

Introduction to **GWAS**

Linkage Disequilibrium and Linear Regression

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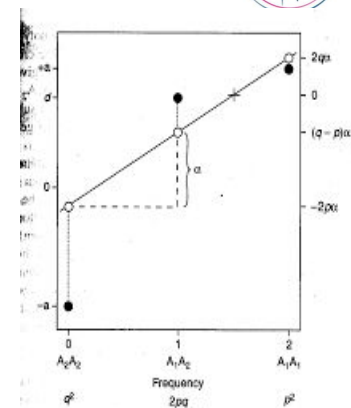
- What marker is associated with the phenotype?
- What individual is genetically more susceptible?

trait	id	snp1	snp2	snp3	snp4	snp5	snp6
9.616	1	Aa	Aa	AA	Aa	Aa	AA
10.140	2	aa	AA	AA	AA	AA	AA
10.687	3	AA	AA	aa	AA	aa	Aa
9.906	4	Aa	aa	Aa	AA	AA	aa

- What marker is associated with the phenotype?
- What individual is genetically more susceptible?

Allele substitution effect

The effect that the presence of a copy of an allele has on the phenotype (regarding the reference allele).



$$f(A) = \text{mean}(Aa) - \text{mean}(aa)$$

snp1	snp2	snp3	snp4	snp5	snp6
-0.4	0	0	0.2	-0.4	0.4

- What marker is associated with the phenotype?
- What individual is genetically more susceptible?

Allele substitution effect

$$P = G + E \text{ (may dominate over } G \text{)}$$

snp1	snp2	snp3	snp4	snp5	snp6
-0.4	0	0	0.2	-0.4	0.4

GWAS aims to discover the loci with causal effect and their magnitude

- What marker is associated with the phenotype?
- What individual is genetically more susceptible?

id (pop. $\mu=10$)	Genetic merit	trait	id
1	0.20	9.616	1
2	0.40	10.140	2
3	0.00	10.687	3
4	-0.80	9.906	4

Predicting the genetic response, the phenotype or the disease susceptibility is approached in Genome-Wide Prediction

Genome-wide association studies (**GWAS**)

- Based on linear regression models (mathematics, linear algebra, statistics)
- Uses linkage disequilibrium between genomic markers and genes

Genome-wide association studies (**GWAS**)

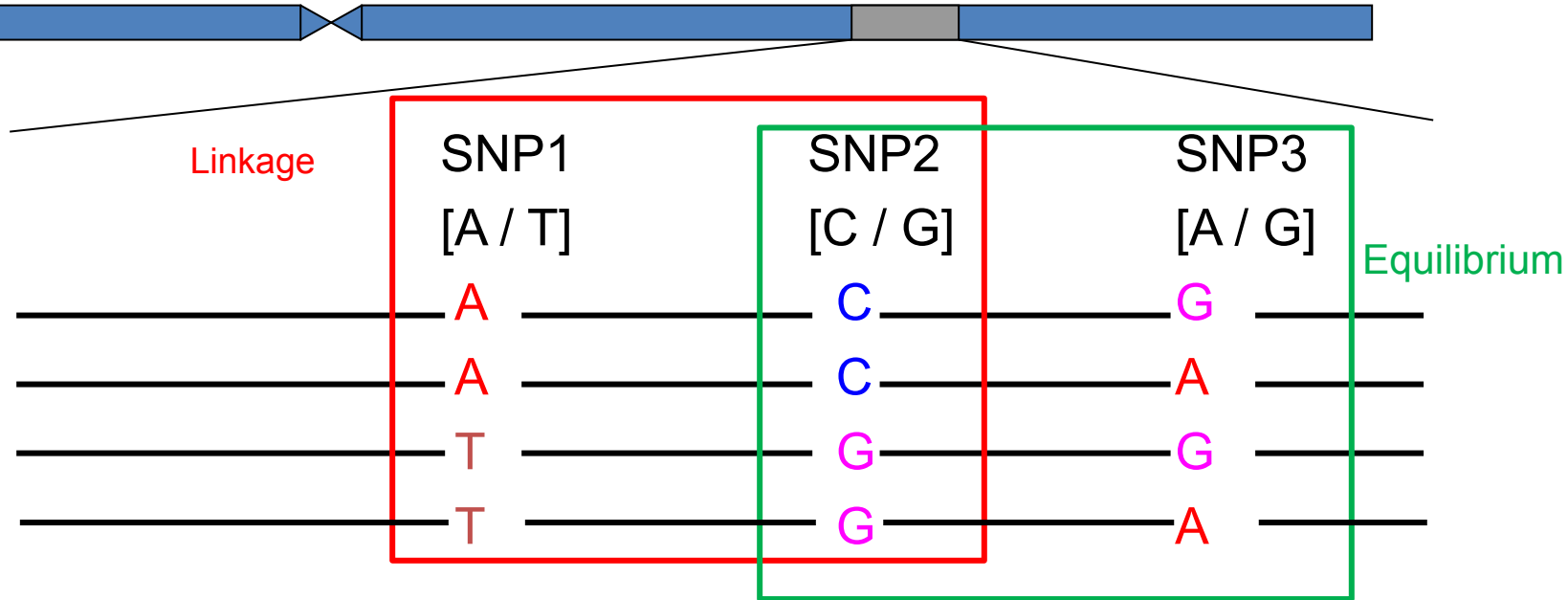
- Linkage disequilibrium
- Linear regression



