## Final Review

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Hypothesis Testing
We make decisions every day in our lives.
Should I believe $A$ or should I believe $B(\operatorname{not} A)$ ?
Two Competing Hypotheses. $A$ and $B$.
Null Hypothesis ( $\mathbf{H}_{0}$ ): No difference, no association, or no effect.
Alternative Hypothesis $\left(\mathbf{H}_{\mathbf{1}}\right)$ : Investigators belief.
The Alternative Hypothesis is always set up to be what you want to build up evidence to prove.
7.1 Introduction to Hypothesis Testing

The hypothesis testing process consists of 5 Steps.
Step 1: Set up the hypotheses and determine the level of significance.

Step 2: Select the appropriate test statistic.

Step 3: Set-up the decision rule.

Step 4: Compute the test statistic.

Step 5: Conclusion.

### 7.5 Tests with Two Independent Samples, Continuous Outcome

The hypothesis testing process consists of 5 Steps.

$$
t=\frac{\left(\bar{X}_{1}-\bar{X}_{2}\right)}{S_{P} \sqrt{1 / n_{1}+1 / n_{2}}}
$$

Step 3: Set-up the decision rule.


Reject $\mathrm{H}_{0}$ if $t \geq t_{\alpha, d f}$
$\mathrm{H}_{0}: \mu_{1}=\mu_{2}$ vs. $\mathrm{H}_{1}: \mu_{1}<\mu_{2}$


Reject $\mathrm{H}_{0}$ if $t \leq t_{a, d f}$ Reject $\mathrm{H}_{0} \leq t_{a / 2, d f}$ or $t \geq t_{\alpha 2, d f}$

### 7.5 Tests with Two Independent Samples, Continuous

Outcome

OUTCOME | New drug | 15 | 195.9 | 28.7 |
| :--- | :--- | :--- | :--- |
| Placebo | 15 | 227.4 | 30.3 |

Example: Is the mean cholesterol of new drug < mean of placebo? Step 1: Null and Alternative Hypotheses.
$\mathrm{H}_{0}: \mu_{1} \geq \mu_{2}$ vs. $\mathrm{H}_{1}: \mu_{1}<\mu_{2}$
Step 2: Test Statistic.

$$
t=\frac{\left(\bar{x}_{1}-\bar{X}_{2}\right)}{S_{p} \sqrt{1 / n_{1}+1 / n_{2}}} \quad d f=n_{1}+n_{2}-2
$$

Step 3: Decision Rule. $\alpha=0.05, d f=15+15-2=28$
Reject $\mathrm{H}_{0}$ if $t \leq-1.701$.
Step 4: Compute test statistic.

$$
t=(195.9-227.4) /(29.5 \sqrt{1 / 15+1 / 15})=-2.92
$$



Step 5: Conclusion

$$
\begin{aligned}
\bar{X}_{i} & =\frac{1}{n} \sum X \quad i=1,2 \\
S_{P} & =\sqrt{\frac{(15-1)(28.7)^{2}+(15-1)(30.3)^{2}}{15+15-2}}=29.5
\end{aligned}
$$

Because $-2.92 \leq-1.701$, reject and conclude mean of drug less than placebo.

### 7.6 Tests with Matched Samples, Continuous Outcome

The hypothesis testing process consists of 5 Steps.

$$
t=\frac{\bar{X}_{d}}{s_{d} / \sqrt{n}}
$$

Step 3: Set-up the decision rule.
$\mathrm{H}_{0}: \mu_{d}=0$ vs. $\mathrm{H}_{1}: \mu_{d}>0$

$$
\mathrm{H}_{0}: \mu_{d}=0 \text { vs. } \mathrm{H}_{1}: \mu_{d}<0
$$

$$
\mathrm{H}_{0}: \mu_{d}=0 \text { vs. } \mathrm{H}_{1}: \mu_{d} \neq 0
$$



Reject $\mathrm{H}_{0}$ if $t \geq t_{\alpha, d f}$


Reject $\mathrm{H}_{0}$ if $t \leq t_{\alpha, d f}$ Reject $\mathrm{H}_{0} t \leq t_{\alpha / 2, d f}$ or $t \geq t_{\alpha / 2, d f}$

### 7.6 Tests with Matched Samples, Continuous Outcome

Example: Is there a difference in mean of new drug from baseline? Step 1: Null and Alternative Hypotheses.
$\mathrm{H}_{0}: \mu_{d}=0$ vs. $\mathrm{H}_{1}: \mu_{d} \neq 0$
Step 2: Test Statistic.

$$
t=\frac{\bar{X}_{d}}{s_{d} / \sqrt{n}} \quad d f=n-1
$$

Step 3: Decision Rule. $\alpha=0.05, d f=15-1=14$
Reject $\mathrm{H}_{0}$ if $t \leq-2.145$ or $t \geq 2.145$.


Step 4: Compute test statistic.

$$
t=-5.3 /(12.8 / \sqrt{15})=-1.60
$$

Step 5: Conclusion

$$
\begin{aligned}
& \bar{X}_{d}=\frac{1}{n} \sum d \\
& s_{d}^{2}=\frac{1}{n-1}\left[\sum d^{2}-\frac{1}{n}\left(\sum d\right)^{2}\right]
\end{aligned}
$$

Because $-2.145 \leq-1.60$, do not reject $\mathrm{H}_{0}$ and conclude no reduction.

### 7.7 Tests with Two Independent Samples, Dichotomous Outcome

We often have two populations that we are studying.

We may be interested in knowing if the proportion $p_{1}$ of population 1 is different (while accounting for random statistical variation) from the proportion $p_{1}$ of population 2.

When we have independent random sample from each population and the sample sizes are large.

