





# Practical intensity-based meta-analysis

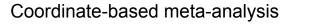


Image-based meta-analysis



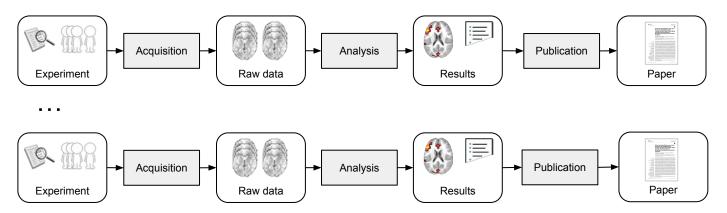
#### **Camille Maumet**

#### **Presented by Thomas Nichols**

OHBM Neuroimaging Meta-Analysis Educational course June 17<sup>th</sup> 2018

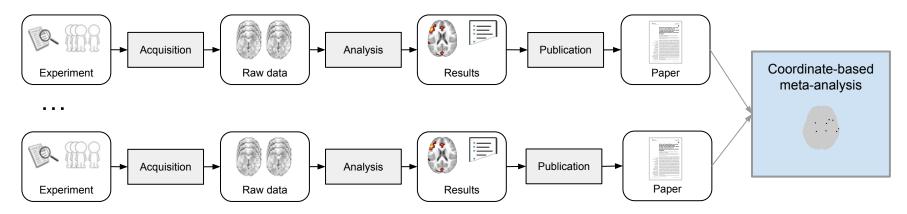
# Coordinate-Based & Image-Based Meta-Analyses

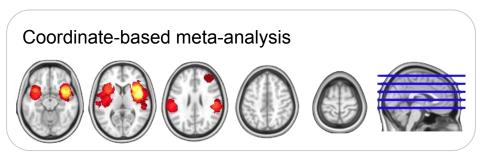
### Neuroimaging meta-analyses



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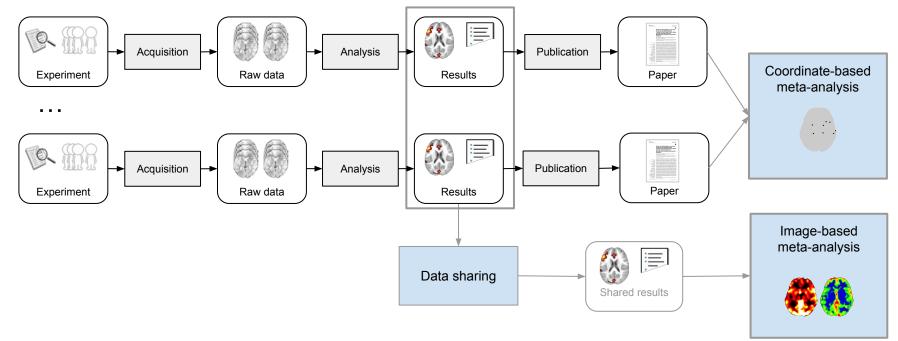
# Neuroimaging meta-analyses

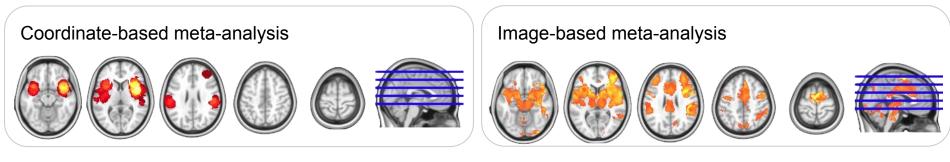




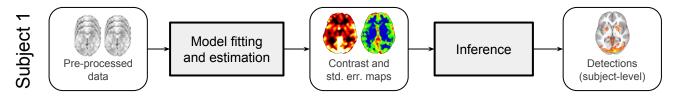
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# Neuroimaging meta-analyses

