

# Preregistering an fMRI study

Helena Hartmann

29.06.2021

*First  
experiences  
and lessons  
learned*



## When you preregister, you...

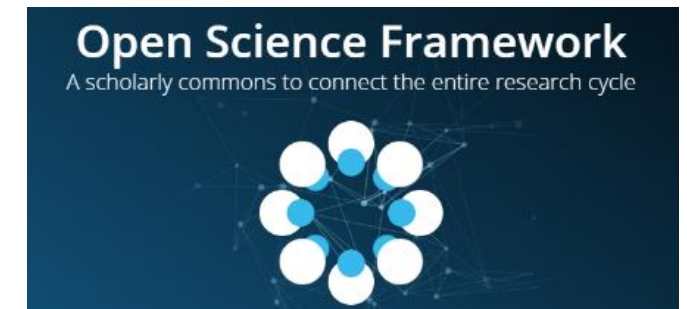
- ... **openly declare the research** you are planning to conduct (public or with embargo)
- ... **commit ahead of time** to hypotheses, methods, and analytic strategy
- ... clearly **distinguish confirmatory from exploratory** research

**Resources** → multiple templates online (e.g. OSF or AsPredicted)

**Claim early authorship** for ideas/designs

**No peer-review** → faster start than with Registered Reports

**Faster dissemination** → a shift in time commitment from post- to pre-study



# Why do we need preregistration of fMRI studies?

## Article

<https://www.narps.info/>

# Variability in the analysis of a single neuroimaging dataset by many teams

<https://doi.org/10.1038/s41586-020-2314-9>

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Published online: 20 May 2020



Check for updates

A list of authors and affiliations appears

Data analysis workflows in many science fields are complex and flexible. Here we assess variability in the analysis of functional magnetic resonance imaging data using the same dataset, testing the same 9 approaches is exemplified by the fact that different teams analyse the data. This flexibility results in different hypothesis tests, even for teams who use the same intermediate stages of the analysis pipeline. We report on several aspects of analysis method variability and aggregated information across team regions. Furthermore, prediction models

## News & views

### Reproducibility

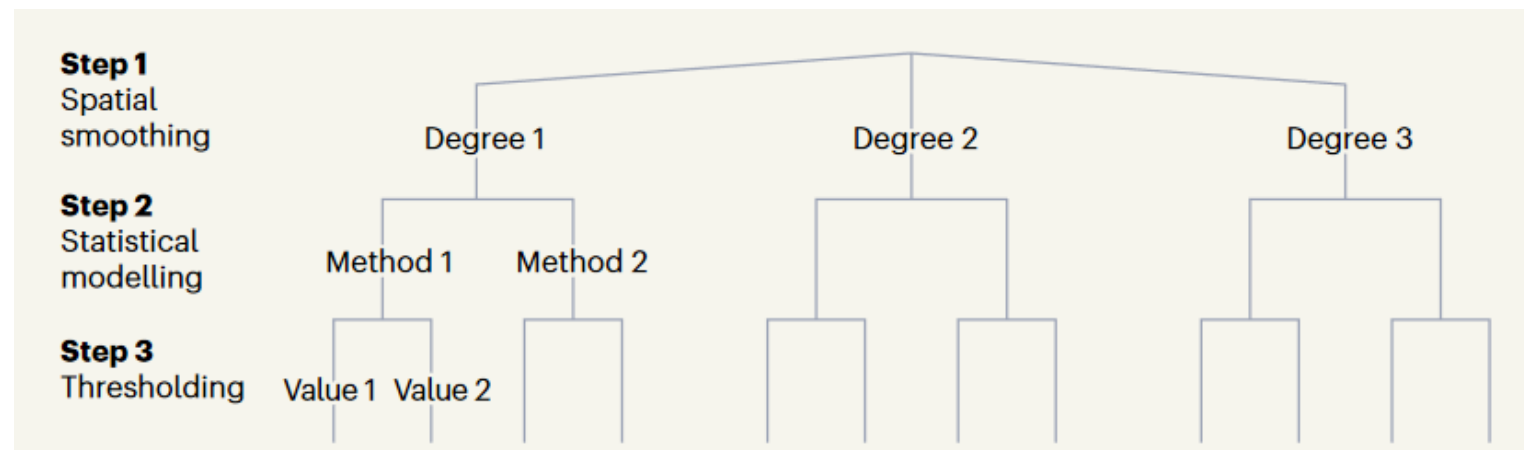
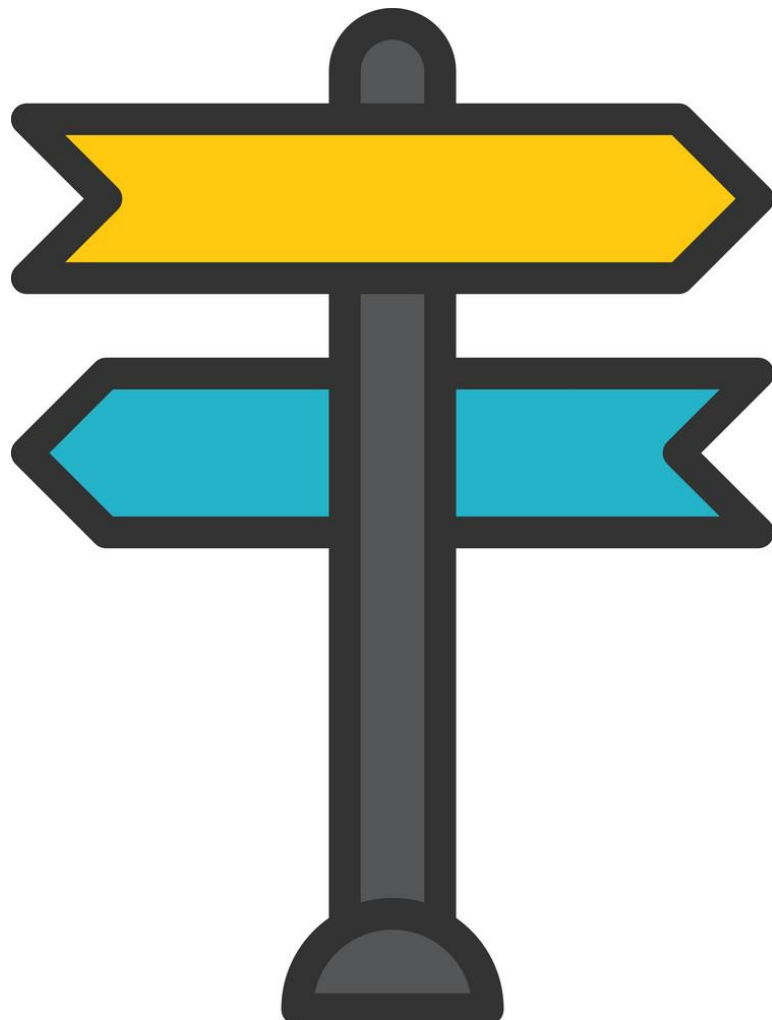
# Pipeline choices alter neuroimaging findings

