation</b>	<b>4</b>
<b>Vector Add, Multiply</b>	<b>5</b>
<b>Bloch and Precession, Bloch Solutions</b>	<b>6</b>
<b>RF Polarization, Signal Equation, Phase-sensitive Detection</b>	<b>13</b>
<b>Spin Echo, Stimulated Echo</b>	<b>17</b>
<b>Gradient Echo</b>	<b>23</b>
<b>Image Contrast</b>	<b>24</b>
<b>Signal-to-Noise</b>	<b>31</b>
<b>Fourier Transform</b>	<b>33</b>
<b>Slice Selection</b>	<b>42</b>
<b>Frequency Encoding</b>	<b>46</b>
<b>Phase Encoding</b>	<b>51</b>
<b>Image Reconstruction</b>	<b>55</b>
<b>Fast Spin Echo (FSE)</b>	<b>61</b>
<b>Fast Gradient Echo (FLASH)</b>	<b>64</b>
<b>Echo Planar Imaging (EPI)</b>	<b>67</b>
<b>Undersampling (GRAPPA, SENSE)</b>	<b>70</b>
<b>Multiband, EVI, Spiral</b>	<b>71</b>
<b>Artifacts (<math>B_0</math>, EPI, Gradients, Navigators)</b>	<b>76</b>
<b>Diffusion (DTI)</b>	<b>84</b>
<b>Perfusion (ASL, pCASL)</b>	<b>86</b>
<b>Off-Resonance, Spectroscopy</b>	<b>88</b>
<b>Phase-encoded Mapping</b>	<b>91</b>
<b>Convolution</b>	<b>92</b>
<b>General Linear Model (GLM)</b>	<b>93</b>
<b>Surface-Based Analysis (segment, filter, tessellate, unfold, align)</b>	<b>95</b>
<b>Origin of EEG/MEG</b>	<b>102</b>
<b>Grad, Div, Curl</b>	<b>104</b>
<b>Current Source Density (CSD)</b>	<b>105</b>
<b>Forward Solution for EEG, MEG</b>	<b>108</b>
<b>Inverse Solution (derive, constrain, normalize, use covariance)</b>	<b>113</b>

## Hardware

### MAGNET HARDWARE

1 -  $B\phi$  field from superconducting magnet

2 - gradient coils

3 - body RF transmit/receive

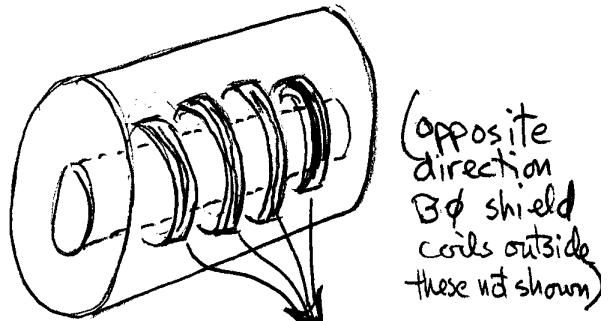
4 - RF receive-only

5 - shim coils  
(in gradients)

$B\phi \rightarrow z$   
(longitudinal)

$B_1 \rightarrow x, y$   
(transverse)

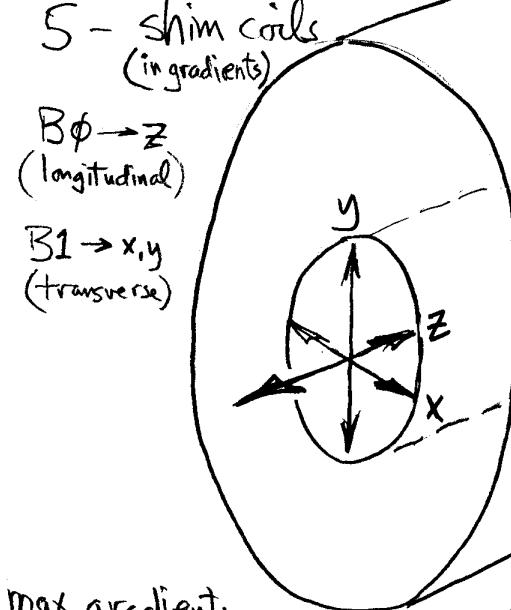
(1)  $B\phi$  field



(opposite direction  
 $B\phi$  shield  
coils outside  
these not shown)

Superconducting  
coils in  
liquid helium

(no power  
required  
after current  
injected to  
bring up  
field using  
induction)



max gradient:

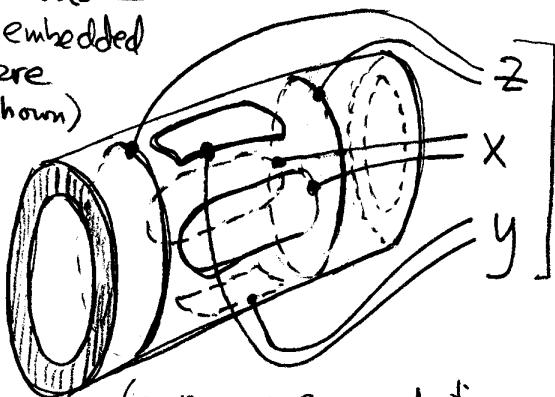
[ $80 \text{ mT/m}$   
 $200 \text{ T/m/sec}$ ]

(4) RF receive-only  
head coils

(3) RF transmit body coil

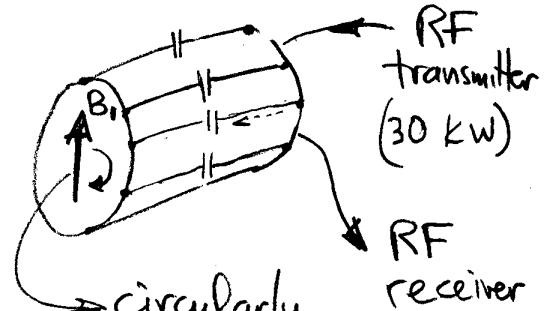
(2) body gradient coils

shim coils  
also embedded  
in here  
(not shown)



non-superconducting  
water-cooled, external shield

three  
1.5 million  
watt amplifiers  
to add ramps  
to  $B\phi$  field



circularly  
polarized  
 $B_1$  field  
rotating  $\perp$   
to  $B\phi$  at  
Larmor freq.

( $B_1$  is  
several  
orders of  
magnitude  
smaller  
than  $B\phi$ )

# SPIN & PRECESSION

- nuclei act like a spinning sphere of matter with an embedded equatorial charge (nuclei w/ odd atomic weight or odd proton numbers)
- moving charge creates magnetic field



classical picture

- current loop from spinning charge (right-hand rule)
- N.B.: classically this would cause EM radiation, spindown

- Stern-Gerlach experiment

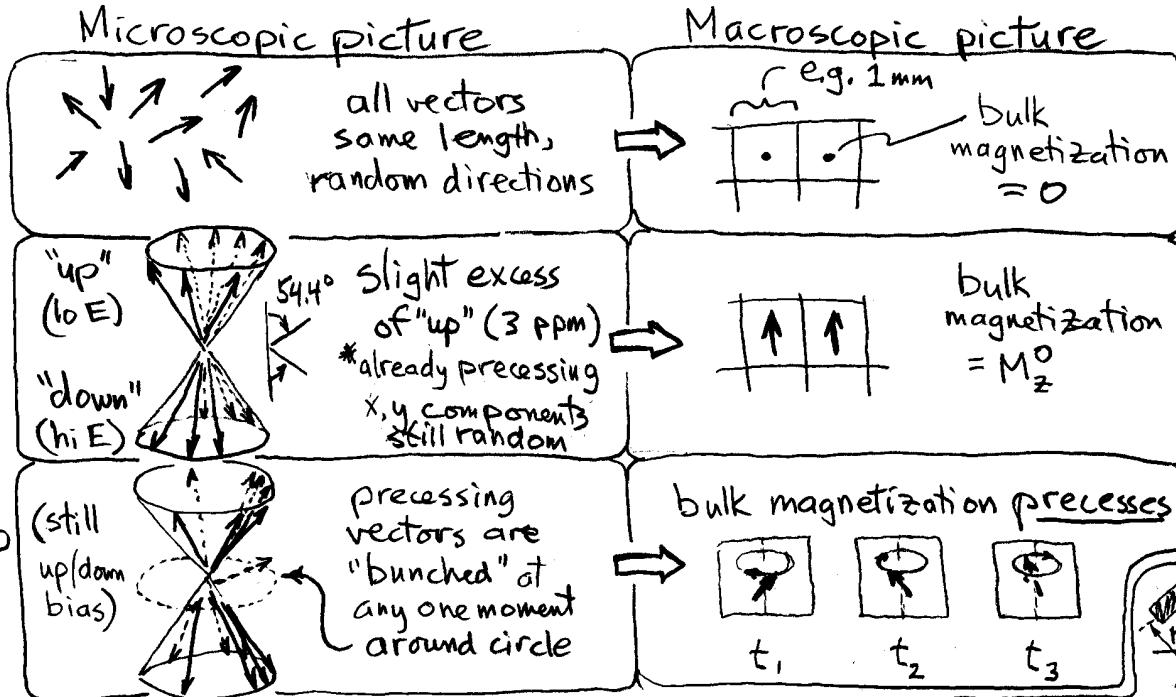


pass silver atoms thru strong mag. field  $\rightarrow$  split into just 2 beams

no strong magnetic field  
 $B\phi = 0$

Strong magnetic field,  $B\phi =$

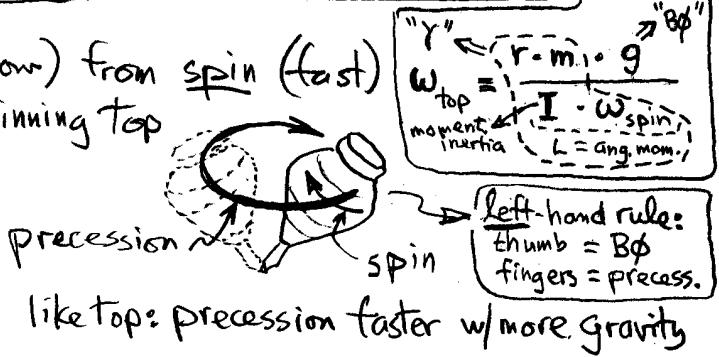
Strong  $B\phi$   
plus oscillating  $B_1$



## Precession

- distinguish precession (slow) from spin (fast)
- treat classically, like spinning top

$$\frac{2\pi f_0}{\omega_0} = \frac{\omega_0}{\gamma B_0} = \frac{\text{Larmor freq. (e.g. } 63 \text{ MHz)}}{\text{Same in radians/sec}} \frac{\text{gyro-magnetic ratio}}{\text{static field (e.g. } 1.5 \text{ T)}}$$



- bulk equilibrium magnetization (parallel to  $B_0$ )

$$M_z^0 = |\vec{M}| = \frac{\gamma^2 h^2 B_0 N_s}{4 K T_s}$$

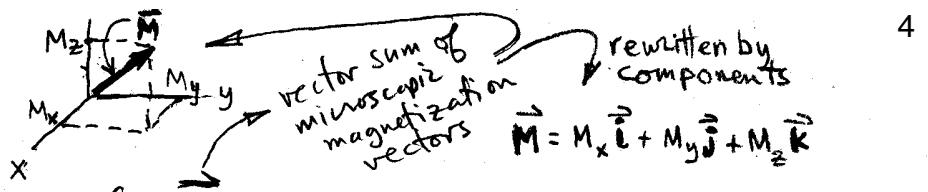
where  $I = \pm \frac{1}{2}$     $\frac{I(I+1)}{3}$

two non-constants

- $\gamma$  = gyromagnetic ratio
- $h$  = Planck's const.
- $B_0 \rightarrow$  i.e.,  $M_z^0$  proportional to  $B_0$  strength
- $N_s \rightarrow$  i.e.,  $M_z^0$  proportional to number spins
- $K =$  Boltzmann const.
- $T_s =$  abs. temperature sample

Block-1

# BLOCH EQUATION



- time-dependent behavior of  $\vec{M}$  in the presence

of an applied magnetic field (excitation & relaxation)

$$\text{(laboratory frame)} \quad \frac{d\vec{M}}{dt} = \vec{M} \times \gamma \vec{B} - \frac{M_x \vec{i} + M_y \vec{j}}{T_2} - \frac{(M_z - M_z^0) \vec{k}}{T_1}$$

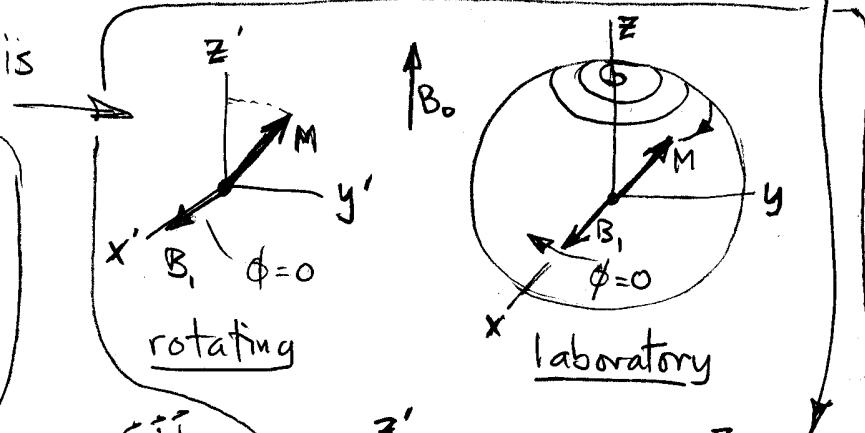
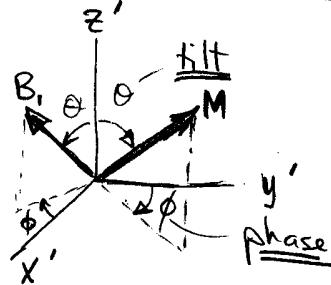
precession:  $B = B_0$   
 excitation:  $B = B_0 + B_1$

existing mag vector sum  
 applied field (typically rotating)

change in mag vector

- in the Larmor-rotating coordinate system, a tilt w/o a phase shift for a standard  $B_1$ , excitation is rotation around  $x$ -axis

general case  
rotating  
(distinguish  $B_1$  &  $M$ !)

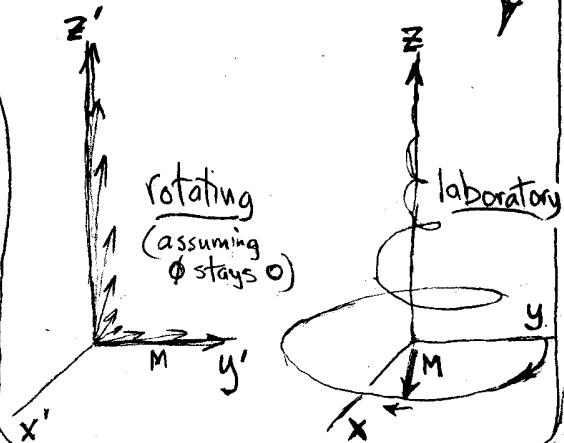


- longitudinal and transverse relaxations

$$\frac{dM_z(t)}{dt} = -\frac{M_z(t) - M_z^0}{T_1}$$

from Bloch equation  
after dropping applied field term

$$\frac{dM_{x'y'}(t)}{dt} = -\frac{M_{x'y'}(t)}{T_2}$$



- solution to equations above: time-dependent free precession eq's

Lab frame: same!

$$M_z(t) = M_z^0 (1 - e^{-t/T_1}) + M_z'(0+) e^{-t/T_1}$$

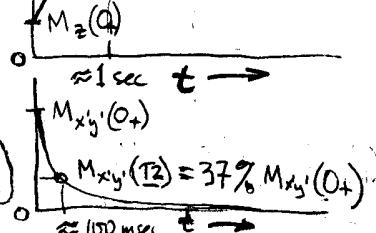
$$[\text{equiv}] = M_z^0 + [M_z'(0+) - M_z^0] e^{-t/T_1}$$

$$M_z'(T_1) = 63\% M_z^0$$

$M_L$   
(rotating frame)

$$M_{x'y'}(t) = M_{x'y'}(0+) e^{-t/T_2}$$

$M_L, M_T$  at time immed. after pulse



Lab frame: times  $e^{-i\omega t}$

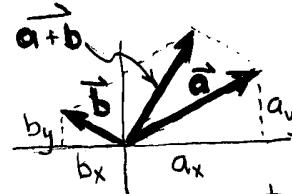
Block-1c

## VECTOR ADD, MULTIPLY

- adding vectors is easy

$$\vec{c} = \vec{a} + \vec{b} = [a_x + b_x, a_y + b_y]$$

- [ - just add components (vector) ]
- [ - applies to complex numbers ]



- generalizes to any D

get length  
 $\|\vec{c}\| = \sqrt{(a_x + b_x)^2 + (a_y + b_y)^2}$

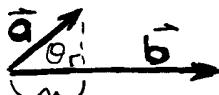
- multiple ways to multiply vectors: here are 3

### dot product

(= inner product)

(= "scaled projection onto")

$$c = \vec{a} \cdot \vec{b} = [b_x \ b_y \ b_z] \begin{bmatrix} a_x \\ a_y \\ a_z \end{bmatrix} = a_x b_x + a_y b_y + a_z b_z$$



$$p = \|\vec{a}\| \cos \theta$$

$$c = p \|\vec{b}\| \quad \text{if } \|\vec{b}\| = 1$$

(scalar)  
 - generalizes to any D

N.B. equals:  $\vec{a} \cdot \vec{a}$

$$\sqrt{a_x^2 + a_y^2 + a_z^2}$$

length of  $\vec{a}$

$$c = \|\vec{a}\| \|\vec{b}\| \cos \theta$$

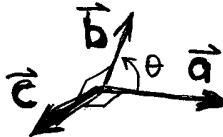
↳ zero if  $\vec{a}, \vec{b}$  orthogonal

### cross product

(= outer product)

(can be generalized: see "geometric algebra")

$$\vec{c} = \vec{a} \times \vec{b} = \begin{bmatrix} 0 & b_z & -b_y \\ -b_z & 0 & b_x \\ b_y & -b_x & 0 \end{bmatrix} \begin{bmatrix} a_x \\ a_y \\ a_z \end{bmatrix} = [a_y b_z - a_z b_y, a_z b_x - a_x b_z, a_x b_y - a_y b_x]$$



skew-symmetry:  $A^T = -A$

right hand rule: curl fingers from  $\vec{a}$  to  $\vec{b}$ : thumb is  $\vec{c}$

- unique  
orthogonal  
specific to 3D

geometric algebra: bivector  $\vec{b}$   
plane area

$$\|\vec{c}\| = \|\vec{a}\| \|\vec{b}\| \sin \theta$$

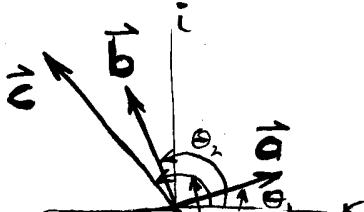
↳ max if orthogonal

### complex multiply

(see also quaternions, geometric algebra generalization)

$$\vec{c} = \vec{a} \cdot \vec{b} = [b_x \ -b_y] \begin{bmatrix} a_x \\ a_y \end{bmatrix} = [a_x b_x - a_y b_y, a_x b_y + a_y b_x]$$

(vector)



- angles add  
- magnitudes multiply

sum of angles:  $\theta_1 + \theta_2$

- specific to 2D

l.c.  
not affected  
by angle  
between

$$\|\vec{c}\| = \|\vec{a}\| \|\vec{b}\|$$

↳ like real nums

Block 1a

# EFFECTS OF $\vec{M}$ , $\vec{B}$ , and $\theta$ ON PRECESSION FREQ.

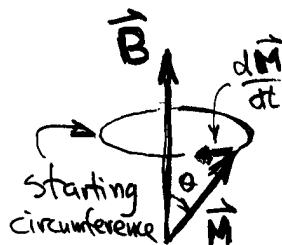
**Block 1st term**

$$\frac{d\vec{M}}{dt} = \vec{\gamma} \vec{B} \times \vec{M}$$



cross prod. properties review

$$\left\| \frac{d\vec{M}}{dt} \right\| = \left\| \vec{M} \right\| \cdot \left\| \vec{\gamma} \vec{B} \right\| \cdot \sin \theta$$

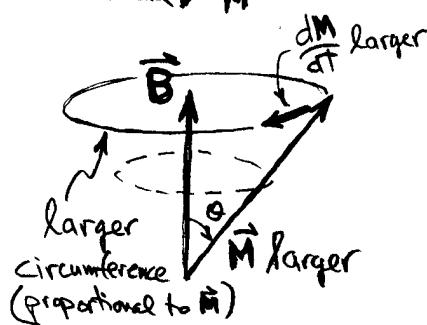


Starting condition

↳ now see effects of changing  $\left\| \vec{M} \right\|$ ,  $\left\| \vec{B} \right\|$ ,  $\theta$

N.B. Length

vectors define this

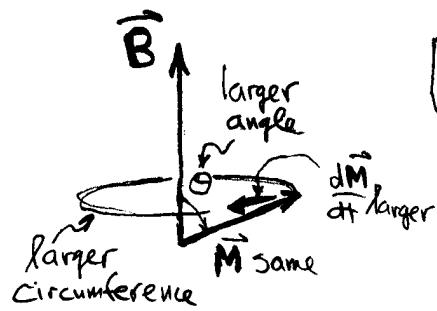


Change  $\vec{M}$  length

↳  $\frac{d\vec{M}}{dt}$  proportionally larger, so cancels effect of larger  $\vec{M}$

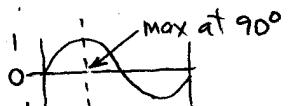
b/c  $\vec{M}$  on both sides eq.

↳ same precession freq. as starting cond.



Change  $\theta$  between  $\vec{M}$  and  $\vec{B}$

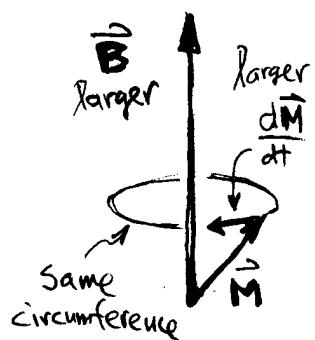
↳  $\frac{d\vec{M}}{dt}$  goes up (then down) as  $\sin \theta$  ↳ cross prod.



but circumference also goes up as  $\sin \theta$ , cancelling again

↳ opposite =  $\sin \theta$  = radius so circumference proportional to  $\sin \theta$

↳ same precession freq.



Change  $\vec{B}$  length

↳  $\frac{d\vec{M}}{dt}$  goes up, proportional to  $\vec{B}$

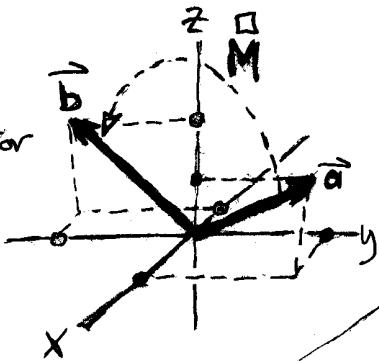
but circumference is same at starting cond.

↳ increased precession freq. ( $\omega = \gamma B \theta$ )

# Block 1e SIMPLE MATRIX OPERATIONS

## Basic idea

- a matrix  $\begin{bmatrix} \text{rotates} \\ \text{scales} \end{bmatrix}$  a vector
- $$\vec{b} = M \vec{a}$$



## 3D example

$$\begin{bmatrix} b_x \\ b_y \\ b_z \end{bmatrix} = \begin{bmatrix} M_{11} & M_{12} & M_{13} \\ M_{21} & M_{22} & M_{23} \\ M_{31} & M_{32} & M_{33} \end{bmatrix} \begin{bmatrix} a_x \\ a_y \\ a_z \end{bmatrix}$$

## Add translate (after rotate/scale)

- commonly used "hack" for aligning vols

$$\begin{aligned} b_x &= M_{11}a_x + M_{12}a_y + M_{13}a_z \\ b_y &= M_{21}a_x + M_{22}a_y + M_{23}a_z \\ b_z &= M_{31}a_x + M_{32}a_y + M_{33}a_z \end{aligned}$$

- a 4D matrix  $\begin{bmatrix} \text{rotates/scales} \\ \text{then} \\ \text{translates} \end{bmatrix}$  a 3D vector  $(4^{\text{th}} D=1)$

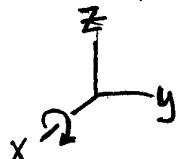
- N.B.: Have to keep track of order!!

rotate/scale then trans  $\neq$  trans then rot/scale  
change rot component: untranslate, rot, retranslate

$$\begin{bmatrix} b_x \\ b_y \\ b_z \\ 1 \end{bmatrix} = \begin{bmatrix} 3 \times 3 \\ \text{rot/scale} \\ \text{translate?} \\ dx \\ dy \\ dz \\ 0 & 0 & 0 & 1 \end{bmatrix} \begin{bmatrix} a_x \\ a_y \\ a_z \\ 1 \end{bmatrix}$$

## 3 special cases (3D): rotate around each major axis without changing length (Scale = 1.0)

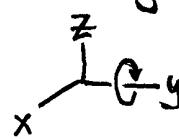
- rotate around x-axis:



$$R_x(\alpha) = \begin{bmatrix} 1 & 0 & 0 \\ 0 & \cos\alpha & \sin\alpha \\ 0 & -\sin\alpha & \cos\alpha \end{bmatrix}$$

e.g.,  $90^\circ$  flip

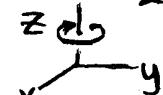
- rotate around y-axis:



$$R_y(\alpha) = \begin{bmatrix} \cos\alpha & 0 & -\sin\alpha \\ 0 & 1 & 0 \\ \sin\alpha & 0 & \cos\alpha \end{bmatrix}$$

e.g.,  $180^\circ$  flip  
to avoid add  
 $180^\circ$  phase after  
 $90^\circ$  flip on x'

- rotate around z-axis:

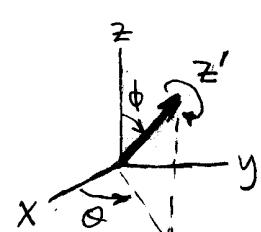


$$R_z(\alpha) = \begin{bmatrix} \cos\alpha & \sin\alpha & 0 \\ -\sin\alpha & \cos\alpha & 0 \\ 0 & 0 & 1 \end{bmatrix}$$

e.g., precession  
with  $B\phi$   
along  $z'$

## General case

- rotate around general  $z'$ -axis:



$$R_{z'}(\alpha) = R_z(-\theta) R_y(-\phi) R_z(\alpha) R_y(\phi) R_z(\theta)$$

unrot. axis to z  
z-rot.  
re-rot.

(quaternions are  
more efficient)

Bloch - 1b

# SOLUTIONS TO SIMPLE DIFFERENTIAL EQ.

diff. eq:

$$dM_{x'y'}(t) = -\frac{M_{x'y'}(t)}{T_2}$$

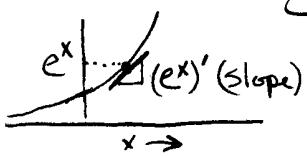
solution:

$$M_{x'y'}(t) = M_{x'y'}(0_+) \cdot e^{-t/T_2}$$

?? How this works,  
and where  $M_{x'y'}(0_+)$   
comes from...

Goal: 1) find eq. whose derivative satisfies diff. eq.

2) also find sol'n (one of many) that passes thru int condition



Since our diff. eq. is: derivative of funct. = const. same funct

try exponential, since derivative  $(e^x) = e^x$  ☺

diff eq.

$$M'(t) = -\frac{1}{T_2} \cdot M(t)$$

N.B. this function is the "unknown" like the x in  $x + 1 = 3$

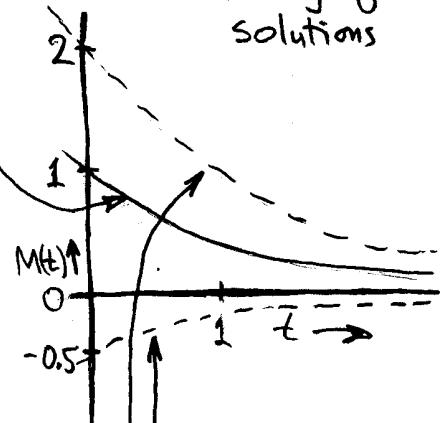
one sol'n

$$M(t) = e^{-t/T_2} = e^{-t/T_2}$$

family of solutions

take deriv.  
to check

$$M'(t) = -\frac{1}{T_2} \cdot e^{-t/T_2} \cdot M(t)$$



diff eq.

$$M'(t) = -\frac{1}{T_2} \cdot M(t)$$

another  
sol'n

$$M(t) = \text{const. } e^{-t/T_2}$$

so: any  
const is OK!

take deriv.  
to check

$$M'(t) = -\frac{1}{T_2} \cdot \text{const. } e^{-t/T_2}$$

for ex.:  
const = 2  
const = -0.5

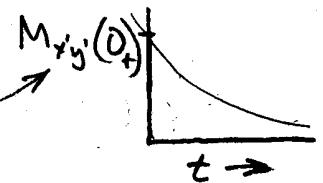
const is "initial condition"

information  
added to sol'n  
(not from  
diff eq!)

$$\text{const} = M_{x'y'}(0_+)$$

magnetization immed.  
after RF (B1) ends

$$M(t) = M_{x'y'}(0_+) \cdot e^{-t/T_2}$$



Bloch-1c2

# VERIFY SOLUTION TO T1 REGROWTH

- slightly more complex T1 sol'n compared to T2 sol'n

**T2 sol'n verify (from prev.)**

**T1 solution verify**

$$\frac{dM}{dt} = \frac{M_{xy}}{\tau_2}$$

original diff eq.

$$\frac{dM}{dt} = -\frac{(M_z - M_z^0)}{\tau_1}$$

$$M'(t) = -\frac{1}{\tau_2} \cdot M(t)$$

make unknown funct  $M(t)$  more visible

$$M'(t) = -\frac{1}{\tau_1} (M(t) - M_z^0)$$

init cond.

$$M(t) = [M_{xy}(0_+)] e^{-t/\tau_2}$$

proposed solution

$$\begin{aligned} M(t) &= M_z^0 (1 - e^{-t/\tau_1}) + M_z(0_+) e^{-t/\tau_1} \\ &= \underbrace{M_z^0}_{\text{const}} - \underbrace{M_z^0 e^{-t/\tau_1}}_{\substack{\text{chain rule} \\ \text{as before}}} + \underbrace{M_z(0_+) e^{-t/\tau_1}}_{\text{chain rule}} \end{aligned}$$

$$M'(t) = -\frac{1}{\tau_2} M_{xy}(0_+) e^{-t/\tau_2}$$

test by take deriv.

$$\begin{aligned} M'(t) &= 0 + \frac{1}{\tau_2} M_z^0 e^{-t/\tau_1} - \frac{1}{\tau_1} M_z(0_+) e^{-t/\tau_1} \\ &= \left[ -\frac{1}{\tau_1} (-M_z^0 e^{-t/\tau_1} + M_z(0_+) e^{-t/\tau_1}) \right] \end{aligned}$$

- derivative in original T1 eq. says  $M(t)$  minus  $M_z^0$

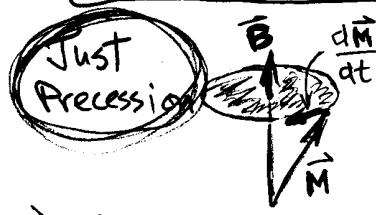
$$M'(t) = -\frac{1}{\tau_1} (M(t) - M_z^0)$$

$$\text{solution} \rightarrow \left[ M_z^0 - M_z^0 e^{-t/\tau_1} + M_z(0_+) e^{-t/\tau_1} \right]$$

- which equals our re-calculated derivative:

$$M'(t) = -\frac{1}{\tau_1} (-M_z^0 e^{-t/\tau_1} + M_z(0_+) e^{-t/\tau_1})$$

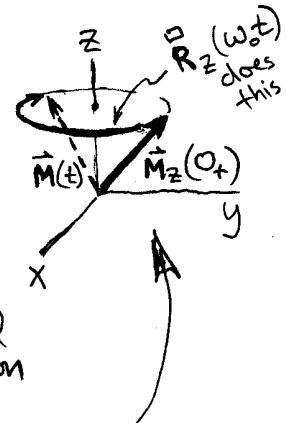
Bloch = 1d

**BLOCH EQ. - MATRIX VERSION**Differential Eq.:

$$\frac{d\vec{M}}{dt} = \begin{bmatrix} \frac{dM_x}{dt} \\ \frac{dM_y}{dt} \\ \frac{dM_z}{dt} \end{bmatrix} = \begin{bmatrix} 0 & \gamma B \phi & 0 \\ -\gamma B \phi & 0 & 0 \\ 0 & 0 & 0 \end{bmatrix} \begin{bmatrix} M_x \\ M_y \\ M_z \end{bmatrix}$$

assume only z component of  $\vec{B}\phi$  present

$$\vec{a} \times \vec{b} = \begin{bmatrix} 0 & b_2 & -b_3 \\ -b_2 & 0 & b_1 \\ b_3 & -b_1 & 0 \end{bmatrix} \begin{bmatrix} a_x \\ a_y \\ a_z \end{bmatrix}$$

Solution:

$$\vec{M}(t) = \begin{bmatrix} M_x(t) \\ M_y(t) \\ M_z(t) \end{bmatrix} = \begin{bmatrix} \cos w_0 t & \sin w_0 t & 0 \\ -\sin w_0 t & \cos w_0 t & 0 \\ 0 & 0 & 1 \end{bmatrix} \begin{bmatrix} M_x(0_+) \\ M_y(0_+) \\ M_z(0_+) \end{bmatrix} = \boxed{R_z(w_0 t)} \vec{M}(0_+)$$

Include Relaxation

$$\frac{d\vec{M}}{dt} = \vec{M} \times \gamma \vec{B} \phi - \frac{M_x \vec{i} + M_y \vec{j}}{T_2} - \frac{(M_z - M_z^0) \vec{k}}{T_1}$$

Differential Eq.:

$$\frac{d\vec{M}}{dt} = \begin{bmatrix} -1/T_2 & \gamma B \phi & 0 \\ -\gamma B \phi & -1/T_2 & 0 \\ 0 & 0 & -1/T_1 \end{bmatrix} \begin{bmatrix} M_x \\ M_y \\ M_z \end{bmatrix} + \begin{bmatrix} 0 \\ 0 \\ M_z^0/T_1 \end{bmatrix}$$

Solution:

$$\vec{M}(t) = \begin{bmatrix} e^{+t/T_2} & 0 & 0 \\ 0 & e^{-t/T_2} & 0 \\ 0 & 0 & e^{-t/T_1} \end{bmatrix} \begin{bmatrix} \cos w_0 t & \sin w_0 t & 0 \\ -\sin w_0 t & \cos w_0 t & 0 \\ 0 & 0 & 1 \end{bmatrix} \begin{bmatrix} M_x(0_+) \\ M_y(0_+) \\ M_z(0_+) \end{bmatrix} + \begin{bmatrix} 0 \\ 0 \\ M_z(0_+) (1 - e^{-t/T_1}) \end{bmatrix}$$

(as above)

Bloch-1f

# EXCITATION IN THE ROTATING FRAME

- original Bloch eq. in laboratory frame

$$\frac{d\vec{M}_{lab}}{dt} = \vec{M} \times \gamma \vec{B}$$

B<sub>0</sub>  
B<sub>1</sub>  
gradients

Remember:

$$\vec{a} \times \vec{b} = \begin{bmatrix} 0 & b_2 & -b_1 \\ -b_2 & 0 & b_3 \\ b_1 & -b_3 & 0 \end{bmatrix} \begin{bmatrix} a_x \\ a_y \\ a_z \end{bmatrix}$$

- add on-resonance B<sub>1</sub> to T<sub>1</sub>φ

$$\vec{B} = B_1(t) (\cos \omega_0 t \hat{i} - \sin \omega_0 t \hat{j}) + B_0 \hat{k}$$

$\gamma B_0 \hat{k}$   
 $\gamma B_1(t)$

lab frame < on-resonance  
\* basic excite  
↳ matrix version

$$\frac{d\vec{M}}{dt} = \begin{bmatrix} \frac{dM_x'}{dt} \\ \frac{dM_y'}{dt} \\ \frac{dM_z'}{dt} \end{bmatrix} = \begin{bmatrix} 0 \\ -\omega_0 \\ 0 \end{bmatrix}$$

$\xrightarrow{\text{b/c}} \begin{bmatrix} -W_1(t) \sin \omega_0 t & -W_1(t) \cos \omega_0 t \\ \hookrightarrow B_1 y \text{ comp.} & \hookrightarrow B_1 z \text{ comp.} \end{bmatrix}$

$\omega_0$   
 $W_1(t) \sin \omega_0 t$   
 $W_1(t) \cos \omega_0 t$   
 $0$   
 $M_x'$   
 $M_y'$   
 $M_z'$

- Substitution to convert to the rotating frame

$$\begin{bmatrix} \vec{M} \\ \vec{B} \end{bmatrix} = \begin{bmatrix} \square \\ \square \end{bmatrix} R_2(\omega_0 t) \cdot \begin{bmatrix} \vec{M}_{rot} \\ \vec{B}_{rot} \end{bmatrix}$$

matrix multiply  
"Subtract off" rotating frame from both  $\vec{M}$  and  $\vec{B}$

- after substitution only off-resonance appears as residual B<sub>0</sub> (B<sub>2</sub>) (see off-res notes page)

$$\frac{d\vec{M}_{rot}}{dt} = \vec{M}_{rot} \times \gamma \vec{B}_{eff}$$

off-res (appears as B<sub>0</sub>)  
B<sub>1</sub> gradients

rotating frame < on-resonance  
\* basic excite, B<sub>1x</sub>-only no gradient  
↳ removes  $\omega_0$ , cos/sin

$$\frac{d\vec{M}_{rot}}{dt} = \begin{bmatrix} 0 & 0 & 0 \\ 0 & 0 & \omega_1(t) \\ 0 & -\omega_1(t) & 0 \end{bmatrix} \begin{bmatrix} M_x' \\ M_y' \\ M_z' \end{bmatrix}$$

$\hookrightarrow B_{1y} = 0$

rotating frame - off-resonance  
\* general, B<sub>1x</sub>-only incl gradients

$$\frac{d\vec{M}_{rot}}{dt} = \begin{bmatrix} 0 & \omega_0 - \omega + \omega(2) & 0 \\ -( \omega_0 - \omega + \omega(2)) & 0 & \omega_1(t) \\ 0 & -\omega_1(t) & 0 \end{bmatrix} \begin{bmatrix} M_x' \\ M_y' \\ M_z' \end{bmatrix}$$

$\xrightarrow{\text{off-res.}}$  gradient

gradient:  $\omega(z) = \gamma G_z z$   
off-res: appears as residual B<sub>z</sub>, lifting  $\vec{B}_1$  vect. out of x-y plane

This means  $\vec{M}$  vect. update will contain component that rotates  $\vec{M}$  around z-axis (in rotating coords  $\equiv$  phase)

rotating frame < on-resonance  
\* small tip approx.  
incl gradient  
small tip  $\frac{M_z}{\frac{dM}{dt}} \approx 0$   
 $\vec{B} \approx 0$

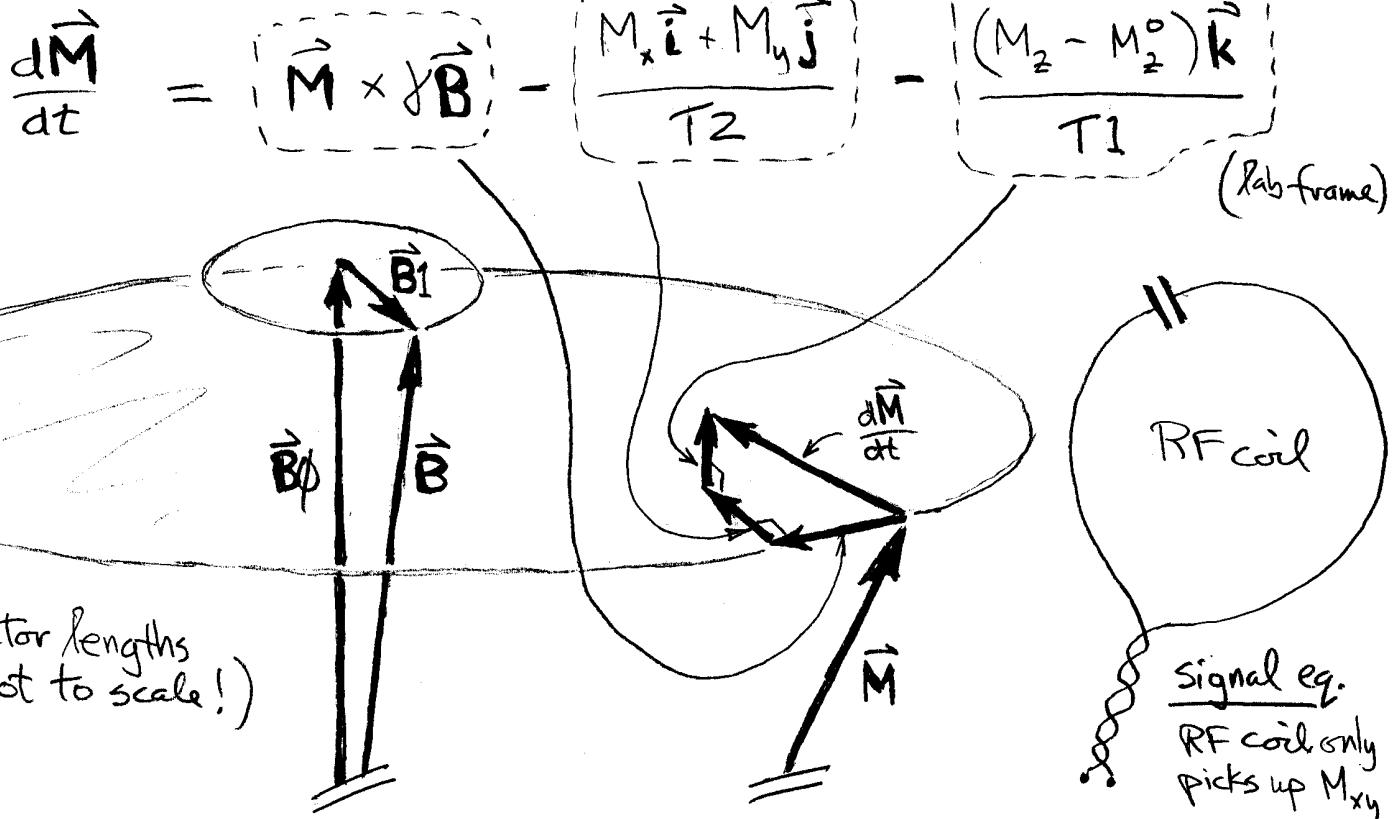
$$\frac{d\vec{M}_{rot}}{dt} = \begin{bmatrix} 0 & \omega(2) & 0 \\ -\omega(2) & 0 & \omega_1(t) \\ 0 & 0 & 0 \end{bmatrix} \begin{bmatrix} M_x' \\ M_y' \\ M_z' \end{bmatrix}$$

$\hookrightarrow$  small tip  $\Rightarrow$  easier to solve!  
small tip

zeros this line in matrix

Bloch-1g

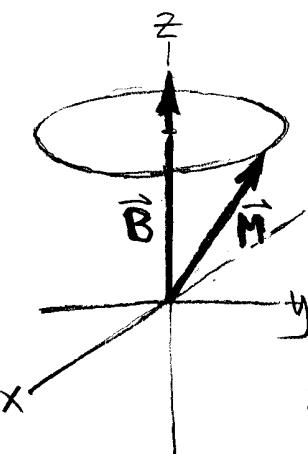
# BLOCH EQ. SUMMARY



- full lab frame picture is complex:
  - 3 components of  $\frac{d\vec{M}}{dt}$  update vector
  - Larmor freq. component 7-9 orders magnitude larger than  $T_2, T_1$  decay
  - $\vec{B}_1$  is also rapidly wiggling
- conceptual simplification in 4 stages:

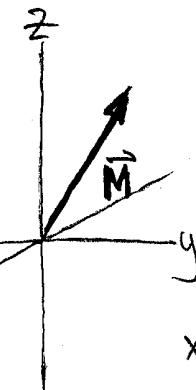
1) lab frame

- just precession

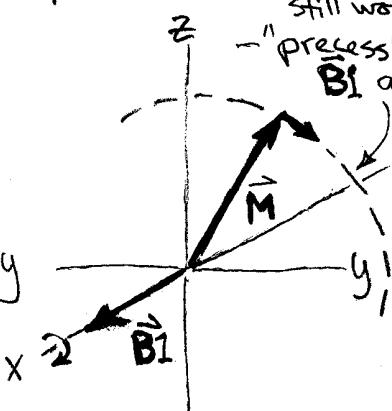


2) rotating frame

- $\vec{M}$  stopped
- that is,  $B_{\phi} = 0$

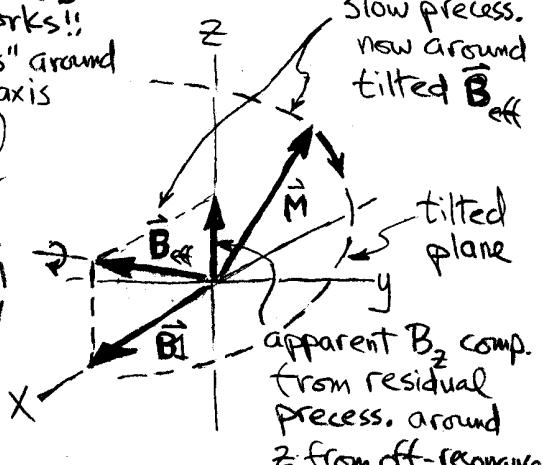
3) add  $\vec{B}_1$ 

- $\vec{B}_1$  also stopped!
- but  $\vec{M} \times \gamma \vec{B}$  still works!!
- "precess" around  $\vec{B}_1$  axis



4) off-resonance

- slow precess. now around tilted  $\vec{B}_{\text{eff}}$
- tilted plane
- apparent  $B_z$  comp. from residual precess. around z from off-resonance



Block-2

## RF FIELD POLARIZATION

- polarization (change of direction) of magnetic field (vs. electric field)

- Linearly polarized field

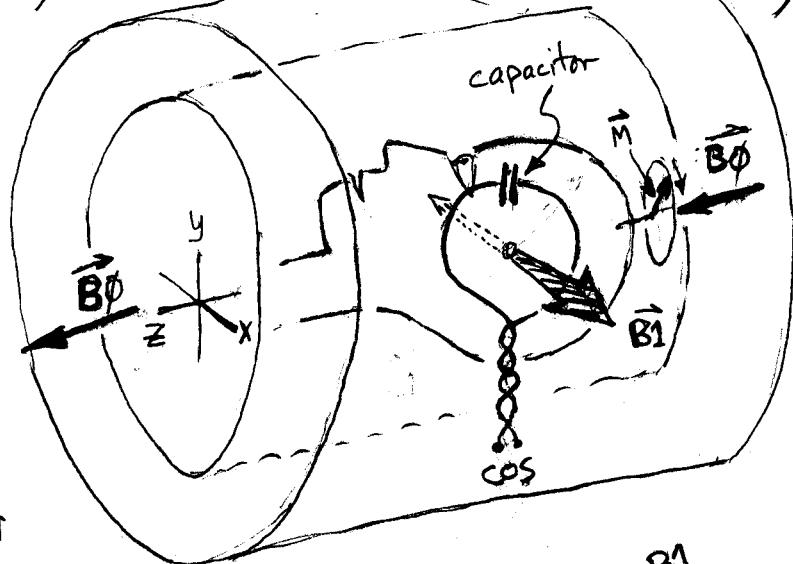
$$\vec{B}_1(t) = B_1 \cdot \cos \omega t \hat{x}$$

rad/sec    sec  
magn. strength: {-1, 1} · 1

- N.B.:  $\vec{B}_1$  adds to much larger  $\vec{B}_0$

$$\vec{B}_0 \rightarrow \vec{B}_0 + \vec{B}_1 \quad \text{one instant in time}$$

$$\vec{B}_0 + \vec{B}_1 \rightarrow \text{wiggles L/R across time}$$

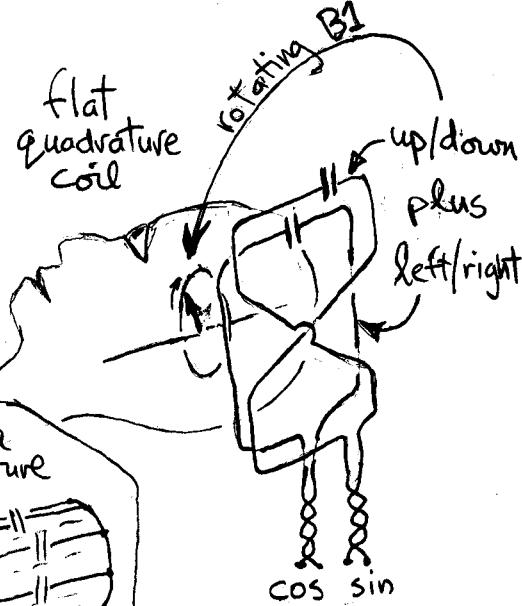
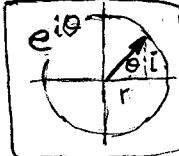


- Circularly polarized field (quadrature)

$$\vec{B}_1^{\text{circ}}(t) = B_1 (\cos \omega t \hat{x} - \sin \omega t \hat{y})$$

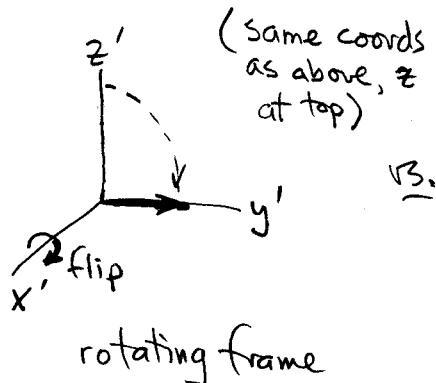
$$= B_1 \cdot e^{-i\omega t}$$

$\vec{B}_0 + \vec{B}_1$  rotates

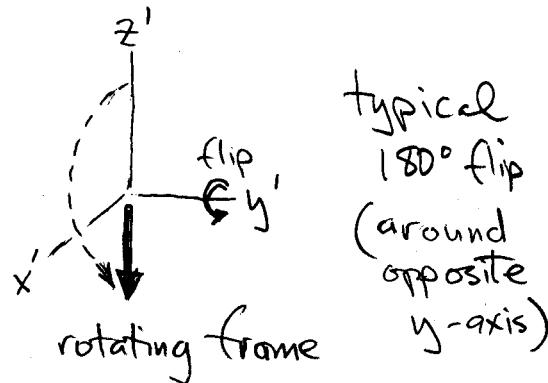


- in the rotating coordinate syst, flipping around x-axis vs. y-axis is just difference in phase of RF stim

typical  
90° flip  
(around  
x-axis)

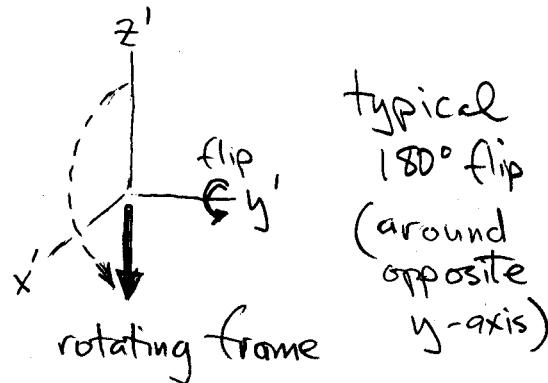


B.



typical  
180° flip  
(around  
opposite  
y-axis)

180° flip  
req's  
~6X power  
of 90°



Bloch-3

# SIGNAL EQUATION

$$\Phi(t) = \int_{\text{obj}} \vec{B}(\vec{r}) \cdot \vec{M}(\vec{r}, t) d\vec{r}$$

one "voxel"

for a particular instant in time vs. Bloch  $M_{x,y}$  which is change of  $M$  w/t

i.e., projection of mag. vector at each point onto coil magnetic field direction at each point, summed across object

magnetic "flux" thru coil  $\rightarrow$  scalar (integral mag. field perpendicular to area)

obj magn. field detected (generated) by coil geometry at each point in object

local magnetization of object (time-dependent)

position:  $\vec{r} \rightarrow x, y, z$

$$V(t) = - \frac{\partial \Phi(t)}{\partial t} = - \frac{\partial}{\partial t} \int_{\text{obj}} \vec{B}(\vec{r}) \cdot \vec{M}(\vec{r}, t) d\vec{r}$$

Faraday Law of Induction

(const retained in deriv.)

- evaluate using free precession eqs. (solution to Bloch) ignoring relaxation

- rewrite w/ complex notation w/ time-dependence from lab frame Bloch

- ignore change in z-comp.  $\vec{M}$  because so slow  $\rightarrow$  i.e., we only see  $M_{xy}$ , not  $M_z$
- substitute  $\vec{M}(t)$  with lab frame  $\vec{M}_{xy}(t) = M_{xy}(0) e^{-T_2 t} e^{-i\omega t}$
- Simplify:
  - ignore decay (assume this  $t=0$ )  $\rightarrow$  so equals 1 (scalar)
  - assume phase-sensitive detection  $\rightarrow$   $\Delta\omega$  (difference from  $\omega_0$ )  $\rightarrow$  rotating frame makes data complex

$$\vec{S}(t) = \int_{\text{obj}} B_{xy}(\vec{r}) M_{xy}(\vec{r}, 0) e^{-i\Delta\omega(\vec{r})t} d\vec{r}$$

omit receive and init excite flip phase offsets

complex: 2  $V(t)$ 's

demodulate

even from one coil!

laboratory frame Bloch solutions:

$M_L \rightarrow$  same

$M_T = M_{xy}(0) e^{-(T_2 - i\omega)t}$

assume homogeneous (ignore)

lab frame transverse magnetization at time  $t=0$

$M_{xy}$  is now scalar, direction is here

$\Delta\omega(\vec{r})$  freq diff from  $\omega_0$

$\omega_0$  subtracted off by PSD

sum across object

spatially-dependent resonant freq in rotating frame — i.e. after subtraction  $\omega_0 = \gamma B_0$

$$\vec{S}(t) = \int_{\text{obj}} M_{xy}(\vec{r}, 0) e^{-i\Delta\omega(\vec{r})t} d\vec{r}$$

i.e., at a single time point, RF signal is vector sum across object of local transverse magnetization vectors

standard Signal expression

phase angle in rotating frame

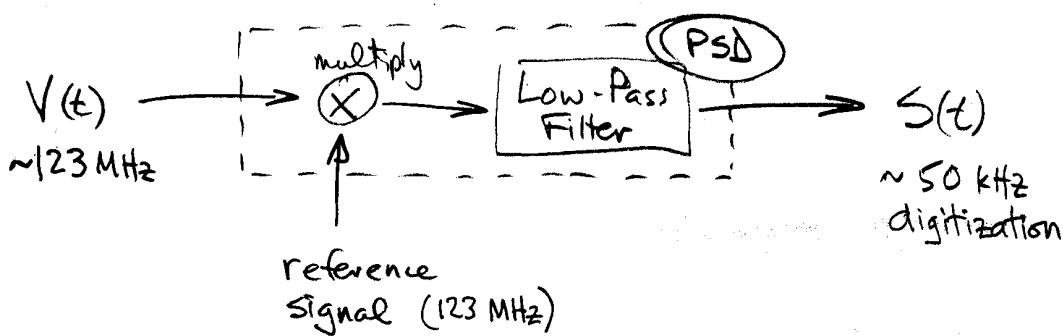
$\omega t = \frac{\text{radians}}{\text{sec}} \times \text{sec} = \text{radians}$  ( $\phi = \int \omega dt$ )

getting difference converts lab  $\rightarrow$  rotating

Block-4

# PHASE-SENSITIVE DETECTION

how we get rotating frame



- method for moving very high frequency Larmor oscillations down to tractable frequency range

demodulated signal  $\propto$  RF coil signal - reference (transmitter)

$$\propto \sin[(\omega_0 + \delta\omega)t] \cdot \sin[\omega_0 t]$$

↓  
trig identity

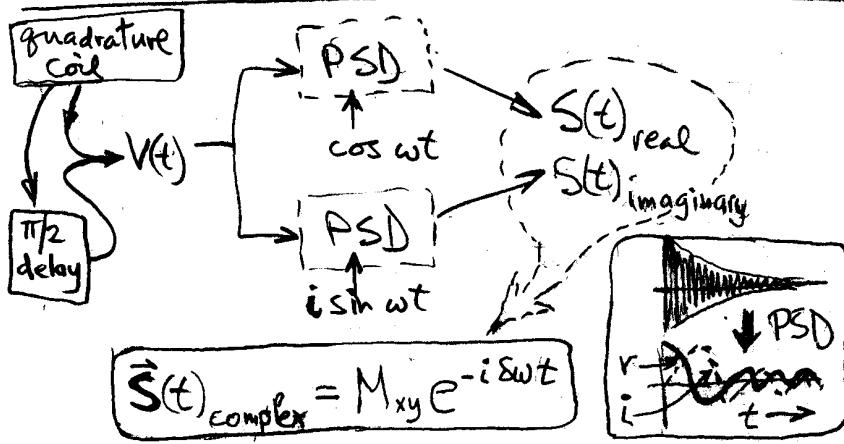
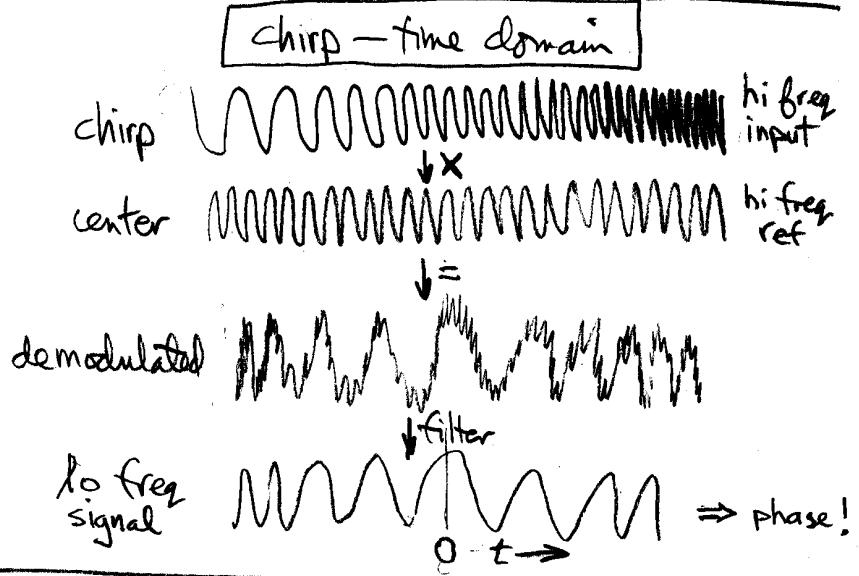
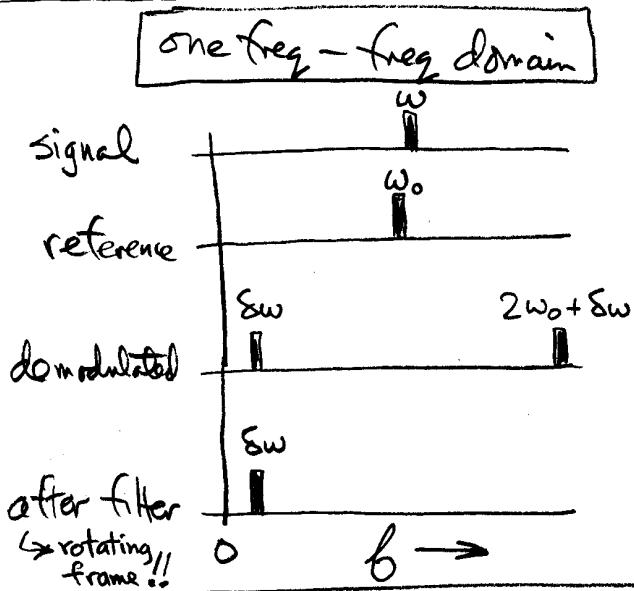
$$\propto \frac{1}{2} [\cos \delta\omega t - \cos(2\omega_0 + \delta\omega)t]$$

freq diff between  
RF input & ref

$\sin a \sin b = \frac{1}{2} [\cos(a-b) - \cos(a+b)]$   
 $\sin a \cos b = \frac{1}{2} [\sin(a-b) + \sin(a+b)]$

filter this one out  
w/ low pass filter

*this signal is digitized*



- two signals are made from a single receiving RF coil
- a quadrature coil can be treated the same way (OK to combine after adding  $\pi/2$  phase, then PSD)
- quadrature coil has better S/N since noise in each part is uncorrelated ( $\sqrt{2}$  better)

echoes - I

# FID - FREE INDUCTION DECAY, $T_2^*$

$T_2$  - unrecoverable (= rapid)  
 $T_2^*$  - add recoverable (= rapid + static)

- Signal (FID) resulting from RF pulse w/ angle  $\alpha$

$$\vec{S}(t) = \underbrace{\sin \alpha}_{\substack{\text{recorded} \\ \text{complex} \\ \text{signal}}} \cdot \underbrace{\rho(\omega)}_{\substack{\text{amount of tip} \\ \text{M}_z^0}} \cdot \underbrace{e^{-t/T_2(\omega)}}_{\substack{\text{spectral} \\ \text{density} \\ \text{funct.}}} \cdot \underbrace{e^{-i\omega t}}_{\substack{\text{time-dep} \\ \text{decay}}} \cdot \underbrace{dw}_{\substack{\text{rapid} \\ \text{oscillations} \\ \text{incr} \\ \text{freq.}}}$$

ignore space in obj

- An example spectral density ("Lorentzian inhomogeneity")

$$\rho(\omega) = M_z^0 \cdot \frac{(\gamma \Delta B \phi)^2}{(\gamma \Delta B \phi)^2 + (\omega - \omega_0)^2}$$

all atoms assumed to interact in same way in freq. range under curve

$\omega_0 \rightarrow \omega$  width is proportional to  $\Delta B \phi$

[subst  $\rho(\omega)$ , rearrange to extract  $\omega_0$ , take integral]

$\omega = \gamma B \phi \text{ (Bloch)}$   
 $\Delta \omega = \gamma \Delta B \phi$

$\rho(\omega)$  behavior:  $y = \frac{C}{C+x^2}$  (fixed height)

$$\vec{S}(t) = \pi \cdot M_z^0 \cdot \gamma \Delta B \phi \cdot \sin \alpha \cdot e^{-t\gamma\Delta B \phi} \cdot e^{-t/T_2} \cdot e^{-i\omega_0 t}$$

from integral of:  $\frac{C}{C+\omega^2}$  (big  $\Delta \rightarrow$  big sig.) (b/c fixed height)

extra decay from static (big  $\Delta \rightarrow$  big decay)

regular  $T_2$  decay

oscillations (complex) (one freq!)