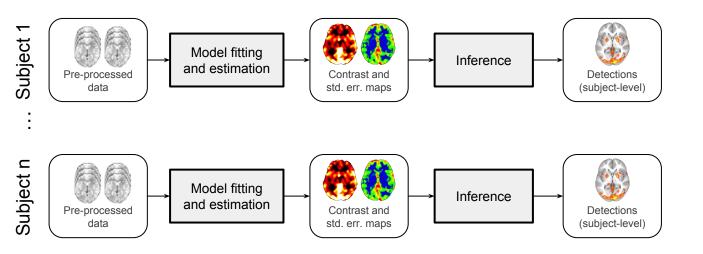


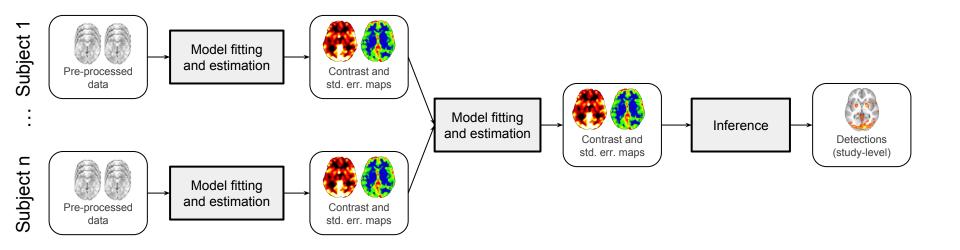


# How to perform an image-based meta-analysis?

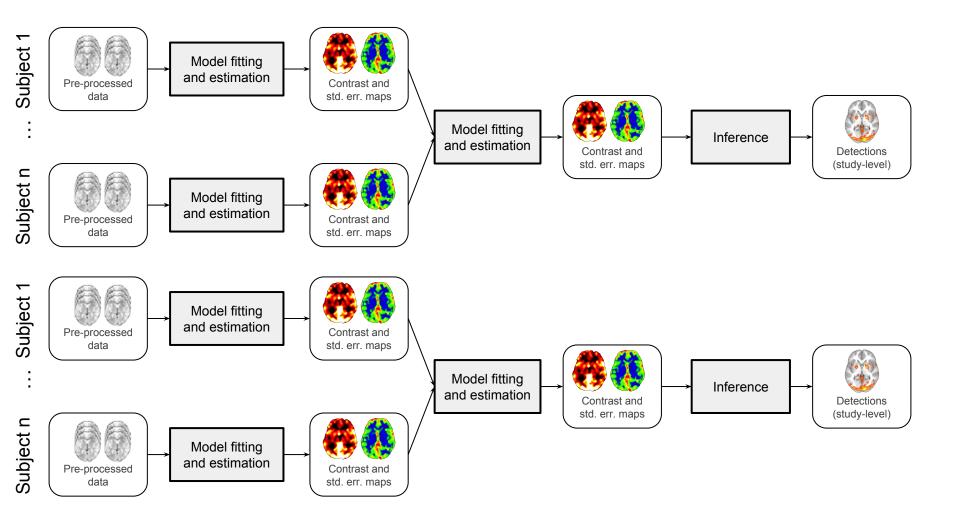


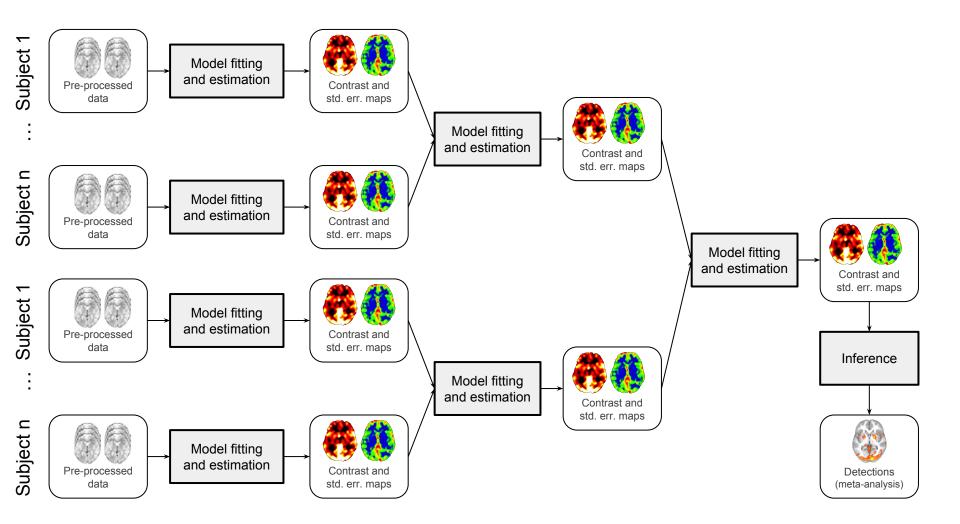
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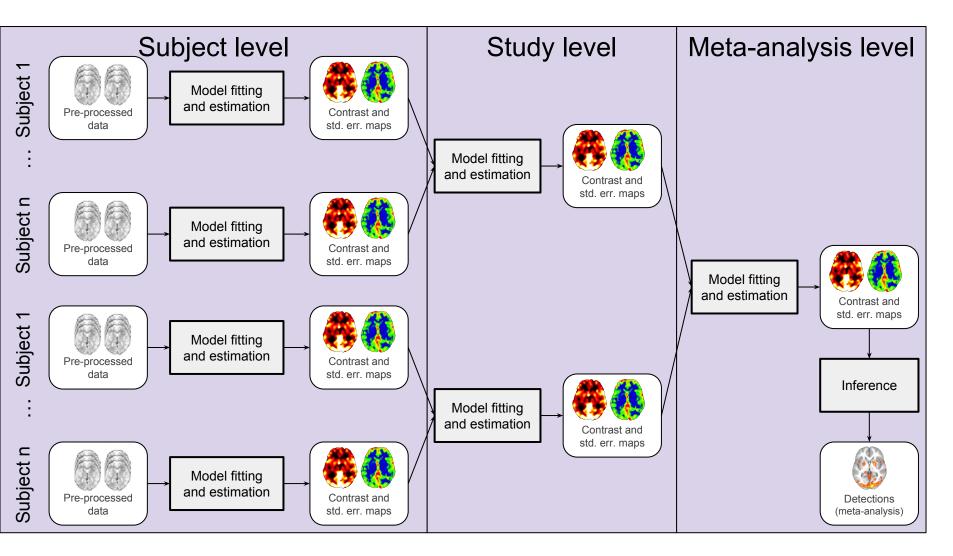