### 7.7 Tests with Two Independent Samples, Dichotomous Outcome

The hypothesis testing process consists of 5 Steps.
Step 3: Set-up the decision rule. Assume $n$ "Large."
$\mathrm{H}_{0}: p_{1}=p_{2}$ vs. $\mathrm{H}_{1}: p_{1}>p_{2}$
$\mathrm{H}_{0}: p_{1}=p_{2}$ vs. $\mathrm{H}_{1}: p_{1}<p_{2}$
$\mathrm{H}_{0}: p_{1}=p_{2}$ vs. $\mathrm{H}_{1}: p_{1} \neq p_{2}$


Reject $\mathrm{H}_{0}$ if $z \geq z_{\alpha}$


Reject $\mathrm{H}_{0}$ if $z \leq z_{\alpha}$


Reject $\mathrm{H}_{0} z \leq z_{\alpha / 2}$ or $z \geq z_{\alpha / 2}$

### 7.8 Tests with More than Two Independent Samples, Continuous Outcome (ANOVA)

The hypothesis testing process consists of 5 Steps.
Step 1: Set up the hypotheses and determine the level of significance $\alpha$.

$$
\begin{aligned}
& \mathrm{H}_{0}: \mu_{1}=\mu_{2} \ldots=\mu_{k} \text { vs. } \mathrm{H}_{1}: \text { at least two } \mu \text { 's different } \\
& \text { reject for "large" disparities or } F=M S B / M S E .
\end{aligned}
$$

We will assume the means are equal and calculate two different variances. If the means are truly equal, the two different variances will be the same. If the means are noy equal, the two different variances will be different.

### 7.8 Tests with More than Two Independent Samples, Continuous Outcome (ANOVA)

The hypothesis testing process consists of 5 Steps.
Step 3: Set-up the decision rule.
$\mathrm{H}_{0}: \mu_{1}=\mu_{2} \ldots=\mu_{k}$ vs. $\mathrm{H}_{1}$ : at least two different


$$
M S B=\frac{\sum n_{j}\left(\bar{X}_{j}-\bar{X}\right)^{2}}{k-1}
$$

$$
\begin{aligned}
& d f_{1}=k-1 \\
& d f_{2}=N-k
\end{aligned}
$$

$$
M S E=\frac{\sum \sum n_{j}\left(X-\bar{X}_{j}\right)^{2}}{N-k}
$$

$$
F=\frac{M S B}{M S E}
$$

Reject $\mathrm{H}_{0}$ if $F \geq F_{o, d f, d f f}$.

### 7.8 Tests with More than Two Independent Samples,

 Continuous Outcome (ANOVA)Example: Find the value of $F_{0.05,3,16}$.
The (critical) value of $F$ that has an area of 0.05 larger than it when we have $d f_{1}=3$ (numerator) and $d f_{2}=16$ (denominator) degrees of freedom is 3.24 .

This is the value we use for a $95 \% \mathrm{HT}$ when $\alpha=0.05, n_{1}=6$, and $n_{2}=11$.
The book only has $\alpha=0.05$, but would have another page for each $\alpha$ value.

### 7.7 Tests with Two Independent Samples, Dichotomous Outcome

The hypothesis test on risk difference $\mathrm{H}_{0}: p_{1}=p_{2}$ vs. $\mathrm{H}_{1}: p_{1} \neq p_{2}$
$\mathrm{H}_{0}: R D=0$ vs. $\mathrm{H}_{1}: R D \neq 0$

$$
z=\frac{\hat{p}_{1}-\hat{p}_{2}}{\sqrt{\hat{p}(1-\hat{p})\left(\frac{1}{n_{1}}+\frac{1}{n_{2}}\right)}}
$$

Is equivalent to the two hypothesis tests

Risk Ratio $R R$
$\begin{array}{ll}\mathrm{H}_{0}: R R=1 \text { vs. } \mathrm{H}_{1}: R R \neq 1 & R R=\frac{\hat{p}_{1}}{\hat{p}_{2}} \\ \text { and }\end{array}$
Odds Ratio $O R$
$\mathrm{H}_{0}: O R=1$ vs. $\mathrm{H}_{1}: O R \neq 1$

$$
O R=\frac{\hat{p}_{1} /\left(1-\hat{p}_{1}\right)}{\hat{p}_{2} /\left(1-\hat{p}_{2}\right)}
$$

## Biostatistical Methods

### 7.8 Tests with More than Two Independent Samples, Continuous Outcome (ANOVA)

Example: Statistical difference in weight loss among 4 diets?
Step 1: Null and Alternative Hypotheses.
$\mathrm{H}_{0}: \mu_{1}=\mu_{2}=\mu_{3}=\mu_{4}$ vs. $\mathrm{H}_{1}$ : at least two different
Step 2: Test Statistic.

$$
F=M S B / M S E \quad d f_{1}=k-1 \quad d f_{2}=N-k
$$

Step 3: Decision Rule. $\alpha=0.05, d f_{1}=4-1=3, d f_{2}=20-4=16$
 Reject $\mathrm{H}_{0}$ if $F \geq 3.24$.
Step 4: Compute test statistic.

$$
\begin{gathered}
F \text { to be } \\
\text { calculated }
\end{gathered} \quad M S B=\frac{\sum n_{j}\left(\bar{X}_{j}-\bar{X}\right)^{2}}{k-1}=25.3
$$

$$
n_{1}=n_{2}=n_{3}=n_{4}=5
$$

$F=25.3 / 3.0=8.43$
Step 5: Conclusion

$$
M S E=\frac{\sum \sum n_{j}\left(X-\bar{X}_{j}\right)^{2}}{N-k}=3.0
$$

Because 8.43 > 3.24, reject $\mathrm{H}_{0}$ and conclude diets mean weight loss different.

### 7.10 Summary

TABLE 7-50 $\quad$ Summary of Key Formulas for Tests of Hypothesis

Outcome Variable, Number of Groups: Null Hypothesis
Continuous outcome, two independent samples: $\mathrm{H}_{0}: \mu_{1}=\mu_{2}$

Continuous outcome, two matched samples: $\mathrm{H}_{0}: \mu_{d}=0$

Continuous outcome, more than two independent samples: $\mathrm{H}_{0}: \mu_{1}=\mu_{2}=\ldots=\mu_{k}$

Dichotomous outcome, one sample: $\mathrm{H}_{0}: p=p_{0}$

Dichotomous outcome, two independent samples:
$H_{0}: p_{1}=p_{2}, R D=0, R R=1, O R=1$

Categorical or ordinal outcome, one sample:
$H_{0}: p_{1}=p_{10}, p_{2}=p_{20}, \ldots, p_{k}=p_{\kappa 0}$
Categorical or ordinal outcome, two or more independent samples:
$\mathrm{H}_{0}$ : Outcome and groups are independent