[combine  $T_2$  + static terms]

$$\vec{S}(t) = \pi \cdot M_z^0 \cdot \gamma \Delta B \phi \cdot \sin \alpha \cdot e^{-t/T_2^*} \cdot e^{-i\omega_0 t}$$

N.B. center freq., not original integration variable

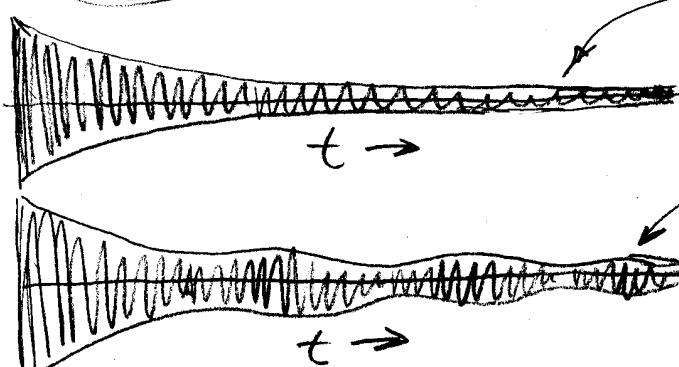
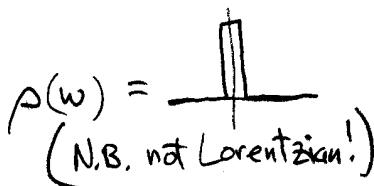
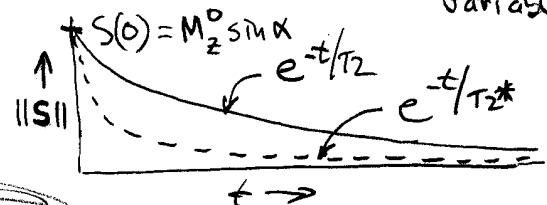
$$\frac{1}{T_2^*} = \frac{1}{T_2} + \frac{1}{T_2'} \rightarrow \gamma \Delta B \phi$$

overall decay rate including inhomogeneous  $B \phi$

unrecoverable "intrinsic" spin-spin

recoverable "static"

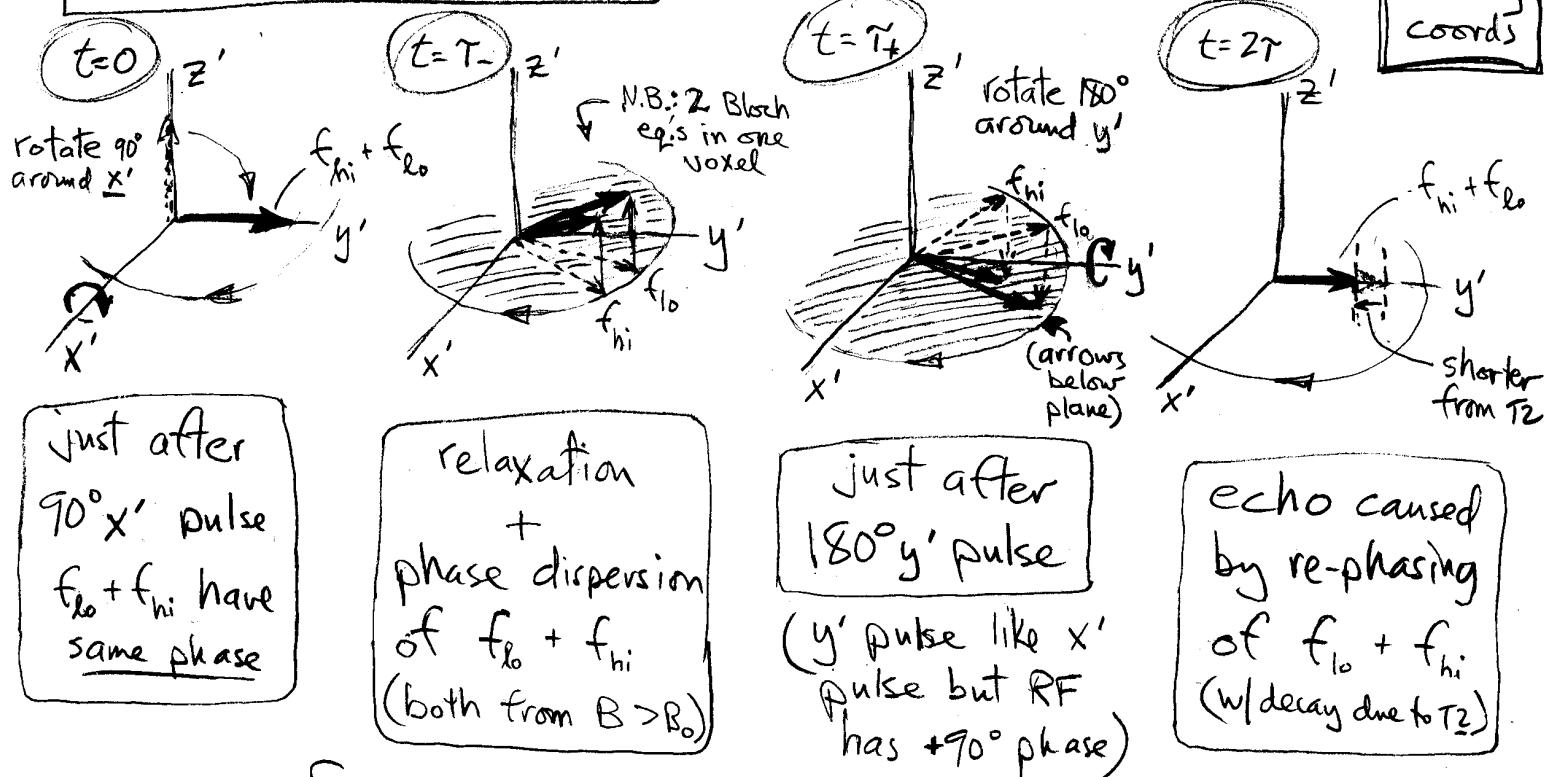
e.g. extra decay from BOLD



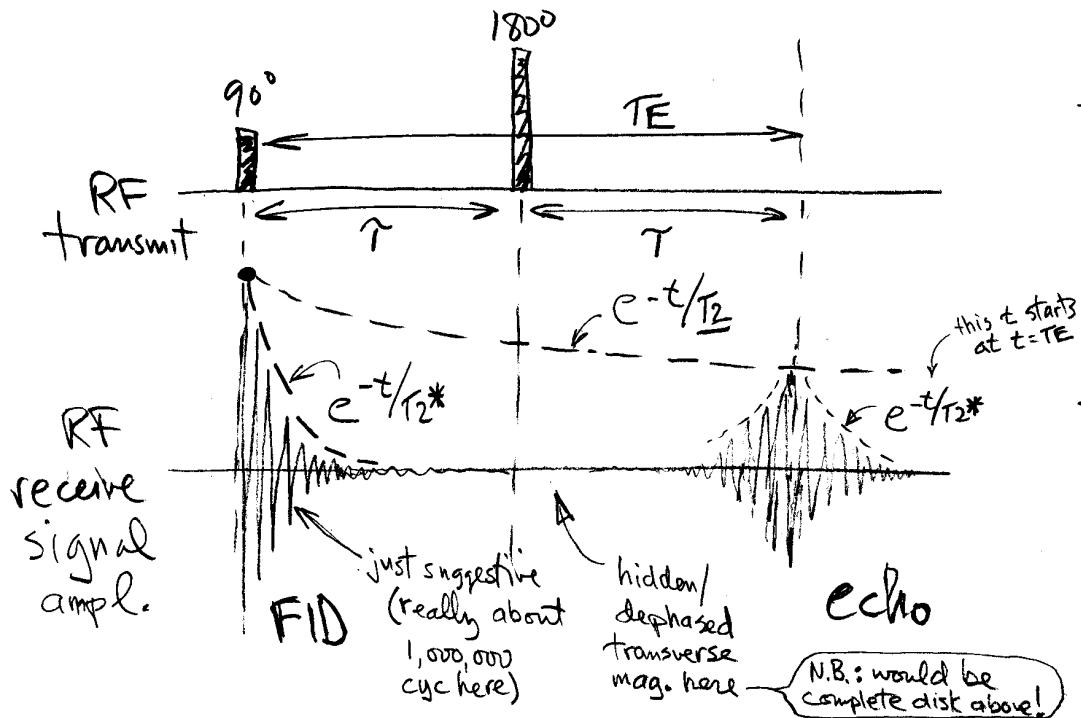
suggestive, since 100 million cycles per second  
 ↳ N.B. complex ↳ (r, i)

only approximated by  $e^{-t/T_2^*}$

## ECHOES — spin echo



- remember [brief RF just tips vectors while retaining length  
relaxation includes tips and shrinks ( $M_T$ ) and grows ( $M_z$ , echo)]
- $180^\circ x'$  pulse works, too, but echo will have  $+\pi$  phase (left side in figs above)
- echo generated even if second pulse not  $180^\circ$  (see next)



- FID decay (and echo growth/decay) described by  $T_2^*$ , from inhomogeneity
- reduction in height of echo compared to initial described by  $T_2$ , echo fixes the "star"

echoes-3-

# ECHOES — Spin echo

$\alpha_1 - \tau - \alpha_2 - \tau$  (both pulses along  
y' for simplicity)

see  $R_y(\alpha)$

## effect of $\alpha_y$ pulse

$$\begin{aligned} M_{x'} &\rightarrow M_{x'} \cos \alpha - M_{z'} \sin \alpha \\ M_{y'} &\rightarrow M_{y'} \\ M_{z'} &\rightarrow M_{x'} \sin \alpha + M_{z'} \cos \alpha \end{aligned}$$

↳ (etc for  $\alpha_x, \alpha_z$ )

General  
transforms  
(operators)

indicates  
rotating  
coord  
system

## effect of $\tau$ delay

see  $R_z(\tau)$

$$\begin{aligned} M_{x'} &\rightarrow (M_{x'} \cos \omega \tau + M_{y'} \sin \omega \tau) e^{-\tau/T_2} \\ M_{y'} &\rightarrow (-M_{x'} \sin \omega \tau + M_{y'} \cos \omega \tau) e^{-\tau/T_2} \\ M_{z'} &\rightarrow M_z^0(1 - e^{-\tau/T_1}) + M_{z'} e^{-\tau/T_1} \end{aligned}$$

immediately after  
 $\alpha_1$  pulse

$$\begin{aligned} M_{x'}(\omega, \alpha_1) &= -M_z^0(\omega) \sin \alpha_1 \\ M_{y'}(\omega, \alpha_1) &= 0 \\ M_{z'}(\omega, \alpha_1) &= M_z^0(\omega) \cos \alpha_1 \end{aligned}$$

for one isochromat  
of freq.  $\omega$

after  $\tau$  delay

$$\begin{aligned} M_{x'}(\omega, \tau) &= -M_z^0(\omega) \sin \alpha_1 \cos \omega \tau e^{-\tau/T_2} \\ M_{y'}(\omega, \tau) &= M_z^0(\omega) \sin \alpha_1 \sin \omega \tau e^{-\tau/T_2} \\ M_{z'}(\omega, \tau) &= M_z^0(\omega) [1 - (1 - \cos \alpha_1) e^{-\tau/T_1}] \end{aligned}$$

immediately after  
 $\alpha_2$  pulse (no effect  
on  $M_{y'}$ ; rewrite y;  
combine x and y eq.s)

$$\begin{aligned} M_{x'y'}(\omega, \tau_+) &= M_z^0(\omega) \sin \alpha_1 \left( \sin^2 \frac{\alpha_2}{2} e^{i\omega\tau} - \cos^2 \frac{\alpha_2}{2} e^{i\omega\tau} \right) e^{-\tau/T_2} \\ &\quad - M_z^0(\omega) [1 - (1 - \cos \alpha_1) e^{-\tau/T_1}] \sin \alpha_2 \end{aligned}$$

$$M_{x'y'}(\omega, t) = M_{x'y'}(\omega, \tau_+) e^{-(t-\tau)/T_2} e^{-i\omega(t-\tau)}$$

$$= M_z^0(\omega) \sin \alpha_1 \sin^2 \frac{\alpha_2}{2} e^{-t/T_2} e^{-i\omega(t-2\tau)}$$

$$- M_z^0(\omega) \sin \alpha_1 \cos^2 \frac{\alpha_2}{2} e^{-t/T_2} e^{-i\omega t}$$

$$- M_z^0(\omega) [1 - (1 - \cos \alpha_1) e^{-\tau/T_1}] \sin \alpha_2 e^{-(t-\tau)/T_2} e^{-i\omega(t-\tau)}$$

— for a large num of freq's:

[terms ② ≠ ③ are dephasing → FID of echo]

term ① rephasing → rephase at  $t = 2\tau$

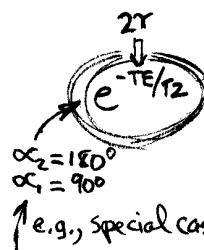
echo  
signal  
from ①

$$S(t) = \sin \alpha_1 \sin^2 \frac{\alpha_2}{2} \int_{-\infty}^{\infty} \rho(\omega) e^{-t/T_2(\omega)} e^{-i\omega(t-TE)} d\omega$$

$\Downarrow t=TE$

peak ampl

$$A_E = \sin \alpha_1 \sin^2 \frac{\alpha_2}{2} \int_{-\infty}^{\infty} \rho(\omega) e^{-TE/T_2(\omega)} d\omega = M_z^0 \sin \alpha_1 \sin^2 \frac{\alpha_2}{2} e^{-TE/T_2}$$



90°<sub>y'</sub> -  $\tau$  - 90°<sub>y'</sub>  
90°<sub>y'</sub> -  $\tau$  - 180°<sub>y'</sub>  
90°<sub>x'</sub> -  $\tau$  - 180°<sub>y'</sub>

$$S_1(t) = \frac{1}{2} \int_{-\infty}^{\infty} \rho(\omega) e^{-t/T_2(\omega)} e^{-i\omega(t-TE)} d\omega$$

S<sub>2</sub>(t) = no 1/2 factor

S<sub>3</sub>(t) = multiply by i → add  $\pi/2$  phase

(echo amplitude, ignoring  
freq dependence of T<sub>2</sub>)

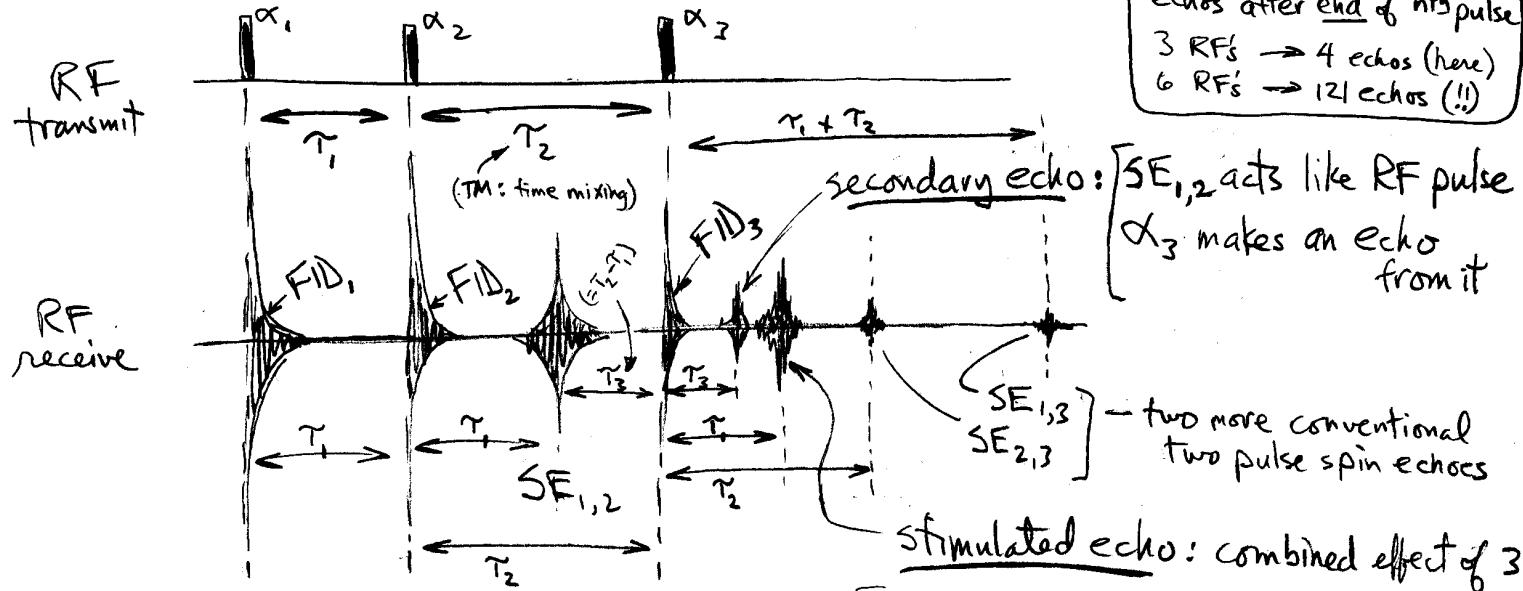
etc for A<sub>E</sub>... like ↑

lectures - 4

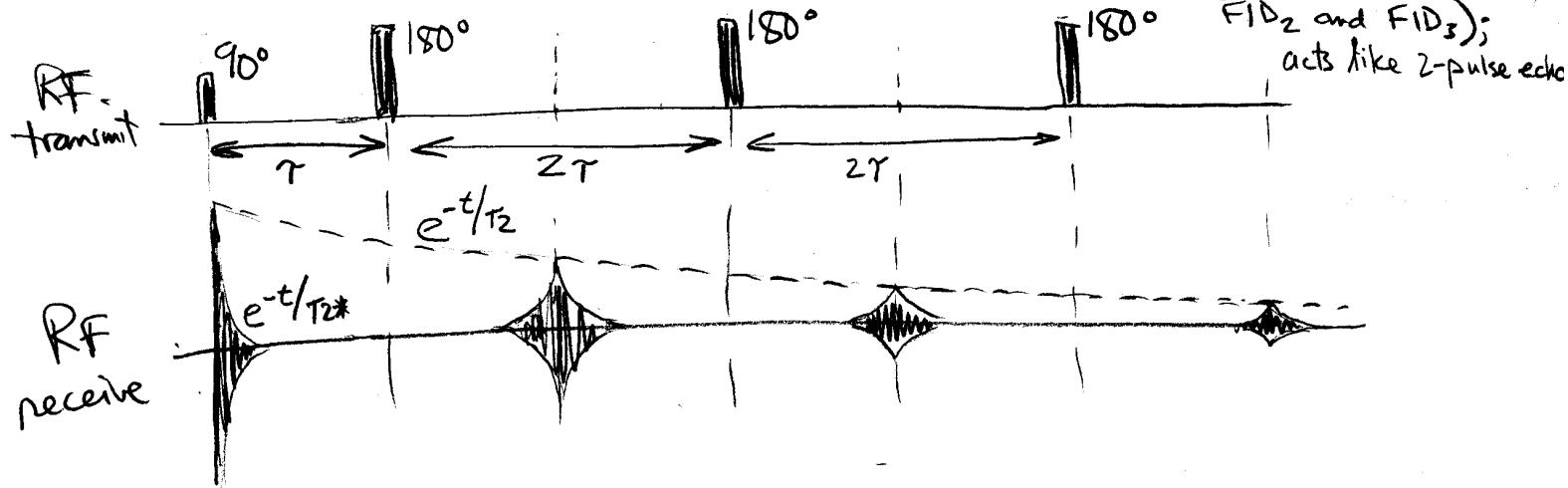
# ECHO TRAINS

- spin-echo trains.

- it's (too) easy to make echos ...



- a useful multi-echo sequence (CPMG) is a  $90^\circ$  followed by  $180^\circ$  at  $2\tau$  spacing



- typically,  $90^\circ$  and  $180^\circ$  applied in different axes ( $x'$ , then  $y', y'', y'''$ ) which reduces phase errors due to imperfect  $180^\circ$  pulses (since slightly-off rotation around  $y'$  affects phase less)

# EXTENDED PHASE GRAPHS

- Using full Bloch eq. solutions is tedious (need 1000 copies)
- Pictorial method for visualizing effects of a series of  $\alpha$  pulses
- Starting point: initial RF creates new transverse

↳ vs. easier to visualize 90/180°

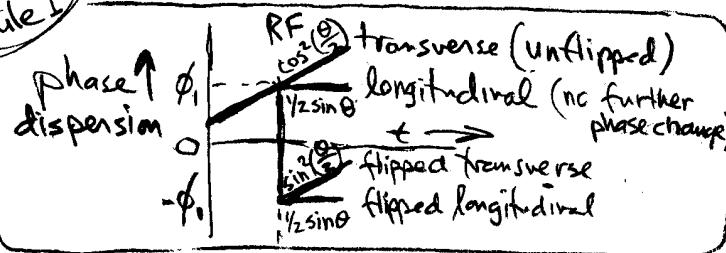
effects on existing

$\alpha$  pulse rotates a portion of existing transverse mag. into a position that results in rephasing and another portion into  $M_L$

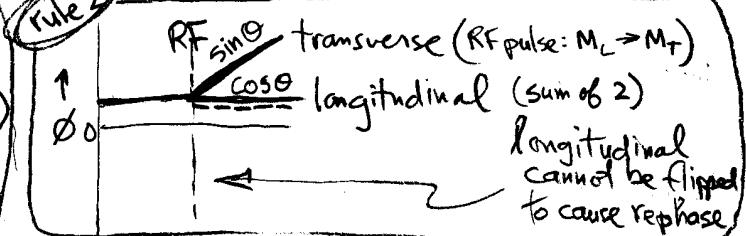
third pulse can uncover and rephase transverse mag. temporarily saved in longitudinal

↳ QM view helps

Rule 1

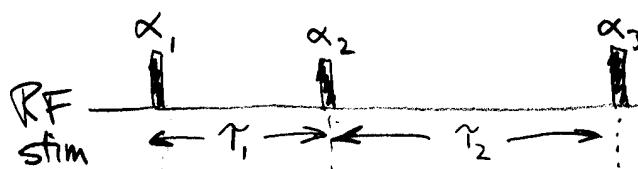


Rule 2



↳ branching rule for effect of  $\alpha$  RF pulse on transverse mag

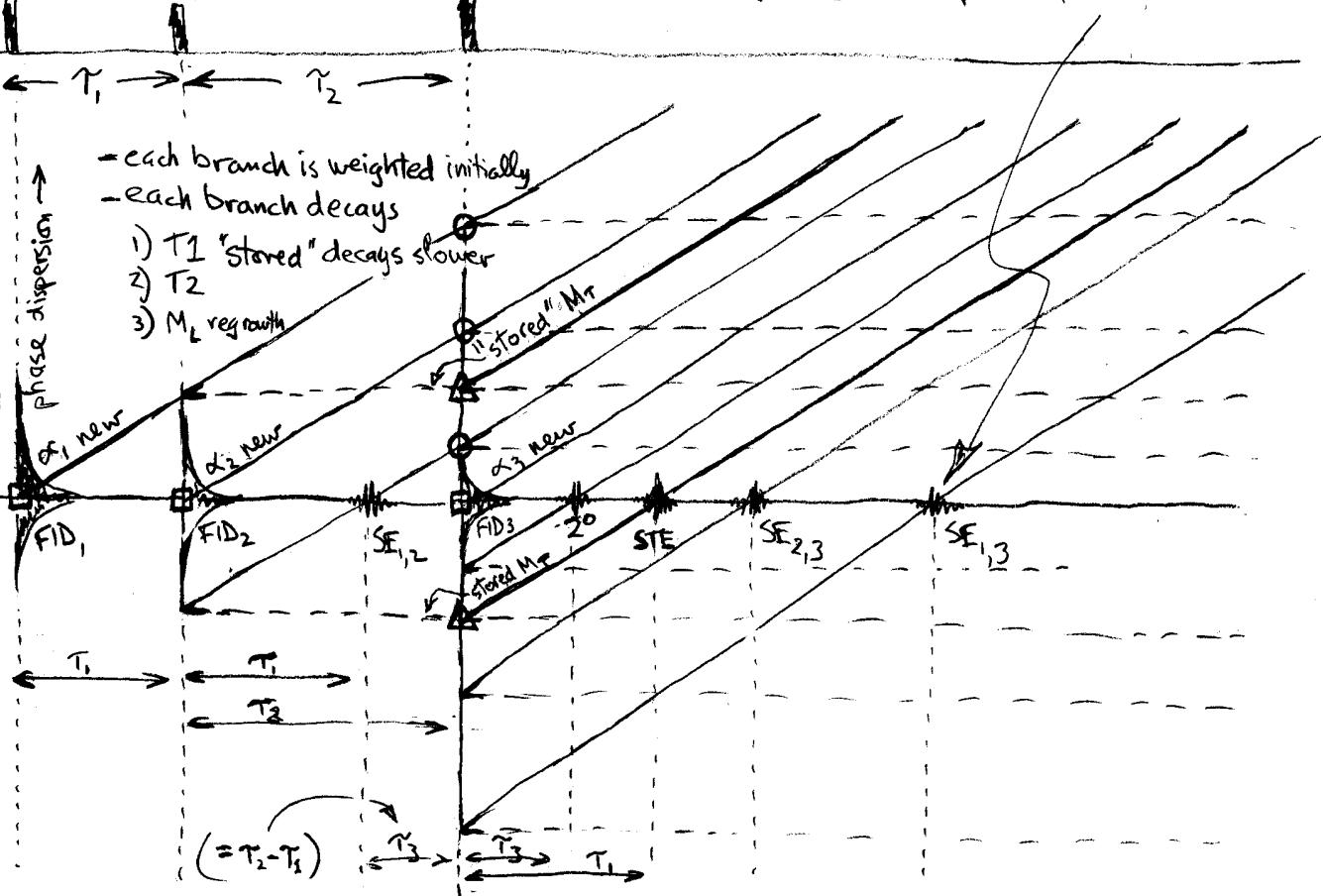
↳ branching rule for effect of  $\alpha$  RF pulse on longitudinal mag



→ echo when phase path crosses zero

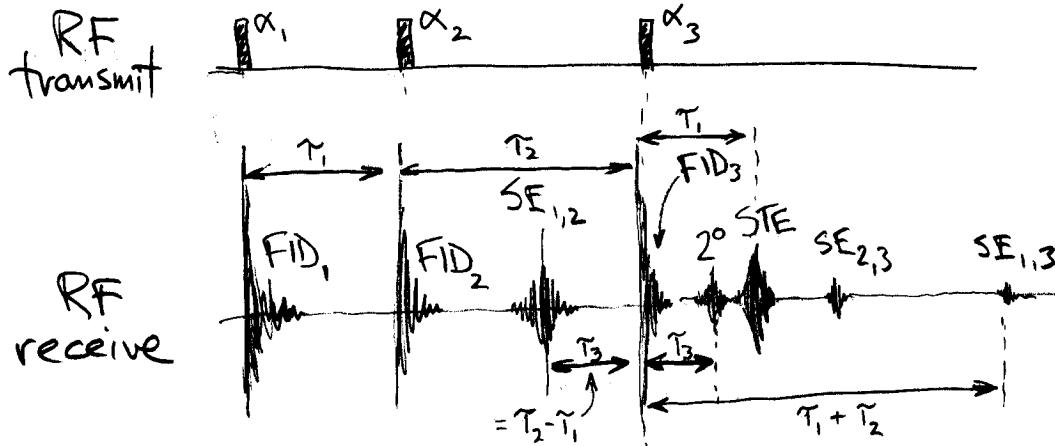
branch points

- new  $M_T$
- O - apply rule 1
- apply rule 2



echoes-4b

### 3-PULSE ECHO AMPLITUDES

— assume  $M_z^0 = 1$ 

echo	time	amplitude
$SE_{1,2}$	$(t = 2T_1)$	$\sin \alpha_1 \sin^2 \frac{\alpha_2}{2} e^{-2T_1/T_2}$ ↳ special cases
$SE_{1,2}$ ("secondary")	$(t = 2T_2)$	$-\sin \alpha_1 \sin^2 \frac{\alpha_2}{2} \sin^2 \frac{\alpha_3}{2} e^{-2T_2/T_2}$
$SE_{1,2}$ ("stimulated")	$(t = 2T_1 + T_2)$	$\frac{1}{2} \sin \alpha_1 \sin \alpha_2 \sin \alpha_3 e^{-\tilde{T}_2/T_1} e^{2T_1/T_2}$
$SE_{2,3}$	$(t = T_1 + 2\tilde{T}_2)$	$[1 - (1 - \cos \alpha_1) e^{-T_1/T_1}] \sin \alpha_2 \sin^2 \frac{\alpha_3}{2} e^{-(T_1 + 2\tilde{T}_2)/T_2}$
$SE_{1,3}$	$(t = 2(T_1 + T_2))$	$\sin \alpha_1 \cos^2 \frac{\alpha_2}{2} \sin^2 \frac{\alpha_3}{2} e^{-2(T_1 + T_2)/T_2}$

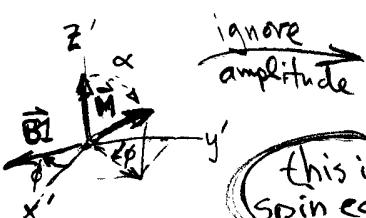
—  $T_1$ -dependence in STE (but also  $SE_{2,3}$ ) from temporary "storage" of  $M_T$  in  $M_L$ , then recovery by third pulse

lectures - 6

**HYPER ECHOES**

(N.B.: coord rot to put z horiz. vs. Block notes)

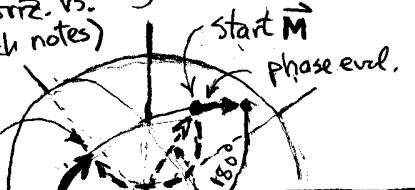
(N.B. right-handed coord syst vs. left-handed in Block eq. notes)



1.

$$\varphi_z - 180^\circ - \varphi_z = 180^\circ$$

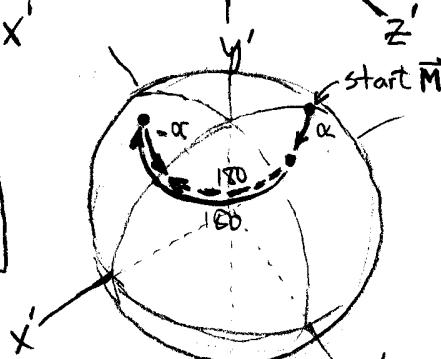
3 solid lines      1 dashed line



3 symmetries

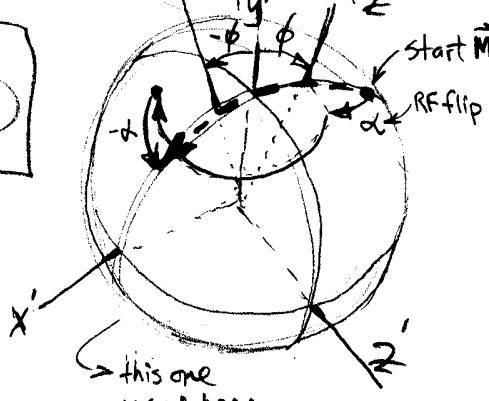
2.

$$\alpha_y - 180^\circ - -\alpha_y = 180^\circ$$



3.

$$\alpha_\phi - 180^\circ - -\alpha_\phi = 180^\circ$$

Practical use

- multi-echo example
- can also use to prepare, then separate read-out

- practical prob: 180° pulses deposit a lot of RF ( $6 \times 90^\circ$ )  
 $\hookrightarrow$  prob at high fields

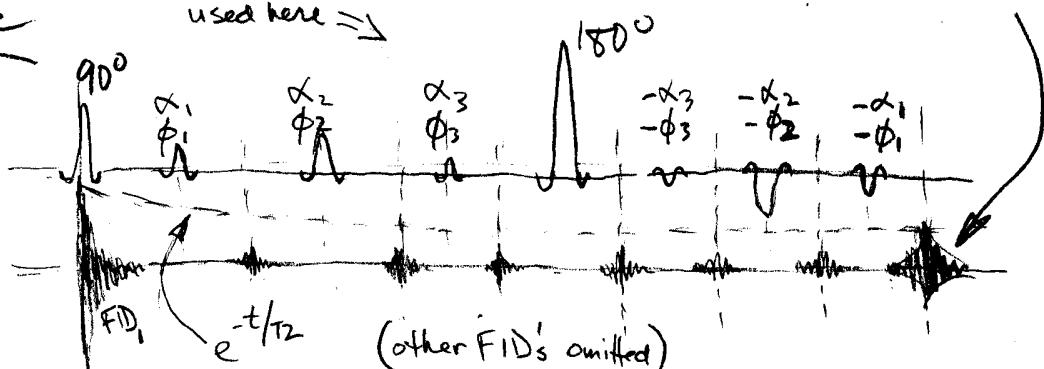
- by arranging to get big echo in middle of k-space  
 can get by with much less RF power

- Hennig & Scheffler (2001)
- normalize  $\vec{M}$  amplitude  $\rightarrow 1$
- sphere surface defines 2D space for  $\vec{M}$  moved by:

  - 1) vect. rotation of  $\vec{M}$  around tilted axis in transverse x-y plane by RF with flip,  $\alpha$ , and phase,  $\phi$ :  $P(\alpha, \phi)$
  - 2) rotation around z by phase evolution due to freq offset,  $\omega$  (B0 offset) and time,  $t$ :  $\varphi(\omega, t)$

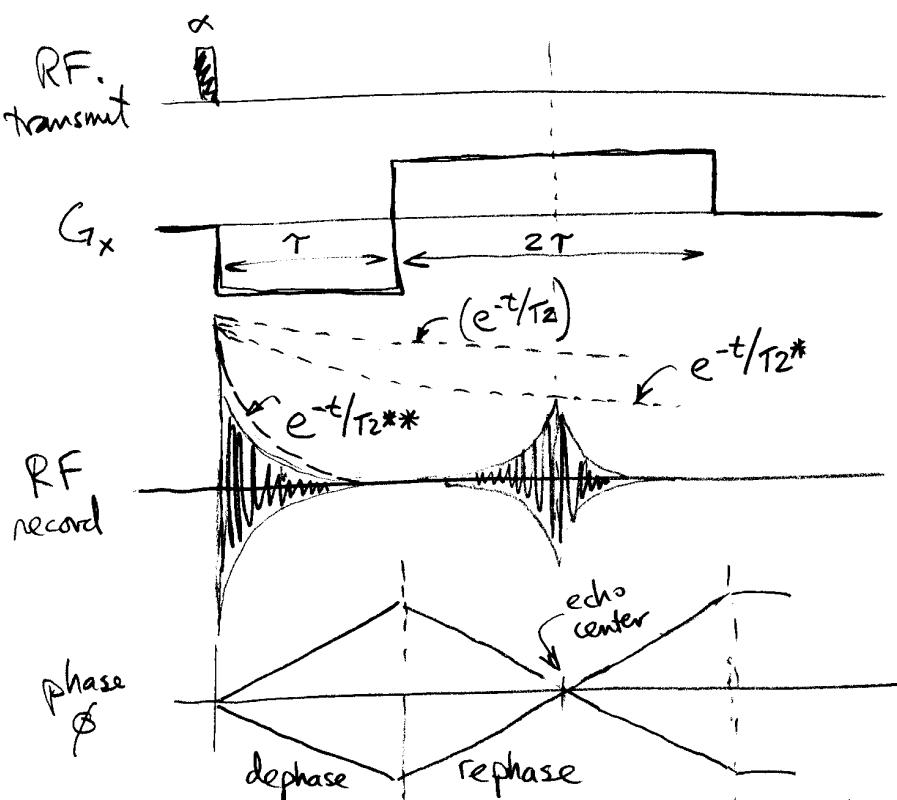
- three symmetries:
  - Solid lines: phase evol or RF flip  $\rightarrow 180^\circ$  eval or RF
  - dashed lines: just 180° equiv.

- by combining long sequences observing these symmetries, can generate as strong echo even w/ many inserted  $\alpha$ -pulses in between



## GRADIENT ECHOES

-  $T_2^*$ , GE chains



- initial negative gradient dephases spins

- after  $t=T$  of positive gradient, spins rephase

- does not correct for  $T_2^*$  inhomogeneities  
So echo amplitude is

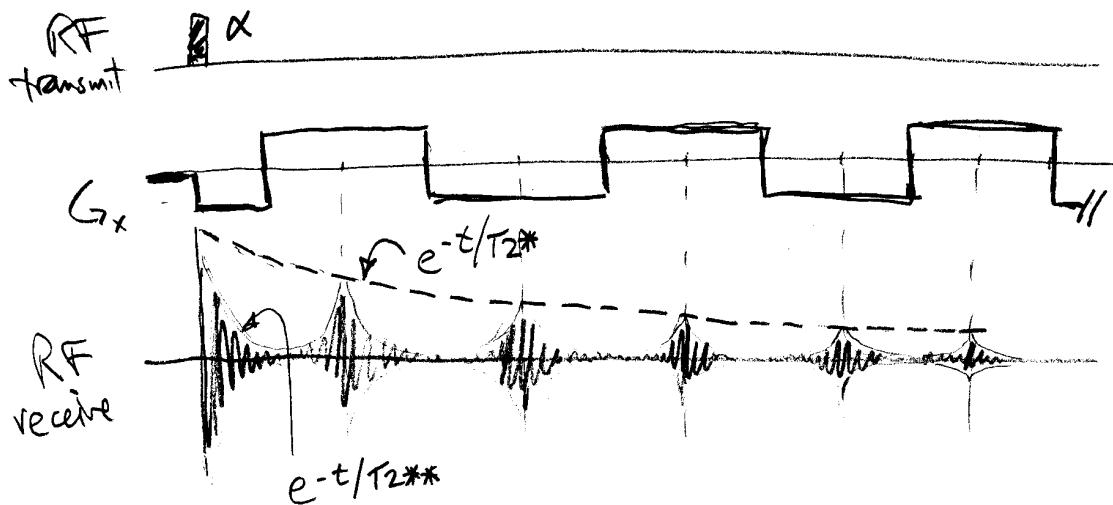
$$A_E = e^{-t/T_2^*}$$

- the initial "FID" is not "free" since it is being actively de-phased by gradient, so FID decay.

$$\frac{1}{T_2} < \frac{1}{T_2^*} < \frac{1}{T_{2**}} \Rightarrow A_F = e^{-t/T_{2**}}$$

- key difference between spin-echoes (SE) and gradient echo (GE) is that  $B_0$  inhomogeneities not corrected
  - ↪ hence, echoes are  $T_2^*$ -weighted, not  $T_2$ -weighted  $\Rightarrow$  more susceptible to inhomogeneities

- echo trains possible w/ gradient echo (CPMG-like)



- the faster the gradients are switched, the more echoes you get

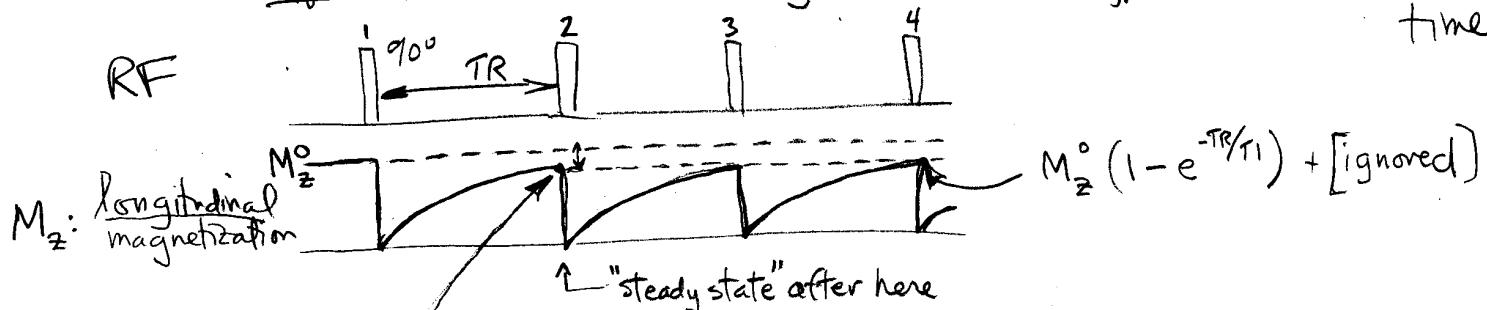
- EPI hardware  
 $\Rightarrow$  64 echoes

Contrast I

# IMAGE CONTRAST

T1 Saturation-recovery (no echo, just FID)

- contrast (PD, T1, T2, T2\*) depends on magnetization not getting back to equilibrium, and then differences in how far away each tissue type is at measurement time



- simple saturation/recovery w/ no echo
- initial conditions:  $[M_z \text{ before first pulse} = M_z^0]$   
 $[M_z = 0 \text{ immmed. after first pulse (i.e., } 90^\circ \text{ pulse)}]$

- from Bloch eq,  $M_z$  just before second pulse:

$$M_z^{(n)}(0_-) = M_z^{(0)} (1 - e^{-\frac{TR}{T1}}) + M_z^{(n-1)}(0_+) e^{-\frac{TR}{T1}}$$

$M_z$  before current pulse       $M_z$  "regrowth-from-zero" term       $M_z$  "left-immmed.-after-pulse" term (N.B. decaying)

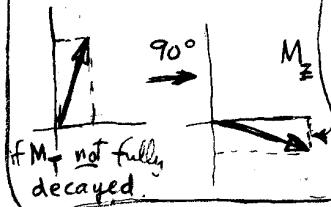
- given  $\begin{cases} (1) 90^\circ \text{ pulse} \\ (2) \text{no } M_{xy} \text{ left} \end{cases}$  → pure tip:  $M_{xy} = M_z$

assume this is 0 because we assume that  $M_{xy}$  (transverse) completely decayed so that a  $90^\circ$  pulse doesn't generate any initial longitudinal

- tip existing mag

$$M_z^{(n)}(0_-) = M_{xy}(0_+) = M_z^0 (1 - e^{-\frac{TR}{T1}})$$

longitudinal mag just before pulse      transverse we can record after pulse      transverse mag depends on T1!



- that is, the not-completely-regrown longitudinal magnetization, which depends on T1, but which we cannot record, is completely converted to recordable transverse magnetization

assume immediate recording of signal

$$I(r) = C \underbrace{\rho(r)}_{\text{rec const. spectral dens.}} \left( 1 - e^{-\frac{TR}{T1(r)}} \right)$$

spectral dens  $\rho(r)$   
 $\approx$  p. density: underlies equilb.  $M_z^0$

Contrast-1b

IMAGE CONTRAST

Why imperfect  $90^\circ$  takes multiple flips  
til steady state

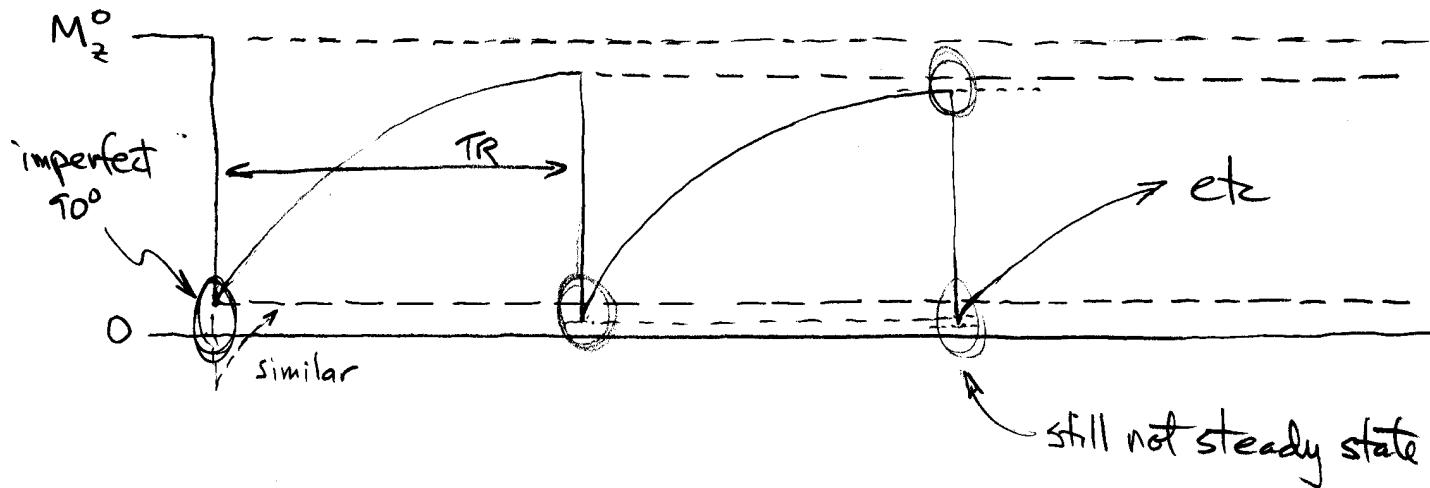
- initial fMRI images are usually discarded (why?)

↳ because they are brighter than all the rest

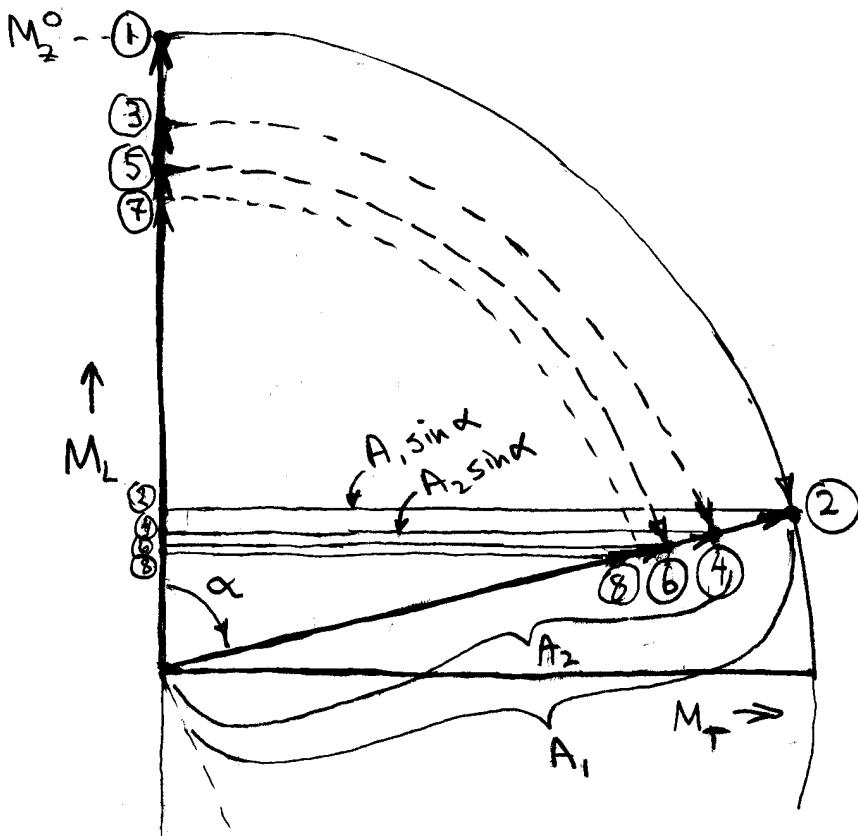
↳ because multiple flip required before steady state

N.B.: B1 imperfections guarantee this situation will occur

(e.g. at 3T, flip angle varies almost 25% across brain)



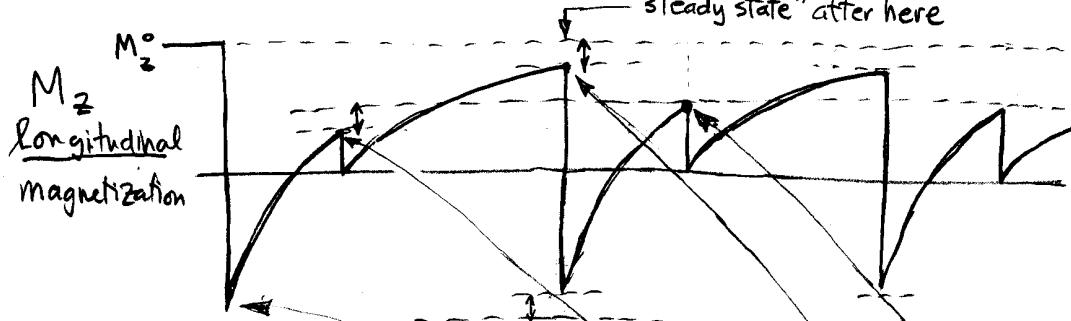
- at 3T, steady state for typical 1-2 sec TR images reached after ~8 images



## IMAGE CONTRAST

IR (still just saturation-recovery — no echo)

- inversion recovery w/ no echo



- $180^\circ$  deg pulse reverses longitudinal magnetization

$$M_z' = -M_z^0$$

- recovery to end of first TI from long. part of Bloch eq.

$$M_z' = M_z^0 \left(1 - 2e^{-\frac{t}{T_1}}\right) \quad \xrightarrow{\text{flipped into transverse by second pulse (1st } 90^\circ)}$$

- longitudinal then regrows from zero

$$M_z' = M_z^0 \left(1 - e^{-(TR-TI)/T_1}\right) \quad \xrightarrow{\text{from first Bloch term only}}$$

- after second  $180^\circ$ , just change sign again

$$M_z' = -M_z^0 \left(1 - e^{-(TR-TI)/T_1}\right)$$

- apply relaxation eq. again

$$M_z' = M_z^0 \left(1 - e^{-TI/T_1}\right) - M_z^0 \left(1 - e^{-(TR-TI)/T_1}\right) e^{-TI/T_1}$$

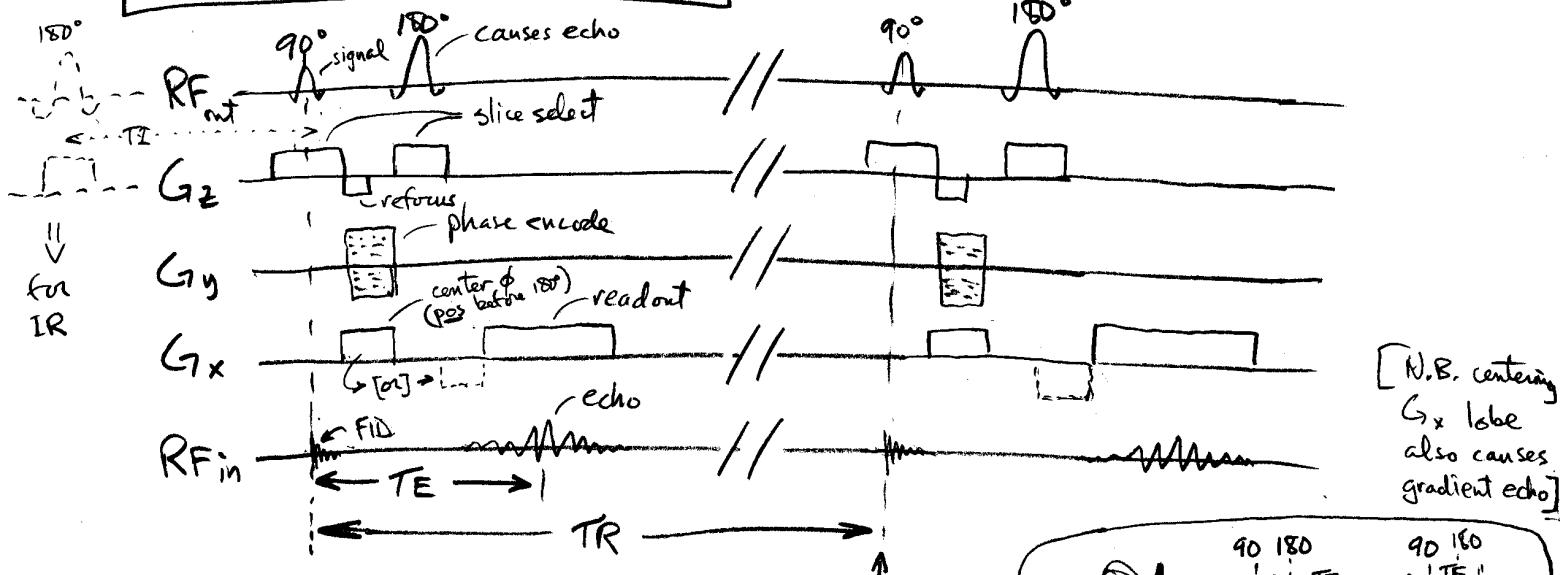
$$M_z' = M_z^0 \left(1 - 2e^{-TI/T_1} + e^{-TR/T_1}\right)$$

→ this is magnetization flipped to transverse, therefore made recordable

Content - 3

# IMAGE CONTRAST

SE, IR-SE



- steady state mag (2nd TR) just before 90°  $\Rightarrow$  sum Bloch eq. for  $M_L$   $\xrightarrow{TR-TE \ll TR}$
- $$M_{z'}(0_-) = M_z^0 \left( 1 - 2e^{-(TR-TE/2)/T_1} + e^{-TR/T_2} \right)$$
- the echo signal ( $M_T$ ) unlike in simple saturation-recovery FID has an additional delay before it is recorded, so we have to take account of transverse mag relaxation  $\xrightarrow{M_L=0 \text{ during max } M_T(\text{echo})}$
- $$A_E = M_z^0 \left( 1 - 2e^{-(TR-TE/2)/T_1} + e^{-TR/T_2} \right) e^{-TE/T_2} \xrightarrow{\text{from spin echo equation here}}$$

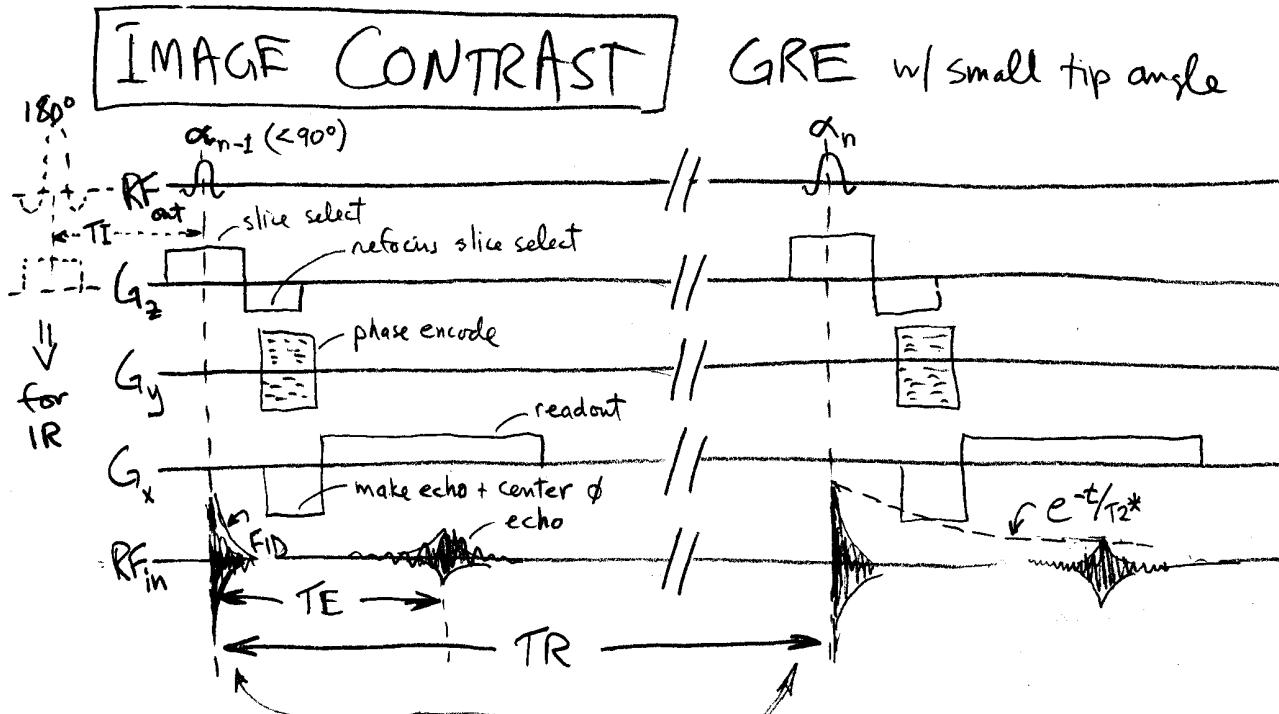
- if we assume TE much less than TR, then we can simplify:

$$A_E = \underbrace{M_z^0}_{\substack{\text{amplitude} \\ \text{echo}}} \underbrace{\left( 1 - e^{-TR/T_1} \right)}_{\substack{\text{proton} \\ \text{density}}} \underbrace{e^{-TE/T_2}}_{\substack{\text{TR controls } T_1 \\ \text{contrast}}} \quad \xrightarrow{\text{TR-TE} \approx \text{TR}}$$

