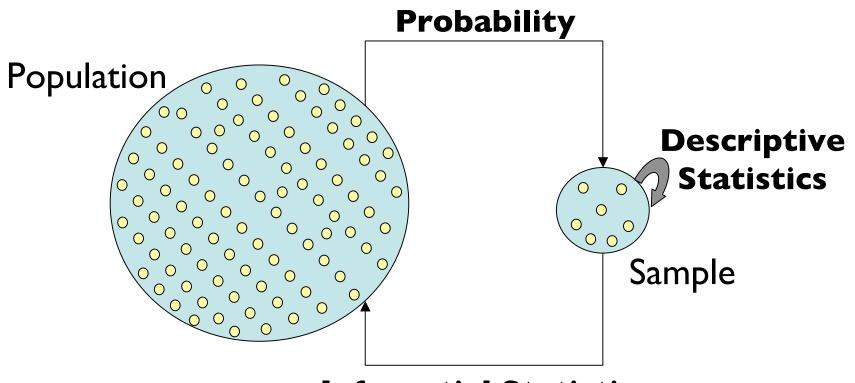
Lecture 5: Estimation

Goals

- Basic concepts of estimation
- Statistical approaches for estimating parameters

- Parametric interval estimation
- Nonparametric interval estimation (bootstrap)

"Central Dogma" of Statistics



Inferential Statistics

Estimation

- Estimator: Statistic whose calculated value is used to estimate a population parameter, $\boldsymbol{\theta}$
- Estimate: A particular realization of an estimator, $\hat{ heta}$

• Types of Estimators:

- point estimate: single number that can be regarded as the most plausible value of $\boldsymbol{\theta}$
- interval estimate: a range of numbers, called a confidence interval indicating, can be regarded as likely containing the true value of θ

Properties of Good Estimators

- In the **Frequentist** world view parameters are fixed, statistics are rv and vary from sample to sample (i.e., have an associated sampling distribution)
- In theory, there are many potential estimators for a population parameter
- What are characteristics of good estimators?

Statistical Jargon for Good Estimators

Good Estimators Are:

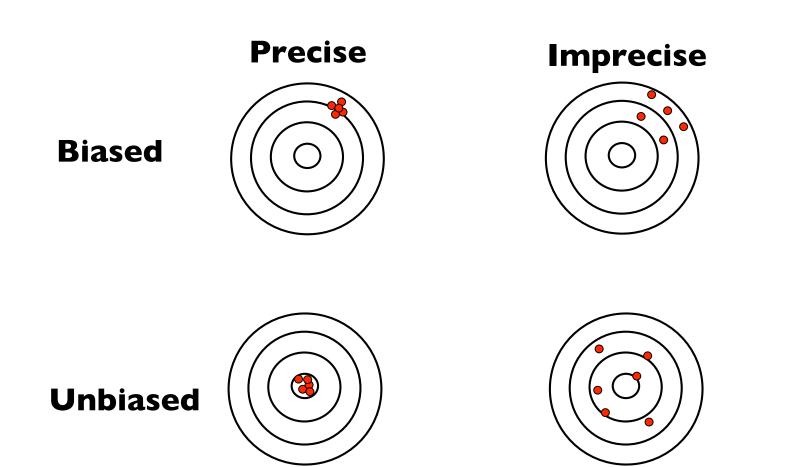
• <u>Consistent</u>: As the sample size increases $\hat{\theta}$ gets closer to θ

$$\lim_{n \to \infty} P(\left|\hat{\theta} - \theta\right| > \varepsilon) = 0$$

• <u>Unbiased</u>: $E[\hat{\theta}] = \theta$

• <u>Precise</u>: Sampling distribution of $\hat{\theta}$ should have a small standard error

Bias Versus Precision



Methods of Point Estimation

I. Method of Moments

2. Maximum Likelihood

3. Bayesian

Methods of Moments

- Advantage: simplest approach for constructing an estimator
- **Disadvantage:** usually are not the "best" estimators possible

