



Introduction to EEG/MEG Source Estimation

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The Basic Problem

What is “the” solution to:

$$x + y = 1$$

If you are not shocked by the EEG/MEG inverse problem...

... then you haven't understood it yet.

(freely adapted from Niels Bohr)

What Can We Hope For?

A rough estimate of spatial resolution:

With n sensors:

-> n independent measurements

-> at best separate activity from n brain regions

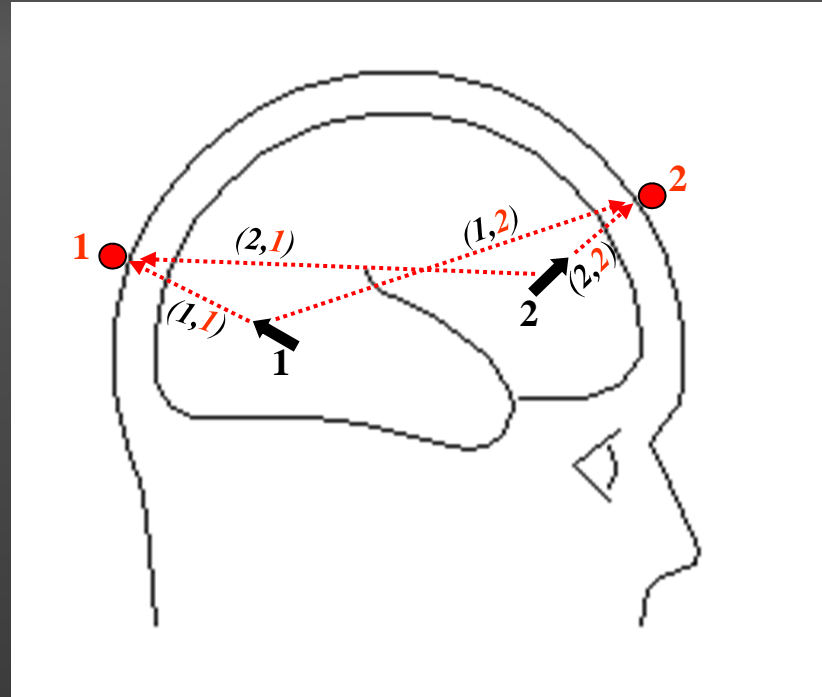
Sensors are not independent -> ~ 50 degrees of freedom

Volume of source space:

Sphere 8cm minus sphere 4 cm: volume $\sim 5600 \text{ cm}^3$

“Resel”: $113 \text{ cm}^3 \rightarrow \underline{4.8}^3 \text{ cm}^3$

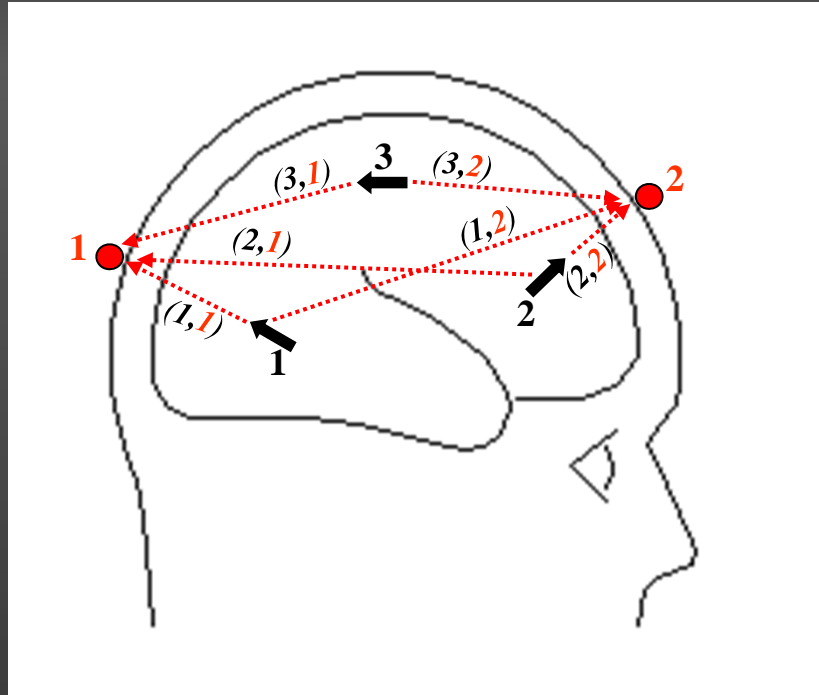
Uniquely Solvable Problem



$$\begin{array}{c}
 \text{data} \quad \text{"leadfield"} \quad \text{dipoles} \\
 \begin{matrix} 1 \\ \bullet \end{matrix} \begin{pmatrix} d_1 \\ d_2 \end{pmatrix} = \begin{pmatrix} 0.5 & 0 \\ 0 & 1 \end{pmatrix} \begin{pmatrix} j_1 \\ j_2 \end{pmatrix} \begin{matrix} \swarrow 1 \\ \nearrow 2 \end{matrix} \\
 \begin{matrix} \bullet 2 \end{matrix}
 \end{array}
 \xrightarrow{\text{inversion}}
 \begin{array}{c}
 \text{dipoles} \quad \text{inverse} \quad \text{data} \\
 \begin{matrix} \swarrow 1 \\ \nearrow 2 \end{matrix} \begin{pmatrix} \hat{j}_1 \\ \hat{j}_2 \end{pmatrix} = \begin{pmatrix} 2 & 0 \\ 0 & 1 \end{pmatrix} \begin{pmatrix} d_1 \\ d_2 \end{pmatrix} \begin{matrix} 1 \\ \bullet 2 \end{matrix}
 \end{array}$$

Assume dipoles 1 and 2 are only visible to electrodes 1 and 2, respectively.

