

Neuroimaging meta-analyses the hard way (manually)

Katie Bottenhorn • Meta-Analysis Educational Course • OHBM 2021

The Plan

- I. Neuroimaging
meta-analysis recap
- II. Why would I do this by
hand?
- III. How to
- IV. “Semi-automated”
- V. Why you should share
your statistical maps

Recap

Neuroimaging meta-analyses are facilitated by conventions in the field:

- Results are presented as statistical parametric maps...
- With brain activation peaks reported as $[x, y, z]$ coordinates...
- (And sometimes images!)
- In common stereotaxic reference space

Recap

Meta-analytic algorithms make it possible to synthesize such results

- To assess convergent activation patterns across a task or paradigm
- To contrast neural recruitment across cognitive systems
- To compare activation across subject groups
- Or to define regions of interest (ROIs) for future studies

Examples

Phan et al., 2002: subsystems of emotional processing

Buhle et al, 2014: support for a role of cognitive control in emotion regulation

Glahn et al., 2005: schizophrenia beyond hypofrontality

Fox et al., 2015: spontaneous thought extends beyond the default mode network

But why?

Manual vs. automated metas

Manual

- Search publication database/online library for publications
- Pull coordinates and metadata from individual publications (or email authors for images)
- Run meta-analysis on extracted coordinates/images

Automated

- Search neuroimaging database (e.g., BrainMap, Neurosynth) for publications
- Pull coordinates/images and metadata from database
- Run meta-analysis on extracted coordinates/images

Pros & cons of two approaches

Manual

Greater access to potentially relevant publications

Not limited by bounds of database

Hands-on quality control

Time consuming

Room for error

Automated

Pros

Easier access to coordinates/images and metadata

Several large databases to choose from

Outsource quality control

Cons

Inherently limited metadata

Size vs. quality trade-off

The case for manual meta-analyses

1. *Fewer metadata limitations*

Not all publication metadata is included in databases

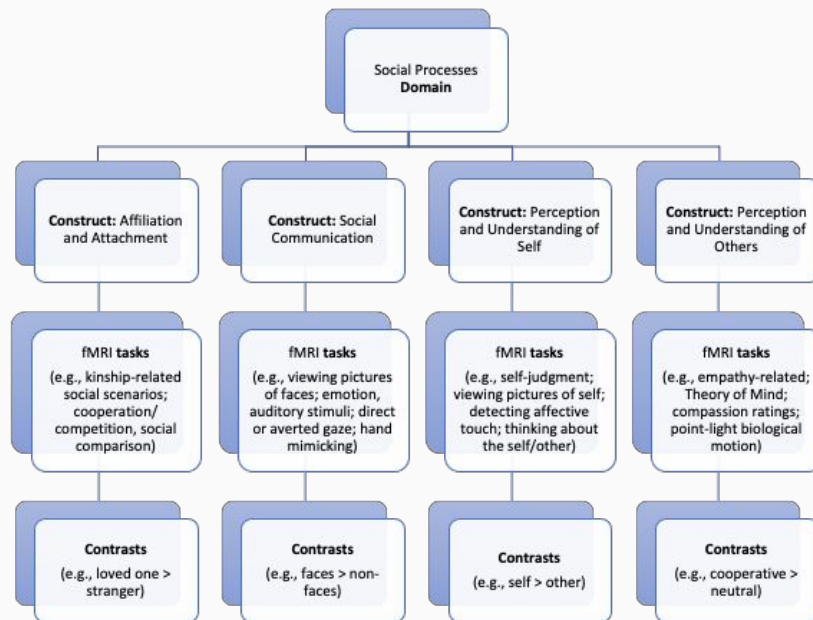
Commonly included:

- Task
- Cognitive construct
- Participant qualities (e.g., age, diagnosis, etc.)

1. Fewer metadata limitations

Examples of meta-analyses requiring uncommon metadata:

- Novel/uncommon task definitions
 - Naturalistic fMRI paradigms (Bottenhorn et al., 2019)
 - RDoC social paradigm groupings (Pintos Lobo et al., in prep)
- Uncommon participant qualities
 - Neuroimaging the menstrual cycle (Dubol et al., 2020)
 - Heart rate variability (Thayer et al., 2012)



2. Greater access to publications

Databases contain a subset of the literature

Which subset depends on how the database is populated

- Manually-populated databases (e.g., BrainMap) are often biased by task, cognitive domain
- Automatically-populated databases (e.g., Neurosynth) can be biased by publication year

Now that you're convinced...

Steps

*Or, How to perform a
manual neuroimaging
meta-analysis*

(for more information, see Müller et al., 2018)

1. Decide on inclusion / exclusion criteria
2. Search for publications
3. Annotate & exclude papers
4. Extract coordinates & metadata
5. Meta-analyze!

*What if we wanted
to do a
meta-analysis of
video games in
fMRI?*

1. *Decide on criteria*

Fundamentals:

1. Brain activation coordinates reported, or images shared, in standard space
2. Whole-brain analysis (i.e., not small-volume, region of interest analysis)

Everything else depends on your use case.

