

Statistical Inference Course Notes

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Overview

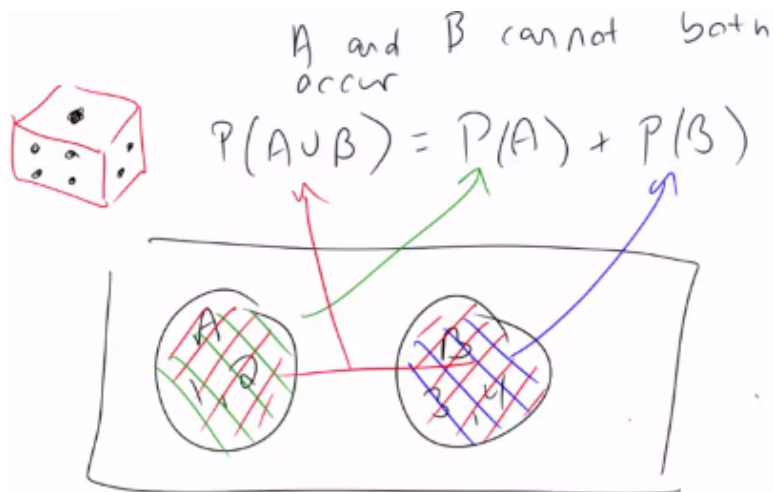
- **Statistical Inference** = generating conclusions about a population from a noisy sample
- Goal = extend beyond data to population
- Statistical Inference = only formal system of inference we have
- many different modes, but **two** broad flavors of inference (inferential paradigms): **Bayesian** vs **Frequentist**
 - **Frequentist** → uses long run proportion of times an event occurs independent identically distributed repetitions
 - * frequentist is what this class is focused on
 - * believes if an experiment is repeated many many times, the resultant percentage of success/something happening defines that population parameter
 - **Bayesian** → probability estimate for a hypothesis is updated as additional evidence is acquired
- **statistic** = number computed from a sample of data
 - statistics are used to infer information about a population
- **random variable** = outcome from an experiment
 - deterministic processes (variance/means) produce additional random variables when applied to random variables, and they have their own distributions

Probability

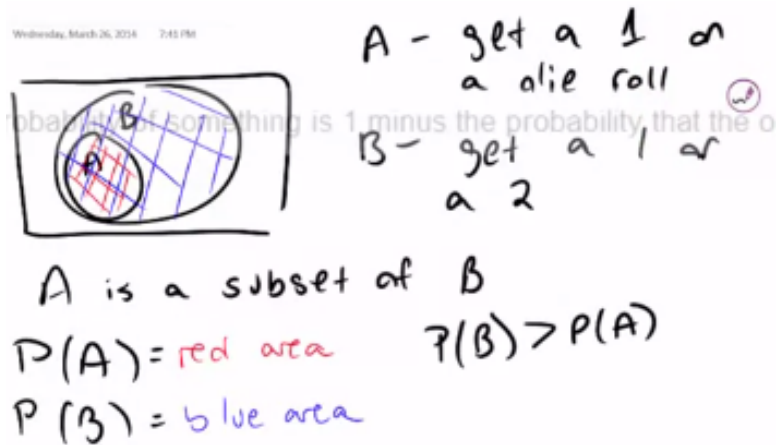
- **Probability** = the study of quantifying the likelihood of particular events occurring
 - given a random experiment, **probability** = population quantity that summarizes the randomness
 - * not in the data at hand, but a conceptual quantity that exist in the population that we want to estimate

General Probability Rules

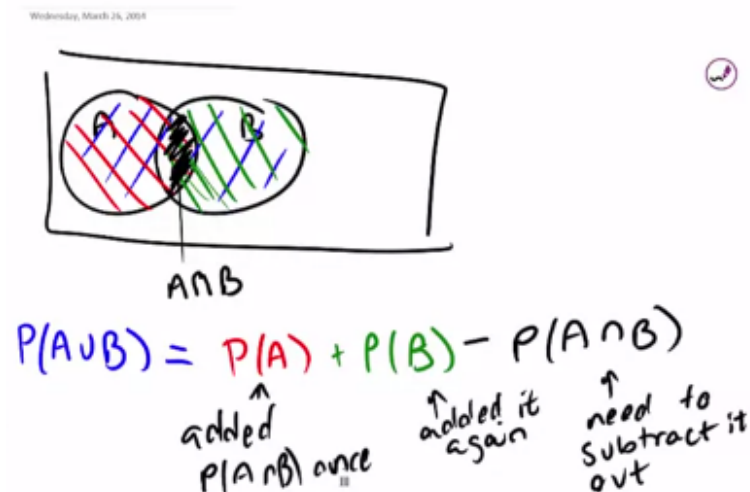
- discovered by Russian mathematician Kolmogorov, also known as “Probability Calculus”
- probability = function of any set of outcomes and assigns it a number between 0 and 1
 - $0 \leq P(E) \leq 1$, where E = event
- probability that nothing occurs = 0 (impossible, have to roll dice to create outcome), that something occurs is 1 (certain)
- probability of outcome or event E, **$P(E)$** = ratio of ways that E could occur to number of all possible outcomes or events
- probability of something = 1 - probability of the opposite occurring
- probability of the **union** of any two sets of outcomes that have nothing in common (mutually exclusive) = sum of respective probabilities



- if A implies occurrence of B, then $P(A) \text{ occurring} < P(B) \text{ occurring}$



- for any two events, probability of at least one occurs = the sum of their probabilities - their intersection (in other words, probabilities can not be added simply if they have non-trivial intersection)



- for independent events A and B, $P(A \cup B) = P(A) \times P(B)$
- for outcomes that can occur with different combination of events and these combinations are mutually exclusive, the $P(E_{total}) = \sum P(E_{part})$

Conditional Probability

- let B = an event so that $P(B) > 0$
- **conditional probability** of an event A, given B is defined as the probability that BOTH A and B occurring divided by the probability of B occurring

$$P(A | B) = \frac{P(A \cap B)}{P(B)}$$

- if A and B are *independent*, then

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

- *example*

– for die roll, $A = \{1\}$, $B = \{1, 3, 5\}$, then

$$P(1 | Odd) = P(A | B) = \frac{P(A \cap B)}{P(B)} = \frac{P(A)}{P(B)} = \frac{1/6}{3/6} = \frac{1}{3}$$

Baye's Rule

- definition

$$P(B | A) = \frac{P(A | B)P(B)}{P(A | B)P(B) + P(A | B^c)P(B^c)}$$

where B^c = corresponding probability of event B, $P(B^c) = 1 - P(B)$

Random Variables

- **random variable** = numeric outcome of experiment
- **discrete** (what you can count/categories) = assign probabilities to every number/value the variable can take
 - coin flip, rolling a die, web traffic in a day
- **continuous** (any number within a continuum) = assign probabilities to the range the variable can take
 - BMI index, intelligence quotients
 - ***Note:** limitations of precision in taking the measurements may imply that the values are discrete, but we in fact consider them continuous*
- `rbinom()`, `rnorm()`, `rgamma()`, `rpois()`, `runif()` = functions to generate random variables from the binomial, normal, Gamma, Poisson, and uniform distributions
- density and mass functions (population quantities, not what occurs in data) for random variables = best starting point to model/think about probabilities for numeric outcome of experiments (variables)
 - use data to estimate properties of population → linking sample to population

Probability Mass Function (PMF)

- evaluates the probability that the **discrete random variable** takes on a specific value
 - measures the chance of a particular outcome happening
 - always ≥ 0 for every possible outcome
 - \sum possible values that the variable can take = 1
- ***Bernoulli distribution example***
 - $X = 0 \rightarrow$ tails, $X = 1 \rightarrow$ heads
 - * X here represents potential outcome
 - $p(X = x) = \left(\frac{1}{2}\right)^x \left(\frac{1}{2}\right)^{1-x}$ for $X = 0, 1$
 - * x here represents a value we can plug into the PMF
 - * general form $\rightarrow p(x) = (\theta)^x (1 - \theta)^{1-x}$
- `dbinom(k, n, p)` = return the probability of getting **k** successes out of **n** trials, given probability of success is **p**

Probability Density Function (PDF)

- evaluates the probability that the **continuous random variable** takes on a specific value
 - always \geq everywhere
 - total area under the must = 1
- **areas under PDFs** correspond to the probabilities for that random variable taking on that range of values (PMF)



- but the probability of the variable taking a specific value = 0 (area of a line is 0)