Non-Uniquely Solvable Problem



“Minimum Norm Solution”

$$\begin{matrix} \bullet \\ \bullet \end{matrix} \begin{pmatrix} d_1 \\ d_2 \end{pmatrix} = \begin{pmatrix} 0.5 & 0 & 0.3 \\ 0 & 1 & -0.3 \end{pmatrix} \begin{pmatrix} j_1 \\ j_2 \\ j_3 \end{pmatrix} \begin{matrix} \leftarrow 1 \\ \nearrow 2 \\ \leftarrow 3 \end{matrix}$$

?
inversion

$$\begin{matrix} \leftarrow 1 \\ \nearrow 2 \\ \leftarrow 3 \end{matrix} \begin{pmatrix} j_1 \\ j_2 \\ j_3 \end{pmatrix} = \begin{pmatrix} 1.5034 & 0.1241 \\ 0.2483 & 0.9379 \\ 0.8276 & -0.2069 \end{pmatrix} * \begin{pmatrix} d_1 \\ d_2 \end{pmatrix} \begin{matrix} \bullet \\ \bullet \end{matrix}$$

Non-Uniqueness

Non-Unique

$$\begin{array}{c} \bullet^1 \\ \bullet^2 \end{array} \begin{array}{c} \text{data} \\ \begin{pmatrix} 1 \\ 1 \end{pmatrix} \end{array} = \begin{array}{c} \text{"leadfield"} \\ \begin{pmatrix} 0.5 & 0 & 0.3 \\ 0 & 1 & -0.3 \end{pmatrix} \end{array} * \begin{array}{c} \text{dipoles} \\ \begin{pmatrix} j_1 \\ j_2 \\ j_3 \end{pmatrix} \end{array} \begin{array}{c} \swarrow 1 \\ \nearrow 2 \\ \leftarrow 3 \end{array}$$

?

inversion

"Minimum norm solution:"

$$\begin{array}{c} \swarrow 1 \\ \nearrow 2 \\ \leftarrow 3 \end{array} \begin{array}{c} \text{dipoles} \\ \begin{pmatrix} 1.62 \\ 1.18 \\ 0.62 \end{pmatrix} \end{array} = \begin{array}{c} \text{inverse} \\ \begin{pmatrix} 1.5034 & 0.1241 \\ 0.2483 & 0.9379 \\ 0.8276 & -0.2069 \end{pmatrix} \end{array} * \begin{array}{c} \text{data} \\ \begin{pmatrix} 1 \\ 1 \end{pmatrix} \end{array} \begin{array}{c} \bullet^1 \\ \bullet^2 \end{array}$$

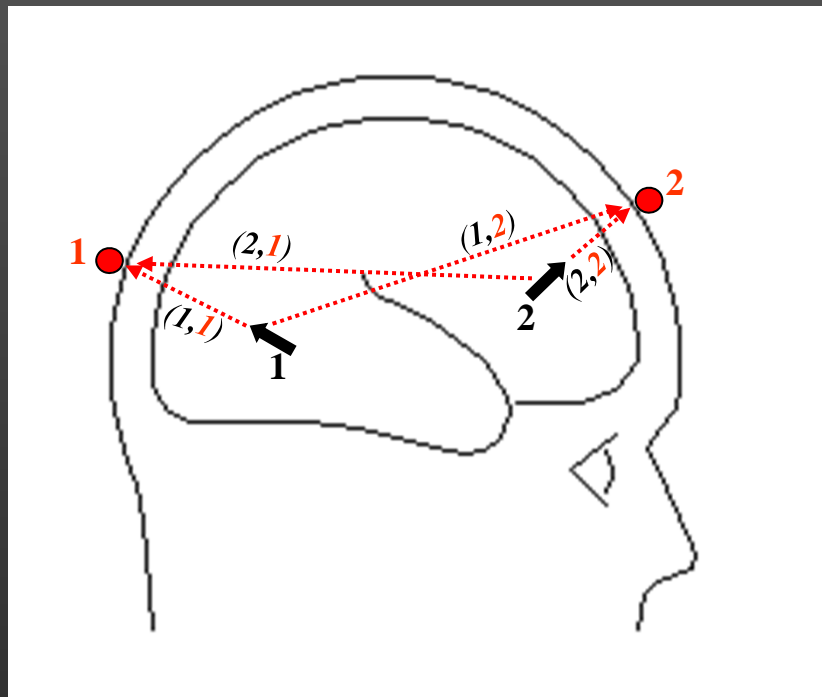
$$\text{or } \begin{pmatrix} 2 \\ 1 \\ 0 \end{pmatrix} \quad \text{or } \begin{pmatrix} 1.81 \\ 1.09 \\ 0.31 \end{pmatrix}$$

are also possible solutions
that fit the data exactly –

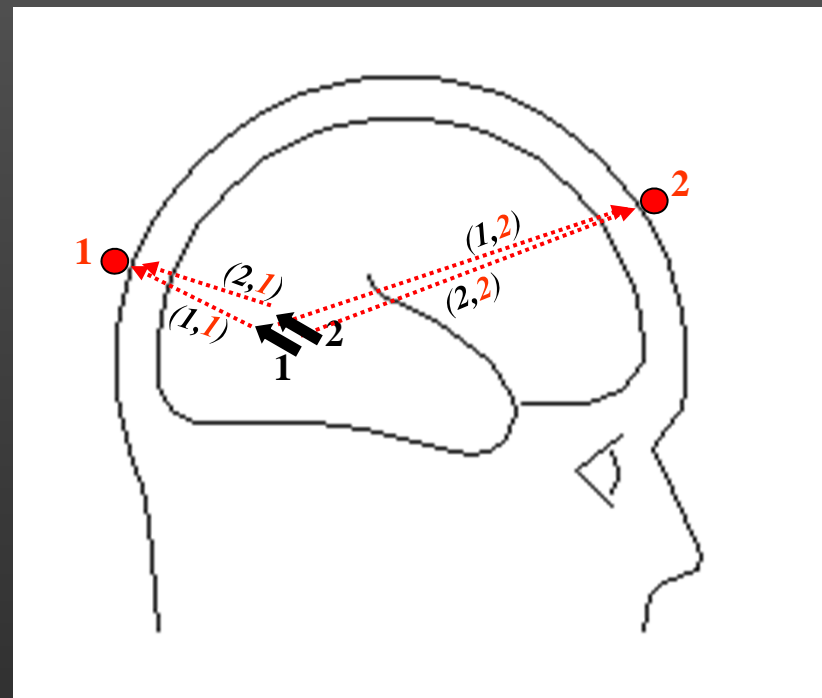
there is no "better" or "worse" solution
Solely on mathematical grounds.