Martin Lindquist

Seventy laboratories that analysed the same data each produced different results, highlighting the potential consequences of variability in analysis pipelines for processing complex data. S

“Notably, no two teams chose identical workflows to analyse the data, resulting in substantial variation in the results.”

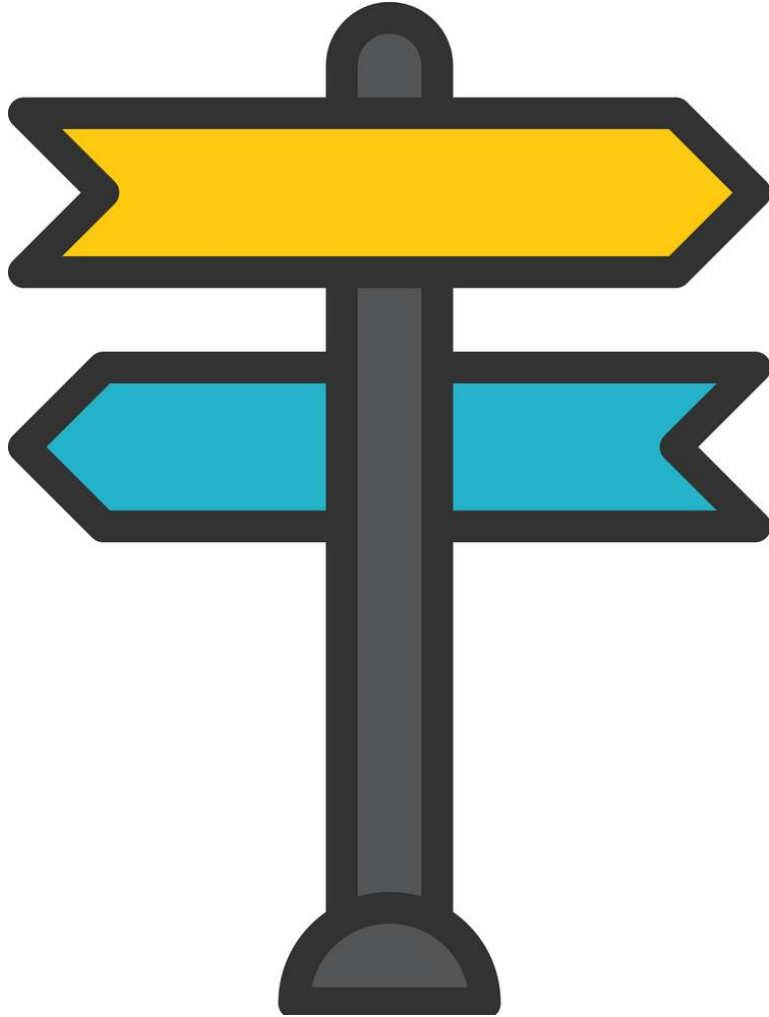
# Because of decisions, decisions...



n possible  
combinations  
(where n = A LOT)