- similar equation for SE-IR

$$A_E = M_z^0 \left( 1 - 2e^{-TI/T_1} + e^{-TR/T_2} \right) e^{-TE/T_2}$$

## Contrast - 4



- use basic longitudinal relaxation from Bloch eq. again  
 $\hookrightarrow$  & assume  $M_{x'y'}^{(n)}(O_-) = 0$   $\rightarrow$  transverse dephased before next pulse

$$M_{z'}^{(n)}(O_-) = M_z^0 (1 - e^{-\text{TR}/T_1}) + M_{z'}^{(n-1)}(O_+) e^{-\text{TR}/T_1} \quad \text{long TR or spoiler}$$

- assume we have a small tip angle:  $M_z \cos \alpha \uparrow \downarrow \frac{\alpha}{M_z} \Rightarrow M_{z'}^{(n)}(O_+) = M_z^{(n)}(O_-) \cos \alpha$

$$M_{z'}^{(n)}(O_-) = [M_z^0 (1 - e^{-\text{TR}/T_1})] + M_{z'}^{(n-1)}(O_-) \cos \alpha e^{-\text{TR}/T_1}$$

- assume we are in dynamic equilibrium:  $(\text{current} = \text{previous})$

$$M_{z'}^{(n)}(O_-) = M_{z'}^{(n-1)}(O_-) \Rightarrow M_{z'}^{ss}(O_-)$$

steady state:  
 - subst. into prev. eq.  
 for both  $M_{z'}(O_-)$ 's  
 - solve for it

prepulse steady state longitudinal

$$M_{z'}^{ss}(O_-) = \frac{M_z^0 (1 - e^{-\text{TR}/T_1})}{1 - \cos \alpha e^{-\text{TR}/T_1}}$$

$$\begin{aligned} M_{ss} &= M_0 + M_{ss} \cos \\ M_{ss} - M_{ss} \cos &= M_0 \\ M_{ss}(1 - \cos) &= M_0 \\ M_{ss} &= M_0 / (1 - \cos) \end{aligned}$$

post-pulse transverse magnetization

$$M_{x'y'}^{ss}(t) = \frac{M_z^0 (1 - e^{-\text{TR}/T_1})}{1 - \cos \alpha e^{-\text{TR}/T_1}} \cdot \sin \alpha e^{-t/T_2^*}$$



gradient echo amplitude

$$A_E = \frac{M_z^0 (1 - e^{-\text{TR}/T_1})}{1 - \cos \alpha e^{-\text{TR}/T_1}} \sin \alpha e^{-\text{TE}/T_2^*}$$

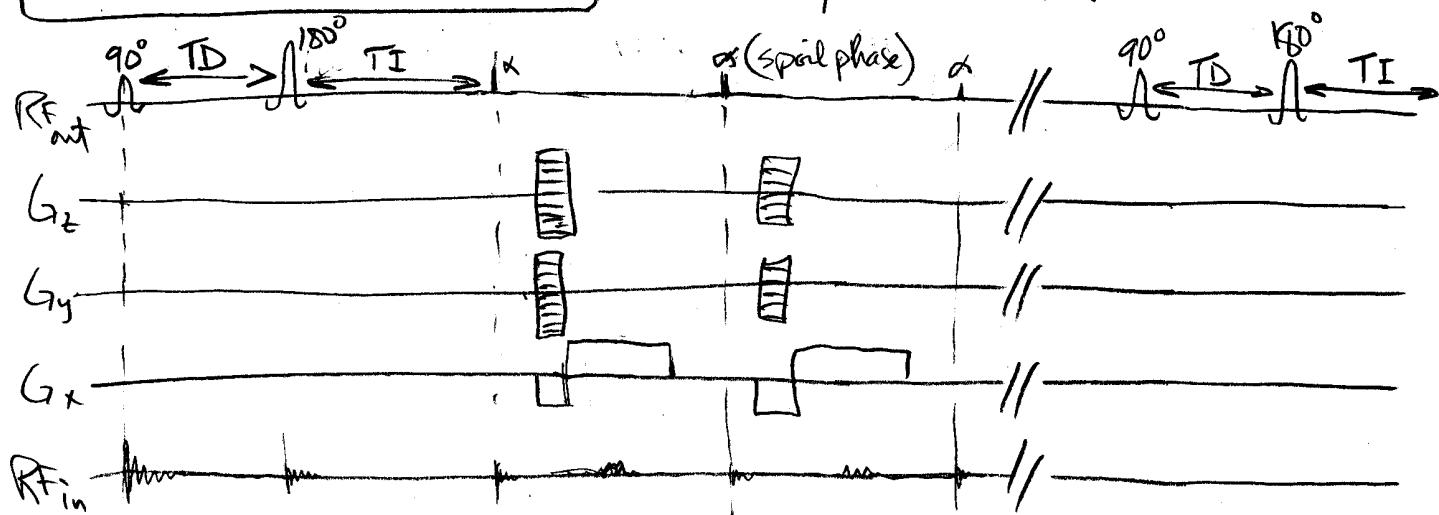
from gradient echo

$T_1$  contrast mainly depends on flip angle, not  $\text{TR} \rightarrow \cos 90^\circ = 1 \rightarrow$  eliminates  $T_1$  weight since denom = numer

Contrast - 4b

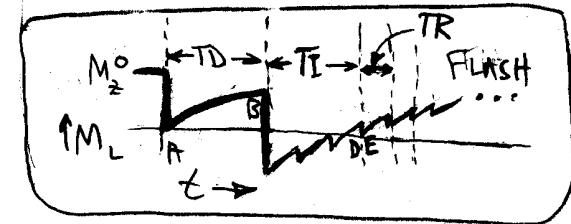
## IMAGE CONTRAST

## MDEFT / 3D FLASH



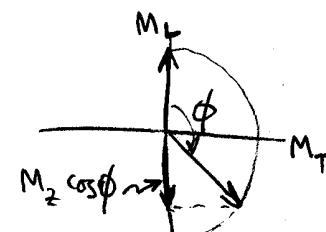
- Saturate, wait for contrast<sub>1</sub>, invert, wait for contrast<sub>2</sub>, FLASH (center out)

A)  $M_z' \left( \begin{smallmatrix} \text{just} \\ \text{after} \\ 90^\circ \end{smallmatrix} \right) = 0$  (perfect 90°)



B)  $M_z' \left( \begin{smallmatrix} \text{after} \\ \text{TD} \end{smallmatrix} \right) = M_z^0 \left( 1 - e^{-\text{TD}/T_1} \right)$  (Bloch term #1)

C)  $M_z' \left( \begin{smallmatrix} \text{just} \\ \text{after} \\ \text{invert} \end{smallmatrix} \right) = \cos \phi M_z^0 \left( 1 - e^{-\text{TD}/T_1} \right)$



D)  $M_z' \left( \begin{smallmatrix} \text{after} \\ \text{TI} \end{smallmatrix} \right) = M_z^0 \left( 1 - e^{-\text{TI}/T_1} \right) + [\cos \phi M_z^0 \left( 1 - e^{-\text{TD}/T_1} \right)] e^{-\text{TI}/T_1}$

=  $M_z^0 \left[ 1 - [1 - \cos \phi (1 - e^{-\text{TD}/T_1})] e^{-\text{TI}/T_1} \right]$  after preparation

Special case TI=TD:

$$= M_z^0 \left[ 1 - e^{-\text{TI}/T_1} \right]^2$$

using hard 180° 'inversion'  
can cancel hard alpha B1  
inhomogeneities (Thomas et al. '05)

- after the first X pulse:

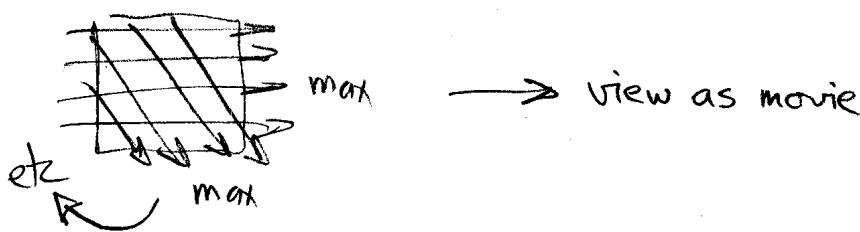
E) \*  $M_z' \left( \begin{smallmatrix} \text{just} \\ \text{after} \\ \text{first X} \end{smallmatrix} \right) = M_z^0 \left[ 1 - [1 - \cos \phi (1 - e^{-\text{TD}/T_1})] e^{-\text{TI}/T_1} \right] \sin \alpha$

contrast-4c

## MAGNETIZATION TRANSFER CONTRAST

- protons in macromolecules & bond to membranes have wide range of resonant freqs ("bound") }  $\rightarrow T_2 = 1 \text{ msec}$
- free protons in blood, CSF, water have narrow range of resonant freqs ("free") }  $\rightarrow T_2 = 50 \text{ msec}$
- mag transfer pulse sequence
  - 1) off center freq. pulse to hit "bound" (but don't hit water too hard)
  - 2) wait for magnetization transfer from saturated longitudinal  $M_z$  of "bound"  $\rightarrow M_z$  of "free"
  - 3) result of transfer  $\rightarrow$  attenuation

$\rightarrow$  N.B. this always happens a little (cf. T1-weighted, T2-weighted)  
Something to keep in mind if hard pulse (wide free)
- used to increase contrast in TOF  
TOF (not explained) bright vessels from inflow fresh spins  
MT - contrast added: suppress tissue but not blood
- view w/ MIP: maximum intensity projection along lines



contrast-5

## SIGNAL-TO-NOISE, CONTRAST-TO-NOISE

- Signal-to-noise defined as:  $\text{SNR} \equiv \frac{\bar{S}}{\sigma_n}$  avg obj signal s.d. non-object region
- temporal SNR:  $t\text{SNR} \equiv \frac{\bar{S}_t}{\sigma_t}$
- "Contrast" is a difference
- contrast-to-noise ratio:

$$\text{CNR}_{AB} \equiv \frac{\bar{C}_{AB}}{\sigma_n} = \frac{\bar{S}_A - \bar{S}_B}{\sigma_n} = \text{SNR}_A - \text{SNR}_B$$

Haacke

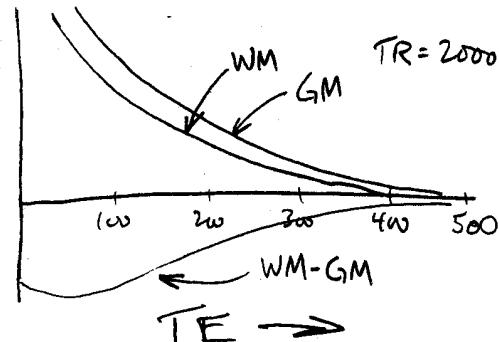
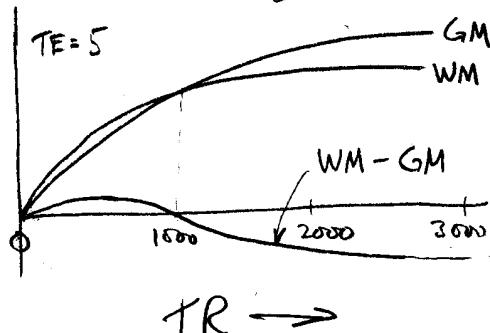
tissue	T1 (1.5T)	T2	PD
GM	950	100	0.80
WM	600	80	0.65
CSF	4500	2200	1.00
blood	1200	100-200	water

Lauterbur tissue	T1	T2	PD
GM	760	77	0.69
WM	510	67	0.61
CSF	2650	280	1.00

- spin-echo:

$$A_E = M_0^o (1 - e^{-TR/T1}) e^{-TE/T2}$$

Signal, contrast



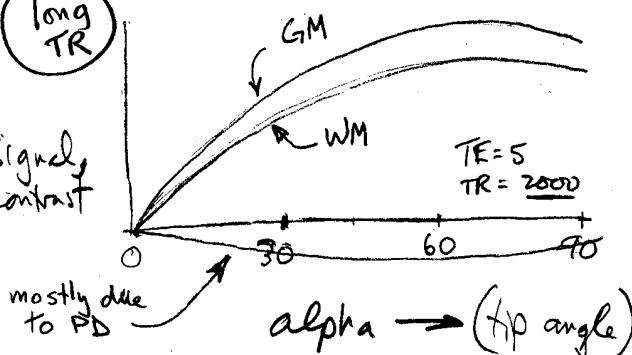
- gradient echo:

$$A_E = \frac{M_0^o (1 - e^{-TR/T1})}{1 - \cos \alpha e^{-TR/T1}} \sin \alpha e^{-TE/T2*}$$

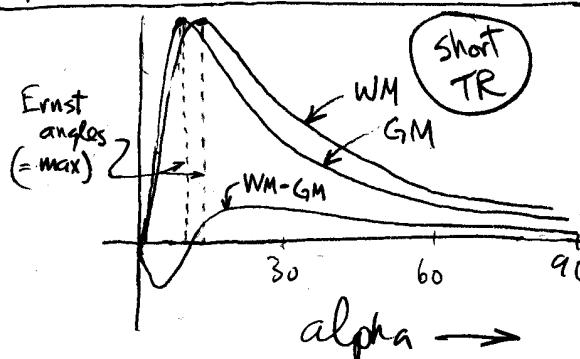
1.5T  
 $T_{2*}^*_{GM} = 50$   
 $T_{2*}^*_{WM} = 40$

long TR

Signal, contrast



mostly due to PD



- general rules: spin-echo, long TR GE

proton-density weighted	TR ↑↑ (no T1 diff)	TE ↓↓ (no T2 diff)
T1-weighted	TR ≈ T1 (big T1 diff)	TE ↓↓ (no T2 diff)
T2-weighted	TR ↑↑ (no T1 diff)	TE ≈ T2 (big T2 diff)

Contrast - b

## SIGNAL-TO-NOISE S/N

- generalized dependence of SNR on 3D imaging parameters

$$\text{SNR/voxel} \propto \Delta x \Delta y \Delta z \sqrt{N_{\text{acq}} N_x N_y N_z \Delta t}$$

[ voxel size (of same number) ]    [ num repeats ]    [ number of voxels (of same size) ]    [ read timestep ]

- size (volume) of voxels (with the number of voxels held constant), linear effect on S/N

↳ e.g.,  $\boxed{3 \times 3 \times 3 \text{ mm} \rightarrow 4 \times 4 \times 4 \text{ mm}} \rightarrow \frac{64}{27} = \underline{2.37 \text{ times better S/N}}$

- more voxels (with size of voxels,  $\Delta t$  per readstep constant)  $\sqrt{n}$  effect on S/N

↳ e.g.,  $\boxed{64 \times 64 \rightarrow 128 \times 128} \rightarrow \frac{\sqrt{128 \times 128}}{\sqrt{64 \times 64}} = \underline{2 \text{ times better S/N}}$

- # acquisitions  $\sqrt{n}$  better S/N

↳ e.g.  $\boxed{1 \text{ acq} \rightarrow 2 \text{ acq}} \rightarrow \frac{\sqrt{2}}{\sqrt{1}} = \underline{1.41 \text{ times better S/N}}$

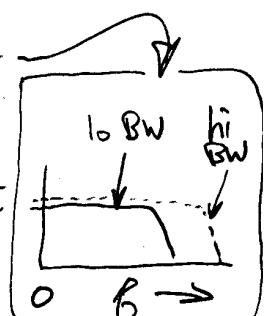
- longer timestep during readout,  $\sqrt{\Delta t}$  better S/N

$$\Delta t = \frac{1}{\text{BW}_{\text{read}}}, \text{ digitization timestep during echo acquisition}$$

-  $\text{BW}_{\text{read}}$  determined by cutoff freq. analog lowpass filter

-  $\Delta t$  controls BW because low-pass cut off has to be set higher for smaller (higher freq-detecting)  $\Delta t$

- must filter out freq's  $> f_{\text{max}} = \frac{1}{2\Delta t}$  because they alias



fourier-0

# COMPLEX ALGEBRA

real / imaginary

$$\text{add: } (r_1, i_1) + (r_2, i_2) = (r_1 + r_2, i_1 + i_2)$$

$$\text{mult: } (r_1, i_1) \times (r_2, i_2) = (r_1 r_2 - i_1 i_2, r_1 i_2 + i_1 r_2)$$

ampl. / phase

$$\text{add: } (A_1, \phi_1) + (A_2, \phi_2) = (A_1 \cos \phi_1 + A_2 \cos \phi_2, A_1 \sin \phi_1 + A_2 \sin \phi_2)$$

$$\begin{array}{l} \text{multiply: } (\text{commutative}) (A_1, \phi_1) \times (A_2, \phi_2) = (A_1 A_2, \phi_1 + \phi_2) \\ \text{divide: } (A_1, \phi_1) \div (A_2, \phi_2) = (A_1 / A_2, \phi_1 - \phi_2) \end{array}$$

$$\text{complex to real power: } (A, \phi)^n = (A^n, n\phi)$$

- don't confuse freq, angle!

$r e^{-i\omega t}$

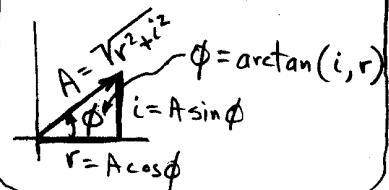
makes linear var. into circular variable

freq. × current = (lin. var.) time

Angle (lin.var.)  $\rightarrow$  complex/circular var.

- how to convert:

$(r, i) \Leftrightarrow (A, \phi)$



$$e^{i\phi} = \begin{bmatrix} \text{expand as series} \\ \text{recognize cos, sin series} \end{bmatrix}$$

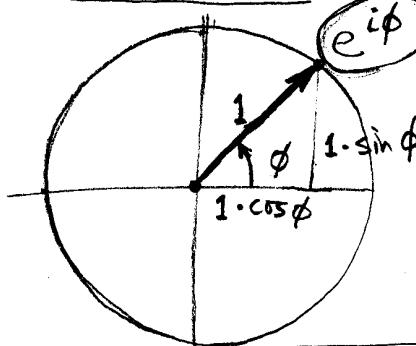
$$= \cos \phi + i \sin \phi$$

$$= \cos \phi, \sin \phi$$

= vector on unit circle

$$e^{i\phi n} = (\cos \phi + i \sin \phi)^n$$

$$= \cos n\phi + i \sin n\phi$$



shorthand for a unit vector (2D)  
pointing in the direction of  $\phi$

→ for arbitrary amplitude, multiply  
 $A e^{i\phi}$

- change in phase is freq  
 $\frac{d\phi}{dt} = \omega$

- phase is integral of freq. variable  
 $\phi = \int \omega dt$

Fourier transform

$$\begin{aligned} H(f) &= \int_{-\infty}^{\infty} h(t) e^{-i2\pi ft} dt \\ H(\omega) &= \int_{-\infty}^{\infty} h(t) e^{i2\pi \omega t} dt \end{aligned}$$

Convolution

$$\begin{aligned} f(x) &= g(x) * h(x) = \int_z g(z) \cdot h(x-z) dz \\ \text{Cross-correlation} \quad f(x) &= g(x) \otimes h(x) = \int_z g(z) \cdot h(x+z) dz \end{aligned}$$

Convolution Theorem

$$\mathcal{F}[g(x) \cdot h(x)] = G(k) * H(k)$$

because of FFT, faster if kernel not small

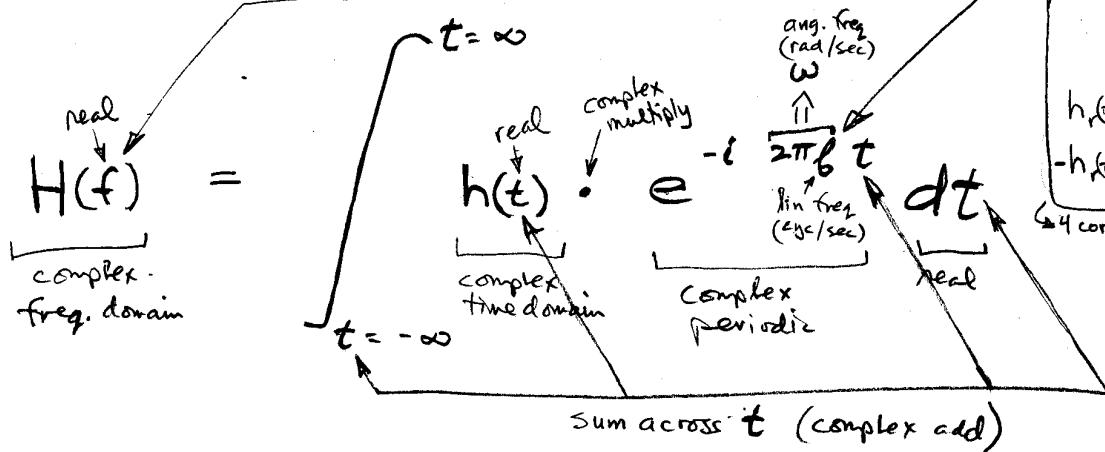
the Fourier transform of two functions multiplied by each other equals the convolution of the Fourier transform of each funct

Fourier-1

# Fourier transform (1)

bt is a phase angle

$$b \cdot t = bt \\ \text{cyc/sec} \cdot \text{sec} = \text{cyc}$$

for one  $f$ 

Fourier integral terms written out

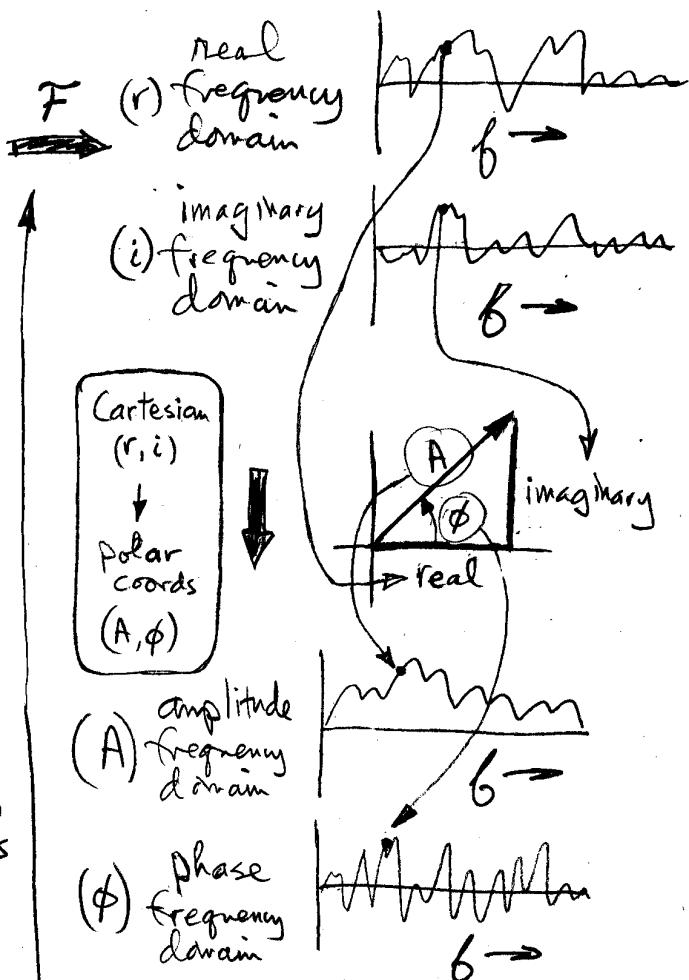
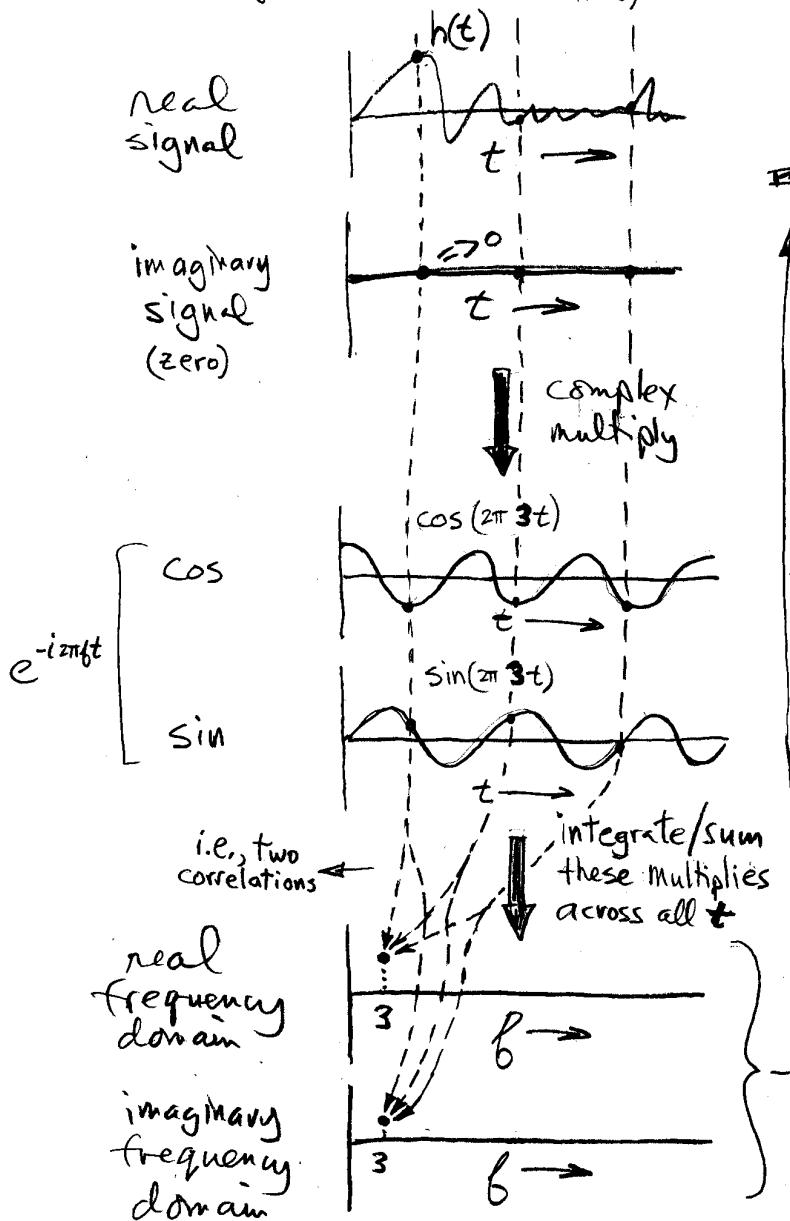
$$h_r(t) \cos(2\pi ft) + h_i(t) \sin(2\pi ft), \\ -h_r(t) \sin(2\pi ft) + h_i(t) \cos(2\pi ft)$$

→ 4 correlations

Fourier integral w/real signal  
 $h_r(t) \cos(2\pi ft),$   
 $-h_r(t) \sin(2\pi ft)$

→ 2 correlations

- How to calculate  $H(f)$  for one  $f$  ( $f=3$ ):  
 (real signal: only need 2 correlations)



like correlating with sin and cos (at each freq.) so we get phase (at each freq.)

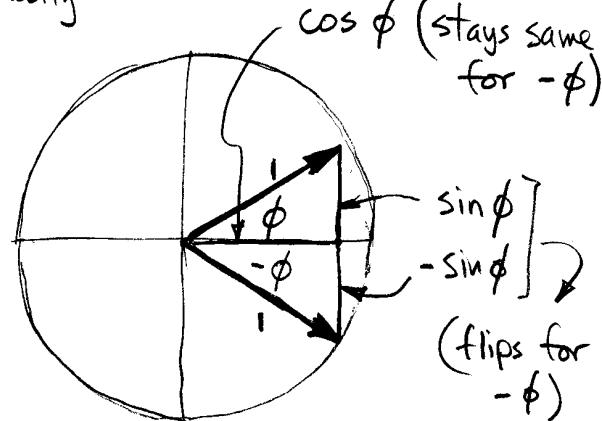
## Fourier transform (1b)

- neg complex exponents/freq
- orthogonality

$$e^{i\phi} = \boxed{\cos \phi + i \sin \phi}$$

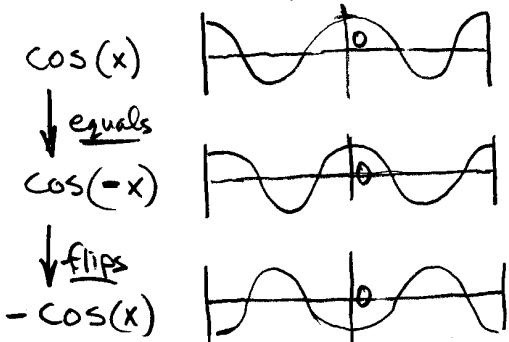
$$\begin{aligned} e^{-i\phi} &= e^{i(-\phi)} \\ &= \cos(-\phi) + i \sin(-\phi) \quad \begin{matrix} \text{no effect} \\ \text{(see below)} \end{matrix} \quad \begin{matrix} \text{move outside} \end{matrix} \\ &= \cos(\phi) - i \sin(\phi) \end{aligned}$$

graphically  $\Rightarrow$



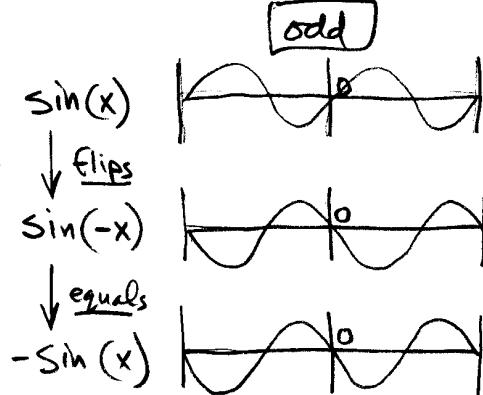
- $\cos$  is an "even" function,  $\sin$  is an "odd" function

even



if a function is mirror-symmetric along the  $x$ -axis around zero:  
 $f(x) = f(-x)$   
it is even

odd



## An orthogonal decomposition

- think of discretely sampled  $\sin(f_1 x)$ ,  $\cos(f_1 x)$  as vectors
- $\text{Corr}(\vec{v}_1, \vec{v}_2) \equiv \text{projection of } v_1 \text{ onto } v_2 \equiv \vec{v}_1 \cdot \vec{v}_2$

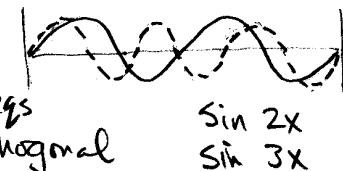
$$\text{Corr}(\cos f_1 x, \sin f_1 x) = 0$$

=  $\sin \frac{\pi}{2} \cos$  of same frequency are orthogonal



$$\text{Corr}(\sin f_1 x, \sin f_2 x) = 0$$

= different integer freqs of  $\sin$  ( $\text{or } \cos$ ) are orthogonal



$$\text{Corr}(\cos f_1 x, \sin f_2 x) = 0$$

[as above]

- in the continuous case, orthogonal functions defined as:

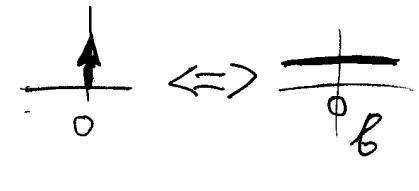
$$\int_{x=l_0}^{x=h_1} f(x) g(x) dx = 0$$

fourier-1c

# UNDERSTANDING INVERSE FOURIER TRANSFORM AS ANOTHER CASE OF CORR w/ COS, SIN

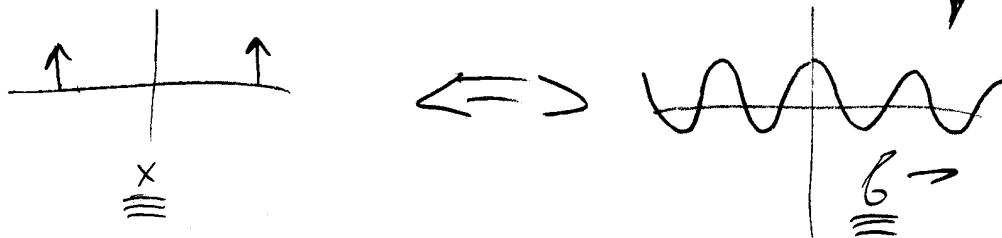
- start with spike in image domain
- take example of spike at  $x=0$

$\cos(x), \cos(2x), \cos(kx)$  all equal 1 there  
all freq's correlate w/ spike at  $x=0$

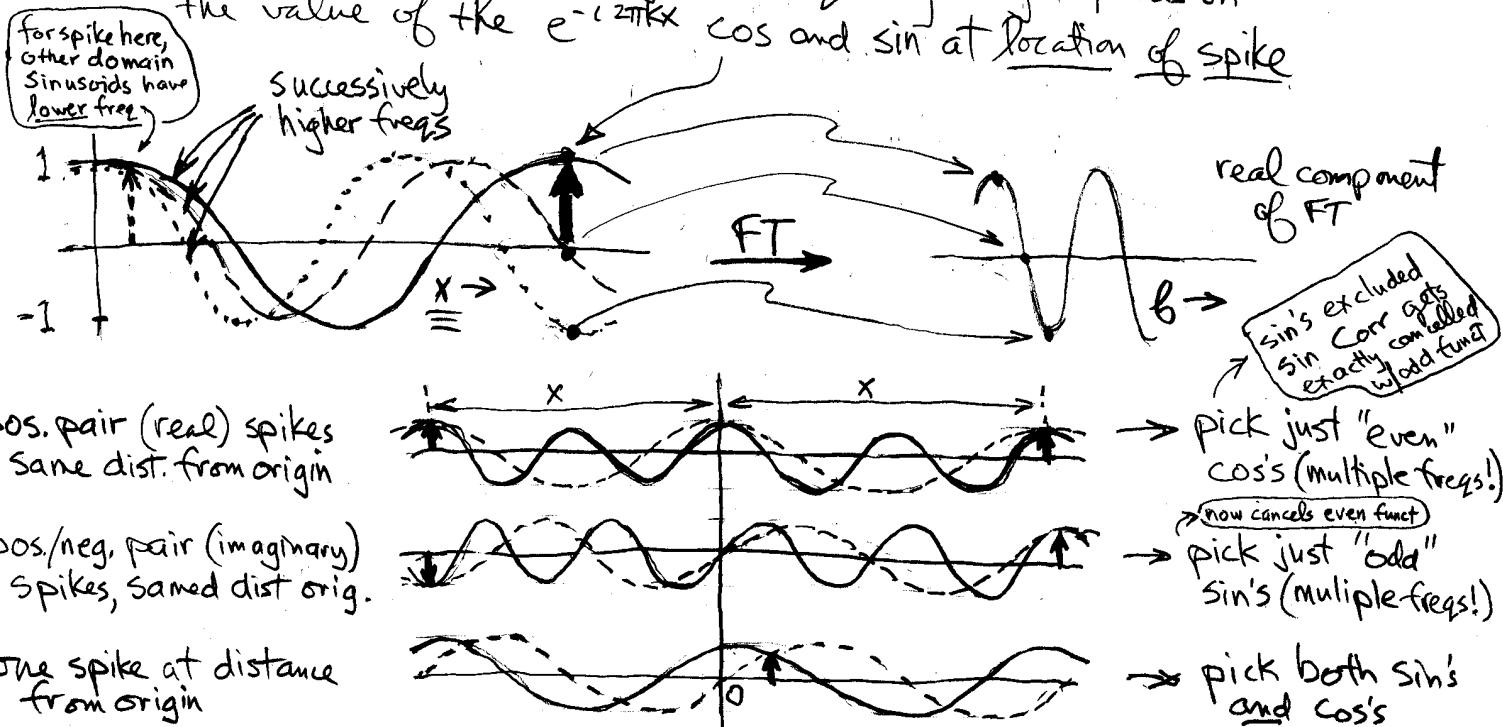


- if spike is moved away from zero, the frequency spectrum obtained by correlating cos with spikes oscillates

N.B. opposite direction sin spike are on imaginary axis



- to see why this is, since spike is all zero except at spike, the dot product for a given frequency only depends on the value of the  $e^{-i2\pi kx}$  cos and sin at location of spike



- pos. pair (real) spikes same dist. from origin

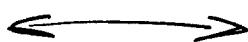
- pos./neg. pair (imaginary) spikes, same dist. orig.

- one spike at distance from origin

→ this is one way of thinking about what one point in k-space means, via correlating it w/ cos's, sin's to get periodic result in image space (inverse FT)

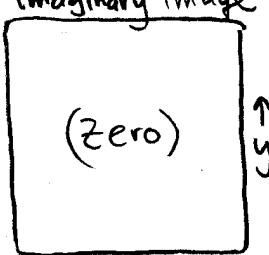
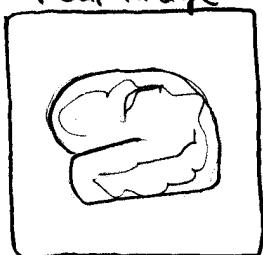
# FOURIER TRANSFORM OF AN IMAGE (2)

(1)



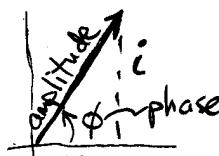
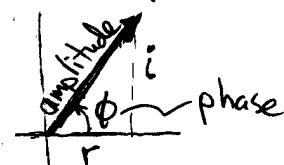
real image

imaginary image



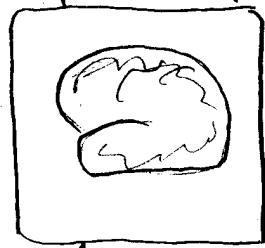
real spat. freq.

imag. spat. freq.

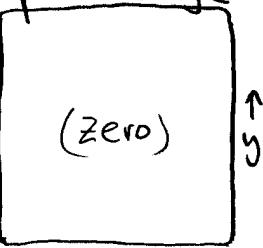
Fourier  
TransformInverse  
Fourier  
Transform

(2)

amplitude image



phase image

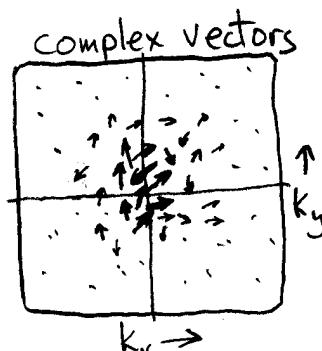
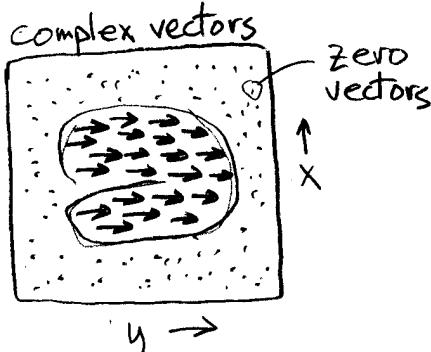


ampl. spat. freq.

phase spat. freq.

view complex  
vectors directlyview complex  
vectors directlywhat you  
see on  
screen

(3)

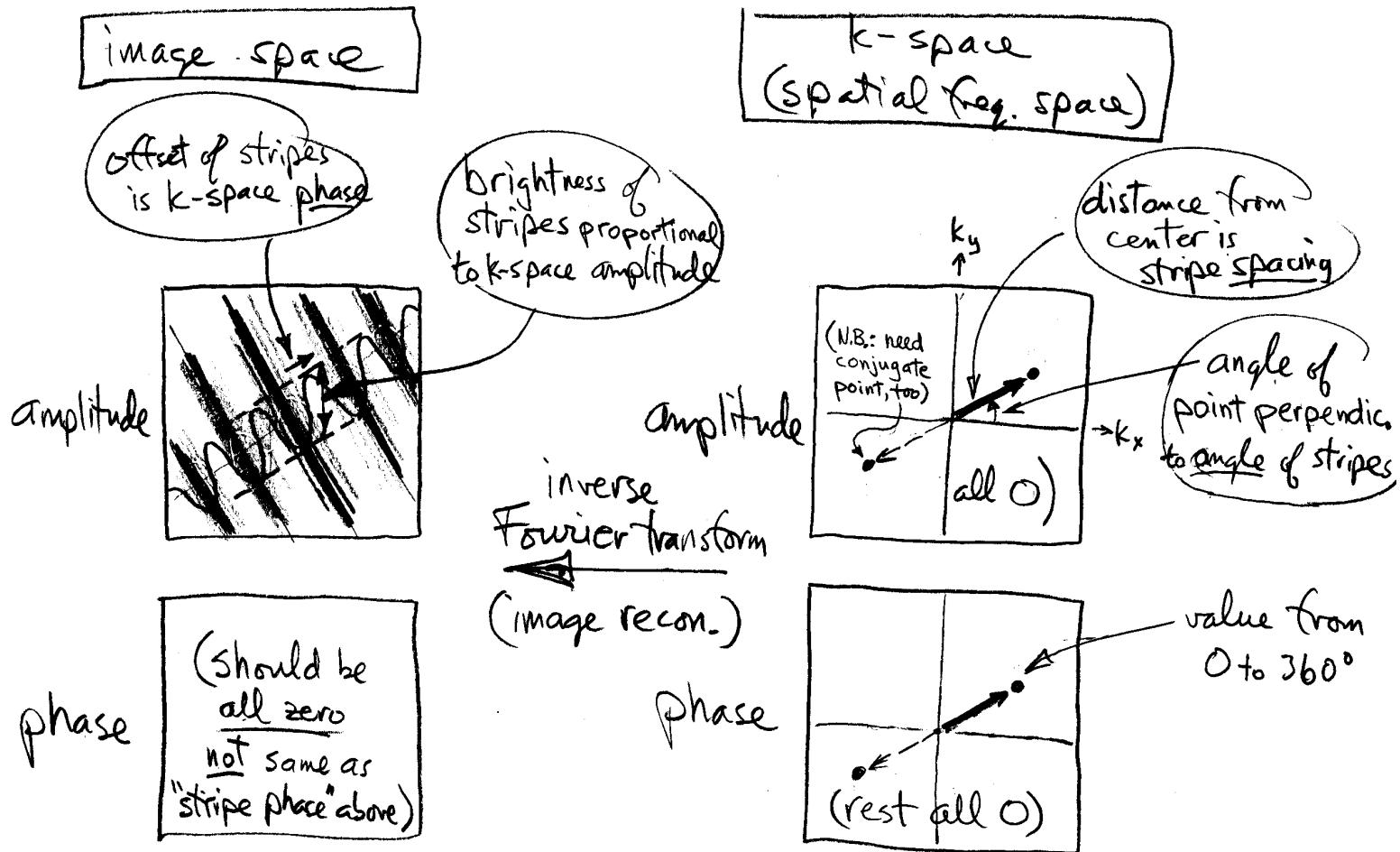
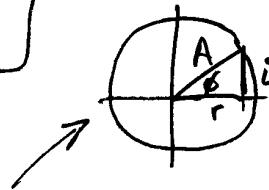


- 3 equivalent representations of image & spat. freq. space

Fourier-3

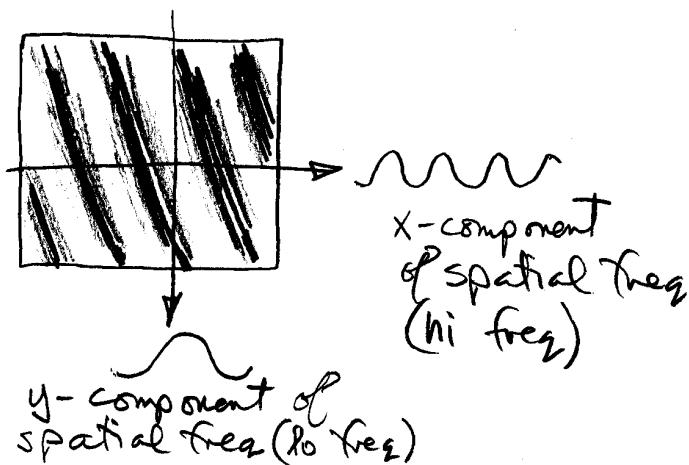
# FOURIER TRANSFORM OF REAL IMAGE (3)

- what a single k-space point looks like  
 for real image (polar coordinates  $A, \phi$  instead of  $r, i$ )  
 (actually pair of points)



- Cartesian dimensions of k-space — x- and y- spatial freq

N.B.: each dimension of spatial freq. space (k-space) from correlation w/  $\sin + \cos$  — don't confuse  $k_x, k_y$  w/  $\sin, \cos$ !

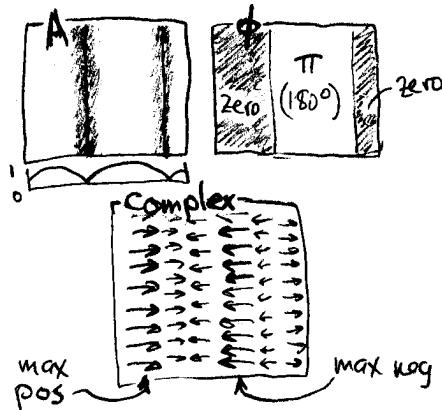
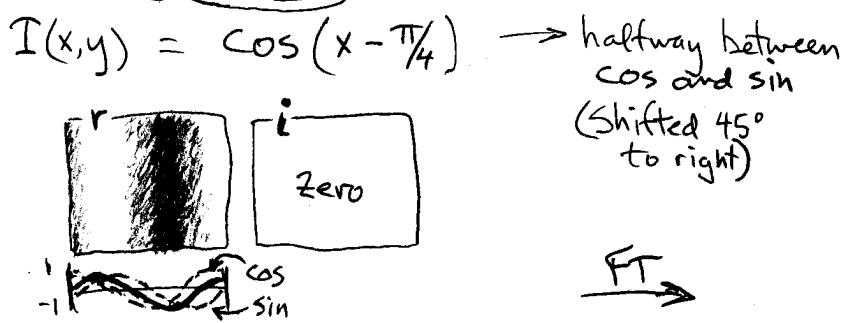
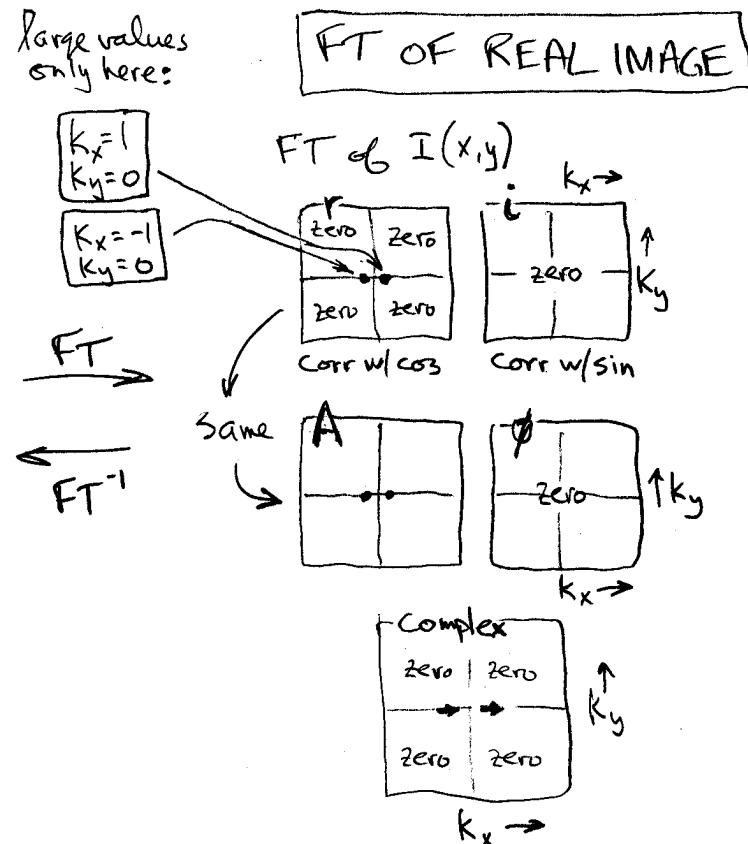
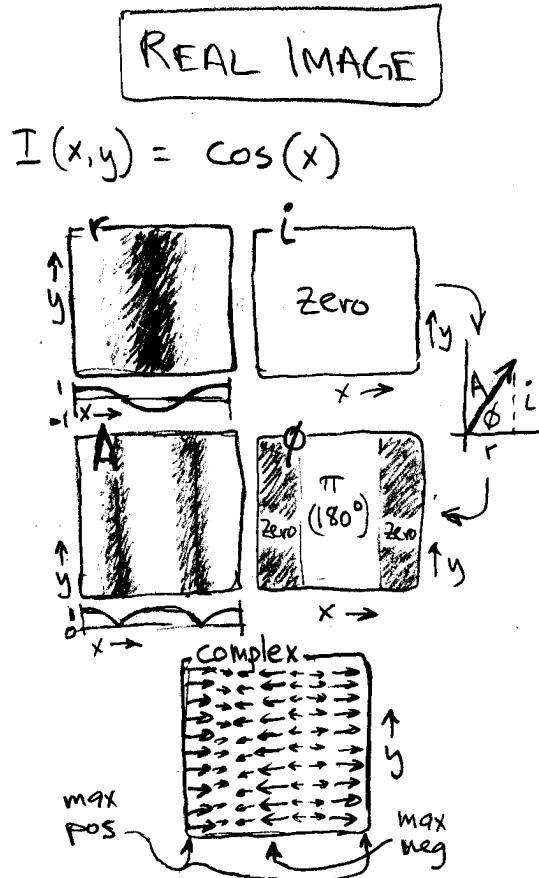


N.B. — increasing one 1D component increases the spatial freq of the 2D wave and rotates it

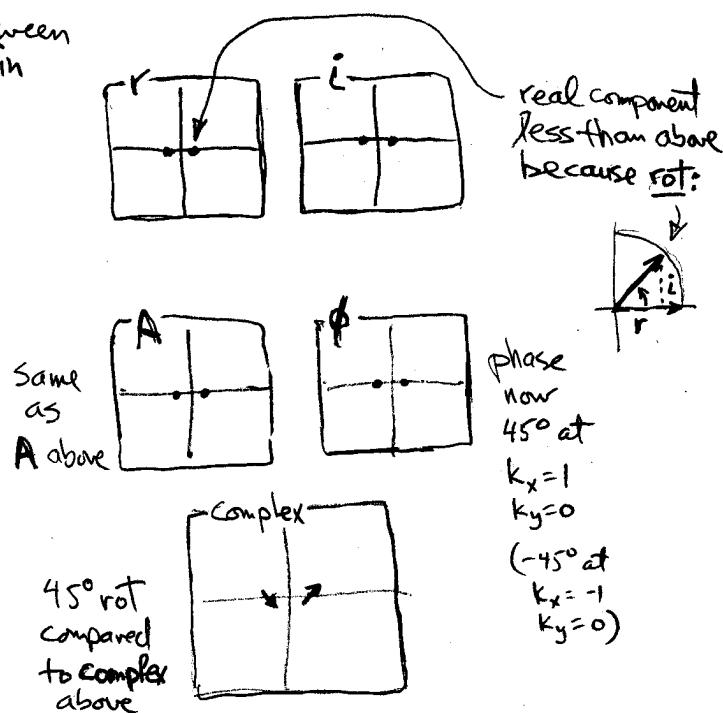
fourier-3b

# FOURIER TRANSFORM OF IMAGE (4)

- 3 equivalent representations of complex numbers in image space and spatial-freq. Space ( $k$ -space)
- example: cosinusoid in image space, then shifted in x-dir



N.B.: an example of the "Fourier Shift Theorem" (see below)



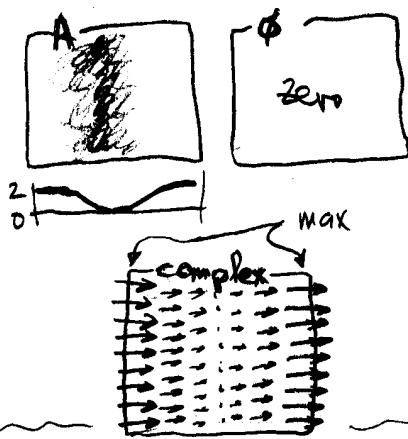
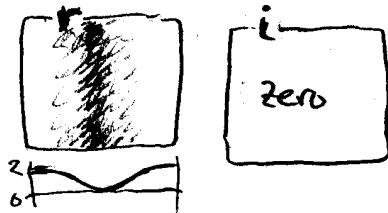
fourier-3c

## FOURIER TRANSFORM OF IMAGE (5)

- (cont.) center of k-space (real image)
- complex image

### REAL IMAGE

$$I(x, y) = \underline{1} + \cos(x)$$



### center of k-space:

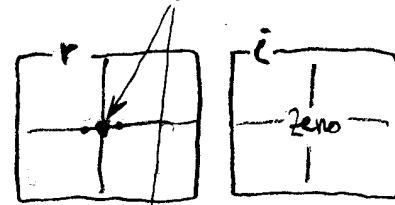
$$H(k) = \int_x h(x) \cdot e^{-i2\pi kx} dx$$

avg image brightness  $\Leftrightarrow$  1 (real)

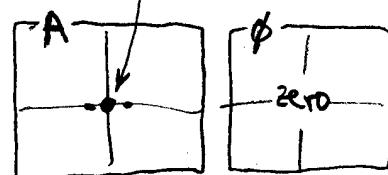
$\xrightarrow{\text{FT}}$

### FT OF REAL IMAGE

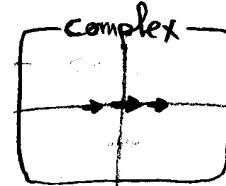
positive center k-space



$\xleftarrow{\text{FT}^{-1}}$

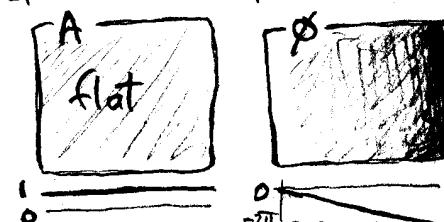
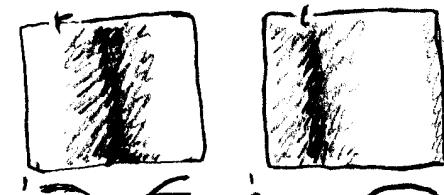


[the center of k-space is zero w/ pure sin or cos image b/c avg. brightness = 0]



### \*COMPLEX IMAGE

$$I(x, y) = \cos(x) - i \sin(x) = e^{-ix}$$



### FT, FT<sup>-1</sup>

(missing "spike" results in single spike correlating with cos and sin)

N.B.: this k-space is non-Hermitian:

K-space will only have Hermitian symmetry if image is real:

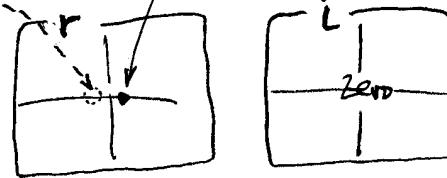
Hermitian symm. when complex conjugate (= complex num w/ sign flipped in imag. part) is equal to funct value w/ neg arg:

$$1D: H(-k) = H^*(k)$$

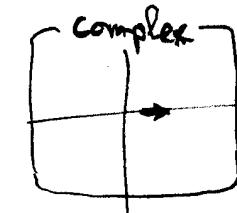
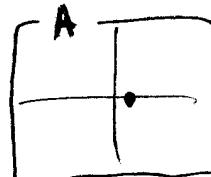
$$2D: H(-k_x, -k_y) = H^*(k_x, k_y)$$

### FT OF COMPLEX IMAGE

spike only on one side of k-space



[N.B. this is like what an artifact "spike" does tho it would have rand. phase]



[N.B. this is also exactly what a gradient does to image space!]

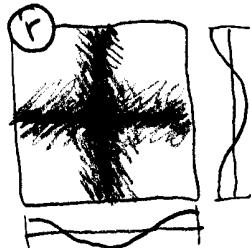
fourier-3d

# FOURIER TRANSFORM OF IMAGE (6)

- (cont.) x- and y-spatial freqs.
- special case: real image from sum of reals

## REAL IMAGE

$$I(x,y) = \cos(x) + \cos(y)$$

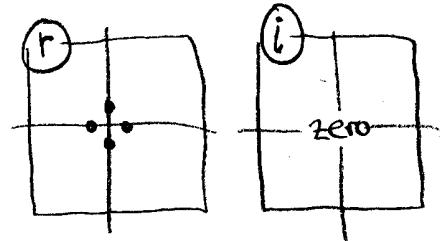


N.B. adds but  
doesn't rotate  
stripes

zero

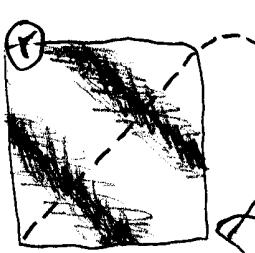
$$\xrightarrow{\text{FT}} \quad \xleftarrow{\text{FT}^{-1}}$$

## FT OF REAL IMAGE



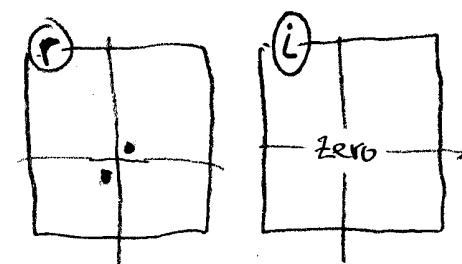
$$I(x,y) = \cos(x+y)$$

rotates  
stripes!



zero

$$\xrightarrow{\text{FT}} \quad \xleftarrow{\text{FT}^{-1}}$$



- Remember, single k-space point transforms to complex img.