- **Note:** the above is true because it is modeling random variables as if they have infinite precision, when in reality they do not
- `dnorm()`, `dgamma()`, `dpois()`, `dunif()` = return probability of a certain value from the normal, Gamma, Poisson, and uniform distributions

Cumulative Distribution Function (CDF)

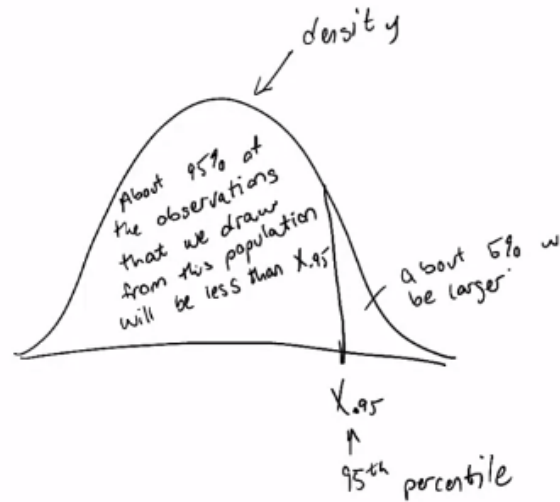
- CDF of a random variable X = probability that the random variable is \leq value x
 - $F(x) = P(X \leq x)$ <- applies when X is discrete/continuous
- PDF = derivative of CDF
 - integrate PDF \rightarrow CDF
 - * `integrate(function, lower=0, upper=1)` \rightarrow can be used to evaluate integrals for a specified range
- `pbinom()`, `pnorm()`, `pgamma()`, `ppois()`, `punif()` = returns the cumulative probabilities from 0 up to a specified value from the binomial, normal, Gamma, Poisson, and uniform distributions

Survival Function

- survival function of a random variable X = probability the random variable $> x$, complement of CDF
 - $S(x) = P(X > x) = 1 - F(x)$, where $F(x)$ = CDF

Quantile

- the α^{th} quantile of a distribution with distribution function F = point x_α
 - $F(x_\alpha) = \alpha$
 - percentile = quantile with α expressed as a percent
 - median = 50th percentile
 - $\alpha\%$ of the possible outcomes lie below it



- `qbeta(quantileInDecimals, 2, 1)` = returns quantiles for beta distribution
 - works for `qnorm()`, `qbinom()`, `qgamma()`, `qpois()`, etc.
- median estimated in this fashion = a population median
- probability model connects data to population using assumptions
 - population median = *estimand*, sample median = *estimator*

Independence

- two events A and B are *independent* if the following is true
 - $P(A \cap B) = P(A)P(B)$
 - $P(A | B) = P(A)$
- two random variables X and Y are *independent*, if for any two sets, **A** and **B**, the following is true
 - $P([X \in A] \cap [Y \in B]) = P(X \in A)P(Y \in B)$
- **independence** = statistically unrelated from one another
- if A is *independent* of B, then the following are true
 - A^c is independent of B
 - A is independent of B^c
 - A^c is independent of B^c

IID Random Variables

- random variables are said to be **IID** if they are *independent and identically distributed*
 - **independent** = statistically unrelated from each other
 - **identically distributed** = all having been drawn from the same population distribution
- IID random variables = default model for random samples = default starting point of inference

Diagnostic Test

- Let $+$ and $-$ be the results, positive and negative respectively, of a diagnostic test
- Let D = subject of the test has the disease, D^c = subject does not
- **sensitivity** = $P(+ | D)$ = probability that the test is positive given that the subject has the disease (the higher the better)
- **specificity** = $P(- | D^c)$ = probability that the test is negative given that the subject does not have the disease (the higher the better)
- **positive predictive value** = $P(D | +)$ = probability that that subject has the disease given that the test is positive
- **negative predictive value** = $P(D^c | -)$ = probability that the subject does not have the disease given the test is negative
- **prevalence of disease** = $P(D)$ = marginal probability of disease

Example

- specificity of 98.5%, sensitivity = 99.7%, prevalence of disease = .1%

$$\begin{aligned}
 P(D | +) &= \frac{P(+ | D)P(D)}{P(+ | D)P(D) + P(+ | D^c)P(D^c)} \\
 &= \frac{P(+ | D)P(D)}{P(+ | D)P(D) + \{1 - P(- | D^c)\}\{1 - P(D)\}} \\
 &= \frac{.997 \times .001}{.997 \times .001 + .015 \times .999} \\
 &= .062
 \end{aligned}$$

- low positive predictive value \rightarrow due to low prevalence of disease and somewhat modest specificity
 - suppose it was know that the subject uses drugs and has regular intercourse with an HIV infect partner (his probability of being $+$ is higher than suspected)
 - evidence implied by a positive test result

Likelihood Ratios

- from Baye's Rules, we can derive the *positive predictive value* and *false positive value*

$$\begin{aligned}
 P(D | +) &= \frac{P(+ | D)P(D)}{P(+ | D)P(D) + P(+ | D^c)P(D^c)} \\
 P(D^c | +) &= \frac{P(+ | D^c)P(D^c)}{P(+ | D)P(D) + P(+ | D^c)P(D^c)}
 \end{aligned}$$

- if we divide the about quantities over each other (same denominator), we get the following

$$\frac{P(D | +)}{P(D^c | +)} = \frac{P(+ | D)}{P(+ | D^c)} \times \frac{P(D)}{P(D^c)}$$

- **odds** = $p/(1 - p)$
 - $\frac{P(D)}{P(D^c)}$ = odds of disease in absence of test
 - $\frac{P(D | +)}{P(+ | D^c)}$ = odds of disease given a positive test result
- **Diagnostic Likelihood Ratio** of a positive test result is defined as

$$DLR_+ = \frac{P(+ | D)}{P(+ | D^c)}$$

- in previous example, $DLR_+ = .997/(1 - .985) = 66$
- $DLR_- = (1 - .997)/.985 = 0.003$
- **post-test odds of D = $DLR_+ \times$ pre-test odds of D**
 - DLR_+ = the factor by which you multiply your odds in the presence of a positive test to obtain your post-test odds

Expected Values/Mean

- useful for characterizing a distribution (properties of distributions)
- **mean** = characterization of the center of the distribution = *expected value*
- expected value operation = **linear** $\rightarrow E(aX + bY) = aE(X) + bE(Y)$
- **variance/standard deviation** = characterization of how spread out the distribution is
- sample expected values for sample mean and variance will estimate the population counterparts
- **population mean**
 - expected value/mean of a random variable = center of its distribution (center of mass)
 - **discrete variables**
 - * for X with PMF $p(x)$, the population mean is defined as: $E[X] = \sum_x xp(x)$ where the **sum** is taken over all possible values of x
 - * $E[X]$ = center of mass of a collection of location and weights $x, p(x)$
 - * *coin example*
 - $E[X] = 0 \times (1 - p) + 1 \times p = p$
 - **continuous variable**
 - * for X with density $f(x)$, the expected value = the center of mass of the density
 - * instead of summing over discrete values, the expectation **integrates** over a continuous function
 - pdf = $f(x)$
 - $\int xf(x) = \text{area under the curve} = \text{mean/expected value of X}$
- **sample mean**
 - sample mean estimates the population mean
 - * sample mean = center of mass of observed data = empirical mean

$$\bar{X} = \sum_x^n x_i p(x_i)$$

where $p(x_i) = 1/n$

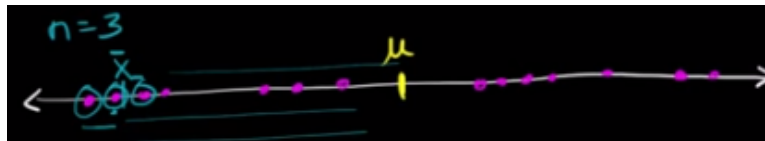
- average of random variables = a random variable and its distribution has an expected value that is the **same** as the original distribution (centers are the same)
 - the mean of the averages = average of the original data \rightarrow estimates average of the population
 - $E[\text{sample mean}] = \text{population mean}$ <- this estimator is **unbiased**
 - * derivation
 - let $X_1, X_2, X_3, \dots, X_n$ be a collection of n samples from the population with mean μ
 - mean of this sample = $\frac{X_1 + X_2 + X_3 + \dots + X_n}{n}$
 - since $E(aX) = aE(X)$, the expected value of the mean, $E[\frac{X_1 + X_2 + X_3 + \dots + X_n}{n}] = \frac{1}{n} \times [E(X_1) + E(X_2) + E(X_3) + \dots + E(X_n)]$
 - since each of the $E(X_i)$ is drawn from the population with mean μ , we expect that the $E(X_i) = \mu$
 - so $\frac{1}{n} \times [E(X_1) + E(X_2) + E(X_3) + \dots + E(X_n)] = \frac{1}{n} \times n \times \mu = \mu$
- **Note:** the more data that goes into the sample mean, the more concentrated its density/mass functions are around the population mean