(In)Stability - Sensitivity to Noise

Stable



Instable



(In)Stability - Sensitivity to Noise

Stable

$$\begin{array}{c} \text{data} \quad \text{"leadfield"} \quad \text{dipoles} \\ \begin{pmatrix} d_1 \\ d_2 \end{pmatrix} = \begin{pmatrix} 0.5 & 0 \\ 0 & 1 \end{pmatrix} \begin{pmatrix} j_1 \\ j_2 \end{pmatrix} \end{array}$$

inversion

$$\begin{array}{c} \text{dipoles} \quad \text{inverse} \quad \text{data} \\ \begin{matrix} \nwarrow 1 \\ \nearrow 2 \end{matrix} \begin{pmatrix} \hat{j}_1 \\ \hat{j}_2 \end{pmatrix} = \begin{pmatrix} 2 & 0 \\ 0 & 1 \end{pmatrix} \begin{pmatrix} d_1 \\ d_2 \end{pmatrix} \begin{matrix} \bullet^1 \\ \bullet^2 \end{matrix} \end{array}$$

Instable

$$\begin{array}{c} \text{data} \quad \text{"leadfield"} \quad \text{dipoles} \\ \begin{pmatrix} d_1 \\ d_2 \end{pmatrix} = \begin{pmatrix} 0.001 & 1 \\ 0 & 1 \end{pmatrix} \begin{pmatrix} j_1 \\ j_2 \end{pmatrix} \end{array}$$

inversion

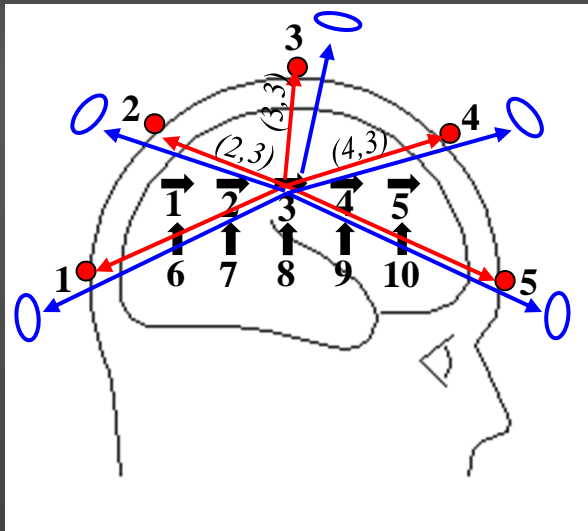
$$\begin{array}{c} \text{dipoles} \quad \text{inverse} \quad \text{data} \\ \begin{pmatrix} \hat{j}_1 \\ \hat{j}_2 \end{pmatrix} = \begin{pmatrix} 1000 & -1000 \\ 0 & 1 \end{pmatrix} \begin{pmatrix} d_1 \\ d_2 \end{pmatrix} \end{array}$$

Addressed by "Regularisation" ("lambda"):

Add smoothness constraint to solution, at the expense of spatial resolution

Recommended to check SNR in source space, "sanity checks"

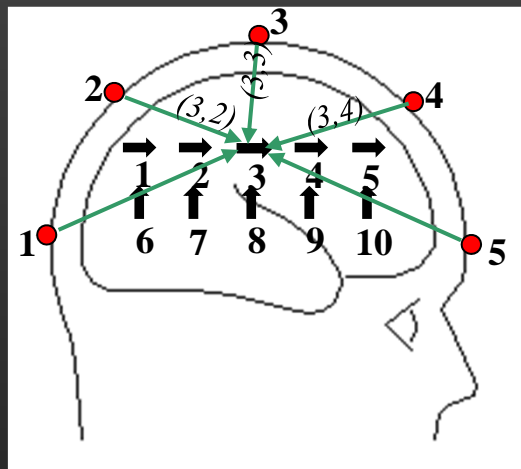
"Forward" and "Inverse" problem



Forward Problem

$$\begin{pmatrix} d_1 \\ d_2 \\ d_3 \\ d_4 \\ d_5 \end{pmatrix} = \mathbf{d} = \sum_{i=1}^{N_s=10} s_i \begin{pmatrix} L_{1i} \\ L_{2i} \\ L_{3i} \\ L_{4i} \\ L_{5i} \end{pmatrix} = \sum_{i=1}^{N_s=10} s_i \mathbf{L}_{\cdot i} = \mathbf{L}\mathbf{s}$$

“Leadfield” matrix
Forward Problem



Inverse Problem

$$\begin{pmatrix} s_1 \\ s_2 \\ s_3 \\ s_4 \\ s_5 \\ s_6 \\ s_7 \\ s_8 \\ s_9 \\ s_{10} \end{pmatrix} = \mathbf{s} = \sum_{i=1}^{N_i=5} d_i \begin{pmatrix} G_{1i} \\ G_{2i} \\ G_{3i} \\ G_{4i} \\ G_{5i} \\ G_{6i} \\ G_{7i} \\ G_{8i} \\ G_{9i} \\ G_{10i} \end{pmatrix} = \sum_{i=1}^{N_i=5} d_i \mathbf{G}_{\cdot i} = \mathbf{G}\mathbf{d}$$

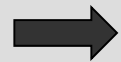
“Inverse” matrix
“Spatial Filter”

Minimum Norm Estimation: Minimal Modelling Assumptions

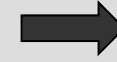
“No frills” solution (Minimum Norm)

$$(\hat{\mathbf{s}} - \hat{\mathbf{s}}_0)^T \mathbf{C}_s (\hat{\mathbf{s}} - \hat{\mathbf{s}}_0) = \min$$

$$(\mathbf{L}\hat{\mathbf{s}} - \mathbf{d})^T \mathbf{C}_d (\mathbf{L}\hat{\mathbf{s}} - \mathbf{d}) = \varepsilon > 0$$



$$\hat{\mathbf{s}} = \hat{\mathbf{s}}_0 + \mathbf{C}_s^{-1} \mathbf{L}^T (\mathbf{L} \mathbf{C}_s^{-1} \mathbf{L}^T + \lambda \mathbf{C}_d^{-1})^{-1} (\mathbf{d} - \mathbf{L} \hat{\mathbf{s}}_0)$$