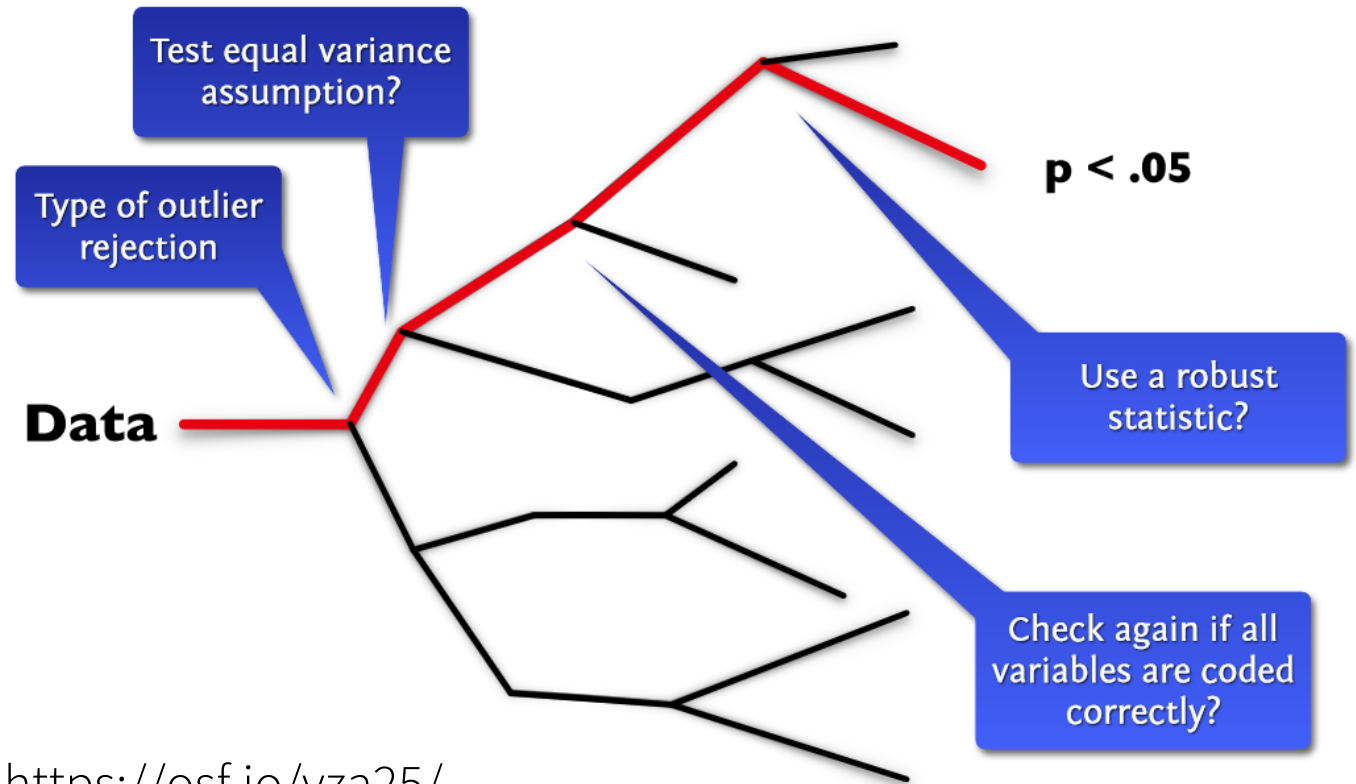
Lindquist et al. (2020)

# Because of decisions, decisions...



## The garden of forking paths

Andrew Gelman & Eric Loken, 2013



<https://osf.io/vza25/>

Inspired by Neuroskeptic's blog: <http://blogs.discovermagazine.com/neuroskeptic/2015/05/18/p-hacking-a-talk-and-further-thoughts/#.VV2TiOePKsN>

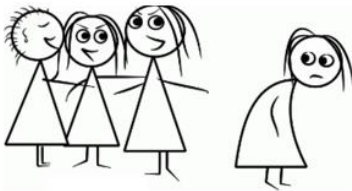
17



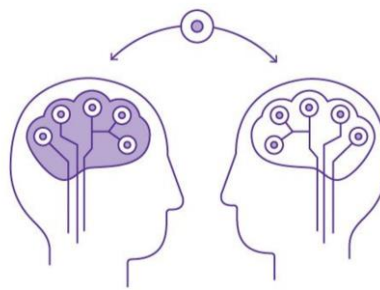
MSc in Clinical  
& Biological Psychology



PhD in Psychology



EMPATHY



NETHERLANDS  
INSTITUTE  
FOR NEUROSCIENCE  
Master the mind



Psychologist  
for MyMind

MyMind  
EMBRACING NEURODIVERSITY



Brain Hero



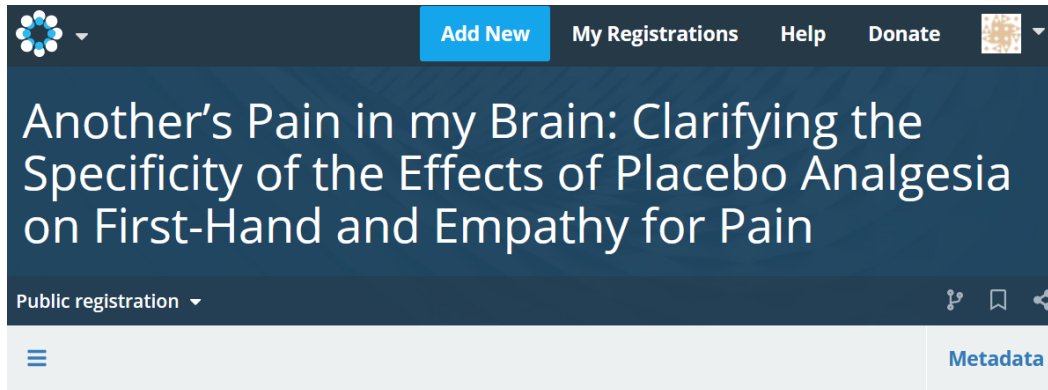
CLINICAL SOCIAL  
NEUROSCIENCE UNIT



SCAN Unit



# How it started



## Study Information <https://osf.io/uwzb5>

### Title

Effects of placebo analgesia on somatosensory responses during first-hand and empathy for pain

### Research Questions

Previous studies on placebo empathy analgesia (Rütgen et al., 2015, PNAS, J Neurosci) did not report any variation of somatosensory activation in the pain-processing network during empathy for pain, but only showed activation changes in the network related to the affective-motivational component. This is surprising given that placebo analgesia generally also affects the sensory-discriminative component of first-hand pain (Benedetti et al., 2005; Wager & Atlas, 2015). However, this mismatch might have resulted from the specifics of the experimental paradigm used in past studies, which did not seem ideally suited to observe activation of somatosensory areas (Keyser, Kaas, & Gazzola, 2010). In fact, it has been suggested that certain types of paradigms showing a specific body part in pain are required to observe modulation of somatosensory areas. Importantly, previous research on placebo empathy analgesia did not use such a setup (but used abstract cues and facial expressions of pain as a more indirect measure).

