# Stimulus Identification from fMRI scans

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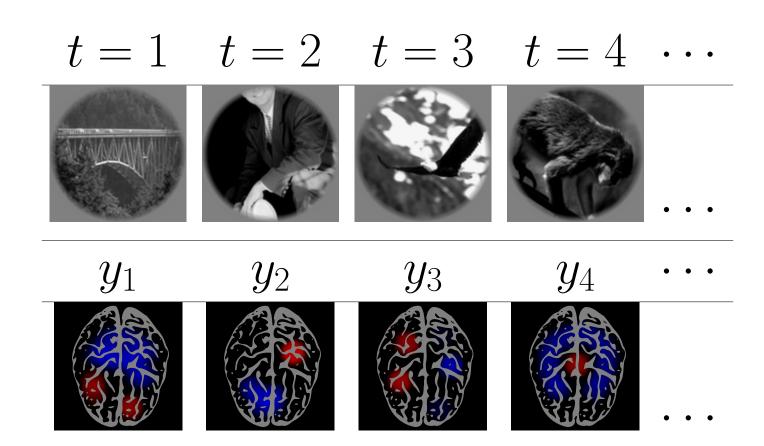
#### Overview

Seeking to explain the processes behind human perception, scientists employ forward models to model the causal relationship between perception of stimuli and neural activity. But how can we measure the quality of these models? Kay et al (2008) introduced the task of identification as a way to demonstrate the fidelity and generalizability of the model.

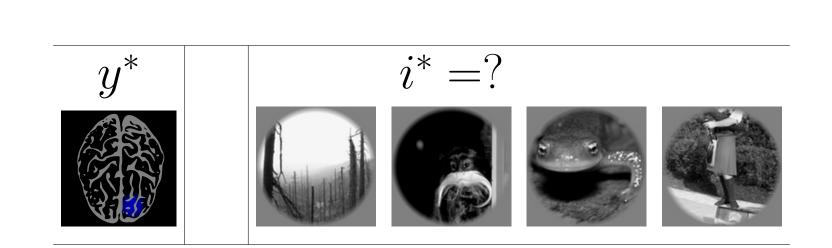
Using the data of Kay et al. as a motivating example, we consider the statistical problem of optimal identification. While identification superficially resembles a classification task (with many classes), it combines the challenge of multivariate regression with high-dimensional discrimination.

### Data

- Sequence of stimuli (pictures) shown at time  $t=1,\ldots,T=3500$
- Record subject's multivariate response  $y_t \in \mathbb{R}^p$ , here  $p \approx 20000$  voxels



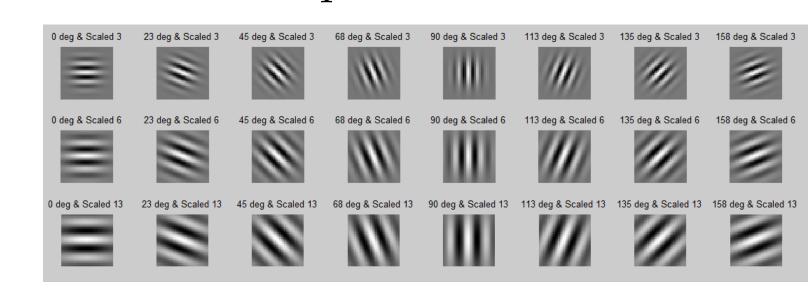
### Identification



- Let S be a set of stimuli, possibly outside the training set! |S| can range from 120 to 10000
- Scientist picks a stimulus  $i^*$  from S and measures the subject's reponse  $y^*$
- Can the statistician  $identify i^* \in S$  from  $y^*$ ?
- Objective: minimize misclassification rate

## Previous approaches

- In order to generalize to new stimuli, we need to find some quantitative representation
- Kay (2008) uses Gabor filters to describe each picture in terms of q=10000 real-valued features



• Notation: write  $Y_{T\times p}$  is a matrix containing the T of recorded responses, and where  $X_{T\times q}$  is the matrix of the *image features* of the corresponding stimuli

Now consider a parametric model

$$Y \sim F_{\theta}(X)$$

Such a *forward model* gives the distribution of the response conditional on the stimuli features; while identification requires the converse.

However, the maximum likelihood (ML) principle can be invoked to identify the stimuli  $i \in S$  "most likely" to have produced  $y^*$ .

Let  $x_i : i \in S$  denote features of the test stimuli, and identify  $y^*$  based on the maximum likelihood (ML) principle

$$i^* = \operatorname{argmax}_i \ell_{\theta}(y^* | x_i)$$

Example. We take the following as a representative approach, combining features of [1] and [2]:

• Assume the normal mutivariate linear model

$$Y \sim N(XB, \Sigma_E)$$
, where  $B \in \mathbb{R}^{q \times p}$ 

- Obtain point estimates of B using elastic net [4], and  $\Sigma_E$  using the sample covariance of residuals with off-diagonal shrinkage
- The ML rule takes the form