Variance

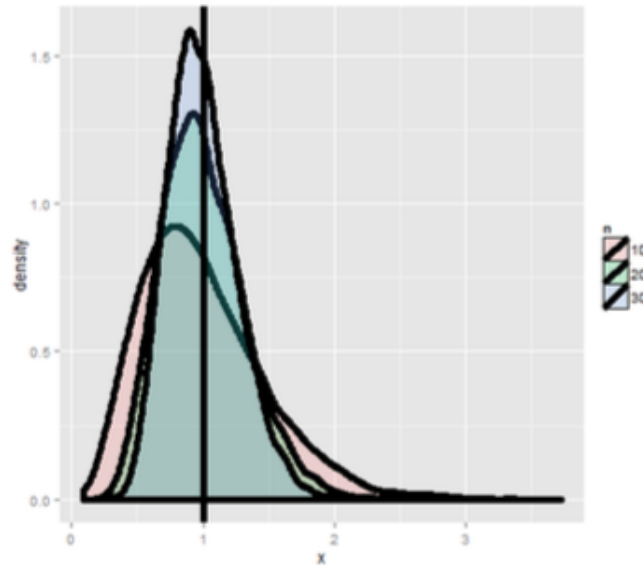
- **variance** = measure of spread, the square of expected distance from the mean (expressed in X's units²)
 - $Var(X) = E[(X - \mu)^2] = E[X^2] - E[X]^2$
 - higher variances \rightarrow more spread, lower \rightarrow smaller spread
 - **standard deviation** = $\sqrt{var(X)}$ \rightarrow has same units as X
 - *example*
 - * for die roll, $E[X] = 3.5$
 - * $E[X^2] = 12 \times 1/6 + 22 \times 1/6 + 32 \times 1/6 + \dots + 62 \times 1/6 = 15.17$
 - * $Var(X) = E[X^2] - E[X]^2 \approx 2.92$
 - *example*
 - * for coin flip, $E[X] = p$
 - * $E[X^2] = 0^2 \times (1 - p) + 1^2 \times p = p$
 - * $Var(X) = E[X^2] - E[X]^2 = p - p^2 = p(1 - p)$

Sample Variance

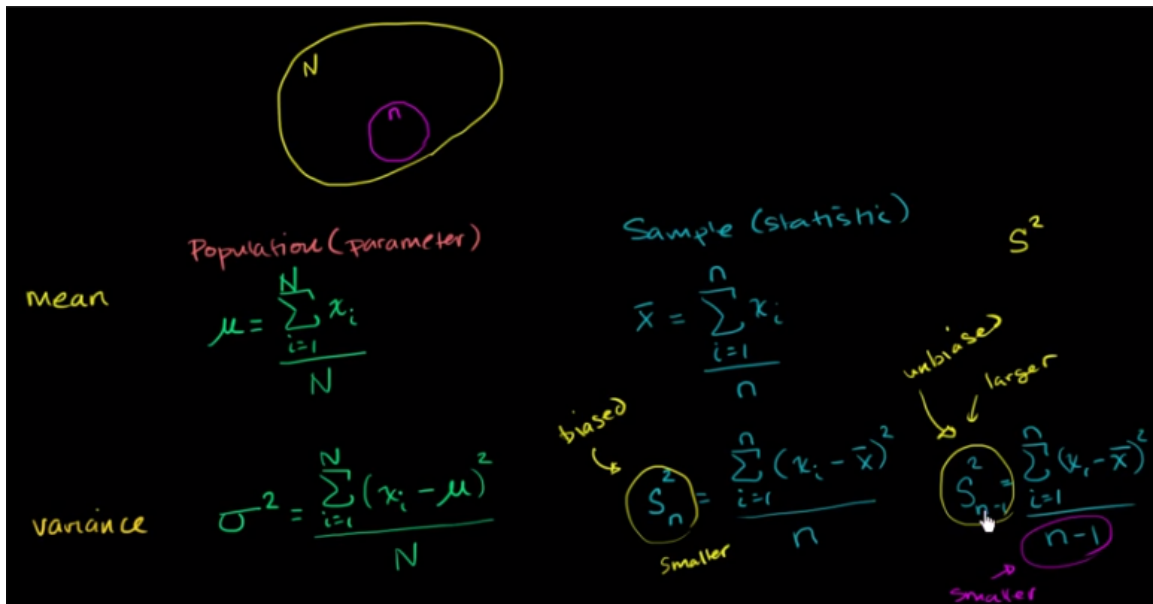
- $S^2 = \frac{\sum_{i=1}^n (X_i - \bar{X})^2}{n-1}$
 - **Note:** samples are much more likely to have variances lower than the population \rightarrow why S^2 is calculated by dividing by $n - 1$



- on the above line, any subset of data will most likely have a variance that is **lower than** the population variance
- dividing by $n - 1$ will make the variance estimator **larger** to adjust for this fact \rightarrow leads to more accurate estimation
 - random variable, and thus has an associate population distribution
 - * $E[S^2] =$ population variance, where S = sample standard deviation
 - * with more data, its distribution gets more concentrated around population variance



- **Note:** for any variable, properties of the population = parameter, estimates of properties for samples = statistic



- distribution for mean of random samples
 - expected value of the mean of this distribution = expected value of the sample = population mean
 - * $E[\bar{X}] = \mu$
 - expected value of the variance of this distribution
 - * $Var(\bar{X}) = \sigma^2/n$
 - * as **n** becomes larger, the mean of random sample → more concentrated around the population mean → variance approaches 0
 - **Note:** normally we only have 1 sample mean (from collected sample) and can estimate the variance σ^2 = so we know a lot about the distribution of the means from the data observed

- **Standard Error (SE)**

- SE of the mean = σ/\sqrt{n} -> effectively the standard deviation of the distribution of a statistic (i.e. mean)
 - * represents variability of means

Entire Estimator-Estimation Relationship

- Start with a sample
- S^2 = sample variance
 - estimates how variable the population is
 - estimates population variance σ^2
 - S^2 = a random variable and has its own distribution centered around σ^2
 - * more concentrated around σ^2 as n increases
- \bar{X} = sample mean
 - estimates population mean μ
 - \bar{X} = a random variable and has its own distribution centered around μ
 - * more concentrated around μ as n increases
 - * variance of distribution of $\bar{X} = \sigma^2/n$
 - * estimate of variance = S^2/n
 - * estimate of standard error = S/\sqrt{n} -> “sample standard error of the mean”
 - * estimates how variable sample means (n size) from the population are

Example - Standard Normal

- variance = 1
- means of **n** standard normals (sample) have standard deviation = $1/\sqrt{n}$

```
# specify number of simulations with 10 as number of observations per sample
nosim <- 1000; n <- 10
sd(apply(matrix(rnorm(nosim * n), nosim), 1, mean))
```

```
## [1] 0.3208222
```

- **rnorm()** -> generate samples from the standard normal
- **matrix()** -> puts all samples into a nosim by n matrix, so that each row represents a simulation with nosim observations
- **apply()** -> calculates the mean of the n samples
- **sd()** -> returns standard deviation
- **Note:** standard uniform -> triangle straight line distribution -> mean = 1/2 and variance = 1/12

Binomial Distribution

- **binomial random variable** = sum of n Bernoulli variables = $\sum X_i$ where $X_i = \text{Bernoulli}(p)$
 - PMF $\rightarrow P(X = x) = \binom{n}{x} p^x (1-p)^{n-x}$
 - * $\binom{n}{x}$ = counts the number of ways selecting x items out of n options without replacement or regard to order
 - $\binom{n}{x} = \frac{n!}{x!(n-x)!}$
 - $\binom{n}{x}, \binom{n}{x} = 1$
- **Bernoulli distribution** \rightarrow binary outcome
 - only possible outcomes
 - * 1 = “success” with probability of p
 - * 0 = “failure” with probability of $1 - p$
 - PMF $\rightarrow P(X = x) = p^x (1-p)^{1-x}$
 - mean = p , variance = $p(1-p)$

Example

- of 8 children, what's the probability of 7 or more girls (50/50 chance)?
- $\binom{8}{7} .5^7 (1-.5)^1 + \binom{8}{8} .5^8 (1-.5)^0 \approx 0.04$
- `choose(8, 7)` \rightarrow R function to calculate n choose x
- `pbinom(6, size=8, prob=0.5, lower.tail=F)` \rightarrow probability of 6 or less out of 8 samples with probability of 0.5
 - `lower.tail = F` returns the complement