Test Statistic*

$$
z=\frac{\bar{X}_{1}-\bar{X}_{2}}{S_{p} \sqrt{1 / n_{1}+1 / n_{2}}}
$$

$$
z=\frac{\bar{X}_{d}-\mu_{d}}{s_{d} / \sqrt{n}}
$$

$$
F=\frac{\Sigma n_{i}\left(\bar{X}_{j}-\bar{X}\right)^{2} /(k-1)}{\left.\Sigma \Sigma(X-\bar{X})^{2}\right)^{2} /(N-K)}
$$

$$
z=\frac{\hat{p}-p_{0}}{\sqrt{\frac{p_{0}\left(1-p_{0}\right)}{n}}}
$$

$$
z=\frac{\hat{p}_{1}-\hat{p}_{2}}{\sqrt{\hat{p}(1-\hat{\rho})\left(1 / n_{1}+1 / n_{2}\right)}}
$$

$$
\chi^{2}=\Sigma \frac{(O-E)^{2}}{E}, d f=k-1
$$

$$
\chi^{2}=\Sigma \frac{(0-E)^{2}}{E}, d f=(r-1)(c-1)
$$

## D.B. Rowe

## Associations

We often are interested in the association between variables.

We often say correlation, with little thought to an actual definition.

We often say trend or linear relationship without defining how determine this relationship.

We define $y$ to be the response or dependent (on $x$ ) variable and $x$ to be the explanatory or independent variable. i.e. $y$ depends on $x$ (or several $x$ 's).

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## Biostatistical Methods

9.3 Introduction to Correlation and Regression Analysis-Correlation

Correlations $r$ are between -1 and $1,-1 \leq r \leq 1$.

$$
\begin{aligned}
& r=\frac{\operatorname{cov}(x, y)}{\sqrt{s_{x}^{2} s_{y}^{2}}} \\
& s_{x}^{2}=\frac{1}{n-1} \sum(X-\bar{X})^{2}=\frac{1}{n-1}\left[\sum X^{2}-\frac{1}{n}\left(\sum X\right)^{2}\right] \\
& s_{y}^{2}=\frac{1}{n-1} \sum(Y-\bar{Y})^{2}=\frac{1}{n-1}\left[\sum Y^{2}-\frac{1}{n}\left(\sum Y\right)^{2}\right]
\end{aligned}
$$

$$
\sum x^{2} \sum^{2}
$$

$$
\text { Variance of } X
$$

$$
\sum X Y
$$

$$
\operatorname{cov}(x, y)=\frac{1}{n-1} \sum(Y-\bar{Y})(X-\bar{X})=\frac{1}{n-1}\left[\sum X Y-\frac{1}{n}\left(\sum Y\right)\left(\sum X\right)\right]
$$

### 9.3 Introduction to Correlation and Regression Analysis-Correlation

## We are going to calculate the correlation in column format with sums.

| n | X | $\mathrm{X}^{2}$ | Y | Y | Y |
| :---: | :---: | :---: | :---: | :---: | :---: |
| 1 | 34.7 | 1204.1 | 1895 | 3591025.0 | 65756.5 |
| 2 | 36.0 | 1296.0 | 2030 | 4120900.0 | 73080.0 |
| 3 | 29.3 | 858.5 | 1440 | 2073600.0 | 42192.0 |
| 4 | 40.1 | 1608.0 | 2835 | 8037225.0 | 113683.5 |
| 5 | 35.7 | 1274.5 | 3090 | 9548100.0 | 110313.0 |
| 6 | 42.4 | 1797.8 | 3827 | 14645929.0 | 162264.8 |
| 7 | 40.3 | 1624.1 | 3260 | 10627600.0 | 131378.0 |
| 8 | 37.3 | 1391.3 | 2690 | 7236100.0 | 100337.0 |
| 9 | 40.9 | 1672.8 | 3285 | 10791225.0 | 134356.5 |
| 10 | 38.3 | 1466.9 | 2920 | 8526400.0 | 111836.0 |
| 11 | 38.5 | 1482.3 | 3430 | 11764900.0 | 132055.0 |
| 12 | 41.4 | 1714.0 | 3657 | 13373649.0 | 151399.8 |
| 13 | 39.7 | 1576.1 | 3685 | 13579225.0 | 146294.5 |
| 14 | 39.7 | 1576.1 | 3345 | 11189025.0 | 132796.5 |
| 15 | 41.1 | 1689.2 | 3260 | 10627600.0 | 133986.0 |
| 16 | 38.0 | 1444.0 | 2680 | 7182400.0 | 101840.0 |
| 17 | 38.7 | 1497.7 | 2005 | 4020025.0 | 77593.5 |
|  | 652.1 | 25173.2 | 49334.0 | 150934928.0 | 1921162.6 |

5 Sums

$$
\begin{aligned}
& \sum X=652.1 \quad \sum X^{2}=25173.2 \\
& \sum Y=49334.0 \quad \sum Y^{2}=150934928.0 \\
& \sum X Y=1921162.6 \\
& \operatorname{cov}(x, y)=1798.0 \\
& s_{x}^{2}=9.9638 \\
& r=\frac{1798.0}{\sqrt{(10.0)(485478.8)}} \\
& s_{y}^{2}=485478.8 \\
& r=0.82
\end{aligned}
$$

### 9.3 Introduction to Correlation and Regression Analysis-Regression

We can estimate the $y$-intercept and slope from what we have already computed for the correlation.

The slope is estimated as $b_{1}=r \frac{s_{y}}{s_{x}}$ and $b_{0}=\bar{Y}-b_{1} \bar{X}$.

$$
\begin{aligned}
s_{x}^{2} & =9.9638 \\
s_{y}^{2} & =485478.8 \\
r & =0.82
\end{aligned}
$$

Line goes through $(\bar{X}, \bar{Y})$. Note $b_{1}$ has same sign as $r$.