Linkage disequilibrium recap

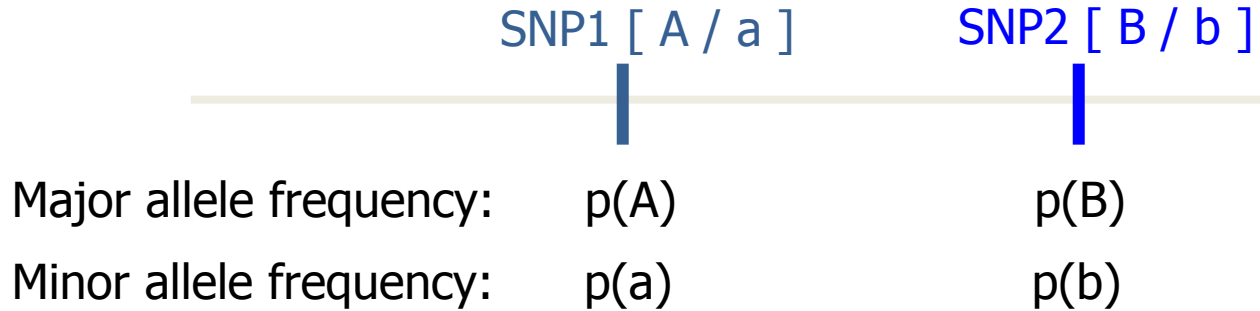


Linkage disequilibrium (**LD**)



Haplotype: specific combination of alleles that appear in the same chromosome or segment (*in-cis*) and are inherited together from a single parental.

Linkage disequilibrium (**LD**)



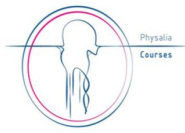
Marker segregation:

Haplotype frequency $p(ab) = p(a) \times p(b)$ Linkage equilibrium

Haplotype frequency $p(ab) \neq p(a) \times p(b)$ Linkage disequilibrium



Linkage disequilibrium (**LD**)



		SNP2 Allele		
		B	b	
SNP1 Allele	A	$p(A)p(B)$ AB	$p(A)p(b)$ Ab	$p(A)$
	a	$p(a)p(B)$ aB	$p(a)p(b)$ ab	$p(a)$
		$p(B)$	$p(b)$	



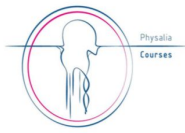
Example:

$$p(\mathbf{ab}) = p(\mathbf{a})p(\mathbf{b}) \quad \text{Expected Haplotype Frequencies}$$

$$p(\mathbf{A})p(\mathbf{B}) + p(\mathbf{a})p(\mathbf{B}) = p(\mathbf{B})\{p(\mathbf{A}) + p(\mathbf{a})\} = p(\mathbf{B})$$



Linkage disequilibrium (**LD**)



		SNP2 Allele			
		B	b		
SNP1 Allele	A	$p(A)p(B)+D$	$p(A)p(b)-D$	$p(A)$	
	a	$p(a)p(B)-D$	$p(a)p(b)+D$	$p(a)$	
		$p(B)$	$p(b)$		

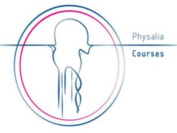
Example:

$$p(ab) \neq p(a)p(b) \rightarrow p(ab) = p(a)p(b) + D$$

**Non-Expected Haplotype
Frequencies (D is LD degree)**

$$\{p(A)p(B)+D\} + \{p(a)p(B)-D\} = p(B)\{p(A)+p(a)\} = p(B)$$

Linkage disequilibrium (LD)



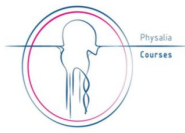
Question: What markers are in LD?

- **A** frequency=0.6; **B** frequency=0.3; haplotype **ab** frequency =0.40
- **a** frequency=0.4; **C** frequency=0.3; haplotype **AC** frequency=0.18
- haplotype **bC** frequency=0.49



Linkage disequilibrium (**LD**)

Most common measurements



$$-1 \longleftarrow D'_{\text{(Lewontin, 1964)}} \longrightarrow 1$$

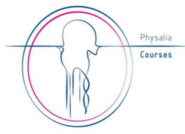
Recombination

$$0 \longleftarrow r^2 \longrightarrow 1$$

Correlation



Linkage disequilibrium (**LD**)