Furthermore, placebo empathy analgesia has only been investigated using first-hand painful and non-painful electrical stimulation and with a paradigm focusing on abstract cues ("cue-based" task). What is not clear yet, is whether also other types of empathic pain can be modulated by placebo analgesia, e.g. merely seeing pictures of

# How it's going



## Another's pain in my brain: No evidence that placebo analgesia affects the sensory-discriminative component in empathy for pain

Helena Hartmann, Markus Rütgen, Federica Riva, Claus Lamm\*

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### ARTICLE INFO

**Keywords:**  
Empathy  
Social  
Electrical pain  
Placebo analgesia  
Somatosensation  
fMRI



### ABSTRACT

The shared representations account of empathy suggests that empathic processes rely on neural processes similar to those engaged when one experiences pain. Recent research corroborated this by showing that placebo analgesia reduces pain empathy and decreased activation in shared pain-processing areas. However, these studies did not report any placebo-related variation of somatosensory activation, which may explain the absence of effects for somatosensation. The mismatch between the experimental paradigms used in these studies did not directly address the sensory-discriminative component of empathy for pain, which may explain the absence of effects for somatosensation. The mismatch between the experimental paradigms used in these studies did not directly address the sensory-discriminative component of empathy for pain. The mismatch between the experimental paradigms used in these studies did not directly address the sensory-discriminative component of empathy for pain.

Free of HARKing or p-hacking!

### 1. Introduction

Empathy is a multifaceted psychological construct fundamental for human social interactions and relationships (e.g. Marsh, 2018 for recent review). While many definitions of empathy have been proposed, here we define empathy as an affective state isomorphic to the state

**Another's Pain in my Social Brain: The Effects of Placebo (Empathy) Analgesia on Social Behavior**

Hartmann & Lamm

**The Effects of Placebo Analgesia on Interoceptive Abilities**

Hartmann, Riva & Lamm

## Experiences when writing up the papers

- **Open Science („21 word“) statement** at the beginning of the methods

### 2.2. Preregistration

We report how we determined our sample size, all data exclusions, all manipulations, and all measures in the study. This study was pre-registered on the OSF prior to any creation of data (Hartmann et al., 2018; preregistration: [osf.io/uwzb5](https://osf.io/uwzb5); addendum: [osf.io/h7v9p](https://osf.io/h7v9p)) and was designed to extend and specify the results of Rütgen et al. (2015b) in regard to somatosensory sharing. Methods reported below are therefore reproduced partly verbatim from the preregistration. Note that the pre-registered plan contains a second research question that is not part of the present paper but will be reported elsewhere. In the following methods and results, we clearly distinguish preregistered procedures and analyses from those added post hoc.

- Clear **distinction between preregistered and post hoc** analyses in methods and results

#### 2.7.1. Preregistered analyses

We implemented a within-subjects, full-factorial design with three factors of two levels each (*treatment*: placebo vs. control hand, *target*: self vs. other, *intensity*: pain vs. no pain). Two parametric repeated-measures analyses of variance (ANOVAs) were used to analyze the results. In the first ANOVA (analysis A1 in Fig. 2), the dependent variable was the self- and other-related pain ratings. A second ANOVA (analysis A2 in Fig. 2) included the unpleasantness ratings as the dependent variable (omitting the factor *target*, as unpleasantness ratings were only collected in the empathy condition). For each ANOVA, we then computed planned comparisons using paired *t*-tests.

#### 2.7.2. Post hoc analyses

Due to the unexpected “null” finding of no transfer of the first-hand placebo effect to empathy, we aimed to gather further relative evidence for the null vs. the alternative hypothesis, using a Bayesian approach



## Experiences when writing up the papers

- **Editor:** „(...) Reviewers **complimented the Authors on the decision to pre-register the study.**“
- **Reviewer 1:** „The fact that the **study was pre-registered** led to **high clarity in the description of the methods**, as well as to an **honest report of the planned analysis and of the additional investigations** - an aspect that is absolutely commendable.“
- **Reviewer 3:** „(...) interesting study design, combined with **rigorous, pre-registered analyses** that are compiled to a well-written manuscript.“
- **Reviewer 4:** “Overall, this study is a **carefully conducted, preregistered study.**”





**Learning by doing:**  
I preregistered one  
fMRI study so far!

aka

„The naive first-year  
PhD student approach“

# The 20% Statistician

A blog on statistics, methods, philosophy of science, and open science.  
Understanding 20% of statistics will improve 80% of your inferences.