Normal Distribution

- normal/Gaussian distribution = random variable X
 - mean = μ , variance = σ^2
 - PMF $\rightarrow (2\pi\sigma^2)^{-1/2} e^{-(x-\mu)^2/2\sigma^2}$
 - notation $\rightarrow X \sim N(\mu, \sigma^2)$
- $X \sim N(0, 1)$ = **standard normal distribution** (standard normal RVs often labeled Z)
 - ~68% of data/normal density \rightarrow between ± 1 standard deviation from μ
 - ~95% of data/normal density \rightarrow between ± 2 standard deviation from μ
 - ~99% of data/normal density \rightarrow between ± 3 standard deviation from μ
 - ± 1.28 standard deviations from $\mu \rightarrow 10^{th}$ (-) and 90^{th} (+) percentiles
 - ± 1.645 standard deviations from $\mu \rightarrow 5^{th}$ (-) and 95^{th} (+) percentiles
 - ± 1.96 standard deviations from $\mu \rightarrow 2.5^{th}$ (-) and 97.5^{th} (+) percentiles
 - ± 2.33 standard deviations from $\mu \rightarrow 1^{st}$ (-) and 99^{th} (+) percentiles
- for any $X \sim N(\mu, \sigma^2)$, calculating the number of standard deviation from the mean **converts the random variable to a standard normal**

$$Z = \frac{X - \mu}{\sigma} \sim N(0, 1)$$

- conversely, **a standard normal can then be converted to any normal distribution** by multiplying by standard deviation and adding the mean

$$X = \mu + \sigma Z \sim N(\mu, \sigma^2)$$

- R Commands:
 - n^{th} percentiles \rightarrow `qnorm(n, mean = mu, sd = sd)`
 - probability larger than $x \rightarrow$ `pnorm(x, mean = mu, sd = sd, lower.tail = F)`

Poisson Distribution

- used to model counts
 - PMF \rightarrow

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

where $X = 0, 1, 2, \dots, \infty$

 - mean = λ , variance = λ
- modeling uses for Poisson distribution
 - count data
 - event-time/survival \rightarrow cancer trials, some patients never develop and some do, dealing with the data for both (“censoring”)
 - contingency tables \rightarrow record results for different characteristic measurements
 - approximating binomials \rightarrow instances where **n** is large and **p** is small (i.e. pollution on lung disease)
 - * $X \sim \text{Binomial}(n, p)$
 - * $\lambda = np$
 - rates $\rightarrow X \sim \text{Poisson}(\lambda t)$
 - * $\lambda = E[X/t] \rightarrow$ expected count per unit of time
 - * $t =$ total monitoring time
 - * example: `ppois(n, lambda = lambda * t)` \rightarrow returns probability of n or fewer events happening given the rate and time

Asymptotics

- **asymptotics** = behavior of statistics as sample size $\rightarrow \infty$
- useful for simple statistical inference/approximations
- form basis for frequency interpretation of probabilities (“Law of Large Numbers”)
- **Law of Large Numbers (LLN)** = IID sample statistic becomes population statistic to what it estimates as **n** increases (sample mean \rightarrow population mean)
- **Note:** an estimator is **consistent** if it converges to what it is estimating
 - sample mean, variance, standard deviation are all consistent for their population counterparts
- **Central Limit Theorem**
 - distribution of means of IID variables \rightarrow standard normal as **n** increases
 - for large values of **n**

$$\frac{\bar{X}_n - \mu}{\sigma/\sqrt{n}} = \frac{\sqrt{n}(\bar{X}_n - \mu)}{\sigma} = \frac{\text{Estimate} - \text{Mean of estimate}}{\text{Std. Err. of estimate}} \rightarrow N(0, 1)$$

- this translates to the distribution of the sample mean \bar{X} is approximately $N(\mu, \sigma^2/n)$
- **Note:** speed at which the normalized coin flips converge to normal distribution depends on how biased the coin is (value of p)
- **Note:** does not guarantee that the normal distribution will be a good approximation, but just that eventually it will be a good approximation as $n \rightarrow \infty$

Confidence Intervals - Z (using Central Limit Theorem)

- Z confidence interval

$$Estimate \pm ZQ \times SE_{Estimate}$$

- ZQ = quantile from the standard normal distribution
- sample mean = $\bar{X} \sim N(\mu, \sigma^2/n)$, with mean = μ and standard deviation = σ^2/n
- **95% confidence interval for $\mu = \bar{X} \pm 2\sigma^2/n$** (1.96 to be more accurate)
 - probability that \bar{X} is larger than $\mu + 2\sigma^2/n$ or smaller than $\mu - 2\sigma^2/n = 5\%$
 - interpretation: if we were to repeated samples of size n from the population and construct this confidence interval for each case, approximately 95% of the intervals will contain μ
- **Note:** Poisson and binomial distributions have exact intervals that don't require CLT
- **Wald confidence interval**
 - for Bernoulli distributions, confidence interval takes the form

$$\hat{p} \pm z_{1-\alpha/2} \sqrt{\frac{p(1-p)}{n}}$$

- p = unknown, so use $\hat{p} = X/n$ to replace
- $p(1-p) \rightarrow$ largest when $p = 1/2$, confidence interval becomes $\hat{p} \pm \frac{1}{\sqrt{n}}$
- this is useful in **roughly estimating confidence intervals**
 - * generally need $n = 100$ for 1 decimal place, 10,000 for 2, and 1,000,000 for 3
- `binom.test(success, n)` = returns confidence interval
- **Agresti/Coull interval** (binomial distribution)
 - for smaller values of n , when n is not large enough for CLT
 - take number of successes, X , add 2
 - take number of failure, add 2
 - $\hat{p} = \frac{X+2}{n+4}$, use this to construct confidence interval (tend to be **conservative**)
- **Poisson Interval**
 - $X \sim Poisson(\lambda t)$
 - * estimate rate $\hat{\lambda} = X/t$
 - * $var(\hat{\lambda}) = \lambda/t$
 - * variance estimate = $\hat{\lambda}/t$
 - for small values of λ (few events larger time interval), should not use the asymptotic interval estimated here
 - as $t \rightarrow \infty$, the interval becomes the true 95% interval

Confidence Intervals - T (small samples)

- t Confidence Interval

$$Estimate \pm TQ \times SE_{Estimate} = \bar{X} \pm \frac{t_{n-1}S}{\sqrt{n}}$$

- TQ = quantile from T distribution
- t_{n-1} = relevant quantile = $qt(0.975, df = n-1)$
- t interval assumes data is IID normal
- works well with data distributions that are roughly symmetric/mound shaped, and **does not** work with skewed distributions
 - * skewed distribution \rightarrow meaningless to center interval around the mean \bar{X}
 - * logs/median can be used instead
- paired observations (multiple measurements from same subjects) can be analyzed by t interval of differences
- as more data collected (large degrees of freedom), t interval \rightarrow z interval
- William Gosset's **t** Distribution ("Student's T distribution")
 - test = Gosset's pseudonym which he published under
 - indexed/defined by **degrees of freedom**, and becomes more like standard normal as degrees of freedom gets larger
 - thicker tails centered around 0, thus confidence interval = **wider** than Z interval (more mass concentrated away from the center)
 - for **small** sample size (value of n), normalizing the distribution by $\frac{\bar{X}-\mu}{S/\sqrt{n}} \rightarrow$ t distribution, **not** the standard normal distribution
 - * S = standard deviation may be inaccurate, as the std of the data sample may not be truly representative of the population std
 - * using the Z interval here thus may produce an interval that is too **narrow**
- Independent Group **t** Intervals - Same Variance
 - compare two groups in randomized trial ("A/B Testing")
 - * cannot use the paired t test because the groups are independent and may have different sample sizes
 - perform randomization to balance unobserved covariance that may otherwise affect the result
 - Confidence Interval for $\mu_y - \mu_x$

$$\bar{Y} - \bar{X} \pm t_{n_x+n_y-2, 1-\alpha/2} S_p \left(\frac{1}{n_x} + \frac{1}{n_y} \right)^{1/2}$$