And hence we have determined our regression line.
$\hat{y}=b_{0}+b_{1} x$

### 9.3 Introduction to Correlation and Regression Analysis-Regression

Example: Continuing the small study ... to investigate the association between gestational age and birth weight.

$$
s_{x}^{2}=9.9638
$$

$$
s_{y}^{2}=485478.8
$$

$$
r=0.82
$$

$$
\hat{Y}=-4029.2+180.5 x
$$



$$
\begin{aligned}
& b_{1}=r \frac{s_{y}}{s_{x}} \\
& b_{1}=0.82 \frac{696.8}{3.2} \\
& b_{1}=180.5 \\
& b_{0}=\bar{Y}-b_{1} \bar{X} \\
& b_{0}=2902-(180.5)(38.4) \\
& b_{0}=-4029.2
\end{aligned}
$$

### 9.4 Multiple Linear Regression Analysis

Example: SBP and BMI, Age, Male Sex, and TFH.

A multiple regression analysis is run and coefficients estimated.

$$
S B P=68.15+0.58 B M I+0.65 A G E+0.94 M L S+6.44 T F H
$$

| Independent Variable | Regression Coefficien | n t | $p$-value | U. |
| :---: | :---: | :---: | :---: | :---: |
| Intercept | $b_{0}=68.15$ | $t_{0}=26.33$ | ${ }^{0.0001}=p_{0}$ | The $t$ statistic is for $\mathrm{H}_{0}: \beta_{j}=0, \mathrm{H}_{1}: \beta_{j} \neq 0$. |
| BMI | $b_{1}=0.58$ | $t_{1}=10.30$ | ${ }^{0.0001}=p_{1}$ | The $p$-value is the probability of getting |
| Age | $b_{2}=0.65$ | $t_{2}=20.22$ | $0.0001=p_{2}$ |  |
| Male sex | $b_{3}=0.94$ | $t_{3}=1.58$ | ${ }^{0.1133}=p_{3}$ | $t_{j}=\frac{b_{j}-0}{}$ |
| Treatment for hypertension | $b_{4}=6.44$ | $t_{4}=9.74$ | $0.0001=p_{4}$ | larger in abs if it were truly $0 .{ }_{d j f=n-p-1}^{j} \sqrt{\operatorname{var}\left(b_{j}\right)}$ |

## Biostatistical Methods

### 9.5 Multiple Logistic Regression Analysis

## Using R,



| Hours $(\boldsymbol{x}) \mathbf{A}(\boldsymbol{y})$ |  |
| :---: | :---: |
| 6 | 0 |
| 8 | 0 |
| 10 | 0 |
| 12 | 0 |
| 14 | 0 |
| 16 | 1 |
| 18 | 0 |
| 20 | 0 |
| 22 | 0 |
| 24 | 0 |
| 26 | 1 |
| 28 | 0 |
| 30 | 0 |
| 32 | 1 |
| 34 | 1 |
| 36 | 1 |
| 38 | 1 |
| 40 | 1 |

\# grade data

```
xx <- c(6, 8,10,12,14,16,18,20,22,24,26,28,30,32,34,36,38,40)
yy <- c(0, 0, 0, 0, 0, 1, 0, 0, 0, 0,1, 0, 0, 1, 1, 1, 1, 1)
#scatter plot plot(x = xx,y = yy,xlab = "Hours",ylab = "Grade",
xlim = c(0,45),ylim = c(0,1),col = "darkred",
    cex = 1.5, main = "Hours vs. Grade", pch = 16)
logistic_model <- glm(yy }\mp@subsup{}{~}{xx}\mathrm{ , family=binomial(link="logit"))
summary(logistic_model)
b0 <- logistic_model$coefficients[1]
b1 <- logistic_model$coefficients[2]
phat <- round(1/(1+exp(-b0-b1*xx)), digits = 4)
O <- round(phat/(1-phat) , digits = 4)
df <- data.frame(xx,yy,phat,O)
df
#scatter plot with curve
xhat <- (1:4500)/100
yhat <- 1/(1+exp(-b0-b1*xhat))
plot(x = xx,y = yy,xlab = "Hours",ylab = "Grade",
    xlim = c(0,45),ylim = c(0,1),col = "darkred",
    cex = 1.5, main = "Hours vs. Grade", pch = 16)
points(xhat,yhat,cex = .1,col = "blue")
```


### 9.5 Multiple Logistic Regression Analysis

Once we have $\hat{\beta}_{0}$ and $\hat{\beta}_{1}$, insert them back into
$\hat{p}_{i}=\frac{1}{1+e^{-\hat{\beta}_{0}-\hat{\beta}_{1} x_{i}}}$ for estimated probabilities
and also for odds $\hat{o}_{i}=\frac{\hat{p}_{i}}{1-\hat{p}_{i}}=e^{\hat{\beta}_{0}+\hat{\beta}_{i_{i}} x_{i}}$
and for odds ratio $\hat{O} R=e^{\hat{\beta}_{0}+\hat{\beta}_{x_{b}} x_{b}} / e^{\hat{\beta}_{0}+\hat{\beta}_{1} x_{a}}=e^{\hat{\beta}_{1} \swarrow}, \Delta=x_{b}-x_{a}$.

$$
\begin{aligned}
& \hat{\beta}_{0}=-5.66 \\
& \hat{\beta}_{1}=0.21
\end{aligned}
$$

$$
\hat{O} R=e^{(0.21)(2)}=1.5220
$$

| Hours $(\boldsymbol{x})$ | $\mathbf{A}(\boldsymbol{y})$ | $\hat{\boldsymbol{p}}$ | $\hat{\boldsymbol{o}}$ |
| :---: | :---: | :---: | :---: |
| 6 | 0 | 0.0120 | 0.0122 |
| 8 | 0 | 0.0181 | 0.0184 |
| 10 | 0 | 0.0272 | 0.0279 |
| 12 | 0 | 0.0406 | 0.0423 |
| 14 | 0 | 0.0603 | 0.0641 |
| 16 | 1 | 0.0886 | 0.0972 |
| 18 | 0 | 0.1284 | 0.1473 |
| 20 | 0 | 0.1824 | 0.2232 |
| 22 | 0 | 0.2527 | 0.3381 |
| 24 | 0 | 0.3388 | 0.5124 |
| 26 | 1 | 0.4371 | 0.7764 |
| 28 | 0 | 0.5405 | 1.1764 |
| 30 | 0 | 0.6406 | 1.7824 |
| 32 | 1 | 0.7298 | 2.7008 |
| 34 | 1 | 0.8036 | 4.0923 |
| 36 | 1 | 0.8611 | 6.2008 |
| 38 | 1 | 0.9038 | 9.3957 |
| 40 | 1 | 0.9344 | 14.2365 |

Study 2 more hours and $O R$ increases by 1.5 .