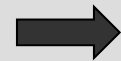
“Minimum Least-Squares Solution”

$$\hat{\mathbf{s}} = \mathbf{L}^T (\mathbf{L} \mathbf{L}^T + \lambda \mathbf{I})^{-1} \mathbf{d}$$

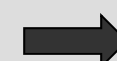
“Most likely” solution (Maximum Likelihood)

$$\mathbf{P}(\mathbf{s}) \sim \exp\{-\hat{\mathbf{s}} - [\mathbf{s}]^T \mathbf{C}_s (\hat{\mathbf{s}} - [\mathbf{s}])\}$$

$$\mathbf{P}(\mathbf{d}, \hat{\mathbf{s}}) \sim \exp\{-\mathbf{d} - \mathbf{L}\hat{\mathbf{s}})^T \mathbf{C}_d (\mathbf{d} - \mathbf{L}\hat{\mathbf{s}})\}$$



$$\hat{\mathbf{s}} = [\mathbf{s}] + \mathbf{C}_s^{-1} \mathbf{L}^T (\mathbf{L} \mathbf{C}_s^{-1} \mathbf{L}^T + \lambda \mathbf{C}_d^{-1})^{-1} (\mathbf{d} - \mathbf{L}[\mathbf{s}])$$

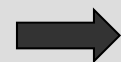


$$\hat{\mathbf{s}} = \mathbf{L}^T (\mathbf{L} \mathbf{L}^T + \lambda \mathbf{I})^{-1} \mathbf{d}$$

“Best focussing” solution (Beamformer)

$$\text{Min}(\mathbf{W}(\mathbf{r}_i - \mathbf{t}_i))^2$$

$$\text{Min}([\mathbf{G}_i \mathbf{n}]^2) \Rightarrow \text{Min}(\mathbf{G}_i \mathbf{C}_n \mathbf{G}_i^T)$$



$$\mathbf{G}_i = (\mathbf{S} + \lambda \mathbf{C}_n)^{-1} \mathbf{u}$$

$$\mathbf{S} = \mathbf{L} \mathbf{L}^T \quad \mathbf{u} = \mathbf{L}_i$$

$$\mathbf{G}_i = (\mathbf{L} \mathbf{L}^T + \lambda \mathbf{I})^{-1} \mathbf{L}_i$$



$$\hat{\mathbf{s}} = \mathbf{L}^T (\mathbf{L} \mathbf{L}^T + \lambda \mathbf{I})^{-1} \mathbf{d}$$

Under the same modelling assumptions,
different approaches converge to the same solution:

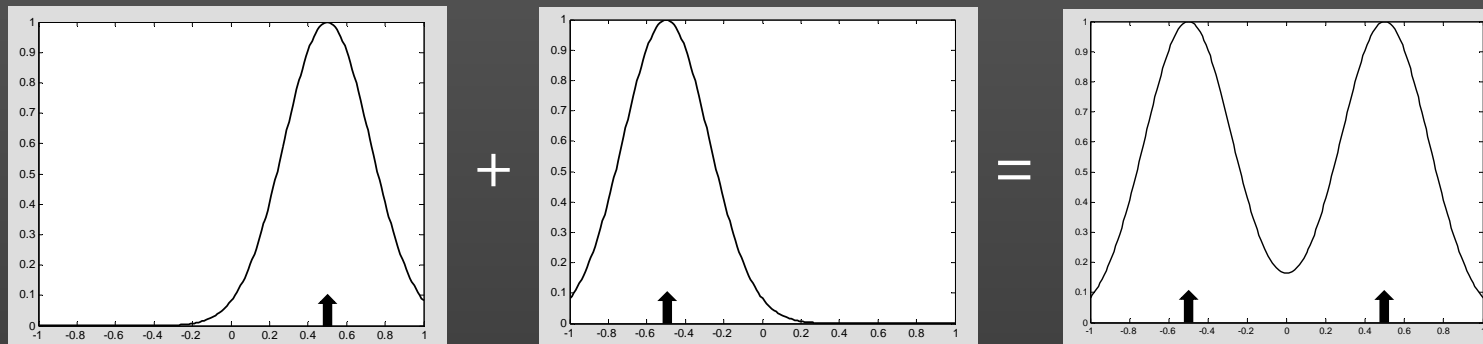
“minimum-norm least-squares” (MNLS), “minimum norm estimate” (MNE)

Advantages of Linear Distributed Solutions

Standard in related areas of signal processing and parameter estimation
("General Linear Model")

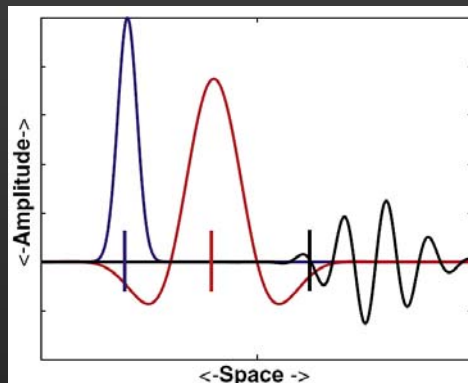
⇒ well-developed theory based on matrix algebra

(Relatively) easy to evaluate, allows generalisable conclusions



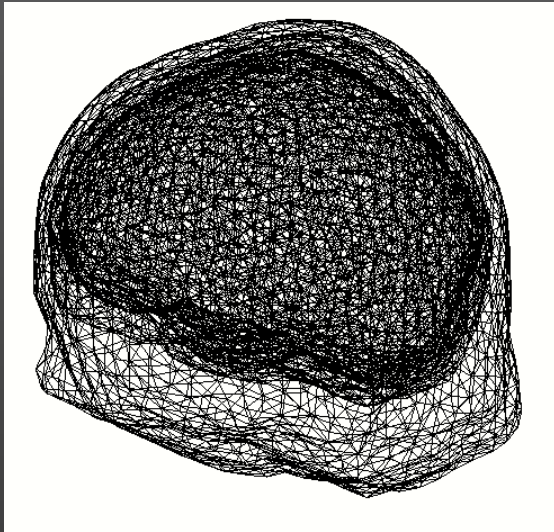
The evil you can evaluate is better than the evil you cannot evaluate

What's the worst than can happen?

