

Flow Effects in Magnetic Resonance (MR) Imaging, MR Angiography, MR Perfusion

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### Part I: Flow in MR Imaging

- 1 Relationship between velocity and phase of the MR signal
- 2 Relationship between acceleration and phase of the MR signal
- 3 Artifacts
- 4 Velocity insensitive sequences

## Overview of flow imaging

MR flow measurements can be distinguished in

- Visualization of flow (Angiography, MRA)
- Quantitative flow measurement (q-flow)

Measurement is done either for phase (phase contrast between moving tissue and stationary tissue), or for magnitude (magnitude contrast between moving and stationary tissue).

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## Q-flow, Phase Contrast

**Principle of PC MRI:** Use the influence of gradient fields on the signal phase to recover position, velocity, and acceleration of the flowing tissue (blood in arteries and veins of body and brain, CSF).

## Relation between velocity and phase for the unipolar gradient

Position of blood  $r(t) = r_0 + v_r(t - t_0) - \frac{1}{2} a(t - t_0)^2$  Taylor 2nd Order

Motion and gradient in x-direction with phase-measurement of the spins at position  $r(t_m) = x(t_m) = x_0$  at time  $t_m = t_0$

The phase at  $t = t_m$  is given by (Substitute  $t \mapsto t' = t - t_m$ )

$$\phi(t_m) = \gamma \int_{T/2}^{T/2+t_m} \mathcal{G}_x dt = \gamma \int_{T/2-t_m}^{T/2} \left( x_0 + v_m t' - \frac{1}{2} a t'^2 \right) \mathcal{G}_x dt'$$

For

$$\phi(t_m) = \gamma x_m A = \gamma v_m A (T - t_m) + \gamma \frac{aT}{2} (T - t_m)^2 - \frac{r_m^2}{12}$$

We get

Source: Vlaardingerbroek et al.

## Relation between velocity and phase for the bipolar gradient

Suppose: Flow of constant velocity  $\phi = \gamma A \mathcal{G} t_1 = \gamma A \mathcal{G} t_2 = \gamma v A (t_2 - t_1) = \gamma v A t'$

With the First Order Moment of the bipolar gradient  $A t' = m_1 = \int \mathcal{G}(t) dt$

The areas of the bipolar gradient have to be same, so that the phase of spins in stationary tissue is not changed:

The Zero Order Moment vanishes  $m_0 = \int \mathcal{G}(t) dt = 0$

Source: Vlaardingerbroek et al.

### Relationship between velocity and phase for the back-to-back bipolar gradient



Two bipolar gradients with opposite First Order Moments lead to a phase independent on velocity and position

$$m_0 = m_1 = 0$$

The back-to-back bipolar gradients can be used for the design of velocity-insensitive sequences

In case of subtraction of an image of a velocity sensitive sequence from an image from a velocity insensitive sequence, only the flow remains (PC-Angio)

Source: Vlaardingerbroek et al.

### Accelerated flow and use of a bipolar gradient

Since velocity depends on time, only velocities at a certain point in time can be measured. Evaluation of the phase evolution for a bipolar gradient yields

$$\varphi(t_m) = \gamma \cdot \nu t_m / 4 (T_1 - T_2) + \gamma \cdot A \cdot \sqrt{4 (T_2 - T_1) + \frac{1}{2} (T_1^2 - T_2^2)}$$

The time of expansion, where the phase gets independent on acceleration, is given by

$$t_p = \frac{1}{2} (T_1 + T_2) \quad \text{Time, where the velocity is determined}$$

There we have

$$\varphi(t_p) = \gamma \cdot \nu t_p / 4 (T_1 - T_2)$$

$\nu(t_p)$  can be found for a corresponding bipolar gradient this way

A bipolar gradient allows to relate the velocity in the center of gravity of the gradient to the phase of the spins

Source: Vlaardingerbroek et al.

### Accelerated flow and the back-to-back bipolar gradient

For a simple back-to-back bipolar gradient with 3 areas we have

$$m_0 = \int G(t) \cdot dt = 0$$

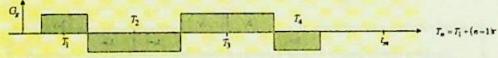
$$m_1 = \int G(t) \cdot t \cdot dt = 0$$

$$m_2 = \int G(t) \cdot t^2 \cdot dt = 4 \cdot \pi \cdot r^2$$

The relationship between phase and velocity given by

$$\varphi = \gamma \cdot \nu \cdot m_2$$

vanishes for multiple back-to-back bipolar gradients with 4 areas



Source: Vlaardingerbroek et al.

