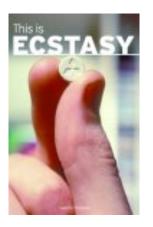
Researcher and statistician error



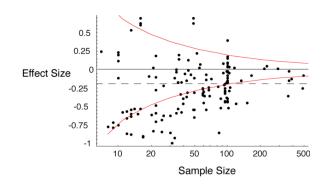
~8% of biomedical papers have substantial statistical flaws

1

Publication bias

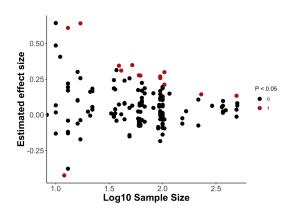
Papers are more likely to be published if P<0.05

This causes a bias in the science reported in the literature.

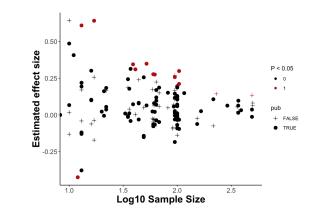


2

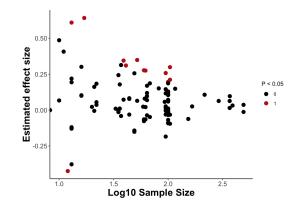
Publication bias with true effect size 0.05



Publication bias with true effect size 0.05



Publication bias with true effect size 0.05



Computer intensive methods

Chapter 19

Computer-intensive methods

Hypothesis testing:SimulationRandomization

Confidence intervals •Bootstrap

Simulation

Simulates the sampling process on a computer many times: generates the null distribution from estimates done on the simulated data

Computer assumes the null hypothesis is true

Example: Social spider sex ratios

Social spiders live in groups



Example: Social spider sex ratios

Groups are mostly females

Hypothesis: Groups have just enough males to allow reproduction

Test: Whether distribution of number of males is as predicted by chance

Problem: Groups are of many different sizes

Binomial distribution therefore doesn't apply

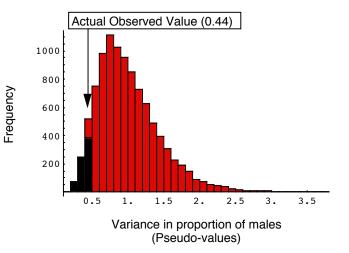
Simulation:

For each group, the number of spiders is known. The overall proportion of males, $p_{\rm m}$, is known.

For each group, the computer draws the real number of spiders, and each has $p_{\rm m}$ probability of being male.

This is done for all groups, and the variance in proportion of males is calculated.

This is repeated a large number of times.



The observed value (0.44), or something more extreme, is observed in only 4.9% of the simulations. Therefore P = 0.049.

Bootstrap

Method for estimation (and confidence intervals)

Often used for hypothesis testing too

"Picking yourself up by your own bootstraps"

Bootstrap

For each group, randomly pick *with replacement* an equal number of data points, from the data of that group.

With this bootstrap dataset, calculate the estimate – the bootstrap replicate estimate.

Revisting the Sage crickets



Sage cricket males sometimes offer their hind-wings to females to eat during mating.

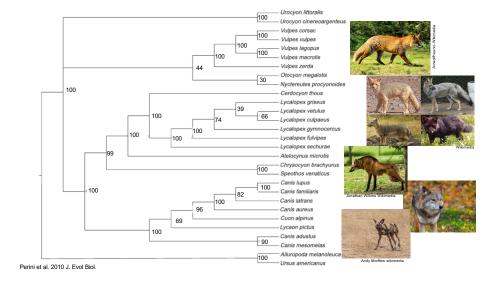
Do females who eat hindwings wait longer to remate? Real data: $\overline{Y}_1 - \overline{Y}_2 = -1.41$

-	_		
Male wingless	Male winged		
0	1.4		
0.7	1.6		
0.7	1.9		
1.4	2.3		
1.6	2.6		
1.8	2.8		
1.9	2.8		
1.9	2.8		
1.9	3.1		
2.2	3.8		
2.1	3.9		
2.1	4.5		
	4.7		

