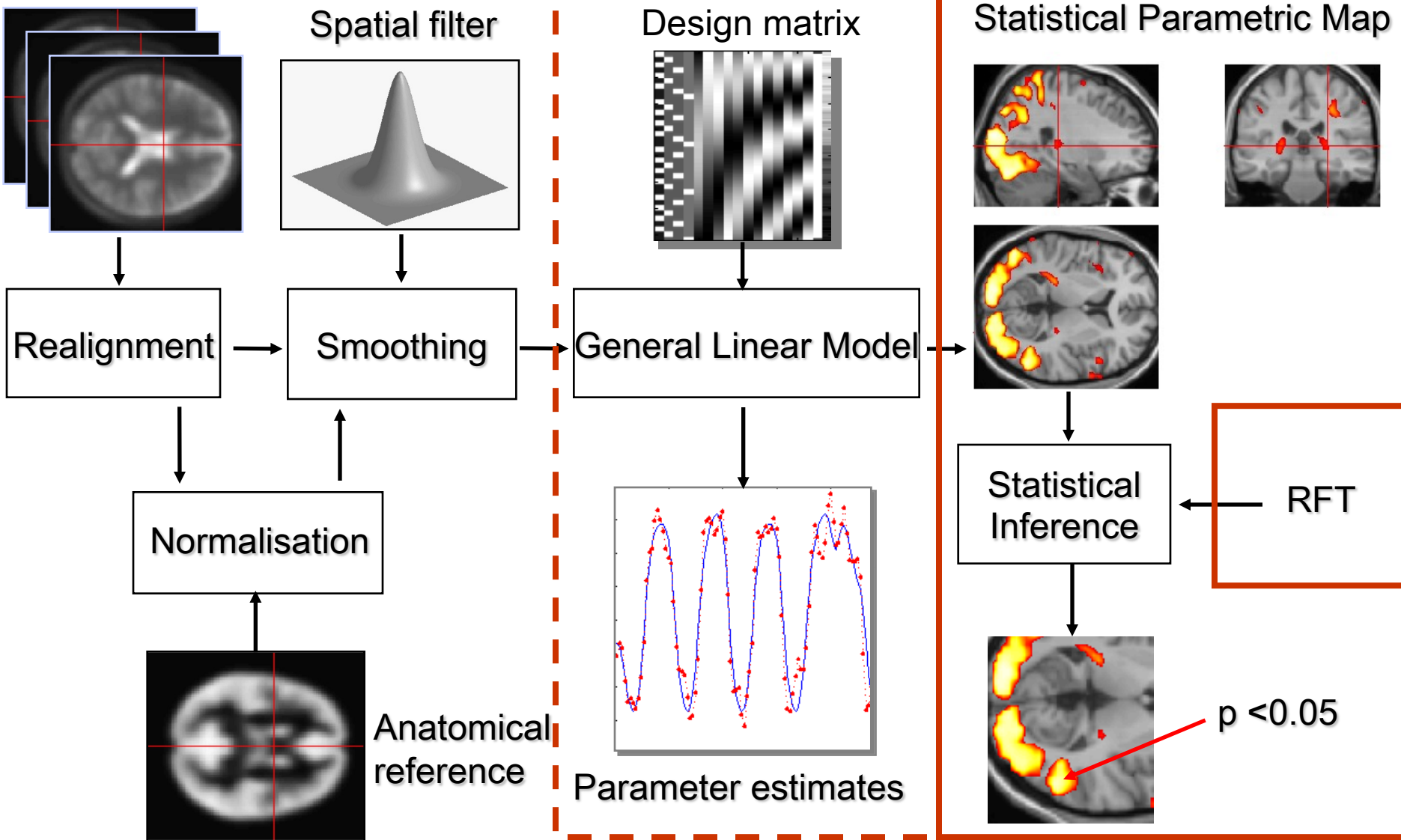


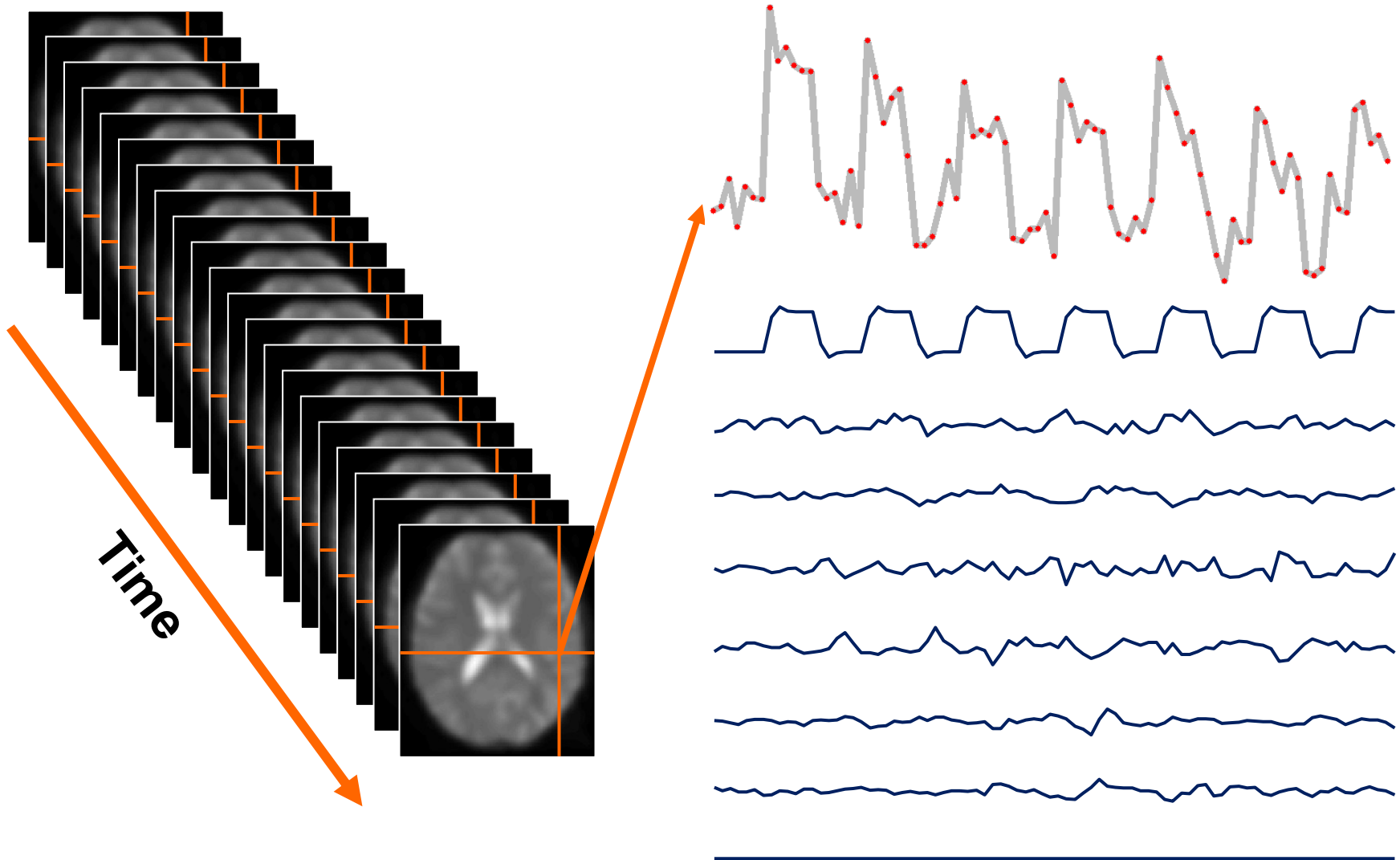
Contrasts & Statistical Inference

Christophe Phillips

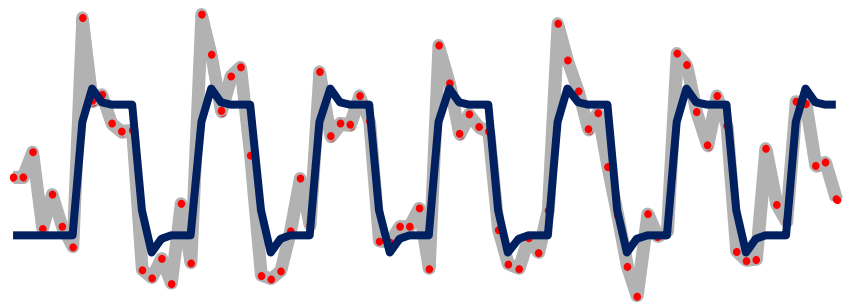
Image time-series



A mass-univariate approach



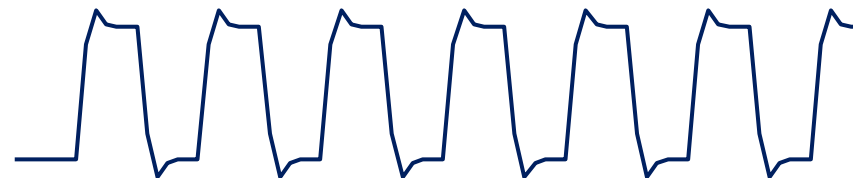
Estimation of the parameters



i.i.d. assumptions: $\varepsilon \sim N(0, \sigma^2 I)$

OLS estimates: $\hat{\beta} = (X^T X)^{-1} X^T y$

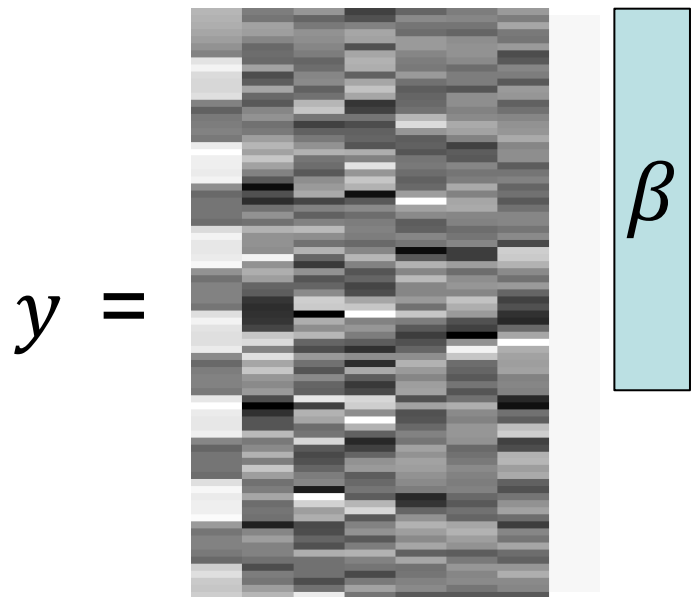
$$\hat{\beta}_1 = 3.9831$$



$$\hat{\beta}_{2-7} = \{0.6871, 1.9598, 1.3902, 166.1007, 76.4770, -64.8189\}$$



$$\hat{\beta}_8 = 131.0040$$



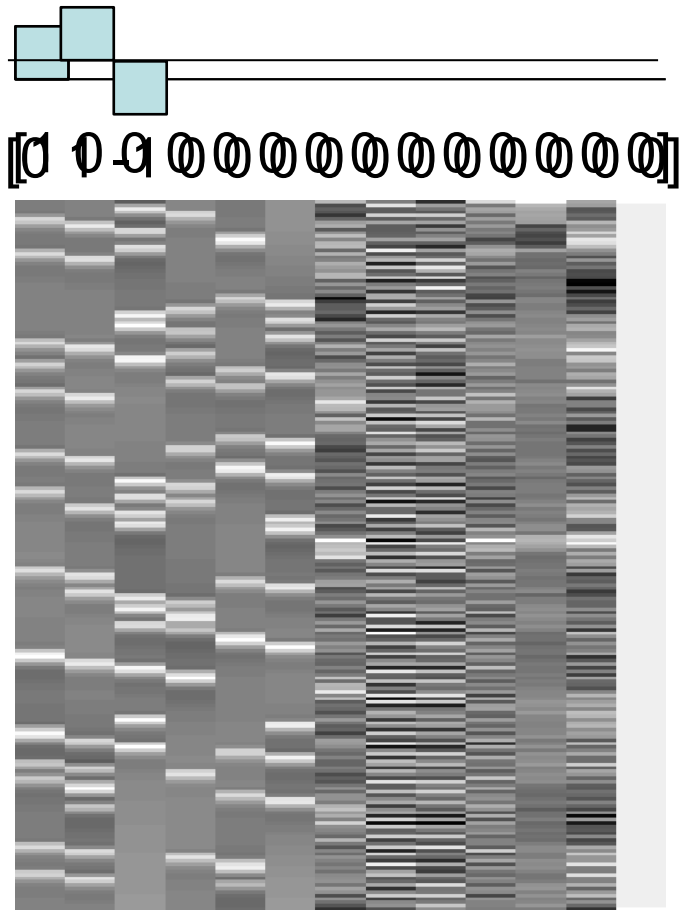
$+\varepsilon$



$$\hat{\beta} \sim N(\beta, \sigma^2 (X^T X)^{-1})$$

$$\hat{\sigma}^2 = \frac{\hat{\varepsilon}^T \hat{\varepsilon}}{N-p}$$

Contrasts



□ A contrast selects a specific effect of interest.

⇒ A contrast c is a vector of length p .

⇒ $c^T \beta$ is a linear combination of regression coefficients β .

$$c = [1 \ 0 \ 0 \ 0 \ \dots]^T$$

$$\begin{aligned} c^T \beta &= \mathbf{1} \times \beta_1 + \mathbf{0} \times \beta_2 + \mathbf{0} \times \beta_3 + \mathbf{0} \times \beta_4 + \dots \\ &= \beta_1 \end{aligned}$$

$$c = [0 \ 1 \ -1 \ 0 \ \dots]^T$$

$$\begin{aligned} c^T \beta &= \mathbf{0} \times \beta_1 + \mathbf{1} \times \beta_2 + \mathbf{-1} \times \beta_3 + \mathbf{0} \times \beta_4 + \dots \\ &= \beta_2 - \beta_3 \end{aligned}$$