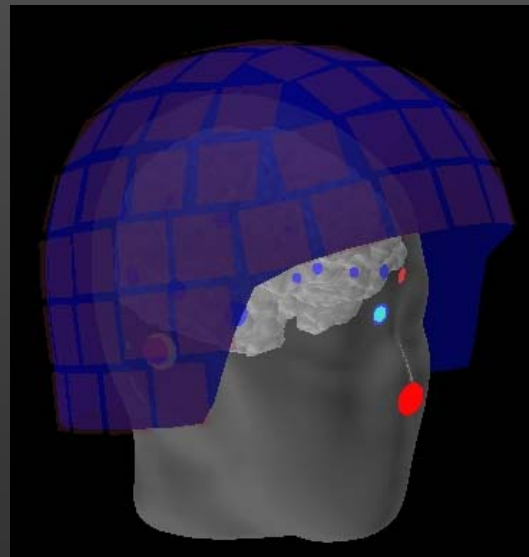


Ingredients for Source Estimation

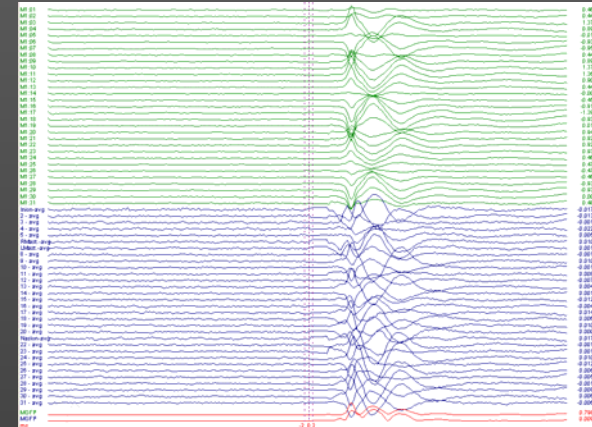
Volume Conductor/
Head Model



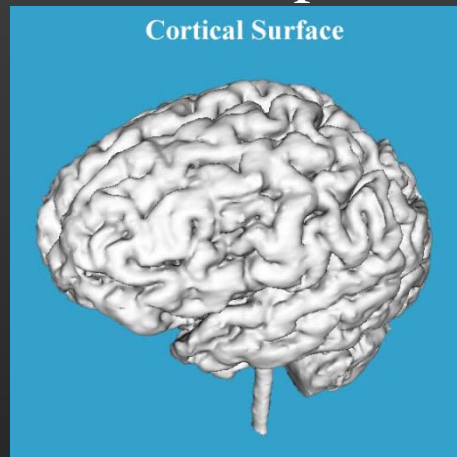
Coordinate
Transformation



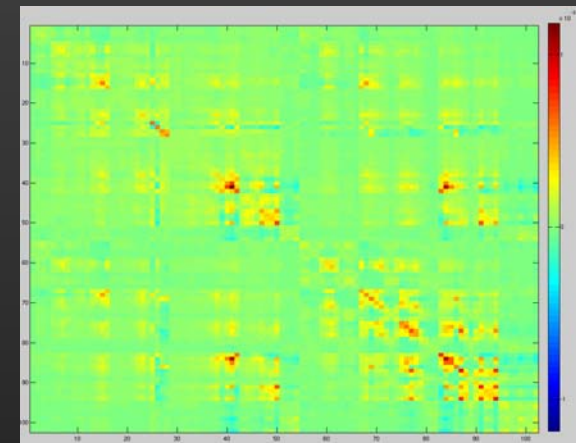
MEG data



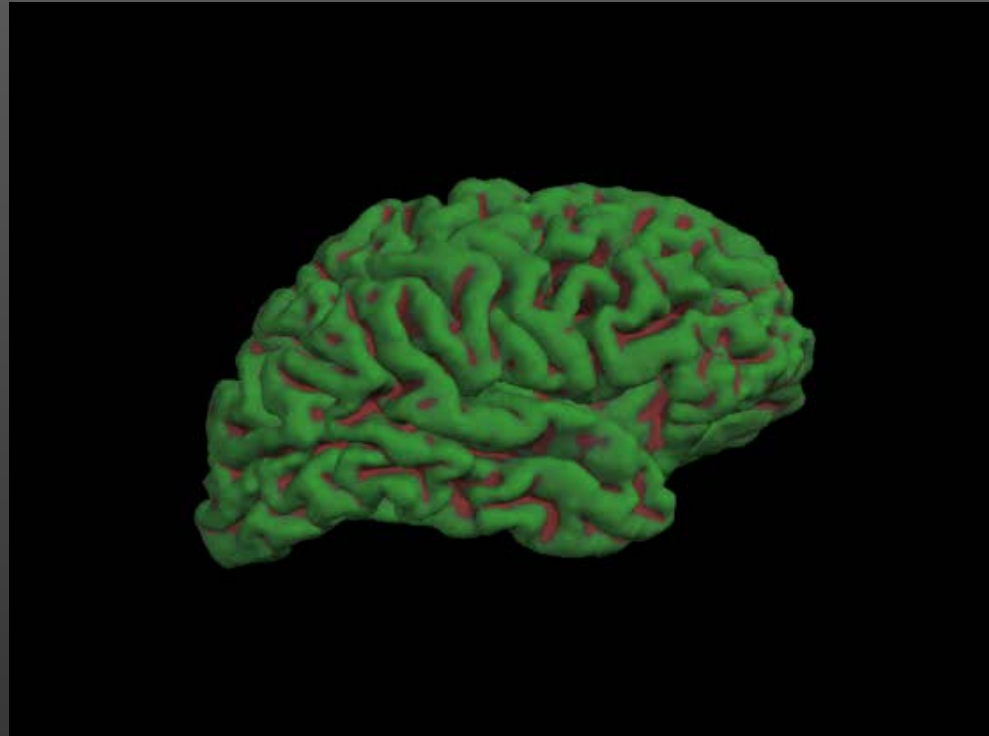
Source Space



Noise/Covariance Matrix



Inflated Cortical Surface



<http://www.cogsci.ucsd.edu/~sereno/movies.html>

Spatial Resolution of Source Estimation

Spatial resolution depends on:

modeling assumptions

number of sensors (EEG/MEG or both)