### Field inhomogeneities and Eddy currents

$$\varphi(t) = \gamma \int (\partial \mathcal{B} + \partial \mathcal{B}_c + \nu G_x(t)) dt$$

Eddy Current

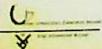
BO Return

Suitable gradient coil  
Measurement 2x with different  $m_1$

Source: Vlaardingerbroek et al.

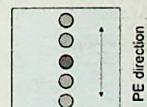
### Artifacts I: Signal extinction

**Flow void:** Signal extinction is provoked by strong velocity gradients. Strong velocity gradients appear beyond stenoses, bifurcations, or in regions with turbulent flow. The intra-voxel dephasing then leads to „flow voids“. This can be reduced or even avoided by the choice of short echo times.

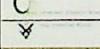


### Artifacts II: Ghosting

**Ghosts:** One finds "ghost images" in phase-encoding direction, since the periodically pulsating flow interferes with the MR-signal.



Artery perpendicular to the image



Source: Vlaardingerbroek et al.

### Artifacts III: Misregistration

The position is encoded when the blood arrives at the center T of the phase encoding gradient. For the unipolar gradient, we then have:

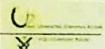
$$\varphi(t_m) = \gamma \cdot t_m \cdot A + \gamma \cdot v_y \cdot \Delta T - t_m \quad \text{especially for} \quad t_m = (n-1) \cdot \Delta T + TE$$

On the other hand, the spins get at position  $y(T)$  a phase-shift by the encoding gradient of

$$\varphi = \gamma \cdot n \cdot t_c \cdot \delta(T) + \gamma \cdot n \cdot A \cdot T$$

It follows the relationship  $y(T) = \delta(T) + v_y(T - TE)$

At the time of the echo, blood at  $[\delta(TE), \delta(TE)]$  is observed actually at  $[\delta(TE) + v_y(T - TE), \delta(TE) + v_y(T - TE)]$



### Flow compensation

#### Subtraction method:

- Use a flow sensitive sequence to get an image
- Use a flow insensitive sequence to get an image
- Compute the difference between the images

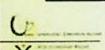
### I Selection gradient



Between  $t = 0$  and  $t = t_1$  bipolar, thus flow sensitive:

$$\varphi(t_m) = \gamma \cdot t_m \cdot A \cdot (T_1 - T_2) + \gamma \cdot A \cdot \left[ t_1 \cdot (T_2 - T_1) + \frac{1}{2} (T_1^2 - T_2^2) \right]$$

Flow insensitivity can be established by adding a second, invers bipolar gradient.



Source: Vlaardingerbroek et al.

### II Readout direction



At the time of the echo the zero order moment of the gradient vanishes, as well for SE as for GRE Imaging

In case of a bipolar gradient, the phase of the moving spins becomes velocity dependent. To get independence, a second bipolar gradient can be added.

Source: Vlaardingerbroek et al.

### III Phase encoding direction

In phase encoding direction flow results in spatial misregistration. Correcting this misregistration is possible while encoding the phase in a way as if it were measured at  $t = TE$ . For each phase encoding step an extra phase

$$\Delta \varphi = \gamma \cdot n \cdot A \cdot v_y \cdot (TE - T)$$

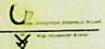
( $\Delta \varphi = \gamma \cdot n \cdot A \cdot t'$  bipolar Gradient)

is to be added. This can be done by adding an extra-bipolar gradient. The corresponding first order moment then can be computed as

$$m_1 = A_2 \cdot (n)'' = \gamma \cdot A \cdot (TE - T)$$

and results in a position encoding that doesn't depend on velocity.

If there are no extra bipolar gradients added, the position encoding of the first echo of imaging sequences becomes velocity dependent.



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#### Part II: MRA (MR Angiography)

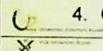
- Phase contrast angiography (PCA)
- Magnitude contrast angiography (MCA)
- Artifacts
- "Time of flight"-method

## PC Angio: Subtraction method



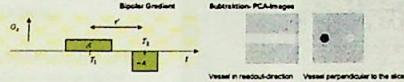
Phase contrast methods are subtraction methods.

1. Take a flow insensitive image.
2. Take 3 flow sensitive images with flow sensitivity in all 3 directions.
3. Subtract the flow insensitive image from the flow sensitive ones.
4. Compute a magnitude image from the 3 images.



Source: Vlaardingerbroek et al.

## Phase-contrast angiography: Quantitative Flow



Subtraction of flow insensitive GRE images from flow sensitive ones allows to determine the velocity of flow and to draw quantitative flow maps.