$$c^T \hat{\beta} \sim N(c^T \beta, \sigma^2 c^T (X^T X)^{-1} c)$$

Hypothesis Testing

To test a hypothesis, we construct “test statistics”.

□ Null Hypothesis H_0

Typically what we want to disprove (no effect).

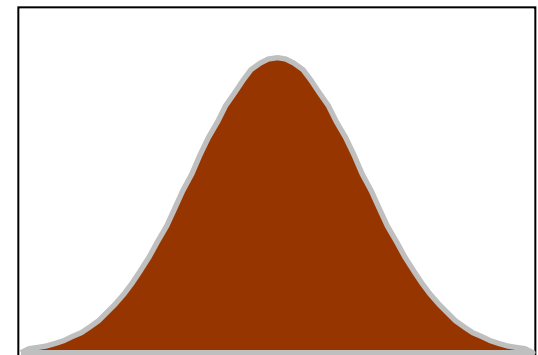
⇒ The Alternative Hypothesis H_A expresses outcome of interest.

□ Test Statistic T

The test statistic summarises evidence about H_0 .

Typically, test statistic is small in magnitude when the hypothesis H_0 is true and large when false.

⇒ We need to know the distribution of T under the null hypothesis.



Null Distribution of T

Hypothesis Testing

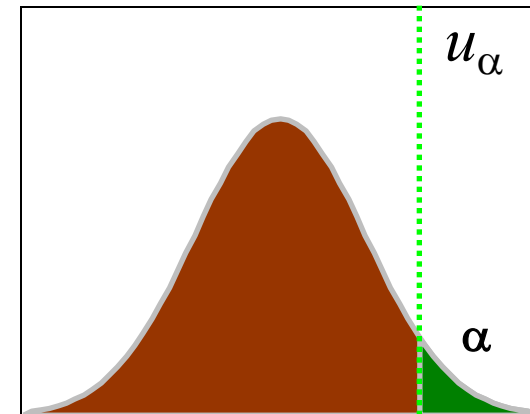
□ Significance level α :

Acceptable *false positive rate* α .

⇒ threshold u_α

Threshold u_α controls the false positive rate

$$\alpha = p(T > u_\alpha | H_0)$$



Null Distribution of T

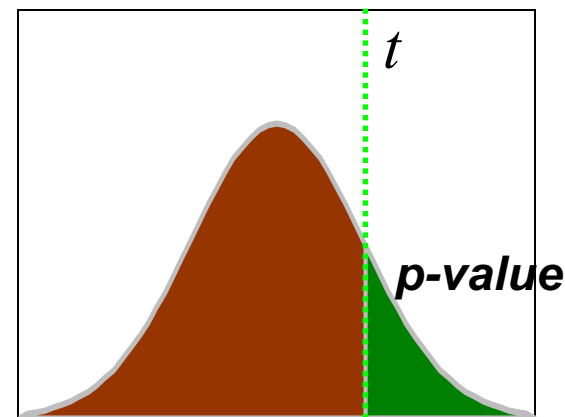
□ Conclusion about the hypothesis:

We reject the null hypothesis in favour of the alternative hypothesis if $t > u_\alpha$

□ *p-value*:

A *p-value* summarises evidence against H_0 .

This is the chance of observing value more extreme than t under the null hypothesis.



Null Distribution of T

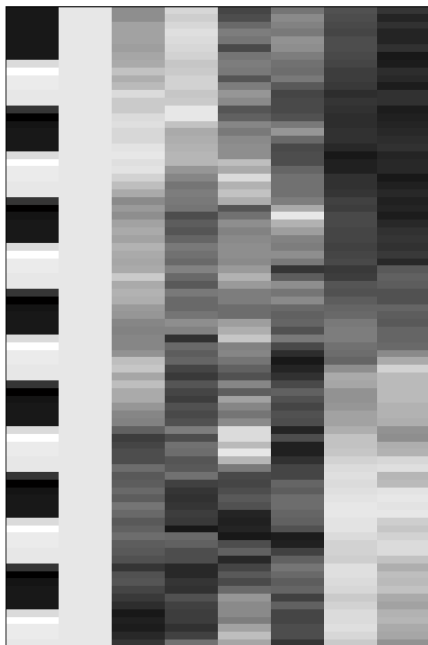
$$p(T > t | H_0)$$

T-test - one dimensional contrasts – SPM{t}

$$c^T = 1\ 0\ 0\ 0\ 0\ 0\ 0\ 0$$



$\beta_1\ \beta_2\ \beta_3\ \beta_4\ \beta_5\ \dots$



Question: box-car amplitude > 0 ?