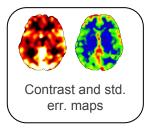
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• Gold standard:

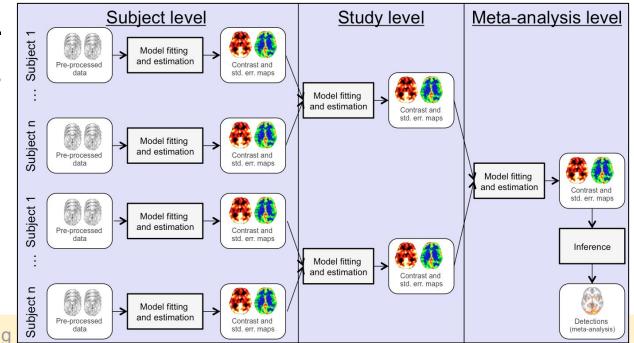


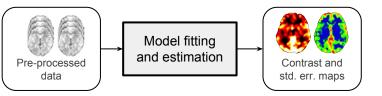
#### **Third-level Mixed-Effects GLM**

- Requirements
  - study-level Contrast estimates and Standard

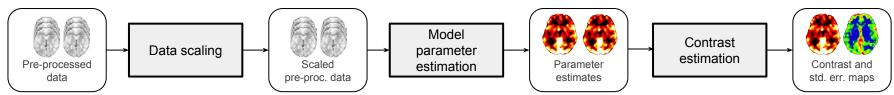
error maps.

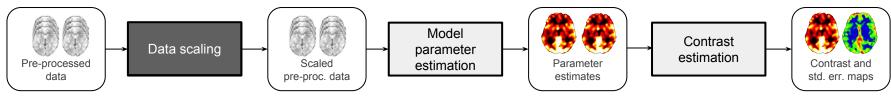
– Same **units** 



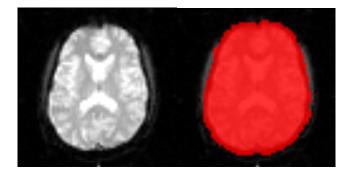


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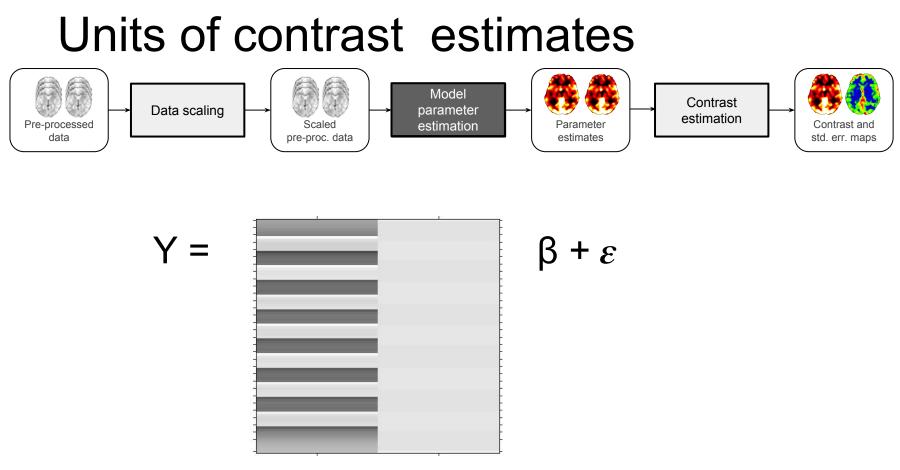