### 9.6 Summary

## Correlation

$$
\begin{aligned}
& \operatorname{cov}(x, y)=\frac{1}{n-1}\left[\sum X Y-\frac{1}{n}\left(\sum Y\right)\left(\sum X\right)\right] \\
& s_{x}^{2}=\frac{1}{n-1}\left[\sum X^{2}-\frac{1}{n}\left(\sum X\right)^{2}\right] \\
& s_{y}^{2}=\frac{1}{n-1}\left[\sum Y^{2}-\frac{1}{n}\left(\sum Y\right)^{2}\right] \\
& r=\frac{\operatorname{cov}(x, y)}{\sqrt{s_{x}^{2} s_{y}^{2}}}
\end{aligned}
$$

## Linear Regression

$$
\begin{array}{ll}
b_{1}=r \frac{s_{y}}{s_{x}} \\
b_{0}=\bar{Y}-b_{1} \bar{X} & \hat{y}=b_{0}+b_{1} x
\end{array}
$$

## Logistic Regression

$$
\begin{aligned}
& \hat{p}=\frac{1}{1+e^{-b_{0}-b_{1} x_{1} \ldots . . b_{p} x_{p}}} \\
& \ln \left(\frac{\hat{p}}{1-\hat{p}}\right)=b_{0}+b_{1} x_{1}+\ldots+b_{p} x_{p} \\
& \hat{O} R=e^{\hat{\beta}_{1} \Delta_{1}+\ldots+\hat{\beta}_{p} \Delta_{p}} \quad \text { logsitic pobababily }
\end{aligned}
$$

### 10.1 Introduction to Nonparametric Testing - Sign Test

Example: Mark is training for 10K. $n=20$ daily runs. Step 1: $\mathrm{H}_{0}: M D=4$ vs. $\mathrm{H}_{1}: M D>4, \alpha=0.05$ Step 2: Test Statistic. $x=$ the number of + 's.

Binomial Distribution, $n=20, p=0.5$

$X_{0.05}=15 \quad$ (or $n-5=15$ )

| $\mathbf{x}$ | $\mathbf{P}(\mathbf{X}=\mathbf{x})$ | CumSum | CumSumR |
| :---: | :---: | :---: | :---: |
| 0 | 0.000 | 0.000 | 1.000 |
| 1 | 0.000 | 0.000 | 1.000 |
| 2 | 0.000 | 0.000 | 1.000 |
| 3 | 0.001 | 0.001 | 1.000 |
| 4 | 0.005 | 0.006 | 0.999 |
| 5 | 0.015 | 0.021 | 0.994 |
| 6 | 0.037 | 0.058 | 0.979 |
| 7 | 0.074 | 0.132 | 0.942 |
| 8 | 0.120 | 0.252 | 0.868 |
| 9 | 0.160 | 0.412 | 0.748 |
| 10 | 0.176 | 0.588 | 0.588 |
| 11 | 0.160 | 0.748 | 0.412 |
| 12 | 0.120 | 0.868 | 0.252 |
| 13 | 0.074 | 0.942 | 0.132 |
| 14 | 0.037 | 0.979 | 0.058 |
| 15 | 0.015 | 0.994 | 0.021 |
| 16 | 0.005 | 0.999 | 0.006 |
| 17 | 0.001 | 1.000 | 0.001 |
| 18 | 0.000 | 1.000 | 0.000 |
| 19 | 0.000 | 1.000 | 0.000 |
| 20 | 0.000 | 1.000 | 0.000 |
|  |  |  |  |


| Two-Sided Test $\alpha$ | . 10 | . 05 | . 02 | . 01 | ata |
| :---: | :---: | :---: | :---: | :---: | :---: |
| One-Sided Test $\alpha$ | . 05 | . 025 | . 01 | . 005 | 5 |
| $n$ |  |  |  |  | 3 |
| 1 |  |  |  |  | 5 |
| 2 |  |  |  |  |  |
| 3 |  |  |  |  | 3 |
| 4 |  |  |  |  | 4 |
| 5 | 0 |  |  |  |  |
| 6 | 0 | 0 |  |  | 4 |
| 7 | 0 | 0 | 0 |  | 6 |
| 8 | 1 | 0 | 0 | 0 | 6 |
| 9 | 1 | 1 | 0 | 0 |  |
| 10 | 1 | 1 | 0 | 0 | 6 |
| 11 | 2 | 1 | 1 | 0 | 4 |
| 12 | 2 | 2 | 1 | 1 |  |
| 13 | 3 | 2 | 1 | 1 | 6 |
| 14 | 3 | 2 | 2 | 1 | 5 |
| 15 | 3 | 3 | 2 | 2 | 5 |
| 16 | 4 | 3 | 2 | 2 | 5 |
| 17 | 4 | 4 | 3 | 2 | 5 |
| 18 | 5 | 4 | 3 | 3 | 4 |
| 19 | 5 | 4 | 4 | 3 |  |
| $\longrightarrow 20$ | 5 | 5 | 4 | 3 | 5 |
| 21 | 6 | 5 | 4 | 4 | 5 |
| 22 | 6 | 5 | 5 | 4 |  |
| 23 | 7 | 6 | 5 | 4 | 5 |
| 24 | 7 | 6 | 5 | 5 | 5 |
| 25 | 7 | 7 | 6 | 5 | 6 |

### 10.1 Introduction to Nonparametric Testing - Sign Test

The hypothesis testing process consists of 5 Steps.
Step 3: Set-up the decision rule.


### 7.1 Introduction to Hypothesis Testing

으른ㄴ
The hypothesis testing process consists of 5 Steps.
Step 3: Set-up the decision rule.