D' (Lewontin, 1964)

$$D' = D / D_{\max}$$

$$D_{\max} = \min (p(A)p(B), p(a)p(b)) \quad D < 0$$

$$D_{\max} = \min (p(A)p(b), p(a)p(B)) \quad D > 0$$

D' sign depends on the reference allele

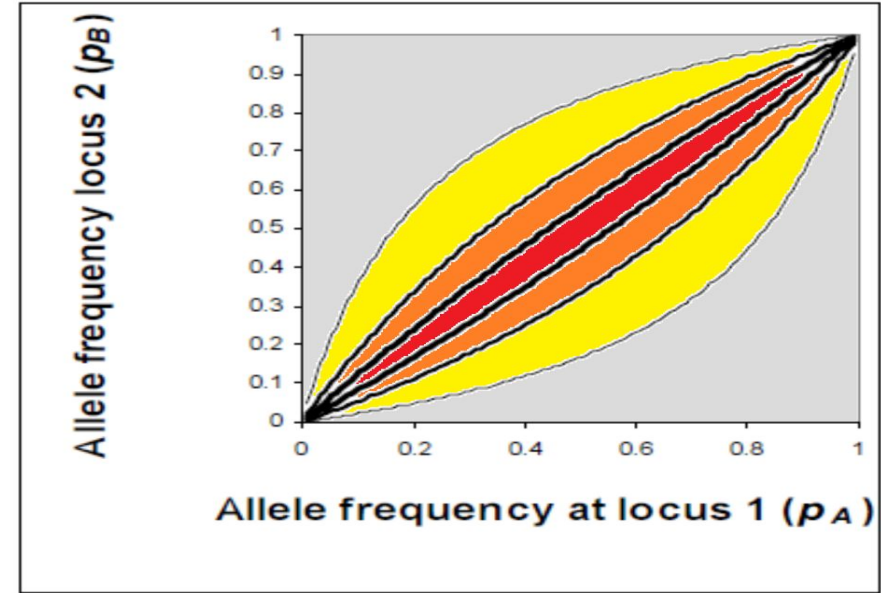


Linkage disequilibrium (**LD**)

r^2 (Hill & Robertson, 1968)

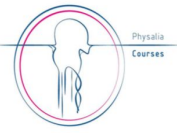
$$r^2 = \frac{D^2}{p(a)p(b)p(A)p(B)}$$

- Between 0 - 1
- Depends on allele frequency
- Most common metric



$r^2 > 0.80$
 $r^2 > 0.50$
 $r^2 > 0.20$
 $r^2 < 0.20$

Linkage disequilibrium (**LD**)



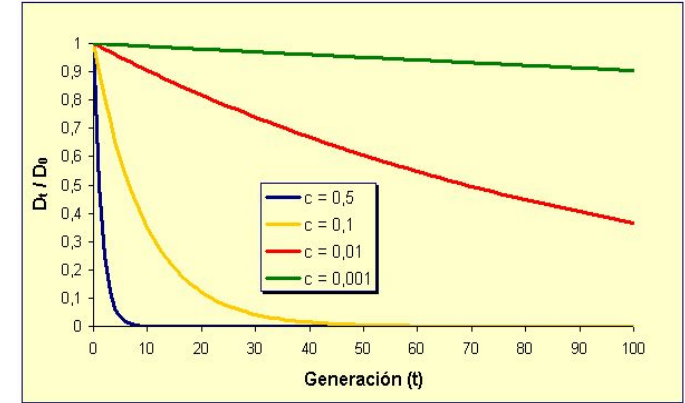
- Non-random allele segregation between 2 or more loci
- Loci on the same chromosome with LD have larger probability of being transmitted together (less likely recombination)
- Physically close loci tend to be in higher LD, as recombination rates are low



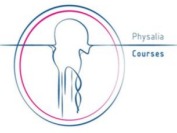
Linkage disequilibrium (**LD**)

Possible causes of LD

- Close location
- Low recombination rates
- Recent populations with low founder size
- Selective breeding (e.g. Holstein Long range LD)
- New mutations
- Inbreeding
- Population bottlenecks
- Population stratification
- Asexual reproduction



Linkage disequilibrium (**LD**)

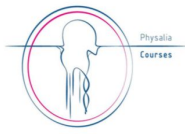


Applications

- We can infer the genotype of one locus given we know the genotype of another locus with LD (Imputation)
- We can predict the effect of a gene, given we know a marker genotype in LD (GWAS – GWP)
- Caution
 - LD may disappear in future generations due to new recombinations. Consider this cautiously when applying results to further apart generations or low LD markers
 - Differentiate between statistical linkage and physical linkage



Linkage disequilibrium (**LD**)



Why do we care about LD?