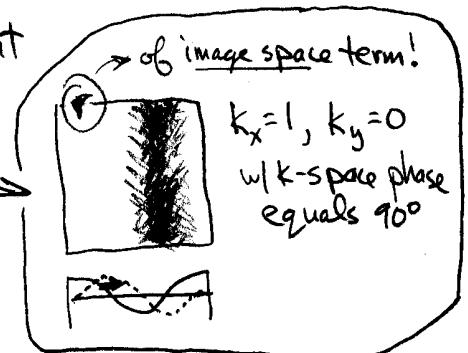
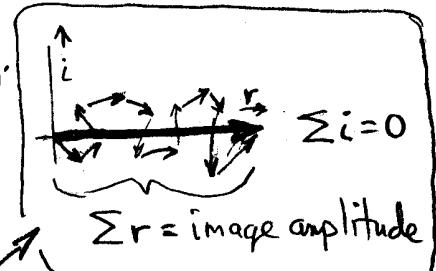
- but if Hermitian Symmetry, imaginary components cancel

- Since all we want in image space reconstruction is real component, can just add real components of complex vectors at each image space point for every complex image corresponding to each k-space point

- N.B.: the k-space phase will affect offset of real valued image space cosinusoid

- therefore for real-valued image, we can visualize inverse FT as real-valued sum of offset real-valued cosinusoids

- N.B. cannot do this with MRI k-space data since phase errors (incl. multiple wraps) mess up real component  $\Rightarrow$  must use amplitude img



$\phi_i$

## GRADIENT COILS

- gradient coils for  $x, y, z$  generate approximately \*  
a linear gradient in the strength of the  
 $\underline{z}$ -component of the magnetic field  $B_z$

all effects of gradients discussed  
before were in  $z$ -direction (parallel  
to  $B_0$ )

- for example, the  $x$  gradient coil induces a ramp in  $z$ -component of the magnetic field when moving in the  $x$ -direction:

$$B_{G,\underline{z}} = G_x x$$

- \* - since a pure linear gradient of  $B_{G,\underline{z}}$  in only the  $x, y$ , or  $z$  directions is not possible according to Maxwell equations,  
each gradient coil also induces a magnetic field that has components in the  $x$ - and  $y$ -direction ( $B_{G,x}$  and  $B_{G,y}$ )

- the other magnetic field components are usu.  
ignored because they are so small relative  
to  $B_{G,z}$ , since  $B_{G,z}$  is added to  $B_0$ , and  
since  $B_0$  is much stronger than  $B_{G,x}, B_{G,y}$ , &  $B_{G,z}$

- since standard reconstruction methods assume  
the existence of "non-Maxwellian" gradient  
fields, spatial distortion is introduced

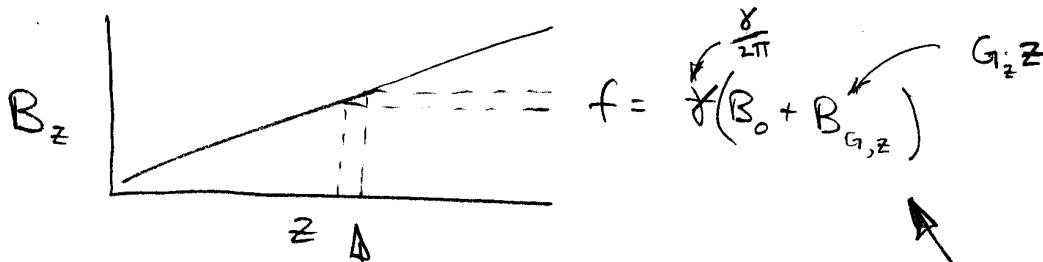
- the Maxwellian terms  $B_{G,x}, B_{G,y}$  are known;  
can be included in the recon. process

e.g. extra phase in  $z$ -dir  $\perp$  to  $x$  grad

$$\Delta\phi G_x(\vec{x}) \approx -\frac{x^2 G_0^2 t}{2B_0}$$

## SLICE SELECTION ( $G_z$ )

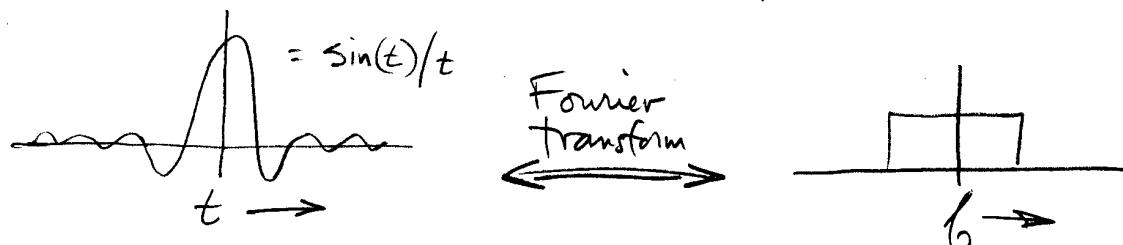
- slice select gradient on during RF stim



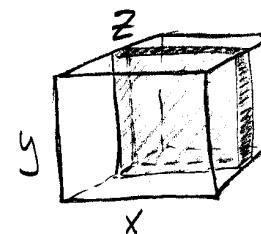
- protons here can only be excited by a narrow band of radio frequencies

- to apply a pulse containing a narrow band of frequencies, we use a sinc pulse envelope (Fourier transform of a narrow freq. band)

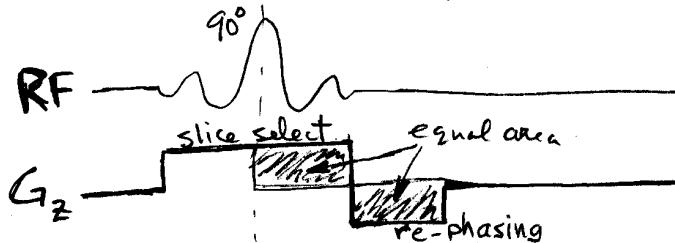
in practice,  
Gaussian  
pulse envelope  
good, too



- this excites protons in a narrow slab



- since the slice-selection gradient introduces (space-dependent) phase shifts (see freq encode) these have to be removed by a post-excitation rephasing  $\pm$ -gradient



- approximation from assuming tip occurs instantaneously in middle
- valid for small tip:  $90^\circ \rightarrow 52\%$
- in practice: adjust to max, use crusher to kill spurious transverse on  $180^\circ$

slice-3

# PULSES FOR SLICE SELECTION

Fourier approach details

- Fourier transform approach to slice-selective pulse (linear approx. even tho tipping is non-linear)

$$\vec{B}_1(t) \propto \int_{f=-\infty}^{f=\infty} p(f) \cdot e^{-i 2\pi f t} df$$

time dependent RF stimulation (complex)

frequency selection function

i.e., time-dependent complex (=quadrature) pulse waveform in Fourier transform of frequency spectrum of RF pulse

Solve with:  $p(f) = \text{frequency band:}$

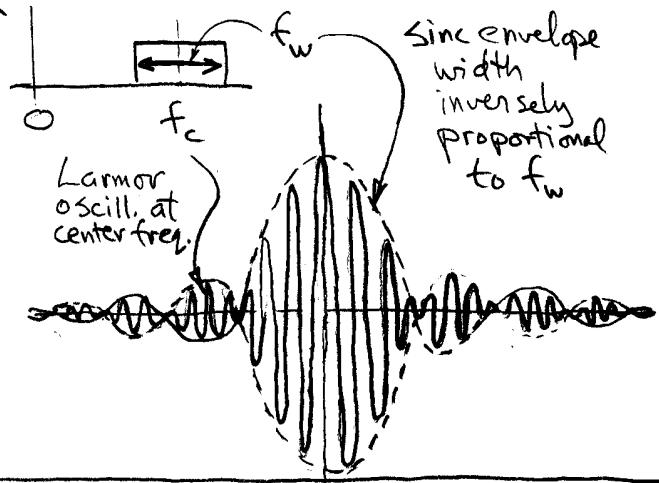
$$\vec{B}_1(t) = A \cdot f_w \cdot \text{sinc}(\pi f_w t) \cdot e^{-i 2\pi f_c t}$$

Amplitude controlling flip angle

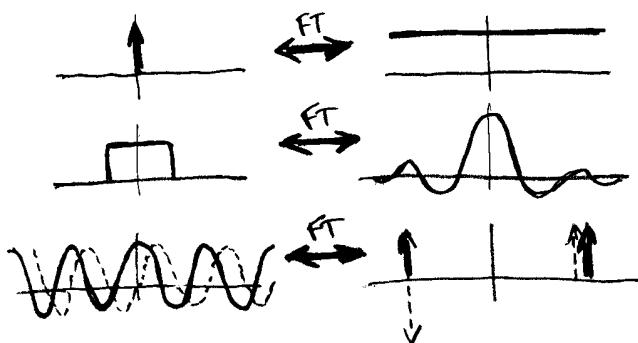
freq width (controls slice width)

sinc envelope determined by freq. width,  $f_w$  (N.B. wider  $f_w$  is narrower sinc)

modulation (complex) at center freq.,  $f_c$



## Fourier Transform Pairs, Rules



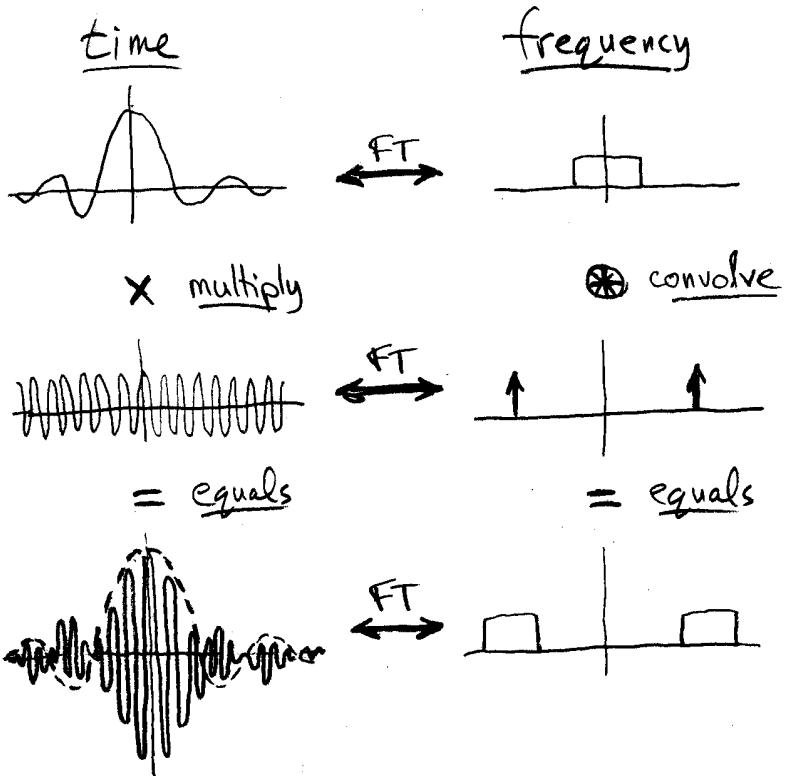
- multiplication in one domain equals convolution in other:

$$F[g(t) \cdot h(t)] = G(f) \otimes H(f)$$

means do FT

- convolution with delta funct impulse moves other function to impulse center

## Fourier Transform Solution to: $\frac{d}{dt} f(t)$



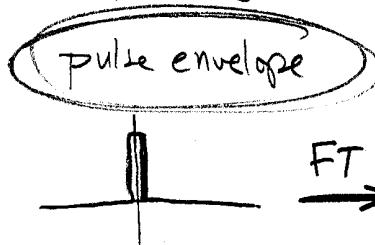
slice-4

## SLICE SELECT RF PULSES

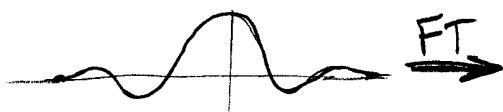
Interleaved Acquisition  $\Rightarrow$  better S/N b/c imperfect slice profile  
 spin history prob if motion

### Common RF pulses

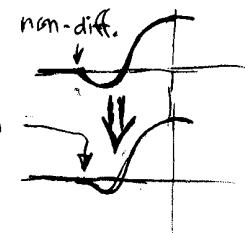
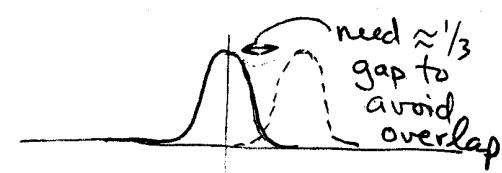
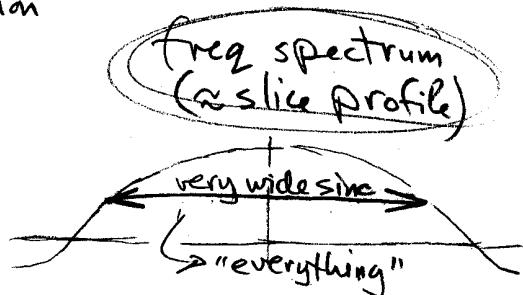
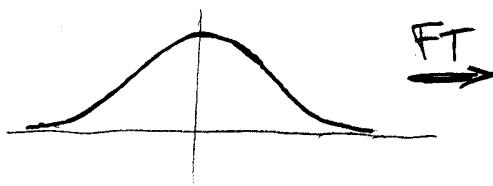
non-selective pulse  
 ("hard" pulse)



standard slice select sinc



Gaussian



### Fat Saturation

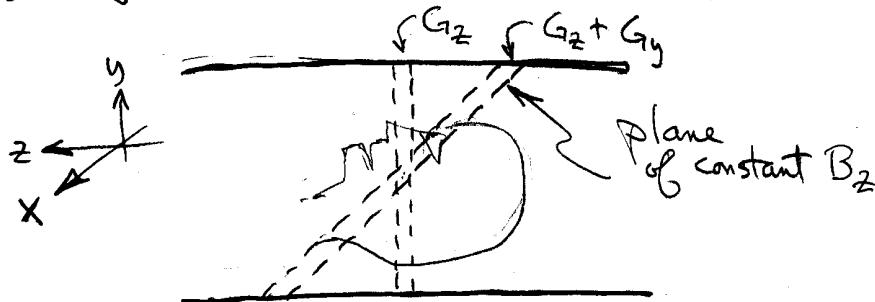
- fat protons have chemical shift causing resonant freq offset
- add phase offset not due to gradients, RF
- fix by off-water-resonance  $90^\circ$  (saturation) pre-pulse centered on fat freq  
 $\hookrightarrow$  need high quality (narrow freq) pulse to avoid saturate water!

HowTO

- 1) fat sat pulse
- 2) wait  $T_2$  so fat signal decays, but no  $T_1$  regrowth of fat
- 3) RF stim for water "protons- $\alpha^2$ -interest"



### Adding Another Gradient Tilts Slice



- with 3 gradients on, can excite arbitrary angle plane
- translate plane by changing either gradient amplitude or RF freq band:  $\frac{FT}{I}$

freq. encode-0

## WHY "FREQUENCY-ENCODING" IS A MISNOMER

- comes from original analogy in Lauterbur (1973):



- 1) chemical shift change freq → gradient changes freq.
- 2) stimulate w/ broadband RF → same
- 3) time-sample FID containing multiple freqs → same
- 4) FT of FID to get spectrum  
 ↪ ~~multiple~~  $\frac{1}{f}$  of  $\Delta f$  offsets → FT of FID to get  $\Delta x$  offsets

- this is technically correct (FT of FID) but highly misleading

↳ e.g., phase-encoding (turning a different gradient ON and OFF before recording FID) seems to be something completely different since OFF gradient can't affect freqs in FID

- the "k-space" perspective is a "Copernican turn"

↳ idea is that data is not a set of samples of a time domain signal generated by multiple chemical-shift like frequencies

→ rather, it is a set of samples of a frequency-domain signal, each sample generated by multiple spatial locations, (which are analogous to multiple time points)

- i.e., the 'direction' of the FT (Fourier transform) is reversed:

	Signal	result
spectroscopy	samples of oscillations in <u>time-domain</u>	FT → frequency-domain spectrum of shifts
MRI	samples of spat. freq. in <u>freq. domain</u>	FT <sup>-1</sup> → spatial object (like a <u>time-domain</u> signal)

N.B. not full set all freqs like gradient makes!

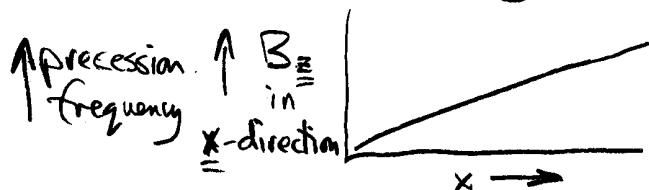
- the original analogy only 'works' because  $FT \approx FT^{-1}$  (except sign change)

freq encode-1

## FREQUENCY ENCODING (1)

avoid this mistaken intuition!

- frequency encode gradient ( $G_x$ ) causes precession rates to vary linearly in x-direction



↳ correct (remember that strength of  $G_x$  causes variation of slope of  $B_z$  in x-direction)

- different frequency signals are mixed together and recorded as a 1-D signal over time

↳ correct, but remember, we are recording summed local magnetization vectors after de-modulation

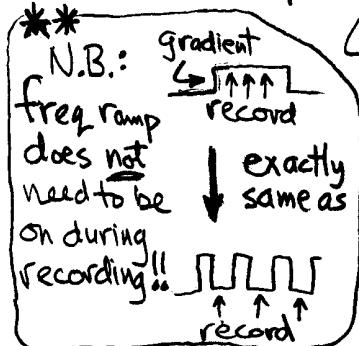
- a Fourier transform, which can convert back and forth between x-position (cf. time) and spatial frequency (cf. temporal freq) is done on signal

↳ correct

- spatial frequencies get confused/confused with precession frequencies

↳ wrong !!

- therefore, the Fourier transform is used to convert Position-dependent precession frequencies into spatial position



conceptually  
wrong !!

inverse

FT actually converts spatial frequencies to spatial position

→ the spatial frequency increases for each time point in the readout

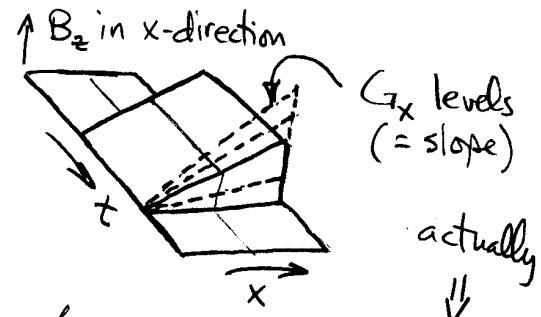
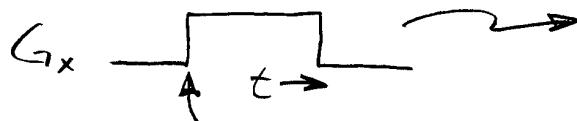
→ the precession freq ramp is constant each timestep

freq encode - 2

## FREQUENCY ENCODING (2)

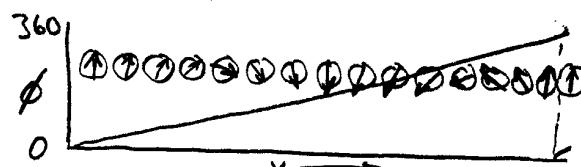
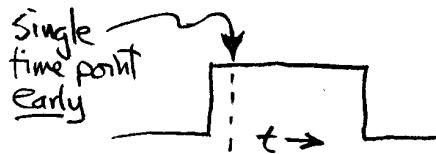
connect intuition - why phase critical

- "frequency"-encode gradient ( $G_x$ ) turned on during echo causes precession rates to immediately vary with  $x$ -position



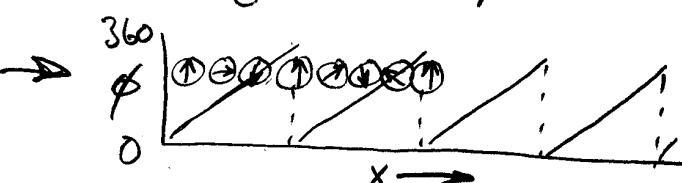
- at beginning of gradient on, the phase of signal coming from each  $x$ -position is the same  
Summed phase angle is what we measure

- early after gradient on, phase advances (because of faster precession frequency) arise with greatest phase advance at largest  $x$ -position



$\Rightarrow$  one cycle of spatial frequency of phase angle  
 $(= \text{low spatial freq.})$

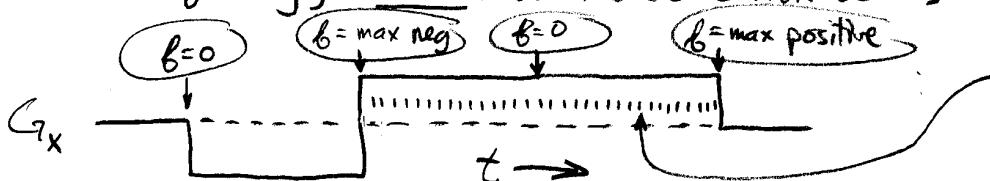
- later during gradient on, phase advances cause multiple wraparounds of phase angle across space



$\Rightarrow$  multiple cycles of spatial frequency of phase angle  
 $(= \text{hi spatial freq.})$

- in practice, the lowest spatial frequency ( $= 0$ ) occurs in the middle of the gradient on time because the phase is "wound" negatively by a preparatory gradient (to the highest negative spatial frequency) before data collection occurs

$b$  is spatial frequency



individual RF data samples (after demodulation)

freq encode - 3

## FREQUENCY ENCODING (3) why each datapoint is 1 spatial freq

Standard Fourier transform : (Temporal freq  $\longleftrightarrow$  time)

$$H(f) = \int_{t=-\infty}^{t=\infty} h(t) \cdot e^{-i 2\pi f t} dt$$

freq domain      time domain

$\hookrightarrow r, i=0$

sum across all time

"k" is often used instead of "f" for the frequency variable

Imaging equation : (Spatial freq.  $\longleftrightarrow$  space)

$$S(f) = \int_{x=-\infty}^{x=\infty} I(x) \cdot e^{-i 2\pi f x} dx$$

↓

one data point (i.e., one spatial freq) during readout (2 components)

↓

Signal strength at one x-position (brightness of image point)

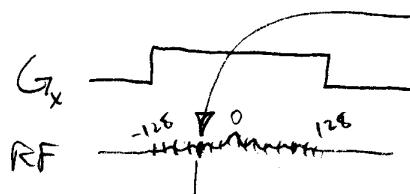
↓

cos (2 $\pi$ f x),  
-sin (2 $\pi$ f x)

↓

Sum across x of object

this is done by RF coil recording sum



get this single readout point by summing signal across x-position (RF coil records sum)

even though variable is  $f$ , it represents one time point during readout

To make image, do inverse Fourier transform of recorded signal  
 $S(f)$

Oscillations come from readout phase wrapping, where  $f$  is single spatial freq (e.g. 5) and  $x$  goes across object

$\downarrow$   $\begin{cases} \text{FID: } G_{xt} \\ \text{SE: } G_x(t-TE) \end{cases}$

$f = G_x t$ , that is, spatial freq depends on amount of time gradient was on (this  $f$  increases with time!)

don't confuse with instantaneously changed precession freqs which are constant across entire readout time (for each  $x$  position)

freq encode - 4

## ALTERNATE DERIVATION (incl. effects of $G_x$ ) SIGNAL EQ

- oscillators at  $\omega = \gamma B$  at each position (just  $x$  for now)

$$S(t) = m(x) e^{-i\phi(x)} dx$$

- by definition, freq,  $\omega$  is rate of change of phase,  $\phi$

$$\frac{d\phi(x,t)}{dt} = \omega(x,t) = \gamma B(x,t) \quad \text{and} \quad \phi(x,t) = \int_0^t \omega(x,t) dt = \gamma \int_0^t B(x,t) dt$$

- assuming phase initially 0,  $B$  affected by gradients

$$B(x,t) = B_0 + G_x(t) \cdot x$$

$$\begin{aligned} \phi(x,t) &= \gamma \int_0^t B_0 dt + \left[ \gamma \int_0^t G_x(t) dt \right] x \\ &= \omega_0 t + (2\pi k_x(t) x) \end{aligned}$$

$k$  is time integral of gradient waveform

- demodulation removes the  $B_0$ -caused carrier frequency  $e^{-i\omega_0 t}$  from the first equation

$$S(t) = \int_x m(x) e^{-i(2\pi k_x(t)x)} dx$$

amplitude of each oscillator      gradient-controlled phase

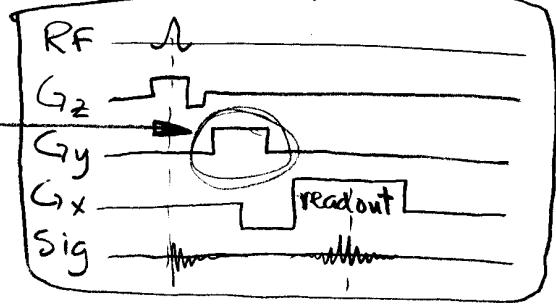
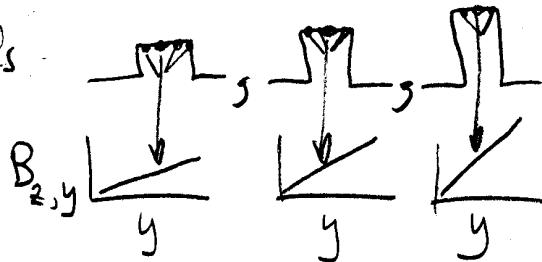
phase-1

## PHASE-ENCODE GRADIENT $G_y$

Pulse Sequence

- Turn on gradient after excitation but before readout

- different levels of  $G_y$



- higher levels of  $G_y$  (slope of  $B_z$  in  $y$ -direction!)  $\hookrightarrow$  higher spatial freq. (more phase wraps) in  $y$ -direction
- phase wraps persist after phase-encode gradient off
- read-out gradient ( $G_x$ ) phase wraps then add to phase-encode phase

## 2D Imaging Equation

$$\underline{S(k_x, k_y)}$$

Signal recorded  
at single time point  
(one readout point)

$\Downarrow$   
complex signal  
(from phase-  
sensitive detection)

$= \iint$   
Sum across  
 $x-y$  plane

$\Downarrow$   
done by  
RF coil

$\underline{I(x, y)}$   
image  
(= strength of  
magnetization  
at each  $x-y$  pt)

$\Downarrow$   
scalar (what  
we try to  
reconstruct)

$$k_x = \frac{G_x t}{\text{readout time}} \quad k_y = \frac{G_y t}{\text{readout time}}$$

fixed strength  
 $\Downarrow$   
 $k_x = \frac{G_x t}{\text{readout time}}$   
 $k_y = \frac{G_y t}{\text{readout time}}$   
 $\Downarrow$   
 $\text{increases each TR}$

$$e^{-i 2\pi (k_x x + k_y y)} dx dy$$

phase (vector of  
unit length and particular  
angle which is function  
of  $G_x$  and  $G_y$ )

$\Downarrow$   
phase angle (of scalar  
magnetization!) in rotating  
frame, set by gradients

- ignoring relaxation, spatial frequency  $k_x$  and  $k_y$  have no "inertia" — they stay wherever the gradients last left them

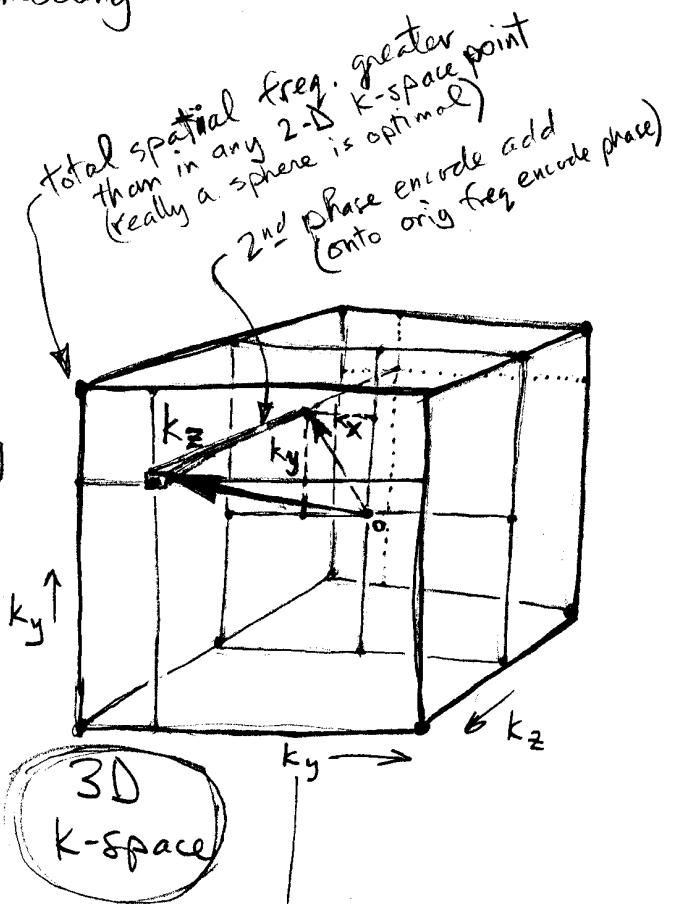
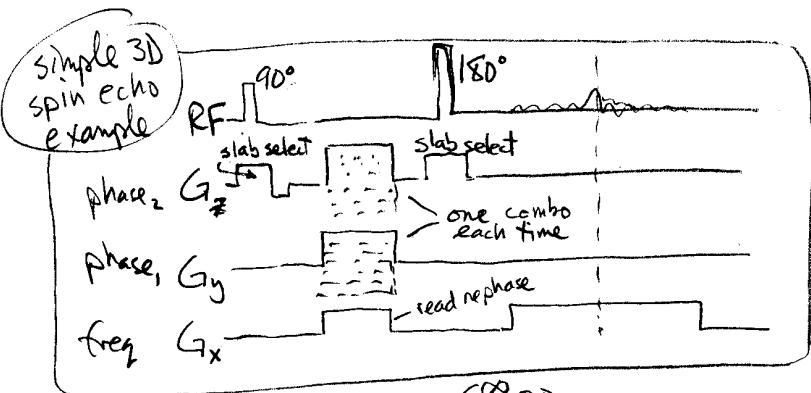
## 3-D IMAGING

- two phase-encode gradients

- use z-gradient for 2<sup>nd</sup> phase-encoding instead of slice selection

- excitation of whole slab (slice-select is whole brain)

- simple spin echo example  
(in real life, usu. done with echo trains [FSE] or small flip angle to allow short TR [SPGR])



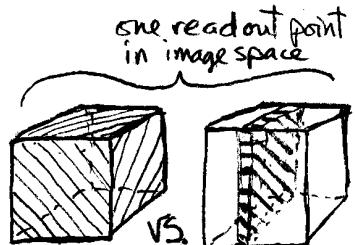
$$S(k_x, k_y, k_z) = \int_{-\infty}^{\infty} \int_{-\infty}^{\infty} \int_{-\infty}^{\infty} I(x, y, z) e^{-i 2\pi (k_x x + k_y y + k_z z)} dx dy dz$$

signal recorded at single point of readout

- i.e., freq-encode phase, first phase-encode phase, and second phase-encode phase all just add (= 3D rotation of phase stripes)

- S/N much better than 2-D because each entire excited volume contributes to signal from each pulse instead of just slice
- phase stripes created throughout volume vs. slice:

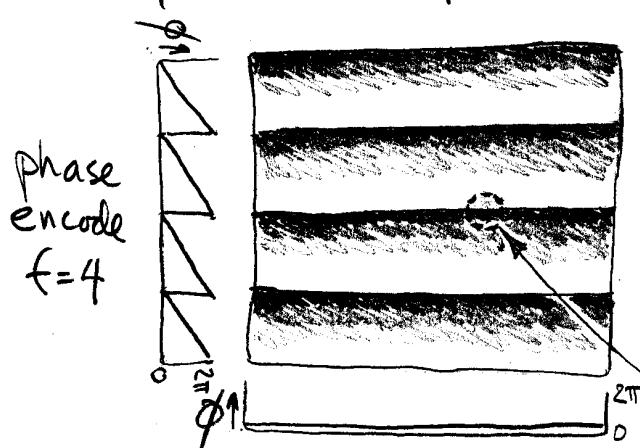
N.B., this ignores relaxation effects for now



phase-3

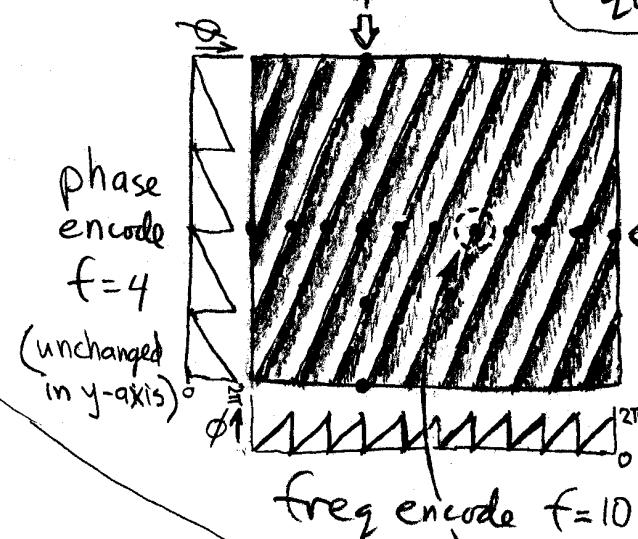
# PHASE & FREQ, 2D & 3D

- Since the phase-encode gradient and the freq encode gradient both affect phase the result is a rotation of phase "stripes" when the two add



Pictures of phase in image space

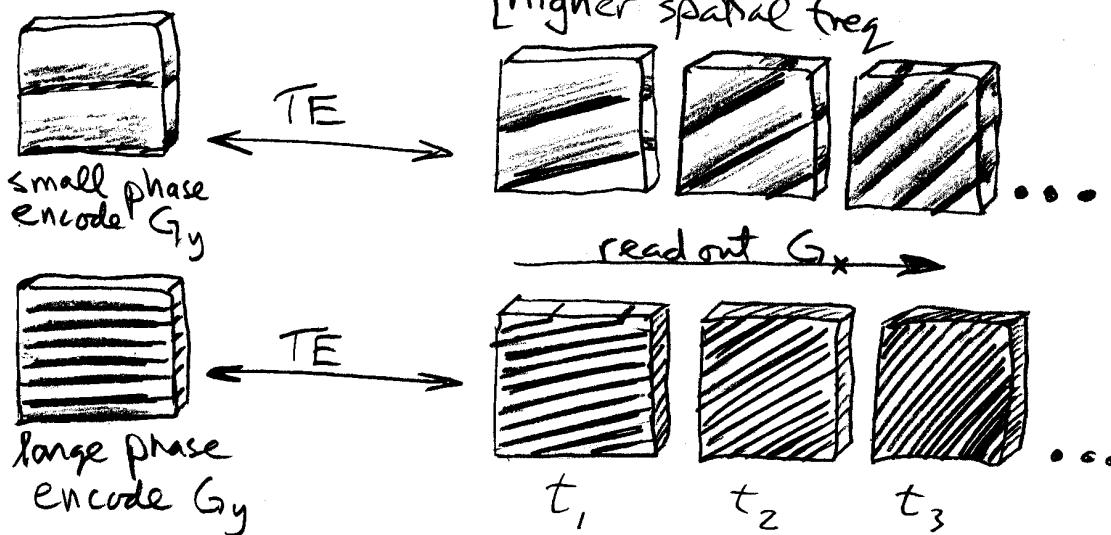
N.B.: Stripes have sharp edges from phase map (not sinusoid since  $\phi$  from 2-comp quadrature!)



stripes here represent complex value

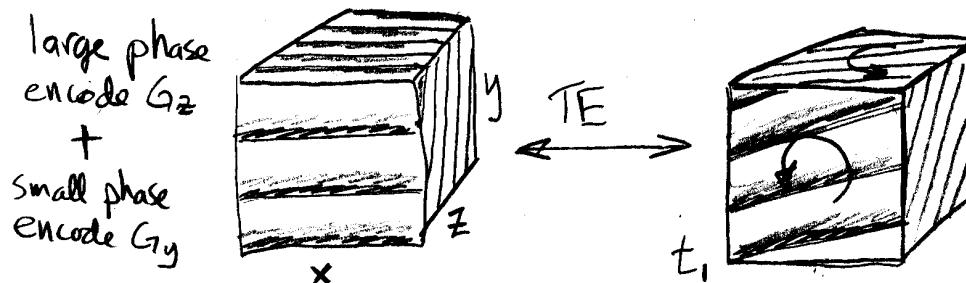
↓  
phase of whole image summed to one(complex) number by RF coils

- successive read out steps:



e.g., after y-gradient, spins at a point might be 2 cyc ahead while after x-gradient spins at same pt 8 cyc ahead: but counting wraps in y-direction, still only 2 ahead

- 3D phase encode w/  $G_y$  and  $G_z$  starts rotated in y-z plane



## GRADIENTS MOVE K-SPACE LOCATION OF DATA POINT

- k-space (spatial-frequency space) location is set by integral of gradient over time up to recording point

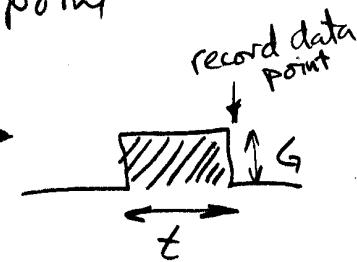
$$k = \star \int_0^{t=\text{recordtime}} G(t) dt$$

spatial freq recorded at t = recordtime  
 gradient strength as function of t

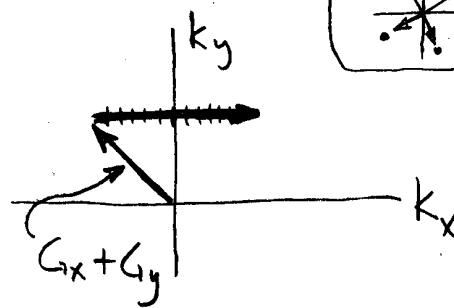
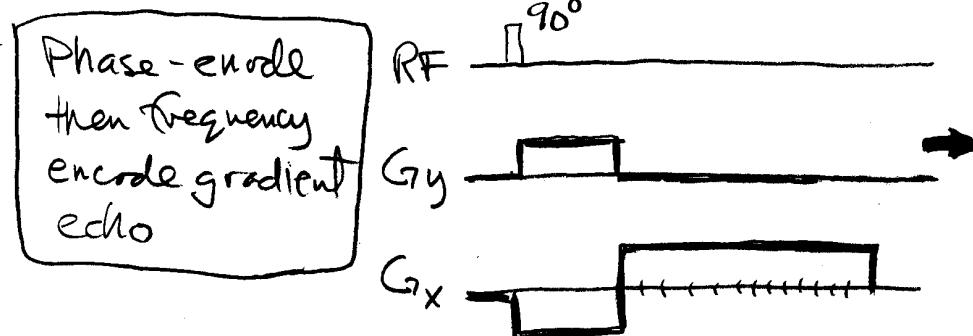
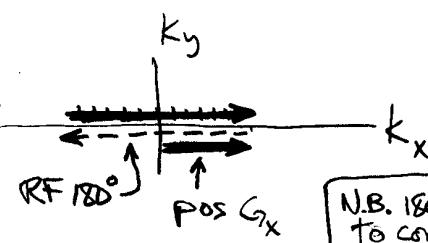
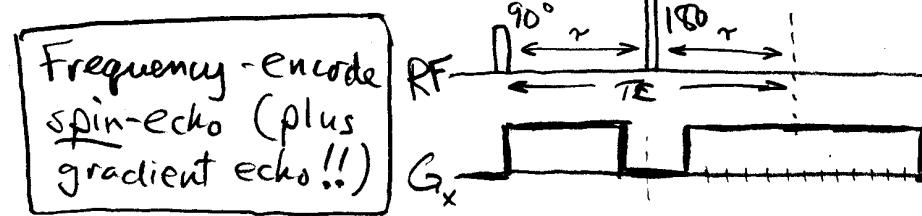
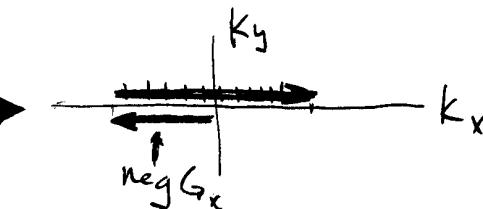
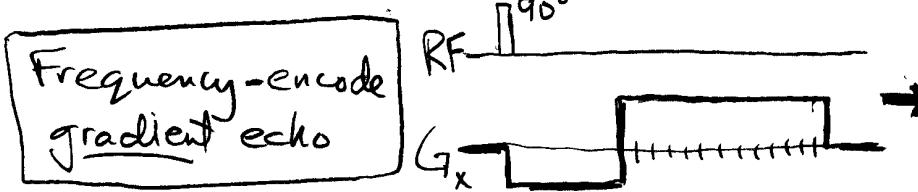
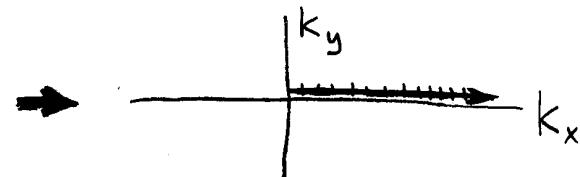
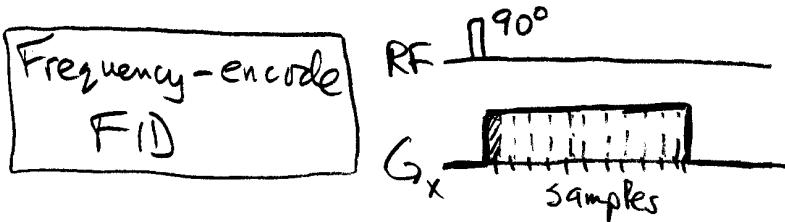
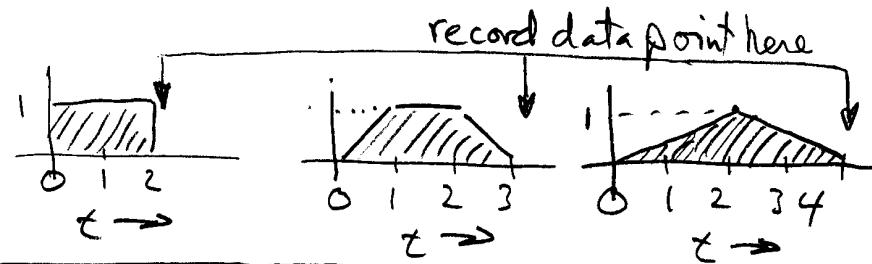
simple form of integral w/ boxcar gradient

$$K = Gt$$

(K is area under curve)



- all of the following gradients end up at the same point in k-space:



# IMAGE RECONSTRUCTION

$$S(k_x, k_y) = \iint I(x, y) e^{i2\pi(k_x x + k_y y)} dx dy$$

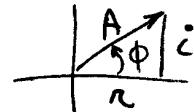
Signal (complex)  
 RF coil sums  
 brain spin density (real)

gradient-caused phase wraps (complex)  
 in transverse magnetization  
 ↳ N.B.: assumes perfect sinusoids! (they're not)

$$I(x, y) = \iint S(k_x, k_y) e^{i2\pi(x k_x + y k_y)} dk_x dk_y$$

screen image  
 $k_y k_x$   
 Signal (complex)  
 sin, cos

ideally → image is real  
 in practice → complex  
 ↳ use amplitude image:



adding exponents  
 same as multiplying  
 two  $e^{i2\pi k_x}$ 's

$$= \iint_{k_y k_x} S(k_x, k_y) e^{i2\pi x k_x} e^{i2\pi y k_y} dk_x dk_y$$

same as two  
 sequential 1D  
 FFT's (actual code)

$$= \int_{k_y} \left[ \int_{k_x} S(k_x, k_y) e^{i2\pi x k_x} dk_x \right] e^{i2\pi y k_y} dk_y$$

- in practice, finite number of samples,  $N$  and  $M$ , are collected  
 $k_x$  and  $k_y$  directions of  $k$ -space (integral → discrete sum)  
 ↳ b/c  $M/2$  belongs to next replica

$$I(x, y) = \sum_{m=-M/2}^{M/2-1} \left[ \sum_{n=-N/2}^{N/2-1} S(n, m) e^{i2\pi \frac{n}{\Delta k_x} \Delta k_x} \right] e^{i2\pi m \Delta k_y y} \Delta k_y$$

$|x| < \frac{1}{\Delta k_x}$   
 $|y| < \frac{1}{\Delta k_y}$

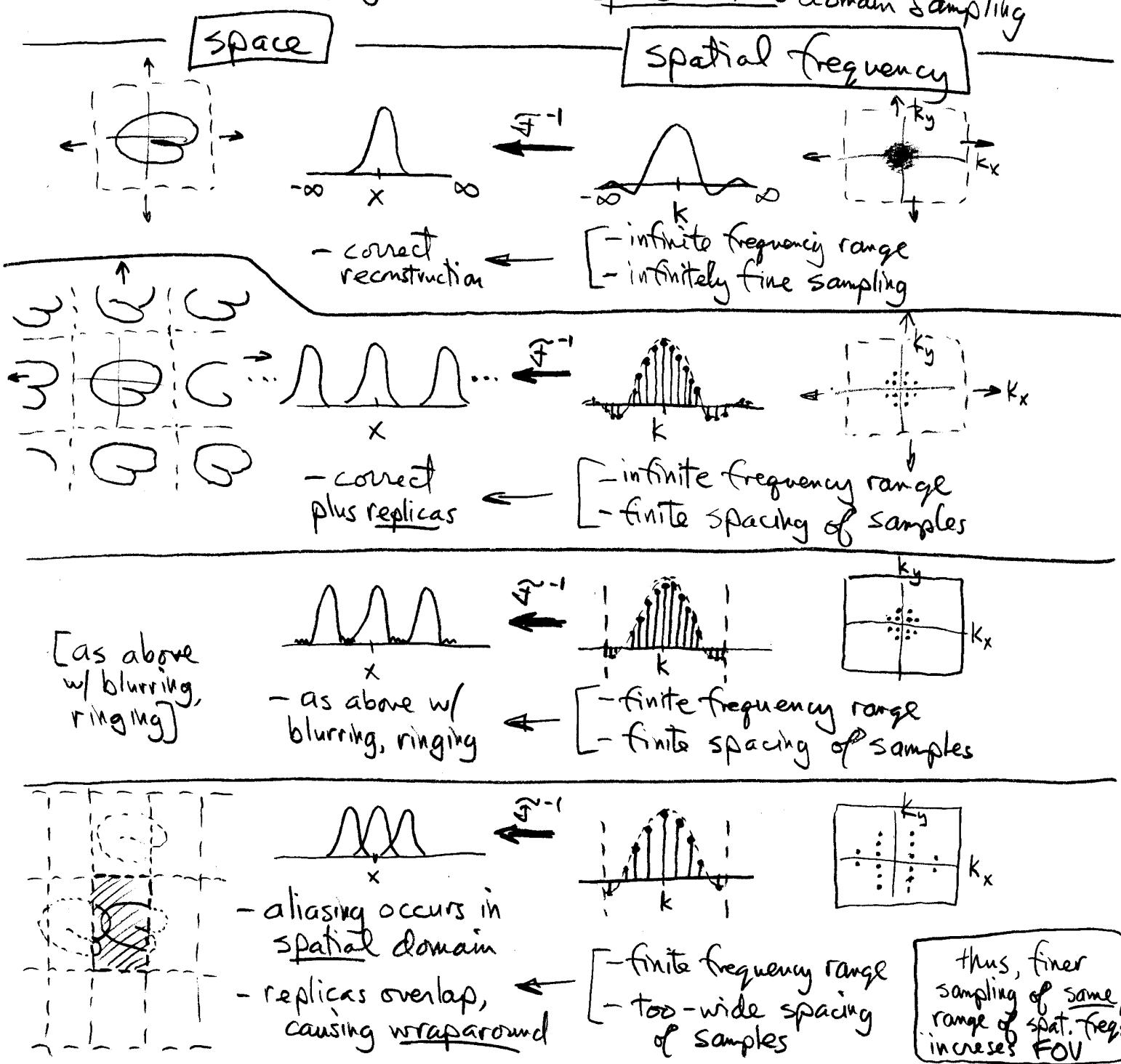
was  $k_x$   
 sampling interval in  $k$ -space

# SAMPLING

aliasing, FOV

aliasing from insufficient samples in time domain

- must consider effects sampling limited points in  $k$ -space
- [limited in range of frequencies sampled ( $k_{\min} \rightarrow k_{\max}$ )]
- [limited in rate of sampling ( $\Delta k$ )]
- N.B. aliasing less familiar when result of limited frequency domain sampling than limited space or time domain sampling



spcon2b

## UNDER/OVER SAMPLE

$$\text{FOV}_x = \frac{1}{\Delta k_x}$$

$$\delta_x = \frac{\text{FOV}_x}{N} = \frac{1}{N \Delta k_x}$$

$\text{FOV}$  (distance to repeat) is reciprocal of spatial frequency sampling interval

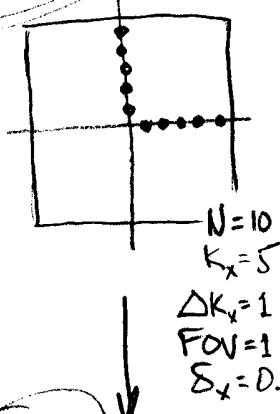
pixel size is  $\text{FOV}$  divided by  $k$ -space sample count

more examples

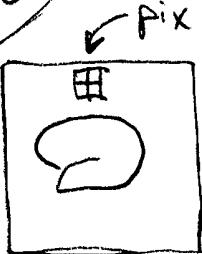
3 more examples (not incl. less samples to same spat. freq [bottom last page])

Basic Image

Spatial freq.  
 $K$ -space

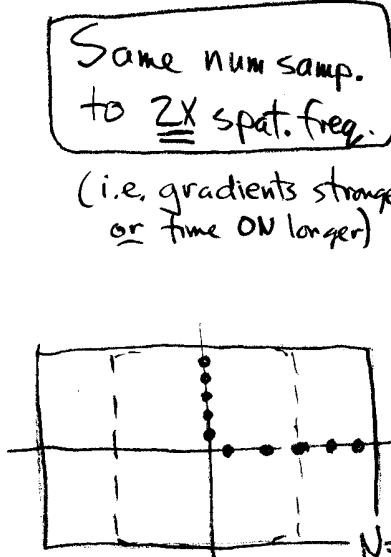


Space



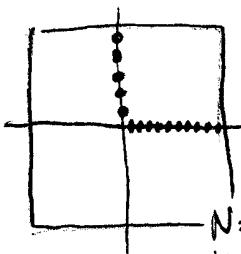
Same num samp.  
to  $\geq$  spat. freq.

(i.e. gradients stronger  
or time ON longer)



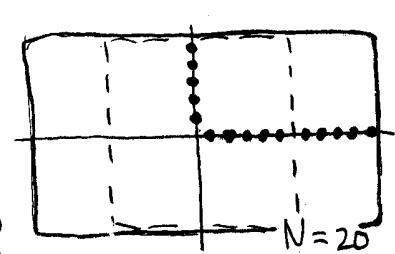
2X num. samples  
to same spat. freq.

(i.e. gradients weaker  
or time ON shorter)



2X number samples  
to  $\geq$  spat. freq.

(i.e. gradients stronger  
or time ON longer)



- basic image
- square pix

- x-pix half width
- replicas intrude

[Scanner makes  
square image  
"wrap" occurs]

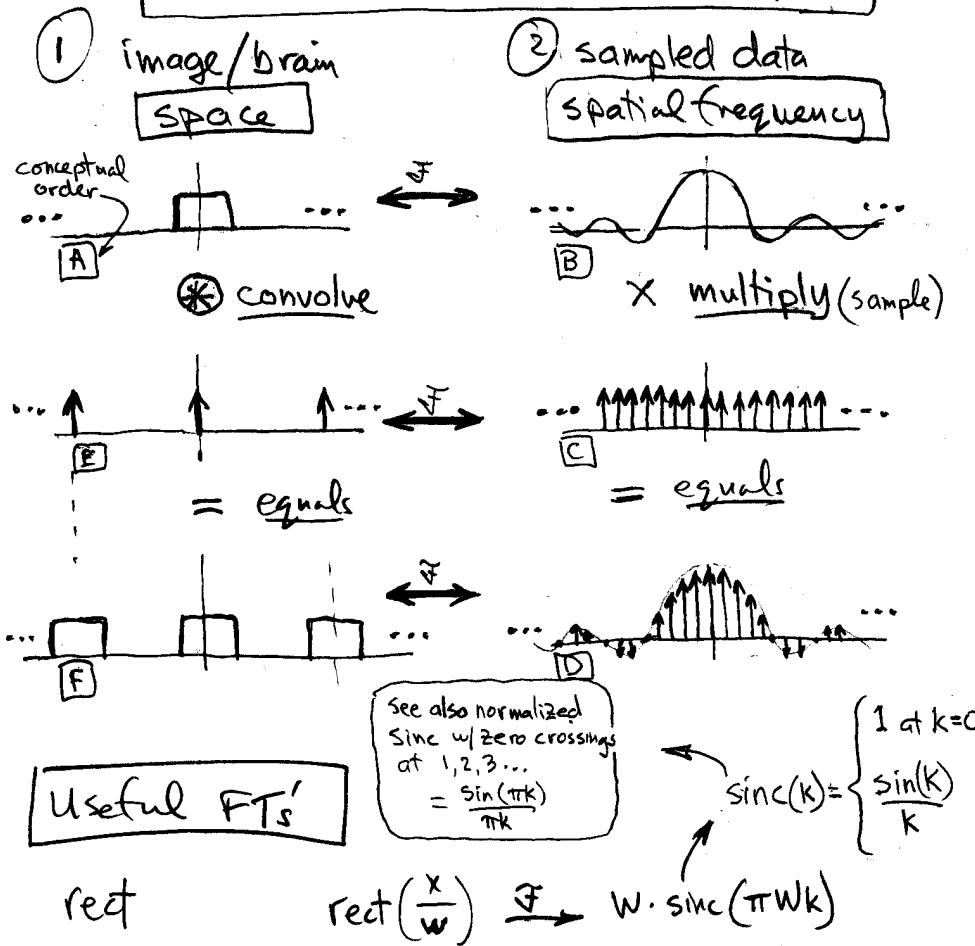
- square pix
- twice x-pix count  
so  $\text{FOV}=2X$
- this is "phase oversamp"

[Scanner crops to square  
replicas move out]

- x-pix half width
- twice x-pix count
- same FOV
- this is decrease  
pixel size w/o  
change FOV

recon-3

### Fourier Transform Solution to Replicas



$$\text{rect} \quad \text{rect}\left(\frac{x}{w}\right) \xrightarrow{\mathcal{F}} W \cdot \text{sinc}(\pi W k)$$

Gaussian  
(special case)

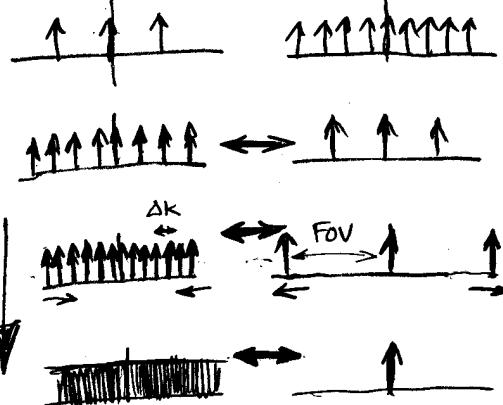
$$e^{-\pi x^2} \xrightarrow{\mathcal{F}} e^{-\pi k^2}$$

larger  $\Rightarrow$  narrower

$$e^{-\alpha x^2} \xrightarrow{\mathcal{F}} \sqrt{\frac{\pi}{\alpha}} e^{-\frac{\pi^2 k^2}{\alpha}}$$

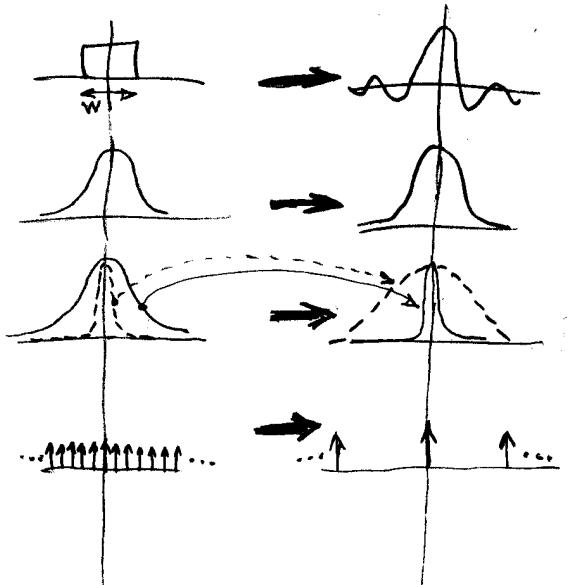
$$\text{comb} \quad \sum_{n=-\infty}^{\infty} \delta\left(x - \frac{n}{\Delta k}\right) \xrightarrow{\mathcal{F}} \Delta k \sum_{p=-\infty}^{\infty} \delta(k - p \Delta k)$$

- limit approach to Fourier transform of Combs



$$\text{FOV} = \frac{1}{\Delta k}$$

$$\Delta k = \frac{1}{\text{FOV}}$$



recon-4

# POINT - SPREAD FUNCTION

$$\hat{I}(x) = \Delta k \sum_{n: \{-N/2, N/2\}} S(n \Delta k) e^{i 2\pi n \Delta k x}$$

- set true image to  $\delta$ -function, then measured signal is:

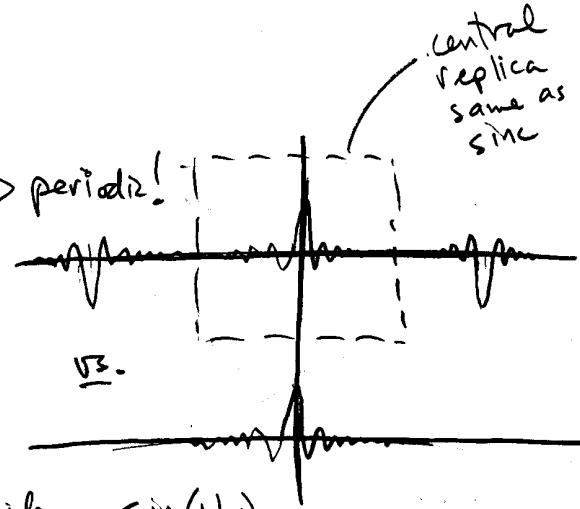
$$S(m \Delta k) = 1$$

- substitute into recon to get PSF:

$$h(x) = \Delta k \sum_{n: \{-N/2, N/2\}} e^{i 2\pi n \Delta k x}$$

- simplify

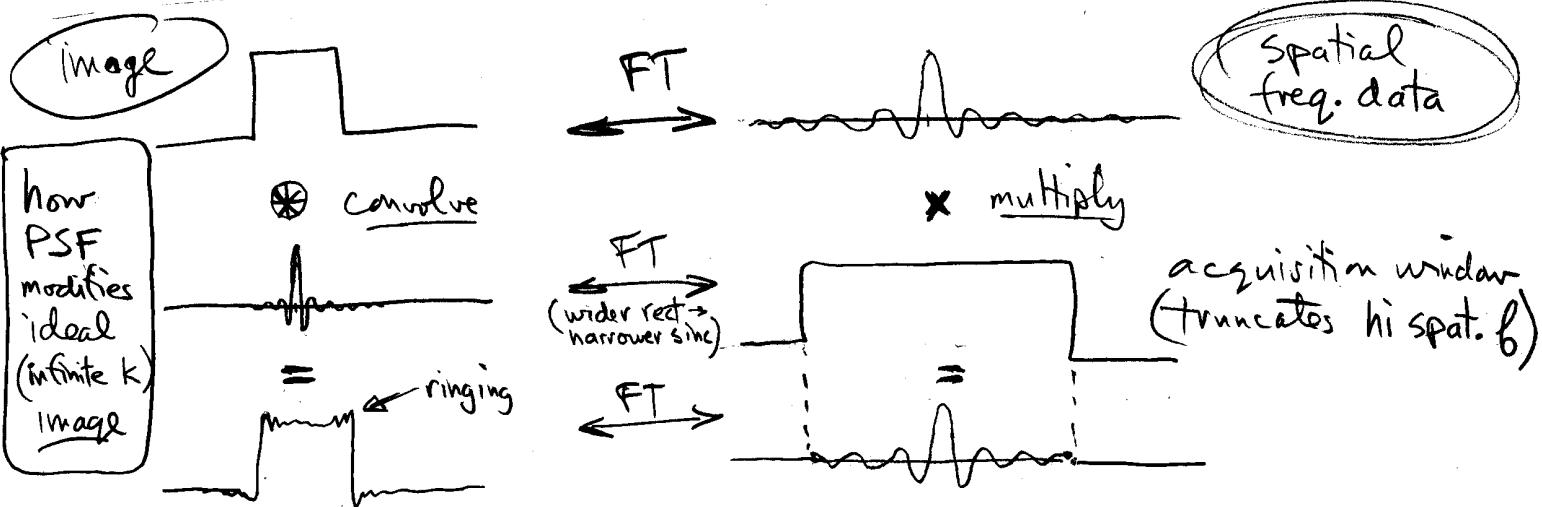
$$h(x) = \Delta k \frac{\sin(\pi N \Delta k x)}{\sin(\pi \Delta k x)} \Rightarrow \text{periodic!}$$



- that is, image is reconstructed from a sum of sinc's, because the FT of a boxcar pixel in  $k$ -space is an ideal sinc

all outside  
center at sinc  
zero-crossings if  
 $K_{max} = X_{max}$

$$\underbrace{\sin(Nx)}_x$$



recon-5

# GENERAL LINEAR INVERSE RECON FOR MRI

$$S(k_x) = \int_x I(x) e^{-i 2\pi k_x x} dx$$

perfect spin phase "stripes"  
modeling real life imperfections

Signal eq.  $\rightarrow$  fwd problem

$$I(x) = \int_{k_x} S(k_x) e^{+i 2\pi x k_x} dk_x$$

perfect "stripes"

Recon eq.  $\rightarrow$  inv. problem

$$\vec{s} = \vec{F} \vec{i}$$

$$\begin{bmatrix} s \\ \vdots \\ k_y \end{bmatrix} = \begin{bmatrix} x, y \rightarrow \\ F \vec{i} \end{bmatrix} \begin{bmatrix} i \\ \vdots \\ i \end{bmatrix}$$