Monday, May 11, 2020

## Red Team Challenge

*by Nicholas A. Coles, Leo Tiokhin, Ruben Arslan, Patrick Forscher, Anne Scheel, & Daniël Lakens*

Today we announce an initiative that we hope can incentivize critical feedback and error detection in science: the Red Team Challenge. Daniël Lakens and Leo Tiokhin are offering a total of \$3,000 for five **individuals to provide critical feedback on the materials, code, and ideas** in the forthcoming preprint titled “**Are facial feedback effects solely driven by demand characteristics? An experimental investigation**”. This preprint examines the role of demand characteristics in research on the controversial facial feedback hypothesis: the idea that an individual’s facial expressions can influence their emotions. This is a project that Coles and colleagues will submit for publication in parallel with the Red Team Challenge. We hope that challenge will serve as a useful case study of the role Red Teams might play in science.



### About Me

Blog by [Daniel Lakens](#),  
experimental psychologist at  
the Human-Technology  
Interaction group at Eindhoven  
University of Technology, The  
Netherlands.

<https://daniellakens.blogspot.com/2020/05/red-team-challenge.html>


## Red Team exercise (1/5)

In what fashion?



### Hypotheses

1) The somatosensory component of the empathic reaction to pain is modulated in a similar fashion by placebo analgesia as the affective component, but only if the attention of participants is explicitly directed to the specific body part in pain. To this end, a new experimental paradigm will be used in which placebo analgesia is induced only for one of the two hands (placebo hand), while no analgesia is induced in the other (control hand). We predict reductions in empathy in the pain and unpleasantness ratings as well as in both pain matrix components. These reductions will be restricted to the hand in which placebo analgesia was induced (in comparison to the control hand).



**What's the pain matrix? What brain regions do you expect?**

<https://osf.io/uwzb5>

## Red Team exercise (2/5)

What on earth constitutes  
extensive movement?

### Data exclusion

If subjects show consistent extensive movement during the fMRI scans, and in particular, if this movement is stimulus-related, they will be excluded from further analysis.

Please explain further  
what this means!

<https://osf.io/uwzb5>



## Red Team exercise (3/5)

Come on, standard  
algorithms and parameters?  
What does that even mean?

### Transformations

Regarding the neuroimaging data, preprocessing will be carried out with the Statistical Parametric Mapping software package (SPM12; Wellcome Trust Centre for Neuroimaging, UCL, London, UK) using standard algorithms and parameters (slice timing correction, VDM calculation from field maps, realign & unwarp, coregistration, segmentation, functional and structural normalization and smoothing). Region of

Nice that you mention all the steps, but  
each of these have multiple parameters  
that can be changed flexibly...

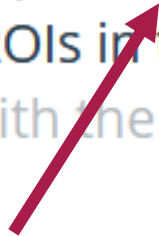
<https://osf.io/uwzb5>

## Red Team exercise (3/5)

### Statistical models

into the analysis as one single factor, with independent ROIs determined from recent findings on pain (e.g. Lamm et al., 2011; Corradi-Dell'Acqua et al., 2011; Rütgen et al., 2015, PNAS and JNeurosci). In addition to previously found areas related to placebo empathy analgesia, the focus will be on ROIs in the first and secondary somatosensory cortex. In the case of significant effects with the factor 'ROI' (pooled activation of all

Independent is great, but what  
are the exact coordinates?  
What's the sphere size?  
Or are they anatomical ROIs?



What areas?



The somatosensory cortex  
is big, so where exactly?

<https://osf.io/uwzb5>

## Red Team exercise (5/5)

### Follow-up analyses

Significant interactions, if not sufficiently explained by the analysis plan and in particular by the planned comparisons, will be followed-up (and corrected for multiple comparisons).

That's nice, but what kind of correction? FDR, FWE, ...?



<https://osf.io/uwzb5>

FROM J. K. ROWLING'S  
WIZARDING WORLD

FANTASTIC

fMRI  
prereg **DILEMMAS**

AND HOW  
TO SOLVE THEM

## Common dilemmas

- I used **OSF Prereg Challenge** and standard **OSF Prereg** templates (more templates can be found here: <https://osf.io/zab38/wiki/home/>)
- **PRP-QUANT Template:**  
<https://www.psycharchives.org/handle/20.500.12034/4042.2>
- Now there is an **fMRI-specific template** available:
  - **OSF:** <https://osf.io/6juft/> (also provides guidance for common difficulties!)
  - Adapted within **MPI-CBS Hackathons:**  
[https://docs.google.com/document/d/1GHkCCvu\\_io56mB8wWwWzuts1hTXyWOC9-DNOHlYSUTw/edit#heading=h.omxd3x3jr2d7](https://docs.google.com/document/d/1GHkCCvu_io56mB8wWwWzuts1hTXyWOC9-DNOHlYSUTw/edit#heading=h.omxd3x3jr2d7) (for questions contact Frauke Beyer at [fbeyer@cbs.mpg.de](mailto:fbeyer@cbs.mpg.de))
- **SIPS 2019 collection of neuroimaging/electrophysiology preregs:**  
<https://docs.google.com/spreadsheets/d/1K-sBCobM1jmTbC3ctWdqaZpkt00RyPVpU1xk95njLU8/edit#gid=1351240212>