=

$$\beta_1 = c^T \beta > 0 ?$$

Null hypothesis:

$$H_0: c^T \beta = 0$$

*contrast of
estimated
parameters*

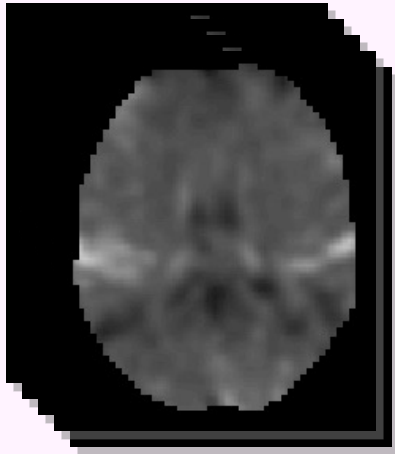
Test statistic:

$$T = \frac{\text{contrast of estimated parameters}}{\sqrt{\text{variance estimate}}}$$

$$T = \frac{c^T \hat{\beta}}{\sqrt{\text{var}(c^T \hat{\beta})}} = \frac{c^T \hat{\beta}}{\sqrt{\hat{\sigma}^2 c^T (X^T X)^{-1} c}} \sim t_{N-p}$$

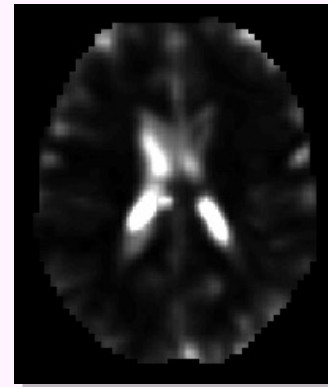
T-contrast in SPM

- For a given contrast c :



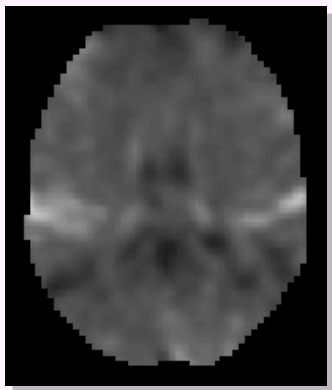
beta_???? images

$$\hat{\beta} = (X^T X)^{-1} X^T y$$



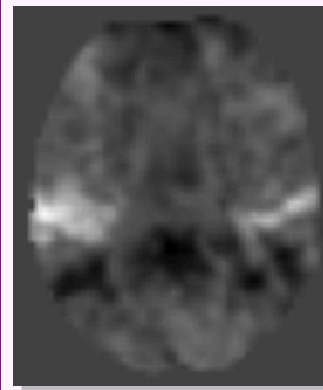
ResMS image

$$\hat{\sigma}^2 = \frac{\hat{\varepsilon}^T \hat{\varepsilon}}{N - p}$$



con_???? image

$$c^T \hat{\beta}$$

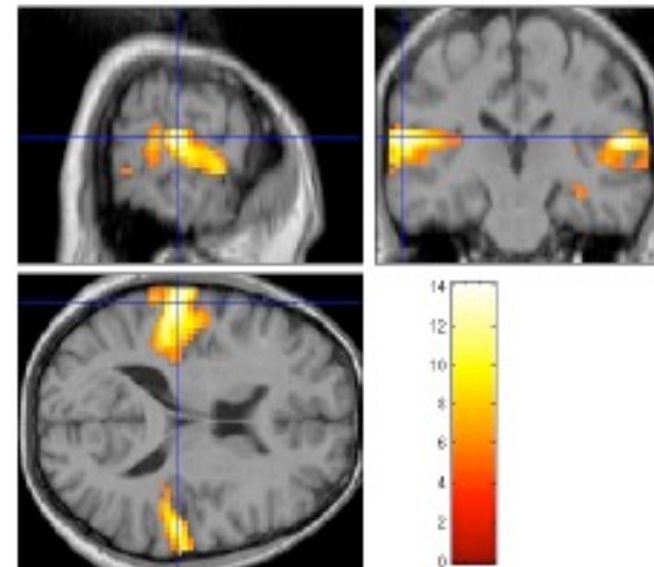


spmT_???? image

SPM{t}

T-test: a simple example

Passive word listening versus rest



SPMresults:

Height threshold $T = 3.2057$ $\{p < 0.001\}$

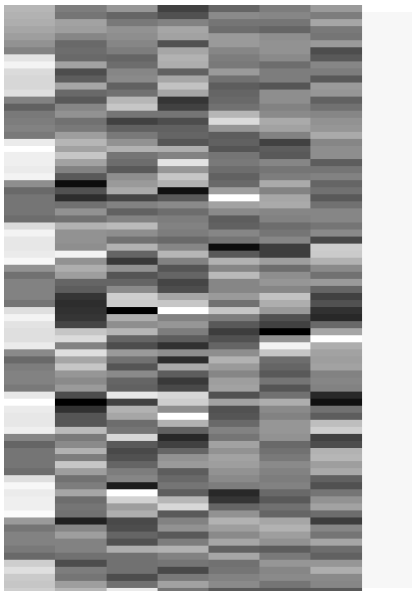
voxel-level			mm mm mm		
T	(Z)	$p_{\text{uncorrected}}$			
13.94	Inf	0.000	-63	-27	15
12.04	Inf	0.000	-48	-33	12
11.82	Inf	0.000	-66	-21	6
13.72	Inf	0.000	57	-21	12
12.29	Inf	0.000	63	-12	-3
9.89	7.83	0.000	57	-39	6
7.39	6.36	0.000	36	-30	-15
6.84	5.99	0.000	51	0	48
6.36	5.65	0.000	-63	-54	-3
6.19	5.53	0.000	-30	-33	-18
5.96	5.36	0.000	36	-27	9
5.84	5.27	0.000	-45	42	9
5.44	4.97	0.000	48	27	24
5.32	4.87	0.000	36	-27	42

Q: activation during listening ?

Null hypothesis: $\beta_1 = 0$

$$t = \frac{c^T \hat{\beta}}{\sqrt{\text{var}(c^T \hat{\beta})}}$$

$$c^T = [1 \ 0 \ 0 \ 0 \ 0 \ 0 \ 0 \ 0 \ 0]$$



T-test: summary

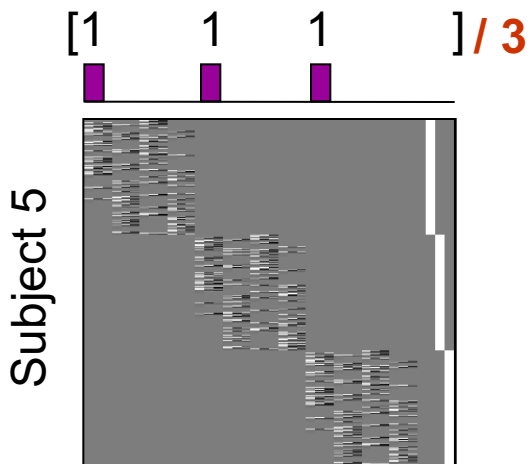
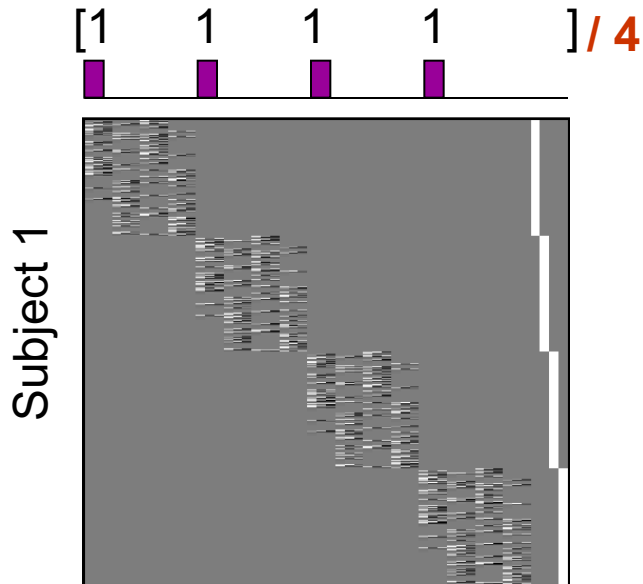
□ T-test is a *signal-to-noise* measure (ratio of estimate to standard deviation of estimate).

□ Alternative hypothesis:

$$H_0: c^T \beta = 0 \quad \text{vs} \quad H_A: c^T \beta > 0$$

□ T-contrasts are simple combinations of the betas; the T-statistic does not depend on the scaling of the regressors or the scaling of the contrast.

Scaling issue



$$T = \frac{c^T \hat{\beta}}{\sqrt{\text{var}(c^T \hat{\beta})}} = \frac{c^T \hat{\beta}}{\sqrt{\hat{\sigma}^2 c^T (X^T X)^{-1} c}}$$

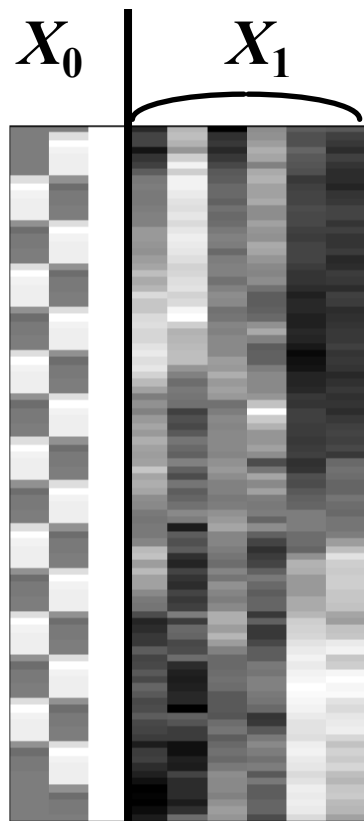
- ❑ The T -statistic does not depend on the scaling of the regressors.
- ❑ The T -statistic does not depend on the scaling of the contrast.
- ❑ Contrast $c^T \hat{\beta}$ depends on scaling.
- Be careful of the interpretation of the contrasts $c^T \hat{\beta}$ themselves (eg, for a second level analysis):

sum \neq average

F-test - the extra-sum-of-squares principle

- Model comparison:

Null Hypothesis H_0 : True model is X_0 (reduced model)



$$RSS$$

$$\sum \hat{\varepsilon}_{full}^2$$



$$RSS_0$$

$$\sum \hat{\varepsilon}_{reduced}^2$$

Test statistic: ratio of explained variability and unexplained variability (error)

$$F \propto \frac{RSS_0 - RSS}{RSS}$$

$$F \propto \frac{ESS}{RSS} \sim F_{v_1, v_2}$$

$$v_1 = \text{rank}(X) - \text{rank}(X_0)$$

$$v_2 = N - \text{rank}(X)$$

Full model ?

or Reduced model?