Linear "forward solution"  
matrix & vectors have complex entries  
can build in any measurable priors

$$F_{x,y,t} = g(x,y) e^{-i\phi(x,y)} e^{-\frac{(nT \pm m\Delta t + TE)/T_2}{\gamma}} e^{-i\gamma\Delta B(x,y, nT \pm m\Delta t)} e^{-i2\pi(m\Delta k_x x + n\Delta k_y y)}$$

coil gain at this location      coil phase      T2 decay      B $\phi$  error (x,y dep.)      freq + phase (complex)

could insert x-y-dependent gradient non-lin.

multi-coil

$$\begin{bmatrix} s \\ \vdots \\ 2 \end{bmatrix} = k_y \begin{bmatrix} x, y \rightarrow \\ F \\ \text{coil 1} \\ \cdots \\ \text{coil 2} \end{bmatrix} \begin{bmatrix} i \\ \vdots \\ i \end{bmatrix}$$

naturally incorporates undistorted field map  
different sensitivity function for each coil  
contains additional info about source loc.  
but, need reference scan, lo-res OK  
(need phase corrections for each coil)

$$\vec{i} = \vec{F}^+ \vec{s}$$

over-determined

More  
Pseudo  
inverse

$$\vec{F}^+ = (\vec{F}^T \vec{F})^{-1} \vec{F}^T \quad (x,y)^2 \rightarrow \text{"small"}$$

$$= \vec{F}^T (\vec{F} \vec{F}^T)^{-1} \quad (x,y \cdot \text{coils})^2 \rightarrow 16 \times \text{bigger}$$

for 4 coils

$$(\begin{bmatrix} \vec{F}^T \\ \vec{F} \end{bmatrix})^{-1} \begin{bmatrix} \vec{F}^T \\ \vec{F} \end{bmatrix}$$

$$\begin{bmatrix} \vec{F}^T \\ \vec{F} \end{bmatrix} \left( \begin{bmatrix} \vec{F} \\ \vec{F}^T \end{bmatrix} \right)^{-1}$$

$$\vec{i} = [(\vec{F}^T \vec{F})^{-1} \vec{F}^T] \vec{s}$$

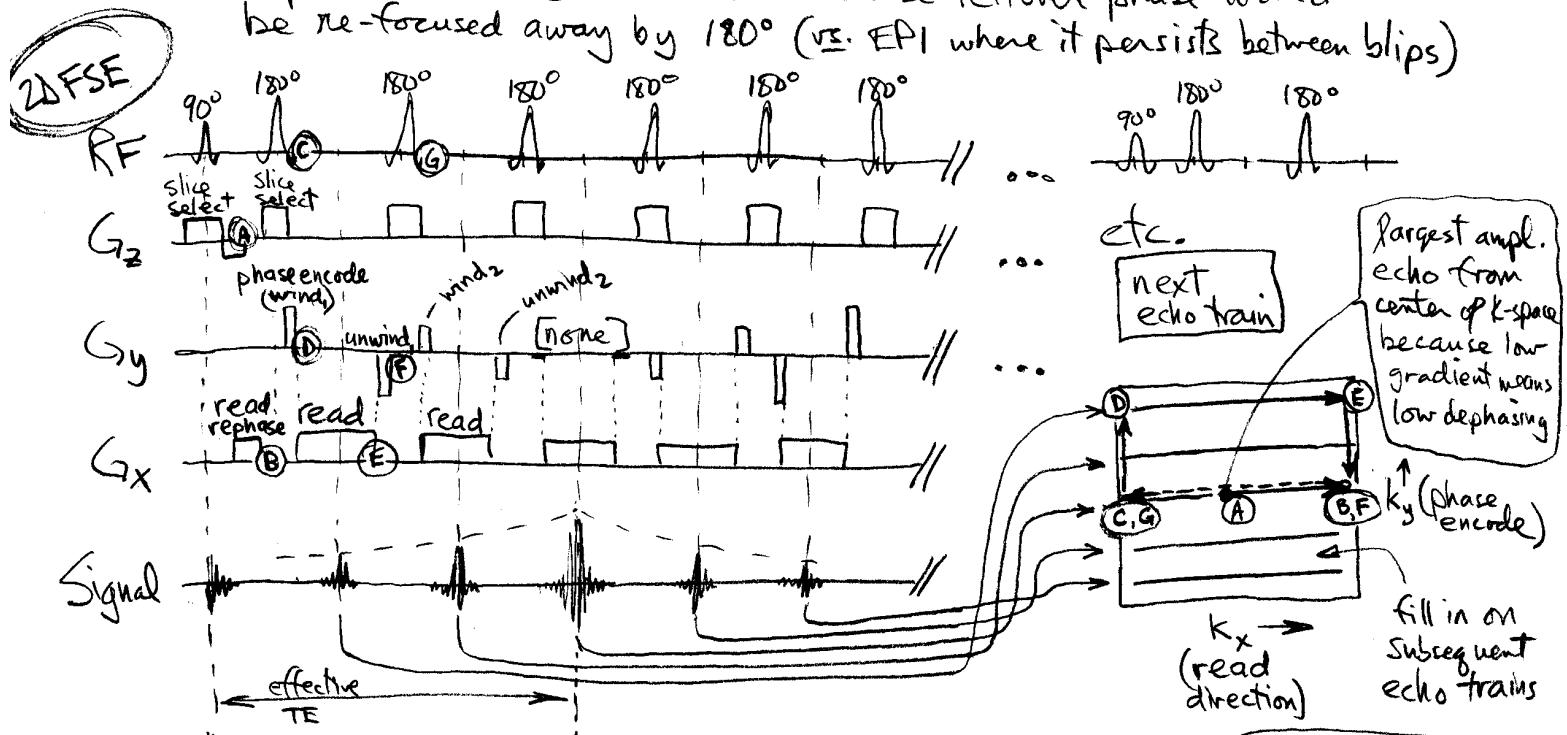
Slice-by-slice  
assume slice select swamps  $\Delta B_0$

# FAST SPIN ECHO (FSE)

RARE, FSE, 3DFSE

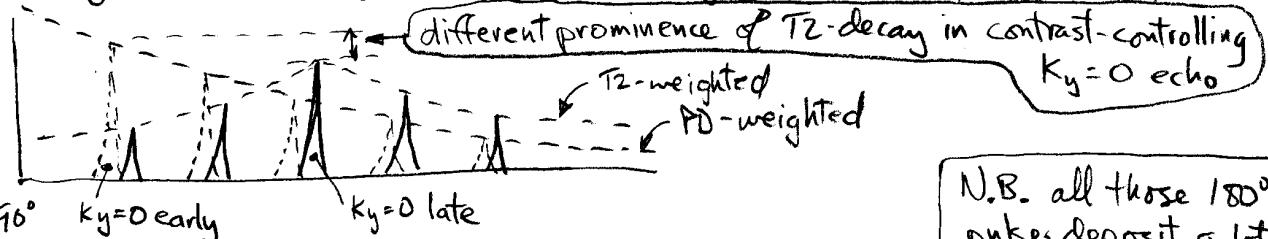
"echo train"

- one  $90^\circ$  pulse followed by multiple  $180^\circ$  pulses (e.g., 8) each with a different phase-encode gradient
- each phase "winder" is "unwound" because leftover phase would be re-focused away by  $180^\circ$  (vs. EPI where it persists between blips)



- the "effective TE" is the TE when center of k-space is collected (largest effect on contrast, largest echo)
- each subsequent echo has more T2 decay:  $E_n = e^{-nTE/T_2}$   $n = 1, 2, \dots, M$

- by arranging to collect  $k_y=0$  early, PD-weighted instead of T2-weighted

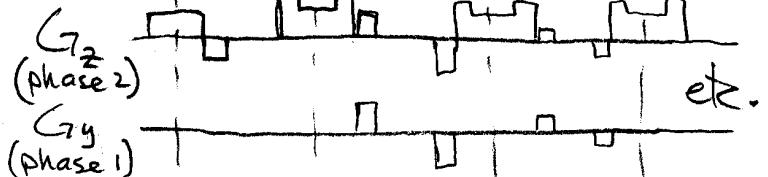


- possible to correct different T2-weighting of echoes by estimating T2 curve from  $G_y=0$  echo train

N.B. all those  $180^\circ$  pulses deposit a lot of RF power:  
 $90^\circ + 180^\circ = 45 \times \text{power } 30^\circ$

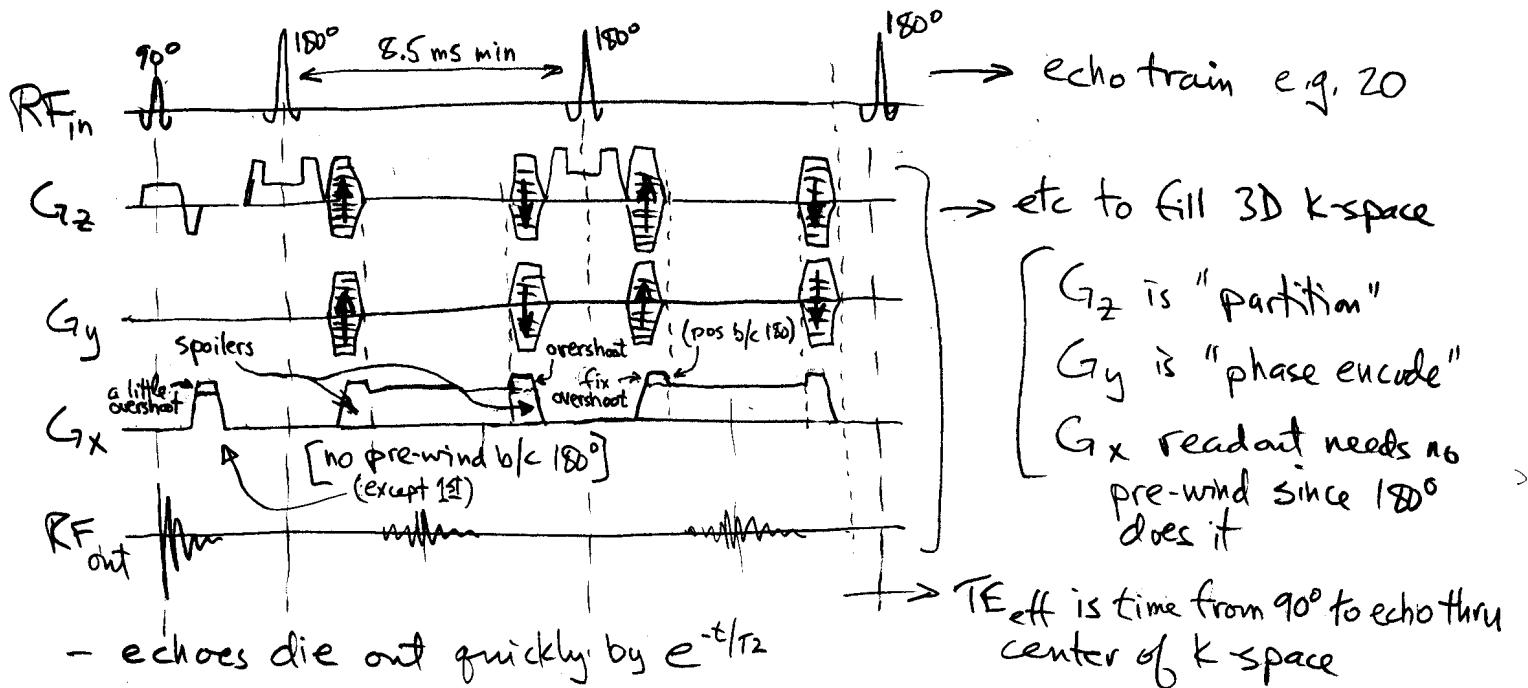
- 3DFSE - like 2D except

wind/unwind added to thick  
Slice select (w/crushers on  $180^\circ$ )

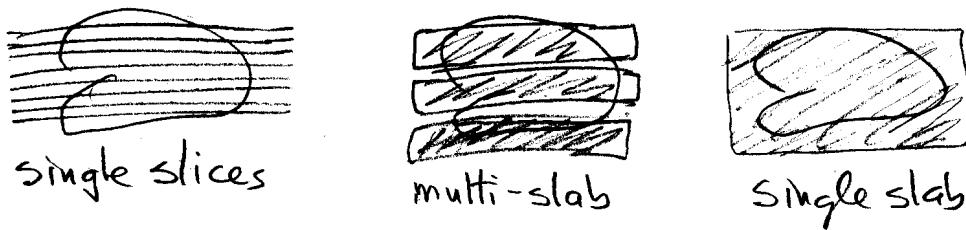


pulse-1b

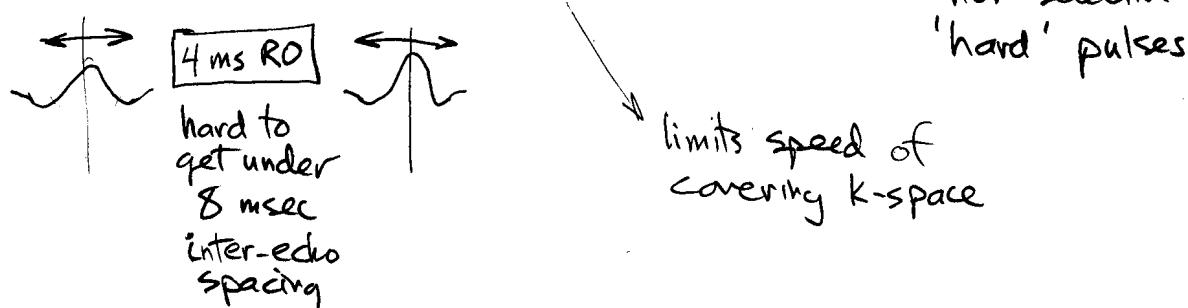
# MULTI-SLAB 3D FSE, PROBLEMS



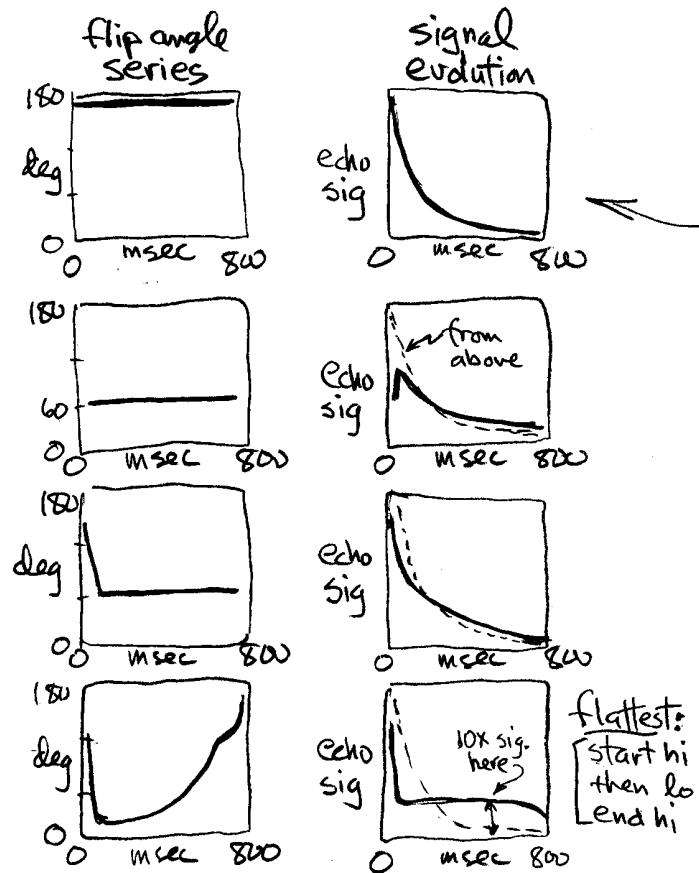
- echoes die out quickly by  $e^{-t/\tau_2}$
- since echoes after 90° limited to <30, can't fill 3-D k-space in a reasonable time
- SAR constraint  $SAR \propto B_0^2 \Omega^2 \Delta f$  rough approx.  
for non-adiabatic  
standard pulses  
 $\hookrightarrow 180^\circ$  pulses deposit 4-6x power of 90°
- "multi-slab" is halfway between slices and single-slab



- problem at slice boundaries — esp. movement
- multislab requires slice selective RF pulses  $\rightarrow$  longer than non-selective 'hard' pulses



pulse-1c  
SINGLE-SLAB 3D FSE



(SPACE)

from Mugler (2014) J Mag Res Img

- regular FSE (180° pulse train)

- Sub 180° pulses cause each successive pulse to also generate a stimulated echo (STE)  
 ↳ this "storage" in z-axis preserves magnetization for longer time

- smaller flip angles allow much longer echo trains  
 ↳ enough to collect whole plane of 3-D k-space +

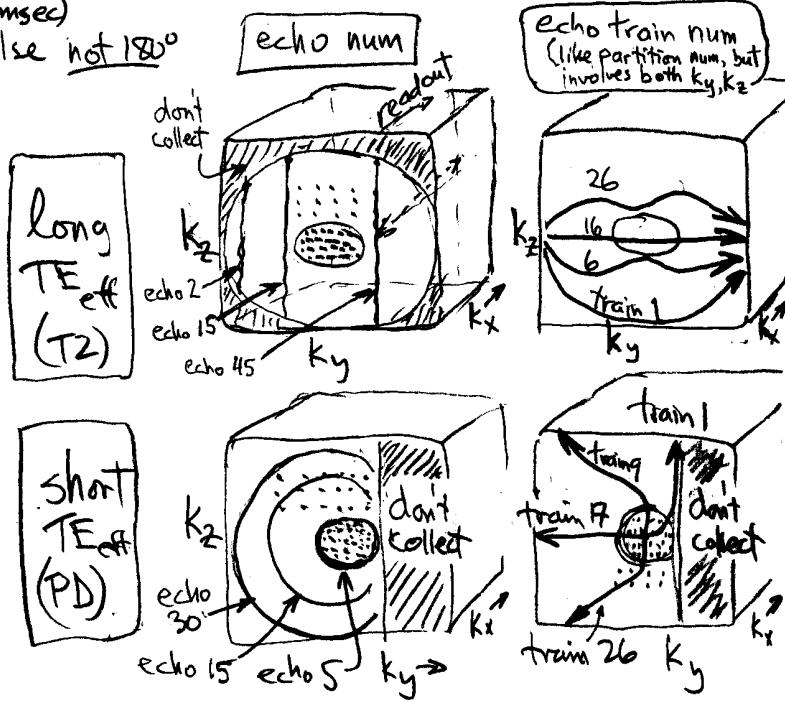
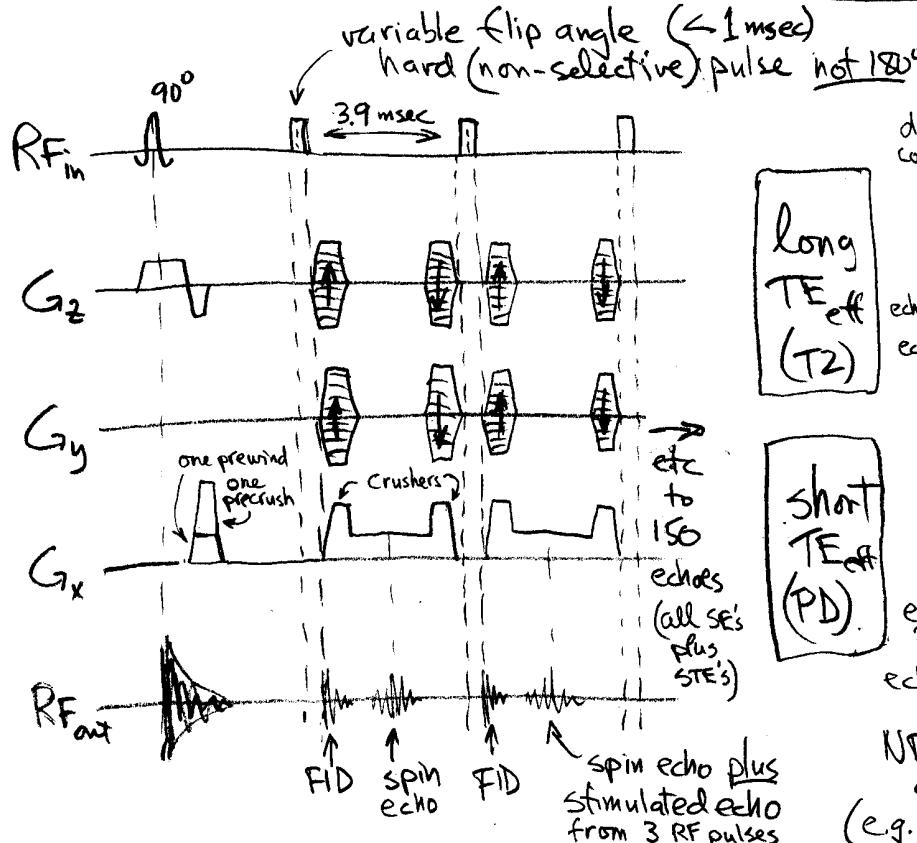
- different than hyper echoes (not symmetric)  
 - contrast must consider STE

N.B.: T1<sup>11</sup>

$$SE = \sin \alpha_1 \sin^2 \frac{\alpha_2}{2} e^{-2\gamma_1/T_2} \zeta$$

$$STE = \frac{1}{2} \sin \alpha_1 \sin \alpha_2 \sin \alpha_3 e^{-T_1/T_1} e^{-2\gamma_1/T_2}$$

- single-slab 3D FSE pulse seq.



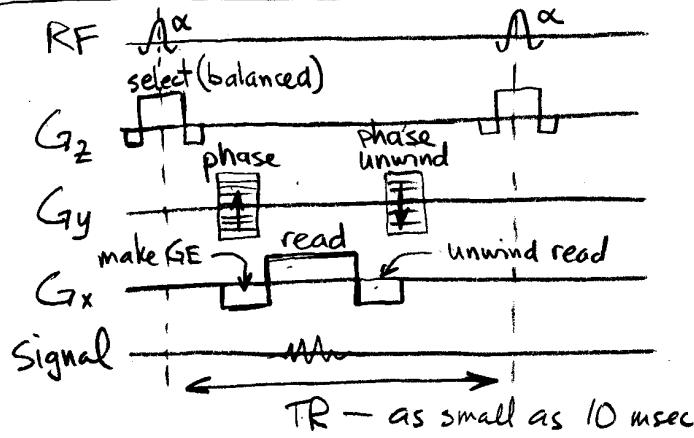
NB: time to cent k-space is  $\approx 5 \times$  apparent contrast time b/c of "storage"  
 (e.g. TE<sub>eff</sub> = 585 ms looks like FSE TE = 140 ms)

Pulse-2

## FAST GRADIENT ECHO

(GRASS | FLASH | MPRAGE)  
 FISP | SPGR |

- small tip so TR can be greatly reduced (e.g. 10 msec, less than  $T_2$ )
- 'leftover' undecayed transverse magnetization
  - "unwound" and re-used
  - "spoiled" before next shot



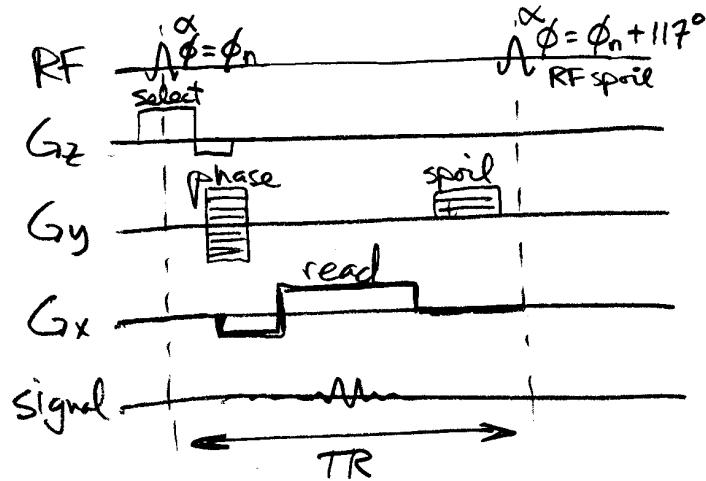
### STEADY-STATE COHERENT (GRASS, FISP)

- unwind phase from phase-encode  $M_T$  before next pulse (there because  $TR < TE$ )
- unwind read gradient, too

$$S = k \sin \alpha \left[ \frac{1}{1 + \cos \alpha + (1 - \cos \alpha) T_1/T_2} \right] e^{-TE/T_2}$$

tissue  $T_2/T_1$   
 brain 0.11  
 fat 0.3  
 CSF 0.7

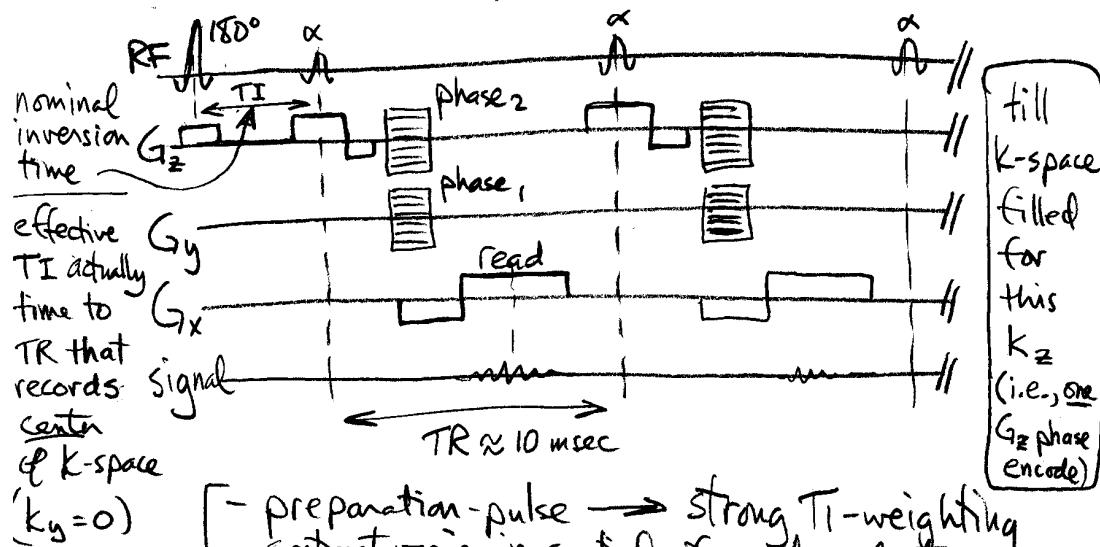
$\hookrightarrow T_2/T_1$  - weighted contrast (bright CSF)



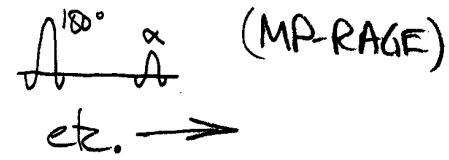
### STEADY-STATE, SPOILED (SPGR, FLASH)

- spoil with random gradient (but this still allows some  $\alpha$  refocusing)
- spoil with gradient plus incremented phase of RF pulses (RF spoiling)
- good gray-white contrast ( $T_1$ -weighted)

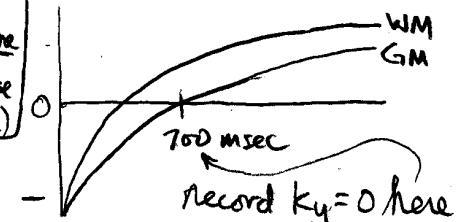
(shown as 3D sequence — possible with ones above, too)



### NON-STEADY STATE, MAGNETIZATION-PREP.



- longitudinal mag. not affect much by low angle pulses



- preparation-pulse  $\rightarrow$  strong  $T_1$ -weighting
- contrast varies in spatial-freq-dependent way

pulse-2b

# QUANTITATIVE TI - INTRO, METHODS

## Motivation

**HOW TO FIX:**  
collect more than one copy of vol.

- image values are arbitrary/relative (diff seq's, manufacturers)
- uncorrected coil fall-off (receive inhomogeneity) can result in 2-3x differences in voxel brightness
- uncorrected variation in local B1 field can cause contrast variation
  - at 3T, B1 can vary by 25% across the brain
  - this can invert contrast in a fast gradient echo

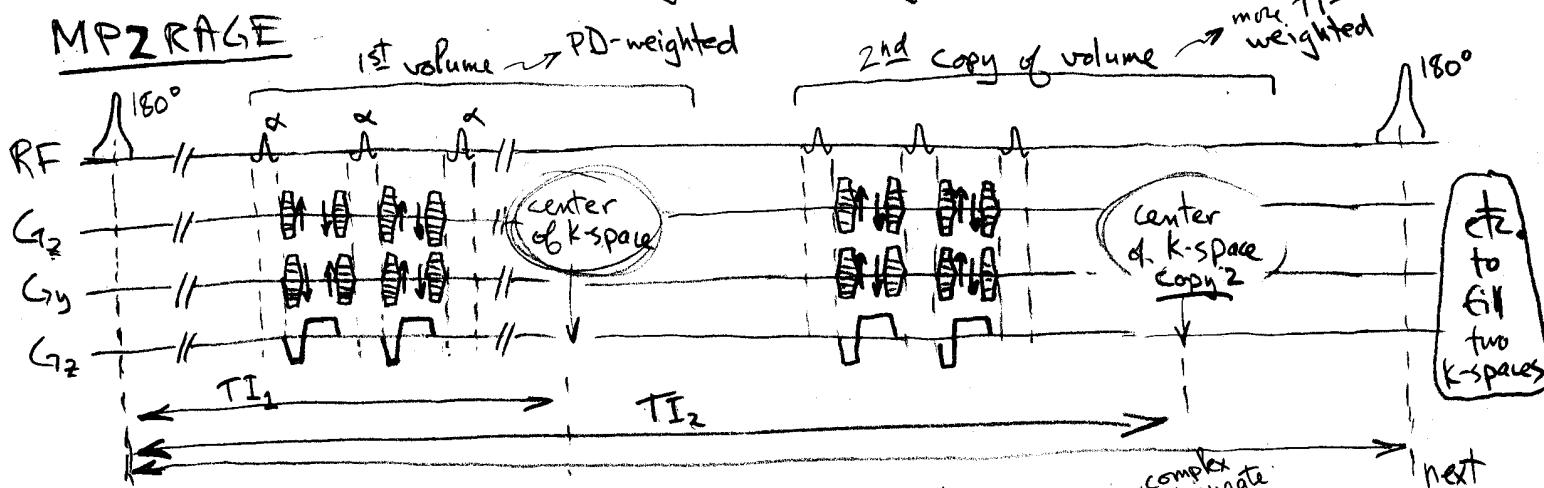
## Pre-scan normalise

- collect lo-res GE image, receive w/ body coil (no coil fall-off)
- set parms. to get low GM/WM contrast
- collect data scan (e.g. MPRAGE) w/ surface coils, strong GM/WM
- use ratio between scans to generate smooth correction field

## T1 divided by T2

- MPRAGE  $\rightarrow$  strong T1-contrast
- SPACE  $\rightarrow$  T2-weighted (no T1 weighting)
- T1/T2 removes coil fall-off
- Problems
  - distortion different in GE (MPRAGE) and SE (SPACE)
  - noise in regions of low signal

## MP2RAGE



- N.B. SSFP-like in partition, phase-encode directions

- convert to -0.5 to 0.5 image:  $S = \text{real} \left[ \frac{\vec{s}_{T1_1}^* \cdot \vec{s}_{T1_2}}{\|\vec{s}_{T1_1}\|^2 + \|\vec{s}_{T1_2}\|^2} \right]$

- calc. PD & T1 from above

cf. 2 flip angles

$$\left[ \frac{\vec{s}_{T1_1}^* \cdot \vec{s}_{T1_2}}{\|\vec{s}_{T1_1}\|^2 + \|\vec{s}_{T1_2}\|^2} \right]$$

pulse-2c

# QUANTITATIVE T<sub>1</sub> — HELMS 2-FLIP ANGLE METHOD

— start w/ gradient echo signal e.g., dropping T<sub>2</sub>-decay  $\rightarrow e^{-TE/T_2}$

$$S_{\text{Ernst}} = A \cdot \sin \alpha \cdot \frac{1 - e^{-TR/T_1}}{1 - \cos \alpha \cdot e^{-TR/T_1}}$$

Ernst eq.

$$(\max: \cos \alpha_E = e^{-TR/T_2})$$

"Ernst angle"

$$(T_E = \cos^{-1}(e^{-TR/T_2}))$$

— simplify/linearize/estimate

$$TR \ll T_1$$

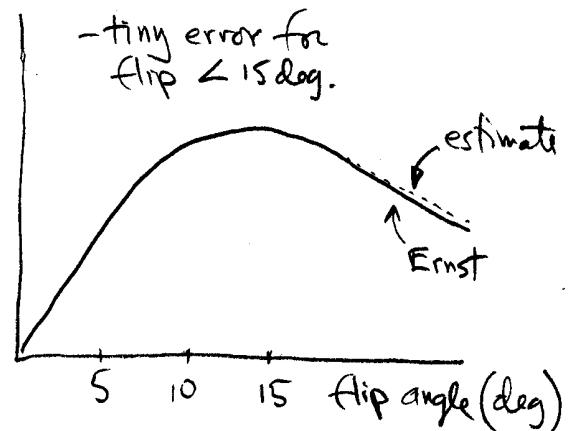
linear approx. of exponentials

Taylor expansion Simplification of sin, cos, drop small terms

Helms et al. (2008)

$$S \approx A \cdot \alpha \cdot \frac{TR/T_1}{\alpha^2/2 + TR/T_1}$$

tiny error for flip &lt; 15 deg.



— solve for T<sub>1</sub> and A (proton-density) given

signals from 2 diff flip angles

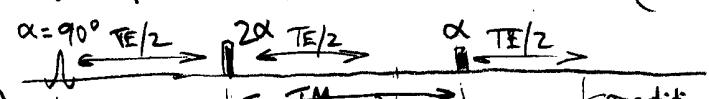
2 flip angles

$$T_1^{\text{est.}} = 2TR \frac{S_1/\alpha_1 - S_2/\alpha_2}{S_2\alpha_2 - S_1\alpha_1}$$

$$A^{\text{est.}} = \frac{S_1 S_2 (\alpha_2/\alpha_1 - \alpha_1/\alpha_2)}{S_2\alpha_2 - S_1\alpha_1}$$

— problem: flip angle varies a lot at 3T (e.g. 25%) from nominal/requested (e.g., flip series)

**B1 map** — acq. spin-echo and stimulated echo (EPI)



estimate T<sub>1</sub>, T<sub>2</sub>  
solve for 2 α's  
insert

$$\left( \sin \alpha \cdot \sin^2 \left( \frac{2\alpha}{2} \right) \right)$$

$$S_{SE} = k \cdot \sin^3 \alpha \cdot e^{-TE/T_2}$$

slice

phase

read

RFout

$$S_{STE} = k_2 \cdot \sin^2 \alpha \cdot \sin 2\alpha \cdot e^{-TE/T_2} \cdot e^{-TM/T_1}$$

$$\alpha = \cos^{-1} \left( \frac{S_{STE} \cdot e^{-TM/T_1}}{S_{SE}} \right)$$

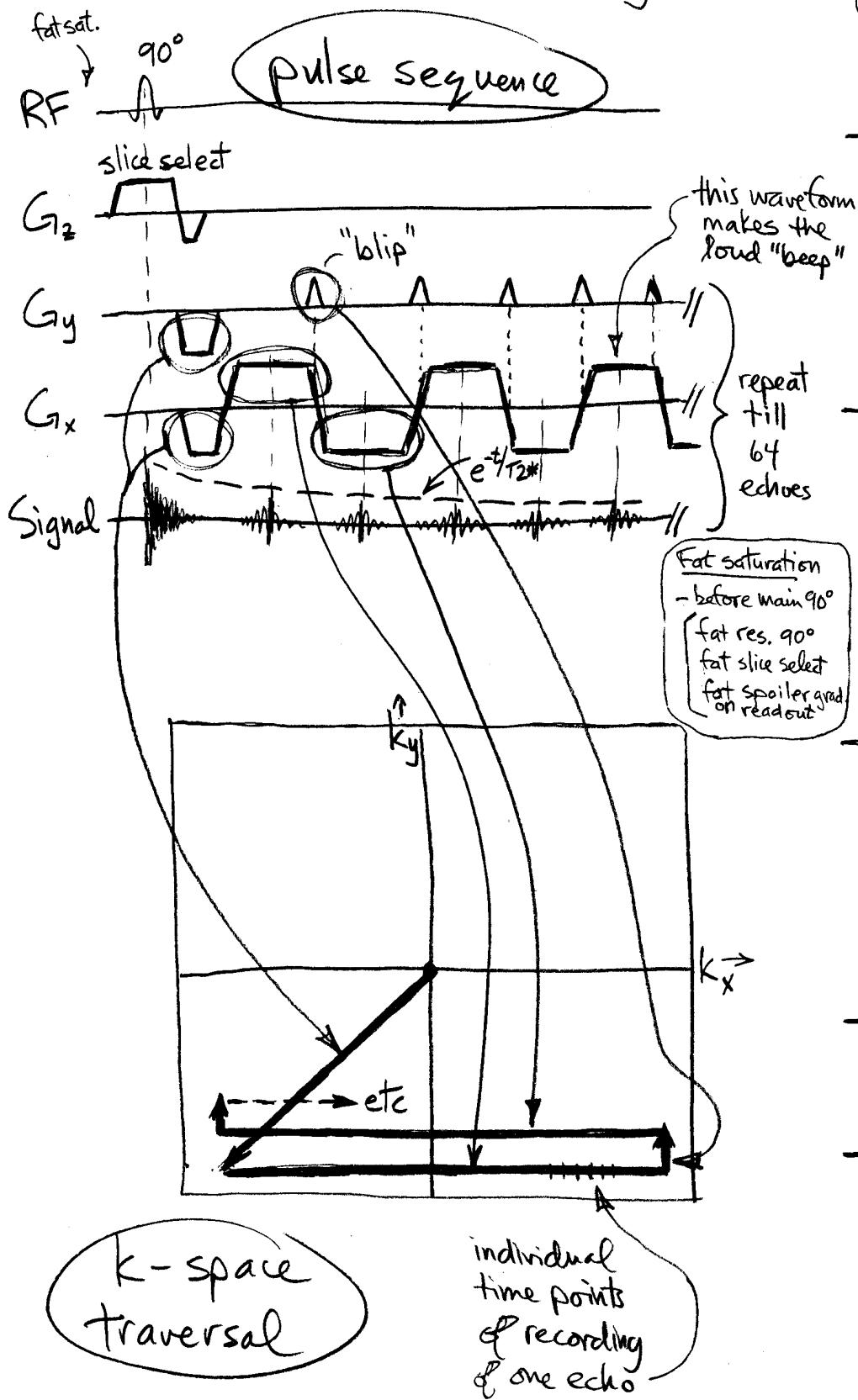
T<sub>2</sub>\*<sub>est</sub>

— add EPI-like echo train to each FLASH excit.

Jiru &amp; Klose (2006)

## ECHO PLANAR IMAGING EPI (another fast gradient echo)

- single shot EPI collects all k-space lines (e.g. 64) after a  $90^\circ$  RF pulse using a train of gradient echoes

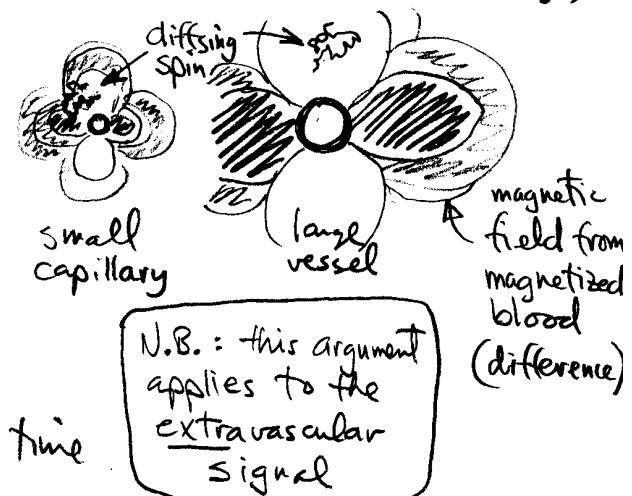


- since there is only one RF pulse per slice, spins never get reset to all-the-same (= zero freq, center of k-space)
- therefore, the recording point ( $\Delta t$ ) in k-space (= spin phase stripe pattern) stays wherever the x and y gradients last left it
- that explains why successive y phase-encode steps are achieved without changing the size of the  $G_y$  "blips"
- echoes are  $T_2^*$ -weighted (gradient echo)
- contrast mainly determined by echoes near center of k-space, which are only recorded after about 32 echoes

## SPIN ECHO EPI

why SE-BOLD may be selective for capillary bed

- Standard EPI is a gradient echo method, which results in  $T_2^*$ -weighting
- deoxyhemoglobin is paramagnetic, which reduces signal in a  $T_2^*$ -weighted image due to greater dephasing
- the excess of oxyhemoglobin (probably the result of the need to drive  $O_2$  into tissue, which requires more  $O_2$  in the blood than is actually used) leads to the positive BOLD effect
- spin echo corrects (cancels) static  $T_2^*$  ( $T_2'$ ) dephasing, incl. deoxy
- if all spins stayed in the same position, spin echo would eliminate the BOLD effect by eliminating dephasing
- diffusion exposes spins to different fields (reducing gradient echo dephasing!)
- magnetic field gradients produced by large vessels are smoother across space than those produced by small vessels
- for  $TE \approx 100$  ms, spins diffuse 10's of  $\mu\text{m}$ , which is larger than diameter of small capillary, meaning that spins will likely experience different fields over time
- therefore, spin echo will be less successful at cancelling BOLD effect near small vessels (BOLD effect will be reduced near large vessels where diffusion is less likely to expose spin to different fields here)

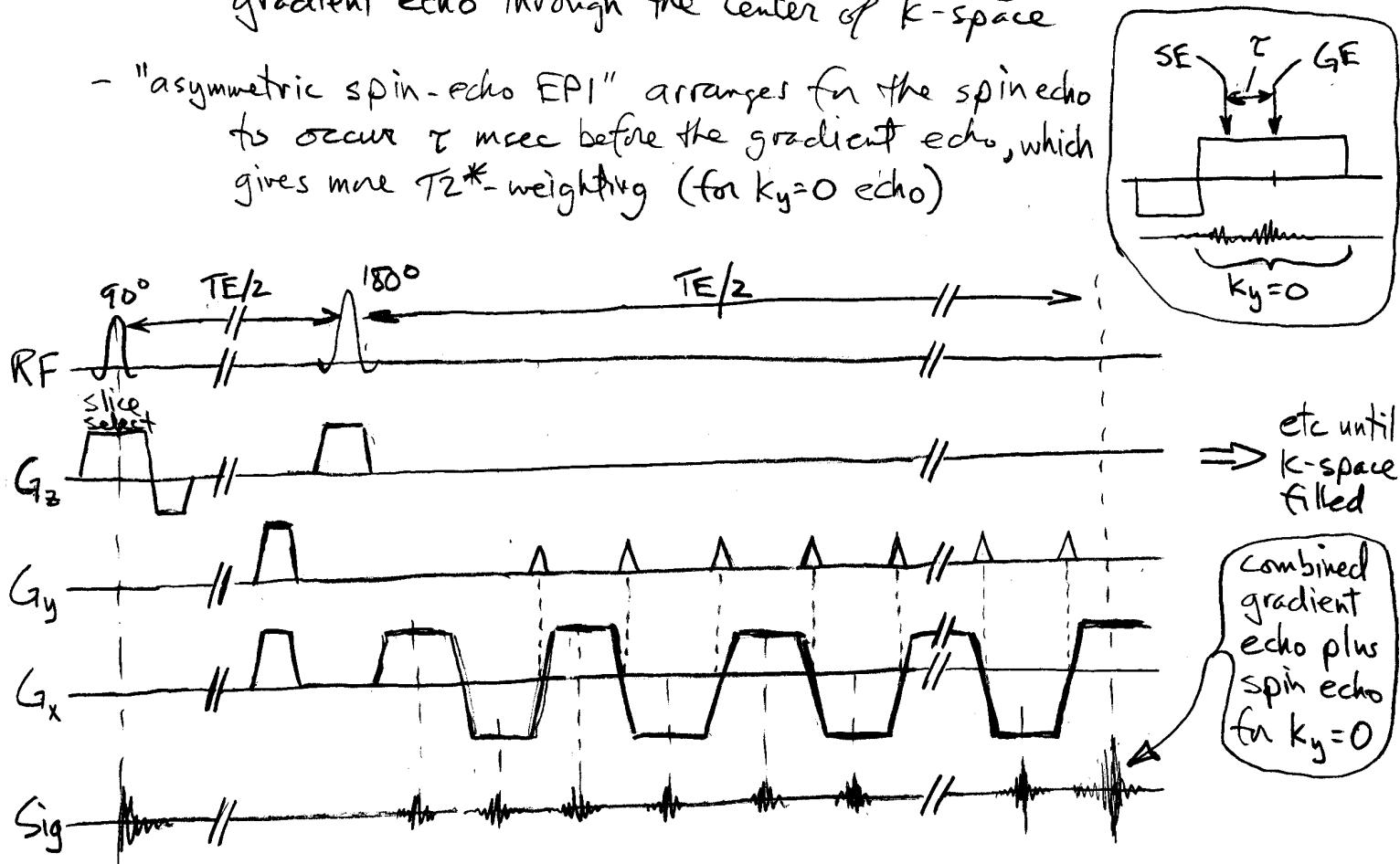


- this argument only works for extravascular spins — intravascular signal in BOLD is large (despite being only 4% by volume) because large gradient produced around red blood cells
  - measure intra/extravascular with bipolar pulse which kills signal in faster moving blood in moderate and larger vessels
  - ⇒ over half of SE-BOLD at LST is venous...

pulse-5

## SPIN ECHO EPI

- EPI is a multi-gradient echo pulse sequence
- "spin-echo EPI" uses a  $180^\circ$  pulse to add a single spin echo to the contrast-controlling gradient echo through the center of k-space
- "asymmetric spin-echo EPI" arranges for the spin echo to occur  $\tau$  msec before the gradient echo, which gives more  $T_2^*$ -weighting (for  $k_y=0$  echo)



- the  $180^\circ$ -pulse rephasing reduces the  $T_2^*$  signal, which is why the partially rephased asymmetric spin echo has been more commonly used

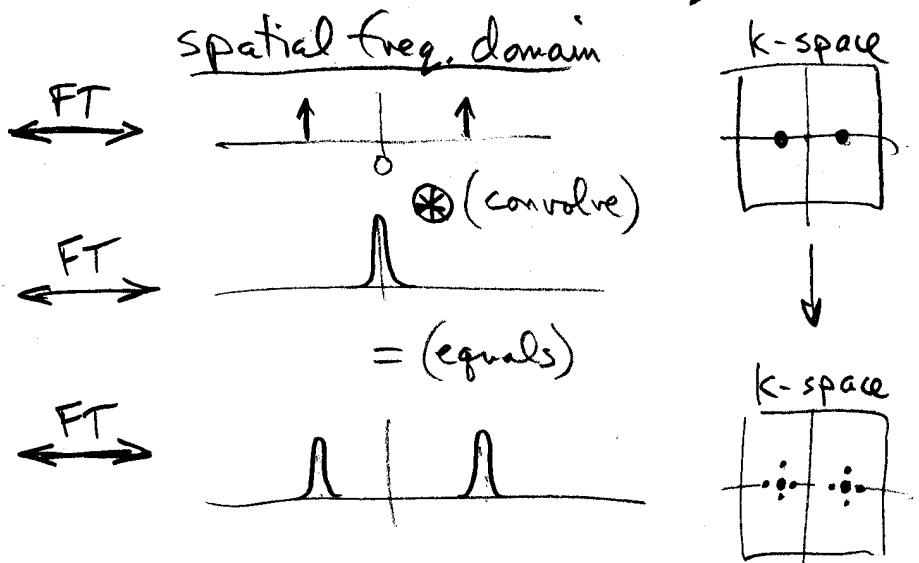
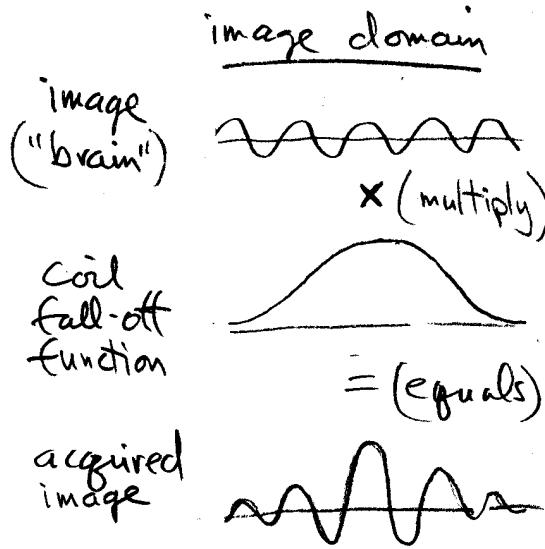
- at higher fields, spin echo EPI is more promising

- signal to noise is higher so we can take spin echo hit
  - contribution from venous blood is reduced, since blood  $T_2$  is so short, we can let it decay away before recording

pulse-5b

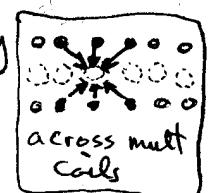
# COIL FALL-OFF / UNDERSAMPLE / GRAPPA / SENSE

- coil fall-off intuitively contains info about location  
if same brain location imaged by different coils w/ diff. fall-offs  
↳ but what does this look like in k-space?
- slow variation in RF-field fall-off (e.g. 1-4 cyc/FOV) causes a blur in acquired data in k-space  
→ (N.B. not addition!)
- to see this, consider multiplication by coil fall-off function in image space, which equals convolution (w/ FT of that function) in k-space - at all spatial frequencies !!
- simple example w/ "brain" consisting of one spatial freq.



- N.B. inverse FT of k-space data "smeared" in spat freq. Space is sharp image w/ fall-off (not blurred img.)
- "smear" means normally orthogonal spat. freq's "leak" to adj. freqs.

**GRAPPA** - construct k-space "kernel" to fill in missing k-space lines by training on fully-sampled data from near k-space center

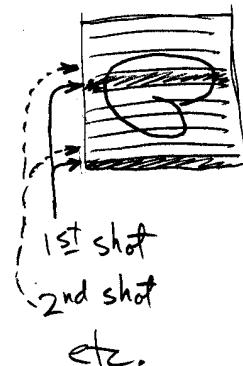
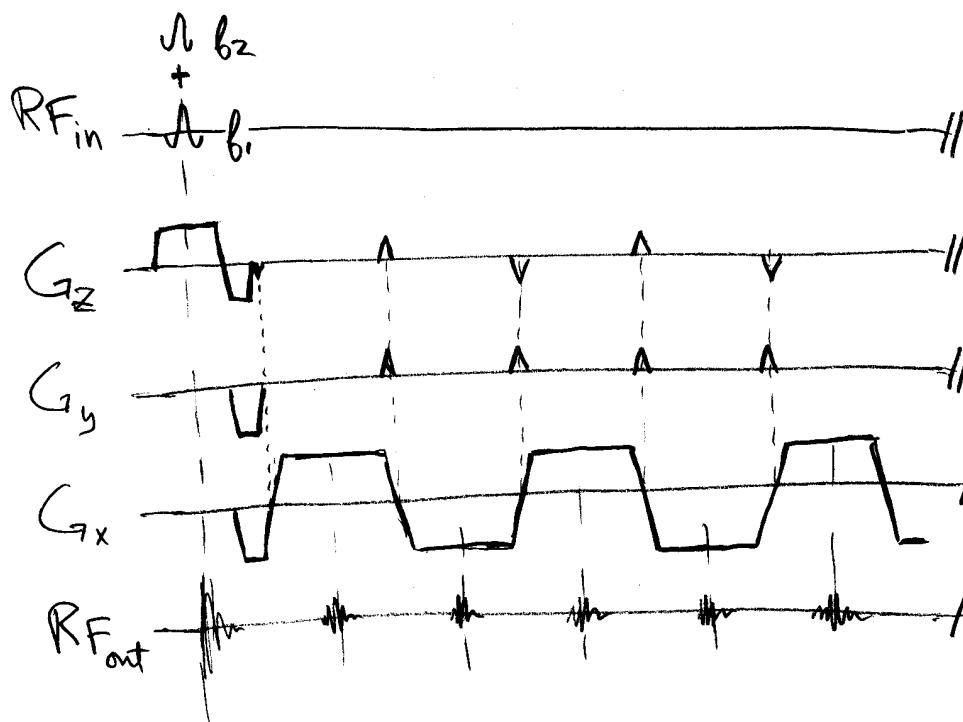


**SENSE** - general linear inverse approach

- N.B.: neither would work unless normally orthog. spat. freqs. blurred!

pulse-SC

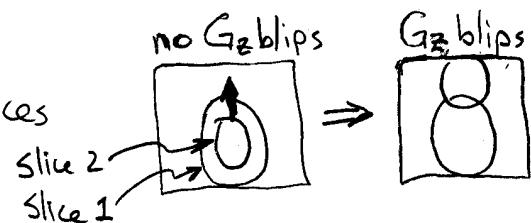
# SMS / MULTIBAND / blipped CAIPI



(or interleaved)

- excite multiple slices at once

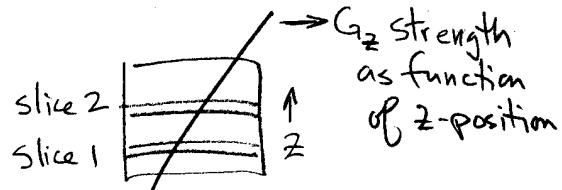
- function of  $G_z$  blips is to shift slices in  $G_y$  direction



- this occurs because for given slice, a phase pedestal is added to the entire slice

↳ this "Fourier Shift Theorem"

↳ [N.B.: different than  $B\phi$  defect-induced incremented phase errors]



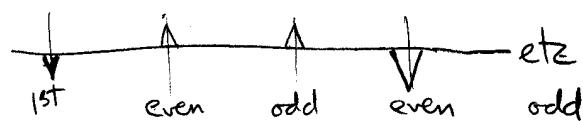
- problem w/ all up  $G_z$  blips → phase error builds up

trick #1 - start w/ 2 slices, one at  $z=0$ , other above

↳ if  $\pi$  ( $180^\circ$ ) phase shift used, blip up/down same! (no effect at  $z=0$ )  
 ↳ i.e., move top or bottom replica

trick #2 - for multiple slices not all at  $z=0$ , phase no longer same for even/odd  
 ↳ but can add phase to equilibrate to k-space before recon.

trick #3 - for more than 2 slices:

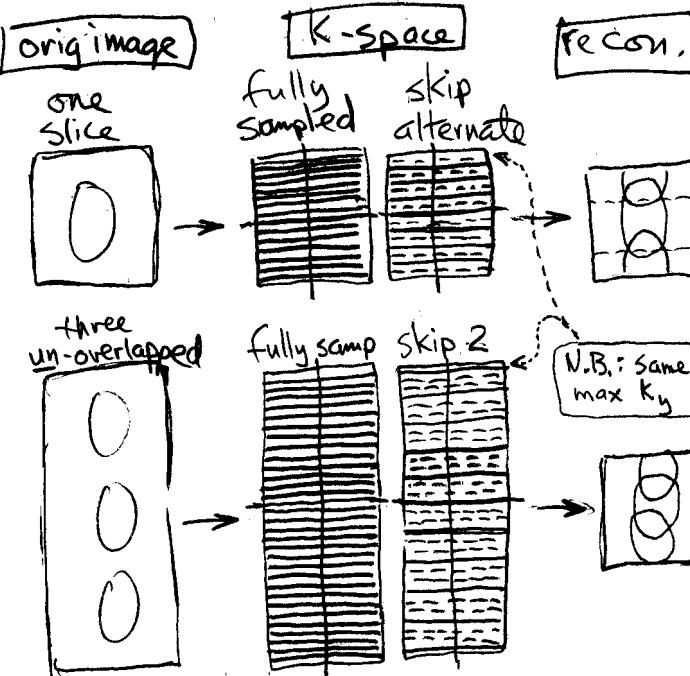


pulse-5d

## MULTI BAND/BLIPPED CAPI

(cont.)