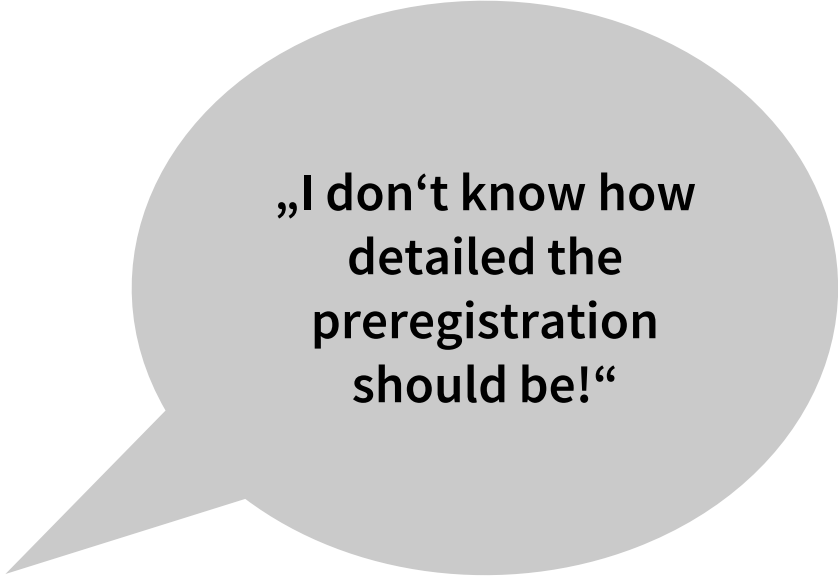
„I have no idea  
where to start or  
which information  
to include!“

<https://osf.io/mtdh2/>



## Common dilemmas

- Be **as detailed as possible**
- BUT very flexible/free in what you include → **It is up to you**, nobody will check it, except (hopefully) the reviewers!
- Who to add as **authors**? → be as sparingly as possible, it's always easier to add new people!
- Upload **additional** method-related **files**: Task/Analysis code, Stimuli, Randomizations, etc.
- Think about all **possible outcomes**, e.g. what analyses to do if the data are not normally distributed?

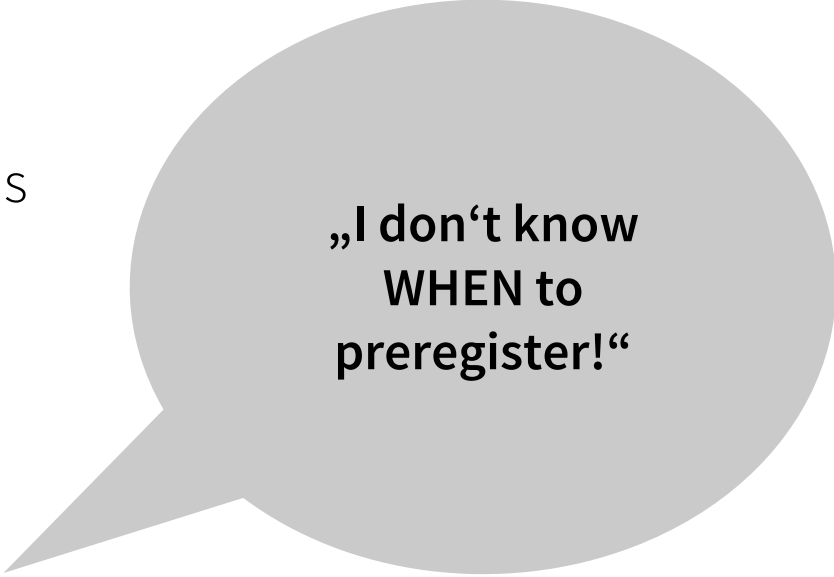


„I don't know how detailed the preregistration should be!“

<https://osf.io/mtdh2/>

## Common dilemmas

- Best **before collection of any data** that you will use in the analysis
- **My approach:** After pilots, before first real participant
- Theoretically possible at all time points, but the closer you get to analysis, the **harder it is to disentangle biasing** by data/analysis/results

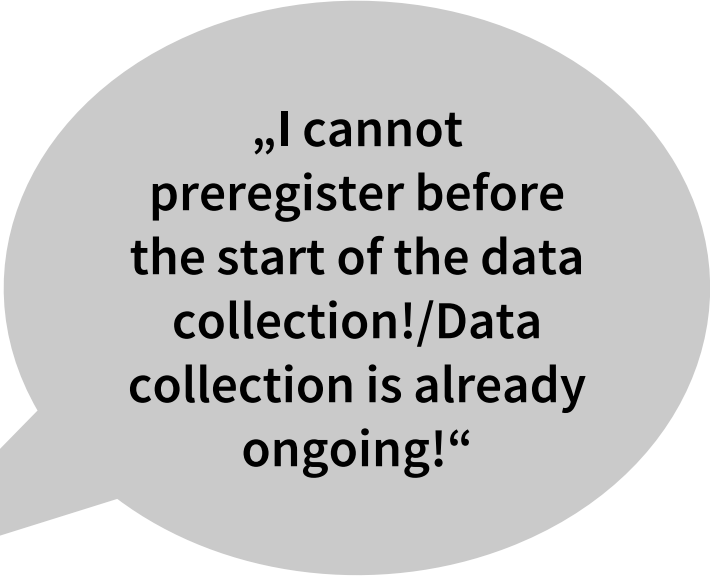


„I don't know  
**WHEN** to  
preregister!“

<https://osf.io/mtdh2/>

## Common dilemmas

- Preregistration is “**a plan not a prison**”
- Keyword **transparency**: State prior knowledge of existing dataset
- **Amendments** always possible



„I cannot  
preregister before  
the start of the data  
collection!/Data  
collection is already  
ongoing!“