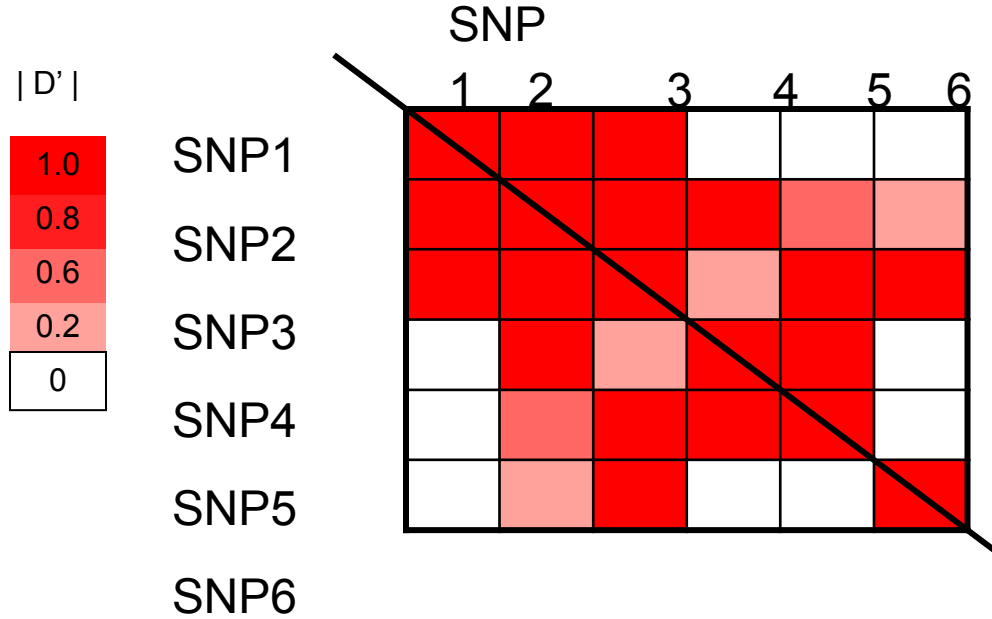
The power of an indirect association study depends on 4 parameters:

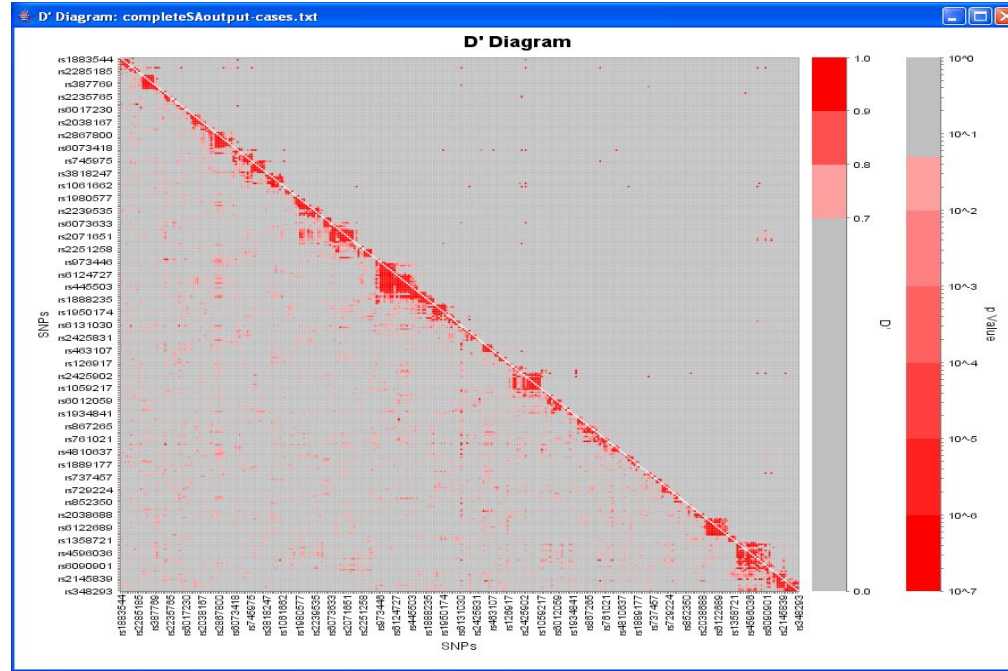
- Disease-allele effect size
- Disease-allele frequency
- Marker-allele frequency
- Extent of LD between marker and disease locus