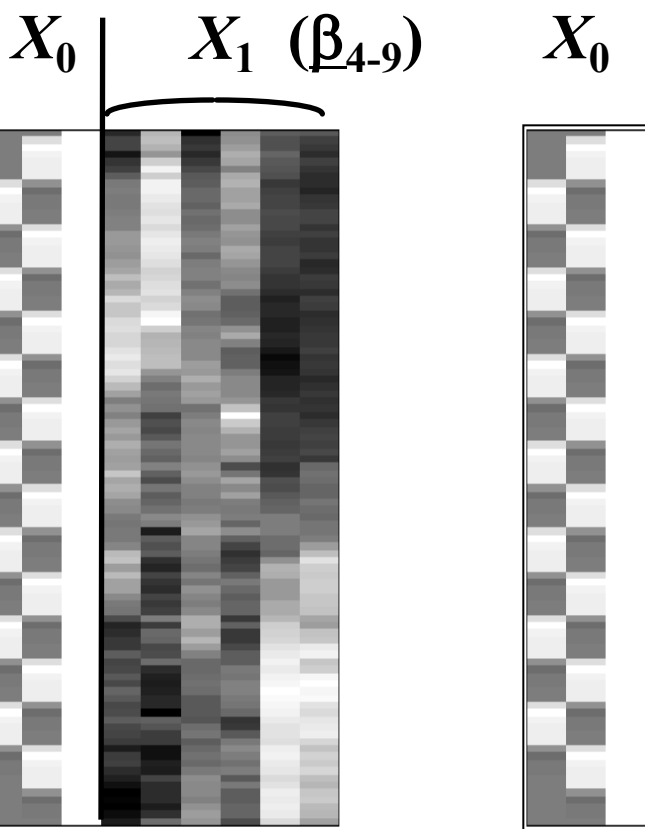
F-test - multidimensional contrasts – SPM{F}

Tests multiple linear hypotheses:

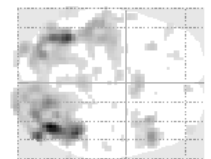
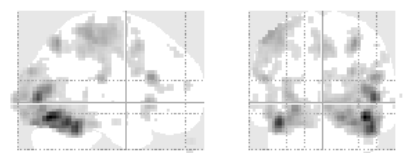
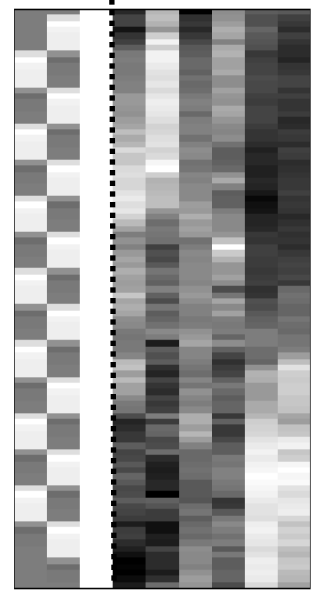
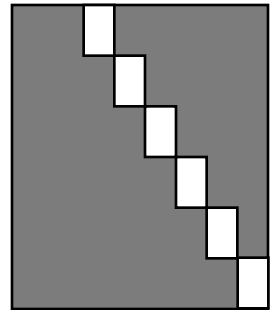
H_0 : True model is X_0

H_0 : $\beta_4 = \beta_5 = \dots = \beta_9 = 0$

test H_0 : $c^T \beta = 0$?



$$c^T = \begin{matrix} 0 & 0 & 0 & 1 & 0 & 0 & 0 & 0 & 0 \\ 0 & 0 & 0 & 0 & 1 & 0 & 0 & 0 & 0 \\ 0 & 0 & 0 & 0 & 0 & 1 & 0 & 0 & 0 \\ 0 & 0 & 0 & 0 & 0 & 0 & 1 & 0 & 0 \\ 0 & 0 & 0 & 0 & 0 & 0 & 0 & 1 & 0 \\ 0 & 0 & 0 & 0 & 0 & 0 & 0 & 0 & 1 \end{matrix}$$

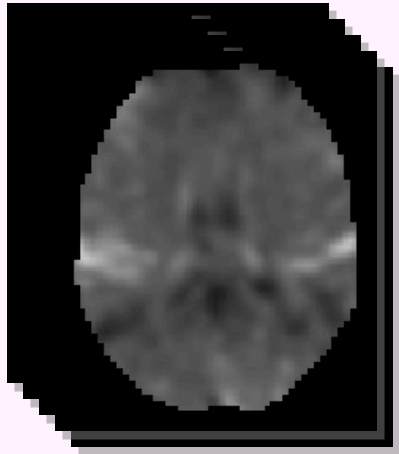


SPM{ $F_{6,322}$ }

Full model?

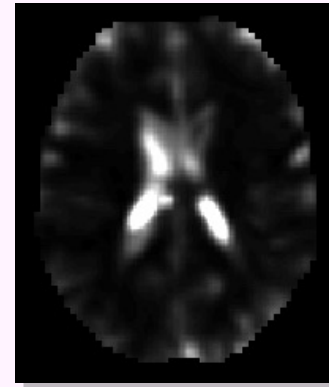
Reduced model?

F-contrast in SPM



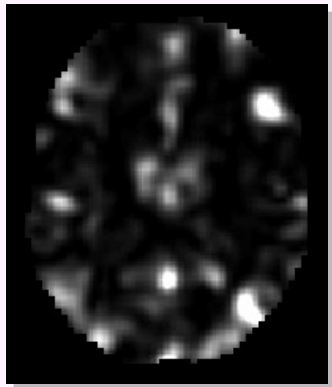
beta_???? images

$$\hat{\beta} = (X^T X)^{-1} X^T y$$



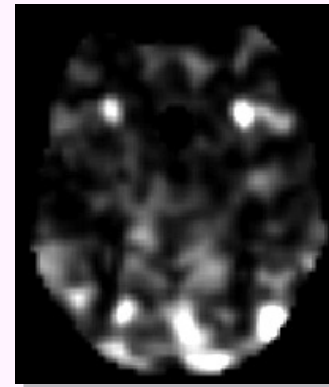
ResMS image

$$\hat{\sigma}^2 = \frac{\hat{\varepsilon}^T \hat{\varepsilon}}{N - p}$$



ess_???? images

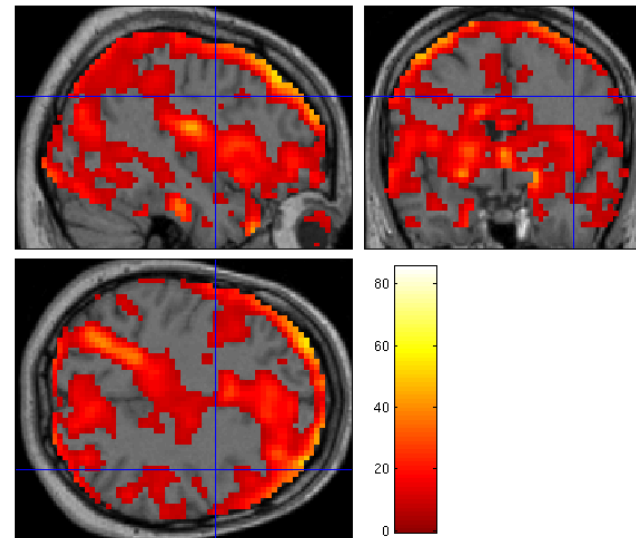
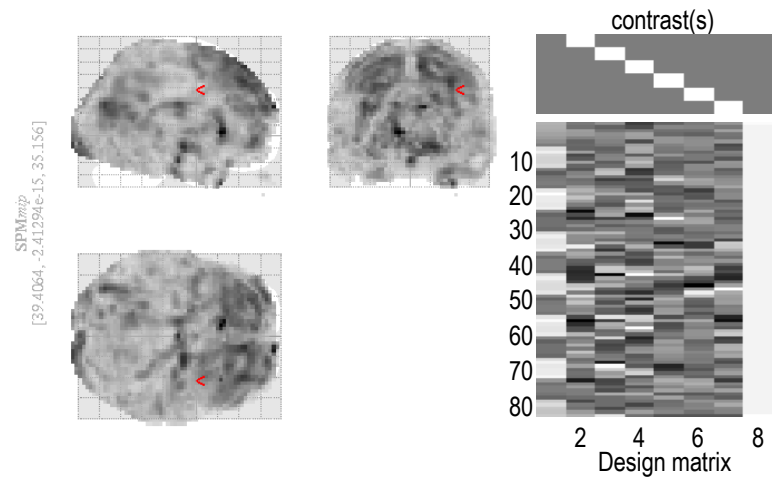
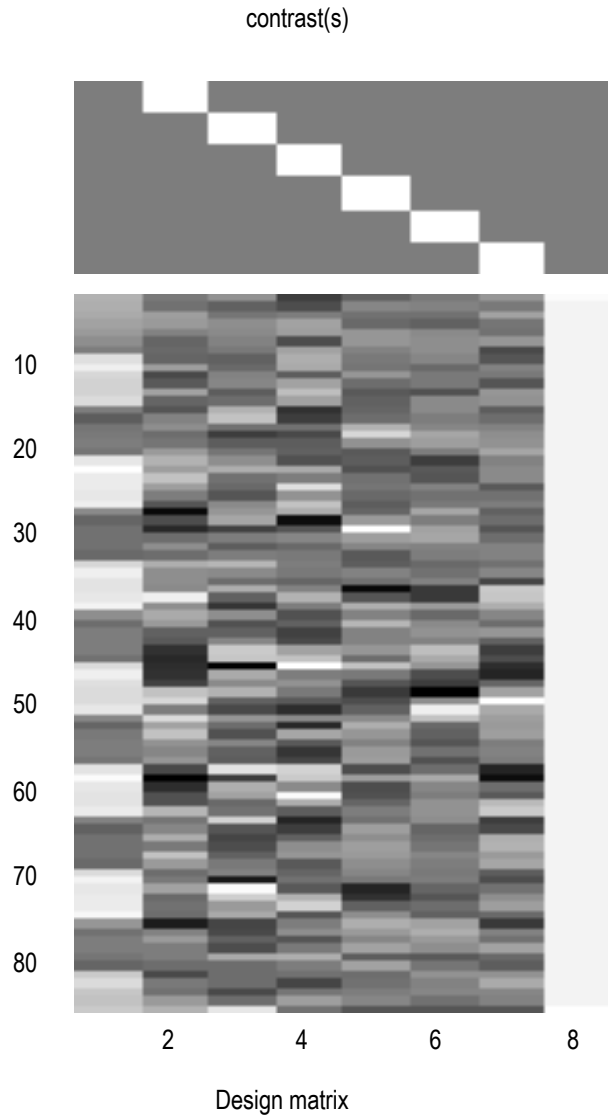
$$(RSS_0 - RSS)$$