$$scaled_data = \frac{data * target}{est_mean}$$

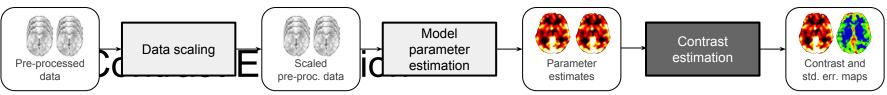


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# Units depend on **mean estimation** and **scaling target**.



# Units depend on scaling of explanatory variables



- Linear combination of parameter estimates
- Final statistics invariant to scale
  - e.g. [1111] gives same T's & P's as [1/4 1/4 1/4 1/4]

#### Units depend on contrast vector

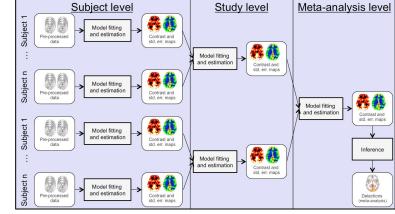
#### Rule for contrasts to preserve units

- Positive elements sum to 1
- Negative elements sum to -1

• Gold standard:

**Third-level Mixed-Effects GLM** 

- But...
  - Units will depend on:
    - The scaling of the data (subject-level)
    - The scaling of the predictor(s) (subject- and study-level)
    - The scaling of the contrast (subject- and study-level).
  - Contrast estimates and standard error maps are rarely shared...



# Which images for IBMA?

SPM

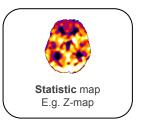




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Contrast & std. err. maps	con_0001.nii [SPM.mat]	cope1.nii varcope1.nii ( <i>squared</i> )	3dMEMA_result+tlrc.BRIK[[0]] [from contrast & stat maps]
Statistic map E.g. Z-map	spmT_0001.nii	tstat1.nii.gz zstat1.nii.gz	3dMEMA_result+tlrc.BRIK[[1]]
Contrast map	con_0001.nii	cope1.nii	3dMEMA_result+tlrc.BRIK[[0]]

# IBMA on Z maps



- Fisher's  $-2\sum_{k}\log P_k \sim \chi^2_{2k}$ 
  - Sum of -log P-values (from T/Z's converted to P's)
- Stouffer's  $\sqrt{K} \times \frac{1}{K} \sum_{k} Z_{k} \sim \mathcal{N}(0, 1)$

- Average Z, rescaled to N(0,1)

"Stouffer's Random Effects (RFX)"

$$\sqrt{K} \times \frac{1}{K} \sum_{k} Z_k \sim \mathcal{N}(0, \sigma_{RFX}^2)$$

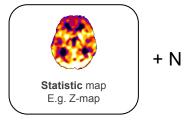
Submit Z's to one-sample t-test

(Slide adapted from Thomas Nichols, OHBM 2015)

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# IBMA on Z maps + N



Weighted Stouffer's

$$\sum_{k} w_k Z_k \sim \mathcal{N}(0, 1), \quad w_k \propto \sqrt{N_k}$$

- Z's from bigger studies get bigger weights

(Slide adapted from Thomas Nichols, OHBM 2015)

# IBMA on Contrast maps

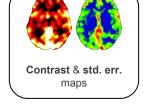


• Random Effects (RFX) GLM  $\frac{1}{K} \sum_{k} c \hat{\beta}_{k} \sim \mathcal{N}(0, \sigma_{\text{RFX}}^{2})$ 

- Analyze per-study contrasts as "data"

# Contrast + standard error maps

• Fixed-Effects (FFX) GLM



(Slide adapted from Thomas Nichols, OHBM 2015)