Reject $\mathrm{H}_{0}$ if $z \geq z_{\alpha}$

$$
\mathrm{H}_{0}: \mu=\mu_{0} \text { vs. } \mathrm{H}_{1}: \mu<\mu_{0}
$$



Reject $\mathrm{H}_{0}$ if $z \leq z_{\alpha}$

$$
\mathrm{H}_{0}: \mu=\mu_{0} \text { vs. } \mathrm{H}_{1}: \mu \neq \mu_{0}
$$



Reject $\mathrm{H}_{0} z \leq z_{\alpha / 2}$ or $z \geq z_{\alpha / 2}$

### 10.1 Introduction to Nonparametric Testing - Sign Test

Step 4: Compute the test statistic.

$$
\begin{aligned}
& x=14 \\
& x=\left(\text { the number of observations }>M D_{0}=4\right)
\end{aligned}
$$

Step 5: Because $x=14<x_{\alpha}=15$, do not reject $H_{0}$.

| X | $P(X=x)$ | CumSum | CumSumR | Two-Sided Test $\boldsymbol{\alpha}$ | . 10 | . 05 | . 02 | . 01 |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 5 | 0.015 | 0.021 | 0.994 | One-Sided Test $\boldsymbol{\alpha}$ | . 05 | . 025 | . 01 | . 005 |
| 6 | 0.037 | 0.058 | 0.979 | 19 | 5 | 4 | 4 | 3 |
| 14 | 0.037 | 0.979 | 0.058 | 20 | 5 | (5) | 4 | 3 |
| 15 | 0.015 | 0.994 | 0.021 | 21 | 6 | 5 | 4 | 4 |
| See also Table 6 |  |  |  | Table 6 |  |  |  |  |

Note: See also Table 6


If we used normal, we would reject $\mathrm{H}_{0}, t=4.07>t_{0.05,19}=2.093$.

$$
t=\frac{\bar{X}-\mu_{0}}{s / \sqrt{n}} \quad d f=n-1 \quad \bar{X}=4.8500 \quad s=0.9333
$$

| data | sorted | sign |
| :---: | :---: | :---: |
| 5 | 3 | -1 |
| 3 | 3 | -1 |
| 5 | 4 | 0 |
| 3 | 4 | 0 |
| 4 | 4 | 0 |
| 4 | 4 | 0 |
| 6 | 5 | +1 |
| 6 | 5 | +1 |
| 6 | 5 | +1 |
| 4 | 5 | +1 |
| 6 | 5 | +1 |
| 5 | 5 | +1 |
| 5 | 5 | +1 |
| 5 | 5 | +1 |
| 4 | 5 | +1 |
| 5 | 6 | +1 |
| 5 | 6 | +1 |
| 5 | 6 | +1 |
| 5 | 6 | +1 |
| 6 | 6 | +1 |

### 10.2 Tests with Two Independent Samples - Mann-Whitney U Test

Example: Phase II clinical trial, $n=10$ children. Difference in episodes?
Step 1: Set up the hypotheses and determine $\alpha$.

$$
\mathrm{H}_{0}: M D_{1}=M D_{2} \text { vs. } \mathrm{H}_{1}: M D_{1} \neq M D_{2}, \quad \alpha=0.05
$$

Group 1 Group 2

| Placebo | NewDrug |
| :---: | :---: |
| 7 | 3 |
| 5 | 6 |
| 6 | 4 |
| 4 | 2 |
| 12 | 10 |
| $n_{1}=5$ | $n_{2}=5$ |

Step 2: Select the appropriate test statistic.
Pool data and assign ranks. Test statistic based on ranks

| Placebo | New Drug | Placebo | New Drug | Placebo | New Drug | Ranks |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  |  |  |  | Placebo | New Drug |  |
| 7 | 3 |  | 1 |  | 1 |  | 1 |  |
| 5 | 6 |  | 2 |  | 2 |  | 2 |  |
| 6 | 4 |  | 3 |  | 3 |  | 3 |  |
| 4 | 2 | 4 | 4 | 4.5 | 4.5 | 4.5 | 4.5 | 7 |
| 12 | 1 | 5 |  | 6 |  | 6 |  |  |
|  |  | 6 | 6 | 7.5 | 7.5 | 7.5 | 7.5 |  |
|  |  | 7 |  | 9 |  | 9 |  | $R_{2}=18$ |
|  |  | 12 |  | 10 |  | 10 |  |  |

### 10.2 Tests with Two Independent Samples - Mann-Whitney U Test

Step 2: Select the appropriate test statistic.
The test statistic is a single (decision) number summarizing information.

$$
\begin{aligned}
& U_{1}=n_{1} n_{2}+\frac{n_{1}\left(n_{1}+1\right)}{2}-R_{1}=(5)(5)+\frac{5(5+1)}{2}-37=3 \\
& U_{2}=n_{1} n_{2}+\frac{n_{2}\left(n_{2}+1\right)}{2}-R_{2}=(5)(5)+\frac{5(5+1)}{2}-18=22 \\
& U=\min \left(U_{1}, U_{2}\right)=\min (3,22)=3
\end{aligned}
$$

| Rankings |  |
| :---: | :---: |
| Group 1 Group 2 | Group 1 Group |
| ${ }^{1}$ | 1 |
| 2 3 | 23 |
| 4 | 4 |
| - 5 | 5 |
| 6 | 6 |
| 7 |  |
| 8 | 8 |
| 10 | 10 |
| $U=0$ | $U=25$ |


complete
Reject $\mathrm{H}_{0}$ for small $U$.


### 10.2 Tests with Two Independent Samples - Mann-Whitney U Test

Step 3: Set-up the decision rule.
$n_{1}=5, n_{2}=5$
If we did Two Sided Test
Reject $\mathrm{H}_{0}$ if $U \leq U_{0.05, n_{1}, n_{2}}$.
Step 4: Compute test statistic.
Already done, $U=3$.