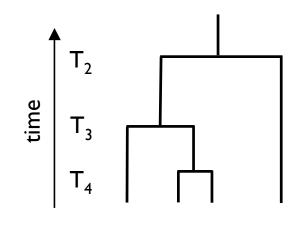
• Principle:

Equate the kth population moment $E[X^k]$ with the kth sample moment $\frac{1}{n}\sum_{k}X_i^k$ and solve for the unknown parameter

• How can I estimate the scaled population mutation rate:

$$\theta = 4N_e\mu$$

• Brief (very brief) expose of coalescent theory:



Coalescent times follow a geometric distribution

$$E[T_i] = \frac{4N}{i(i-1)}$$

$$T_c = \sum_{i=2}^n iT_i$$

$$E[T_c] = \sum_{i=2}^n iE[T_i]$$

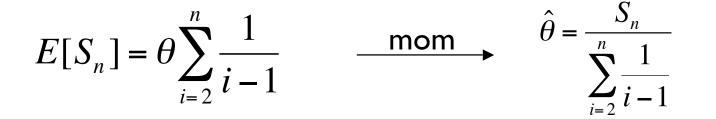
$$E[T_c] = \sum_{i=2}^{n} iE[T_i] = \sum_{i=2}^{n} \frac{4Ni}{i(i-1)}$$

$$E[T_c] = \sum_{i=2}^n iE[T_i] = \sum_{i=2}^n \frac{4Ni}{i(i-1)} = 4N\sum_{i=2}^n \frac{1}{i-1}$$

$$E[T_c] = \sum_{i=2}^n iE[T_i] = \sum_{i=2}^n \frac{4Ni}{i(i-1)} = 4N\sum_{i=2}^n \frac{1}{i-1}$$

 $E[S_n] = \mu E[T_c]$

$$E[S_n] = \mu \bullet 4N \sum_{i=2}^n \frac{1}{i-1}$$



Methods of Point Estimation

I. Method of Moments

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Introduction to Likelihood

 Before an experiment is performed the outcome is unknown.
 Probability allows us to predict **unknown** outcomes based on **known** parameters:

$P(Data | \theta)$

• For example:

$$P(x \mid n, p) = \binom{n}{x} p^x (1-p)^{n-x}$$

Introduction to Likelihood

 After an experiment is performed the outcome is *known*. Now we talk about the *likelihood* that a parameter would generate the observed data:

 $L(\theta \mid Data)$

 $L(\theta \mid Data) = P(Data \mid \theta)$

• For example:

$$L(p \mid n, x) = \binom{n}{x} p^{x} (1-p)^{n-x}$$

• Estimation proceeds by finding the value of θ that makes the observed data most likely

Let's Play T/F

- True or False: The maximum likelihood estimate (mle) of $\,\theta\,$ gives us the probability of $\hat{\theta}\,$
- False why?
- True or False: The mle of θ is the most likely value of $\hat{\theta}$
- False why?
- True or False: Maximum likelihood is cool

Formal Statement of ML

- Let $x_1, x_2, ..., x_n$ be a sequence of n observed variables
- Joint probability:

$$P(\mathbf{x}_{1}, \mathbf{x}_{2}, ..., \mathbf{x}_{n} | \theta) = P(X_{1} = \mathbf{x}_{1})P(X_{2} = \mathbf{x}_{2})...P(X_{n} = \mathbf{x}_{n})$$
$$= \prod_{i=1}^{n} P(X_{i} = x_{i})$$

• Likelihood is then:

$$L(\theta | \mathbf{x}_1, \mathbf{x}_2, ..., \mathbf{x}_n) = \prod_{i=1}^n P(X_i = x_i)$$

Log L(
$$\theta | \mathbf{x}_1, \mathbf{x}_2, ..., \mathbf{x}_n$$
) = $\sum_{i=1}^n \log[P(X_i = x_i)]$

MLE Example

• I want to estimate the recombination fraction between locus A and B from 5 heterozygous (AaBb) parents. I examine 30 gametes for each and observe 4, 3, 5, 6, and 7 recombinant gametes in the five parents. What is the mle of the recombination fraction?