The method is given by the velocity vector

$$I_{Qflow} = I_1(t_{1a}) - I_1(t_1) + I_2(t_{2a}) - I_2(t_2) + I_3(t_{3a}) - I_3(t_3)$$

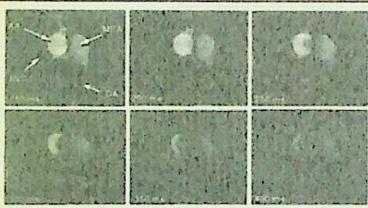
or, expressed by gradients and flow velocities

$$I_{Qflow} = I_1 A_x v_x + I_2 A_y v_y + I_3 A_z v_z$$

SE cannot be used due to cutoff between the excitation and the refocusing pulse.

Source: Vlaardingerbroek et al.

## Phase-contrast angiography Quantitative Flow



Ascending aorta  
Main pulmonary artery  
Descending aorta  
Superior vena cava

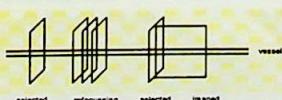
Source: Edelman et al.

## Magnitude contrast angiography

- I. 'Time-of-flight' or 'Inflow' Angiography
- II. Contrast-Enhanced (CE) Angiography
- III. Magnetisation Preparation
- IV. 'Black-Blood' Angiography
- V. Artifacts in MCA

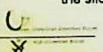
## I. TOF or 'Inflow' Angiography

Time-of-flight Method:  
(a) Refocusing slices parallel to the selected slice.  
(b) Refocusing slice parallel to the vessel.



Chose a refocusing rf-pulse in a refocusing slice. Using several refocusing slices at various positions, this allows to assess a velocity profile of the flow.

The velocity profile allows to compute the amount of blood flowing to the slice.



Source: Vlaardingerbroek et al.

## I. TOF or 'Inflow' Angiography

3 D Time-of-flight (TOF) image of the circle of Willis.

TR/TE/FA = 40/6/23°

Distal flow is better retained with smaller FA (see the arrows).

Acquisition with higher FA (45°) leads to background suppression.



Source: Edelman et al.

## II Contrast-Enhanced Angiography

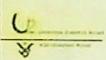
CE-MRA Techniques use the shortening of T1 in blood by CA. T1-FFE spoiled gradient-recalled echo-Sequences are used. The transversal magnetization in steady-state (after a few RF-Pulses) is given by

$$M_T(x, y) = M_0(x, y) \cdot \frac{\sin(\alpha) \cdot (1 - E_1) \cdot \exp\left(-\frac{T_E}{T_1}\right)}{(1 - E_1 \cdot \cos\alpha)}$$

Steady-State Contrast  $C(T1) = (MT(\text{blood}) - MT(\text{background}))$   
 Blood+CA ( $T1 = 40 \text{ ms}$ ;  $T2 = 28 \text{ ms}$ )  
 Stationary Tissue ( $T1 = 700 \text{ ms}$ ,  $T2 = 60 \text{ ms}$ )  
 for GRE sequence parameters  $TR = 7 \text{ ms}$ ,  $TE = 4 \text{ ms}$

Flip Angle [Deg]

The time between injection and measurement is important!



## II Contrast-Enhanced Angiography

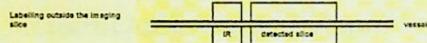
Maximum Intensity Projection (MIP) of two consecutive acquisitions of the carotid arteries.

A: Appropriate timing with strong arterial and low venous signal.

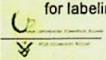


Source: Edelman et al.

## III Magnetisation Preparation



- Preparation pulses can be used to enhance the contrast between stationary and moving tissue
- Contrast for inflow-MRA can also be enhanced by MT (Magnetization Transfer) - Pulses
- Labeling inflowing blood by IR together with the subtraction technique is another technique
- The fact that oxygenation enhances T2 of blood can also be used for labeling



Source: Vlaardingerbroek et al.

## IV Black-Blood Angiography

Black blood: The SE technique can be used for the "black blood" signal extinction of flowing blood. The blood therefore has flown out of the imaging region at the time the second (inversion) pulse is applied. The remaining blood has not experienced the first (excitation) pulse and therefore does not give a signal. The signal thus disappears, and the flowing blood appears black.

The SNR remains somewhat limited with this technique.



## IV Black-Blood Angiography

Black blood technique: The arrow shows an aortic dissection.



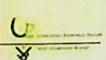
Source: Edelman et al.

## V Artifacts in MC Angiography

Pulsating flow: Artifacts are present in the change of vessel diameter (10-15%) due to pulsating flow.

