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 (not A)?
- Two Competing Hypotheses. A and B.
- Null Hypothesis (H_{n}): No difference, no association, or no effect.
- Alternative Hypothesis (H_1) : 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.

Step 3: Set-up the decision rule.

 $H_0: \mu_1 = \mu_2 \text{ VS. } H_1: \mu_1 > \mu_2 \qquad H_0: \mu_1 = \mu_2 \text{ VS. } H_1: \mu_1 < \mu_2$ 0.4 rejection rejection rejection 0.35 0.35 0.35 region region region 0.3 0.3 0.3 0.25 0.25 0.25 0.2 0.2 0.2 $_{0.15}$ $\alpha/2$ level α level α level 0.15 0.15 (0.05) (0.05) (0.025)0.1 0.1 0.1 0.05 0.05 0.05 0 **-**4 -2 $-t_{\alpha}$ 0 └ -4 3 2 З -3 -2 -1 0 1 2 -3 -1 0 1 4 $\bar{t}_{\alpha/2}$ (1.645)(-1.645) (-1.960)Reject H_0 if $t \ge t_{\alpha, df}$ Reject H₀ if $t \le t_{\alpha,df}$ Reject H₀ $\le t_{\alpha/2,df}$ or $t \ge t_{\alpha/2,df}$

Table 2 in book







7.5 Tests with Two Independent Samples, Continuous Outcome

Example: Is the mean cholesterol of new drug < mean of placebo?

Step 1: Null and Alternative Hypotheses.

$$H_0: \mu_1 \ge \mu_2 \text{ vs. } H_1: \mu_1 < \mu_2$$

Step 2: Test Statistic. $(\overline{X}_1 - \overline{X}_2)$

$$t = \frac{1}{S_P \sqrt{1 / n_1 + 1 / n_2}} \qquad df = n_1 + n_2 - 2$$

Step 3: Decision Rule. α =0.05 , *df*=15+15-2=28

Reject H₀ 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

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



$$\overline{X}_i = \frac{1}{n} \sum X$$
$$S_P = \sqrt{\frac{(15-1)(2)}{2}}$$



$\frac{28.7)^2 + (15-1)(30.3)^2}{15+15-2} = 29.5$

i=1.2

	Sample Size	Mean	Standard Deviation
w drug	15	195.9	28.7
acebo	15	227.4	30.3

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7.6 Tests with Matched Samples, Continuous Outcome

The hypothesis testing process consists of 5 Steps.

Step 3: Set-up the decision rule.



Table 2 in book



 $t = \frac{X_d}{S_d / \sqrt{n}}$



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. rejection 0.35 $H_0: \mu_d = 0$ vs. $H_1: \mu_d \neq 0$ region 0.25 Step 2: Test Statistic. $\alpha/2$ level 0.15 $t = \frac{X_d}{S_d \sqrt{n}} \qquad df = n-1$ (0.025)0.1

Step 3: Decision Rule. α =0.05 , df=15-1=14

Reject H₀ if $t \le -2.145$ or $t \ge 2.145$.

Step 4: Compute test statistic.

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

Step 5: Conclusion

Because $-2.145 \le -1.60$, do not reject H₀ and conclude no reduction.





0.05

 $s_{d}^{2} = \frac{1}{n-1} \left[\sum d^{2} - \frac{1}{n} \left(\sum d \right)^{2} \right]$

 $\overline{X}_d = \frac{1}{n} \sum d$

-3

 $\bar{t}^2_{\alpha,df/2}$

Number	Baseline	6 Weeks	Difference
1	215	205	10
2	190	156	34
3	230	190	40
4	220	180	40
5	214	201	13
6	240	227	13
7	210	197	13
8	193	173	20
9	210	204	6
10	230	217	13
11	180	142	38
12	260	262	-2
13	210	207	3
14	190	184	6
15	200	193	7
		$\overline{\overline{X}_d} = \cdot$	-5.30