Probability of observing X = r recombinant gametes for a single parent is binomial:

$$P(X = r) = {\binom{\mathsf{n}}{\mathsf{r}}}\theta^r (1 - \theta)^{n-r}$$

MLE Example: Specifying Likelihood

Probability:

$$P(r_{1}, r_{2}, ..., r_{n} | \theta, n) = P(R_{1} = r_{1})P(R_{2} = r_{2})...P(R_{5} = r_{5})$$

$$P(r_{1}, r_{2}, ..., r_{n} | \theta, n) = {\binom{n}{r_{1}}}\theta^{r_{1}}(1-\theta)^{n-r_{1}} \cdot {\binom{n}{r_{2}}}\theta^{r_{1}}(1-\theta)^{n-r_{2}} \cdot ... \cdot {\binom{n}{r_{5}}}\theta^{r_{1}}(1-\theta)^{n-r_{5}}$$

Likelihood:

$$L(\theta \mid \mathbf{r}_1, \mathbf{r}_2, \dots, \mathbf{r}_n, \mathbf{n}) = \prod_{i=1}^{5} {\binom{\mathbf{n}}{\mathbf{r}_i}} \theta^{r_i} (1-\theta)^{n-r_i}$$

$$\log L = \sum_{i=1}^{5} \log\binom{n}{r_i} + r_i \cdot \log \theta + (n - r_i) \log(1 - \theta)$$

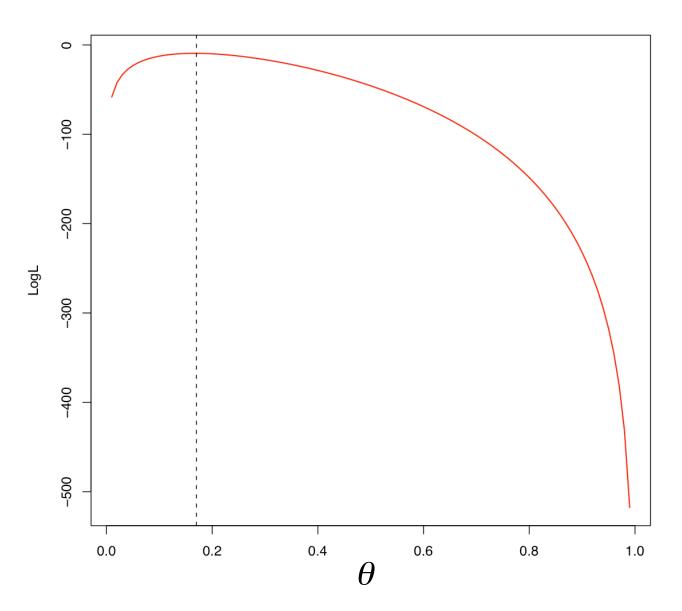
MLE Example: Maximizing the Likelihood

Want to find p such that Log L is maximized

$$Log L = \sum_{i=1}^{5} \log\binom{\mathbf{n}}{\mathbf{r}_{i}} + r_{i} \cdot \log\theta + (n - r_{i})\log(1 - \theta)$$

- How?
 - 1. Graphically
 - 2. Calculus
 - 3. Numerically

MLE Example: Finding the mle of p



Methods of Point Estimation

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World View According to Bayesian's

• The classic philosophy (frequentist) assumes parameters are **fixed** quantities that we want to estimate as precisely as possible

 Bayesian perspective is different: parameters are *random variables* with probabilities assigned to particular values of parameters to reflect the degree of evidence for that value