- relation between leave-one-out aliasing and nominally fully-sampled SMS

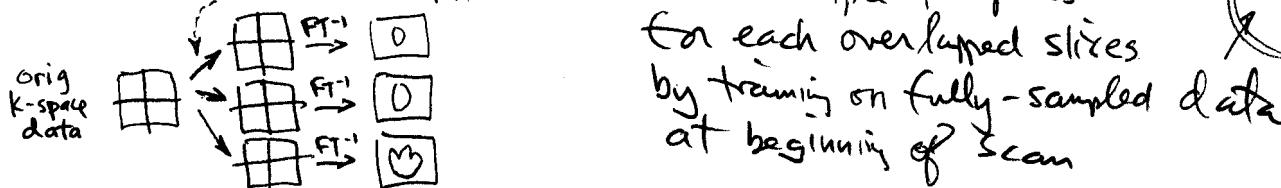


- leave alternate lines out wraps image
- SENSE/GRAPPA can fix (fill in) b/c local coil view smears k-space data to adjacent spatial frequencies
- nominally, w/ SMS we record every line of k-space
- but equivalent to leave alternate out b/c our multi-slice FOV was not big enough

### - slice-GRAPPA

↳ reg GRAPPA → fill in missing lines

↳ slice GRAPPA → fill in multiple k-spaces



i.e. not  
SMS

### - interslice "leakage block"

↳ when training GRAPPA kernel on fully-sampled data, also minimize interslice leakage (split-slice-GRAPPA)

↳ can also do regular GRAPPA on top of this

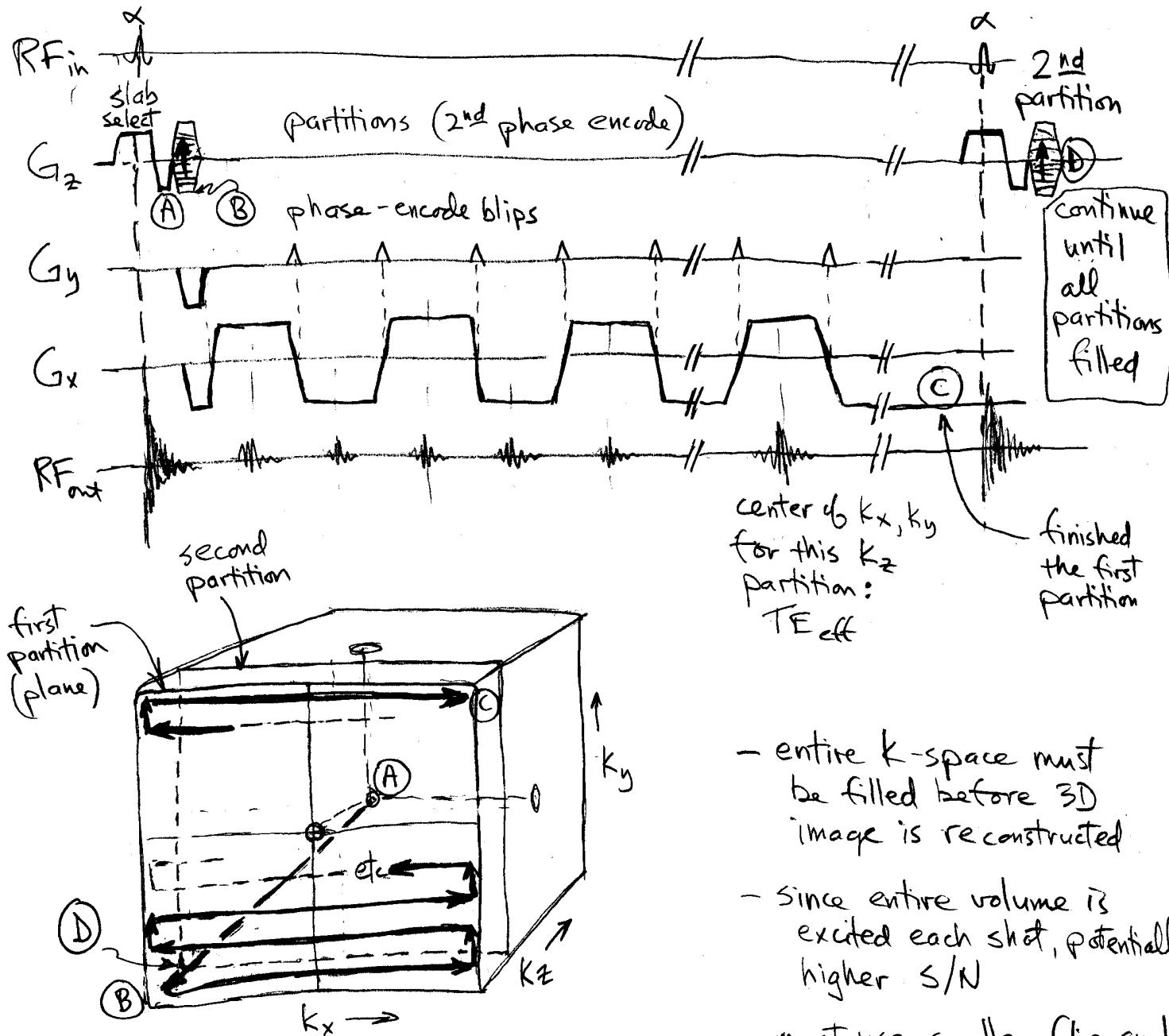
Reason: for diffusion, loss in S/N from undersample cancelled by shorter TE readout (bigger signal)

↳ also, gain from reduced image distortion from shorter  $k_y$  readout

pulse-se

## ECHO-VOLUME IMAGING EVI

- multi-shot (like FLASH) but acquiring one plane of 3-D k-space per shot (can do spin echo, too)



- main issue is movement artifact since data assembled from many shots over several secs
- breathing-induced B<sub>0</sub> problems in different partitions may cause blur

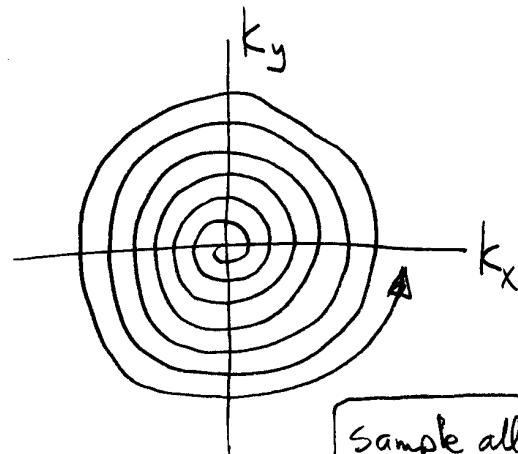
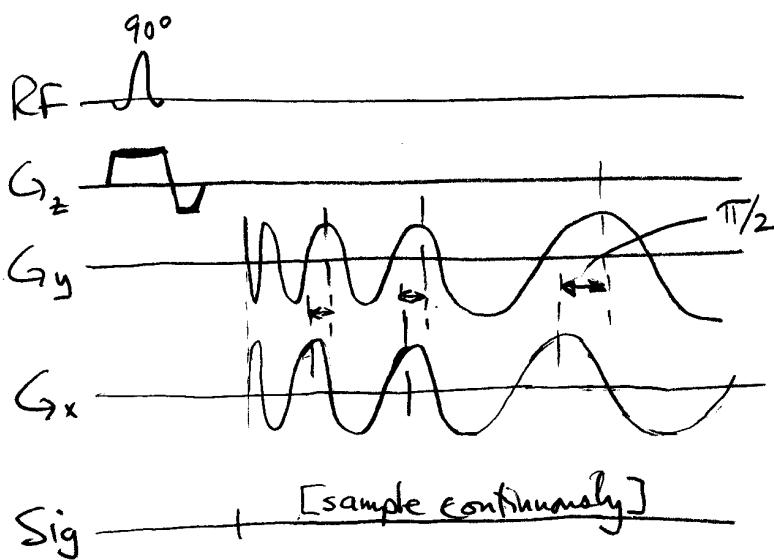
- entire k-space must be filled before 3D image is reconstructed
- Since entire volume is excited each shot, potentially higher S/N
- must use smaller flip angle to avoid killing M<sub>L</sub> since entire volume excited every partition (e.g. every 80 msec)

# SPIRAL IMAGING

- by using smoothly changing gradients (sinusoids) less gradient power required than w/trapezoids (less eddy currents)

earlier EPI hardware like this: sinusoidal gradient waveform from resonant circuit w/non-uniform sampling to get constant  $\Delta k_x$

- sinusoids in both  $G_x$  and  $G_y$  give spiral k-space trajectory



Sample all orientations of each spatial frequency while slowly increasing spatial freq.

- constant angular velocity goes too fast at large  $k_x, k_y$
- constant linear velocity better but impractical near  $k_x=0, k_y=0$
- compromise: start constant angular, end constant linear  $\rightarrow$  

## Constant angular velocity

$$\omega(t) = \omega_0 t$$

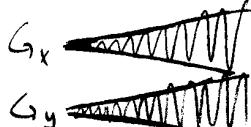
$$\mathbf{k}(t) = A t e^{i \omega_0 t}$$

$$\mathbf{G}(t) = \frac{1}{\tau} \frac{d}{dt} \mathbf{k}(t)$$

$$= A e^{i \omega_0 t} + i A \omega_0 e^{i \omega_0 t}$$

$$G_x(t) = A \cos \omega_0 t - A t \omega_0 \sin \omega_0 t$$

$$G_y(t) = t \sin \omega_0 t + A t \omega_0 \cos \omega_0 t$$



## Constant linear velocity

$$\omega(t) = \omega_0 \tau t$$

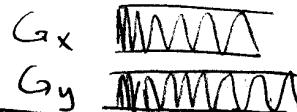
$$\mathbf{k}(t) = A \tau t e^{i \omega_0 \tau t}$$

$$\mathbf{G}(t) = \frac{1}{\tau} \frac{d}{dt} \mathbf{k}(t)$$

$$= \frac{A}{2t} e^{i \omega_0 \tau t} + \frac{A}{2} \omega_0 e^{i \omega_0 \tau t}$$

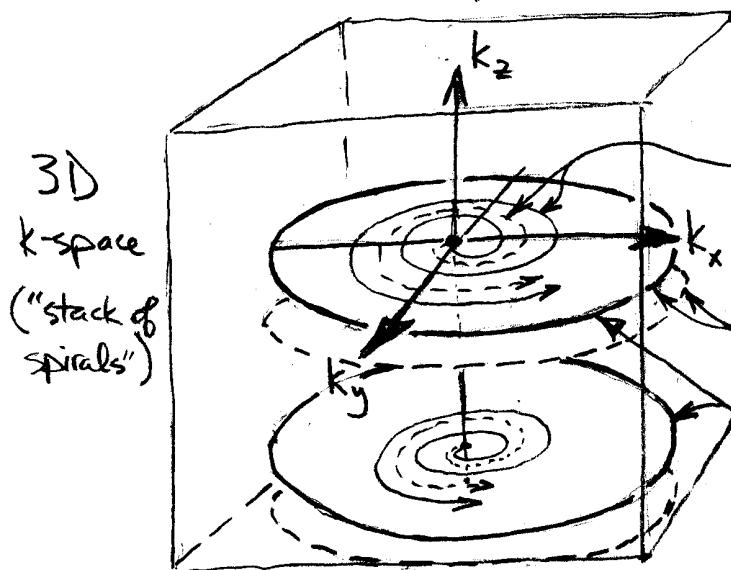
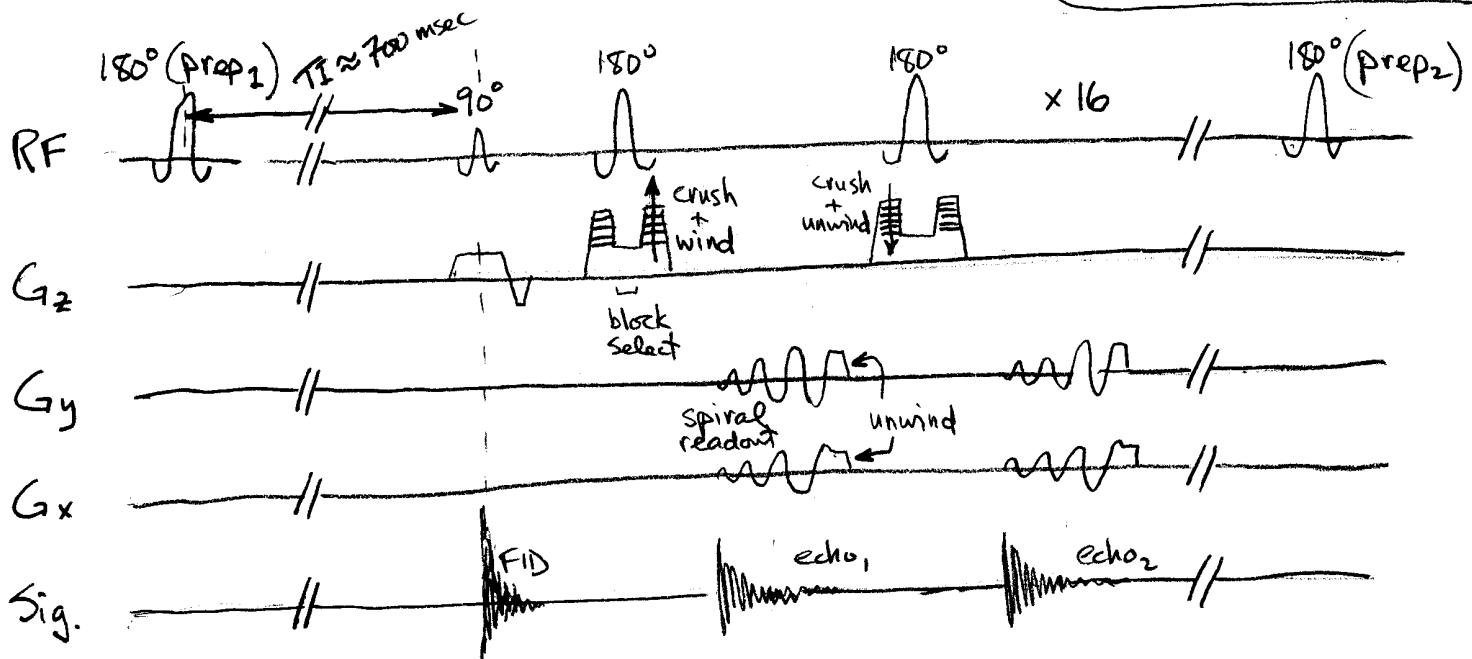
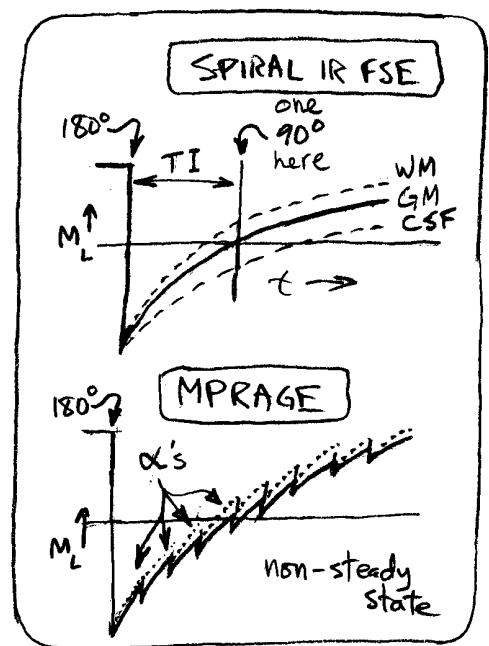
$$G_x(t) = \frac{A}{2t} \cos \omega_0 \tau t + \frac{A}{2} \omega_0 \cos \omega_0 \tau t$$

$$G_y(t) = \frac{A}{2t} \sin \omega_0 \tau t + \frac{A}{2} \omega_0 \sin \omega_0 \tau t$$



# SPIRAL 3D IR FSE (from Eric Wong)

- 3D: block select vs. slice select
- FSE: multiple echoes from one  $90^\circ$
- spiral: multiple spirals vs. multiple lines
- interleaved spirals (like FSE interleaves)
- true IR (vs. MPRAGE)
  - all echoes after  $90^\circ$  derive from mag w/ same T1 contrast (vs. non-steady-state)
- possible to preserve sign
- high, uniform contrast, but lots of waiting (TI), high BW

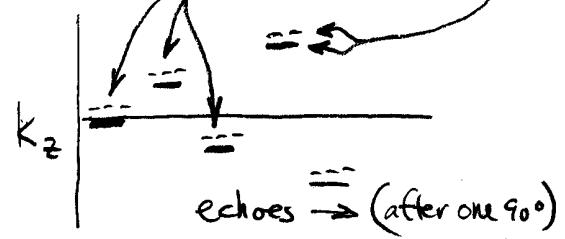


loop order

spiral interleaves

$k_z$  interleaves

$k_z$  echoes

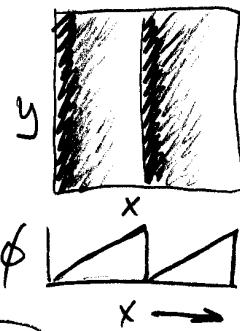


# PHASE ERRORS & ECHO-CENTERING ERRORS

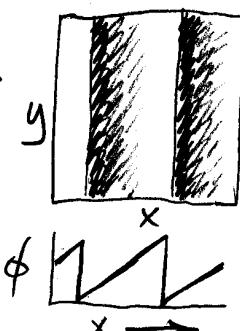
## Phase Errors

- anything that causes a deviation of the  $B_2$  field strength from the expected value ( $B_{0,z} + G_{x,z}x + G_{y,z}y + G_{z,z}z$ ) changes precession frequency and therefore, expected phase angle
- incorrect phase of spatial frequency stripes results in a shift in space in the magnitude image after reconstruction

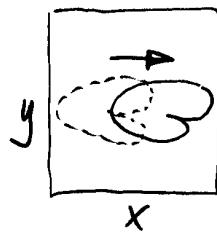
phase stripes in image domain (one k-space point)



phase error



add other spatial freqs in



(phase) Fourier shift theorem  
phase shift in spatial freq. domain causes spatial shift in image domain

- first defense: freq prescan

- refine w/ shimming and  $B_0$ -mapping/phase unwrapping before reconstruction

$$I(x - \textcircled{x}_0) = \int_{k_x} e^{-i2\pi k_x(x - \textcircled{x}_0)} S(k_x) e^{i2\pi k_x x} dk_x$$

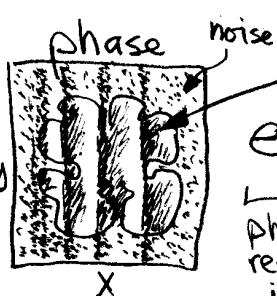
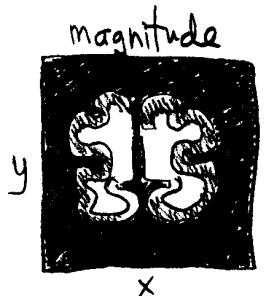
x-offset in image      phase shift in signal      k-space signal

multiply by complex same as add to exponent:  $x - x_0$

N.B.: this is a "pedestal" of phase, not a gradient

## Echo Centering Error

- if realignment of all spins ( $k_x = k_y = 0$ ) doesn't occur at the middle of read gradient, echo is shifted
- Since echo is in spatial frequency domain, this is frequency shift
- spatial frequency shift results in wrapping in phase image after reconstruction  
 ↳ magnitude image unchanged



$e^{i2\pi k_x x}$   
phase shift in reconstructed image

$I(x)$  =  $\int_{k_x} S(k_x - \textcircled{k}_x) e^{i2\pi k_x x} dk_x$   
image (complex)

Fourier freq. shift theorem  
freq shift in freq domain causes phase shift in spatial

offset in spatial freq. space

artifacts - 1

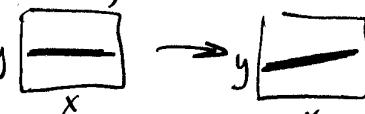
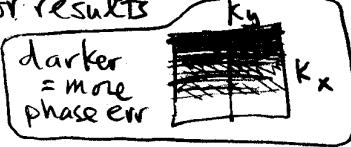
## FAST SCAN ARTIFACTS

### EPI vs. Spiral

brain-induced field defects lead to phase errors

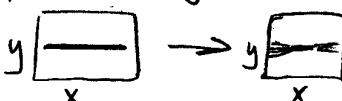
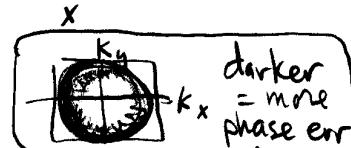
#### EPI

- $G_x$  readout gradient strong → [field defects smaller percentage less deformation of  $k_x$  (vertical stripe components)]
- $G_y$  "blips" are weak and total  $G_y$  record time much longer (5 times) than standard readout (50 ms vs. 10 ms)
- an extra gradient in the x-direction, for example, maps and unmaps phase as a function of x-position
- but  $G_x$  big, so effect on freq.-encode direction is much less than on phase-encode direction, where error accumulates
- for a given x-position, the strength of the spurious gradient is constant, so the accumulation of phase error results in a shift in the y-direction (= k-space spin-stripe displ.)
- the phase error causes a shift in the y-direction proportional to x-gradient strength (= shear) but no blurring  
(N.B. Shift varies w/x-position)



#### Spiral

- with center-out spirals phase errors accumulate in a radial direction
- thus, higher spatial frequencies have more error (= more shearing)
- for spurious x-direction gradient as above, there is a radial blurring, rather than a vertical shift because higher frequency phase stripes misaligned relative to low spatial freq



- for defects with more complex contours in the y-direction (than linear, as above) the vertical shifts (for EPI) will vary with y-position, and may result in signals from different y-positions being reconstructed on top of each other

artifacts-1b

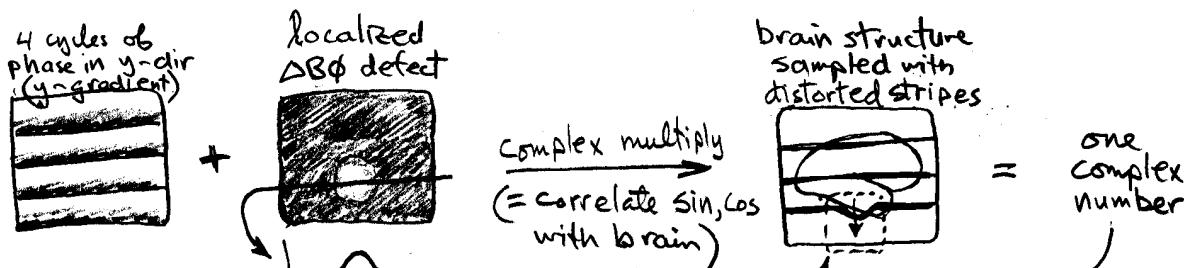
# IMAGE-SPACE VIEW OF LOCALIZED $\Delta\phi$ DEFECT, EFFECT ON RECON

- localized  $\Delta\phi$  defects often arise from air pockets embedded in tissue

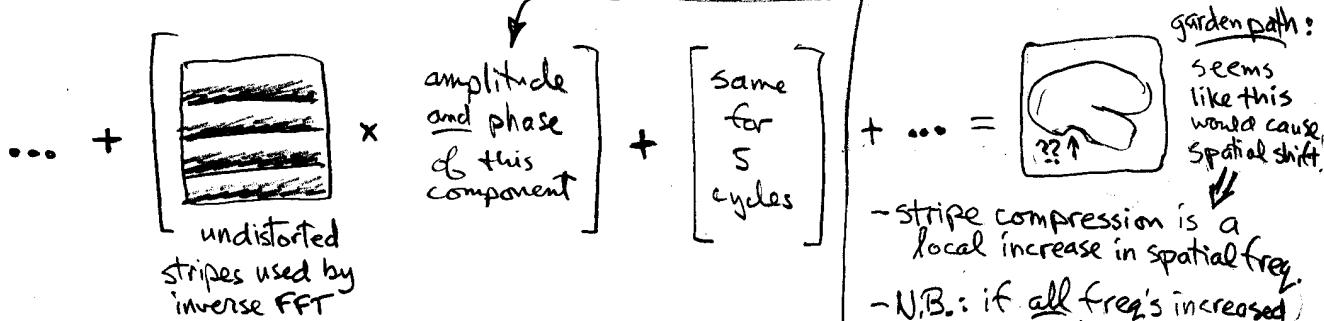
[air in middle/outer ear  $\rightarrow$  indentation in inferior temporal lobe]

[air under olfactory epithelium  $\rightarrow$  orbitofrontal ctx, ant. thal. compression]

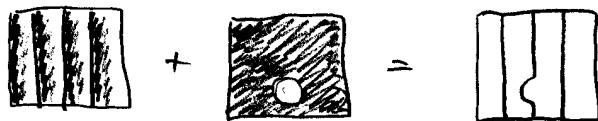
Collect one data (k-space) point



reconstruction from distorted data points

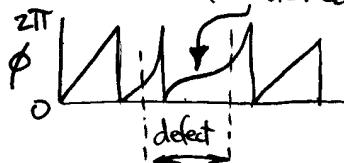


- same defect makes leftward dent in vertical phase stripes



- stripe compression is a local increase in spatial freq.
- N.B.: if all freq's increased same amount, however, this is ex. of Fourier Freq. Shift theorem  
↳ i.e. no amp. image shift! (only  $\phi$ )
- But if phase error (stripe compression) incremented for successive freq's, shift occurs (See next page)

- spatial information can be lost when continuous changes in phase are flattened by  $\Delta\phi$  defect

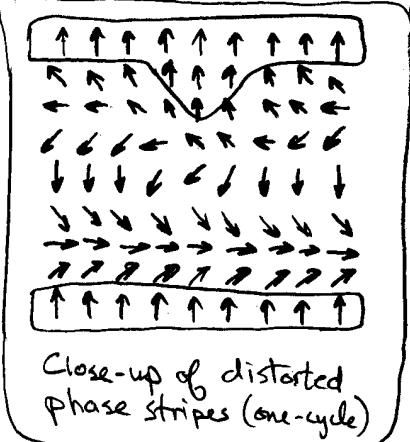


- shifts can pile multiple pixels on top of each other into one bright pixel

- local estimates of  $\Delta\phi$  needed to correct images

1) fieldmap method: multiple TE's to est. local  $\Delta\phi$  from  $\phi/TE$  slope  
shift each image pixel proportional to  $\Delta\phi$  in phase-encoded dir.

2) point-spread-function: extra phase encode to estimate P.S.F. (should be δ-function)  
deconvolve distorted image in phase-encode direction



artifacts- 1c

## LOCALIZED $B\phi$ DEFECT, EFFECT ON RECON (2)

- when local  $B\phi$  defect disturbs image space phase stripes during signal acquisition, estimates of local spatial freq. are affected (compressed stripes = higher spat. freq.)
- if each successive  $k_y$  line recorded w/ same echo time (e.g., w/ single line phase encoding) this will correspond to constant spat. freq. offset in  $k$ -space
- a  $k$ -space freq. offset only results in image space phase shift (Fourier freq. shift theorem), which is invisible in amplitude image (cf. echo cent. error)
- however, with w/EPI, static  $B\phi$  defect causes more and more local displacement of image phase stripes for each additional  $k_y$  line
  - that is, later lines have greater spat. freq. offset
  - effectively stretches  $k$ -space in  $k_y$  direction
  - same num samples to higher spatial freq. shrinks FOV (squishes voxels – see FOV page)
- when image is reconstructed, region with local  $B\phi$  defect shifted oppositely
- Thus, local shift effect due to combination of 3 things:
  - 1) static local  $\Delta B\phi$  defect
  - 2) successive increases in phase error for successive spat. freq. measurements during long EPI readout
  - 3) small size of  $k_y$  phase encode blips relative to  $B\phi$  defect (much less of this effect in freq. encode direction)
- Respiration (which affect  $B\phi$ ) in 3D FLASH might cause similar effect within  $k_z$  partition (if successive spat. freqs.)

N.B.  $B\phi$  affects image phase of all spatial frequencies. If we add, e.g., 90° phase, this means higher freq. image stripes move less since each cycle covers less space:  $\Delta B\phi$

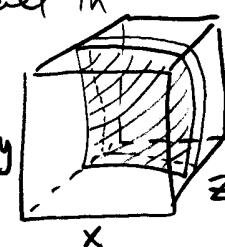
artifacts - 2

## GRADIENT NON-LINEARITIES

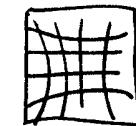
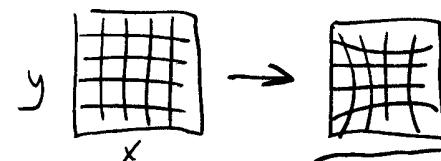
$B_z$   
 $B_x, B_y$

- ideally the  $G_x$ ,  $G_y$ , and  $G_z$  gradient coils attempt to impress a linear variation onto the  $z$ -component of the  $B$  field —  $B_z$  — in the  $x$ ,  $y$ , and  $z$ -directions
- in practice, gradient coils are non-linear (esp. printed-circuit-like)
- non-linearities are worse in smaller coils, but also in higher performance coils, designed for post-processing correction of distortions
- non-linearities result in phase errors, which result in 3-D image distortion

these effects do not build up over time in phase-encode direction since they only occur when gradients are turned on.

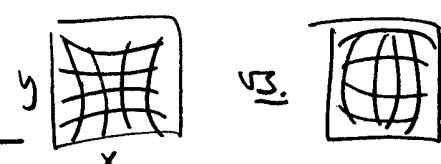


Fourier Shift theorem



these distortions are predictable and can be corrected

- some scanners correct these differently for 3-D scans (all directions), 2-D scans (just in-plane), and EPI scans (no corrections!)
- this can result in errors approaching 1 cm in funct-struct overlays
- different coil designs have different directions of distortion (!)

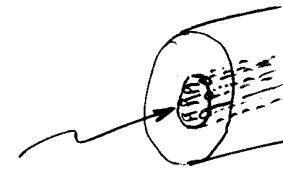


- the assumption of non-Maxwellian gradients result in additional phase errors
- these can also be corrected since the  $B_x$  and  $B_y$  components are known

that is, the assumption that gradients cause no field in the  $B_x$  and  $B_y$  direction

artifacts - 3

## SHIMMING AND $B_0$ -MAPPING



- passive iron shims inserted to flatten  $B_0$  field
- additional coils (usu. in the gradient coils) can be statically energized in an attempt to flatten the  $B_0$  field (a few ppm good)
- primary use is to compensate for defects in flatness present without a sample in the magnet (geometric imperfections, impurities in metal, etc) (= several hundred ppm)

[ linear shim coils impose gradients in x, y, and z  
higher order shims impose higher order spherical harmonic field components (e.g.  $z^2$ ) ]

- secondary use is to compensate for inhomogeneities caused by introducing the sample into the  $B_0$  field

- local resonance offsets caused by  $B_0$  defects estimated from images  
 ↳ e.g., sample phase at multiple echo times



- fit defective field using combination of fields generated by shim coils, then add these corrections to base shim currents

↳ this only corrects spatially gradual field defects  
 ↳ local defects due to air in sinuses much higher order than shims

- after shimming, field map measured again

- image voxel displacements calculated from resonance offset map are used to un warp the reconstructed magnitude image

- for EPI images, assume displacements all in phase-encode direction (since freq encode gradient is strong relative to defects)

because:  
 longer time to echo means more time for phase to be retarded (or advanced) by  $B_0$  defects

artifacts-4

## NAVIGATOR ECHOES

### - 1D navigator

#### B $\phi$ drift problem

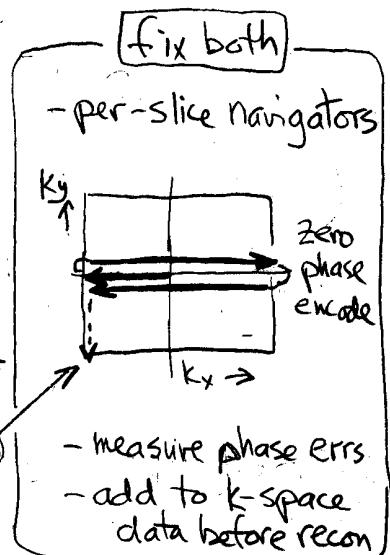
- slow up/down drifts in B $\phi$  continuously occur
- a pedestal in B $\phi$  is pedestal in phase (not gradient) which causes spatial shift (Fourier shift theorem)
- in EPI, mainly affects phase-encode dir b/c small blips long readout
- result is successive volumes drift in phase encode dir

[daily B $\phi$  decay (e.g. 0.05 ppm/hr)  
passive shim heating by gradients]

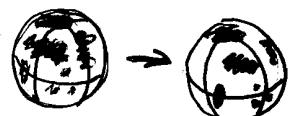
#### Gradient balance problem

- unequal L/R readout gradients cause L/R shift in position of even/odd lines in k-space causing N/2 (Nyquist) ghosting  $\Rightarrow$  another phase error

begin  
std EPI



### - 3D navigator: collect 3D sphere in k-space



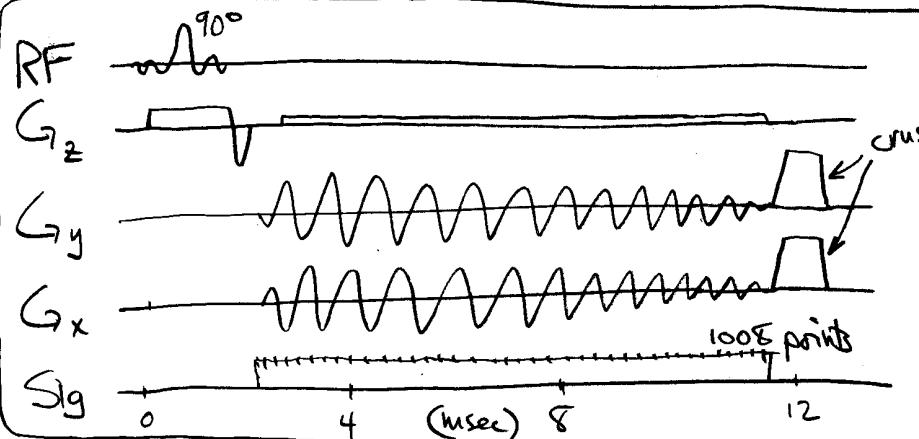
[rotation of object  $\rightarrow$  rotation of k-space amplitude pattern]

[translation of object  $\rightarrow$  phase shift of k-space phase (Fourier shift)]

- sample at sufficient <sup>k-space</sup> radius to pick up high spatial freq features
- N.B.: excite whole volume
- do N,S Hemispheres separately (less T<sub>2</sub><sup>\*</sup>, cancel EPI-like error accumulation)

Welch et al. (2002) MRM  $\rightarrow$

$\hookrightarrow$  equator  $\rightarrow$  up, equator  $\rightarrow$  down



$$z(n) = \frac{2n - N - 1}{N} \text{ total pts}$$

$$y(n) = \cos(T\pi \sin^{-1} z(n)) \sqrt{1-z^2(n)}$$

$$x(n) = \sin(T\pi \sin^{-1} z(n)) \sqrt{1-z^2(n)}$$

(skip pokes — slew rate too high)

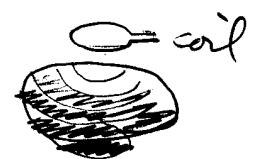
- can be used for prospective motion correction (rotate, translate w/ gradients)
- better estimate, because of speed, than full TR of EPI images (27 ms vs. 2-4 sec)
- May need to smooth rot/trans estimates across time (e.g. Kalman filter)

artifacts-5

## RF FIELD INHOMOGENEITIES

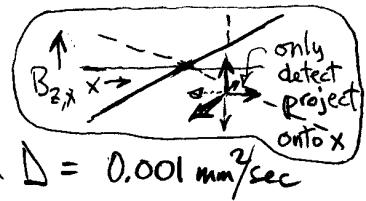
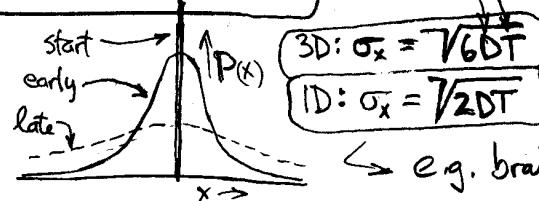
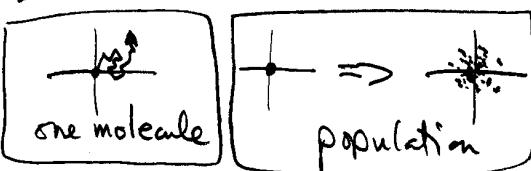
$B_1$ , inhomogeneities

- receive coil inhomogeneities alter the amplitude of the received signal, altering the reconstructed proton density in a spatially varying way
  - ↳ variations can be used (cf. GRAPPA, SENSE) and/or corrected
- transmit coil inhomogeneities affect the flip angle in a spatially varying way (can affect contrast: FLASH)
  - ↳ potentially worse (why local transmit is still in progress)
  - ↳ usu. fixed by using a large transmit coil (e.g. body coil)
- RF penetration at higher fields (= higher RF frequencies) is less uniform:
  - 1) decreased RF wavelength (closer to size of head) at higher freq.
  - 2) increased permittivity ( $\epsilon$ ) and conductivity ( $\sigma$ ) at higher field
- 2<sup>nd</sup> advantage of the fall-off in signal recorded with a small, receive-only RF coil is better signal-to-noise (less noise received from other parts of brain)
- different sensitivity functions from different coils can be used to scan less lines in k-space (GRAPPA/SENSE/SPACE-RIP) normalization ("pre-scan normalize")
  - record low-res volume (b/c coil fall-off is smooth) through both body coil and small coil(s)
  - divide  $\frac{\text{small coil}}{\text{body coil}}$  at each voxel to determine receive field
  - use receive field to normalize main image(s)
    - [ see also: qT1, MP2RAGE, T1/T2 ]



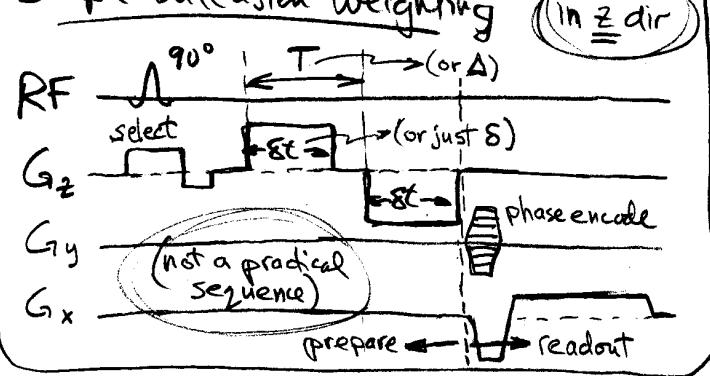
classical diffusion coefficient  
( $\propto$  time for diffusion)

# DIFFUSION - WEIGHTED IMAGING



e.g. brain  $D = 0.001 \text{ mm}^2/\text{sec}$

## Simple diffusion weighting



- "apparent diffusion coefficient" → calculate from  $b=0$  image and at least 6  $b="large"$  (e.g. 1000) images
- after obtaining  $3 \times 3$  tensor, calc eigenvectors/values to find orientation & shape of diffusion ellipsoid
- two useful scalar values from 3 eigenvalues:

apparent diffusion coefficient  
mean diffusivity

$$\text{ADC} = MD = (\lambda_1 + \lambda_2 + \lambda_3)/3$$

fractional anisotropy

$$FA = \sqrt{\frac{(\lambda_1 - \lambda_2)^2 + (\lambda_2 - \lambda_3)^2 + (\lambda_1 - \lambda_3)^2}{2(\lambda_1^2 + \lambda_2^2 + \lambda_3^2)}}$$

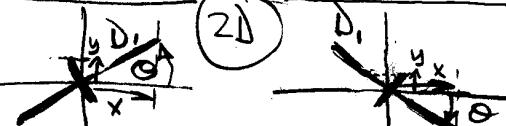
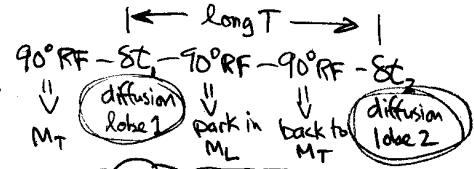
- to get large  $b$ , need large  $G$ ,  $\delta t$ ,  $T$

## 1) Anisotropic Diffusion (Gaussian)

- measure  $D$  along multiple axes

- have to measure tensor, not scalar  
↳ even for determining one primary direction

long  $\delta t$ ,  $T$  can give spurious T2-weighting  
can use stimulated echo to get long  $T$  w/ less T2-weighting



$$D = [U_x \ U_y \ U_z] \begin{bmatrix} D_{xx} & D_{xy} & D_{xz} \\ D_{yx} & D_{yy} & D_{yz} \\ D_{zx} & D_{zy} & D_{zz} \end{bmatrix} \begin{bmatrix} U_x \\ U_y \\ U_z \end{bmatrix}$$

$D_{xx} = \lambda_1, D_{yy} = \lambda_2, D_{zz} = \lambda_3$   
get eigenvectors & eigenvalues,  $\lambda$

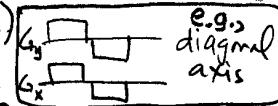
$$D = U^T \cdot D \cdot U$$

scalar diffusion      diffusin tensor      measurement direction

- isotropic: off diag=0, diag=equal

- since  $D$  is symmetric, need minimum of 6 diff. measurement directions

(more is better!)



- without 3rd number ( $\theta$ )  
 $x$  &  $y$  projections same

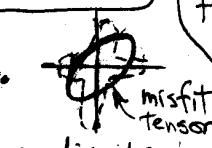
voxels	Tract Tracing
0 1 0 0	1) Markov process
0 0 0 0	2) crossing fibers
0 0 1 0	3) "freeway ramp" prob
0 0 0 1	4) sharp turns into gyri

## Diffusion Surface (non-Gaussian)

fit w/ spherical harmonics  
- need to measure diffusion in many directions (>6) to properly characterize even 2 main directions

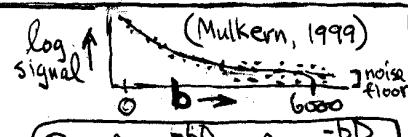


VS.

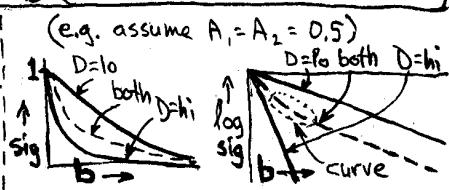


## 2) Length Scale by multiple b-values

- fit line to semilog signal as funct of  $b$
- if not straight line: multi-exponential, e.g., hi ADC/fast/extr. vs. lo ADC/slow/intracellular
- hi ADC/fast/extr. vs. lo ADC/slow/intracellular



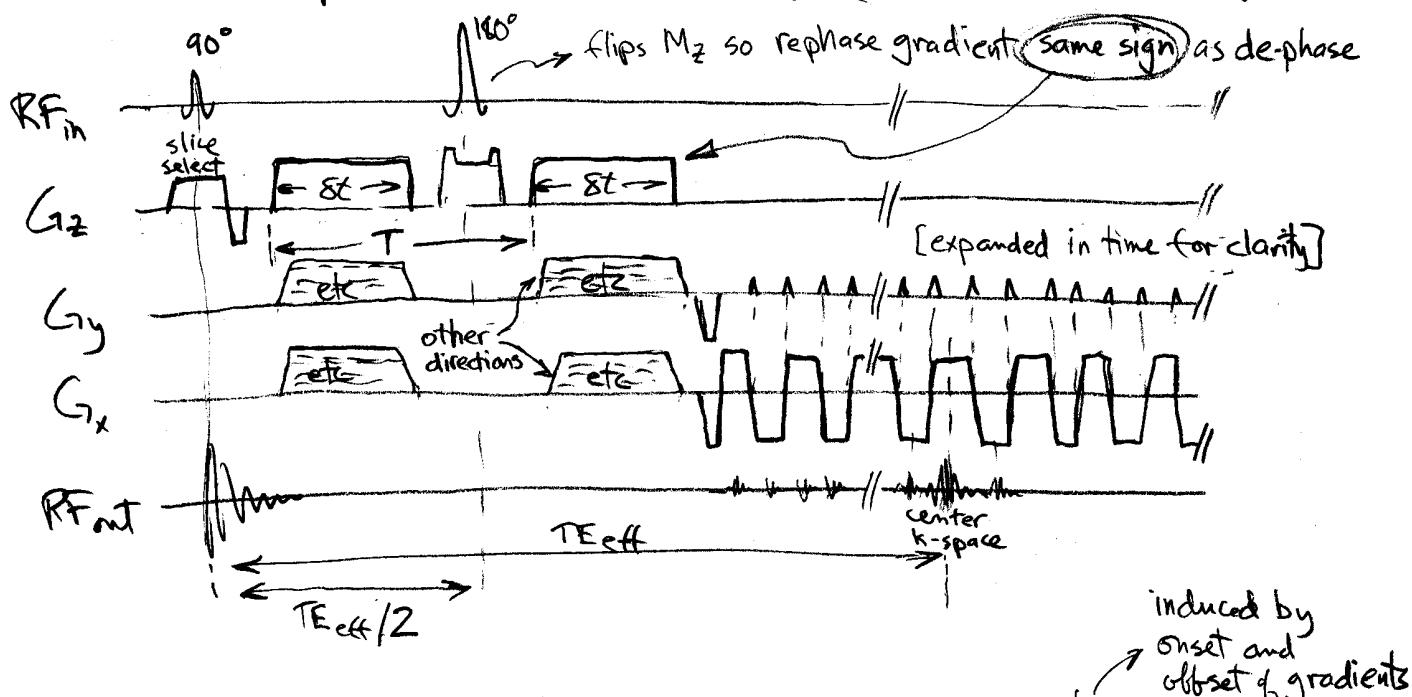
$$S = A_1 e^{-bD_1} + A_2 e^{-bD_2}$$



diffusion-2

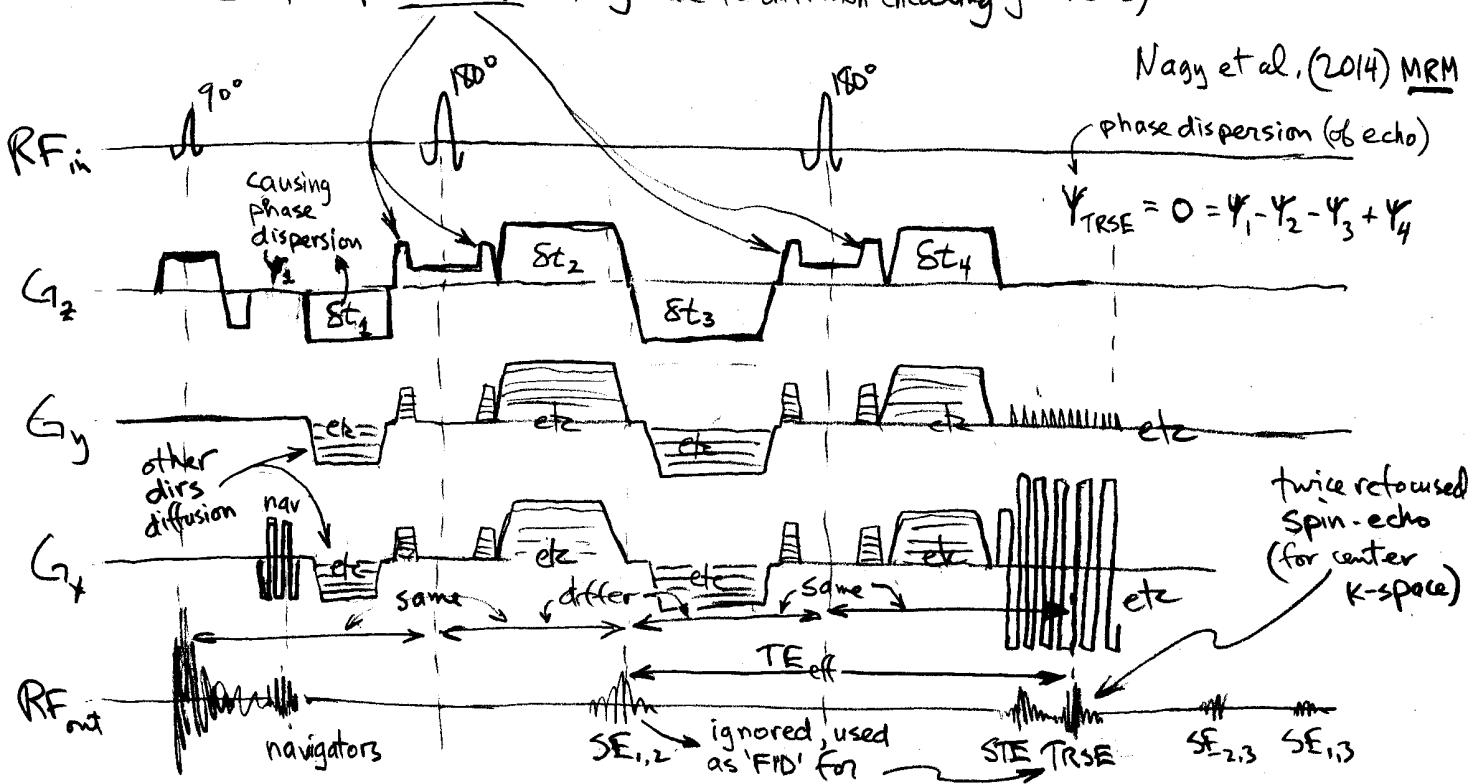
## PRACTICAL DIFFUSION-WEIGHTED PULSE SEQ

- spin-echo 'Stejskal-Tanner' EPI seq. (standard on scanners)
  - ↳ allows longer TE



- eddy-currents are long time-constant currents in metal of scanner that distort B field → spatial image distortion

- "doubly-refocused" spin echo sequence (DSE) can cancel the effects of eddy current (w/partic. time constants)
  - ↳ (also, keep crushers orthogonal to diffusion-encoding gradients)

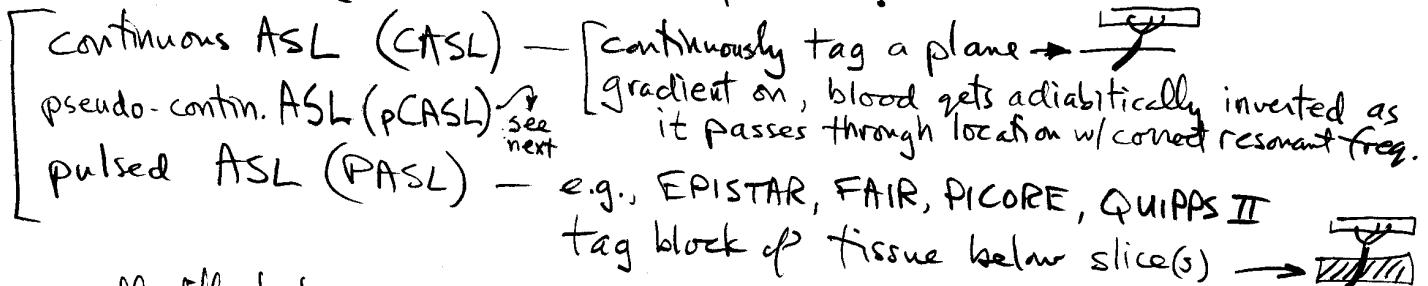


## PERFUSION - ARTERIAL SPIN LABEL

- basic idea: tag blood below area of interest  
collect control & tagged image  
assume directional input flow!

→ (like inversion recovery)

- tag is 180° pulse
- sign not problem when delay long enough (see below)



- small diff's between control and tag (~1%)  
 ↳ requires accurate balancing of control & tag images, control mag. transfer

- contrast problems: transit delays → biggest confounding factor  
relaxation rate diff's  
venous clearance (vs. microspheres, which get stuck!)

→ transit delays

TI	500	1000	1500 msec
	large vessels	frontal temporal parietal	occipital

### solutions for quantitative

- insert delay so all spins arrive into low velocity capillaries
- kill end of tag to reduce spatial variation of tag

### - QUIPPS II - quantitative perfusion

#### TURBO ASL

- use TI longer than TR
- omit QUIPPS tag ending

tag pulse

control img.

control pulse

Tag image

tag pulse

control img.

2X faster but limited slice #

- 1) pre-saturate spins in target slices → adiabatic pulse (freq sweep)

- 2) tag - 180° pulse below slices

- 2) control - 180° pulse above slices (to control off-resonance)

- 3) saturate tagged block to end tag (TI<sub>1</sub>)

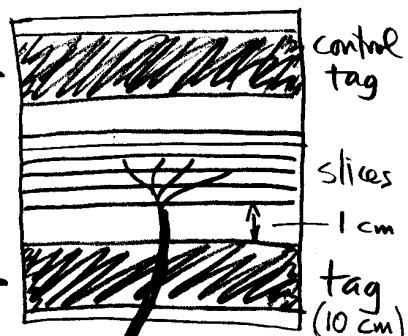
[both tag and control]

can use train of thin slices pulses at top of tag band

- 4) EPI or spiral images of target slices (TI<sub>2</sub>)

↳ image most distal slice last to cancel delays

↳ fast between slice so imaging excitations don't get interpreted as flow



$$\Delta M \cong \text{flow} \times [2M_0 \cdot T_{I_1} e^{-T_{I_2}/T_{1A}}]$$

can extract flow and BOLD

adjacent subtractions minimize movement artifact

- 1) alternate tag and control, GRE TE = 30 ms

control - tag → flow

control + tag → BOLD

↳ both BOLD-weighted

Tag - control - tag - control ...

- 2) dual echo spiral

k=0 early → hi S/N flow  
 TE = 30 ms → BOLD

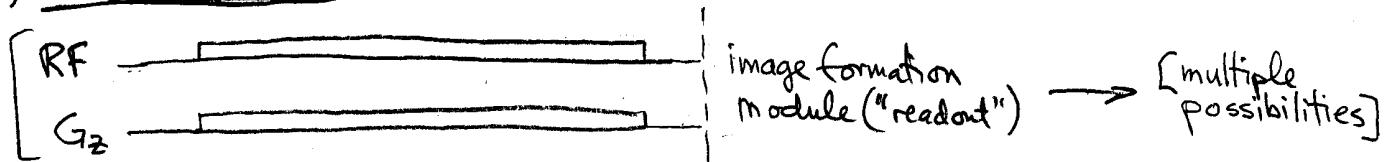
perfusion-2

## PERFUSION — pCASL

- original CASL (continuous arterial spin labeling) requires RF on continuously to adiabatically invert blood flowing through one plane

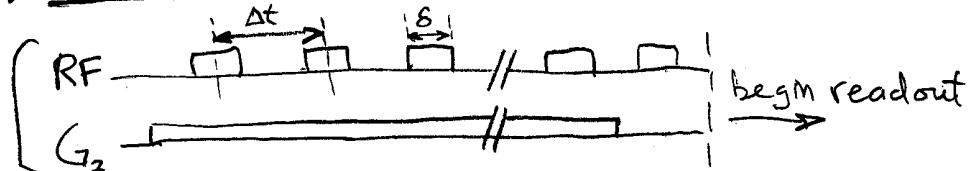
↳ can only image one slice (b/c dephasing from gradient)  
 ↳ hard to keep RF on continuously on modern scanner (esp. BC)  
 ↳ can use special purpose RF transmit (separate xmt channel)

### A) original CASL



### B) pCASL — pseudo continuous arterial spin labeling

Dai, Alsop (2008)



— problem: multiple pulsers create aliased slice planes

$$RF(t) = \frac{1}{\Delta t} \text{comb}\left(\frac{t}{\Delta t}\right) \otimes \text{rect}\left(\frac{t}{\delta}\right)$$

[use: convolution of 2 funct equals multiplying their FT's]

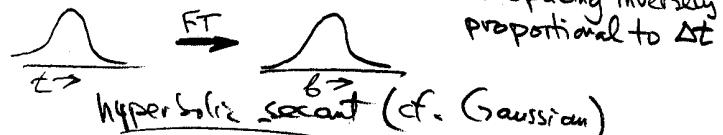
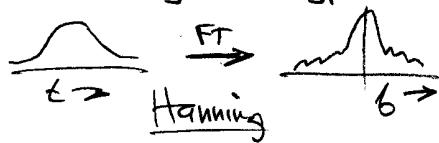
$$F[RF(t)] = \text{Comb}(f\delta t) \cdot \delta\left(\frac{t}{\Delta t}\right)$$

$$\text{comb}(t) = \sum_{n=-\infty}^{\infty} \delta(t-n)$$

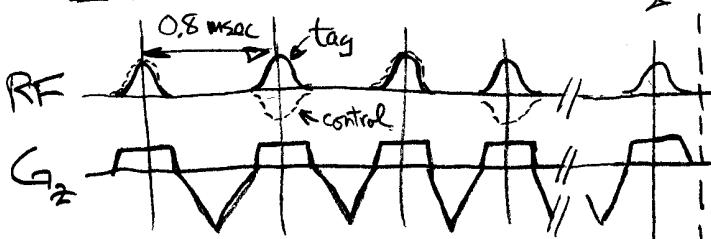
$$\text{rect}(t) = \begin{cases} 1 & \text{if } |t| < \frac{1}{2} \\ 0 & \text{otherwise} \end{cases}$$

↳ aliased labeling planes at:  $f = n/\Delta t$  in frequency space, modulated by broad sinc()

— use Hanning or hyperbolic secant to reduce replicas



### C) pCASL w/ shaped gradients



(control called a "transparent" pulse)

readout

FLASH

EPI

SMS

3D

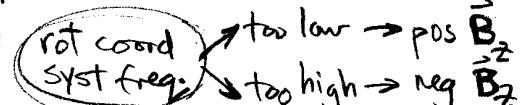
- tag pulses have phase offset respecting gradient

- control identical except every other has  $+\pi$  phase  
 ↳ no net flip

Perfusion-3

## OFF RESONANCE EXCITATION

adiabatic  
RF pulse

- main idea: examine evolution of  $\vec{M}$  vector in rotating coord syst set to "off-resonance"  $\vec{B}_1$  field freq ( $\omega_{rf}$ ), not Larmor freq of  $\vec{M}$  ( $\omega_0$ )
- normally, if rotating coord syst freq set to Larmor freq ( $\omega_{rf} = \omega_0$ ), an actually precessing  $\vec{M}$  will be stationary (ignoring decay)  $\rightarrow$  implies effective  $B_2 = 0$  in rotating
- now, move  $\vec{M}$  to rotating coord syst at  $\vec{B}_1$  freq lower than  $\omega_0$  (assume  $\vec{B}_1 = 0 = \vec{B}_0$ ): existing  $\vec{M}$  will now appear to precess around z-axis:
  - 
  - N.B.: this is precession in already rotating coordinate system! (slow relative to  $\omega_0$ )
  - $$\Delta\omega_0 = \omega_0 - \omega_{rf}$$
    - freq of precession in rotating coordinate syst
    - Larmor freq of  $\vec{M}$
    - rotation freq  $\vec{B}_1$  (= "incorrectly set rotating coord syst frequency")
- thus, viewing  $\vec{M}$  vector in off-resonance rotating coord syst makes it look like additional  $\vec{B}_2$  field is causing "extra" precession
- "extra"  $\vec{B}_2$  component is proportional to  $\Delta\omega_0$  offset
  - $\hookrightarrow$  can be pos or neg: 
- extra  $\vec{B}_2$  adds to  $\vec{B}_1$  resulting in slow precession around tipped axis:  $\vec{B}_{eff}$  (effective)
- extra  $\vec{B}_2$  from any gradient  $\rightarrow$  same effect on  $\Delta\omega_0$  (changes  $\omega_0$  instead of changing  $\omega_{rf}$ )

$$\vec{B}_{eff} = \left( \frac{\Delta\omega_0}{\gamma} \right) \hat{k} + \vec{B}_{Gz} \hat{k} + \vec{B}_1 \hat{i}$$

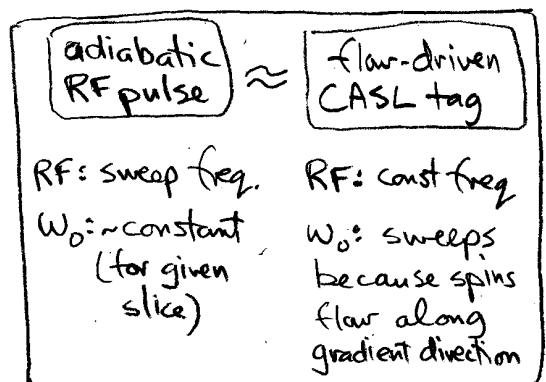
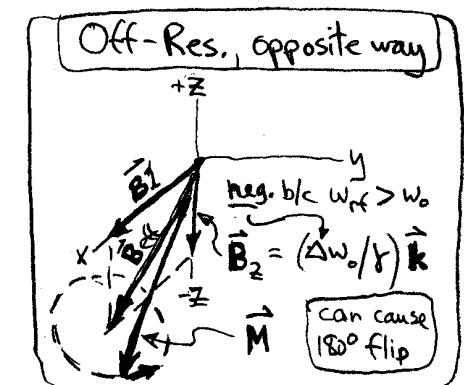
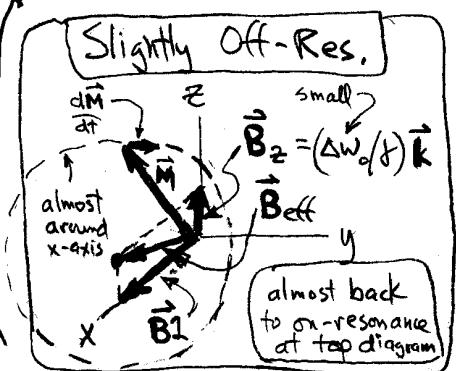
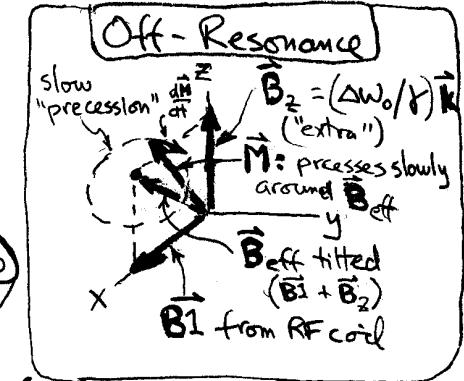
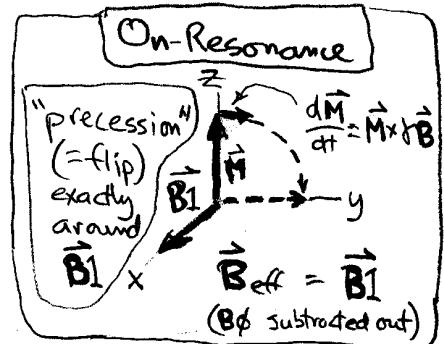
$\nwarrow$  x-axis

effective  $\vec{B}$  in rotating frame set to  $\vec{B}_1$  freq

apparent "extra"  $\vec{B}_2$  from Larmor- $\vec{B}_1$  freq mismatch (pos or neg) (if on-res.  $\rightarrow 0$ )

extra  $\vec{B}_2$  from optional gradient (pos or neg) (additional source of  $\vec{B}_{eff}$  tilt)

transverse RF stim (here, around x-axis)

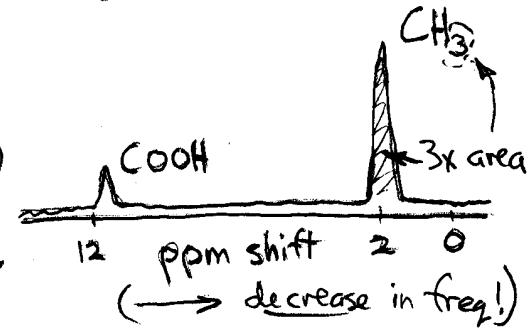
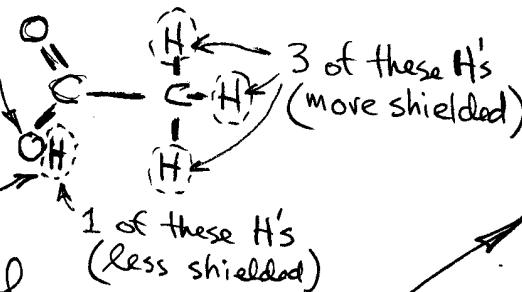


spect-1

# SPECTROSCOPY + IMAGE

- chemical shift: small displacement resonant freqs due to variable shielding of target nucleus (e.g.  $^1\text{H}$ ) by surrounding electron orbitals
- e.g., acetic acid:

Oxygen attracts electron so less shielding of target nucleus



- how we get chemical shift spectrum:



$$\Delta f_1: \Delta f_2: \Delta f_{\text{num}}$$

Larmor oscillations are multiplied (PSD) by center freq to obtain  $\Delta f$  (not MHz high freq)

- data before FT is a series of time-domain samples of the mix of shifted-freq offsets

- FT turns data into "shift spectrum"

- N.B.: opposite "direction" of FTs!