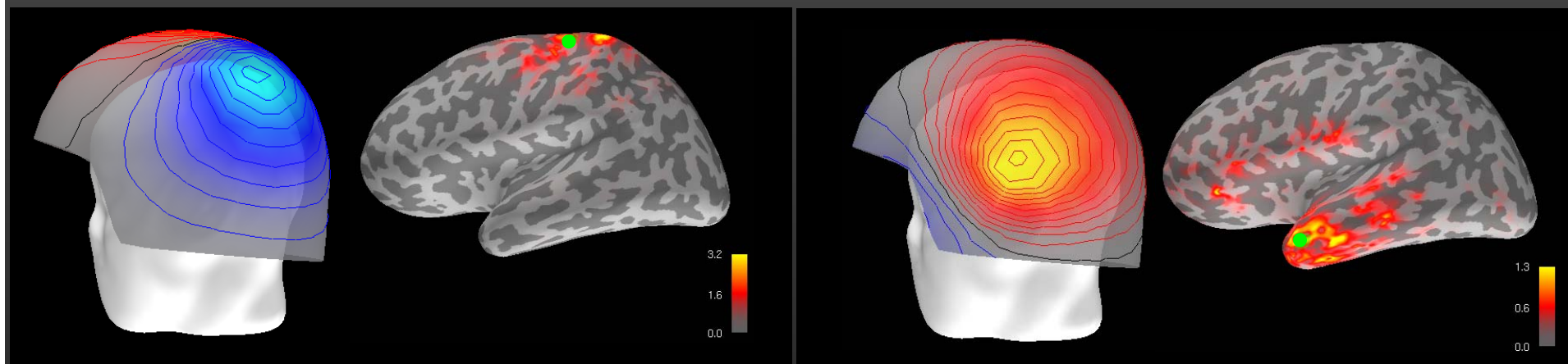
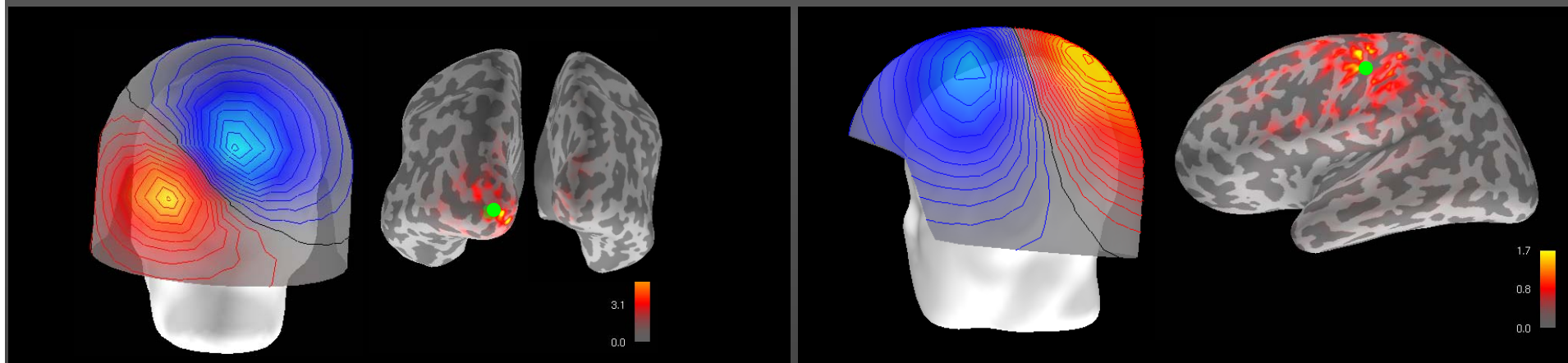
source location

source orientation

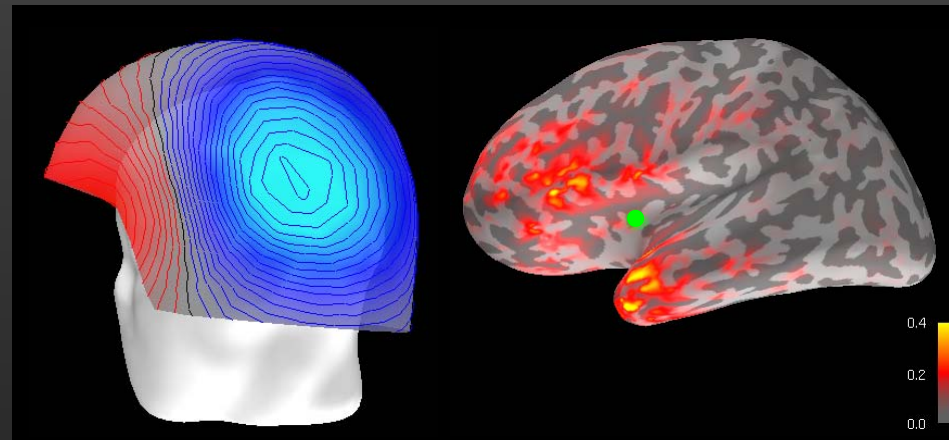
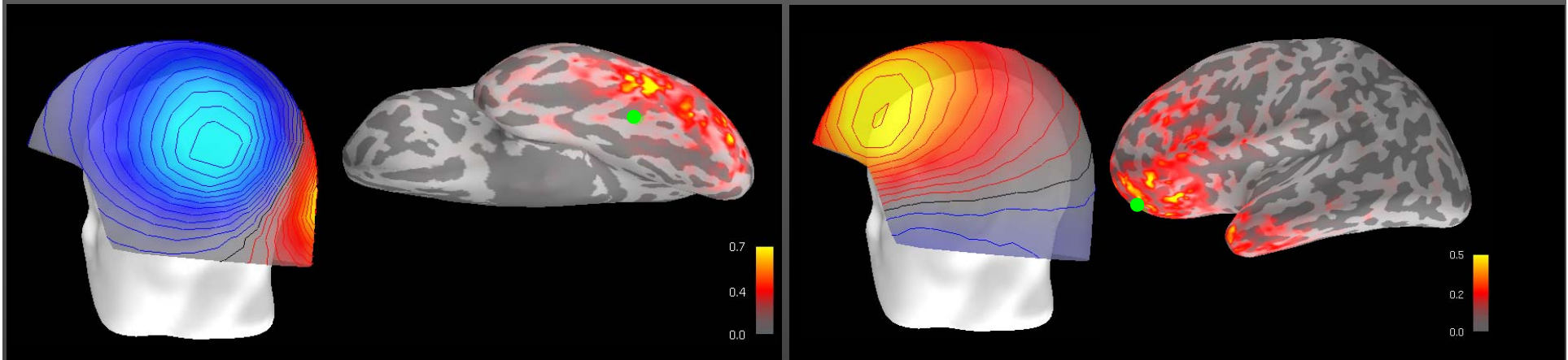
signal-to-noise ratio