Contrast Agent (CA) inflow: Imaging during inflow of CA to the imaging region.

Turbulent flow: Turbulent flow leads to signal extinction.



Magnetic Resonance Perfusion and Diffusion Imaging, functional MRI

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## Perfusion Sensitive Sequences

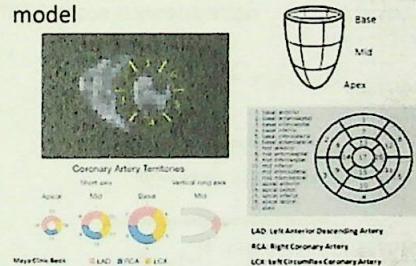
- In MRI: Measurement of AIF, TRF using CA, T1w MR Seq. with  $SI \propto CA$
- First Pass, Dual Bolus, Fast/Ultrafast Imaging, SSFP and EPI, Ex. Turbo FLASH

## Tracers

- Contrast Agent (CA), Transit Times
- In the past: Radioactive Tracers as  $O^{17}$ ,  $H^2$  (Deut) (Endogenous)
- Exogenous like Gd-DTPA (Magnevist), Molecular Weight
- Without CA App., Ex. ASL

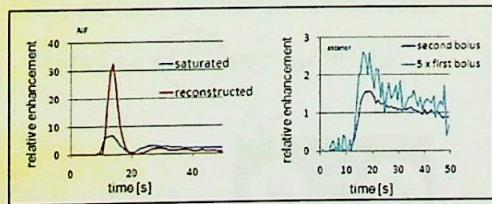
## Example outside Brain

The 17 segment heart model

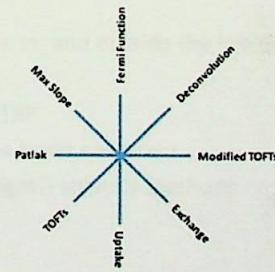


## Data acquisition and analysis

### AIF and TRF



## The 8 perfusion flow analysis methods



## Functional Analysis

- Functional or Operator
- Lives in Sobolev or Hilbert Spaces
- Example: The antiderivative

$$F(x) = \int_a^x f(\xi) d\xi$$

$$F' = f$$

## Convolution

- Definition

$$F(k) \otimes G(k) = \frac{1}{\sqrt{2\pi}} \int_{-\infty}^{+\infty} F(k - k') G(k') dk'$$

'k-space'

- Then it holds that

$$f(x) \cdot g(x) \leftrightarrow F(k) \otimes G(k)$$

Used for solving DEs

## Algorithm

Examples of algorithms giving a number:

- Babylonian Algorithm (1700 BC)
- Newton-Raphson
- Levenberg-Marquardt

Stopping problem: An algorithm always terminates!

## Fermi Function

- Total tissue concentration

$$C = F_P R \otimes C_{P,A}$$

Fermi-Dirac Distribution Function

$$R(t) = \{\exp((t - t_0 - t_d)k + 1)\}^{-1} u(t - t_d)$$

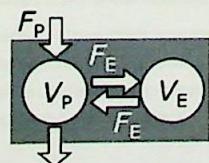
Heaviside or Integrated Dirac Function

$$u(t - t_d) = \begin{cases} 0 & t < t_d \\ 1 & t > t_d \end{cases}$$

## Modified TOFTs

- General form of modified TOFTs model:

$$C = V_P C_{P,A} + K^{\text{trans}} \exp(-k_{ep}t) \otimes C_{P,A}$$



## Resume

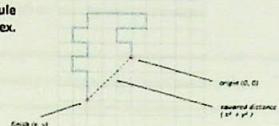
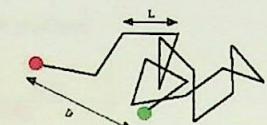
- Perfusion in- and outside the brain
- Tracers
- AIF and TRF
- Examples heart and brain
- The 8 (eight!) analysis methods

## Random Walk

Brownian motion



Unlimited motion of one molecule  
in an ensemble of molecules (ex.  
AIR)



## Brownian motion

The Einstein Diffusion Equation describes microscopic transport of particles and heat:

$$\partial_t p(r, t | r_0, t_0) = \frac{\sigma^2}{2\gamma^2} \nabla^2 p(r, t | r_0, t_0)$$

## Brownian motion

From the EDE, we obtain as spatial distribution of particles in ex. AIR:

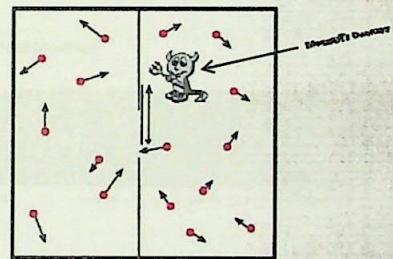
$$\begin{aligned} \langle (r(t) - r(t_0))^2 \rangle &= 6 \frac{\sigma^2}{2\gamma^2} t & \langle r^2 \rangle &= 2 D t \\ D &= \frac{\sigma^2}{2\gamma^2} & \langle r^2 \rangle &= 4 D t \\ & & \langle r^2 \rangle &= 6 D t \end{aligned}$$

During diffusion, entropy increases.