## Summary

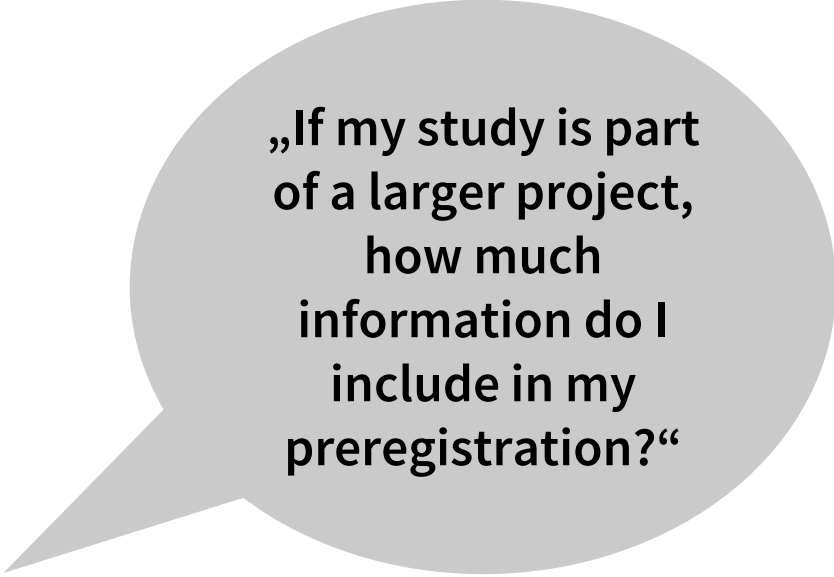
**Provide a narrative summary of what is contained in this registration, or how it differs from prior registrations.**

This is an addendum to the preregistration of this project, uploaded on August 3rd 2018. There, we wrote:

<https://osf.io/mtdh2/>

## Common dilemmas

- **Link to all previous publications/preregistrations**, possibly all under an OSF project
- State **how prior information will influence hypotheses** or how you will not allow prior knowledge to influence your hypotheses
- Include **descriptions of measurements you plan to use**

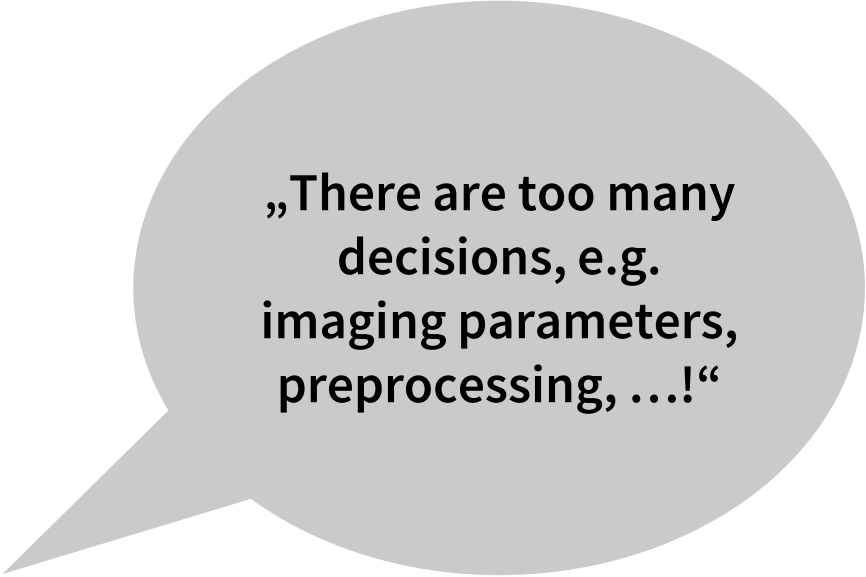


„If my study is part of a larger project, how much information do I include in my preregistration?“

<https://osf.io/mtdh2/>

## Common dilemmas

- It's crucial to **think through those decisions** prior to data analysis!
- Make use of **available data structures and pipelines**:
  - **BIDS** & BIDS apps which allow you to quickly share your exact analysis pipeline, with the specific software and correct software versions
  - **Many Labs** have standard operating procedures and pipelines
- If you don't know a decision yet, it's a good reminder to think about what you want to do!



„There are too many decisions, e.g. imaging parameters, preprocessing, ...!“

<https://osf.io/mtdh2/>



## Common dilemmas

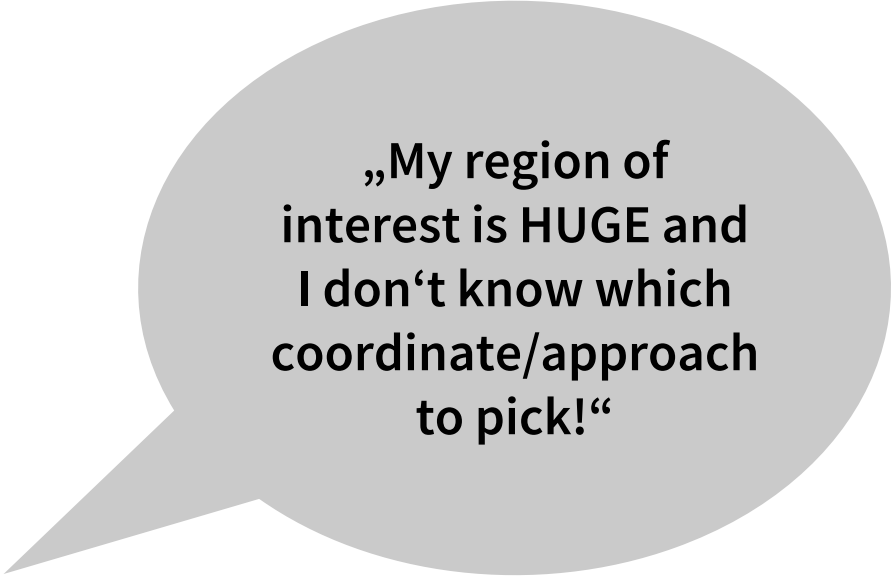
- **NeuroPowerTools**: <http://neuropowertools.org/> (You can use data that you upload or that's already on Neurovault)
- **fMRIPower**: <http://fmripower.org/>