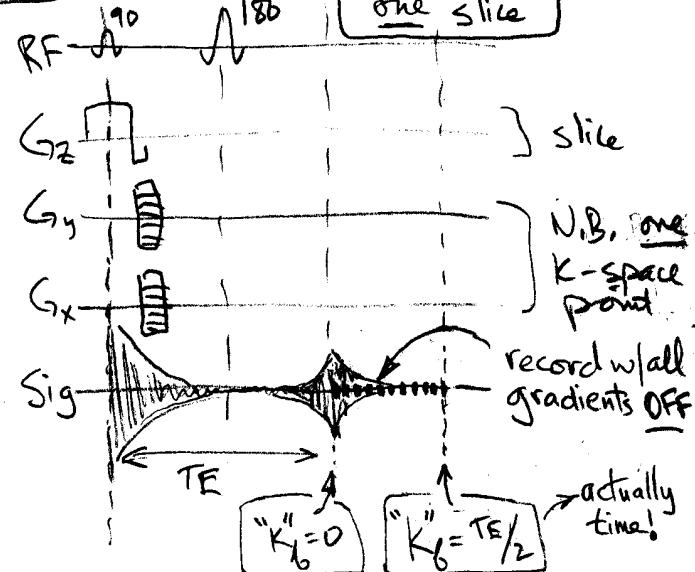
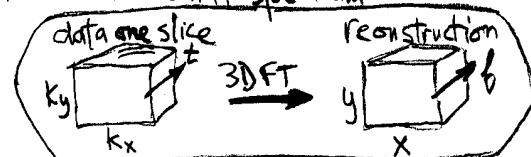
Signal	result
NMR	time domain oscillation samples (shift) $\xrightarrow{\text{FT}}$ shift freq. spectrum
MRI	spatial freq. samples $\xrightarrow{\text{FT}}$ spatial object (like time domain signal)

## Pulse Sequence

- since we are already using phase (=freq) encoding for space, we need an "extra dimension" w/ all gradients OFF!

- use spin-echo to undo built-up chemical shifts, then record chemical-NMR-like signal

$\hookrightarrow$  and FT-it like chemists do!

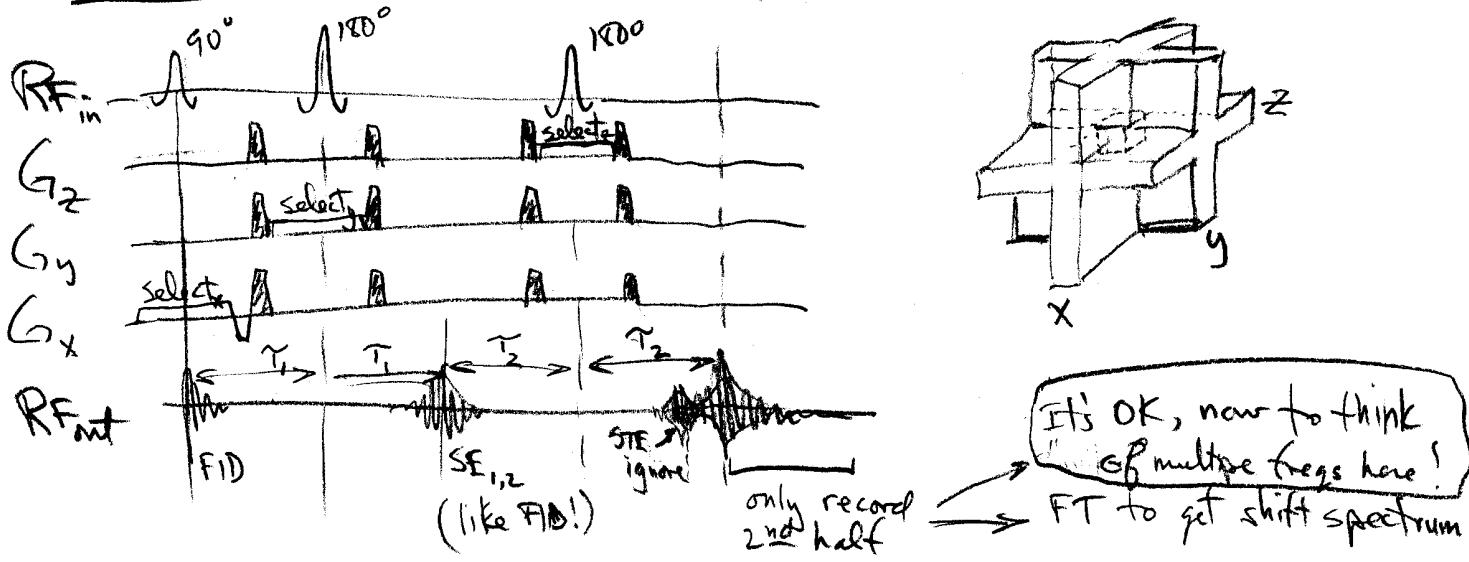


Spec-2

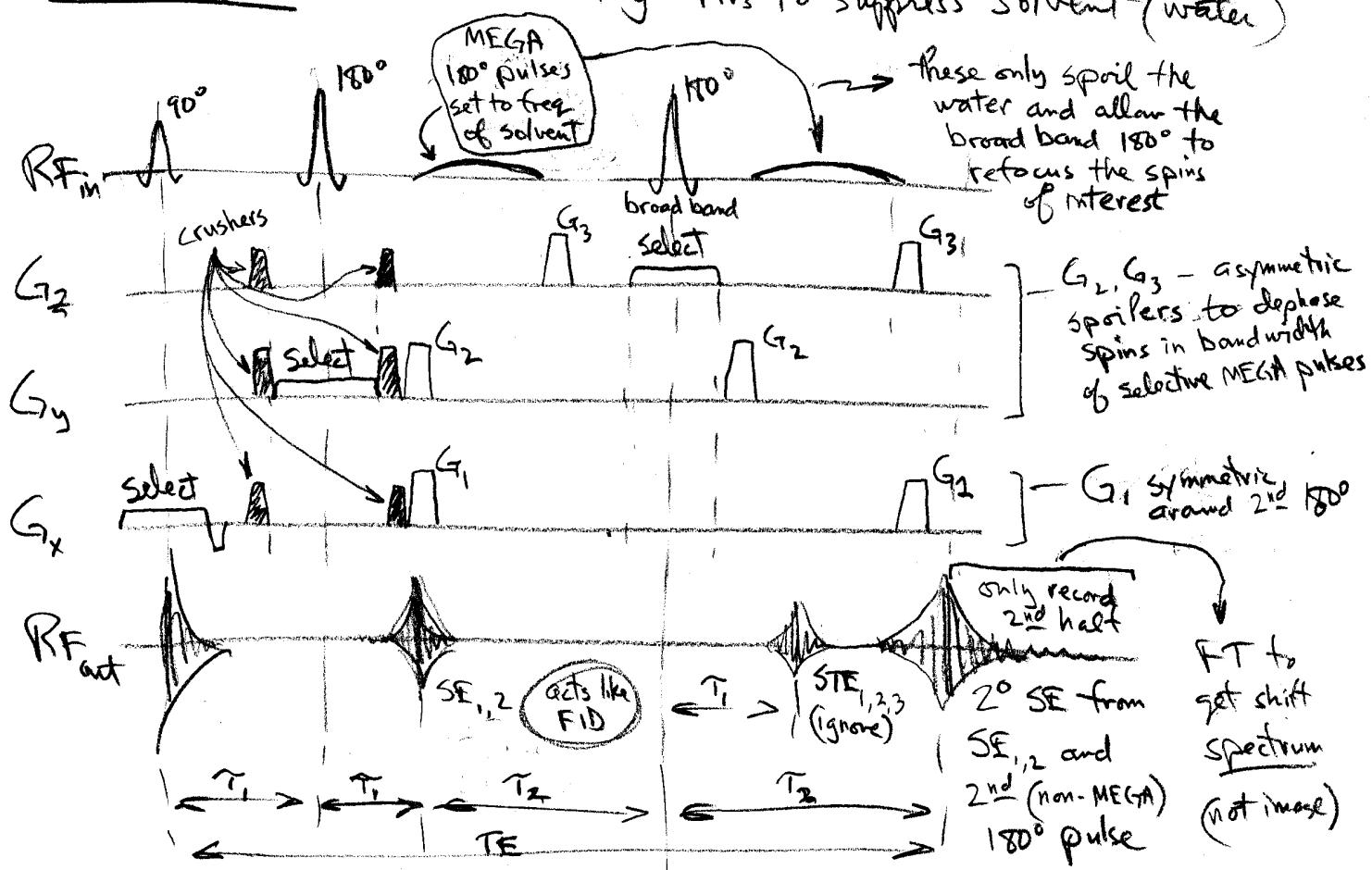
## PRESS, MEGA-PRESS

- usu. single voxel by using 3 orthog. slice selects  
(tho can add PE gradients & more excitations to get multiple vox)

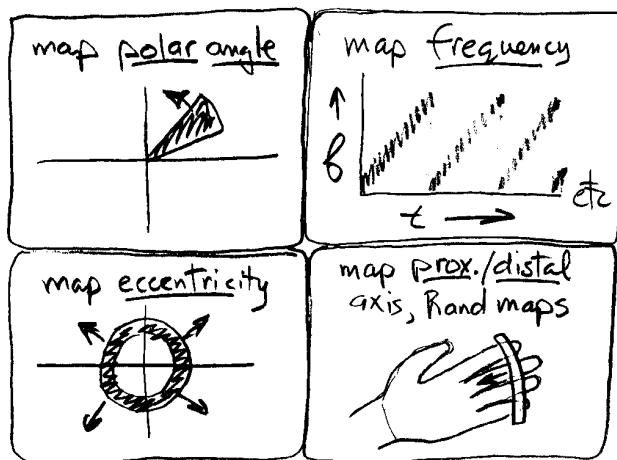
- PRESS ← 3 orthog slice select



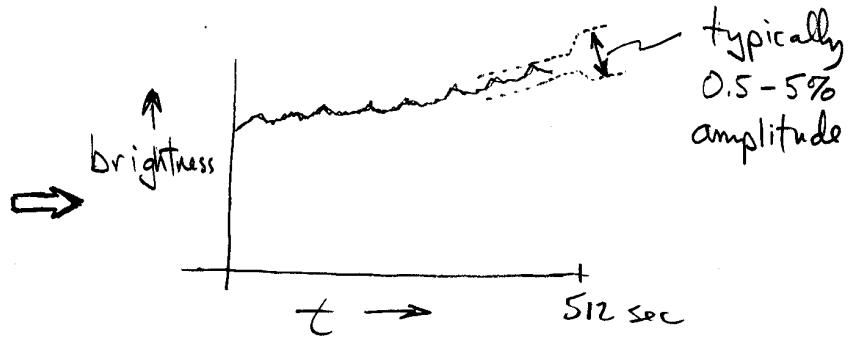
- MEGA-PRESS — add "editing" RFs to suppress Solvent (water)



# PHASE-ENCODED STIMULUS & ANALYSIS



Periodic stimuli (phase-encoded) — e.g., 8 cycles at 64 sec/cycle



strongly periodically activated single voxel time course

remove constant (avg) and linear trend

## calculate significance

- ratio between amplitude at stimulus frequency (= signal) and average of amplitudes at other frequencies (= noise)
- ignore harmonics, low freq (= movement)

## Smooth

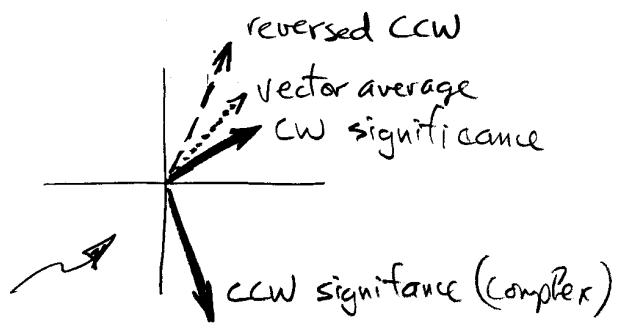
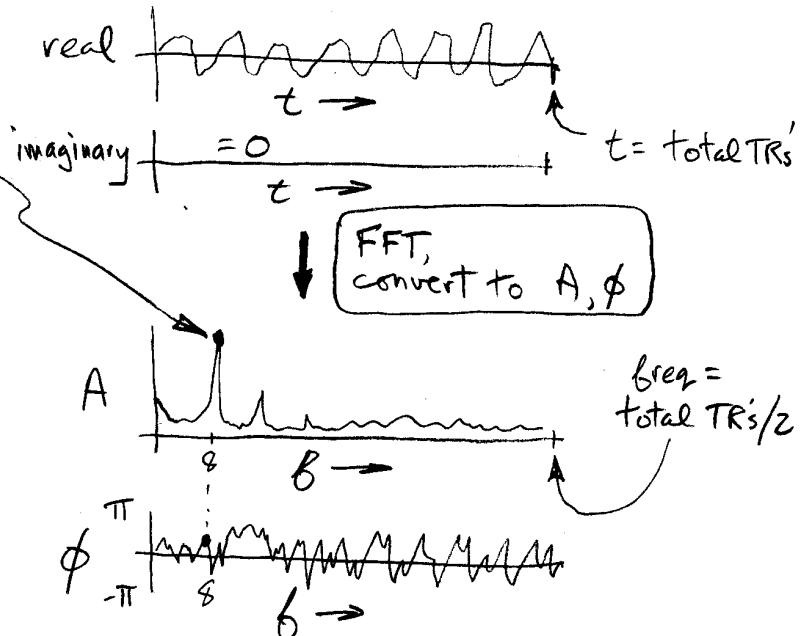
- vector average of complex significance ( $A, \phi$ ) with that at nearest neighbor Surface Points

## display

- plot phase using hue and saturation to indicate significance

## delay correction

- record responses to opposite directions of stimulus (ccw/cw, in/out, up/down)
- vector average after reversing angle of one  
→ penalizes inconsistent more than just avg of angles

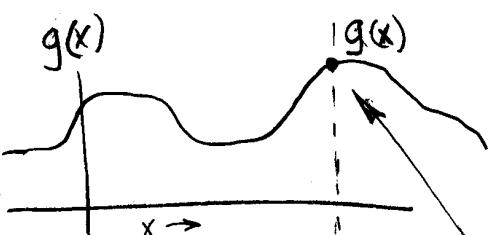


# CONVOLUTION

$$h(x) = f(x) \otimes g(x)$$

or  
 $(f * g)(x)$

- definition of convolution
- commutative



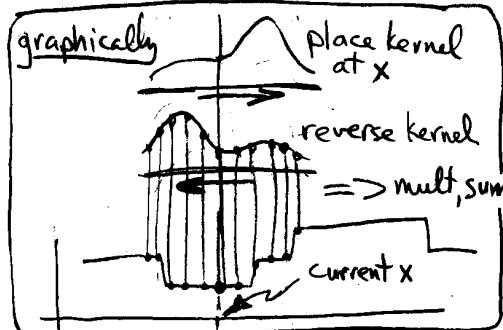
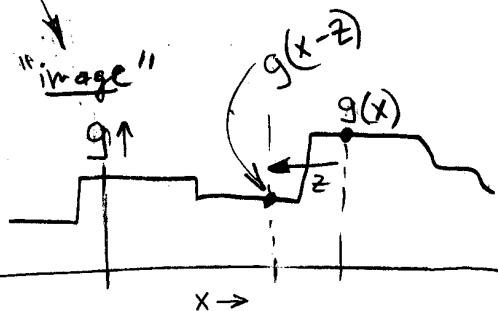
for one  $x$

$$= \int_{z=-\infty}^{z=+\infty} f(z) \cdot g(x-z) dz$$

Sum this product across all  $z$

"kernel"

linear time/space invariant system



- how to calculate one term
- sum across all  $z$  to get the value of the convolution at point  $x$
- move kernel to calculate next  $x$

## Why reverse makes sense

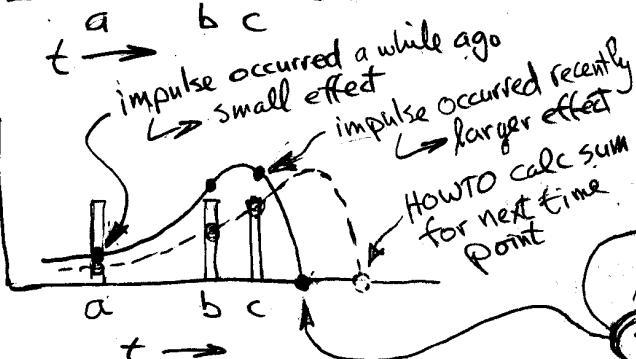
b/c commutative, can think like:  $h(x) = \int g(z) \cdot f(x-z) dz$



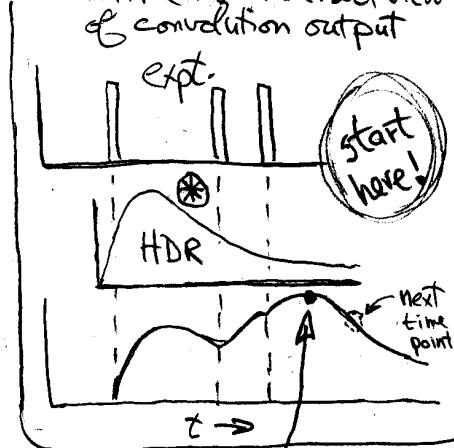
impulse response function (HDR)



impulses (expt. design)



\*\*intuitive non-reversed view of convolution output expt.



N.B. cross-corr same as convolution except no reversal  $g(x+z)$  instead of  $g(x-z)$

N.B. auto-corr Same, except no-reversal and use same-funct for both  $f, g$

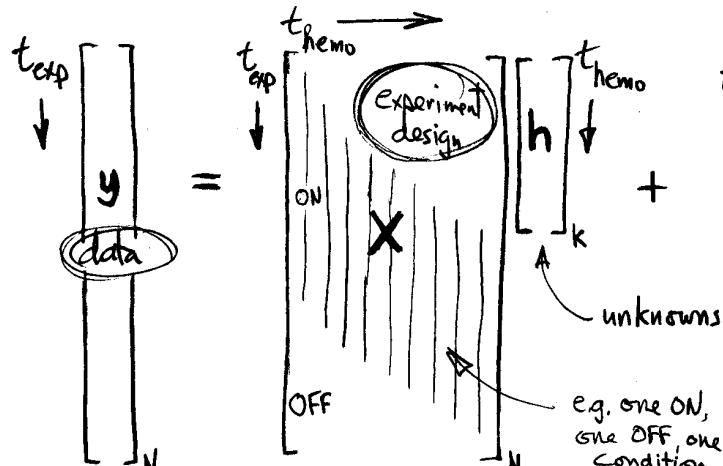
How to calculate convolution output for this time point (only 3 terms in sum; all other zero)

stats-3

# GENERAL LINEAR MODEL

$$\vec{y} = \vec{X}\vec{h} + \vec{S}\vec{b} + \vec{n}$$

data = design · HDR + drifts · weights + noise



"block design"  
better to detect resp.  
"event-related design"  
better to measure HDR shape

→ goal: solve for the hemodynamic response functions,  $\vec{h}$

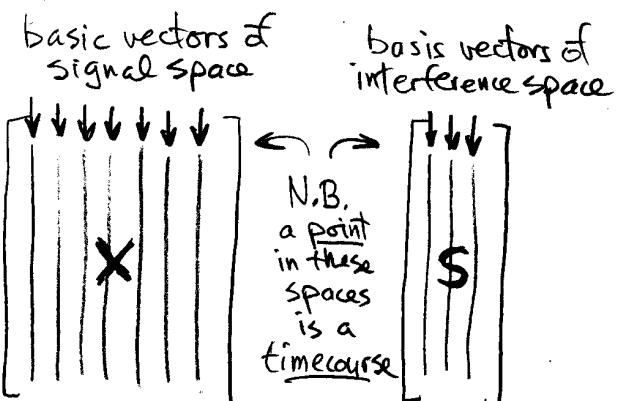
can concat  $X, S$

$$y = [X|S] [h|b] + n$$

simpler: preconvolve  
1) conv.  $X$  w/ fixed  $\vec{h}$   
2) solve for  $\beta$   
 $\vec{y} = \vec{X}\vec{\beta} + \vec{n}$   
 $y$  is scalar

	Cond1	Cond2	
Cond1	1	0	$[h_1]$ Cond1
Cond2	0	1	$[h_2]$ Cond2
Cond1	1	0	...
Cond2	0	1	
Cond1	0	0	
Cond2	1	0	
Cond1	0	0	
Cond2	1	0	

matrix notation for discrete convolution of stimulus pattern with hemodynamic resp. functs.



maximum likelihood estimate (Liu et al. 2001 Neuroimage)

[orthogonal cols' most efficient minimize trace  $[(X^T X)^{-1}]$  to get

1) assume white noise, solve for  $\vec{h}$

$$2) \hat{h} = (X^T P_s^\perp X)^{-1} X^T P_s^\perp y \quad \text{where } P_s^\perp = I - S(S^T S)^{-1} S^T$$

or

$$= (X_\perp^T X_\perp)^{-1} X_\perp^T y \quad \text{where } X_\perp = P_s^\perp X$$

→ projection matrix that removes part of vector that lies in  $S$  space

3) Significance (how to construct F-ratio)

→ design matrix w/ nuisance effects removed from col's

$$F = \frac{N-K-l}{K} \left[ \frac{y^T (P_{XS} - P_S) y}{y^T (I - P_{XS}) y} \right]$$

$P_{XS}$  - projects data on expt + nuisance subspace

$P_S$  - projects data onto nuisance subspace

→ see diagram next page for geometric interp

stats-4

## GEOMETRIC INTERPRETATION OF GENERAL LINEAR MODEL

- with no nuisance functions ( $S$ ), we could look at orthogonal projection of data onto experimental design and compare that to error to determine significance

$$\vec{y} = \vec{X}\vec{h} + \vec{\epsilon}$$

↑ data  
 ↑ expt HDr error  
 ↑ noise  
 ↓ signal

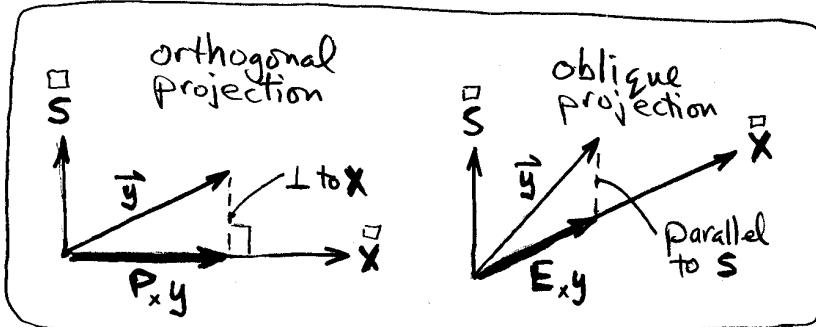
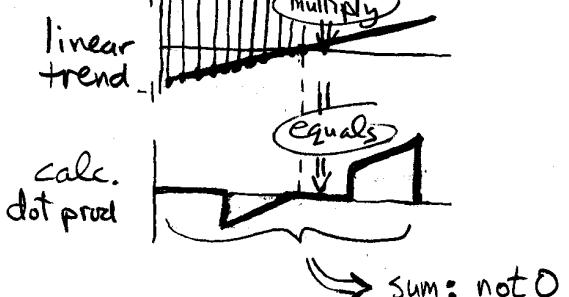
$$\vec{X}\vec{h} = \vec{P}_x \vec{y}$$

Projection matrix,  $\vec{P}_x$ , operates on  $\vec{y}$  to give projection of data into experiment space,  $\vec{X}$

- when nuisance functions,  $S$ , are considered, problem:  $S$  may not be orthogonal to  $X$

↳ for example: linear trend not orthogonal to std. block design

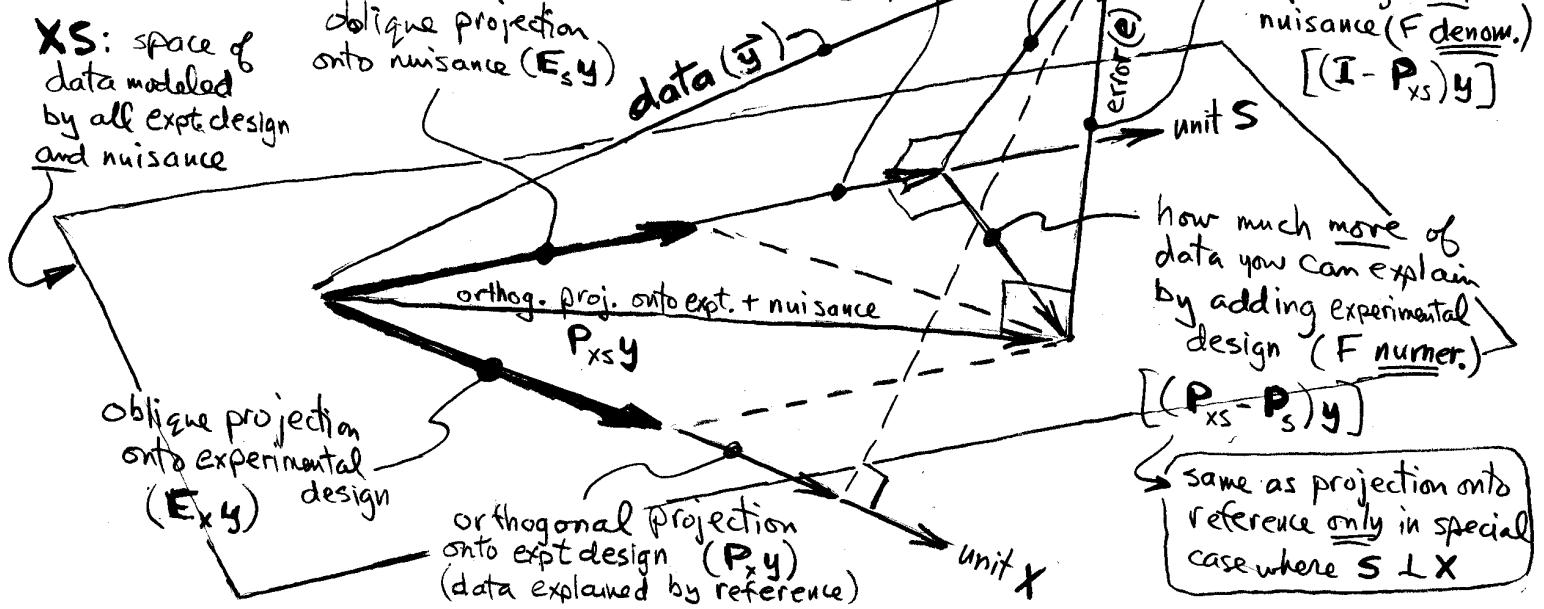
[remember: "orthogonal" means [dot prod. = 0 corr = 0]]



### Geometric Picture

(Liu et al., 2001, *Neuroimage*)

(orig from Schaff & Friedlander, 1994 IEEE)

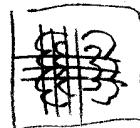


Surf-O

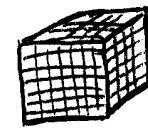
# WHY USE SURFACES?

- raw MRI data is a 2D flat slice or a 3D volume

( $\hookrightarrow I(x,y)$  or  $I(x,y,z)$ )



or



but...

- 1) the neocortex (and cerebellar cortex) are thin, folded 2D sheets

[cortex starts as smooth "balloon"  $\rightarrow$  ①  
major sulci, temporal lobe form  $\rightarrow$  ②  
great size increase, "crinkles" form  $\rightarrow$  ③]



- 2) neocortex contains many topological maps along its surface

[retinotopy  
tonotopy  
somatotopy  
musculotopy]

plus higher level maps  $\rightarrow \sim 2/3$  of its area

- 3) surface displays allow seeing (almost) all of data at once



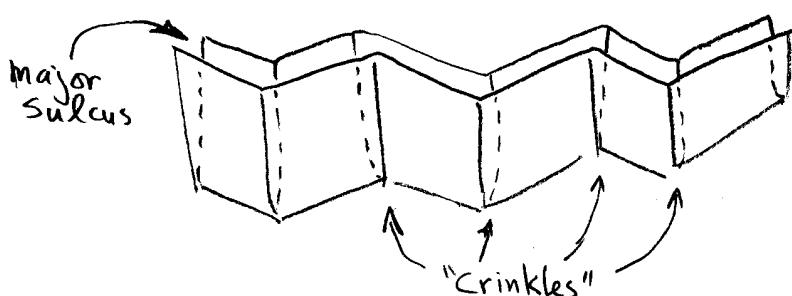
- 4) differences in major Sulci make 3D-based alignment difficult



e.g. STS, monkey-like IPS vs. postcentral plus extra

$\hookleftarrow$  extremely anisotropic def.

- 5) idiosyncratic sulcal crinkles



- these introduce additional noise into alignment in 3D  
- exact position of crinkles unlikely to have functional implications (tho 3D align might respect them)

# SEGMENTATION & SURFACE RECON

Talairach, Normalize, StripSkull

## 1) MNI auto-Talairach → generates $4 \times 4$ matrix

$$\begin{bmatrix} 3 \times 3 \\ (\text{rotate} \& \text{scale}) \end{bmatrix} \begin{bmatrix} x \\ y \\ z \\ 0 \ 0 \ 0 \ 1 \end{bmatrix}$$

translate

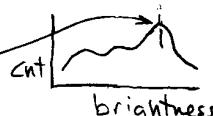
- make average brain target (blurry)
- blur target (further), blur single brain (a lot), gradient descent on  $x_{corr}$
- repeat w/ less blurring of avg target and current brain
- problems: variable neck cutoff  
 ↳ but much better than standard! only 2 points near center of brain!  
 ↳ fit to bounding box

## 2) Intensity Normalization (output: "T1")

- histogram of pixel values in 10 mm thick HOR slices



- smooth histogram



- peak find to get initial estimate of white matter



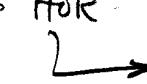
- discard outlier peaks across slices



- fit splines to peaks across slices

↳ interpolates scaling factor  $\perp$  to HOR

- scale each pixel so WM peak is 110



- refine estimate to interpolate in 3D

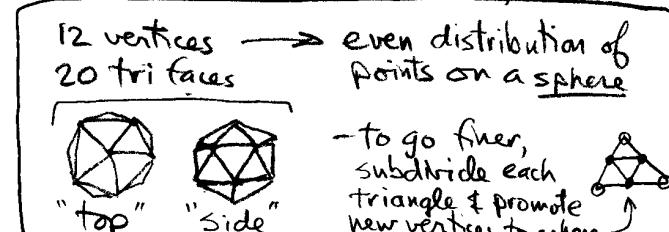


5-10 times

→ find points in  $5 \times 5 \times 5$  within 10% of WM, get new scale for them  
 build Voronoi to interpolate scales unset above  
 soap-bubble-smooth Voronoi boundaries (3 iterations)  
 re-scale each voxel

## 3) Skull Stripping (output: "brain")

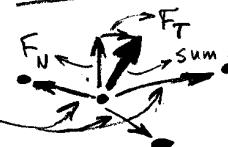
- "shrink-wrap" algorithm



- start with ellipsoidal template → sub-tesselated icosahedron

- minimize brain penetration and curvature

- curvature: spring force  
 (from center-to-neighbor vec sum)



- brain penetration

apply force along surface normal that prevents surface from entering gray matter

- decompose into  $\perp$  and tangential (local normal from summed normed cross products)

# SEGMENTATION & SURFACE RECON

Spring force in detail

- implementing a "force" is like directly constructing the operator that minimizes something (without first defining the "Something")
- more formally, we would define cost function, then take its derivative (gradient) to minimize it

Shrinkwrap update eq. (skull strip, original Dale & Sereno surface refinement)

$$\underline{r_{\text{center}}(t+1)} = \underline{r_{\text{center}}(t)} + \underline{\mathbf{F}_{\text{Smooth}}(t)} + \underline{\mathbf{F}_{\text{MRI}}(t)}$$

for one vertex      previous      local quantities

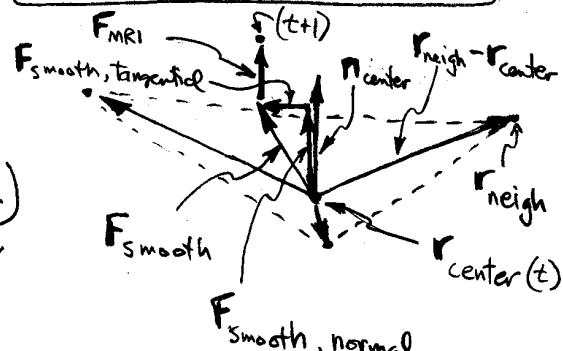
rule for each vertex,  $\vec{r}_{\text{center}}$

HOWTO

$$\mathbf{F}_{\text{smooth}} = \lambda_{\text{tang.}} \sum_{\text{neigh}} \left( \mathbf{I} - \mathbf{n}_{\text{center}} \mathbf{n}_{\text{center}}^T \right) \cdot (\mathbf{r}_{\text{neigh}} - \mathbf{r}_{\text{center}})$$

$\mathbf{n}^T = x^2 + y^2 + z^2$   
 $\mathbf{n} \mathbf{n}^T = \begin{bmatrix} x^2 & xy & xz \\ yx & y^2 & yz \\ zx & zy & z^2 \end{bmatrix}$   
 $(\mathbf{n} \mathbf{n}^T) \mathbf{v} = \mathbf{n}(\mathbf{n}^T \mathbf{v}) =$  dot prod.  
 $d\mathbf{n} = \text{"project } \mathbf{v} \text{ onto } \mathbf{n}"$  if  $\|\mathbf{n}\|=1$

identity  $3 \times 3$   
 stronger than normal ( $0.5$ )  
 expand by distribution to: neighbor vector minus projection of neighbor onto normal  $\rightarrow$  tangential!  
 project second onto first  
 vector to neighbor vertex



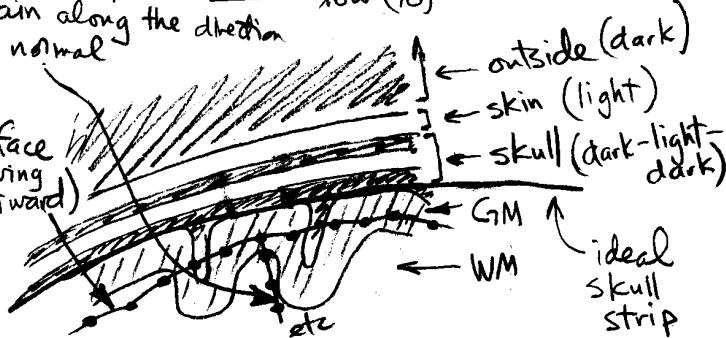
$$+ \lambda_{\text{normal}} \left[ \sum_{\text{neigh}} \left( \mathbf{n}_{\text{center}} \mathbf{n}_{\text{center}}^T \right) \cdot (\mathbf{r}_{\text{neigh}} - \mathbf{r}_{\text{center}}) - \frac{1}{\# \text{Vertices}} \sum_{\text{neigh}} \sum_{\text{v}} (\mathbf{n}_v \mathbf{n}_v^T) \cdot (\mathbf{r}_{\text{neigh}} - \mathbf{r}_v) \right]$$

weaker than tangential ( $0.1$ )  
 projection of a neighbor or vertex vector onto normal in the direction of the normal ( $\mathbf{n}$  is squared (as above) so we get a vector out (not a scalar))  
 average normal component  
 vector subtract off

$$\mathbf{F}_{\text{MRI}} = \lambda_{\text{MRI}} \mathbf{n}_{\text{center}} \max_{d=30} \left[ 0, \tanh \left[ I \left( \mathbf{r}_{\text{center}} - d \mathbf{n}_{\text{center}} \right) - I_{\text{thresh}} \right] \right]$$

intensity MRI data  
 multiply all terms  $\rightarrow$  don't allow neg product = 1.0  
 $\mathbf{F}_{\text{MRI}} = 0$  if any are zero

snapshot of Surface (moving outward) and "core sample" from one vertex



surf-3

## SEGMENTATION & SURFACE RECON

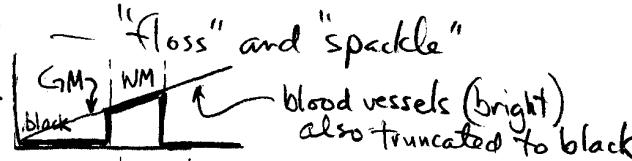
filter, cut, tessellate

### 4) Non-isotropic filtering (output: "win")

- preliminary hard threshold: output
- find ambiguous/boundary voxels

↳ 20% or more of 26 immediate neighbors different

- find plane of least variance



↳ to avoid expensive calc below...

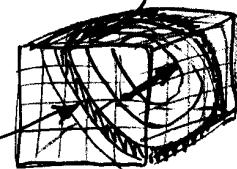
for each ambiguous voxel

[for each direction (from icosahedral supertessellation)]

consider 5x5x5 volume around 1 voxel

find plane of least variance in this hemisphere

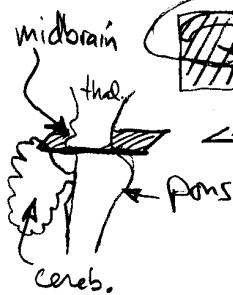
median filter w/ hysteresis (1 vox thick)



↳ if 60% of within-slab differ, reverse classification

↳ "flosses" sulci without blurring

### 5) Find cutting planes



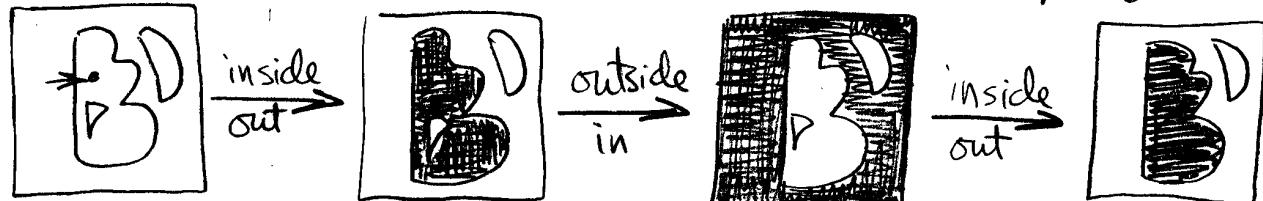
midbrain  
the  
callosum, to separate hemispheres (SAG)  
midbrain, to avoid fill into cerebellum (HOR)

Talairach to start;  
fill WM in SAG  
or HOR till min area

### 6) Region-growing to define connected parts (output: "filled")

- inside-out, outside-in, inside-out — for each hemisphere

- up/down cycles within each plane  
- plane-by-plane



- "wormhole filter" ( $3 \times 3 \times 3 = \text{center} + 26$ )

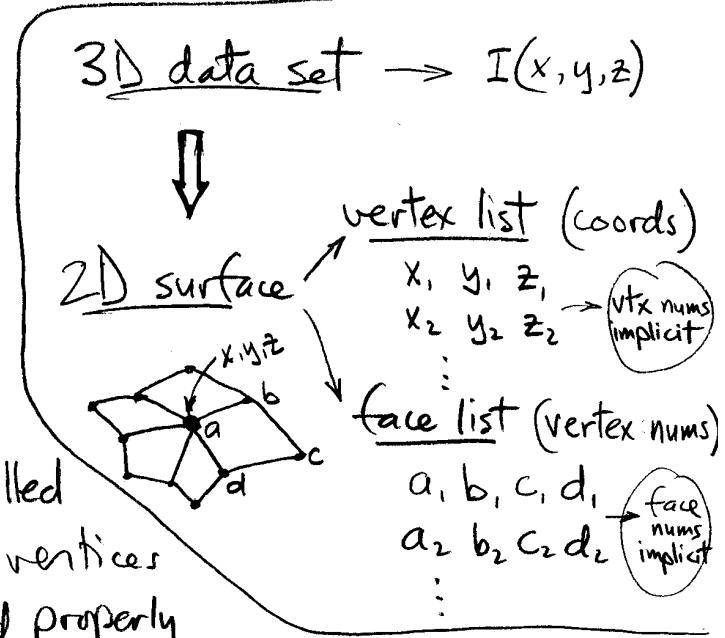
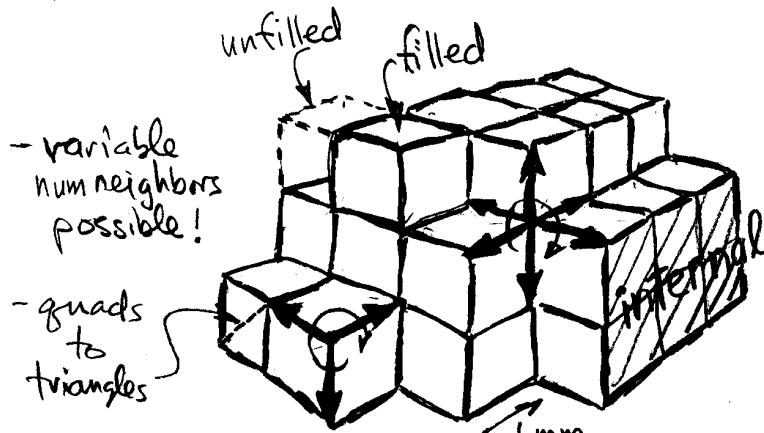
↳ fill (unfilled) voxel if 66% neighbors differ → eliminates structures w/thin, 1-D structure

surf-4

# SEGMENTATION & SURFACE RECON

3D → 2D

## 7) Surface Tessellation (output: rh.orig, lh.orig)



- find filled voxels bordering unfilled
- make ordered list of neighbouring vertices  
↳ so cross-products oriented properly
- long list of values associated with each numbered vertex

e.g.

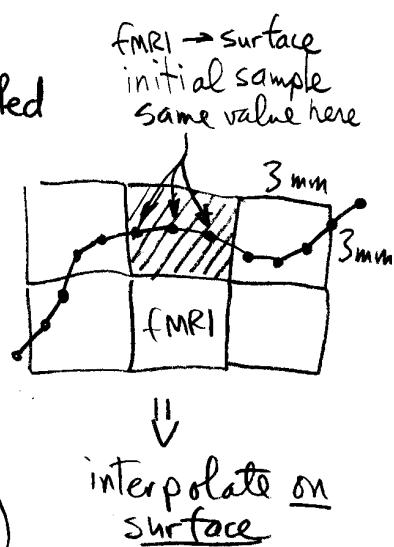
position (orig, morphed)
area (orig, morphed)
curvature (intrinsic, Gaussian)
"Sulcussness" (summed L movement during unfolding)
cortical thickness
fMRI data
EEG/MEG dipole strength

- separate fMRI data set must be aligned, sampled

fMRI voxels larger  
Sample at each surface vertex  
nearest-neighbor "soap bubble" smoothing  
to interpolate data onto hi-res mesh

- some quantities only well-defined on surface

↳ gradient of magnitude of cortical map measure (e.g., eccentricity)



## SEGMENTATION & SURFACE RECON

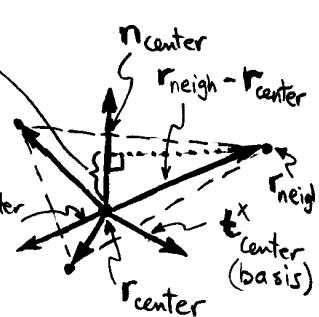
Smooth, inflate, final surfaces

- smoothing/inflation/WM, pial done as derivative of energy functional

$$J = J_{\text{tangential}} + \lambda_{\text{normal}} J_{\text{Normal}} + \lambda_{\text{image}} J_{\text{Image}}$$

total scalar error to minimize  
 scalar tangential error (fixed by redistributing vertices)  
 small (0.25) scalar curvature error (fixed by reducing curvature)  
 smaller (0.075) scalar image error (fixed by moving toward target image value)

$J_{\text{normal}} = \frac{1}{2 \# \text{vert}} \sum_{\text{centers}} \sum_{\text{neighbors}} [n_{\text{center}} \cdot (r_{\text{neigh}} - r_{\text{center}})]^2$



across all vertices, curvature error  
 1/2 so no coefficient on derivative  
 across all vertices neighbors of one vertex  
 vertex unit normal project onto normal  
 vector from current center to one neighbor (position vector diff.)

$J_{\text{tangential}} = \frac{1}{2 \# \text{vert}} \sum_{\text{centers}} \sum_{\text{neighbors}} [t_x^{\text{center}} \cdot (r_{\text{neigh}} - r_{\text{center}})]^2 + [t_y^{\text{center}} \cdot (r_{\text{neigh}} - r_{\text{center}})]^2$

"squishing" of mesh  
 centers neighbors  
 x-direction in tangent plane project vector to neighbor onto x & y  
 y-direction in tangent plane

$J_{\text{image}} = \frac{1}{2 \# \text{vert}} \sum_{\text{centers}} [I_{\text{center}}^{\text{targ}} - I(r_{\text{center}})]^2$

image error  
 target brightness at current location

$t_x, t_y$  are first 2 eigenvect. of neighbor vector cloud ( $n$  is third)

$I^{\text{targ}}$  for WM: mean of voxels labeled WM in 5 mm neighborhood  
 $I^{\text{targ}}$  for pia: global - small num for C.S.F.-like

- take directional derivative of energy functional (to find steepest uphill)
- move each vertex in the opposite (negative) direction w/ self-intersect test

N.B.: eq. 9 in Dale, Fischl & Sereno different - and incorrect!

$\frac{\partial J}{\partial r_{\text{center}}} = \lambda_{\text{image}} [I_{\text{center}}^{\text{targ}} - I(r_{\text{center}})] (-\nabla I(r_{\text{center}})) + \sum_{\text{neighbors}} \lambda_{\text{normal}} [n_{\text{center}} \cdot (r_{\text{neigh}} - r_{\text{center}})] (-n_{\text{center}})$

go the opposite direction (vector) of largest scalar error for one vertex movement  
 direction (vector) of largest scalar error for one vertex movement  
 N.B.: cancels neg. on LHS so  $\lambda$  says which way to go  
 scalar

HOWTO derivative:  
 constants  
 $\frac{\partial J}{\partial r} = \partial(C_1 \cdot (C_2 - V))$   
 var  
 $= \partial(C_1 C_2 - C_1 V)$   
 $= -C_1$

vector - calculate gradient on image (first blur w/ Gaussian)  
 scaled by unit normal vector

x-component of tangential  
 $+ \sum_{\text{neighbors}} [t_x^{\text{center}} \cdot (r_{\text{neigh}} - r_{\text{center}})] (-t_x^{\text{center}}) + [t_y^{\text{center}} \cdot (r_{\text{neigh}} - r_{\text{center}})] (-t_y^{\text{center}})$   
 y-component

surf-6

## SULCUS-BASED CROSS-SUB. ALIGN

- use summed perpendic. vertex move during inflation as vtx measure of "sulcus-ness"
- add term to error function,  $J$ : "sulcus-ness" error

$$J_{\text{sulc}} = \frac{1}{2 \# \text{vert}} \sum_{\text{centers}} [S_{\text{subj}}^{\text{subj}} - S_{\text{targ}}^{\text{targ}}]_{\text{(center)}}^2$$

find neg of steepest uphill direction  
of change in sulcus-ness of target

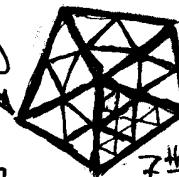
take deriv.

$$\frac{\partial J_{\text{sulc}}}{\partial r_{\text{cent}}} = \lambda_{\text{sulc}} [S_{\text{subj}}^{\text{subj}} - S_{\text{targ}}^{\text{targ}}]_{\text{(center)}} (-\nabla S_{\text{targ}})_{\text{(center)}}$$

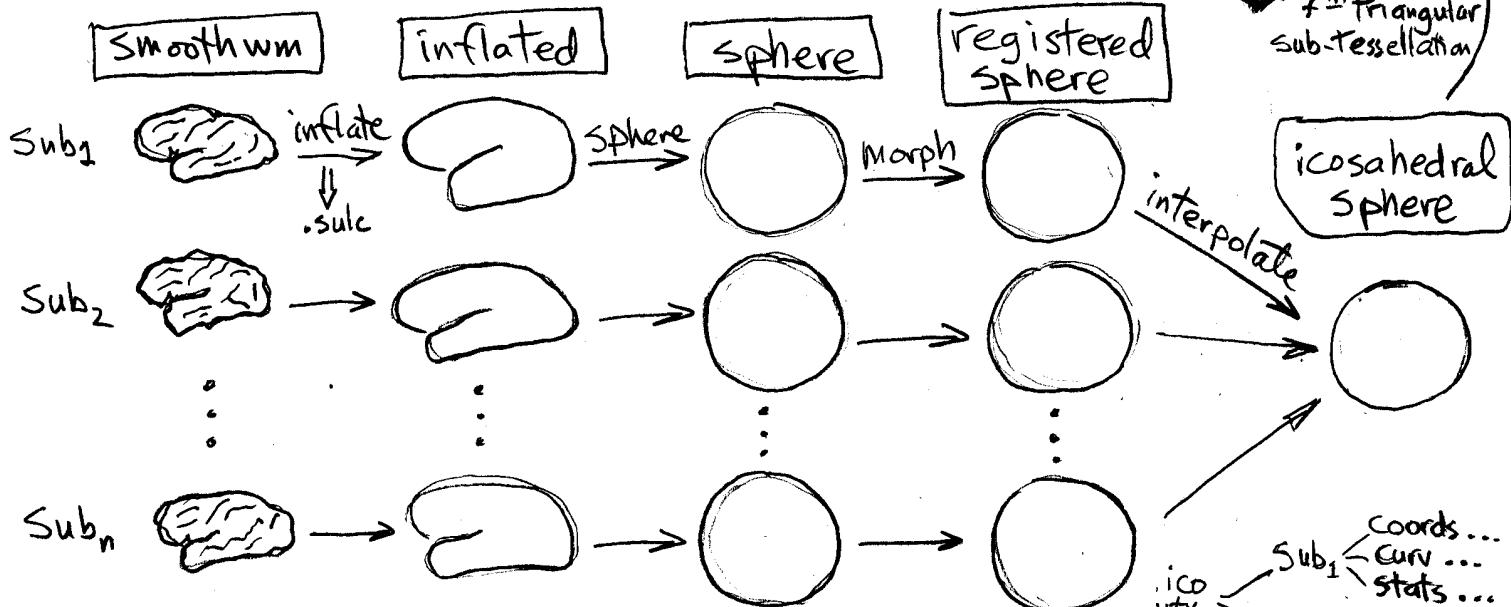
- bootstrap

grad descent 2D  
morph to one brain  
make avg targ  
re-morph to avg targ

icosahedron  
(5-fold symmetry)



7th triangular  
sub-tessellation



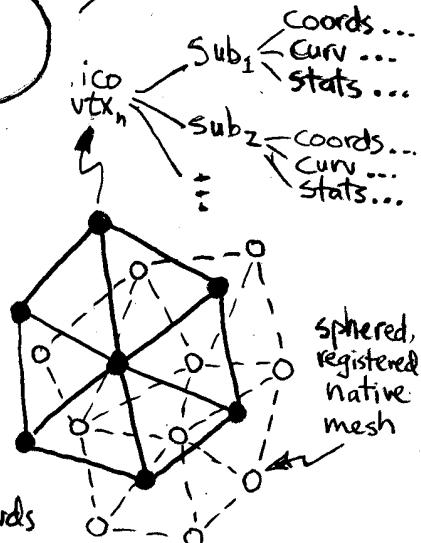
- each sub's native surf has diff # vertices

- interpolate values (coords, surface measures, stats) to each icosahedral vertex from neighboring vertices of native mesh (dashed lines)

- average surface made from folded/inflated avg coords  
 folded: loses area from sulcal crinkles ("average" "inflated")  
 inflated: retains orig area, correct sulc/gyrus ratio ("inflated-avg")

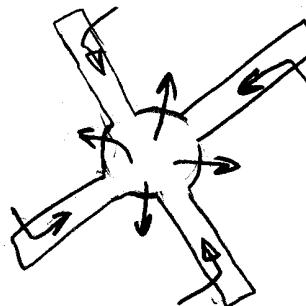
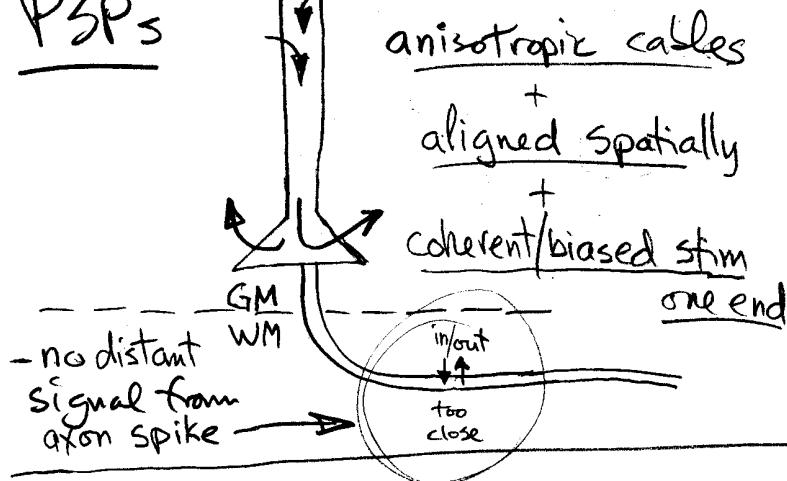
- can use sampled-to-icos individual subj coords to draw icosahedral surface in shape of an indiv. brain

→ N.B.: morph will have changed local vertex density compared to more uniform native mesh (use native for sing. sub.)



source-1

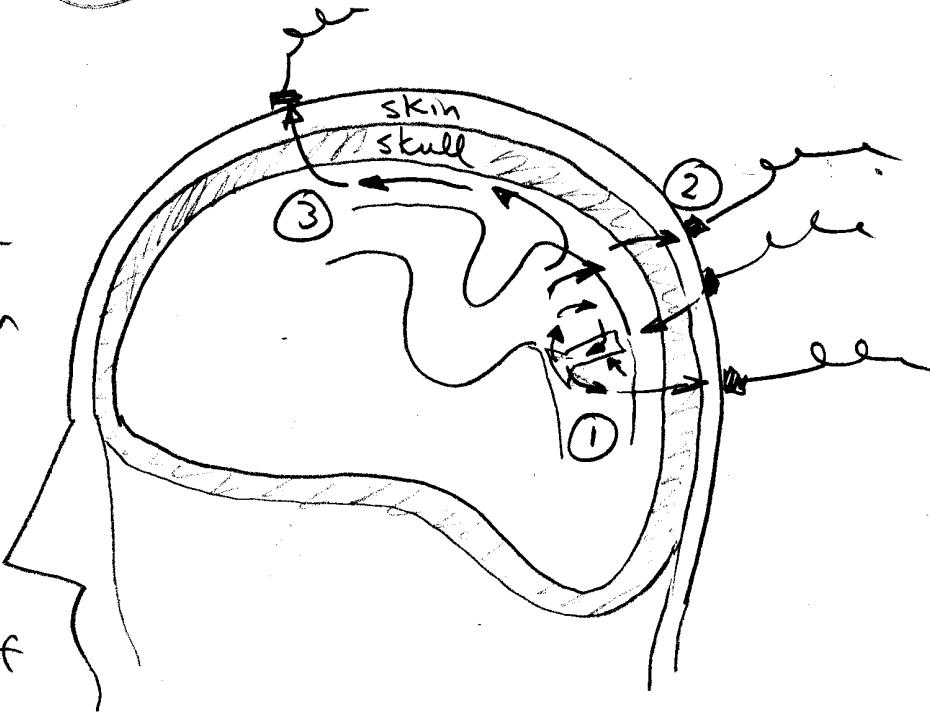
# SOURCE OF EEG / MEG

PSPsisotropic

"closed field"  
(invisible at distance)

Head

- 1) - local dipole
- 2) - EEGs through skull, skin
- 3) - smearing because skull  $\frac{1}{100}$  conductivity of brain

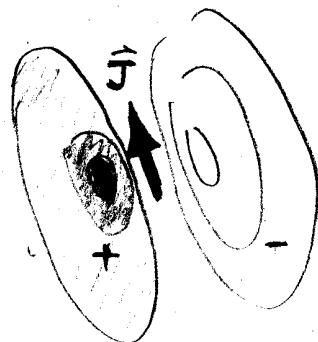
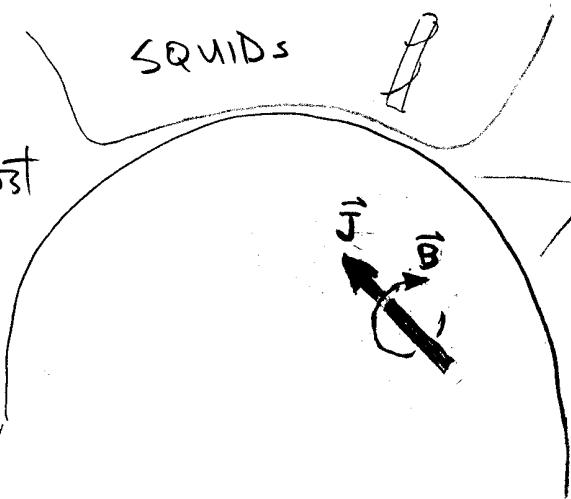


the effect on current flow is like a higher order (larger) "cable"

MEG

- radial dipoles lost
- tangential dipole generates Gabor-like scalp distrib. of  $\vec{B}$  field

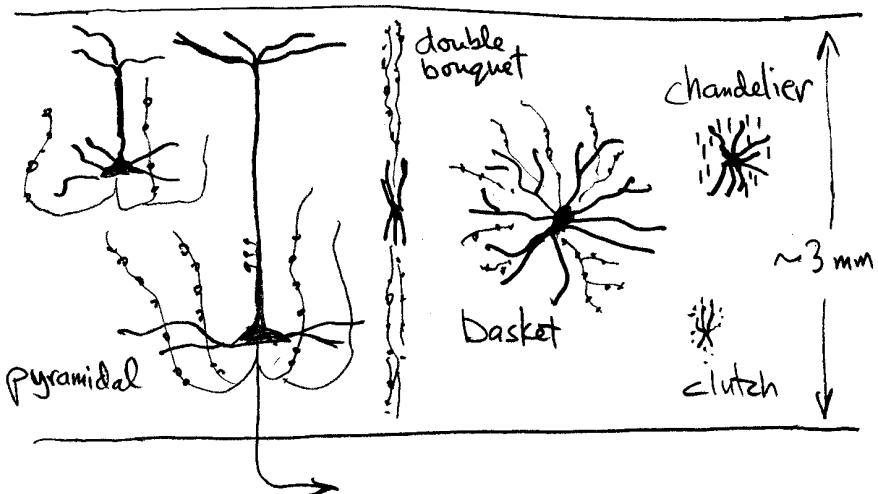
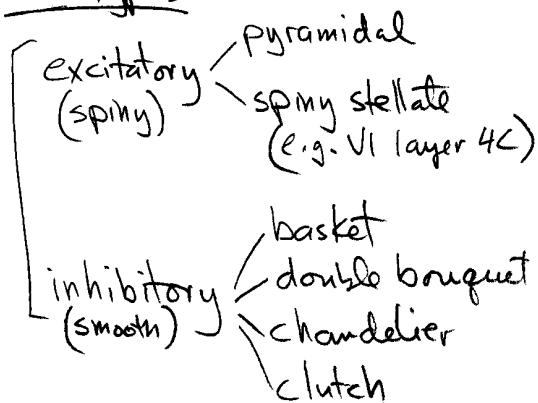
SQUIDS