## Entropie

Entropie is a measure of disorder

Linked terms:  
Energie  
Probability  
Information



## Cybernetics

...society can only be understood through a study of the messages and communication facilities which belong to it; and that in the future development of these messages and communication facilities, messages between man and machines, between machines and man, and between machine and machine, are destined to play an ever increasing part. (p.15)

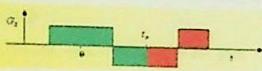
from the article "Cybernetics In History", Wiener (1954), cited after Maggie Mc Garry, "Norbert Wiener's Cybernetic Theory and Parental Control", URL [http://www.colorado.edu/communication/meta-discourses/Papers/App\\_Papers/McGarry.htm](http://www.colorado.edu/communication/meta-discourses/Papers/App_Papers/McGarry.htm), Feb 14 2013.

## Mathematics for flow imaging

- Flow independent sequences
- Flow compensation
- Subtraction method
- Phase or Magnitude

## Mathematics for flow imaging

### Example: Slice selection gradient



Between  $t = 0$  und  $t = t_1$ , bipolar, thus flow sensitive:

The signal phase depends on velocity and acceleration.

Flow insensitivity can be established by adding a second, invers bipolar gradient.

## Mathematics for diffusion

### The decay of the Transverse Magnetization (TM)

#### Bloch Equation for TM:

$$\frac{dM_T}{dt} = -\gamma \cdot G \cdot r \cdot M_T - \frac{M_T}{T_2}$$

#### Transport Process:

$$j_{M_T} = D \nabla M_T$$

## Mathematics for diffusion

### Solution for the Transverse Magnetization (TM)

$$M_T(t) = M_T(0) \exp \left[ - \int_0^t (\zeta^T(t') D \cdot \zeta(t') dt') \right] \exp \left( -J \cdot \zeta(t) r - \frac{t}{T_2} \right)$$

#### In case 1D

In this case we can write the TM as

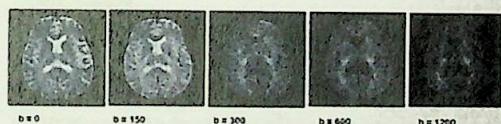
$$M_T(T_2) = M_T(0) \exp \left( -b_1 D_{xx} \exp \left[ -\frac{T_2}{T_2} \right] \right)$$

with

$$b_1 = \int_0^{T_2} k_x^2(t) dt = \int_0^{T_2} \left[ \int_0^t \partial_x(r) M_T \right]^2 dr$$

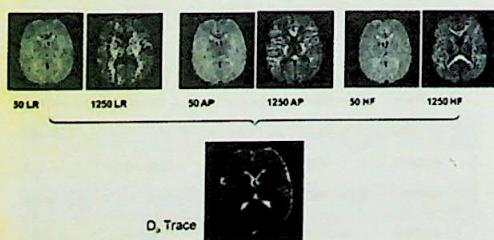
## Measurement of diffusion

### Images for different b-values

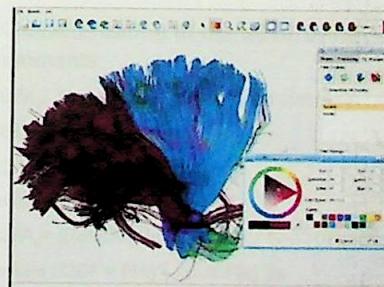


## Measurement of diffusion

### Diffusion weighted images



## Fiber Tracking in White Matter



## Multiple Fibers

Diffusion measures dispersion of water molecules within a few tens of milliseconds described by a probability function:  $p$

White matter axon radii: 0.1 - 10  $\mu\text{m}$

voxel: 1 - 5 mm

Heuristic rule  $b = \text{ADC}^{-1}$

Fiber orientation distribution function: fODF (describes the fraction of fibers in orientation)

Diffusion orientation distribution function: dODF (describes probability of diffusion in orientation)