 $\frac{1}{K} \sum \hat{\theta}_k \sim \mathcal{N}(0, \sum \sigma_{\text{FFX},k}^2 / K^2)$ 

### Implementations

Not all of these options are easily used

Meta-Analysis Method	Inputs	Neuroimaging Implementation
'Gold Standard' MFX	Con's + SE's	FSL'S FEAT SPM spm_mfx AFNI 3dMEMA
RFX GLM Stouffer's RFX	Con's Z's	FSL, SPM, AFNI, etc
FFX GLM Fisher's Stouffer's Stouffer's Weighted	Con's +SE's Z's Z's Z's + N's	n/a

(Slide from Thomas Nichols, OHBM 2015)

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# Self Promotion Alert: IBMA toolbox

- SPM Extension
- Still in beta!
  - But welcome all feedback

NeuroimagingMetaAnaly	sis / <b>ibma</b>	
Image-Based Meta-Analysis toolbox	c — Edit	
25 commits	🖗 <b>1</b> branch	<b>0</b> release
👔 🕼 branch: master 🗸 ibma / +		
Config definition for contrast variance input	t	
Cmaumet authored on Feb 13		
example	example: simulations using IBM	A
in test	Rename stouffer	
.gitignore	Update README and gitignore	
README.md	Update README and gitignore	
ibma_config_analysis_dir.m	Mega-analysis (with FFX at third	l level)
ibma_config_contrast_files.m	Mega-analysis FFX-OLS	
ibma_config_fishers.m	fix:typo	
ibma_config_mega_ffx.m	Config definition for contrast vari	iance input
ibma_config_mega_rfx.m	Rename MFX-OLS into RFX	
ibma_config_nsubjects.m	Mega-analysis (with FFX at third	l level)
	Image-Based Meta-Analysis toolbox	Image: Sector of the sector

• Available on GitHub

https://github.com/NeuroimagingMetaAnalysis/ibm

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# Meta-analysis of 21 pain studies

- Results
  - GLM methods similar
  - Z-based methods similar
    - But FFX Z methods more sensitive (as expected)

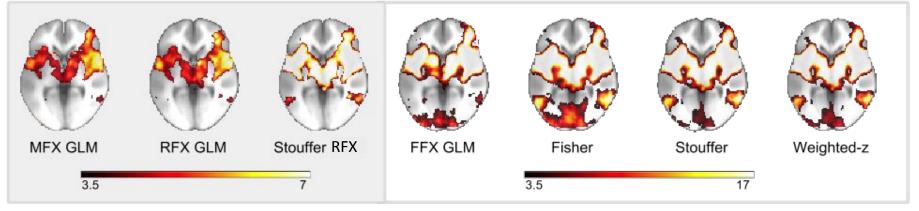


Fig. 1: Result of a meta-analysis of 21 pain studies for 4 fixed-effects (FFX GLM, Fisher, Stouffer, weighted-z) and 2 random-effects (RFX GLM, Stouffer MFX) meta-analytic approaches compared to the reference (MFX GLM) at a threshold of p<0.05 FDR corrected.

#### Data: Tracey pain group, FMRIB, Oxford.

# Self Promotion Alert: Robustness of the meta-analytic estimators

Α Small sample sizes 5 studies 10 studies 25 studies 25 studies 50 studies 20 subjects 20 subjects 20 subjects 100 subjects 20 subjects Within-study 1.5 variance 1.0 Cumulative P - 0.25 FFX 0.5 -- 0.5 0.0 -0.5 -2 Observed P -1.5 varying: 02 1.0 varying: 04 MFX 0.5 varying: 08 0.0 varying: 16 -0.5 2 4 6 2 6 2 4 6 2 6 2 4 6 4 Cumulative P В Heteroscedasticity С Heterogeneity - Cumulative P RFX FFX Stouffer's Weighted Z Perm. E Fisher's 10 -1.5 -1.0 -5 0.5 -Observed P 0.0 -5--0.5 --10-2 2 6 4 6 2 6 2 4 6 Cumulative P Cumulative P