- * $t_{n_x+n_y-2, 1-\alpha/2}$ = relevant quantile
- * $n_x + n_y - 2$ = degrees of freedom
- * $S_p \left(\frac{1}{n_x} + \frac{1}{n_y} \right)^{1/2}$ = standard error
- * $S_p^2 = \{(n_x - 1)S_x^2 + (n_y - 1)S_y^2\} / (n_x + n_y - 2)$ = pooled variance estimator
 - This is effectively a weighted average between the two variances, such that different sample sizes are taken in to account
- * **Note:** this interval assumes **constant variance** across two groups; if variance is different, use the next interval
- Independent Group **t** Intervals - Different Variance
 - Confidence Interval for $\mu_y - \mu_x$

$$\bar{Y} - \bar{X} \pm t_{df} \times \left(\frac{s_x^2}{n_x} + \frac{s_y^2}{n_y} \right)^{1/2}$$

- * t_{df} = relevant quantile with df as defined below
- * **Note:** normalized statistic does not follow t distribution but can be approximated through the formula with df defined below

*

$$df = \frac{(S_x^2/n_x + S_y^2/n_y)^2}{\left(\frac{S_x^2}{n_x}\right)^2/(n_x - 1) + \left(\frac{S_y^2}{n_y}\right)^2/(n_y - 1)}$$

$$* \left(\frac{s_x^2}{n_x} + \frac{s_y^2}{n_y} \right)^{1/2} = \text{standard error}$$

- Comparing other kinds of data
 - binomial → relative risk, risk difference, odds ratio
 - binomial → Chi-squared test, normal approximations, exact tests
 - count → Chi-squared test, exact tests
- R commands
 - t Confidence Intervals
 - * `mean + c(-1, 1) * qt(0.975, n - 1) * std / sqrt(n)`
 - `c(-1, 1)` = plus and minus, \pm
 - Difference Intervals (all equivalent)
 - * `mean2 - mean1 + c(-1, 1) * qt(0.975, n - 1) * std / sqrt(n)`
 - `n` = number of paired observations
 - `qt(0.975, n - 1)` = relevant quantile for paired
 - `qt(0.975, nx + ny - 2)` = relevant quantile for independent
 - * `t.test(mean2 - mean1)`
 - * `t.test(data2, data1, paired = TRUE, var.equal = TRUE)`
 - ***paired*** = whether or not the two sets of data are paired (same subjects different observations for treatment) <- **TRUE** for paired, **FALSE** for independent
 - ***var.equal*** = whether or not the variance of the datasets should be treated as equal <- **TRUE** for same variance, **FALSE** for unequal variances
 - * `t.test(extra ~ I(relevel(group, 2)), paired = TRUE, data = sleep)`
 - ***relevel(factor, ref)*** = reorders the levels in the factor so that “ref” is changed to the first level → doing this here is so that the second set of measurements come first (1, 2 → 2, 1) in order to perform `mean2 - mean1`
 - ***I(object)*** = prepend the class “AsIs” to the object
 - **Note:** `I(relevel(group, 2))` = explanatory variable, must be **factor** and have **two levels**

Hypothesis Testing

- Hypothesis testing = making decisions using data
 - **null** hypothesis (H_0) = status quo
 - assumed to be **true** → statistical evidence required to reject it for **alternative** or “research” hypothesis (H_a)
 - * alternative hypothesis typically take form of $>$, $<$ or \neq
 - Results

Truth	Decide	Result
H_0	H_0	Correctly accept null
H_0	H_a	Type I error
H_a	H_a	Correctly reject null
H_a	H_0	Type II error

- α = Type I error rate
 - probability of **rejecting** the null hypothesis when the hypothesis is **correct**
 - $\alpha = 0.5$ → standard for hypothesis testing
 - **Note:** as Type I error rate increases, Type II error rate decreases and vice versa
- for large samples (large n), use the **Z Test** for $H_0 : \mu = \mu_0$
 - H_a :
 - * $H_1 : \mu < \mu_0$
 - * $H_2 : \mu \neq \mu_0$
 - * $H_3 : \mu > \mu_0$
 - Test statistic $TS = \frac{\bar{X} - \mu_0}{S/\sqrt{n}}$
 - Reject the null hypothesis H_0 when
 - * $H_1 : TS \leq Z_\alpha$ OR $-Z_{1-\alpha}$
 - * $H_2 : |TS| \geq Z_{1-\alpha/2}$
 - * $H_3 : TS \geq Z_{1-\alpha}$
 - **Note:** In case of $\alpha = 0.5$ (most common), $Z_{1-\alpha} = 1.645$ (95 percentile)
 - α = low, so that when H_0 is rejected, original model → wrong or made an error (low probability)
- For small samples (small n), use the **T Test** for $H_0 : \mu = \mu_0$
 - H_a :
 - * $H_1 : \mu < \mu_0$
 - * $H_2 : \mu \neq \mu_0$
 - * $H_3 : \mu > \mu_0$
 - Test statistic $TS = \frac{\bar{X} - \mu_0}{S/\sqrt{n}}$
 - Reject the null hypothesis H_0 when
 - * $H_1 : TS \leq T_\alpha$ OR $-T_{1-\alpha}$
 - * $H_2 : |TS| \geq T_{1-\alpha/2}$
 - * $H_3 : TS \geq T_{1-\alpha}$
 - **Note:** In case of $\alpha = 0.5$ (most common), $T_{1-\alpha} = qt(.95, df = n-1)$
 - R commands for T test:

- * `t.test(vector1 - vector2)`
- * `t.test(vector1, vector2, paired = TRUE)`
 - `alternative` argument can be used to specify one-sided tests: `less` or `greater`
 - `alternative` default = `two-sided`
- * prints test statistic (`t`), degrees of freedom (`df`), **p-value**, 95% confidence interval, and mean of sample
 - confidence interval in units of data, and can be used to interpret the practical significance of the results
- **rejection region** = region of TS values for which you reject H_0
- **power** = probability of rejecting H_0
 - power is used to calculate sample size for experiments
- **two-sided tests** $\rightarrow H_a : \mu \neq \mu_0$
 - reject H_0 only if test statistic is too larger/small
 - for $\alpha = 0.5$, split equally to 2.5% for upper and 2.5% for lower tails
 - * equivalent to $|TS| \geq T_{1-\alpha/2}$
 - * example: for T test, `qt(.975, df)` and `qt(.025, df)`
 - **Note:** *failing to reject one-sided test = fail to reject two-sided*
- **tests vs confidence intervals**
 - $(1 - \alpha)\%$ confidence interval for μ = set of all possible values that fail to reject H_0
 - if $(1 - \alpha)\%$ confidence interval contains μ_0 , fail to reject H_0
- **two-group intervals/test**
 - Rejection rules the same
 - Test $H_0: \mu_1 = \mu_2 \rightarrow \mu_1 - \mu_2 = 0$
 - Test statistic:

$$\frac{\text{Estimate} - H_0\text{Value}}{SE_{\text{Estimate}}} = \frac{\bar{X}_1 - \bar{X}_2 - 0}{\sqrt{\frac{S_1^2}{n_1} + \frac{S_2^2}{n_2}}}$$
 - R Command
 - * `t.test(values ~ factor, paired = FALSE, var.equal = TRUE, data = data)`
 - `paired = FALSE` \rightarrow independent values
 - `factor` argument must have only two levels
- **p values**
 - most common measure of statistical significance
 - **p-value** = probability under the null hypothesis of obtaining evidence as extreme or more than that of the obtained
 - * Given that H_0 is true, how likely is it to obtain the result (test statistic)?
 - **attained significance level** = smallest value for α for which H_0 is rejected \rightarrow equivalent to p-value
 - * if p-value $< \alpha$, reject H_0
 - * for two-sided tests, double the p-values
 - if p-value is small, either H_0 is true AND the observed is a rare event **OR** H_0 is false
 - R Command
 - * `p-value = pt(statistic, df, lower.tail = FALSE)`
 - `lower.tail = FALSE` = returns the probability of getting a value from the t distribution that is larger than the test statistic

- * Binomial (coin flips)
 - probability of getting x results out of n trials and event probability of p = `pbinom(x, size = n, prob = p, lower.tail = FALSE)`
 - two-sided interval (testing for \neq): find the smaller of two one-sided intervals ($X < \text{value}$, $X > \text{value}$), and double the result
 - ***Note:** `lower.tail = FALSE` = strictly greater*
- * Poisson
 - probability of getting x results given the rate r = `ppois(x - 1, r, lower.tail = FALSE)`
 - $x - 1$ is used here because the upper tail includes the specified number (since we want greater than x , we start at $x - 1$)
 - r = events that should occur given the rate (multiplied by 100 to yield an integer)
 - ***Note:** `lower.tail = FALSE` = strictly greater*