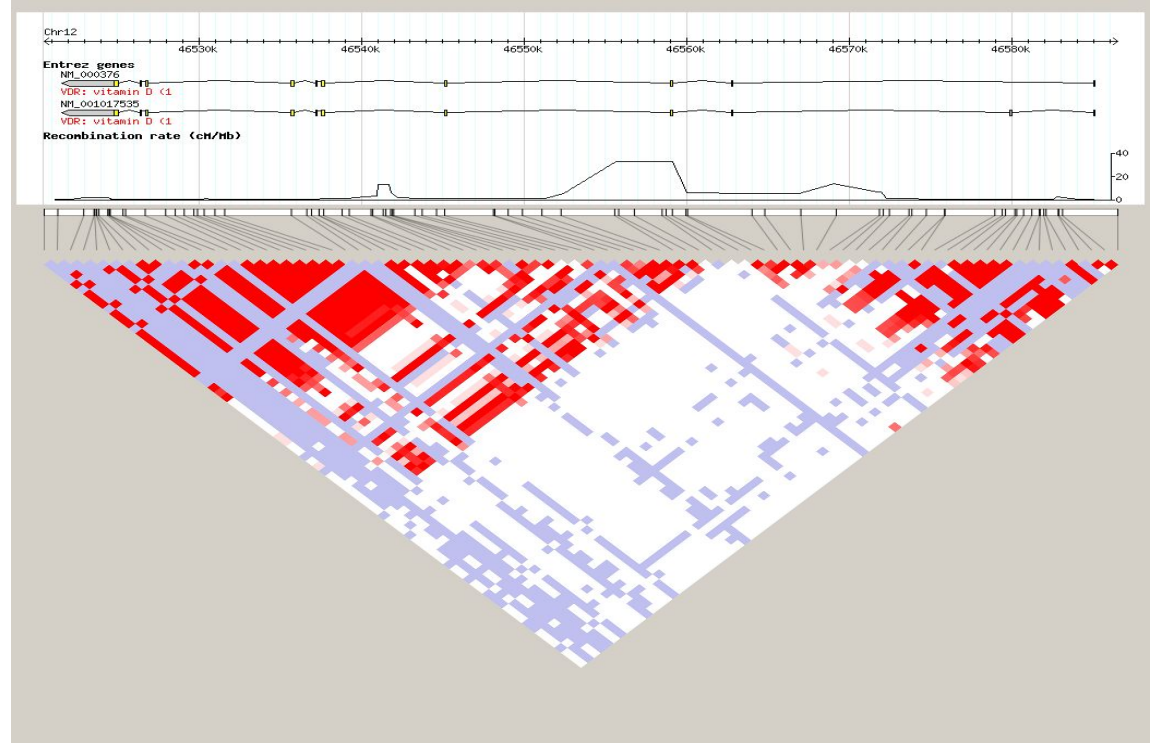
Visualizing LD







Haploview: TCN2 (r^2)



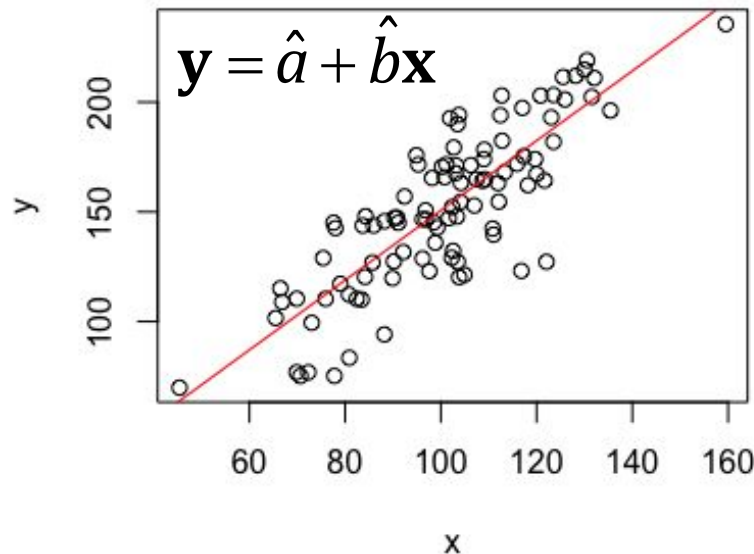
Linear regression recap



Linear regression

$$\mathbf{y} = a + b\mathbf{x} + \mathbf{e}$$

- Estimate the line that minimizes the MSE $\text{Sum}(y - \hat{y})^2$.



$$\hat{b} = \text{cov}(\mathbf{x}, \mathbf{y}) / \text{Var}(\mathbf{x})$$

$$\hat{a} = \bar{y} - \hat{b}\bar{x}$$

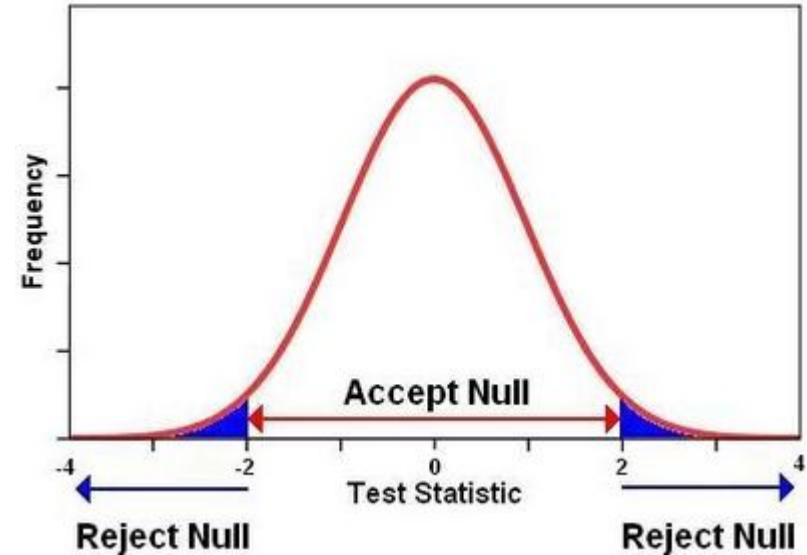
Is the b estimate statistically significant? → Real effect (association)

Linear regression – Hypothesis test

$$H_0 : \beta_1 = 0$$

$$H_1 : \beta_1 \neq 0$$

- Continuous variable → t-Student
- Binary variable → Chi-squared, log-ratio test



Significance level α (e.g. 0.05, 0.01,...)