Step 5: Conclusion.
Do not reject $\mathrm{H}_{0}$ because
$U=3>U_{0.05,5,5}=2$. Interpret.

| Two-Sided Test $\alpha=0.05$ |  |  |  |  |  | $n_{1}$ |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  |  |  | $\downarrow$ |  |  |  |  |  |  |  |  |  |  |  |  |  |  |  |
| $n_{2}$ | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 | 12 | 13 | 14 | 15 | 16 | 17 | 18 | 19 | 20 |
| 2 |  |  |  |  |  |  |  | 0 | 0 | 0 | 0 | 1 | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 |
| 3 |  |  |  |  | 0 | 1 | 1 | 2 | 2 | 3 | 3 | 4 | 4 | 5 | 5 | 6 | 6 | 7 | 7 | 8 |
| 4 |  |  |  | 0 | 1 | 2 | 3 | 4 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 | 11 | 12 | 13 | 13 |
| 5 |  |  | 0 | 1 | (2) | 3 | 5 | 6 | 7 | 8 | 9 | 11 | 12 | 13 | 14 | 15 | 17 | 18 | 19 | 20 |
| 6 |  |  | 1 | 2 | 3 | 5 | 6 | 8 | 10 | 11 | 13 | 14 | 16 | 17 | 19 | 21 | 22 | 24 | 25 | 27 |
| 7 |  |  | 1 | 3 | 5 | 6 | 8 | 10 | 12 | 14 | 16 | 18 | 20 | 22 | 24 | 26 | 28 | 30 | 32 | 34 |
| 8 |  | 0 | 2 | 4 | 6 | 8 | 10 | 13 | 15 | 17 | 19 | 22 | 24 | 26 | 29 | 31 | 34 | 36 | 38 | 41 |
| 9 |  | 0 | 2 | 4 | 7 | 10 | 12 | 15 | 17 | 20 | 23 | 26 | 28 | 31 | 34 | 37 | 39 | 42 | 45 | 48 |
| 10 |  | 0 | 3 | 5 | 8 | 11 | 14 | 17 | 20 | 23 | 26 | 29 | 33 | 36 | 39 | 42 | 45 | 48 | 52 | 55 |
| 11 |  | 0 | 3 | 6 | 9 | 13 | 16 | 19 | 23 | 26 | 30 | 33 | 37 | 40 | 44 | 47 | 51 | 55 | 58 | 62 |
| 12 |  | 1 | 4 | 7 | 11 | 14 | 18 | 22 | 26 | 29 | 33 | 37 | 41 | 45 | 49 | 53 | 57 | 61 | 65 | 69 |
| 13 |  | 1 | 4 | 8 | 12 | 16 | 20 | 24 | 28 | 33 | 37 | 41 | 45 | 50 | 54 | 59 | 63 | 67 | 72 | 76 |
| 14 |  | 1 | 5 | 9 | 13 | 17 | 22 | 26 | 31 | 36 | 40 | 45 | 50 | 55 | 59 | 64 | 67 | 74 | 78 | 83 |
| 15 |  | 1 | 5 | 10 | 14 | 19 | 24 | 29 | 34 | 39 | 44 | 49 | 54 | 59 | 64 | 70 | 75 | 80 | 85 | 90 |
| 16 |  | 1 | 6 | 11 | 15 | 21 | 26 | 31 | 37 | 42 | 47 | 53 | 59 | 64 | 70 | 75 | 81 | 86 | 92 | 98 |
| 17 |  | 2 | 6 | 11 | 17 | 22 | 28 | 34 | 39 | 45 | 51 | 57 | 63 | 67 | 75 | 81 | 87 | 93 | 99 | 105 |
| 18 |  | 2 | 7 | 12 | 18 | 24 | 30 | 36 | 42 | 48 | 55 | 61 | 67 | 74 | 80 | 86 | 93 | 99 | 106 | 112 |
| 19 |  | 2 | 7 | 13 | 19 | 25 | 32 | 38 | 45 | 52 | 58 | 65 | 72 | 78 | 85 | 92 | 99 | 106 | 113 | 119 |
| 20 |  | 2 | 8 | 13 | 20 | 27 | 34 | 41 | 48 | 55 | 62 | 69 | 76 | 83 | 90 | 98 | 105 | 112 | 119 | 127 |

### 10.3 Tests with Matched Samples - Wilcoxon Signed Rank Test

An alternative for the Sign Test for matched samples median difference is the Wilcoxon Signed Rank test.

## Step 1:

$\mathrm{H}_{0}: \delta \leq 0$ vs. $\mathrm{H}_{1}: \delta>0$
$\mathrm{H}_{0}$ : The median difference is zero $\left(\mathrm{H}_{0}: \delta=0\right)$
$\mathrm{H}_{1}$ : The median difference is positive $\left(\mathrm{H}_{1}: \delta>0\right)$
We will calculate a test statistic $W$ the smaller of W+ and W-.
$W+=$ sum of positive ranks
$W_{-}=$sum of negative ranks
$\longrightarrow \quad W=\min \left(W+, W_{-}\right)$
If the median difference of the matched pairs is zero, then the sum of the positive ranks should be the same as the sum of the negative ranks.

### 10.3 Tests with Matched Samples - Wilcoxon Signed Rank Test

An alternative for the Sign Test for matched samples median difference is the Wilcoxon Signed Rank test.

Step 1:
$\mathrm{H}_{0}: \delta \leq 0$ vs. $\mathrm{H}_{1}: \delta>0$
$\delta$ is population version of $d$.
Step 2: Select the test statistic.
$W_{+}=$sum of positive ranks $=32$
$W_{-}=$sum of negative ranks $=4$
$W=\min \left(W_{1}, W_{2}\right)=\min (4,32)=4$
Reject $\mathrm{H}_{0}$ for small $W$.

| $\mathbf{b}$ | $\mathbf{a}$ | d | sorted sign rankS SgnRnk |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 85 | 75 | 10 | -10 | -1 | 3 | -3 |
| 70 | 50 | 20 | -5 | -1 | 1 | -1 |
| 40 | 50 | -10 | 10 | +1 | 3 | 3 |
| 65 | 40 | 25 | 10 | +1 | 3 | 3 |
| 80 | 20 | 60 | 15 | +1 | 5 | 5 |
| 75 | 65 | 10 | 20 | +1 | 6 | 6 |
| 55 | 40 | 15 | 25 | +1 | 7 | 7 |
| 20 | 25 | -5 | 60 | +1 | 8 | 8 |

IF Signed Ranks

| SgnRnk | SgnRnk | SgnRnk | SgnRnk |
| :---: | :---: | :---: | :---: |
| 1 | -4 | -7 | -8 |
| 2 | -3 | -5 | -7 |
| 3 | -2 | -3 | -6 |
| 4 | -1 | -1 | -5 |
| 5 | 5 | 2 | 2 |
| 6 | 6 | 4 | 4 |
| 7 | 7 | 6 | 6 |
| 8 | 8 | 8 | 8 |

Possible Examples
10.3 Tests with Matched Samples - Wilcoxon Signed Rank Test

Step 3: Set-up the decision rule.
$n=8, \alpha=0.05$
If we did One Sided Test
Reject $\mathrm{H}_{0}$ if $W \leq W_{\alpha, n}$.
Step 4: Compute test statistic.
Already done, $W=4$.