## Multiple Fibers

Approach (model-based): Describe the dispersion by a weighted sum of gaussians:

$$p(x) = \sum_{i=1}^n a_i G(x, \mathbf{D}_i, t)$$

volume fraction      Displacement  
Gaussian with covariance      Diffusion time  
2  $D_i t$

The normalized diffusion weighted signal then is given by

$$A(\mathbf{q}) = \sum_{i=1}^n a_i \exp\left\{-\frac{1}{2} \mathbf{q}^T \mathbf{D}_i \mathbf{q}\right\}$$

wavevector in direction of grad B

E.g.  $n = 2: 2 \times 6 + 1$  ( $a_1 = 1 - a_2$ )  
parameters

## Multiple Fibers

$$p(x) = \sum_{i=1}^n a_i G(x, \mathbf{D}_i, t)$$

volume fraction      Displacement  
Gaussian with covariance      Diffusion time  
2  $D_i t$

$$A(\mathbf{q}) = \sum_{i=1}^n a_i \exp\left\{-\frac{1}{2} \mathbf{q}^T \mathbf{D}_i \mathbf{q}\right\}$$

wavevector in direction of grad B

Solution by additional constraints on the diffusion:

- Enforce positive definiteness
- Enforce cylindrical symmetry
- Fix the DT's Eigenvalues
- Ensure voxel-to-voxel coherence by spatial regularization
- Isotropic/ One fiber/ two fiber automatic model selection

## Multiple Fibers

Non parametric approaches (model-free): Distinguish fanning or bending fibers from straight ones.

Diffusion Spectrum Imaging: For infinitely short pulses we have  $p = \text{FT}(A)$ .

Q-Ball Imaging: Interpolate the dODF found by using a linear basis of spherical functions.

Spherical Deconvolution: Measure  $A$  (Diffusion weighted Signal) and  $R$  (Convolution of the measurement). The fiber orientation can be obtained by deconvolving  $A(\mathbf{q}) = \int f(x) R(\mathbf{q}, x) dx$ .

Persistent Angular Structure: Find the angular structure by a maximum entropy principle. PAS is deconvolution with  $R(\mathbf{q}, x) = r^2 \cos(r\mathbf{q}, \mathbf{x})$ .

Diffusion Orientation Transform: The DOT is a single contour of  $p$  at fixed radius  $R_0$ . It calculates a variant of the dODF.

## Multiple Fibers

Method	Acquisition requirement	Computation time	Accuracy	Bias
Two tensor	low/medium	medium	medium	low
Ball and stick	low	medium	medium	low
PAS MRI	medium	high	high	low
SD (low-pass)	medium	low/ medium	medium	medium
SD (cSD)	medium	medium	medium	low
DSI	very high	medium	high	medium
Q-Ball	medium/ high	low/ medium	medium/ low	medium
DOT	medium/ high	medium	medium	medium

## Multiple Fibers

Characterization of the dODF by its moments:

$GFA_2$ : Generalized fractional anisotropy

$GFA_3$ : Generalized skewness

$GFA_4$ : Generalized kurtosis

with moment  $n$  being given by

$$GFA_n = \frac{\int ((dODF(x) - \bar{dODF})^n dx)}{\int (dODF(x))^n dx}^{1/n}$$

where  $DODF = (4\pi)^{-1} \int dODF(x) dx$

## Fiber Tractography in White Matter

1. Brain function relies on white matter pathways.
2. Diffusion is preferred along fiber orientation in axon nerve-fibers.
3. White matter pathways can be found in the living brain by Diffusion Tractography.

## Fiber Tractography in White Matter

**Tractography Algorithms** for the reconstruction of diffusion in white matter pathways:

Local <> Global

Deterministic <> Probabilistic

Model Based <> Model Free

Simple <> Complex

## Fiber Tractography in White Matter

### Example: Streamline Tractography

A streamline is a 3D curve in space with the tangent being the first eigenvector of the diffusion tensor.

$$\frac{d\mathbf{r}}{ds} = \varepsilon_1(\mathbf{r}(s)) \quad \text{Differential eqn.}$$

The solution is based on interpolation (Runge-Kutta).

## Fiber Tractography in White Matter

Errors in Streamline Tractography can be classified in

1. Imaging noise due to poor estimation of diffusion directions
2. Modeling errors by choosing the wrong model for the voxel
3. (Numerical) integration errors

## Fiber Tractography in White Matter

The stopping problem in Streamline Tractography can be solved by

1. Minimum FA allowed
2. Maximum curvature allowed

Probabilistic Tractography can be used to avoid stopping in regions with high uncertainty:

(model, data) → Compute the x% confidence that the path of least hindrance to diffusion is from A to B.