Linear regression – Hypothesis test

$$H_0 : \beta_1 = 0$$

$$H_1 : \beta_1 \neq 0$$

	One-tail (left)	Two-tails	One-tail (right)
Contrast	$H_0 : \beta_1 = b_1$ $H_1 : \beta_1 < b_1$	$H_0 : \beta_1 = b_1$ $H_1 : \beta_1 \neq b_1$	$H_0 : \beta_1 = b_1$ $H_1 : \beta_1 > b_1$
Statistic (t-Student)	$t = \frac{\hat{\beta}_1 - b_1}{\sqrt{\hat{s}_R^2 / SS_{xx}}} \text{ con } \hat{s}_R^2 = \frac{SS_{yy} - \hat{\beta}_1 SS_{xy}}{n-2}$		
Reject Region	$t < t_{\alpha, n-2}$	$ t > t_{1-\alpha/2, n-2}$	$t > t_{1-\alpha, n-2}$

$$SS_{xy} = \sum_{i=1}^n (x_i - \bar{x})(y_i - \bar{y}) = \sum_{i=1}^n x_i y_i - n\bar{x}\bar{y},$$

$$SS_{xx} = \sum_{i=1}^n (x_i - \bar{x})^2 = \sum_{i=1}^n x_i^2 - n\bar{x}^2 = n\sigma_x^2$$

$$\hat{s}_R^2 = \frac{SSE}{n-2} = \frac{\sum_{i=1}^n e_i^2}{n-2} = \frac{SS_{yy} - \hat{\beta}_1 SS_{xy}}{n-2} \quad SSE = \sum_{i=1}^n e_i^2 = \sum_{i=1}^n y_i - (\beta_0 + \beta_1 x_i)^2$$

$$t = \frac{\hat{\beta}_1 - b_1}{\sqrt{\hat{s}_R^2 / SS_{xx}}}$$

Does my *t*-statistic value belong to the null hypothesis distribution



Linear regression – Hypothesis test

$$H_0 : \beta_1 = 0$$

$$H_1 : \beta_1 \neq 0$$

$$t = \frac{\hat{\beta}_1 - b_1}{\sqrt{\hat{s}_R^2 / SS_{xx}}}$$

Does my *t*-statistic value belong to the null hypothesis distribution



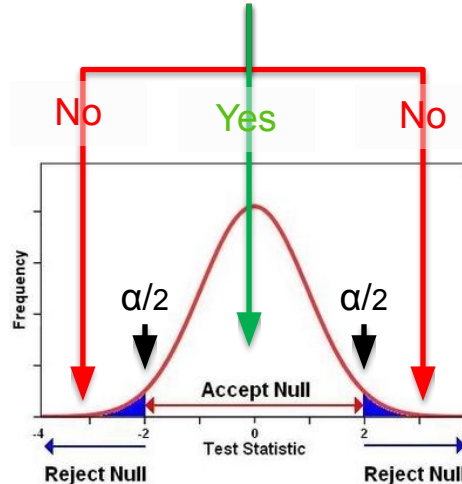
Calculate the *p-value*: probability of obtaining a value of the statistic at least as extreme as the one obtained in the experiment, if we repeat the experiment an infinite number of times

i.e. two-tails test

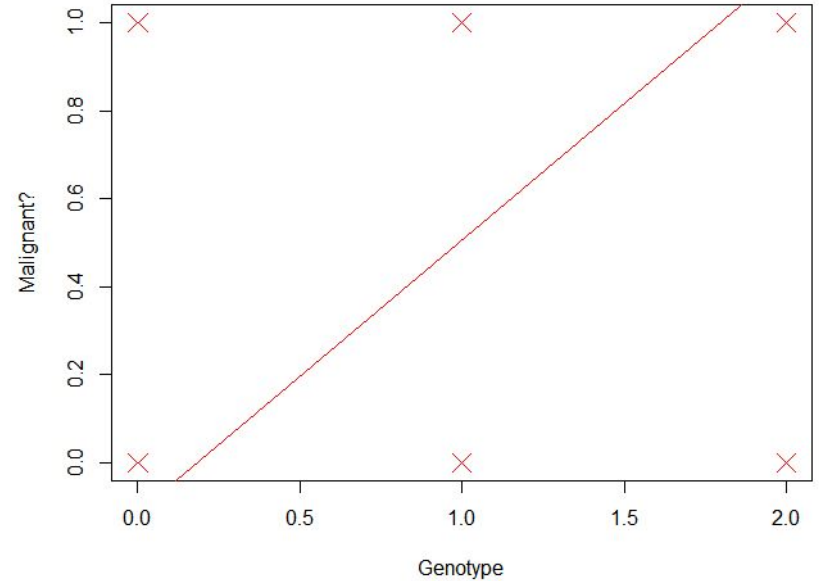
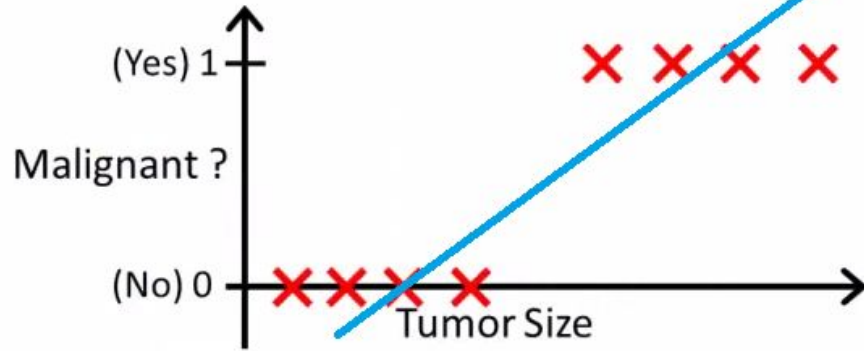
$$2 \cdot (1 - p(z \leq t))$$