Bootstrap data:	\overline{Y}_1 –	\overline{Y}_2	= -1.78
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Male		
winged		
4.7		
4.7		
3.1		
3.9		
2.8		
2.8		
4.7		
4.7		
2.8		
4.5		
3.1		
1.4		
1.4		

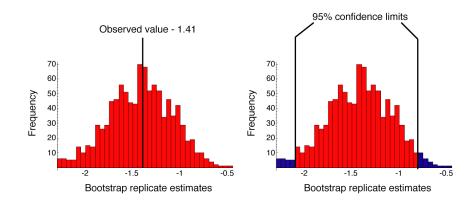
Bootstraps are often used in evolutionary trees





 $L(hypothesis A \mid data) = P[data \mid hypothesis A]$

Likelihood considers many possible hypotheses, not just one



Likelihood

Chapter 20

Law of likelihood

A particular data set supports one hypothesis better than another if the likelihood of that hypothesis is higher than the likelihood of the other hypothesis.

Therefore we try to find the hypothesis with the maximum likelihood.

"Simple" example

Using likelihood to estimate a proportion

Data: 3 out of 8 individuals are male.

Question: What is the maximum likelihood estimate of the proportion of males?

All estimates we have learned so far are also maximum likelihood estimates.

Likelihood

$$L(p = x) = P[3 \text{ males out of } 8 | p = x]$$

where *x* is a hypothesized value of the proportion of males.

e.g., L(p=0.5) is the likelihood of the hypothesis that the proportion of males is 0.5.

For this example only...

The probability of getting 3 males out of 8 independent trials is given by the binomial distribution.

$$L(p = x) = \Pr[data | p = x]$$

=
$$\Pr[3out of 8 | p = x]$$

=
$$\binom{8}{3}x^{3}(1-x)^{8-3}$$

How to find maximum likelihood hypothesis

1. Calculus

or

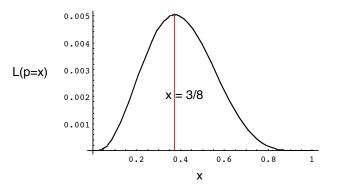
2. Computer calculations

By calculus...

Maximum value of L(p=x) is found when x = 3/8.

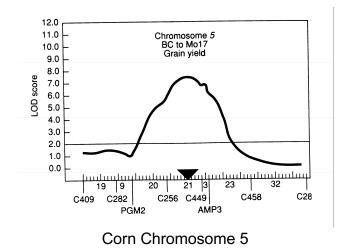
Note that this is the same value we would have gotten by methods we already learned.

By computer calculation...



Input likelihood formula to computer, plot the value of L for each value of x, and find the largest L.

Finding genes for corn yield:



Hypothesis testing by likelihood

Compares the likelihood of maximum likelihood estimate to a null hypothesis

Log-likelihood ratio =

ln Likelihood [Maximum likelihood hypothesis] Likelihood [Null hypothesis]

Test statistic

 $\chi^2 = 2 \log likelihood ratio$

With *df* equal to the number of variables fixed to make null hypothesis

Example:3 males out of 8 individuals

H₀: 50% are male

Maximum likelihood estimate

$$\hat{p} = \frac{3}{8}$$

$$L[p = 3/8] = {\binom{8}{3}} (3/8)^3 (1 - 3/8)^5 = 0.2816$$

Likelihood of null hypothesis

$$L[p=0.5] = {\binom{8}{3}} (0.5)^3 (1-0.5)^5 = 0.21875$$

Log likelihood ratio

$$\ln\left[\frac{L[p=3/8]}{L[p=0.5]}\right] = \ln\left[\frac{0.2816}{0.21875}\right] = 0.2526$$

$$\chi^2 = 2(0.2526) = 0.5051$$

We fixed one variable in the null hypothesis (p), So the test has df = 1.

 $\chi^2_{0.05,1} = 3.84$, so we do not reject H₀.