spmF_???? images

$$SPM\{F\}$$

F-test example: movement related effects

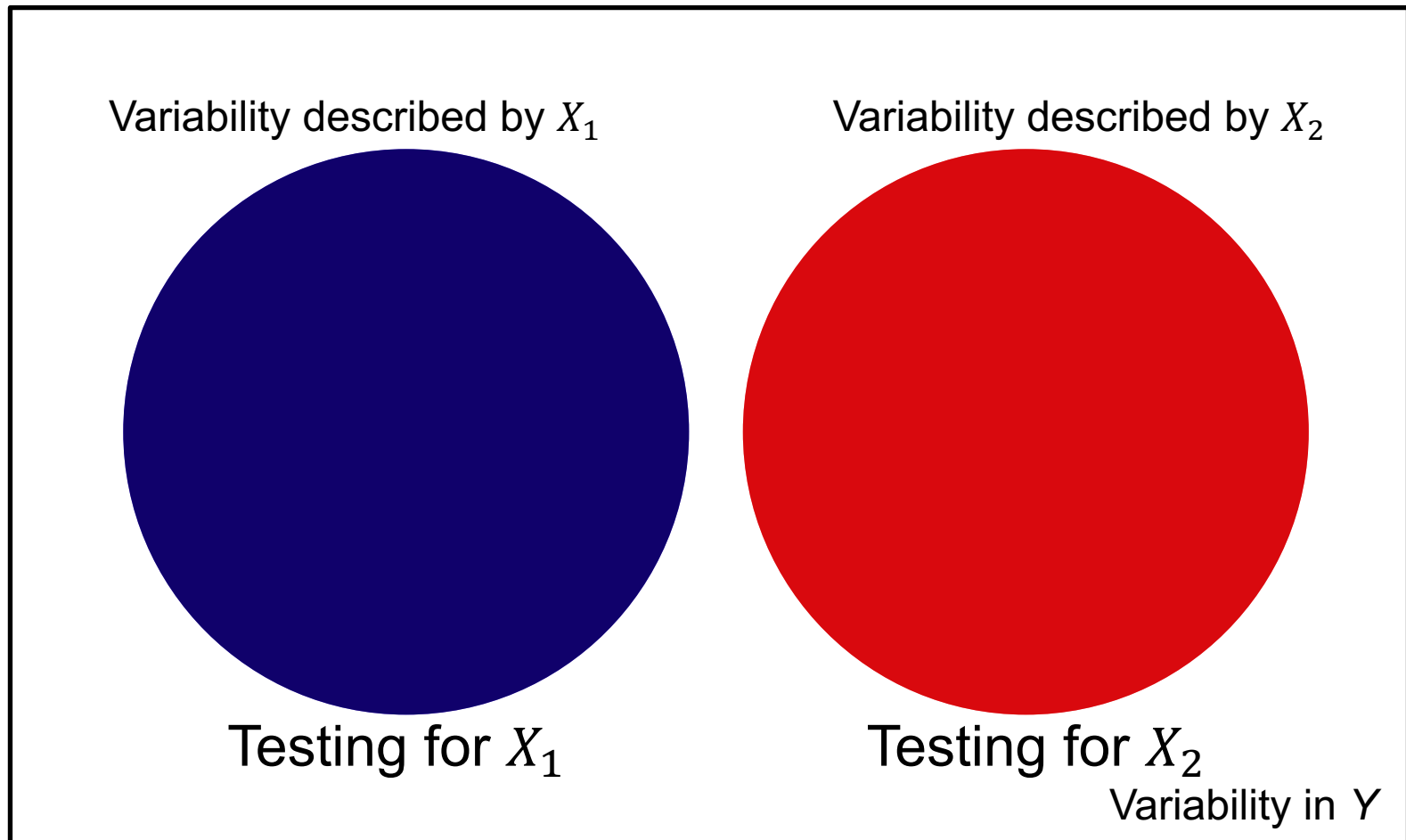


F-test: summary

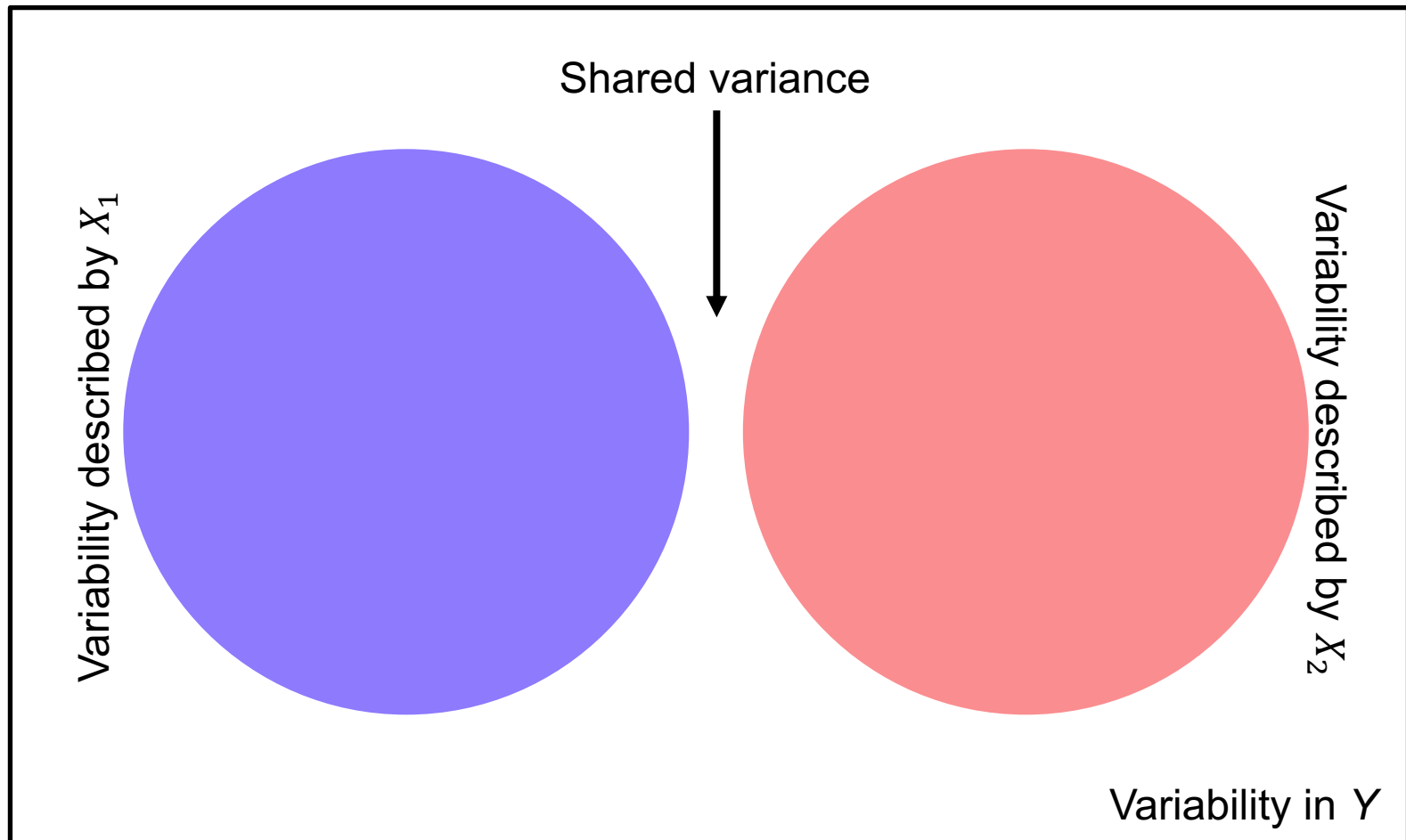
- ❑ F-tests can be viewed as testing for the additional variance explained by a larger model w.r.t. a simpler (***nested***) model
 ⇒ ***model comparison***.
- ❑ F tests a weighted **sum of squares** of one or several combinations of the regression coefficients β .
- ❑ In practice, we don't have to explicitly separate X into $[X_1 X_2]$ thanks to **multidimensional contrasts**.
- ❑ Hypotheses:

$\begin{bmatrix} 1 & 0 & 0 & 0 \\ 0 & 1 & 0 & 0 \\ 0 & 0 & 1 & 0 \\ 0 & 0 & 0 & 0 \end{bmatrix}$	<p>Null Hypothesis $H_0 : \beta_1 = \beta_2 = \beta_3 = 0$</p> <p>Alternative Hypothesis $H_A : \text{at least one } \beta_k \neq 0$</p>
--	--
- ❑ In testing uni-dimensional contrast with an F -test, for example $\beta_1 - \beta_2$, the result will be the same as testing $\beta_2 - \beta_1$. It will be exactly the square of the t -test, testing for both positive and negative effects.