If *p-value* < $\alpha \rightarrow$ **reject** null-hypothesis

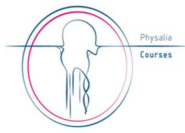
area under curve	0.005	0.010	0.025	0.050	0.100	0.250	0.500	1.000	2.000	5.000	10.000	20.000	50.000	100.000
area under curve	0.005	0.010	0.025	0.050	0.100	0.250	0.500	1.000	2.000	5.000	10.000	20.000	50.000	100.000
1	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
2	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
3	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
4	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
5	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
6	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
7	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
8	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
9	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
10	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
11	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
12	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
13	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
14	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
15	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
16	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
17	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
18	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
19	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
20	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
21	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
22	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
23	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
24	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
25	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
26	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
27	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
28	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
29	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
30	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
31	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
32	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
33	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
34	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
35	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
36	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
37	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
38	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
39	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
40	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
41	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
42	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
43	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
44	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
45	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
46	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
47	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
48	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
49	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000
50	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000	0.000



Logistic regression



Logistic regression $y = a + bx + e$



- Uses the sigmoid function

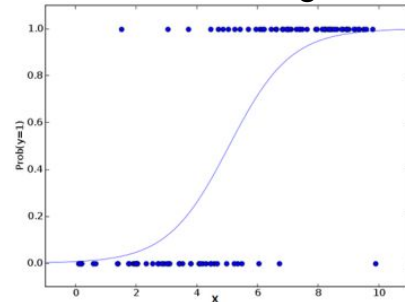
$$\log_{base} \frac{P(Y=1)}{1 - P(Y=1)} = a_0 + a_1 x$$

$$\frac{P(Y=1)}{1 - P(Y=1)} = base^{(a_0 + b_1 x)} \Rightarrow P(Y=1) = base^{(a_0 + b_1 x)} - P(Y=1) base^{(a_0 + b_1 x)} \Rightarrow$$

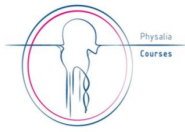
$$(1 + base^{(a_0 + b_1 x)}) P(Y=1) = base^{(a_0 + b_1 x)} \Rightarrow$$

$$P(Y=1) = \frac{base^{(a_0 + b_1 x)}}{1 + base^{(a_0 + b_1 x)}} = \frac{1}{1 + base^{-(a_0 + b_1 x)}}$$

Sigmoid function



Logistic regression $y = a + bx + e$



- Estimation of coefficients a and b

The intercept a is the log of the odds for the control group (AA)

$$a = \log \frac{AA_1 / AA_{\text{totales}}}{AA_0 / AA_{\text{totales}}} = \log \frac{AA_1}{AA_0}$$

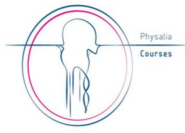
	0(control)	1(case)
AA	AA_0	AA_1
Aa	Aa_0	Aa_1

The regression coefficient b is the log of the odds ratio between the Aa group and the AA group:

$$b = \log \frac{\frac{Aa_1 / Aa_0}{AA_1 / AA_0}}{\frac{Aa_0 / Aa_1}{AA_0 / AA_1}} = \frac{Aa_1 / AA_0}{Aa_0 / AA_1}$$

Logistic regression

$$y = a + bx + e$$



- How to interpret the Odd Ratio
 - It is a measurement of the degree of association between 2 variables, and is estimated with respect to a given base
 - Ranges between 0 and ∞
 - $OR=1 \rightarrow$ lack of association; $OR<1 \rightarrow$ negative association; $OR>1 \rightarrow$ positive association
 - If $OR<1$, estimate the inverse relationship is preferred for easiness of interpretation

	0(control)	1(case)
AA	AA_0	AA_1
Aa	Aa_0	Aa_1

Logistic regression

$$y = a + bx + e$$



- Two examples

- Colorectal cancer and meat consumption.

USA male (30-55yr) and male (40-75yr)

High vs low levels of processed meat intake

Population prevalence: 1%

OR=1.22


Prevalence in the high intake levels of meat ?

Meta-Analysis > Int J Cancer. 2006 Dec 1;119(11):2657-64. doi: 10.1002/ijc.22170.