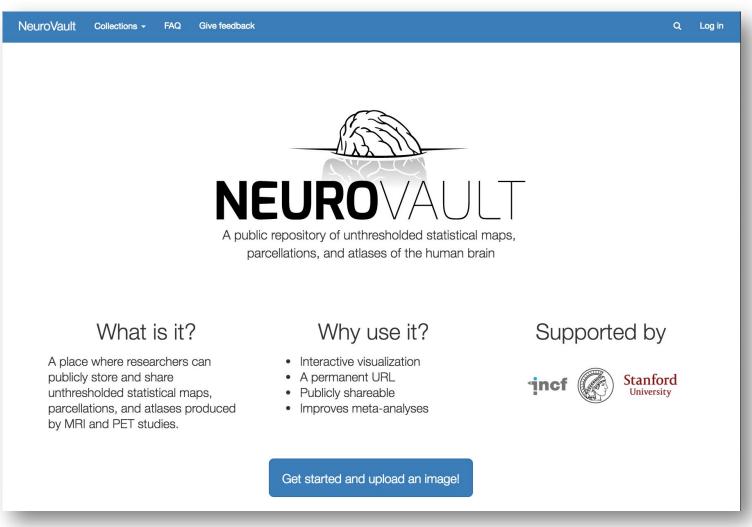
Poster 2653

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# How to publish your statistic maps?

### Share your statistic maps



#### http://neurovault.org

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#### Share your statistic maps

NeuroVault Collections - FAQ Give feedback

Q Log in

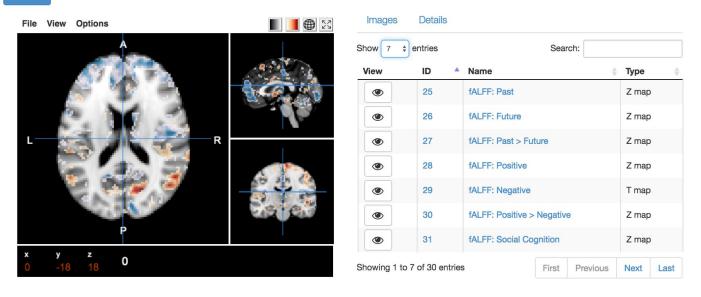
#### A Correspondence between Individual Differences in the Brain's Intrinsic Functional Architecture and the Content and Form of Self-Generated Thoughts

Contributed by ChrisFiloGorgolewski

Krzysztof J. Gorgolewski, Dan Lurie, Sebastian Urchs, Judy A. Kipping, R. Cameron Craddock, Michael P. Milham, Daniel S. Margulies, Jonathan Smallwood

#### Link to the paper

3D View

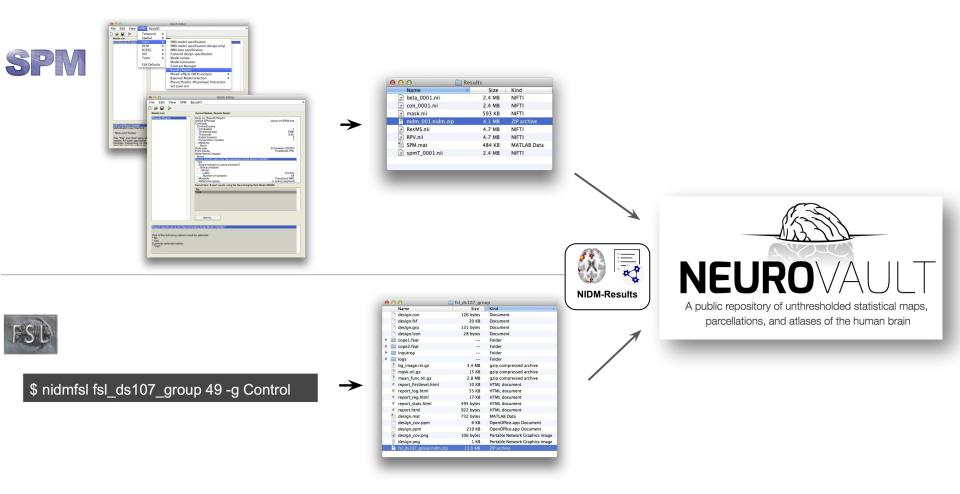


#### http://neurovault.org

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# From SPM & FSL



#### http://nidm.nidash.org/getting-started/

 When data available, Image-Based preferred to Coordinate-Based meta-analysis

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- In practice, it is difficult to use the gold standard Mixed-Effects GLM

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- Few tools for Z-based IBMA, but underway...

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- In practice, it is difficult to use the gold standard Mixed-Effects GLM
- When only contrast estimates are available,
  RFX GLM is a practical & valid approach
- Few tools for Z-based IBMA, but underway...
- Data sharing tools: NeuroVault, NIDM-Results

# Thank you!