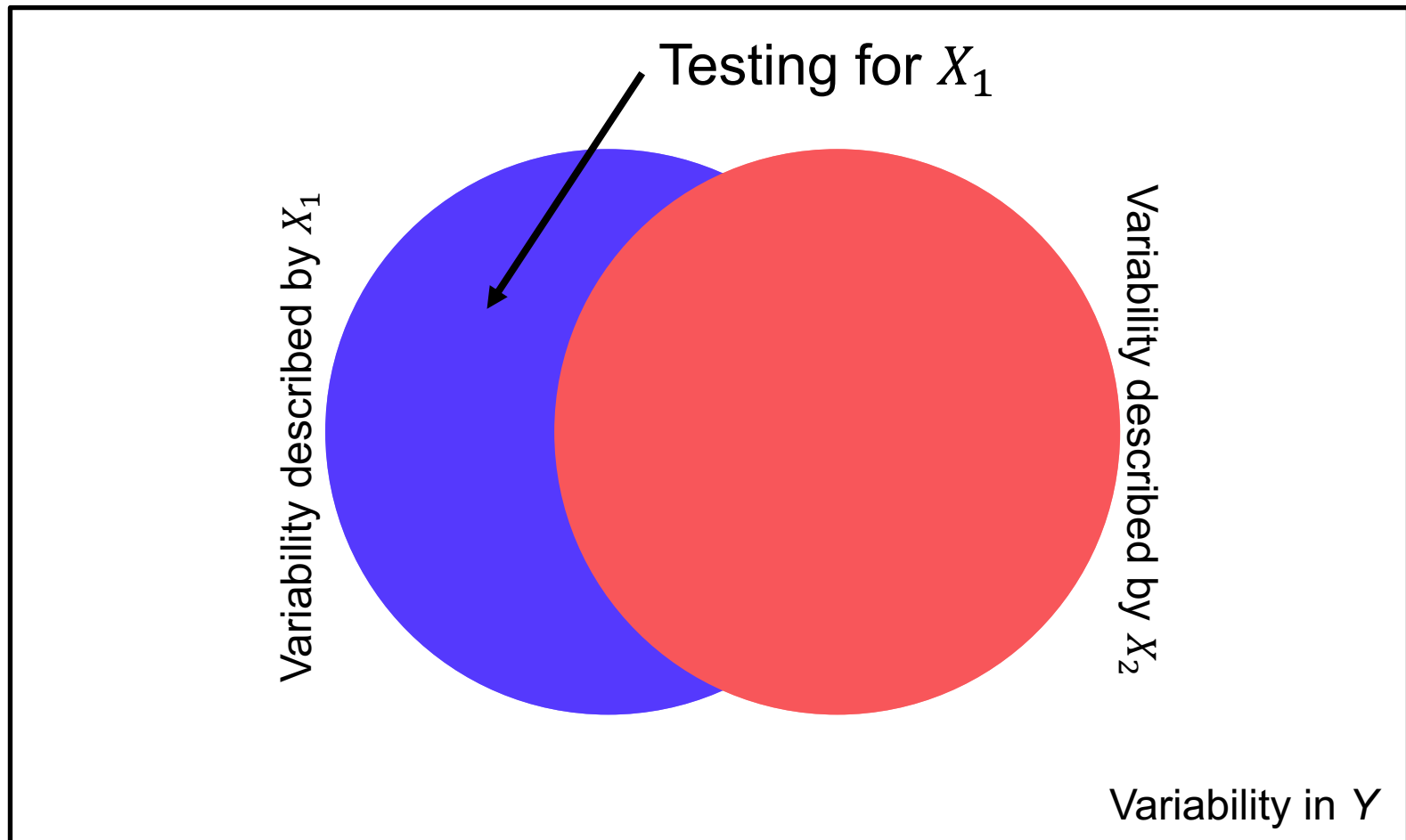
Orthogonal regressors



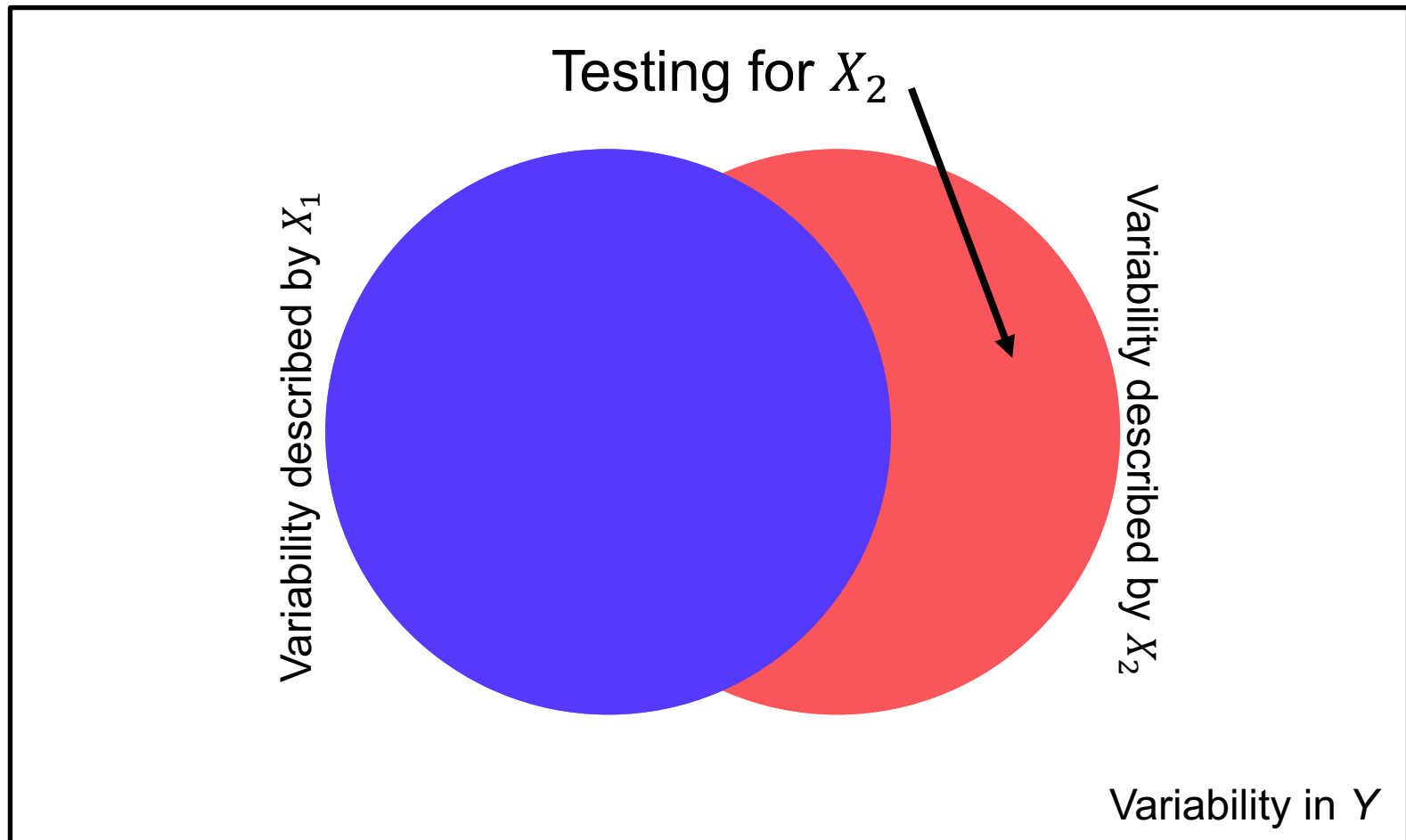
Correlated regressors



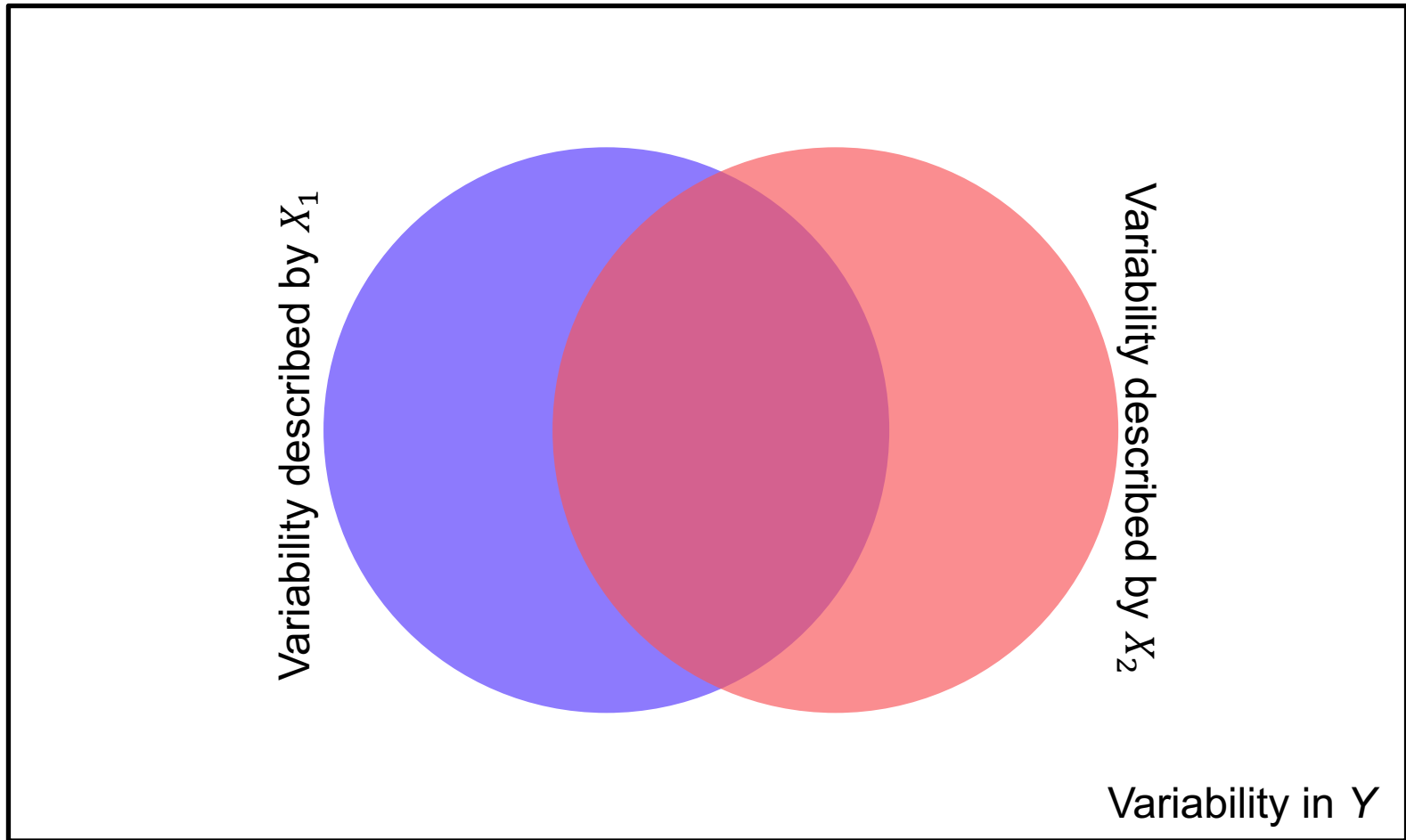
Correlated regressors



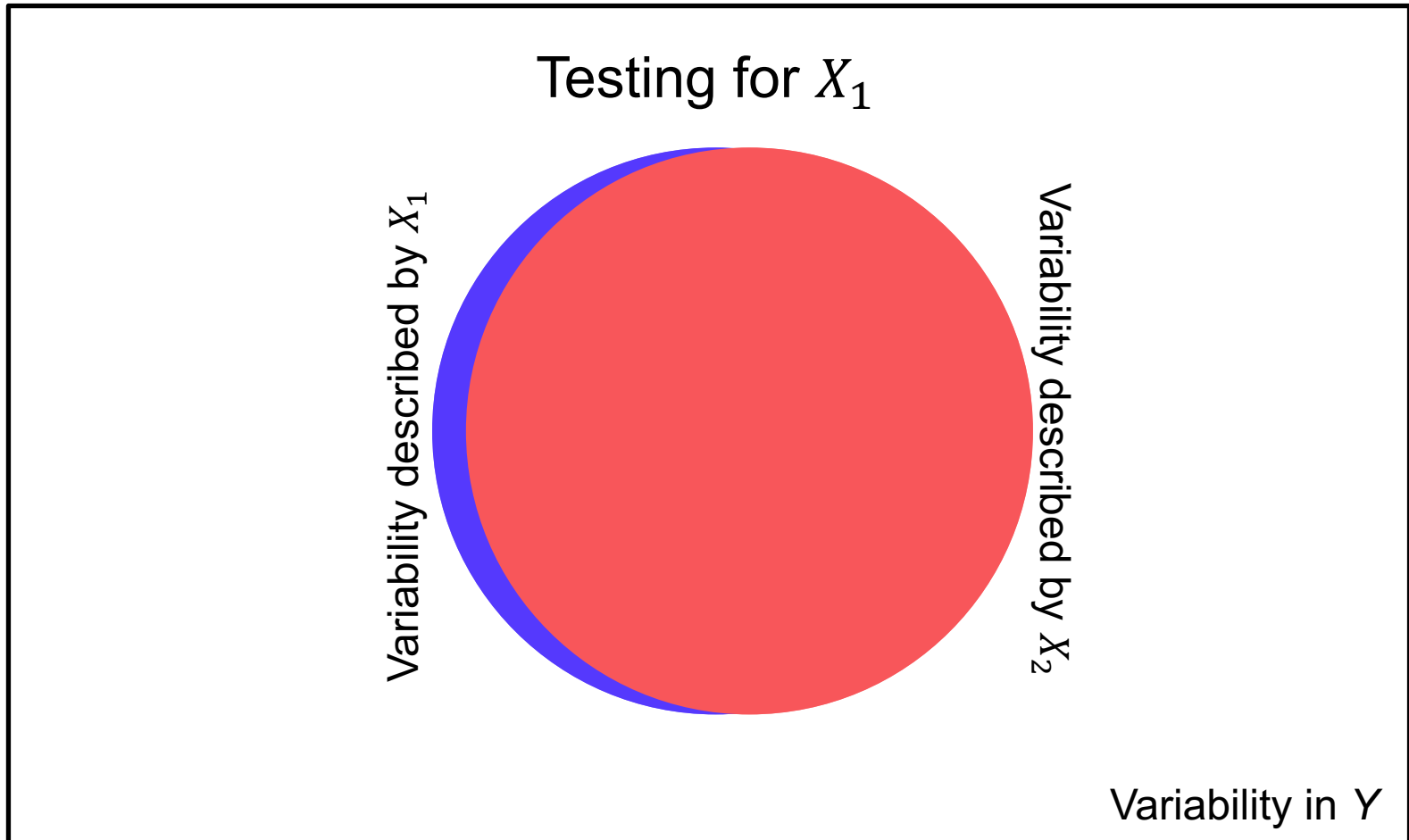
Correlated regressors



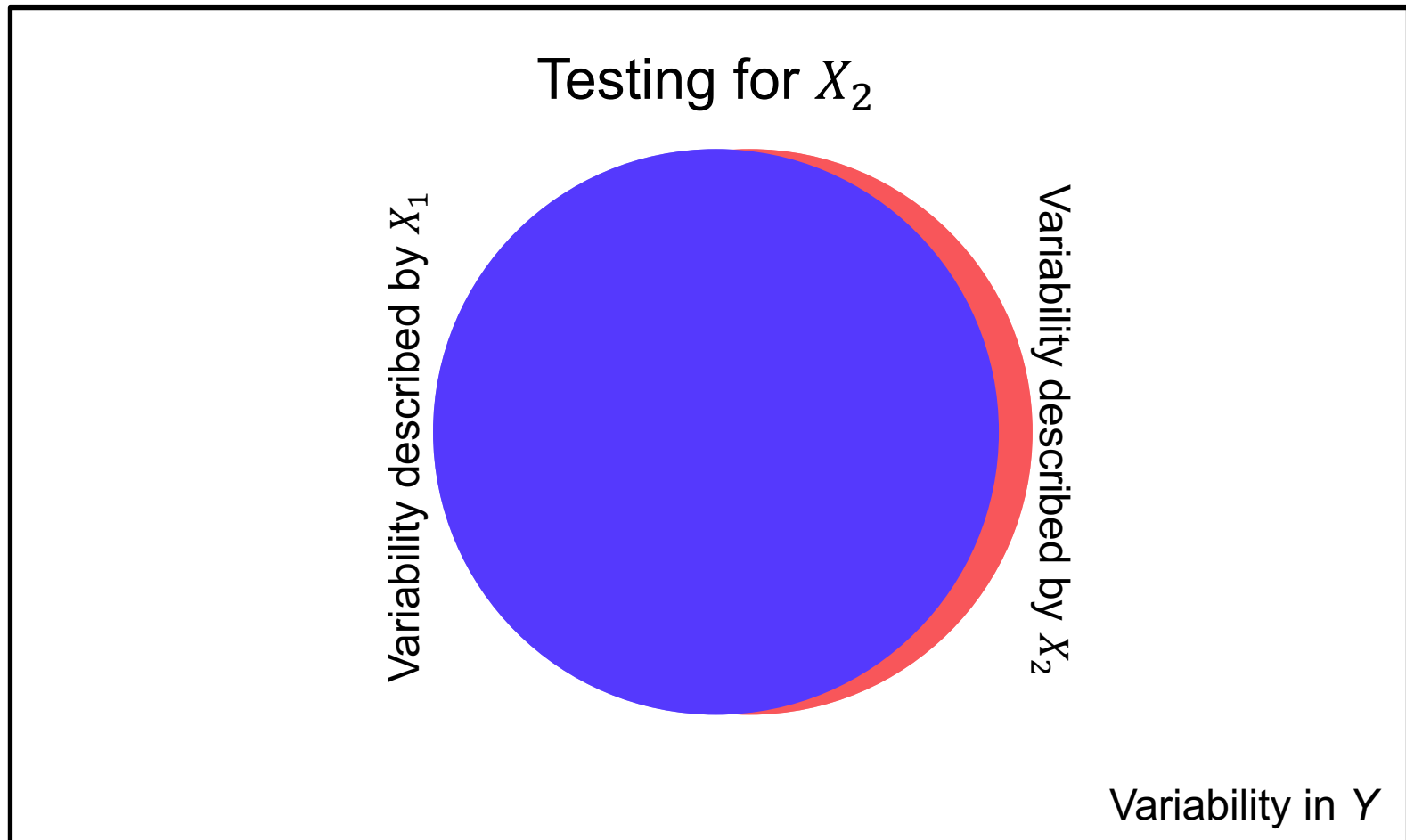
Correlated regressors



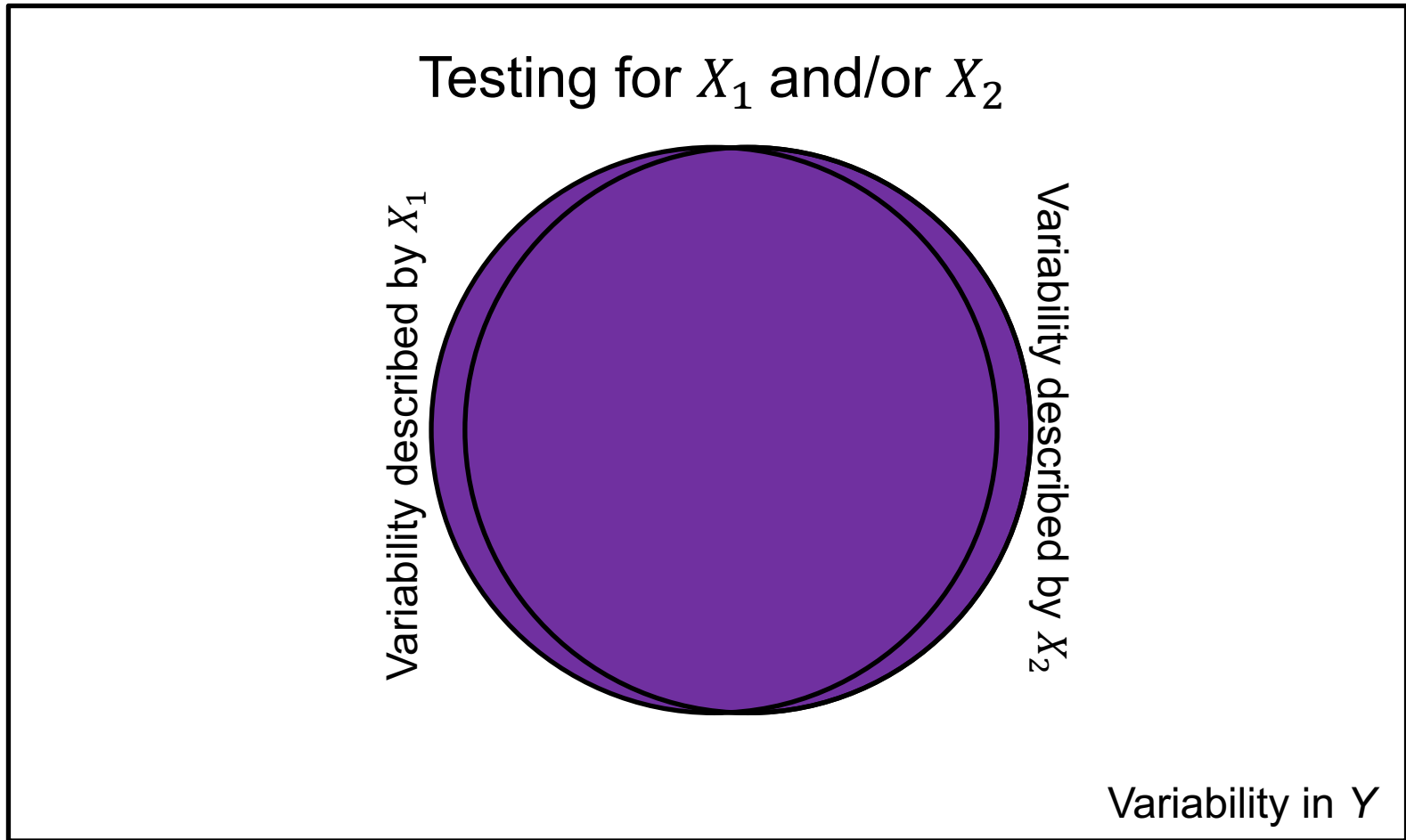
Correlated regressors



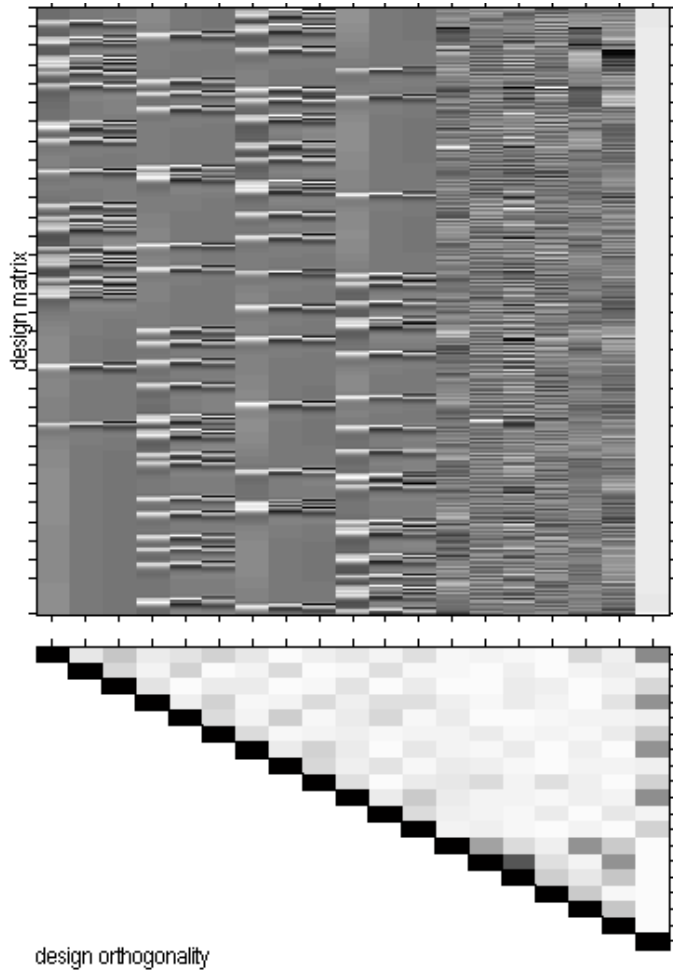
Correlated regressors



Correlated regressors



Design orthogonality

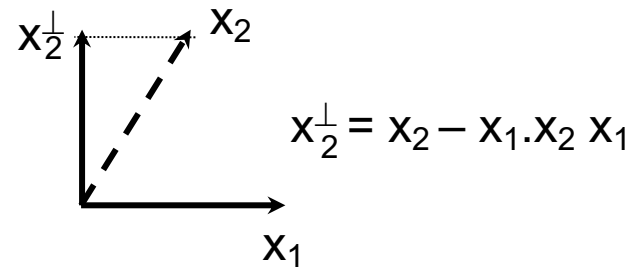
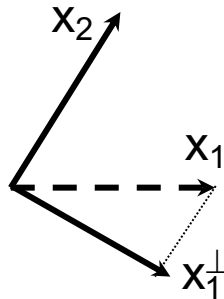
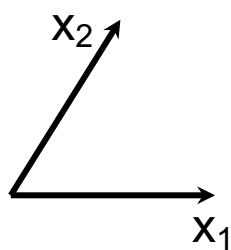


- For each pair of columns of the design matrix, the orthogonality matrix depicts the magnitude of the **cosine of the angle** between them, with the range 0 to 1 mapped from white to black.