Introduce an uncertainty Orientation Distribution Function: The uODF can be computed using various methods as bootstrapping, Bayesian methods, functional approximation and calibration.

## Fiber Tractography in White Matter

Choice of local description:

- Use many different gradient orientation (~100 000) and a complex description for the local model.

Designing a tractography study:

- Reconstruct many pathways
- Relate functional information and related white matter pathways

Future advances:

- Global tractography
- Use the whole FODF
- Combine tractography with quantitative models

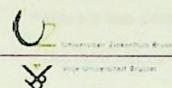
## Resume

- Random Walk and Brownian Motion
- Entropy and Cybernetics
- Flow Imaging
- Diffusion Imaging
- White Brain Matter

## fMRI of the brain



Bernd Müller-Bierl, PhD, Dipl. Phys.



## Historical approach

1936 Magnetic state of hemoglobin changes with its state of oxygenation (Pauling and Coryell)

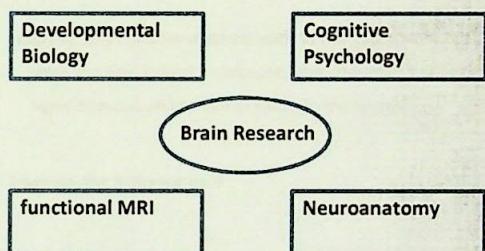
1945 Nuclear magnetic resonance effect by Purcell, Torrey and Pound, and, independently, by Bloch

1982 Transverse relaxation rate (T2) changes in blood depending on the oxygenation (Thulborn)

1990 Signal enhancement in rat brain during blood oxygen level increase (Ogawa)

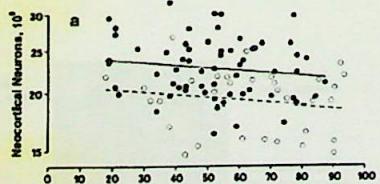
1992 fMRI signal changes during visual stimulation (Menon, Ogawa, Kim, Ellermann, Merkle, Tank, Ugurbil)

## Brain Research



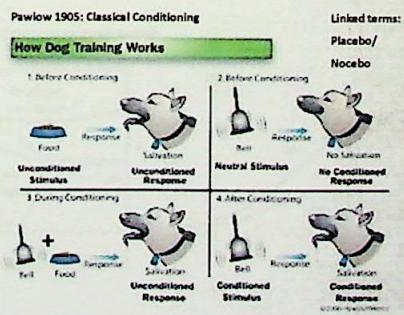
Source: Müller-Bierl (2013)

## Developmental Biology



Number of neurons in human brain, depending on age [years]. Black circles are men, white circles are women (Pakkenberg & Gundersen, THE JOURNAL OF COMPARATIVE NEUROLOGY, 1997).

## Cognitive Psychology



## Neuroanatomy

Human brain – white matter and grey matter



Source: Wikipedia

## fMRI: The BOLD effect

### Neural activity in brain

- local cerebral blood flow (CBF) increases more than the cerebral metabolic rate of oxygen (CMRO<sub>2</sub>)
- oxygen extraction fraction (E) decreases with activation of the brain
  - local blood is more oxygenated
  - there is less deoxyhaemoglobin present in nearby tissue
  - the local MR signal increases

## Susceptibility effect for GRE and SE

### Susceptibility Artifacts in Spin-Echo and Gradient-Echo Imaging

STEFAN POSSE AND WALTER P. AUE

NMR Unit of the Medical Faculty, University of Berne, Bühlerstrasse 28, 3012 Berne, Switzerland

Received August 9, 1990; revised January 4, 1990

A general model for the quantitation of spin-echo and gradient-echo artifacts in samples with macroscopic susceptibility inhomogeneities is developed. Its usefulness is tested by comparing experimental images and computer simulations of cylindrical susceptibility inhomogeneities. Positional displacements, gradient-echo phase shifts, and motion and sampling time effects, which are responsible for the signal distortions and signal losses, are analyzed. The model provides a quantitative comparison of susceptibility artifacts under various imaging conditions. The formulation will be useful for the interpretation of susceptibility-related signal losses in gradient-echo imaging and for the analysis of spin-echo susceptibility artifacts at high field strengths. In addition susceptibility effects in NMR spectroscopy and contrast properties of particulate contrast agents may be simulated. © 1990 Academic Press Inc.