Source-2

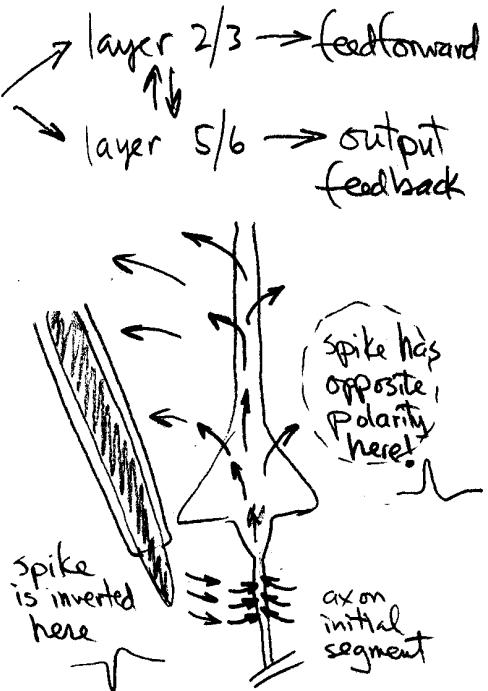
# INTRACORTICAL CIRCUITS & ORIGIN OF EEG

## Cell types

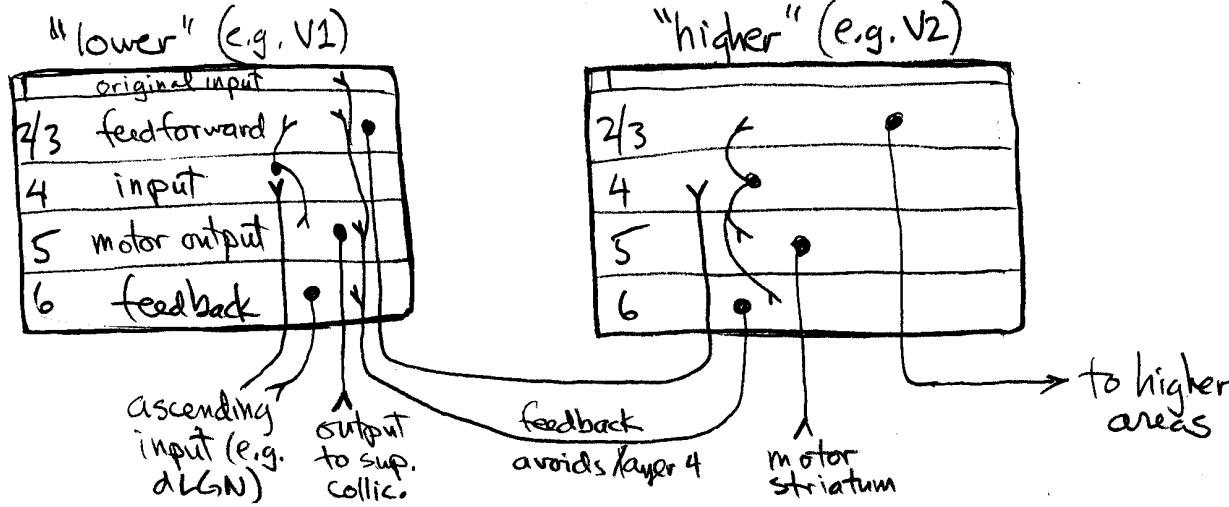


## Circuits

- huge complexity
- first principal components: input  $\rightarrow$  layer 4  $\rightarrow$  layer 2/3  $\rightarrow$  feedforward
- micro-electrode recording (e.g. 10  $\mu\text{m}$  tip)
  - [high pass  $\rightarrow$  spikes]
  - [low pass  $\rightarrow$  local field potentials]
- spikes only recordable in gray matter
- white matter spikes only recordable with pipette w/ very fine tip b/c inward & outward currents so spatially close in axon/spike ( $> 1 \mu\text{m}$ )



## Intra/inter cortical connections cartoon



graddivcurl-1

# GRADIENT, DIVERGENCE, CURL

gradient ( $\nabla$ ) (generalized deriv.)

$$\nabla s(\vec{r}) = \frac{\partial s(\vec{r})}{\partial x} \hat{i} + \frac{\partial s(\vec{r})}{\partial y} \hat{j} + \frac{\partial s(\vec{r})}{\partial z} \hat{k}$$

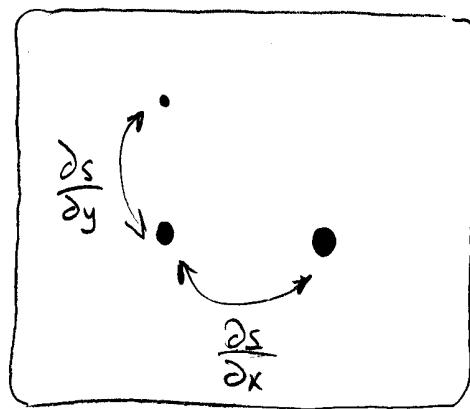
$x, y, z$

turns scalar field into vector field

scalar funct defined at each  $x, y, z$  point,  $\vec{r}$

change of  $s$  in  $x$  direction at point  $\vec{r}$

unit vector in  $x$ -dir



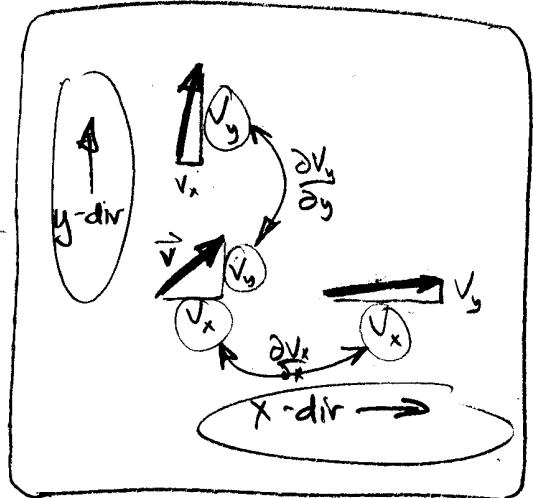
divergence ( $\nabla \cdot \vec{v}$ ) (deriv. "dot prod")

$$\nabla \cdot \vec{v}(\vec{r}) = \frac{\partial v_x(\vec{r})}{\partial x} + \frac{\partial v_y(\vec{r})}{\partial y} + \frac{\partial v_z(\vec{r})}{\partial z}$$

turns vector field into scalar field

vector funct defined at each  $x, y, z$  point,  $\vec{r}$

change of just  $x$ -component of  $\vec{v}$  in  $x$ -direction at point  $\vec{r}$



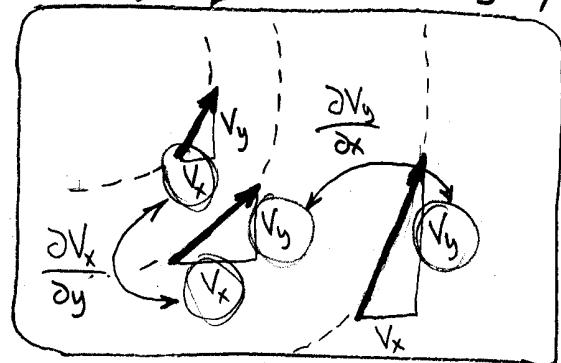
curl ( $\nabla \times \vec{v}$ ) (deriv. "cross product")

$$\nabla \times \vec{v}(\vec{r}) = \left( \frac{\partial v_z(\vec{r})}{\partial y} - \frac{\partial v_y(\vec{r})}{\partial z} \right) \hat{i} + \left( \frac{\partial v_x(\vec{r})}{\partial z} - \frac{\partial v_z(\vec{r})}{\partial x} \right) \hat{j} + \left( \frac{\partial v_y(\vec{r})}{\partial x} - \frac{\partial v_x(\vec{r})}{\partial y} \right) \hat{k}$$

turns vector field into another vector field

vector funct defined at each  $x, y, z$  point,  $\vec{r}$

change of just  $z$  component of  $\vec{v}$  in  $y$ -direction at point  $\vec{r}$



## Vector identities

$$\nabla \times \nabla s = 0$$

curl of the gradient of any scalar field is zero

$$\nabla \cdot (\nabla \times \vec{A}) = 0$$

divergence of the curl of any vector field is zero

$$\vec{A} \cdot (\vec{B} \times \vec{C}) = (\vec{A} \times \vec{B}) \cdot \vec{C}$$

N.B.  $\hat{k}$   $\perp$  to this plane

# POTENTIAL ( $\Phi$ ), ELECTRIC FIELD ( $\nabla \Phi$ ) & C.S.D. ( $\nabla \cdot (-\nabla \Phi) = \nabla^2 \Phi$ )

## low-frequency field approximation

- electric fields uncoupled from magnetic (vs. electromagnetic radiation)
  - ↳ pre-Maxwellian approx. (EEG freq's  $\ll 1 \text{ MHz}$ )
- calculate electric fields as if magnetic fields don't exist
- calculate magnetic fields strictly from distribution of currents
- ignore capacitative effects, too

## scalar potential, $\Phi$ (what we measure with electrode)

$$\vec{E} = -\nabla \Phi - \frac{\partial \vec{A}}{\partial t} \approx -\left[ \frac{\partial \Phi}{\partial x} \vec{i} + \frac{\partial \Phi}{\partial y} \vec{j} + \frac{\partial \Phi}{\partial z} \vec{k} \right] + 0$$

gradient of scalar field

ignore coupling:  $\vec{B} = \nabla \times \vec{A}$  vector potential  $\Rightarrow$  because EEG  $\ll 1 \text{ MHz}$

scalar potential at one position

electric field vector

turns scalar field ( $\Phi$ ) into vector field ( $\vec{E}$ )

↳ (1)  $\vec{E}$  defined as force (vector) acting on unit charge at a given point in space (as result of arbitrary distribution of other charges)

CSD is Laplacian of  $\Phi$  ( $= \operatorname{div} \vec{E}$ )

divergence

(2) Current density,  $\vec{J}$  (not curr. source dens.!) is proportional to  $\vec{E}$   $\rightarrow$  still a vector!

$$\vec{J} = \sigma \vec{E} \quad (\text{Ohm's law}) \quad \sigma \text{ is conductivity}$$

(3) Two def's of  $\vec{A}$ :

$$\vec{B} = \nabla \times \vec{A}$$

$$\vec{A} = \frac{\mu_0}{4\pi} \int_{\text{vol}} \frac{\vec{J}}{\|\vec{r}\|} d\text{vol}$$

dist-weighted sum of directional currents

$$\nabla \cdot (-\nabla \Phi) = \text{scalar field} = -\left[ \frac{\partial^2 \Phi}{\partial x^2} + \frac{\partial^2 \Phi}{\partial y^2} + \frac{\partial^2 \Phi}{\partial z^2} \right] \equiv -\nabla^2 \Phi$$

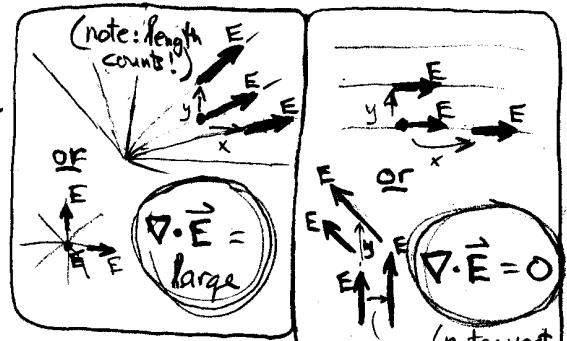
measures rate of change of vector field across space

makes scalar!  $\nabla$  turns vector field into scalar field

meaning of  $\nabla \cdot$  ↑↓

find radiating and contracting regions of  $\vec{E}$  vector field

sum of:  
change in x comp. in x direction  
+ change in y comp. in y dir.  
⋮



## 3D CSD gold standard (rat SPAER paper)

$\Phi$  data (360 points)

$\nabla \rightarrow \nabla \Phi$   
electric vector field

$\rightarrow \nabla \cdot (-\nabla \Phi)$

scalar field source/sink movie as function of  $t$

IC

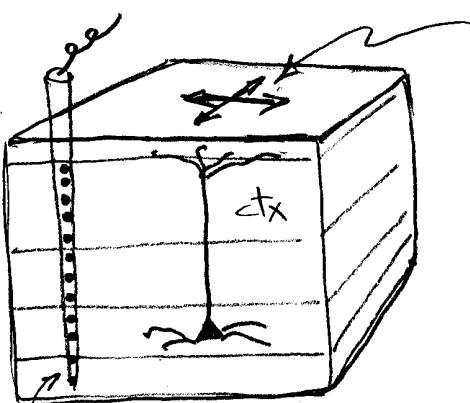
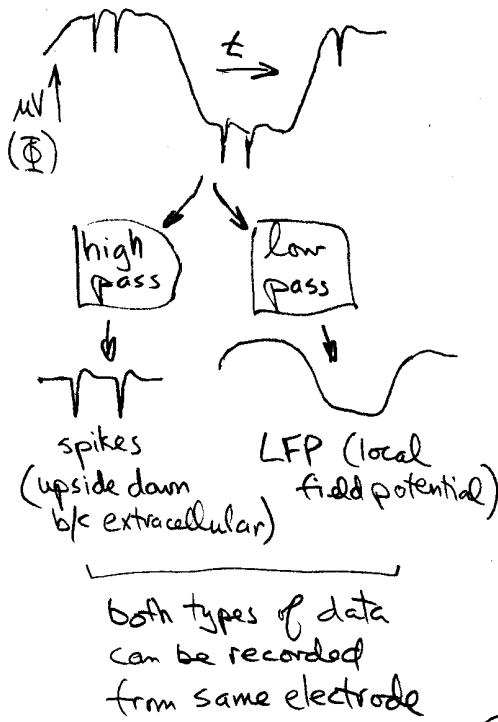
NLL

CSD-2

# 1D AND 2D CURRENT SOURCE DENSITY EXPTS.

## 1D CSD

- raw, event-related  
Signal relative  
to ground,  $\frac{dV}{dt}$  (e.g. skull)

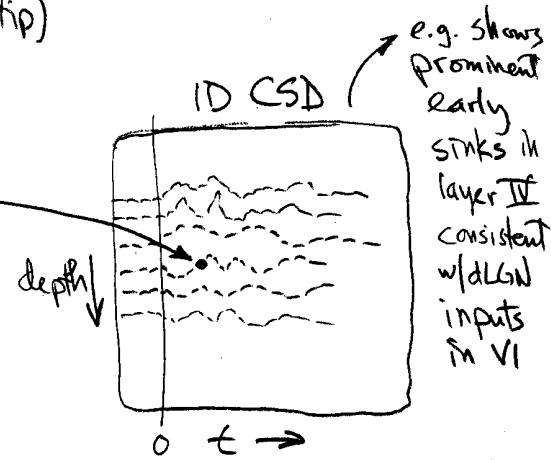
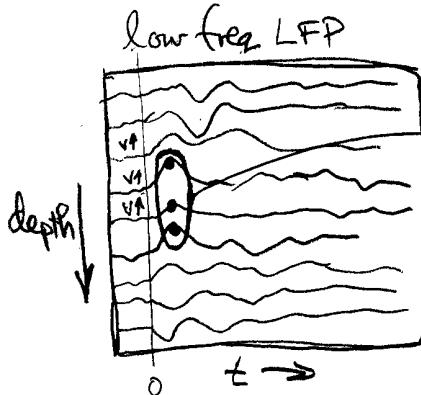


ignore these dimensions parallel to cortical sheet

Rationale: CSD changes much more slowly parallel to cortex than perpendicular to cortical sheet

→ assume approx constant ( $\approx 0$ ) parallel to ctx

recording sites (e.g., by slowly withdrawing electrode tip)

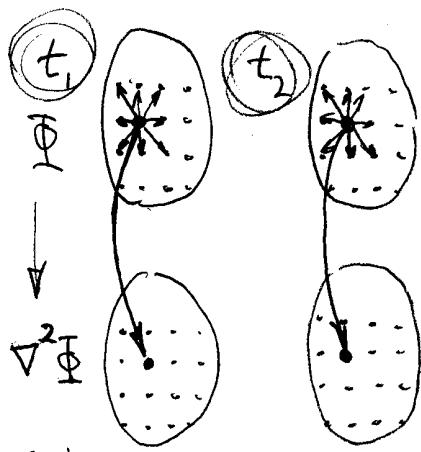


## 2D CSD

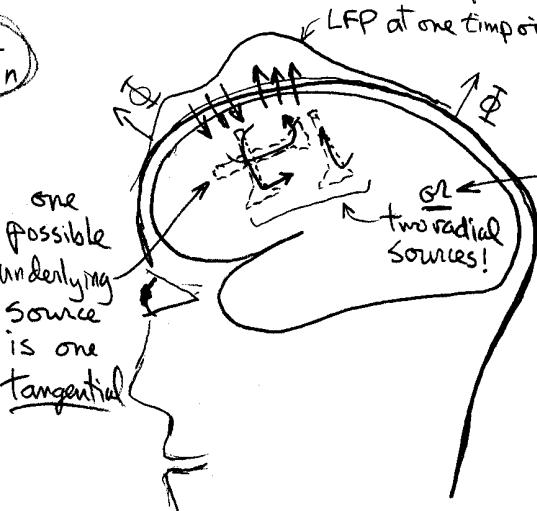
- 2D array of electrodes on pial surface or on scalp

- $\nabla^2$  means find spatial (i.e., 1D depth) curvature of potential
- discrete approx: center -  $\frac{\text{above} + \text{below}}{2}$
- N.B. in example above, even though all 3 potentials are positive, smaller value of center point implies sink!

Rationale: all electrodes record along same surface so assume depth profiles are constant

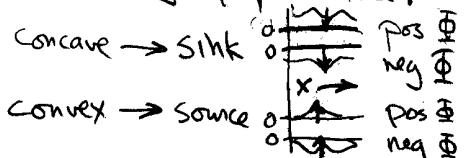


one possible underlying source is one tangential



- for scalp recordings, sources and sinks are at the scalp (not a depth loc method unless done in 3D)

- can't tell curvature from sign of potential!

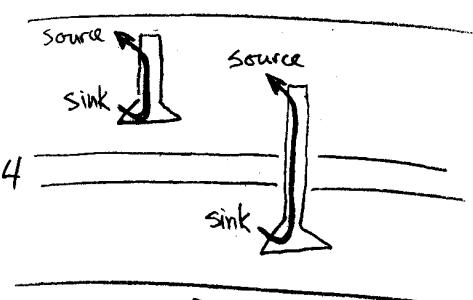
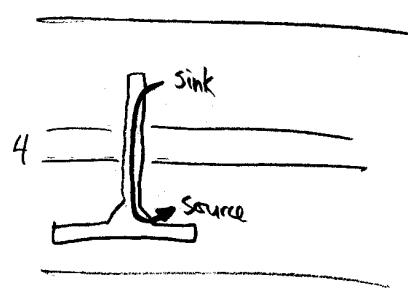
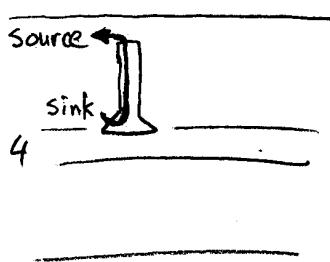
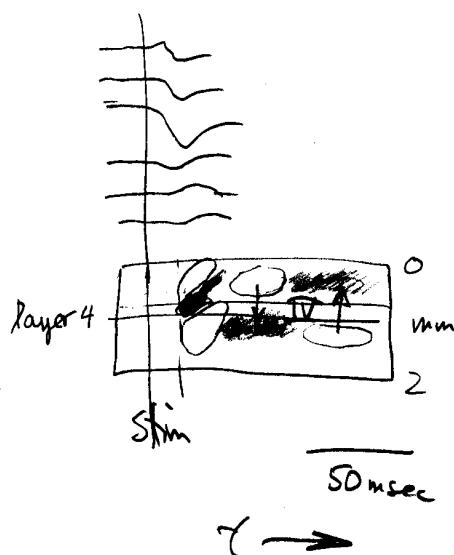


CSD-3

# INTRACORTICAL C.S.D.

- e.g. click evoked rat A-I  
(Sukov & Barth, 1998)

CSD-1D



P1

N1

P2

- phase-locked CSD of gamma shifts w/ each cycle



Surface electrode is typical ref

forward +

# MAXWELL EQUATIONS

Electrostatics, Magnetostatics  
low freq limit

Ohm's law:  $\sigma \vec{E}_v = \vec{J}$

$$\nabla \cdot (\sigma \nabla \Phi) = \nabla \cdot \vec{J}^i$$

divergence  
conductivity  
constant (or  
tensor constant  
if inhomogeneous  
in different directions)

gradient  
of scalar  
potential  
(what we  
measure  
invasively)  
at each point

like CSD but times  
 $\sigma$ , conductivity  
(cf.  $gV = I$ ,  $V = IR$ )

N.B. these are all  
defined at a (every)  
point in space

impressed currents

currents due to ionic flow  
that "appear out of nowhere"  
(Nernst batteries)

$$\nabla \cdot \vec{B} = 0$$

divergence  
magnetic  
field vector  
at each point

[incidentally, this  
Maxell equation  
violated by a  
linear gradient  
in  $B_z$  in x,  
y or z dir]

$$\nabla \times \vec{B} = \mu_0 (\vec{J}^i - \sigma \nabla \Phi)$$

curl  
magnetic  
field

permeability

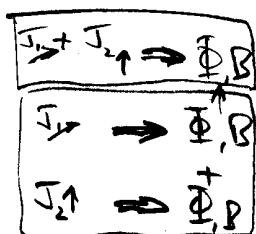
impressed  
currents

conducting

gradient of  
scalar potential  
 $(= \vec{E})$

- propagation of potentials, magnetic fields instantaneous (no capacitance)
- Simultaneous eqs to solve:  $\vec{J}$  are sources,  $\Phi, \vec{B}$  are data
- linear

Potential ( $\Phi$ ) and magnetic fields ( $\vec{B}$ ) produced by a weighted sum of two current source distributions are equal to weighted sum of fields produced by each current source distribution by itself



forward-2

# WHY WE CAN IGNORE MAGNETIC INDUCTION

$$\vec{E} = -\nabla \Phi - \frac{\partial \vec{A}}{\partial t}$$

(from Nunez, 1981)

electric field

field component  
due to charge  
distributionfield component due  
to coupling between electric  
& magnetic fields

$$\vec{B} \equiv \nabla \times \vec{A}$$

"vector potential"

$$\nabla \times \vec{H} = \vec{J} + \frac{\partial \vec{D}}{\partial t}$$

magnetic field  
(in given medium)  
induced

currents  
(in given medium)

time varying  
electric fields  
(in given medium)

this induced magnetic field  
induces an electric  
field in turn

$$\begin{aligned}\vec{H} &= \frac{1}{\mu} \vec{B} \\ \vec{J} &= \sigma \vec{E} \\ \vec{D} &= \epsilon \vec{E}\end{aligned}$$

permeability  $\mu$   
conductivity  $\sigma$   
permittivity  $\epsilon$ characterize  
substance  
linear  
in all  
three

$$\nabla \times \vec{E} = -\frac{\partial \vec{B}}{\partial t}$$

take  $\nabla \times$  of both sides  
use  $\vec{B} = \mu \vec{H}$   
substitute this into this

$$\nabla \times \nabla \times \vec{E} = -\mu \frac{\partial}{\partial t} \left( \vec{J} + \frac{\partial \vec{D}}{\partial t} \right)$$

if linear in conductivity and dielectric, too, and fields periodic w/f

$$\nabla \times \nabla \times \vec{E} = -2\pi f i \mu (\sigma + 2\pi f \epsilon) \vec{E}$$

to neglect:  $\frac{2\pi f i \mu (\sigma + 2\pi f \epsilon) |\vec{E}|}{|\nabla \times \nabla \times \vec{E}|} \ll 1$

- 1)  $|\nabla \times \nabla \times \vec{E}| \approx |\vec{E}|/L^2$  where  $L$  is dist over which  $\vec{E}$  varies significantly
- 2)  $\mu$  of tissue similar to empty space
- 3) assume conservative (large)  $\sigma$ , dielectric const., and EEG freq

↪ number is about  $10^{-6} \rightarrow$  small

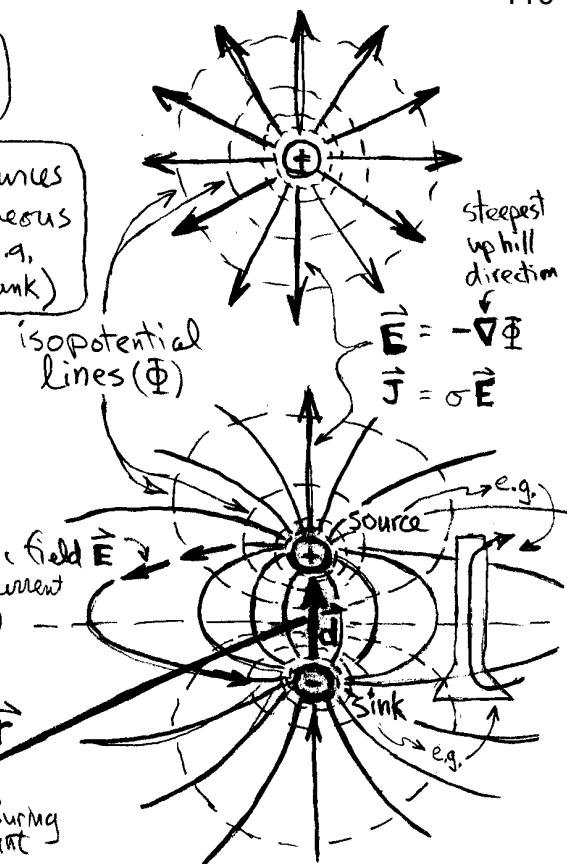
forward-3

# MONPOLE, DIPOLE FORWARD SOL'N

$$\underline{\Phi}_1 = \frac{s}{4\pi\sigma r}$$

potential recorded for source monopole  
 strength of source  
 distance, source to measuring point ( $= \|\vec{r}\|$ )

current sources in homogeneous medium (e.g., saltwater tank)



$$\underline{\Phi}_2 = \frac{s}{4\pi\sigma} \left( \frac{1}{r_1} - \frac{1}{r_2} \right)$$

potential recorded for source-sink pair ("near field")  
 dipole source to measuring distance  
 dipole sink to measuring distance

$$\text{scalar } \underline{\Phi}_2 \approx \left( \frac{1}{4\pi\sigma} \right) \frac{\vec{s} \cdot \vec{d} \cdot \vec{r}}{r^3}$$

scalar strength  
 dipole vector, Sink to Source  
 N.B.: reverse of current flow +-

now: also explicitly include dipole orientation and measuring point in equations

approximations for "far enough away" measurements (subtracting two 1/r's gives inverse square)

$$\text{vector } \underline{\vec{B}}_2 \approx \left( \frac{\mu_0}{4\pi} \right) \frac{\vec{s} \cdot \vec{d} \times \vec{r}}{r^3}$$

N.B.: both assume inside infinite isotropic conductor

Since  $\vec{r}$  also in numerator, this now inverse square

$$\underline{\Phi}_i(t) = e_i \underline{s}(t)$$

electrode  
 gain  
 one source strength

$$\vec{b}_i(z) = \vec{m}_i s(t)$$

$$\vec{b}_i(z) = m_i s(t) \rightarrow \text{Squids measure component of } \vec{B}$$

$$\underline{\Phi}_i(t) = \sum_j e_{ij} \underline{s}_j(t)$$

all sources

$$\vec{b}_i(z) = \sum_j m_{ij} s_j(t)$$

linear superposition with fixed electrodes and sensors

$$\underline{\vec{x}}(t) = \sum_j \underline{g}_j \underline{s}_j(t)$$

elec & mag. measures  
 gain vect  
 strength

$$\underline{\vec{x}}(t) = \underline{G} \underline{s}(t)$$

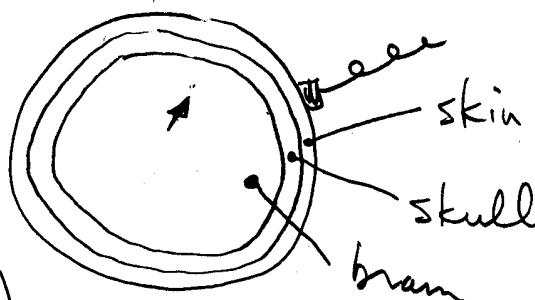
# forward-4 Forward Solution (( ))

- well-posed (one answer)
- linear:  $f(A) + f(B) = f(A+B)$
- approximations due to unknown electrical properties of head

## 3-shell spherical analytic

- skull  $\neq$  brain  
conductivity  
of brain

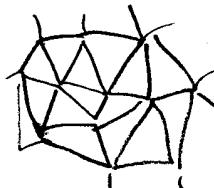
- "smearing"  
(cf. cable theory)



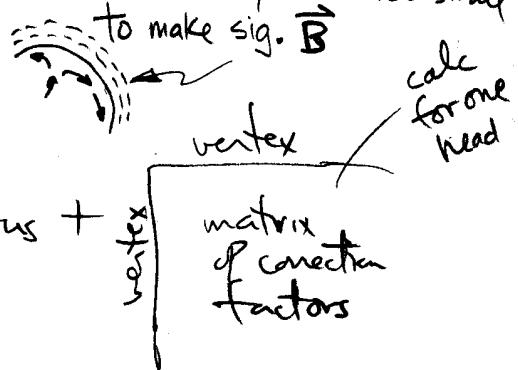
- remember, we only need to be able to calc. weight for each dipole/electrode pair independently

## 3-shell boundary element

[arbitrary shape  
homogeneous  
conductivity]

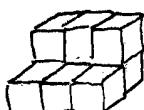


- for magnetic, only need one shell b/c currents thru skin/skull too small to make sig.  $\vec{B}$



$$\text{solution} = \text{infinite homogeneous} + \left[ \begin{array}{c} x \\ y \\ z \end{array} \right]$$

## finite element



[most general  
computational intensive w/ small grid  
many unknown parameters to estimate]

forward-S

# FORWARD SOLN (2)

recording  
sensor

$$v_i = \sum_j e_{ij} s_j + n_i$$

sources  
forward  
source strength at:  
position ( $x, y, z$ )  
angle ( $\theta, \phi$ )  
in brain

noise

matrix form

usu.  $\gg$  than sensors

$$\begin{bmatrix} v \\ \vdots \end{bmatrix} = \begin{bmatrix} \text{sensors} \\ \vdots \end{bmatrix} \begin{bmatrix} \text{sources} \\ \vdots \end{bmatrix} E \begin{bmatrix} \text{sensors} \\ \vdots \end{bmatrix} \begin{bmatrix} s \\ \vdots \end{bmatrix} + \begin{bmatrix} n \\ \vdots \end{bmatrix}$$

weights for one source

weights for one sensor

$$v = Es + n$$

$E$  fixed across time  
 $v, s, n$  vary

lower case bold  $\rightarrow$  vector  
upper case bold  $\rightarrow$  matrix

electric recordings  $\rightarrow$   $v$

magnetic recordings  $\rightarrow$   $m$

$$= \begin{bmatrix} \text{electric forward solution} \\ \hline \text{magnetic forward solution} \end{bmatrix} B \begin{bmatrix} s \\ \vdots \end{bmatrix} + \begin{bmatrix} n \\ \vdots \end{bmatrix}$$

current source dipole amplitudes (same length as above)

$$\vec{x} = \vec{A} \vec{s} + \vec{n}$$

note: only one current source for each column in the  $E+B$  matrix!

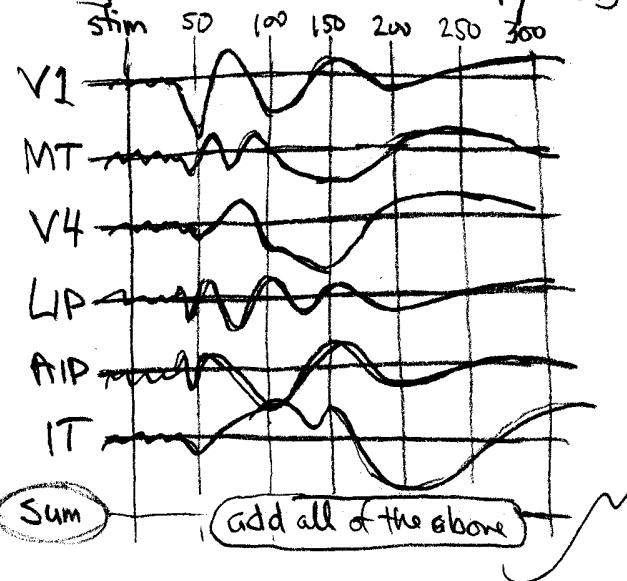
inverse-φ

## WHY LOCALIZE?

- most of ERP literature based (instead) on temporal "components"
- but:
  - 1) underlying local cortical generators (from microelectrode LFP, CSD)
    - extended in time (400 msec), visible from every scalp electrode
    - multiphasic in every cortical area
    - temporally non-static depending on stimulus
      - ↳ e.g. simple contrast, brightness diffs can modulate retinal delay by 50 msec!
  - 2) thus, any "component" consists of sum of activity from multiple cortical areas at different hierarchical levels
  - 3) stimulus manipulations will change temporal overlap
    - ↳ may cause "component" peak to disappear without changing cortical areas being activated
  - 4) verified by intra cortical LFP/CSD (Schroeder et al, 1998)

macaque monkey  
intra cortical data

→ these areas span the visual system from bottom to top, accounting for roughly 50% of the entire macaque monkey CTX



LFP's from approx. layer 4 in cortex ("input layer")

psychologists now identify a few temporal "component" peaks...

↳ but each peak comes from every one of these cortical areas!!

- by contrast, the spatial signature of the signal from one cortical area is static — a better area-based "component"
- temporal "components" should be retired!!
  - easier to record more temporal points (EEG started w/ few electrodes, many time points)
  - easier to "paste" high level psychological functions onto a few waveform deflections
- their original reason for being no longer relevant!!

## inverse1 Derivation of Ill-posed Inverse

(from Dale & Sereno, 1993)

$$\mathbf{x} = \mathbf{As} + \mathbf{n}$$

solve for inverse operator

$$\text{Err}_w = \langle \|W\mathbf{x} - \mathbf{s}\|^2 \rangle$$

$\mathbf{x}$  = sensor data vector

$\mathbf{A}$  = forward soln matrix ( $\mathbf{E} + \mathbf{B}$ )

$\mathbf{s}$  = source vector

$\mathbf{n}$  = sensor noise vector

probability that rand. var. value is  $k$

$$\text{expectation: } \left[ = \sum_k p_k k \right] \text{ rand var. value}$$

assume  $\mathbf{n}, \mathbf{s}$  normal, zero-mean w/ corresponding covar. matrices  $\mathbf{C}, \mathbf{R}$

$$\text{Err}_w = \langle \|W(\mathbf{As} + \mathbf{n}) - \mathbf{s}\|^2 \rangle$$

$$= \langle \|(WA - I)\mathbf{s} + W\mathbf{n}\|^2 \rangle$$

$$= \langle \|Ms + Wn\|^2 \rangle \quad \text{where } M = WA - I$$

$$= \langle \|Ms\|^2 \rangle + \langle \|Wn\|^2 \rangle \quad \text{diag is noise variance (already squared)}$$

$$= \text{tr}(M^T M) + \text{tr}(W^T W) \quad \text{trace is sum of diag elements}$$

$$(\text{re-expand}) = \text{tr}(WARAT^T - RAT^T - WAR + R) + \text{tr}(WCW^T)$$

~~Explicitly minimize by taking derivative w.r.t.  $W$ , set to zero, solve for  $W$~~

$$0 = 2WARAT - 2RAT^T + 2WC$$

$$WARAT + WC = RAT$$

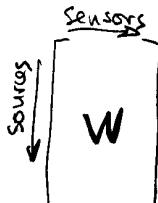
$$W(ARA^T + C) = RAT$$

$$W = RAT (ARA^T + C)^{-1}$$

equivalent to minimum norm and Tikhonov regularized

Inverse if  $\mathbf{C}, \mathbf{R}$  are proportional to identity matrix (i.e., sensor noise & sources independent and equal variance)

$W$  is inverse solution operator:



inverse-2

# INVERSE Sol'n (2)

w/ liberal priors

sources uncorrelated ( $R$ )  
 noise uncorrelated ( $C$ )

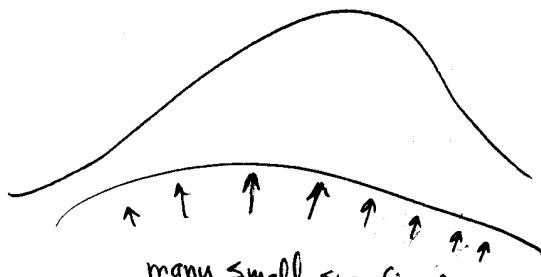
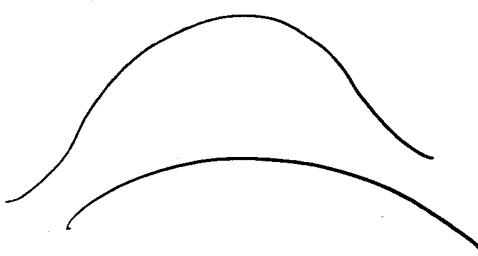
$$W = RA^T(ARA^T + C)^{-1}$$

↳ "minimum norm" solution  
 (find  $\vec{s}$  w/ smallest norm =  $\|\vec{s}\|$ )

- the minimum norm solution appropriately downplays deeper (=weaker scalp signal) sources since these are more likely to fall into the noise floor

- "problems" of minimum norm:

deeper sources get displaced to the surface



achieves smaller norm w/ equivalent fit/error

also tiny dipoles in correct place  
(see noise normalization)

- small superficial sources "win" because of approx. inverse square form of fwd solution

↳ smaller norm of distributed superficial soln

- can't fix by increasing priors of deep sources!!

↳ that will give deep sources given noise as input!!

inverse-3

# INVERSE SOLUTIONS TO ILL-POSED (COMPARED)

$$\begin{matrix} \text{Sources} \\ \downarrow \\ \mathbf{s} \end{matrix} = \mathbf{W} \mathbf{x}$$

sensors

$$\mathbf{s} = \begin{matrix} \text{sources} \\ \downarrow \\ \mathbf{W} \end{matrix} \begin{matrix} \text{sensors} \\ \downarrow \\ \mathbf{x} \end{matrix}$$

sensor data

- how to use the inverse solution,  $\mathbf{W}$
- same  $\mathbf{W}$  for all time points

"minimum norm" solution

i.e., norm  $\|\mathbf{s}\|$  of solution is smallest of infinitely many alternate solutions

Over-determined  
↳ least squares  
Under-determined  
↳ minimum norm

linear inverse operator  $\downarrow$

$$\mathbf{W} = \mathbf{R} \mathbf{A}^T (\mathbf{A} \mathbf{R} \mathbf{A}^T + \mathbf{C})^{-1}$$

spatial covariance  
sources      forward soln

spatial covariance  
sensor noise

from error minimization derivation

$\begin{matrix} \text{Sensors} \\ \downarrow \\ \mathbf{W} \end{matrix} = \begin{bmatrix} \mathbf{R} & \mathbf{A}^T \end{bmatrix} \left( \begin{bmatrix} \mathbf{A} & \mathbf{R} \end{bmatrix} \begin{bmatrix} \mathbf{A}^T & \mathbf{R} \end{bmatrix} + \mathbf{C} \right)^{-1}$

$\Downarrow$

$\begin{bmatrix} \mathbf{A} \mathbf{R} \mathbf{A}^T \end{bmatrix} \Rightarrow$  square in # of sensors (small)

$\begin{bmatrix} \mathbf{A}^T \mathbf{C}^{-1} \mathbf{A} + \mathbf{R}^{-1} \end{bmatrix} \Rightarrow$  square in # of sources (large)

$\begin{bmatrix} \mathbf{A}^T \mathbf{C}^{-1} \mathbf{A} \end{bmatrix}$  easier inverse

linear inverse operator

$$\hookrightarrow \mathbf{W} = (\mathbf{A}^T \mathbf{C}^{-1} \mathbf{A} + \mathbf{R}^{-1})^{-1} \mathbf{A}^T \mathbf{C}$$

alternate, algebraically  
equivalent Bayesian derivation  
(w/ bigger inverses!)

$\begin{bmatrix} \mathbf{W} \end{bmatrix} = \underbrace{\begin{bmatrix} \mathbf{A}^T & \mathbf{C} \end{bmatrix} \begin{bmatrix} \mathbf{C}^{-1} & \mathbf{A} \end{bmatrix}}_{\text{easy}} + \begin{bmatrix} \mathbf{R} & \mathbf{A}^T \end{bmatrix} \begin{bmatrix} \mathbf{C} \end{bmatrix}$

$\Downarrow$

$\begin{bmatrix} \mathbf{A}^T \mathbf{C}^{-1} \mathbf{A} \end{bmatrix} \Rightarrow$  both square in # of sources (large)

hard inverses

use inverse-1

# PROBLEMS W/ SURFACE NORMAL

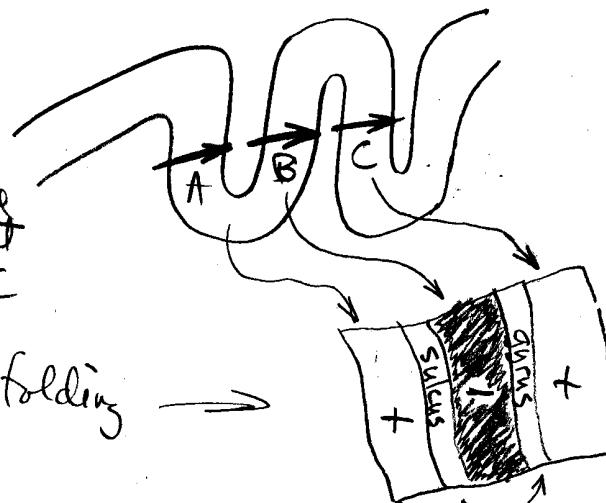
- Since nearby points on surface often have different orientation, surface normal constraint can help (since fwd soln A,B very different)



- but, since point spread function: point  $\xrightarrow{\text{fwd}}$  data spread  $\xleftarrow{\text{invers}}$   
typically extends across sulci, artificial sign reversals occur

positive current at A indistinguishable from negative current at B, or pos at C

→ after unfolding



## - Solutions

- 1) ignore sign → saves useful orientation info !

- 2) solve onto 3 orthogonal dipoles at each cortical point instead of a single oriented dipole

→ more appropriate when averaging across subjects, since detailed orientations vary a lot

→ also, fills in bottom of sulcus (else unsigned stripes)

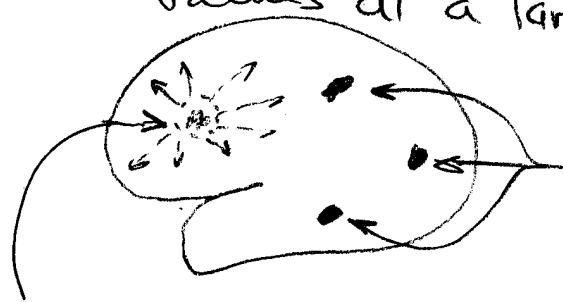


use inverse-2

## fMRI Constrained Inverse

(could be combined w/ MUSIC!)

- insert fMRI values for  $R_{ii}$ 's
- but still allow other sites to have non-zero  $R_{ii}$ 's
- pathologies occur if solution restricted completely to fMRI points by setting non fMRI  $R_{ii}$ 's to zero → set to small number instead!
- this allows extracting time course from sources visible in EEG/MEG and fMRI
- N.B.: sources that are only visible in EEG/MEG will be dispersed to small distributed values at a large number of vertices



visible in both EEG/MEG  
and fMRI

visible only  
in EEG/MEG

and not fMRI → distributed at small amplitude across many vertices

use inverse-3

# NOISE SENSITIVITY NORMALIZATION (1)

forward:  $\mathbf{x} = \mathbf{Ax}$  well-posed (Liu, Dale, and Belliveau, 2002)

inverse:  $\mathbf{s} = \mathbf{Wx}$  ill posed

Solve:  $\mathbf{x} = \mathbf{As} + \mathbf{n}$  for  $\mathbf{s}$

$$\mathbf{W} = \mathbf{R}\mathbf{A}^T(\mathbf{A}\mathbf{R}\mathbf{A}^T + \mathbf{C})^{-1}$$

Variance of dipole strength estimate due to additive sensor noise  
 $\text{Var}(S_i) = \langle (W_i n(t))^2 \rangle = W_i C W_i^T$   
 that is, sensor noise run through inverse

- multiply inverse operator by noise sensitivity matrix,  $\mathbf{D}$  (diagonal)

$$D_{ii} = \frac{1}{\text{diag}_i \sqrt{WCW^T}}$$

$$W^{\text{ns\_norm}} = DW$$

normalize each row of inverse operator by noise sensitivity at that location

$$S_i^{\text{ns\_norm}} = (W^{\text{ns\_norm}} \mathbf{x})_i = (DW\mathbf{x})_i = \frac{W_i \mathbf{x}}{\sqrt{(WCW^T)_i}} = \sqrt{\frac{(W\mathbf{x} \mathbf{x}^T W^T)_i}{(WCW^T)_i}}$$

if assume Gaussian white noise, noise covariance,  $C$ , is multiple of  $\mathbf{I}$ , so

$$W_i^{\text{ns\_norm}} = \frac{W_i^{\text{orig}}}{\|W_i^{\text{orig}}\|}$$

$$\|w_i\| = \sqrt{(WCW^T)_i}$$

$$[w][\epsilon][w^T]$$

$\Rightarrow WW^T \Rightarrow$  each entry is sum of squares

i.e., scale each row of  $W$  by single value — the norm of that row

row of  $W$  is:

one source  
one row sensors

$$S_i = \frac{W_i^{\text{orig}} \cdot \mathbf{x}}{\|W_i^{\text{orig}}\|}$$

$$\begin{bmatrix} S \\ \vdots \\ S \end{bmatrix} = \begin{bmatrix} & & & \\ & W & & \\ & & & \\ & & & \end{bmatrix} \begin{bmatrix} X \\ \vdots \\ X \end{bmatrix}$$

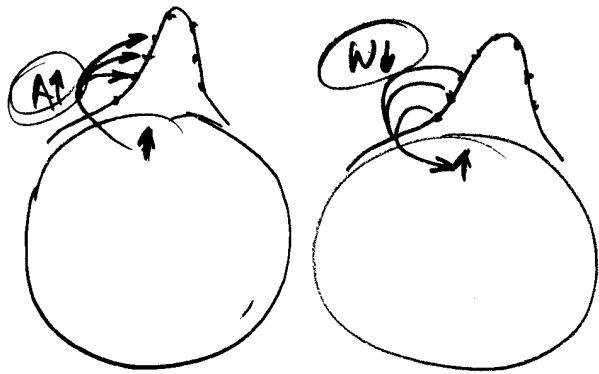
inverse solution  
coefficients for  $\Rightarrow$  scale(divide) by one source norm of this row

that is, if inverse sol'n for deep source is reduced by interaction of inverse square nature of fwd and min norm, dividing by norm of row of inverse (1 source) will increase/rescue deep source

use inverse-4

# NOISE SENSITIVITY NORMALIZATION (2)

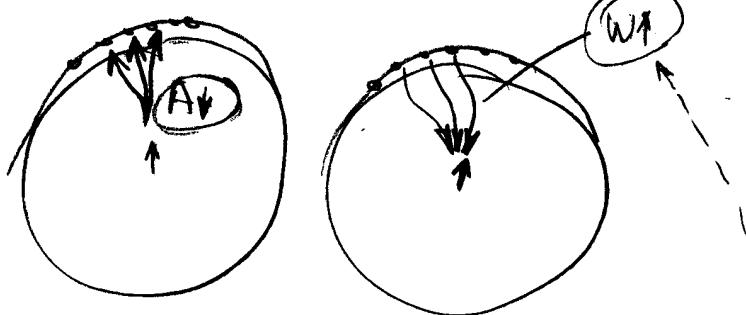
Shallow source (unit strength)



fixed big

inv small

deep source (unit strength)



fixed small

inv reduced because  
of minimum norm

(N.B. W should be  
bigger than for  
superficial source --  
but, min norm reduces  
b/c of inverse square)

$$S_i = \overrightarrow{W_i^{\text{orig}}} \cdot \overrightarrow{x}$$

$$\| \overrightarrow{W_i^{\text{orig}}} \|$$

shallow: small

deep & even smaller!

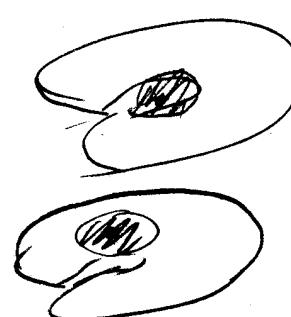
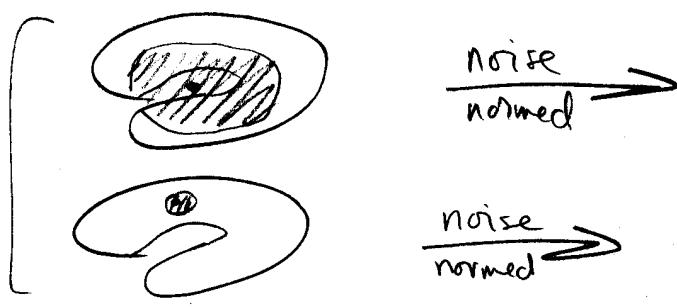
therefore, fixed  
estimate increased  
relative to shallow

- effect on inverse solution  $\rightarrow$  more like significance vs actual power

- effect on point-spread function is to equalize shallow & deep

[Shallow spread out more than min norm  
deep shrunk to same as shallow]

point  
spread  
functions



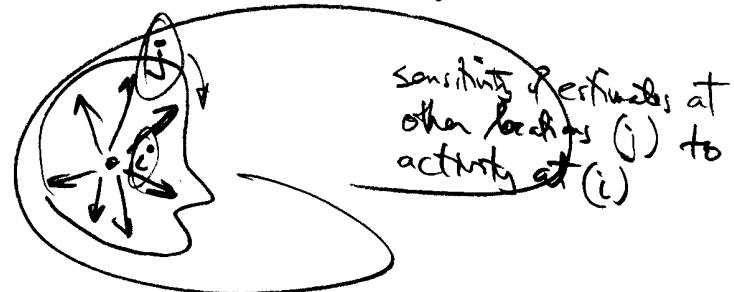
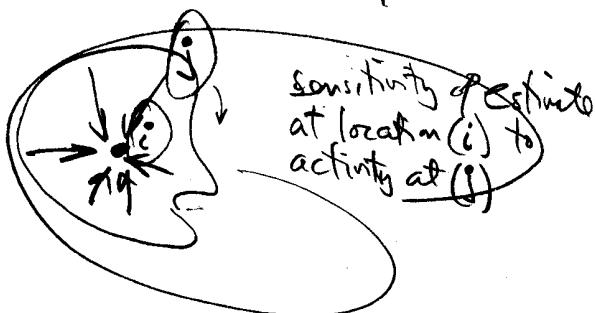
point-spread  
Point  $\xrightarrow{\text{fixed}}$  data  
spread  $\leftarrow$  in

use inverse-S

# POINT-SPREAD / CROSS TALK

point  $\xrightarrow{\text{fwd}} \xrightarrow{\text{inv}} \text{blur}$

thus,  $WA$  measures blur!  
just "take inverse of fwd"



Crosstalk (rows of resolution matrix,  $WA$ )

$$\xi_{ij}^2 = \frac{|(W\tilde{A})_{ij}|^2}{|(W\tilde{A})_{ii}|^2} = \frac{|w_i \tilde{a}_j|^2}{|w_i \tilde{a}_i|^2}$$

Point spread (columns of resolution matrix,  $WA$ )

$$P_{ij}^2 = \frac{|(W\tilde{A})_{ji}|^2}{|(W\tilde{A})_{ii}|^2} = \frac{|w_j \tilde{a}_i|^2}{|w_i \tilde{a}_i|^2}$$

avg crosstalk

$$ACM_i = \frac{\sum_j \xi_{ij}^2}{j}$$

avg point spread

$$APSF_i = \frac{\sum_j P_{ij}^2}{j}$$

- PSF & crosstalk maps identical for standard inverse ( $WA$ , the "resolution matrix" is symmetric)
- PSF affected by noise-normalized; crosstalk same  
 $((DW)A$  not symmetric)

## Conclusions

- more EEG or More MEG better
- EEG better than MEG (cf. radial) (EEG fwd<sub>real</sub> currently less accurate)
- biggest gain from adding small # EEG (or MEG) (e.g. 30)  
to many MEG (or EEG) (e.g. 150)
- easier to add many MEG, so: optimal
  - 30 EEG
  - 300 MEG
- EEG/MEG forward-solution-scaling-factor error  
Causes  $\rightarrow$  more cross talk

# music-1

## MUSIC

(1)

(from Dale &amp; Sereno, 1993) (cf. Mosher &amp; Leahy)

— using sensor covariance

$$\mathbf{D} = \underbrace{\langle \mathbf{x} \mathbf{x}^T \rangle}_{\substack{\text{average} \\ \text{value of covariance}}} = \underbrace{\sigma^2 \mathbf{I}}_{\substack{\text{sensor} \\ \text{noise}}} + \sum_i \sum_j \sigma_i \sigma_j \text{Corr}(i, j) \mathbf{A}_i \mathbf{A}_j^T$$

recording vectors at one time point

$$\approx \frac{[\mathbf{x}_1 \dots \mathbf{x}_n] [\mathbf{x}_1 \dots \mathbf{x}_n]^T}{n}$$

recording time points

can take this estimate over short time epochs  
 $t \rightarrow$

$$\mathbf{D} = \mathbf{U} \Lambda \mathbf{U}^T = \left[ \begin{matrix} \mathbf{u}_1 & \mathbf{u}_2 & \dots & \mathbf{u}_n \end{matrix} \right] \left[ \begin{matrix} \lambda_1 & & & \\ & \ddots & & \\ & & \lambda_n & \end{matrix} \right] \left[ \begin{matrix} \mathbf{u}_1 & \mathbf{u}_2 & \dots & \mathbf{u}_n \end{matrix} \right]^T$$

↑ eigenvector      ↑ Corresponding eigenvalue

\* Columns of  $\mathbf{U}$  matrix are orthogonal basis vectors of "spatial pattern" space (one per unit spatial pattern across sensors)

= find, order most significant spatial patterns in sensors over time

Project forward solns onto these spatial patterns  
 ("project" = dot prod = similarity)  
 (fwd defines spatial sensor pattern for unit source)

$$\xi_i = \underbrace{\mathbf{A}_i^T \mathbf{U} \Lambda \mathbf{U}^T \mathbf{A}_i}_{\substack{\text{eigenvals} \\ \text{all eigenvectors of sensor spatial patterns}}} \Rightarrow$$

one num for each source

fwd for  $\rightarrow$  1 column  
 one source ("gain vector") of  $\mathbf{A}$  matrix

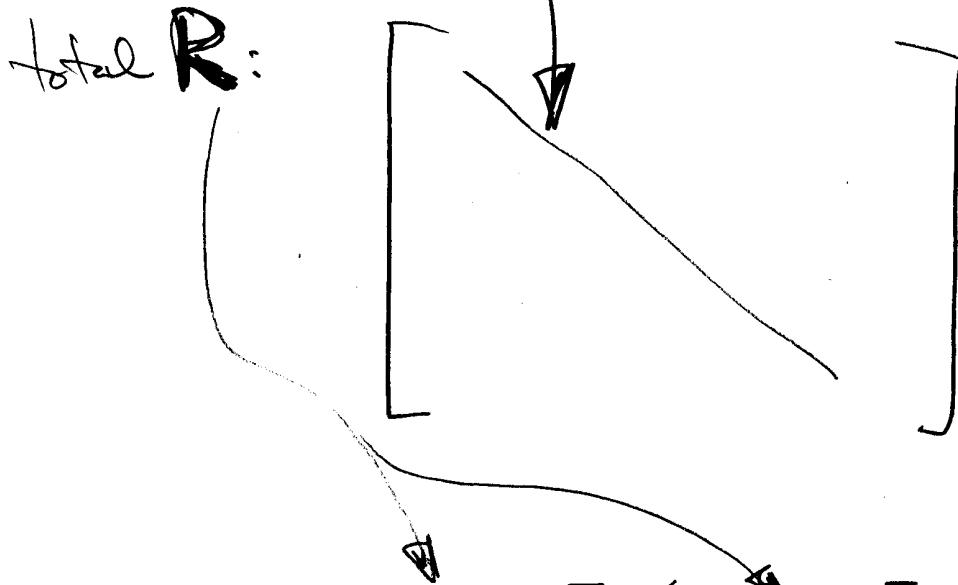
big single number if forward soln looks like  $\mathbf{U}_i$

MUSIC-2  
MUSIC

(2)

how to weight the  
minimum norm inverse

$$R_{ii} \approx \frac{A_i^T A_i}{A_i^T U \Lambda^{-1} U^T A_i}$$



$$W = R A^T (A R A^T + C)^{-1}$$

cf.

- like parallel resistance

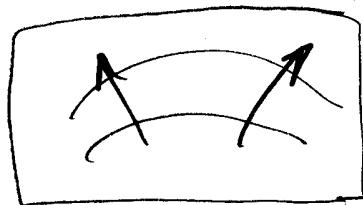
$$R_{\text{parallel}} = \frac{1}{\frac{1}{R_1} + \frac{1}{R_2} + \dots}$$

any low resistance ( $\xi_i$ ) decreases overall resistance (overall  $R_{ii}$ )

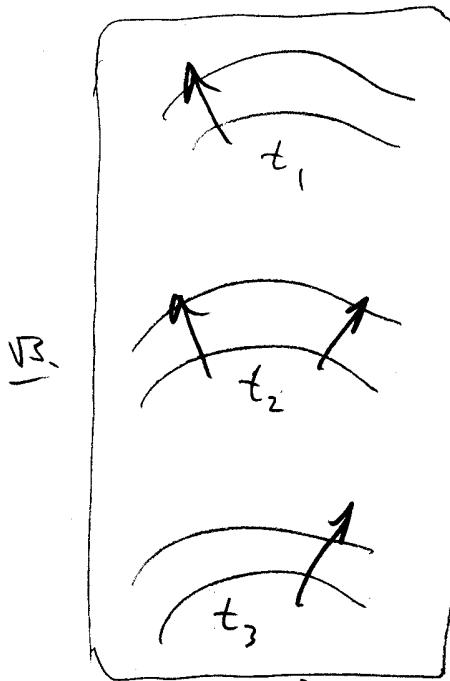
i.e., if forward sol'n has appearance like any low eigenvalue spatial pattern, it gets devalued

music<sup>3</sup>  
MUSIC (?)

- how it works: take advantage of spatial information that changes over time



one time  
point



multiple  
time points

N.B.:

Problem if assumption about lack of perfect correlation is violated

e.g., if two widely separated sources (i.e. diff fird solns) are highly correlated, MUSIC will eliminate both since no single fird soln will look like that "2-separated dipole" pattern (e.g., L/R A-I)  
↳ "dual MUSIC" hack to fix...

- how it fixes min norm problem