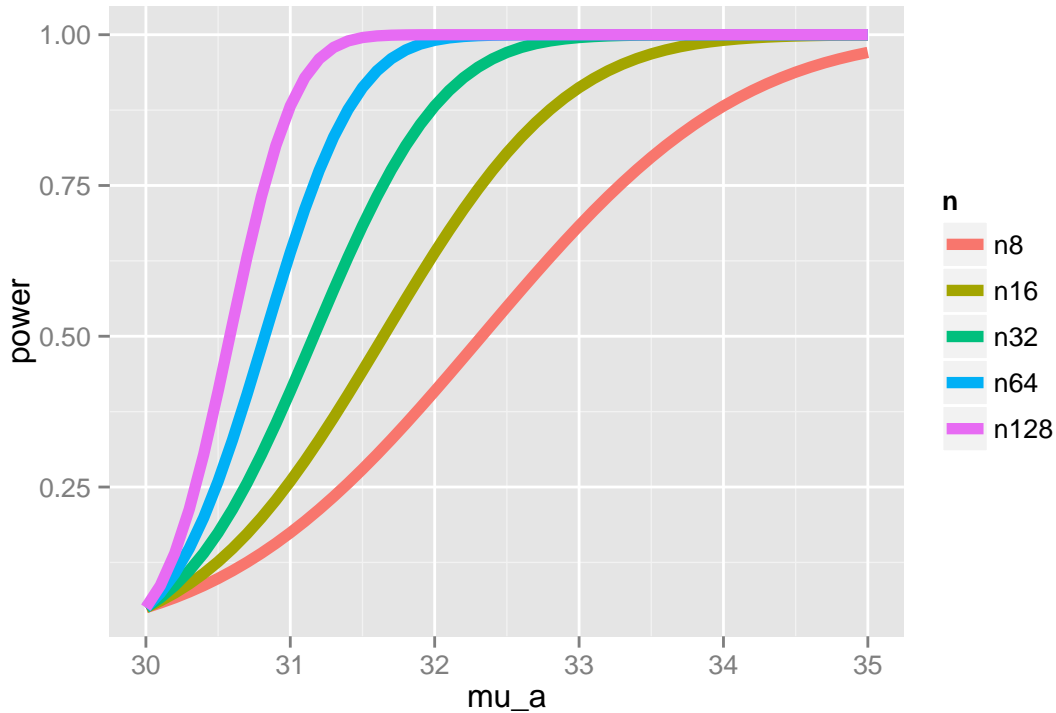
Power

- **Power** = probability of rejecting the null hypothesis when it is false (the more power the better)
 - most often used in designing studies so that there's a reasonable chance to detect the alternative hypothesis if the alternative hypothesis is true
- β = probability of type II error = failing to reject the null hypothesis when it's false
- power = $1 - \beta$
- **example**
 - $H_0 : \mu = 30 \rightarrow \bar{X} \sim N(\mu_0, \sigma^2/n)$
 - $H_a : \mu > 30 \rightarrow \bar{X} \sim N(\mu_a, \sigma^2/n)$
 - Power:

$$\text{Power} = P\left(\frac{\bar{X} - 30}{s/\sqrt{n}} > t_{1-\alpha, n-1} ; \mu = \mu_a\right)$$

- * **Note:** the above function depends on value of μ_a
- * **Note:** as μ_a approaches 30, power approaches α
- assuming the sample mean is normally distributed, H_0 is rejected when $\frac{\bar{X} - 30}{\sigma/\sqrt{n}} > Z_{1-\alpha}$
- or, $\bar{X} > 30 + Z_{1-\alpha} \frac{\sigma}{\sqrt{n}}$
- R commands:
 - `alpha = 0.05; z = qnorm(1-alpha)` -> calculates $Z_{1-\alpha}$
 - `pnorm(mu0 + z * sigma/sqrt(n), mean = mua, sd = sigma/sqrt(n), lower.tail = FALSE)` -> calculates the probability of getting a sample mean that is larger than $Z_{1-\alpha} \frac{\sigma}{\sqrt{n}}$ given that the population mean is μ_a
 - * **Note:** using `mean = mu0` in the function would = `alpha`
 - Power curve behavior
 - * Power increases as μ_a increases -> we are more likely to detect the difference in μ_a and μ_0
 - * Power increases as **n** increases -> with more data, more likely to detect any alternative μ_a

```
library(ggplot2)
mu0 = 30; mua = 32; sigma = 4; n = 16
alpha = 0.05
z = qnorm(1 - alpha)
nseq = c(8, 16, 32, 64, 128)
mu_a = seq(30, 35, by = 0.1)
power = sapply(nseq, function(n)
  pnorm(mu0 + z * sigma / sqrt(n), mean = mu_a, sd = sigma / sqrt(n),
    lower.tail = FALSE)
)
colnames(power) <- paste("n", nseq, sep = "")
d <- data.frame(mu_a, power)
library(reshape2)
d2 <- melt(d, id.vars = "mu_a")
names(d2) <- c("mu_a", "n", "power")
g <- ggplot(d2,
  aes(x = mu_a, y = power, col = n)) + geom_line(size = 2)
g
```



- **Solving for Power**

- When testing $H_a : \mu > \mu_0$ (or $<$ or \neq)

$$Power = 1 - \beta = P\left(\bar{X} > \mu_0 + Z_{1-\alpha} \frac{\sigma}{\sqrt{n}}; \mu = \mu_a\right)$$

where $\bar{X} \sim N(\mu_a, \sigma^2/n)$

- Unknowns = μ_a, σ, n, β
- Knowns = μ_0, α
- Specify any 3 of the unknowns and you can solve for the remainder; most common are two cases
 1. Given power desired, mean to detect, variance that we can tolerate, find the **n** to produce desired power (designing experiment/trial)
 2. Given the size **n** of the sample, find the power that is achievable (finding the utility of experiment)
- **Note:** for $H_a : \mu \neq \mu_0$, calculated one-sided power using $z_{1-\alpha/2}$; however, the power calculation here excludes the probability of getting a large TS in the opposite direction of the truth, but this is only applicable when μ_a and μ_0 are close together

- **Power Behavior**

- Power increases as α becomes larger
- Power of one-sided test $>$ power of associated two-sided test
- Power increases as μ_a gets further away from μ_0
- Power increases as **n** increases (sample mean has less variability)
- Power increases as σ decreases (again less variability)
- Power usually depends only $\frac{\sqrt{n}(\mu_a - \mu_0)}{\sigma}$, and not μ_a, σ , and n
 - * **effect size** = $\frac{\mu_a - \mu_0}{\sigma}$ \rightarrow unit free, can be interpreted across settings

- **T-test Power**

- for Gossett's T test,

$$Power = P\left(\frac{\bar{X} - \mu_0}{S/\sqrt{n}} > t_{1-\alpha, n-1}; \mu = \mu_a\right)$$

- * $\frac{\bar{X} - \mu_0}{S/\sqrt{n}}$ does not follow a t distribution if the true mean is μ_a and NOT μ_0 → follows a non-central t distribution instead
- **power.t.test** → evaluates the non-central t distribution and solves for a parameter given all others are specified
 - * **power.t.test**(n = 16, delta = 0.5, sd = 1, type = "one.sample", alt = "one.sided")\$power
→ calculates power with inputs of n, difference in means, and standard deviation
 - delta = argument for difference in means
 - **Note:** since effect size = *delta/sd*, as *n*, *type*, and *alt* are held constant, any distribution with the same effect size will have the same power
 - * **power.t.test**(power = 0.8, delta = 0.5, sd = 1, type = "one.sample", alt = "one.sided")\$n → calculates size n with inputs of power, difference in means, and standard deviation
 - **Note:** n should always be rounded up (ceiling)

Multiple Testing

- Hypothesis testing/significant analysis commonly overused
- correct for multiple testing to avoid false positives/conclusions (two key components)
 1. error measure
 2. correction
- multiple testing is needed because of the increase in ubiquitous data collection technology and analysis
 - DNA sequencing machines
 - imaging patients in clinical studies
 - electronic medical records
 - individualized movement data (fitbit)