Step 5: Conclusion.
Reject $\mathrm{H}_{0}$ because
$W=4 \leq W_{0.05,8}=6$. Interpret.
10.4 Tests with More than Two Independent Samples - Kruskal-Wallis Test

The hypothesis testing process consists of 5 Steps.
Step 1: Set up the hypotheses and determine the level of significance $\alpha$.
$\mathrm{H}_{0}: M D_{1}=M D_{2} \ldots=M D_{k}$ vs. $\mathrm{H}_{1}$ : at least two $M D$ 's different reject for "large" disparities $H$.

We will assume the medians are equal and see how different from equal.
7.8 Tests with More than Two Independent Samples, Continuous Outcome (ANOVA)


The hypothesis testing process consists of 5 Steps.
Step 1: Set up the hypotheses and determine the level of significance $\alpha$.

$$
\begin{array}{ll}
\mathrm{H}_{0}: \mu_{1}=\mu_{2} \ldots=\mu_{k} \quad \text { vs. } \mathrm{H}_{1}: \text { at least two } \mu \text { 's different } \\
\text { reject for "large" disparities } F=M S B / M S E .
\end{array}
$$

We will assume the means are equal and calculate two different variances. If the means are truly equal, the two different variances will be the same. If the means are noy equal, the two different variances will be different.

### 10.4 Tests with More than Two Independent Samples - Kruskal-Wallis

Example: Statistical difference in albumin for 3 diets?
Step 1: Null and Alternative Hypotheses.
$\mathrm{H}_{0}: M D_{1}=M D_{2}=M D_{3}$ vs. $\mathrm{H}_{1}$ : at least two different

$$
H=\left(\frac{12}{N(N+1)} \sum_{j=1}^{k} \frac{R_{j}^{2}}{n_{j}}\right)-3(N+1)
$$

Step 2: Test Statistic.

$$
H=\left(\frac{12}{N(N+1)} \sum_{j=1}^{k} \frac{R_{j}^{2}}{n_{j}}\right)-3(N+1)
$$

Step 3: Decision Rule. $\alpha=0.05, n_{1}=3, n_{2}=5, n_{3}=4$
Reject $\mathrm{H}_{0}$ if $H \geq 5.656$.
Step 4: Compute test statistic.

Table 8

| Three groups |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: |
| $n_{1}$ | $n_{2}$ | $n_{3}$ |  | $\alpha=.05$ | $\alpha=.01$ |
| 5 | 4 | 3 |  | 5.656 | 7.445 |
|  |  |  |  |  |  |

Sample size order doesn't matter.

$$
H=7.52
$$

Step 5: Conclusion
Reject $\mathrm{H}_{0}$ because $7.52>5.656$, and conclude difference in median albumin.
10.5 Summary

Sign Test (one sample)
$x=$ number of observations $>M D_{0}$

## Mann-Whitney U Test

$U_{1}=n_{1} n_{2}+\frac{n_{1}\left(n_{1}+1\right)}{2}-R_{1}$
$U_{2}=n_{1} n_{2}+\frac{n_{2}\left(n_{2}+1\right)}{2}-R_{2}$
$U=\min \left(U_{1}, U_{2}\right)$
Sign Test (two sample)
$x=$ number of observations $>0$

## Wilcoxon Signed Rank Test

$W=\min (W+, W-)$
$W+=$ sum of positive ranks
$W-=$ sum of negative ranks

## Kruskal-Wallis Test

$$
H=\left(\frac{12}{N(N+1)} \sum_{j=1}^{k} \frac{R_{j}^{2}}{n_{j}}\right)-3(N+1)
$$

### 11.1 Introduction to Survival Data

Survival analysis measures two pieces of information

1) Whether the event occurred, $1=y e s, 0=$ no
2) Last follow-up time, from enrollment.

The survival function is the probability a person survives past a time $t$.

$t=0.0$ : survival probability=1.00
$t=2.0$ : survival probability= 0.83
$t=8.5$ : survival probability=0.50 (Median)
$t=10.0$ : survival probability $=0.47$

### 11.2 Estimating the Survival Function

There are several parametric and nonparametric ways to estimate survival Let's examine nonparametric step survival curves. Time on $x$ axis and survival (percentage) at risk on $y$ axis.


$$
\begin{aligned}
& t=0.0 \text { : survival probability }=1.00 \\
& t=2.0 \text { : survival probability }=0.90 \\
& t=9.0 \text { : survival probability }=0.50 \text { (Median) } \\
& t=10.0 \text { : survival probability }=0.45
\end{aligned}
$$

### 11.6 Summary

The survival function is the probability a person survives past a time $t$.

## Actuarial Life Table

$N_{t}=\underset{\text { \# event free }}{\text { (Numberat risk) }}$ during interyal $t$
$D_{t}=\#$ who die in interval $t$
$C_{t}=$ \# censored in interval $t$
$N_{t^{*}}=$ avg. \#at risk in intervał $t, N_{t^{*}}=N_{t}-C_{t} / 2$
$q_{t}=$ prop. die in interval $t, q_{t}=D_{t} / T$
$p_{t}=$ prop. survive in interval $t, p_{t}=1-q_{t}$
$S_{t}=$ prop. survive past interval $t$ Can plot $S_{t}$ vs. $t$.

Kaplan-Meier Life Table
$S_{t+1}=S, \begin{aligned} & \frac{N_{t}-D_{t}}{N_{t}} \\ & S E\left(S_{t}\right)=S_{t} \sqrt{\frac{D_{t}}{N_{t}\left(N_{t}-D_{t}\right)}}\end{aligned}$

## Chi-Square Test

## Cox Proportional Hazards Model

$$
h(t)=h_{0}(t) \exp \left(b_{1} x_{1}+b_{2} x_{2}+\ldots+b_{p} x_{p}\right)
$$

## Questions?

Bring pencil/eraser, calculator, caffeinated beverage. Will hand out exam and formula sheet/tables.