Meat consumption and risk of colorectal cancer: a meta-analysis of prospective studies

Susanna C Larsson¹, Alija Wolk

Affiliations + expand

PMID: 16991129 DOI: 10.1002/ijc.22170 

Free article

Abstract

Accumulating epidemiologic evidence indicates that high consumption of red meat and of processed meat may increase the risk of colorectal cancer. We quantitatively assessed the association between red meat and processed meat consumption and the risk of colorectal cancer in a meta-analysis of prospective studies published through March 2006. Random-effects models were used to pool study results and to assess dose-response relationships. We identified 15 prospective studies on red meat (involving 7,367 cases) and 14 prospective studies on processed meat consumption (7,903 cases). The summary relative risks (RRs) of colorectal cancer for the highest vs. the lowest intake categories were 1.28 (95% confidence interval (CI) = 1.15-1.42) for red meat and 1.20 (95% CI = 1.11-1.31) for processed meat. The estimated summary RRs were 1.28 (95% CI = 1.18-1.39) for an increase of 120 g/day of red meat and 1.09 (95% CI = 1.05-1.13) for an increase of 30 g/day of processed meat. Consumption of red meat and processed meat was positively associated with risk of both colon and rectal cancer, although the association with red meat appeared to be stronger for rectal cancer. In 3 studies that reported results for subsites in the colon, high consumption of processed meat was associated with an increased risk of distal colon cancer but not of proximal colon cancer. The results of this meta-analysis of prospective studies support the hypothesis that high consumption of red meat and of processed meat is associated with an increased risk of colorectal cancer.

Logistic regression

$$y = a + bx + e$$



- Two examples

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Prevalence in the high intake levels of meat ?


answer: 1.22%

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Logistic regression

$$y = a + bx + e$$



- Two examples

- COVID-19 and Neardenthal gene (Chr3).

UK Biobank (679531 controls), severe COVID (1128 cases)

Table 1 | Lead variants from independent genome-wide significant regions

SNP	Chr.: pos.	Risk	Alt.	RAF _{gsc}	RAF _{ukb}	OR	CI	P _{gsc:ukb}	P _{gsc:gs}	P _{gsc:100k}	Locus
rs73064425	3: 45,901,089	T	C	0.15	0.07	2.1	1.88–2.45	4.8×10^{-30}	2.9×10^{-37}	3.6×10^{-32}	LZTFL1
rs9380192	6: 29,798,794	A	G	0.74	0.69	1.3	1.18–1.43	3.2×10^{-8}	0.00091	1.8×10^{-8}	HLA-G
rs143334143	6: 31,121,426	A	G	0.12	0.07	1.8	1.61–2.13	8.8×10^{-18}	2.6×10^{-24}	5.8×10^{-18}	CCHCR1
rs3131294	6: 32,180,146	G	A	0.9	0.86	1.5	1.28–1.66	2.8×10^{-8}	1.3×10^{-10}	2.3×10^{-8}	NOTCH4
rs10735079	12: 113,380,008	A	G	0.68	0.63	1.3	1.18–1.42	1.6×10^{-8}	2.8×10^{-9}	4.7×10^{-8}	OAS1–OAS3
rs2109069	19: 4,719,443	A	G	0.38	0.32	1.4	1.25–1.48	4×10^{-12}	4.5×10^{-7}	2.4×10^{-8}	DPP9
rs74956615	19: 10,427,721	A	T	0.079	0.05	1.6	1.35–1.87	2.3×10^{-8}	2.2×10^{-13}	3.9×10^{-8}	TYK2
rs2236757	21: 34,624,917	A	G	0.34	0.28	1.3	1.17–1.41	5×10^{-8}	8.9×10^{-5}	8.3×10^{-7}	IFNAR2

As this is a meta-analysis of all available data, external replication cannot be attempted, so SNPs are included if they meet a more stringent P-value threshold of $P < 10^{-4}$. SNP, the strongest SNP in the locus. Chr.: pos., chromosome and position of the top SNP (build 37); Risk, risk allele; Alt., alternative allele; RAF, risk allele frequency; OR, effect size (odds ratio) of the risk allele in the GenOMICC European analysis; CI, 95% confidence interval for the odds ratio in the GenOMICC European cohort; P, P value; Locus, gene nearest to the top SNP. Subscript identifiers indicate the cohorts used for cases (gsc, GenOMICC European cohort) and controls (ukb, UK Biobank; gs, Generation Scotland; 100k, 100,000 Genomes Project).

Population prevalence: 0.16%

OR=2.1

Prevalence in the Risk Allele group ?

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Article | Published: 11 December 2020

Genetic mechanisms of critical illness in COVID-19

Erola Pairo-Castineira, Sara Clohisey, [...], Kenneth Baillie

Nature 591, 92–98 (2021) | Cite this article

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Population prevalence: 0.16%

OR=2.1

Prevalence in the Risk Allele group ?

answer: 0.34%

(needs to be considered carefully, because control group did not report COVID19; we don't know what would have been the progress of the disease)

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Article | Published: 11 December 2020

Genetic mechanisms of critical illness in COVID-19

Erola Pairo-Castineira, Sara Clishsey, [...], Kenneth Baillie

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Now you!

Compute the t-student statistic given some data, and accept or reject the null-hypothesis

1. `Basis_of_linear_regression.R`
2. `Basis_of_logistic_regression.Rmd`
3. Homework: Analyze data in `Exercise1.exampleData.xlsx`,
 - a) Determine SNPs associated to the phenotype, and calculate the Odds-Ratio
 - b) If you got 2 significant SNPs, that's wrong. Figure out why.