Type of Errors

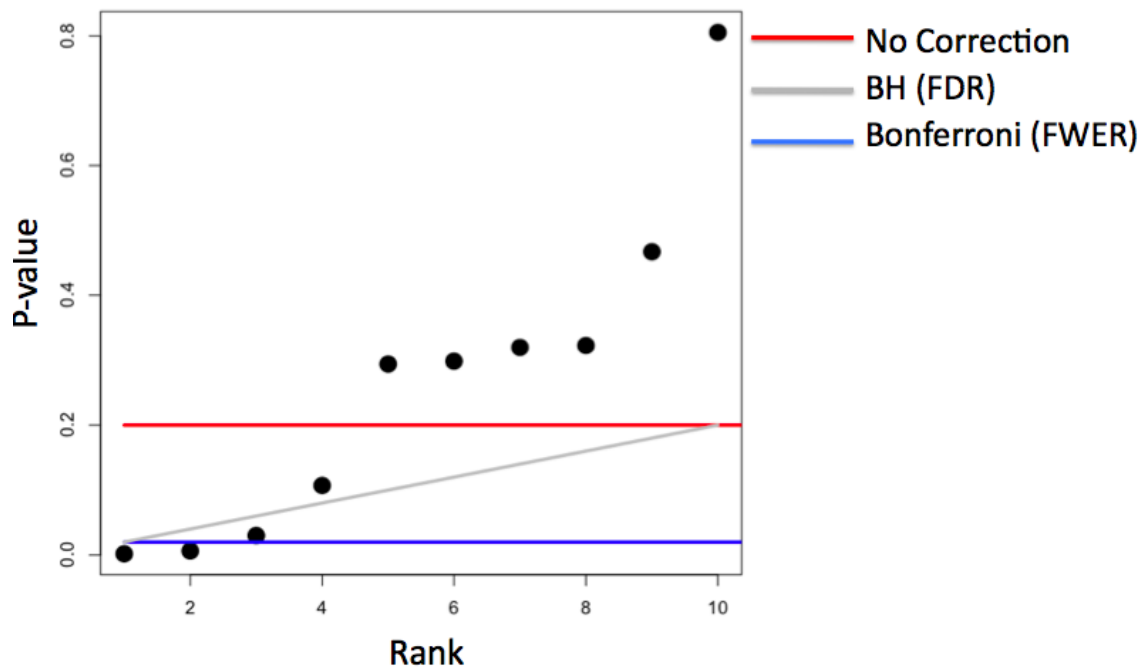
	Actual H_0 = True		Actual H_a = True		Total
Conclude H_0 = True (non-significant)	U	T	$m - R$	Conclude H_a = True (significant)	V
H_a = True (significant)	V	S	R	Total	m_0
					$m - m_0$
					m

- m_0 = number of true null hypotheses, or cases where H_0 = actually true (unknown)
- $m - m_0$ = number of true alternative hypotheses, or cases where H_a = actually true (unknown)
- R = number of null hypotheses rejected, or cases where H_a = concluded to be true (measurable)
- $m - R$ = number of null hypotheses that failed to be rejected, or cases where H_0 = concluded to be true (measurable)
- V = Type I Error / false positives, concludes H_a = True when H_0 = actually True
- T = Type II Error / false negatives, concludes H_0 = True when H_a = actually True
- S = true positives, concludes H_a = True when H_a = actually True
- U = true negatives, concludes H_0 = True when H_0 = actually True

Error Rates

- **false positive rate** = rate at which false results are called significant $E[\frac{V}{m_0}] \rightarrow$ average fraction of times that H_a is claimed to be true when H_0 is actually true
 - **Note:** mathematically equal to type I error rate \rightarrow false positive rate is associated with a post-prior result, which is the expected number of false positives divided by the total number of hypotheses under the real combination of true and non-true null hypotheses (disregarding the “global null” hypothesis). Since the false positive rate is a parameter that is not controlled by the researcher, it cannot be identified with the significance level, which is what determines the type I error rate.
- **family wise error rate (FWER)** = probability of at least one false positive $Pr(V \geq 1)$
- **false discovery rate (FDR)** = rate at which claims of significance are false $E[\frac{V}{R}]$
- **controlling error rates (adjusting α)**
 - false positive rate
 - * if we call all $P < \alpha$ significant (reject H_0), we are expected to get $\alpha \times m$ false positives, where m = total number of hypothesis test performed
 - * with high values of m , false positive rate is very large as well

- family-wise error rate (FWER)
 - * controlling FWER = controlling the probability of even one false positive
 - * *bonferroni* correction (oldest multiple testing correction)
 - for m tests, we want $Pr(V \geq 1) < \alpha$
 - calculate P-values normally, and deem them significant if and only if $P < \alpha_{fewer} = \alpha/m$
 - * easy to calculate, but tend to be very **conservative**
- false discovery rate (FDR)
 - * most popular correction = controlling FDR
 - * for m tests, we want $E[\frac{V}{R}] \leq \alpha$
 - * calculate P-values normally and sort some from smallest to largest $\rightarrow P_{(1)}, P_{(1)}, \dots, P_{(m)}$
 - * deem the P-values significant if $P_{(i)} \leq \alpha \times \frac{i}{m}$
 - * easy to calculate, less conservative, but allows for more false positives and may behave strangely under dependence (related hypothesis tests/regression with different variables)
- **example**
 - * 10 P-values with $\alpha = 0.20$



- **adjusting for p-values**

- **Note:** changing P-values will fundamentally change their properties but they can be used directly without adjusting /alpha
- *bonferroni* (FWER)
 - * $P_i^{fewer} = \max(mP_i, 1) \rightarrow$ since p cannot exceed value of 1
 - * deem P-values significant if $P_i^{fewer} < \alpha$
 - * similar to controlling FWER

Example

```

set.seed(1010093)
pValues <- rep(NA,1000)
for(i in 1:1000){
  x <- rnorm(20)
  # First 500 beta=0, last 500 beta=2
  if(i <= 500){y <- rnorm(20)}else{ y <- rnorm(20,mean=2*x)}
  # calculating p-values by using linear model; the [2, 4] coeff in result = pvalue
  pValues[i] <- summary(lm(y ~ x))$coeff[2,4]
}
# Controls false positive rate
trueStatus <- rep(c("zero","not zero"),each=500)
table(pValues < 0.05, trueStatus)

```

```

##           trueStatus
##           not zero zero
##  FALSE           0  476
##  TRUE           500   24

```

```

# Controls FWER
table(p.adjust(pValues,method="bonferroni") < 0.05,trueStatus)

```

```

##           trueStatus
##           not zero zero
##  FALSE           23  500
##  TRUE           477    0

```

```

# Controls FDR (Benjamin Hochberg)
table(p.adjust(pValues,method="BH") < 0.05,trueStatus)

```

```

##           trueStatus
##           not zero zero
##  FALSE           0  487
##  TRUE           500   13

```

Resample Inference

- **Bootstrap** = useful tool for constructing confidence intervals and calculating standard errors for difficult statistics
 - *principle* = if a statistic's (i.e. median) sampling distribution is unknown, then use distribution defined by the data to approximate it
 - *procedures*
 1. simulate n observations **with replacement** from the observed data → results in 1 simulated complete data set
 2. calculate desired statistic (i.e. median) for each simulated data set
 3. repeat the above steps B times, resulting in B simulated statistics
 4. these statistics are approximately drawn from the sampling distribution of the true statistic of n observations
 5. perform one of the following
 - * plot a histogram
 - * calculate standard deviation of the statistic to estimate its standard error
 - * take quantiles (2.5th and 97.5th) as a confidence interval for the statistic (“*bootstrap CI*”)
 - *example*
 - * Bootstrap procedure for calculating confidence interval for the median from a data set of n observations → approximate sampling distribution

```
# load data
library(UsingR); data(father.son)
# observed dataset
x <- father.son$sheight
# number of simulated statistic
B <- 1000
# generate samples
resamples <- matrix(
  sample(x,                # sample to draw from
        n * B,            # draw B datasets with n observations each
        replace = TRUE), # cannot draw n*B elements from x (has n elements) without replacement
  B, n)                  # arrange results into n x B matrix
                        # (every row = bootstrap sample with n observations)
# take median for each row/generated sample
medians <- apply(resamples, 1, median)
# estimated standard error of median
sd(medians)
```