head modeling

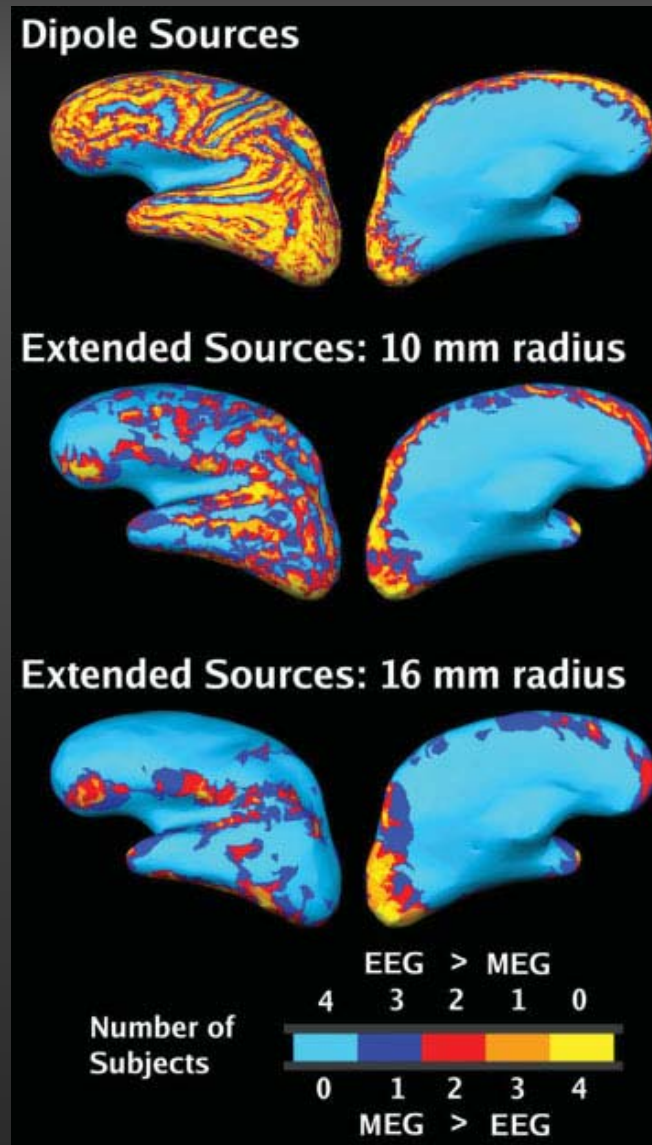
Localisation for Some ROIs



Localisation for Some ROIs



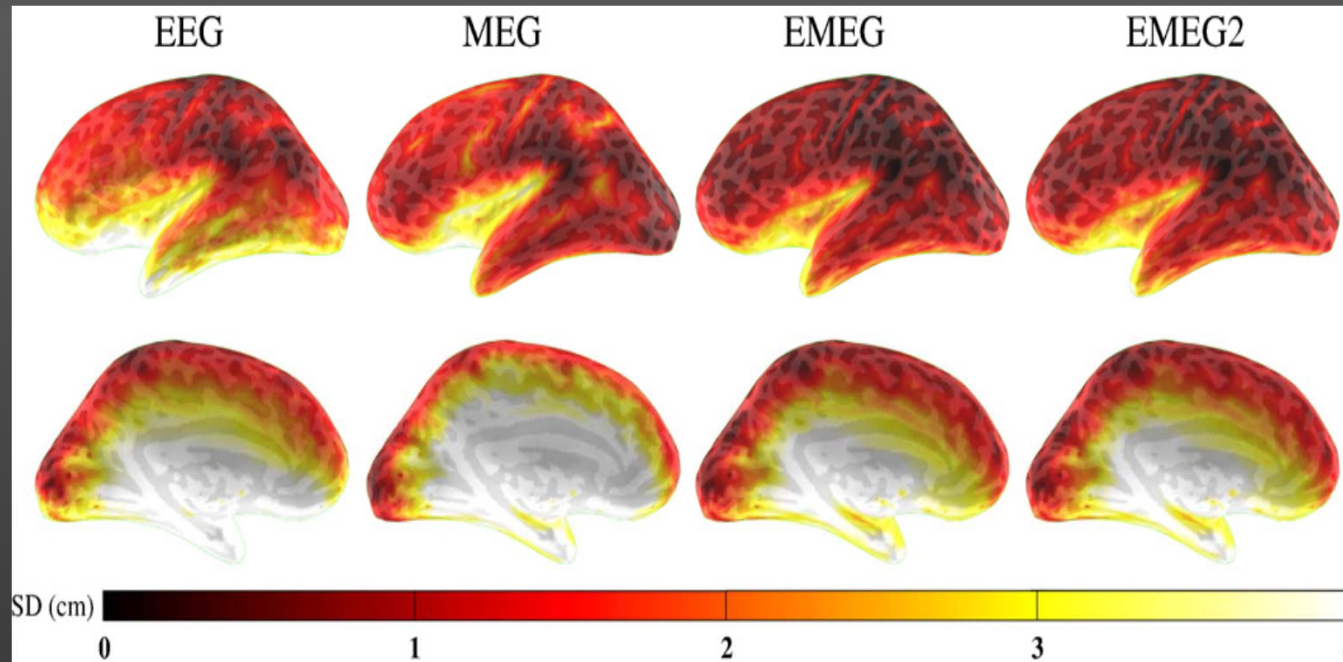
Combining EEG and MEG Increases Sensitivity



