

Blood vessel modelling

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The problem in my words

Big picture questions

- How do age and disease affect brain vessel health?
- What are precapillary sphincters for?

Experimental protocol

Baseline → Hypertension → Recovery → Ablation → Hypertension 2

Measurements

- Diameter before & after whisker stimulation (slow, measures responsiveness)
- Diameter and center point (fast, FTed to measure pulsatility)
- Blood pressure at femoral artery
- Speed (measures velocity and total flux)

Specific questions 1: whisker stimulation

How do ageing and sphincter ablation affect whisker stimulation response?

- a) Does the response depend on age and/or vessel type?
- b) Does the effect of sphincter ablation on the response depend on age?

Specific questions 2: pulsatility

How do age, treatment, pressure and vessel type affect pulsatility of diameter (P_d) and center position (P_c)?

- a) Is there an overall age difference? If so, is it explained by pressure?
- b) Does the effect of sphincter ablation depend on age?
- c) What is the effect of pressure?
- d) Does the effect of treatment (also on diameters) vary by vessel type?

Modelling approach

Bayesian multilevel regression modelling

1. A generative model expressing topologically how measurables \hat{y} depend on parameters θ . i.e. f s.t. $\hat{y} = f(\theta)$
2. A measurement model probabilistically connecting measurables with measurements y . i.e. $p(y \mid \hat{y})$
3. A prior model probabilistically expressing non-experimental information about parameters. i.e. $p(\theta)$

Multilevel models

In a multilevel model, some parameters express second or higher order information.

For example, in this model τ represents information about the parameters α .

$$y_i \sim N(\hat{y}_i, \sigma)$$

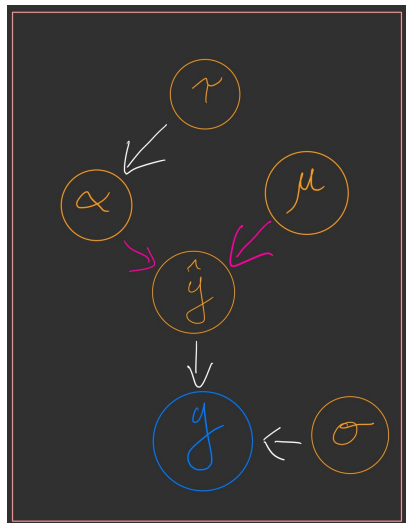
$$\hat{y}_i = \mu + \alpha_{mouse(i)}$$

$$\alpha \sim N(0, \tau)$$

Multilevel models generalise random/mixed effect models.

Representing Bayesian multilevel models with graphs

- Node = parameter (orange) or measurement (blue)
- Edge = full dependence (pink) or probabilistic dependence (white)
- N.B. **Not** the same as “Bayesian networks”
- See [1] for more about graphical models.



Reading about this area

[1] Bayesian graphical models case study with general introduction/discussion.

[2] Paper about “Bayesian workflow”.

https://betanalpha.github.io/assets/case_studies/hierarchical_modeling.html Abstract
mathemtical discussion of multilevel modelling in general.

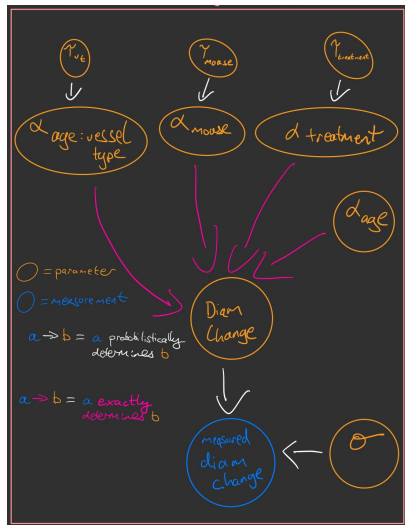
Why is this approach appropriate here?

- We are interested in group-level effects (treatment, mouse, vessel type) and don't know the population parameters (e.g. how much to expect mouse effects to vary).
- We have ample but not massive data, so regularisation from priors is useful.
- We need a fairly complex model; Bayesian multilevel models scale better with complexity.