- If both vectors have **zero mean** then the cosine of the angle between the vectors is the same as the **correlation** between the two variates.

Measure : abs. value of cosine of angle between columns of design matrix
Scale : black - colinear (cos=+1/-1)
 white - orthogonal (cos=0)
 gray - not orthogonal or colinear

Correlated regressors: summary

- We implicitly test for an **additional** effect only. When testing for the first regressor, we are effectively removing the part of the signal that can be accounted for by the second regressor:
 ⇒ **implicit orthogonalisation**.



- Orthogonalisation = decorrelation. Parameters and test on the non modified regressor change.
 Rarely solves the problem as it requires assumptions about which regressor to uniquely attribute the common variance.
 ⇒ change regressors (i.e. design) instead, e.g. factorial designs.
 ⇒ use F-tests to assess overall significance.
- Original regressors may not matter: it's the contrast you are testing which should be as decorrelated as possible from the rest of the design matrix

Bibliography:

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- ❑ *Plane Answers to Complex Questions: The Theory of Linear Models*. R. Christensen, Springer, 1996.

- ❑ *Statistical parametric maps in functional imaging: a general linear approach*. K.J. Friston et al, Human Brain Mapping, 1995.

- ❑ *Ambiguous results in functional neuroimaging data analysis due to covariate correlation*. A. Andrade et al., NeuroImage, 1999.

- ❑ Estimating efficiency a priori: a comparison of blocked and randomized designs. A. Mechelli et al., NeuroImage, 2003.