## Susceptibility effect

Magnetic **susceptibility** material leads to **misregistration**, this leads to **signal intensity distortion**, that expresses itself as **signal attenuation and compression** in the image.

Example: the following slide

## Susceptibility effect for GRE and SE

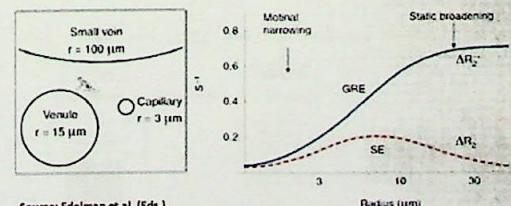
Comparison between measured and computed effect.



Posse and Aue 1990

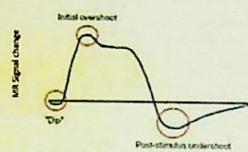
## Vessels in the brain ('Vein effect'):

Technical term: Blood Brain Barrier



## BOLD-model ('Brain effect')

**Balloon model (Buxton 1998):** The venous compartment is treated as a distensible balloon.



A typical BOLD response for a block design stimulus. Transients are an occasional brief signal dip at the beginning, an initial overshoot, and a prolonged post-stimulus undershoot.

Source: Edelman et al. (Eds.)

## BOLD-model

[Density]: The total venous blood content is given by the variable  $q(t) = V(t) \cdot \rho_0$

[Volume]: The total volume is given by the variable  $v(t) = V(t) \cdot \rho_0$

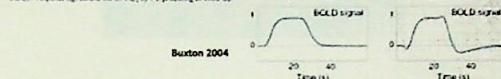
[Flow]:  $f_{in}$  is passed to the model.  $f_{out}(t) = f_{out}(v(t))$  is also passed to the model

[BOLD]: The BOLD signal is obtained

$$\Delta B = k_1 \left[ f_{in}(t) - q(t) - k_2 (t - \tau) + k_3 (1 - v) \right]$$

where  $k_1 = 2.8$ ,  $k_2 = 2$  and  $k_3 = 0.6$  for  $B_0 = 1.5T$ ,  $H_0 = 10ms$

Deriving curves for CBF, CMRO<sub>2</sub>, CBF and the resulting BOLD signal showing a typical response in which the physiology associated with a stimulus increases, and signal decreases due to a CBF change that is slow to respond, creating a post-stimulus undershoot. In addition, the CBF response lags behind the CMRO<sub>2</sub> by 1 s, producing an initial dip.



Buxton 2004

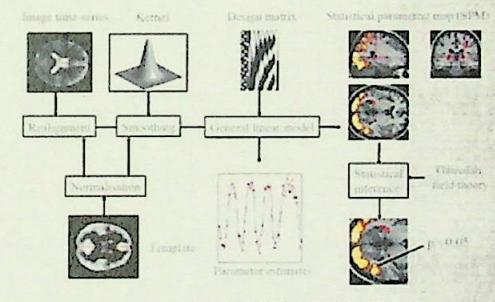
## Activation maps

We need to do subtraction measurement on stimulated and non-stimulated brain.

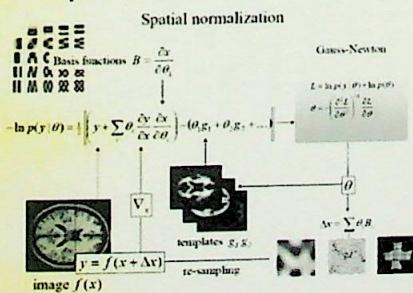
How to compute activation maps from the subtraction measurement?

By statistical evaluation of the data

## Data analysis



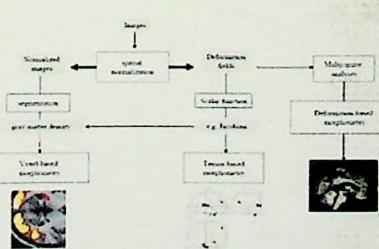
## Spatial Normalization



Friston, Experimental Design and Statistical Parametric Mapping

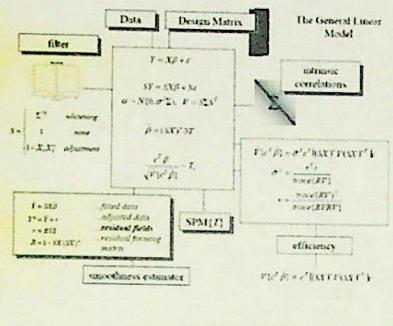
## Computational Neuroanatomy

### Computational neuroanatomy



Friston, Experimental Design and Statistical Parametric Mapping

## General Linear Model



Thank you very much!