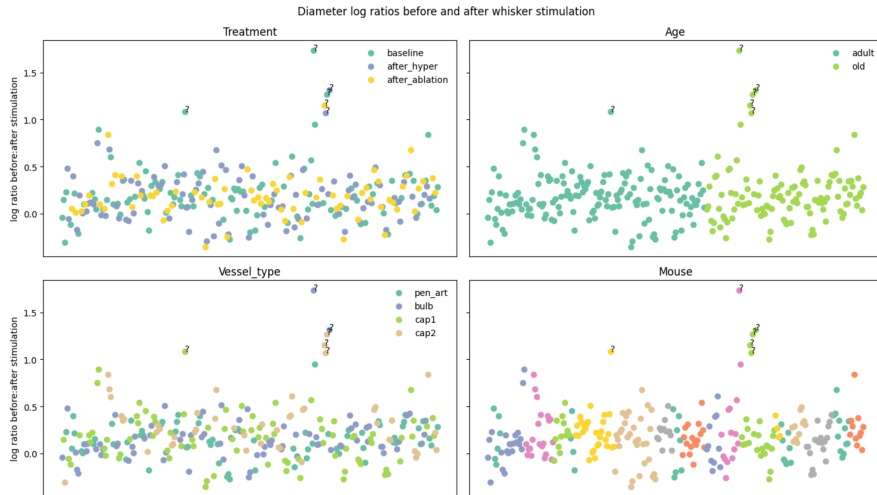
Plan

Whisker stimulation model

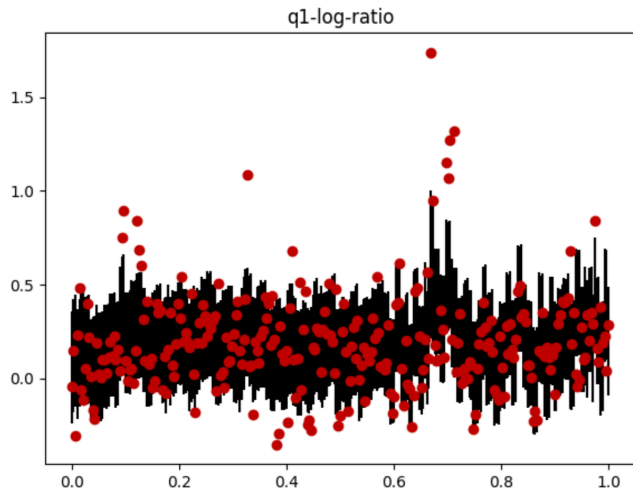
- Measurable: $\ln \frac{diam_{after}}{diam_{before}}$
- Measurement model: linear regression
- Parameters: Note that $\alpha_{age:vessel\ type}$ depends on age



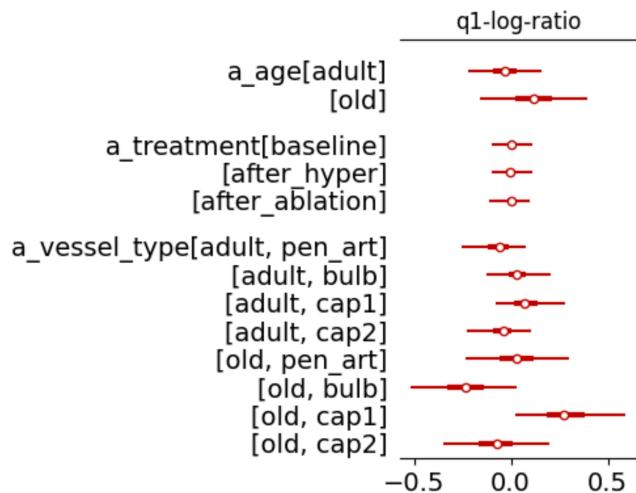
Whisker stimulation log ratios



Model fit

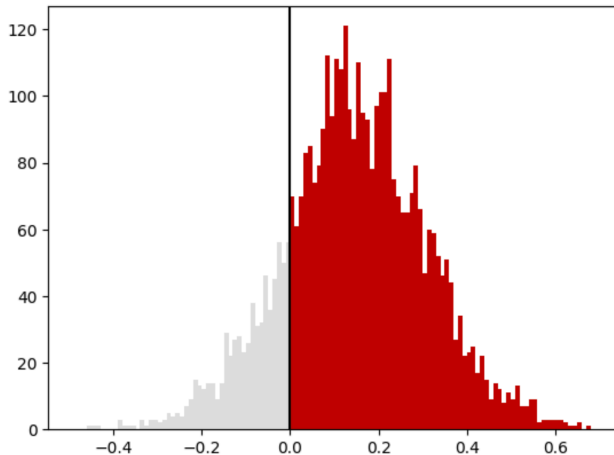


Non-mouse effects



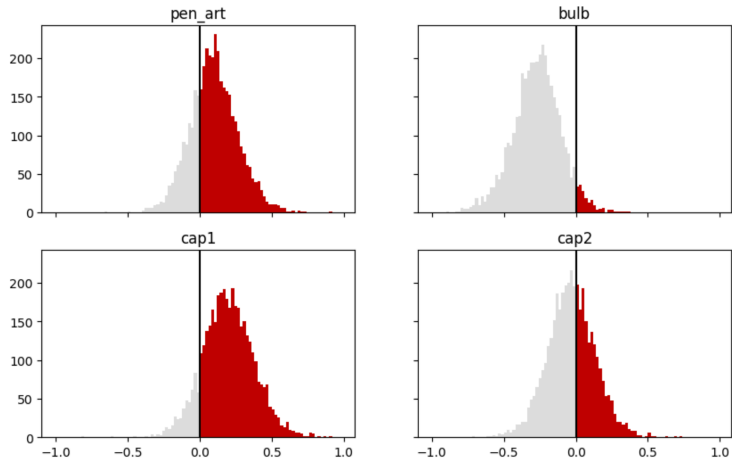
Age effect

Probability of positive effect: 0.82525



Vessel type effect by age

Distributions of effect-size differences (old - adult)



References

- [1] D. J. Spiegelhalter, “Bayesian Graphical Modelling: A Case-Study in Monitoring Health Outcomes,” *Journal of the Royal Statistical Society Series C: Applied Statistics*, vol. 47, no. 1, pp. 115–133, Mar. 1998, doi: [10.1111/1467-9876.00101](https://doi.org/10.1111/1467-9876.00101).
- [2] A. Gelman *et al.*, “Bayesian workflow,” *arXiv:2011.01808 [stat]*, Nov. 2020, Available: <https://arxiv.org/abs/2011.01808>