„I cannot really  
calculate my power,  
which makes sample  
size calculations  
hard!“

<https://osf.io/mtdh2/>

## Common dilemmas

- Definitely **needs to be specified**, this is where a lot of flexibility comes from!
  - Anatomical definition
  - Prior study cluster
  - Neurosynth definition (make sure to be specific here!)
  - Parcellation definition
- Use **previous studies/lab experience** as a guideline

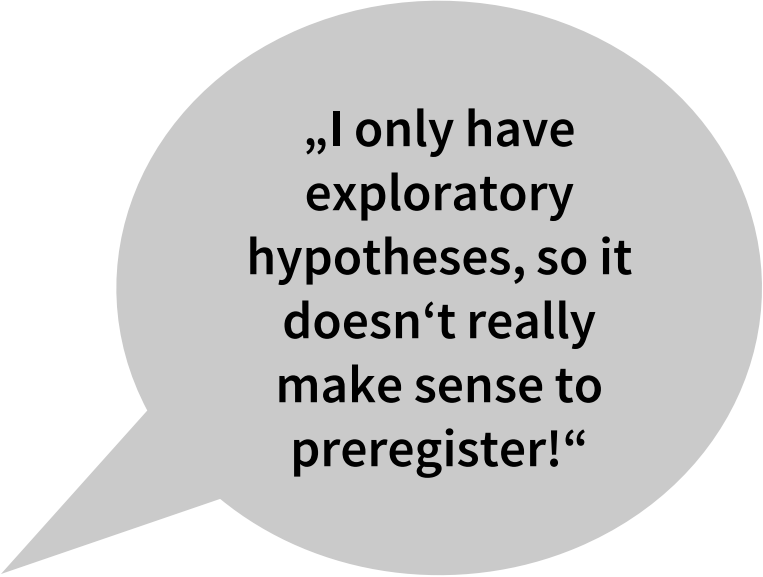


„My region of interest is HUGE and I don't know which coordinate/approach to pick!“

<https://osf.io/mtdh2/>

## Common dilemmas

- **Yes, it does!** Preregistration just clearly distinguishes confirmatory from exploratory research!
- See “**Exploratory Reports**” from Cortex:  
[https://www.elsevier.com/\\_\\_data/promis\\_misc/Exploratory\\_Reports\\_Guidelines.pdf](https://www.elsevier.com/__data/promis_misc/Exploratory_Reports_Guidelines.pdf)



„I only have  
exploratory  
hypotheses, so it  
doesn't really  
make sense to  
preregister!“

<https://osf.io/mtdh2/>

## Personal lessons learned

- Choose a template that is detailed and gives you a good **framework/structure**
- Have a look at **other fMRI preregistrations**
- **Read Team:** Let colleagues **evaluate/review** it before submitting
- Don't be afraid to make mistakes/**amendments**
- **IMO:** Better to have an only 80% perfect preregistration than none!
- Key: **Transparency!**
- Ask yourself: Would somebody else be able to **replicate your study just using the preregistration?**
- **When in doubt:** Write to the COS, they are super helpful ([contact@cos.io](mailto:contact@cos.io))!



## More information


- **OHBM Open Science room** (fMRI prereg template): <https://ohbm.github.io/osr2021/>
- For **EEG**-interested people: Hackathon @ SIPS (<https://www.improvingpsych.org/SIPS2021/>)
- **Nichols et al. (2017)** – best practices in neuroimaging studies
- **Poldrack et al. (2007)** – what to report in an fMRI study
- **Gentili et al. (2020)** – Preregistering ROI analyses



**SIPS**  
@improvingpsych

Happening at **#SIPS2021** on 24 June 2021 15:00 CEST:  
Hackathon on 'Finalizing a Preregistration Template for ERP Studies'

### Best practices in data analysis and sharing in neuroimaging using MRI

Thomas E Nichols , Samir Das, Simon B Eickhoff, Alan C Evans, Tristan Glatard, Michael Hanke, Nikolaus Kriegeskorte, Michael P Milham, Russell A Poldrack, Jean-Baptiste Poline, Erika Proal, Bertrand Thirion, David C Van Essen, Tonya White & B T Thomas Yeo

### Guidelines for reporting an fMRI study

Russell A. Poldrack,<sup>a,\*</sup> Paul C. Fletcher,<sup>b</sup> Richard N. Henson,<sup>c</sup> Keith J. Worsley,<sup>d</sup> Matthew Brett,<sup>c</sup> and Thomas E. Nichols<sup>e</sup>

<sup>a</sup>Department of Psychology, Department of Psychiatry and Biobehavioral Sciences, and Brain Research Institute, University of California Los Angeles, Los Angeles, CA 90095, USA

### The case for preregistering all region of interest (ROI) analyses in neuroimaging research

#### Abstract

In neuroimaging studies, small sample sizes and the resultant reduced statistical power to detect effects that are not

paired with voxel- or cluster-wise corrections for multiple comparisons. Though voxel-wise corrections are more reliable in terms of controlling the false-positive rate, they have





# Any other questions?

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🌐 [www.helenahartmann.com](http://www.helenahartmann.com)

🐦 [@helenahhartmann](https://twitter.com/helenahhartmann)

# Thank you for your attention!