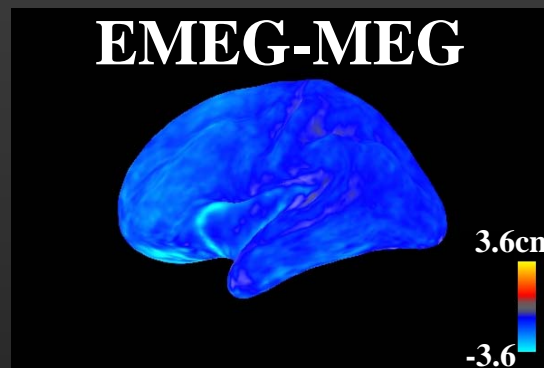
EEG is more sensitive to spatially extended sources

Combining EEG and MEG Improves Resolution

Spatial Extent

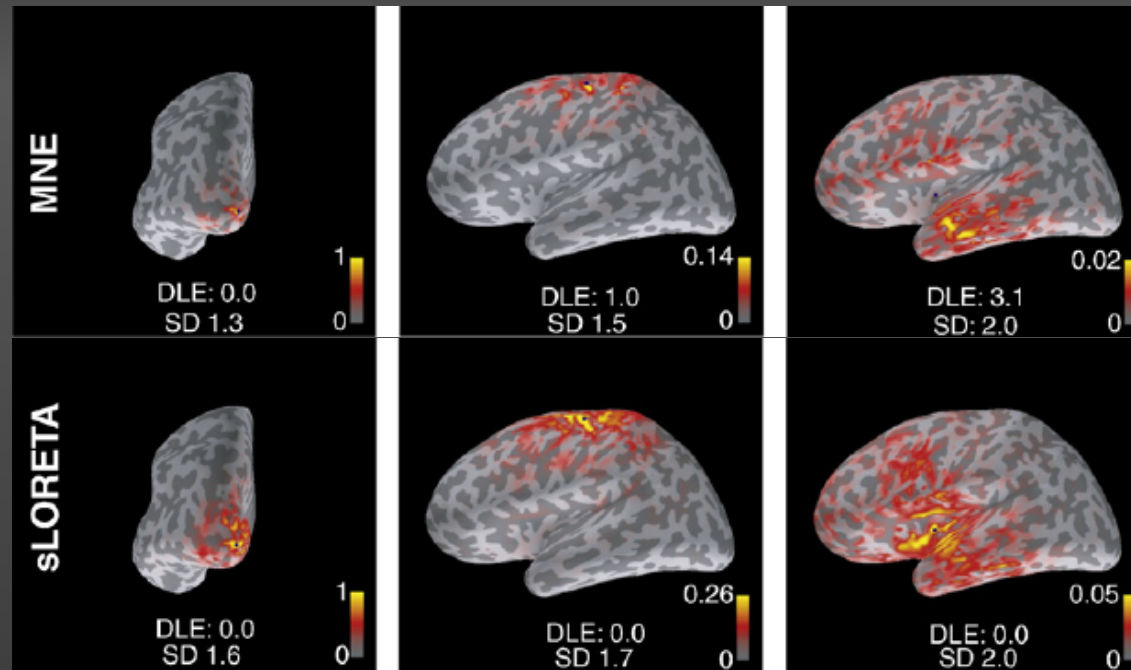


Molins et al., Neuroimage 2008



Stenroos&Hauk, in prep

Source Estimation



“zero dipole localisation error”

Different methods make different compromises

There is no “best” method – best for what?

Source Estimation Approaches

“Dipole Fitting”

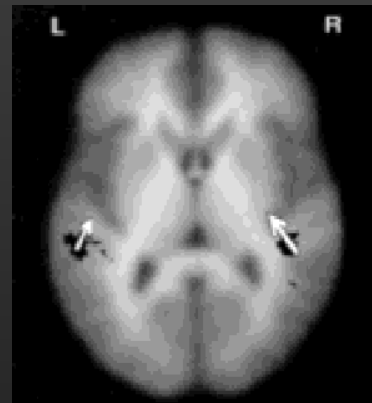
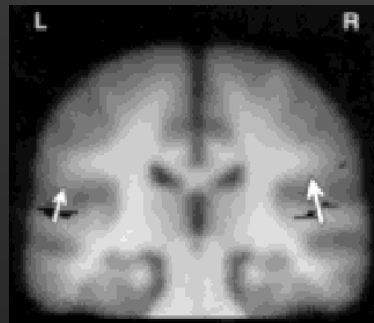
1. Assume there are only a few distinct sources
2. Iteratively adjust the location, orientation and strength of a few dipoles...
3. ...until the result best fits the data

Critical parameter:

Residual Variance or Goodness-of-Fit
(ideally taking into account degrees of freedom of the model)

Good for:

**Hypothesis testing or
precise localisation of well-known
sources**



Source Estimation Approaches

“Beamforming”

1. Create spatial filter that projects maximally on source of interest...

No GOF measure

Spatial resolution difficult to evaluate since estimator is data-dependent

2. ...while minimally projecting on data covariance matrix (incl. signal and noise covariance)

Not suitable when source topographies or time courses highly correlated

3. “Dipole scan”

Applications for spontaneous brain activity (resting state, oscillations), but difficult to justify for evoked responses