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." $H_0: p_1 = p_2 \text{ VS. } H_1: p_1 > p_2 \qquad H_0: p_1 = p_2 \text{ VS. } H_1: p_1 < p_2$ 0.4 rejection rejection rejection 0.35 0.35 0.35 region region region 0.3 0.3 0.3 0.25 0.25 0.25 0.2 0.2 0.2 $_{0.15} \alpha/2$ level α level α level 0.15 0.15 (0.05)(0.05)(0.025)0.1 0.1 0.1 0.05 0.05 0.05 0 └--4 3 -3 -2 -1 0 1 2 -4 -3 -2 -1 0 1 2 З 4 -3 -*Z*α $Z_{\alpha/2}$ (1.645)(-1.645)(-1.960)Reject H_0 if $z \ge z_\alpha$ Reject H₀ if $z \leq z_a$ Table 1 in book







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 α .

 $H_0: \mu_1 = \mu_2 \dots = \mu_k$ vs. $H_1:$ at least two μ 's different reject for "large" disparities or F=MSB/MSE.

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.

 $H_0: \mu_1 = \mu_2 \dots = \mu_k vs. H_1:$ at least two different



See Chapter 07b worksheet for details

Reject H₀ if $F \ge F_{\alpha, df_1, df_2}$.

Table 4 in book



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

Example: Find the value of $F_{0.05,3,16}$. $\alpha \int df_1 = n_1 - 1 df_2 = n_2 - 1$

The (critical) value of F that has an area of 0.05 larger than it when we have $df_1=3$ (numerator) and $df_2 = 16$ (denominator) degrees of freedom is 3.24.

	$P(F_{dr_1, dr_2} > F) = 0.05,$ e.g., $P(F_{3,20} > 3.10) = 0.05$														
	$\bigvee df_1$														
	df 2	1	2	3	4	5	6	7	8	9	10	20	30	40	50
	1	161.4	199.5	215.7	224.6	230.2	234.0	236.8	238.9	240.5	241.9	248.0	250.1	251.1	251.8
	2	18.51	19.00	19.16	19.25	19.30	19.33	19.35	19.37	19.38	19.40	19.45	19.46	19.47	19.48
	3	10.13	9.55	9.28	9.12	9.01	8.94	8.89	8.85	8.81	8.79	8.66	8.62	8.59	8.58
	4	7.71	6.94	6.59	6.39	6.26	6.16	6.09	6.04	6.00	5.96	5.80	5.75	5.72	5.70
	5	6.61	5.79	5.41	5.19	5.05	4.95	4.88	4.82	4.77	4.74	4.56	4.50	4.46	4.44
	6	5.99	5.14	4.76	4.53	4.39	4.28	4.21	4.15	4.10	4.06	3.87	3.81	3.77	3.75
	7	5.59	4.74	4.35	4.12	3.97	3.87	3.79	3.73	3.68	3.64	3.44	3.38	3.34	3.32
	8	5.32	4.46	4.07	3.84	3.69	3.58	3.50	3.44	3.39	3.35	3.15	3.08	3.04	3.02
	9	5.12	4.26	3.86	3.63	3.48	3.37	3.29	3.23	3.18	3.14	2.94	2.86	2.83	2.80
	10	4.96	4.10	3.71	3.48	3.33	3.22	3.14	3.07	3.02	2.98	2.77	2.70	2.66	2.64
	11	4.84	3.98	3.59	3.36	3.20	3.09	3.01	2.95	2.90	2.85	2.65	2.57	2.53	2.51
	12	4.75	3.89	3.49	3.26	3.11	3.00	2.91	2.85	2.80	2.75	2.54	2.47	2.43	2.40
	13	4.67	3.81	3.41	3.18	3.03	2.92	2.83	2.77	2.71	2.67	2.46	2.38	2.34	2.31
	14	4.60	3.74	3.34	3.11	2.96	2.85	2.76	2.70	2.65	2.60	2.39	2.31	2.27	2.24
	15	4.54	3.68	3.29	3.06	2.90	2.79	2.71	2.64	2.59	2.54	2.33	2.25	2.20	2.18
				\frown											
→	16	4.49	3.63	3.24	3.01	2.85	2.74	2.66	2.59	2.54	2.49	2.28	2.19	2.15	2.12
	17	4.45	3.59	3.20	2.96	2.81	2.70	2.61	2.55	2.49	2.45	2.23	2.15	2.10	2.08
	18	4.41	3.55	3.16	2.93	2.77	2.66	2.58	2.51	2.46	2.41	2.19	2.11	2.06	2.04
	19	4.38	3.52	3.13	2.90	2.74	2.63	2.54	2.48	2.42	2.38	2.16	2.07	2.03	2.00
	20	4.35	3.49	3.10	2.87	2.71	2.60	2.51	2.45	2.39	2.35	2.12	2.04	1.99	1.97

