Introduction to statistics

Computational biology week - London, April 2022

Jesús Urtasun Elizari, PhD





Imperial College London

Outline

- Probability in a nutshell
 - Discrete and continuous probability
 - Mean and variance of a distribution
 - Exercises in R
- 2 Linear models
 - What is a linear model
 - Fitting a linear model
 - Prediction vs inference
 - Exercises in R
- Hypothesis testing
 - Probability distributions
 - Statistical tests
 - Real genetics problem
 - Exercises in R

Chapter I Probability in a nutshell

Discrete and continuous probability

- Discrete output → probability well defined, compute by counting
- Continuous output → need for probability distributions
- Definition of probability in both scenarios
- Computing probabilities in both scenarios

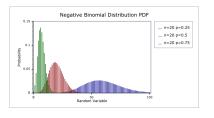
Discrete probability

Compute probability of getting 4 heads in a row

Discrete probability

Compute probability of getting a specific genotype

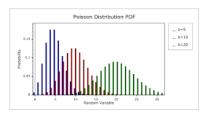
Binomial distribution



Binomial distribution: probability of getting k events in n trials

$$B(x=k) = \binom{n}{x} p^k (1-p)^{n-k}$$

Poisson distribution



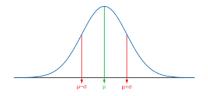
 Poisson distribution: probability of counting k events in a given interval

$$P(x=k) = \frac{\lambda^k e^{-\lambda}}{k!}$$

Continuous probability

Compute probability of measuring a particular height Used when comparing means

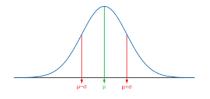
Gaussian distribution



 Gaussian distribution: continuous distribution for a real-valued random variable

$$f(x) = \frac{1}{\sigma\sqrt{2\pi}}e^{-\frac{1}{2}\left(\frac{x-\mu}{\sigma}\right)^2}$$

Mean and variance



- Momenta of a distribution
- Mean, variance, skewness, kurtosis

Standard Deviation vs Standard Error

Performing multiple sets of measurements and compare means

- Standard deviation (σ) quantifies variation over one set of measurements (second momentum of the probability distribution)
- Standard error (SE) quantifies variation of means from multiple sets!

$$SE = \frac{\sigma}{\sqrt{N}}$$

 Source of confusion: SE can be estimated from single set of measurements, even though it describes variation of means

Exercises in R

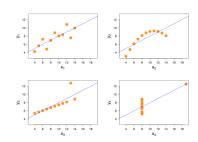
 $https://Imsbioinformatics.github.io/LMS_StatisticsInR/course/CBW_StatisticsInR_course$

Chapter II Linear models

What is a linear model

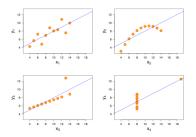
Find a function that describes a set of observations

$$y(x) = b_0 \cdot x + b_1$$



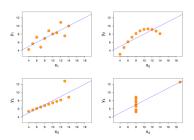
Fitting a linear model

- Get the overall mean
- 2 Compute sum of residuals (SS) over the mean
- \bigcirc Find tine with the smallest SS \rightarrow fit
- Evaluate fit correlation coefficient R²



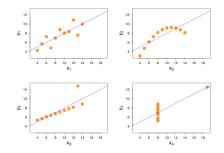
Generalized linear models

- Fit count data (non-linearly distributed)
- 2 Poisson regression (assume variance mean
- Negative binomial



Advanced topics

- ANOVA
- **2** (...)



Exercises in R

 $https://Imsbioinformatics.github.io/LMS_StatisticsInR/course/CBW_StatisticsInR_course$

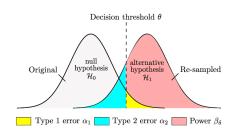
Chapter III
Hypothesis testing

Introduction

- Null hypothesis and alternative hypothesis
- Statistic tests and p-values
- χ^2 -test, *t*-test, Wald test
- Exercises in R

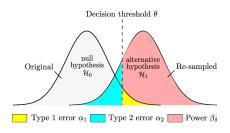
Compare different models

Null and alternative hypothesis



Statistic test

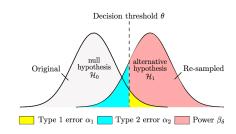
- Quantify the significance of an observation
- Certainty when accepting / rejecting a hypothesis



23 / 26

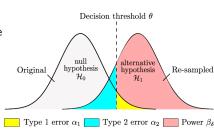
p-values

p-values



Power calculations

- Do not just add samples until getting a good p-value → increases chance of reporting a false positive
- Power": probability that a test will correctly give a small p-value
- 4 factors affect power (effect size, variation in data, sample size, test used)



Exercises in R

 $https://Imsbioinformatics.github.io/LMS_StatisticsInR/course/CBW_StatisticsInR_course$