Example: video games in the scanner

Inclusion:

- Participants play video games in MR bore

Exclusion:

- Coordinates not reported OR no access to images
- Results reported not from video game paradigm
- ROI analysis (i.e., not whole-brain)

2. *Search for publications*

Good search engine options:

- PubMed/MEDLINE
- Web of Science

Less good, but reasonable search options

- Google Scholar
- Individual journals

Many meta-analytic researchers peruse published reviews and reference lists in other publications for additional papers

Video games in the scanner

PubMed search:

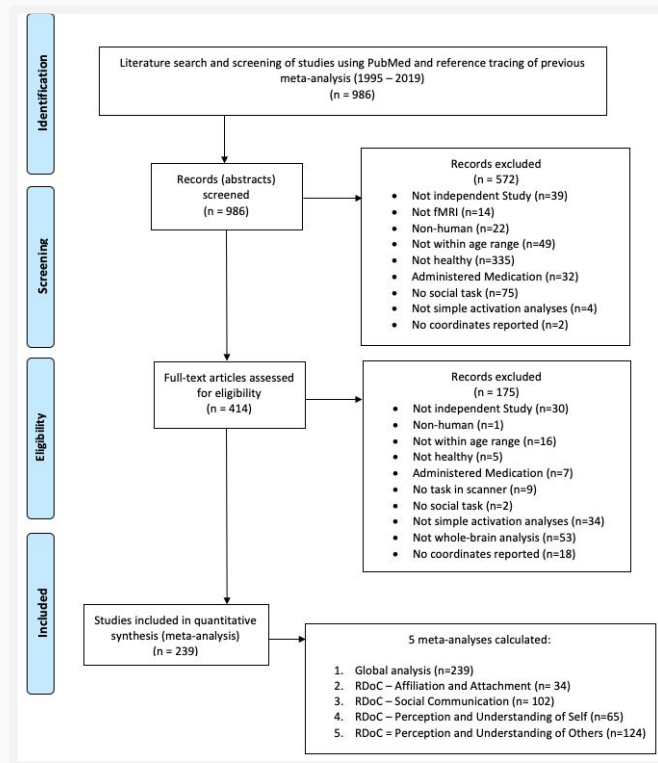
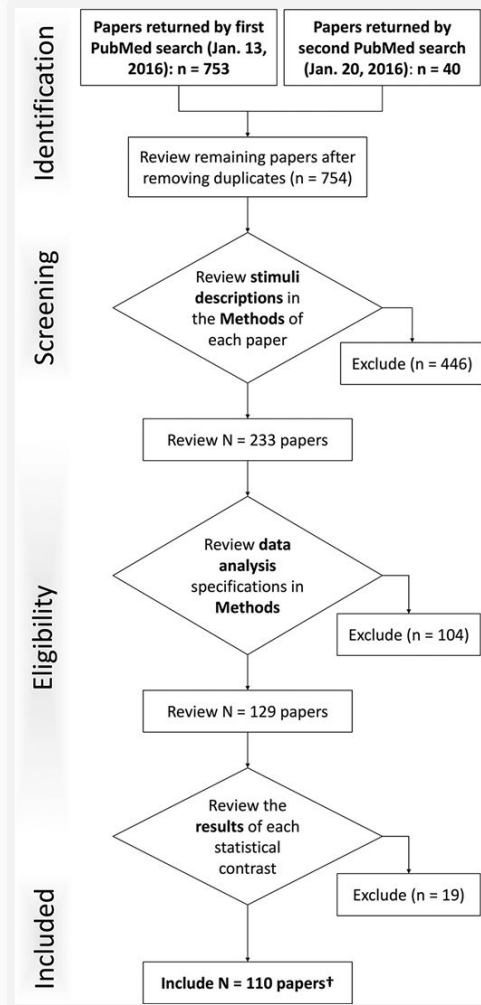
```
((video game*) OR (videogame*)) AND  
((fmri) OR (functional magnetic  
resonance imaging))
```

3. Annotate & exclude

Not all papers turned up by your search will meet criteria

See preferred reporting items for systematic reviews and meta-analyses (PRISMA) Checklist for guidance

Review papers in stages and keep track of reason for exclusion



4. *Acquire data!*

Coordinate-based meta-analyses:

- Extract coordinates from publication
- Separate by statistical analysis (or, *contrast*)

Image-based meta-analyses:

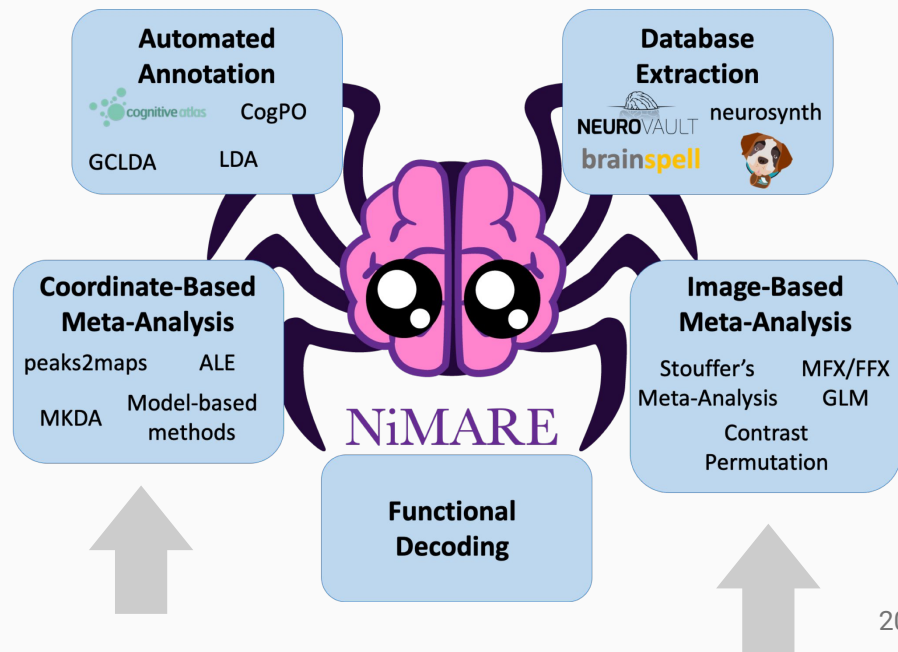
- Fetch images from repository
- Or: email authors to request statistical images

5. Meta-analyze!

This step is almost agnostic to the method of data acquisition/curation

Requirements:

- Coordinate or image data
- Any necessary metadata (i.e., # subjects for most CBMA)
- Meta-analytic algorithm



But what if there were shortcuts?

Semi-automated meta-analysis

1. Decide on inclusion / exclusion criteria
2. Search for publications
3. Annotate & exclude papers
4. Extract coordinates & metadata
5. Meta-analyze!



NEUROVAULT



metaCurious.

neurosynth.org₂

Neurosynth

Coordinate-based repository

Can help with

- Searching for publications
- Extracting coordinates & metadata

neurosynth.org

NeuroVault

Image-based repository

Can help with

- Searching for publications
- Extracting coordinates & metadata



metaCurious

Web-based dataset curation tool
(WIP)

Includes data from Neurosynth

Can help with

- Searching for publications
- Annotating & exclude papers
- Extracting coordinates & metadata



PSA:

*Share your
statistical results
as images*

Thanks!

- Bottenhorn, K. L., Flannery, J. S., Boeving, E. R., Riedel, M. C., Eickhoff, S. B., Sutherland, M. T., & Laird, A. R. (2018). Cooperating yet distinct brain networks engaged during naturalistic paradigms: A meta-analysis of functional MRI results. *Network Neuroscience*, 3(1), 27-48. https://doi.org/10.1162/netn_a_00050
- Buhle, J. T., Silvers, J. A., Wager, T. D., Lopez, R., Onyemekwu, C., Kober, H., Weber, J., & Ochsner, K. N. (2014). Cognitive Reappraisal of Emotion: A Meta-Analysis of Human Neuroimaging Studies. *Cerebral Cortex*, 24(11), 2981-2990. <https://doi.org/10.1093/cercor/bht154>
- Dubol, M., Epperson, C. N., Sacher, J., Pletzer, B., Derntl, B., Lanzenberger, R., Sundström-Poromaa, I., & Comasco, E. (2021). Neuroimaging the menstrual cycle: A multimodal systematic review. *Frontiers in Neuroendocrinology*, 60, 100878. <https://doi.org/10.1016/j.yfrne.2020.100878>
- Glahn, D. C., Ragland, J. D., Abramoff, A., Barrett, J., Laird, A. R., Bearden, C. E., & Velligan, D. I. (2005). Beyond hypofrontality: A quantitative meta-analysis of functional neuroimaging studies of working memory in schizophrenia. *Human Brain Mapping*, 25(1), 60-69. <https://doi.org/10.1002/hbm.20138>
- Müller, V. I., Cieslik, E. C., Laird, A. R., Fox, P. T., Radua, J., Mataix-Cols, D., Tench, C. R., Yarkoni, T., Nichols, T. E., Turkeltaub, P. E., Wager, T. D., & Eickhoff, S. B. (2018). Ten simple rules for neuroimaging meta-analysis. *Neuroscience & Biobehavioral Reviews*, 84, 151-161. <https://doi.org/10.1016/j.neubiorev.2017.11.012>
- Phan, K. L., Wager, T., Taylor, S. F., & Liberzon, I. (2002). Functional Neuroanatomy of Emotion: A Meta-Analysis of Emotion Activation Studies in PET and fMRI. *NeuroImage*, 16(2), 331-348. <https://doi.org/10.1006/nimg.2002.1087>
- Pintos Lobo et al., (forthcoming). *Neural systems underlying RDoC social constructs: An activation likelihood estimation meta-analysis.*
- Thayer, J. F., Åhs, F., Fredrikson, M., Sollers, J. J., & Wager, T. D. (2012). A meta-analysis of heart rate variability and neuroimaging studies: Implications for heart rate variability as a marker of stress and health. *Neuroscience & Biobehavioral Reviews*, 36(2), 747-756. <https://doi.org/10.1016/j.neubiorev.2011.11.009>