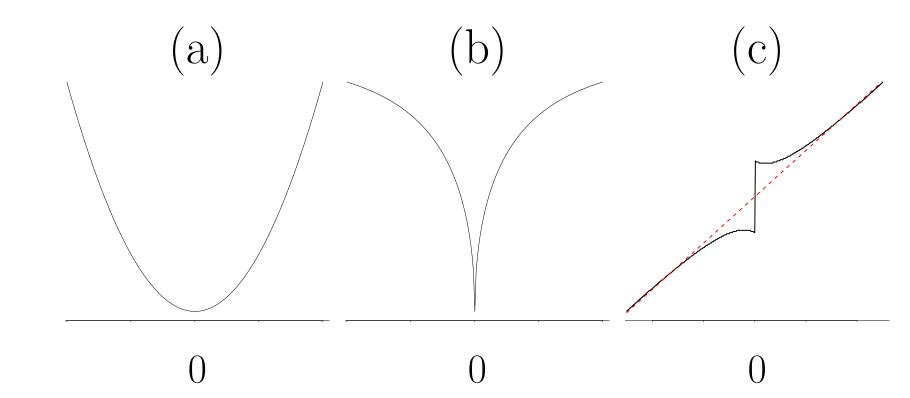
$$i^* = \operatorname{argmin}_i(x_i^T \hat{B} - y^*)^T \hat{\Sigma}_E^{-1}(x_i^T \hat{B} - y^*)$$

## Initial Questions

- Can ML (e.g. the method above) be considered an optimal method for identification in any sense?
- If not, how can we do better? Can we find a good method which is computationally tractable?

#### Limitations of ML

- ML is consistent given the correct model, but can be rather poor in finite samples
- The point estimates  $\hat{B}$  is generally obtained by minimizing  $prediction\ error$ , but the loss function for identification is different
- Perhaps we can find point estimates tailored for the identification loss function. But the nonconvexity makes it difficult!

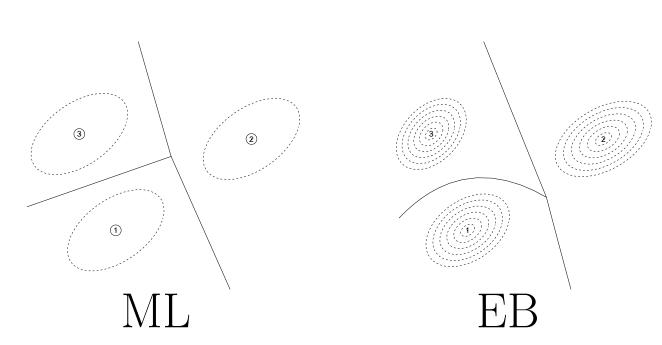


(a) Squared error loss and (b) loss function for identification, as a function of the difference between the true mean signal and the predicted signal.

(c) A p=q=1-dimensional example where ML (actually MAP) fails. The Bayes estimate for identification (black) and the MAP estimate (red) diverge sharply when  $B \sim N(0,1)$ . The same phenomenon can be found in higher dimensions.

## **Empirical Bayes**

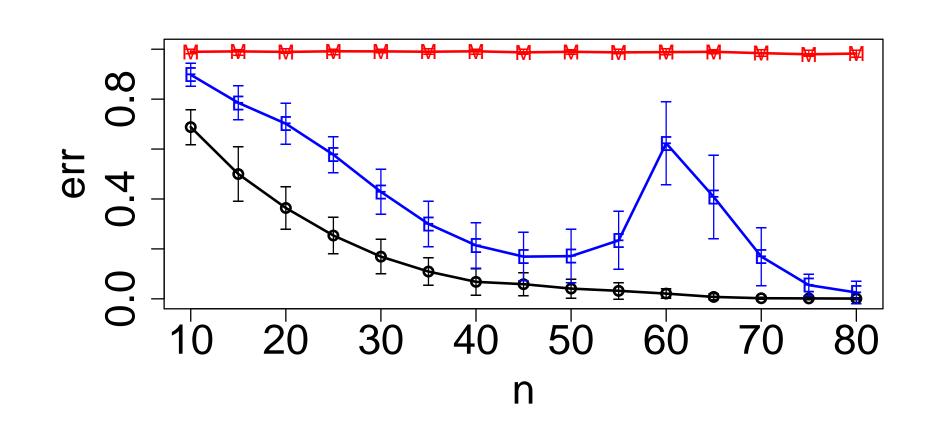
- *Idea*: Unlike ML, the Bayes rule surely optimizes the "correct" objective function. Can we approximate the Bayes rule?
- Empirical Bayes: use the data to estimate the covariances  $\Sigma_B$  and  $\Sigma_E$ , then compute posterior distribution of B
- Assume coefficients of B independent; diagonals of  $\Sigma_B$  can be estimated using any estimate of signal strength, e.g. Eigenprism [3].
- Similar decision rule to ML  $\min(x_i^TB-y^*)^T(\operatorname{Cov}(x_i^TB)+\hat{\Sigma}_E)^{-1}(x_i^TB-y^*)$  but with "added noise" due to uncertainty of B



• Computation: requires inverting  $pq \times pq$  matrix

## Simulation Results

- Parameters p=q=60, random B and  $\Sigma_E$
- Empirical bayes outperforms ML when n < q... however, still unstable!



(E) Empirical Bayes, (M) Maximum likelihood, (o) Bayes risk (knowing true  $\Sigma_B$ ,  $\Sigma_E$ )

## Ongoing Work

- Why does error *increase* with sample size!?
- Refine the crucial step of estimating  $\Sigma_B$
- Required cost of  $O((pq)^3)$  hinders application to real data... develop tractable approximations

## Conclusions

- ML-based approaches rely on point estimates, and hence optimize the wrong objective function
- Empirical bayes achieves better performance by approximating the Bayes rule, but the "empirical" part remains tricky
- Better theoretical understanding is needed to explain why EB succeeds (and sometimes fails)

#### References

- [1] Kay et al. *Nature* (2008)
- [2] Vu et al. Annals of Applied Statistics (2011)
- [3] Janson et al. (2015) http://arxiv.org/abs/1505.02097
- [4] Zou et al. J. R. Statist. Soc. B (2005)

## Acknowledgements

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