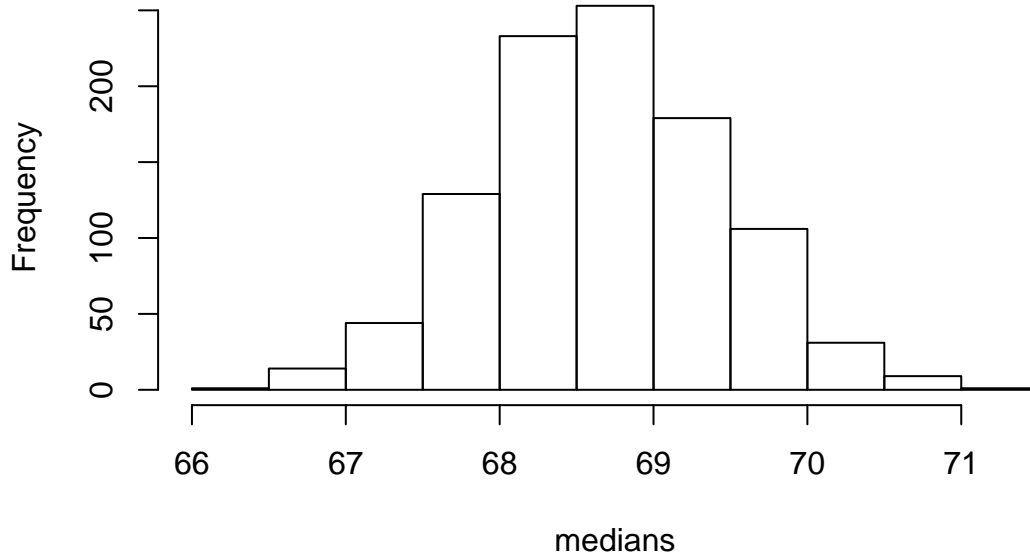
```
## [1] 0.76595
```

```
# confidence interval of median
quantile(medians, c(.025, .975))
```

```
##      2.5%      97.5%
## 67.18292 70.16488
```

```
# histogram of bootstrapped samples
hist(medians)
```

Histogram of medians



- **Note:** better percentile bootstrap confidence interval = “bias corrected and accelerated interval” in *bootstrap* package

- **Permutation Tests**

- *procedures*

- * compare groups of data and test the null hypothesis that the distribution of the observations from each group = same
 - **Note:** if this is true, then group labels/divisions are irrelevant
- * permute the labels for the groups
- * recalculate the statistic
 - Mean difference in counts
 - Geometric means
 - T statistic
- * Calculate the percentage of simulations where the simulated statistic was more extreme (toward the alternative) than the observed

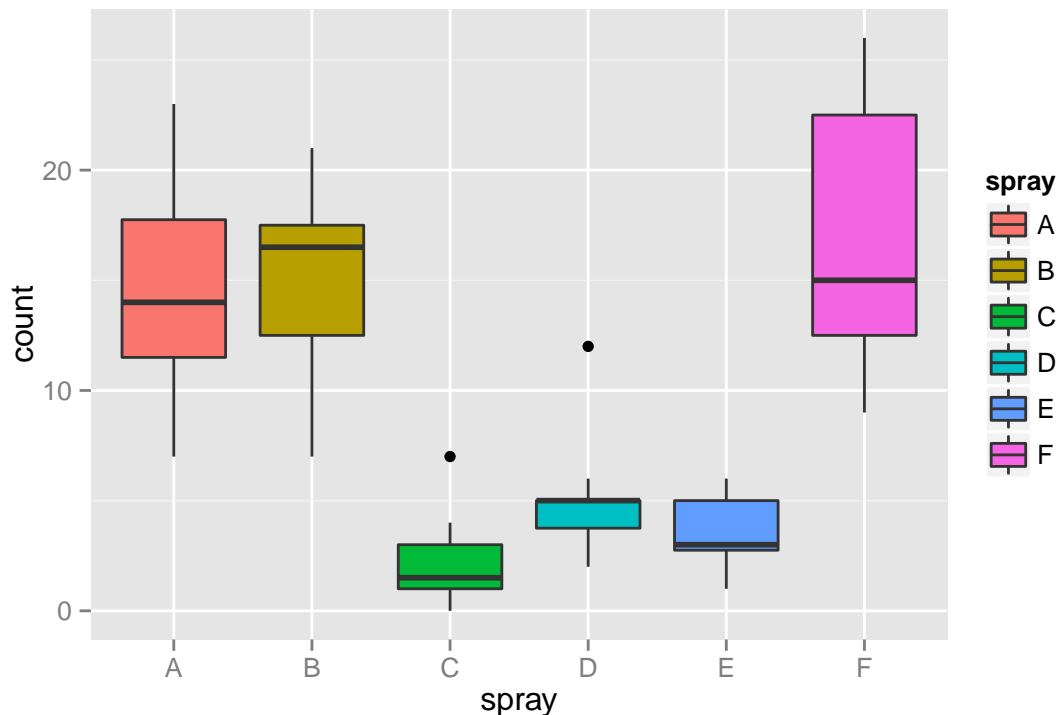
- *variations*

Data type	Statistic	Test name
Ranks	rank sum	rank sum test
Binary	hypergeometric prob	Fisher's exact test
Raw data		ordinary permutation test

- * **Note:** randomization tests are exactly permutation tests, with a different motivation
- * For matched data, one can randomize the signs
- * For ranks, this results in the **signed rank test**
- * Permutation strategies work for regression by permuting a regressor of interest
- * Permutation tests work very well in multivariate settings

– *example*

- * we will compare groups **B** and **C** in this dataset for null hypothesis H_0 : there are no difference between the groups



- we will compare groups **B** and **C** in this dataset for null hypothesis H_0 : there are no difference between the groups

```
# subset to only "B" and "C" groups
subdata <- InsectSprays[InsectSprays$spray %in% c("B", "C"),]
# values
y <- subdata$count
# labels
group <- as.character(subdata$spray)
# find mean difference between the groups
testStat <- function(w, g) mean(w[g == "B"]) - mean(w[g == "C"])
observedStat <- testStat(y, group)
observedStat
```

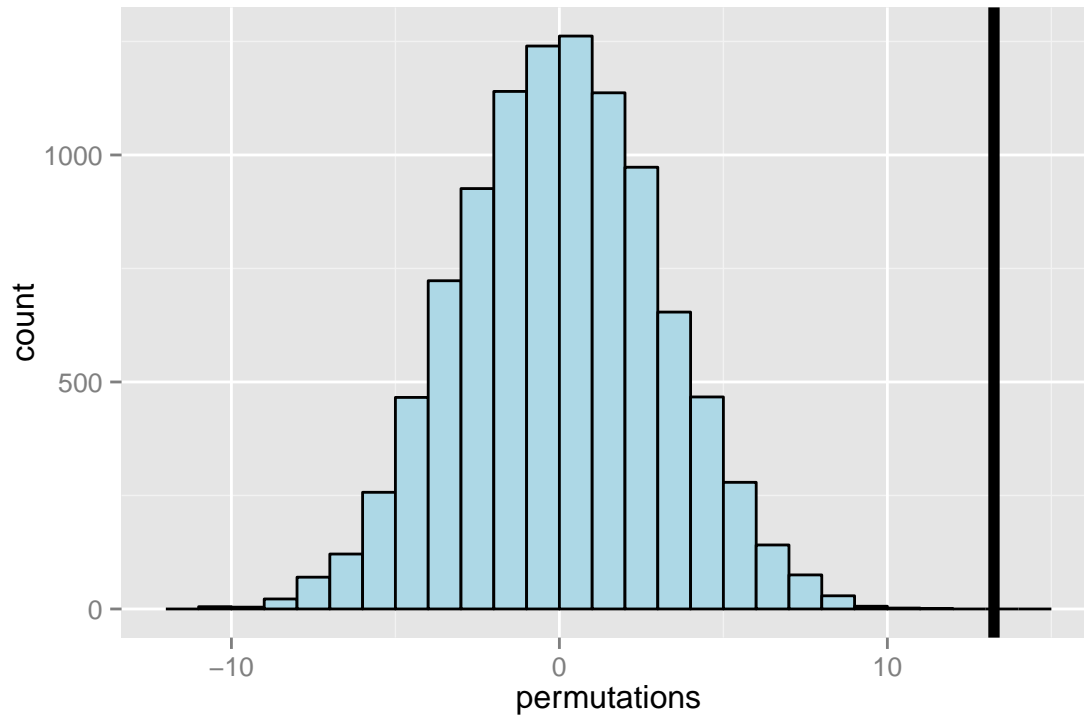
```
## [1] 13.25
```

- the observed difference between the groups is 13.25
- now we changed the resample the lables for groups **B** and **C**

```
# create 10000 permutations of the data with the labels' changed
permutations <- sapply(1 : 10000, function(i) testStat(y, sample(group)))
# find the number of permutations whose difference that is bigger than the observed
mean(permutations > observedStat)
```

```
## [1] 0
```

- we created 1000 permutations from the observed dataset, and found ***no datasets*** with mean differences between groups **B** and **C** larger than the original data
- therefore, p-value is very small and we can ***reject the null*** hypothesis with any reasonable α levels
- below is the plot for the null distribution/permutations



- as we can see from the black line, the observed difference/statistic is very far from the mean \rightarrow likely 0 is ***not*** the true difference
 - with this information, formal confidence intervals can be constructed and p-values can be calculated