Revisiting Bayes Theorem

$$P(A \mid B) = \frac{P(B \mid A)P(A)}{P(B)}$$

Discrete
$$P(B) = \sum_{i=1}^{n} P(B \mid A_i) P(A_i)$$

Continuous $P(B) = \int P(B \mid A)P(A)dA$

Bayesian Estimation

• In order to make probability statements about θ given some observed data, D, we make use of Bayes theorem

$$f(\theta \mid D) = \frac{f(\theta)f(D \mid \theta)}{f(D)} = \frac{f(\theta)L(\theta \mid D)}{\int_{\theta} f(\theta)f(D \mid \theta)d\theta}$$

Posterior ∝ likelihood x prior

The prior is the probability of the parameter and represents what was thought before seeing the data.

The likelihood is the probability of the data given the parameter and represents the data now available.

The posterior represents what is thought given both prior information and the data just seen.

Bayesian Estimation: "Simple" Example

- I want to estimate the recombination fraction between locus A and B from 5 heterozygous (AaBb) parents. I examine 30 gametes for each and observe 4, 3, 5, 6, and 7 recombinant gametes in the five parents. What is the mle of the recombination fraction?
- Tedious to show Bayesian analysis. Let's simplify and ask what the recombination fraction is for parent three, who had 5 observed recombinant gametes.

Specifying The Posterior Density

$$f(\theta \mid n = 30, r = 5) = \frac{f(\theta)f(r = 5 \mid \theta, n = 30)}{\int_{0}^{0.5} f(r = 5 \mid \theta, n = 30)f(\theta) \ d\theta}$$

prior = $f(\theta)$ = uniform[0, 0.5] = 0.5

likelihood =
$$P(r = 5 \mid \theta, n = 30) = {\binom{30}{5}} \theta^{r_i} (1 - \theta)^{n - r_i}$$

normalizing
constant =
$$\int_{0}^{0.5} P(r=5 \mid \theta, n=30) f(\theta) d\theta$$

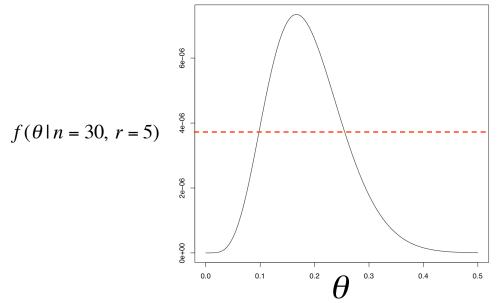
$$= 0.5 \bullet \binom{30}{5} \int_{0}^{0.5} \theta^5 (1-\theta)^{25} d\theta \approx 653 I$$

Specifying The Posterior Density

$$f(\theta \mid n = 30, r = 5) = \frac{f(\theta)f(r = 5 \mid \theta, n = 30)}{\int_{0}^{0.5} f(r = 5 \mid \theta, n = 30) f(\theta) \ d\theta}$$

$$f(\theta \mid n = 30, r = 5) = \frac{0.5 \cdot {\binom{30}{5}} \theta^5 (1 - \theta)^{25}}{6531}$$

Ta **d**a...



Interval Estimation

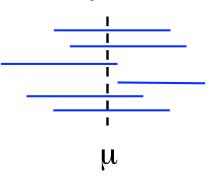
- In addition to point estimates, we also want to understand how much uncertainty is associated with it
- One option is to report the <u>standard error</u>

• Alternatively, we might report a <u>confidence interval</u>

• **Confidence interval:** an interval of plausible values for the parameter being estimated, where degree of plausibility specifided by a "confidence level"

Interpreting a 95% CI

- We calculate a 95% CI for a hypothetical sample mean to be between 20.6 and 35.4. Does this mean there is a 95% probability the true population mean is between 20.6 and 35.4?
- **NO!** Correct interpretation relies on the long-rang frequency interpretation of probability



• Why is this so?