This is the value we use for a 95% HT when α =0.05, n_1 =6, and n_2 =11.

The book only has $\alpha = 0.05$, but would have another page for each α value.

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7.7 Tests with Two Independent Samples, Dichotomous Outcome

The hypothesis test on risk difference $H_0: p_1=p_2$ vs. $H_1: p_1\neq p_2$ $H_0: RD=0$ vs. $H_1: RD\neq 0$



Is equivalent to the two hypothesis tests

Risk Ratio *RR* H₀: *RR*=1 *vs.* H₁: *RR* \neq 1 and Odds Ratio *OR* H₀: *OR*=1 *vs.* H₁: *OR* \neq 1

 $RR = \frac{\hat{p}_1}{\hat{p}_2}$

$$OR = \frac{\hat{p}_1 / (1)}{\hat{p}_2 / (1)}$$

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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. 0.7 0.6 $H_0: \mu_1 = \mu_2 = \mu_3 = \mu_4$ vs. $H_1:$ at least two different 0.5 0.4 **Step 2:** Test Statistic. 0.3 0.2 F = MSB / MSE $df_1 = k-1$ $df_2 = N-k$ $\hat{F}_{\alpha,df_1,df_2}$ (3.24) **Step 3:** Decision Rule. α =0.05, df_1 =4-1=3, df_2 =20-4=16 $n_1 = n_2 = n_3 = n_4 = 5$ Reject H₀ if $F \ge 3.24$. *F* to be calculated **Step 4:** Compute test statistic. F = 25.3 / 3.0 = 8.43**Step 5:** Conclusion Because 8.43 > 3.24, reject H₀ and conclude diets mean weight loss different.





7.10 Summary

ABLE 7-50	Summary of Key Formulas for Tests of Hypothesis	
Ou	tcome Variable, Number of Groups: Null Hypothesis	Test Statistic*
Co	ntinuous outcome, two independent samples: $H_0: \mu_1 = \mu_2$	$Z = \frac{\overline{X}_{1} - \overline{X}_{2}}{S_{p}\sqrt{1/n_{1} + 1/n_{2}}}$
Co	ntinuous outcome, two matched samples: H_0 : $\mu_d = 0$	$z = \frac{\overline{\chi}_d - \mu_d}{s_d / \sqrt{n}}$
Co H	ntinuous outcome, more than two independent samples: $\mu_0: \mu_1 = \mu_2 = \dots = \mu_k$	$F = \frac{\sum n_i (\overline{X}_i - \overline{X})^2 / (k-1)}{\sum (X - \overline{X}_i)^2 / (N - K)}$
Die	chotomous outcome, one sample: $H_0: p = p_0$	$Z = \frac{\hat{\rho} - \rho_0}{\sqrt{\frac{\rho_0 (1 - \rho_0)}{n}}}$
Die H	chotomous outcome, two independent samples: $I_0: p_1 = p_2$, RD = 0, RR = 1, OR = 1	$z = \frac{\hat{\rho}_1 - \hat{\rho}_2}{\sqrt{\hat{\rho}(1 - \hat{\rho})(1/n_1 + 1/n_2)}}$
Ca H	tegorical or ordinal outcome, one sample: $p_0: p_1 = p_{10}, p_2 = p_{20}, \dots, p_k = p_{k0}$	$\chi^2 = \Sigma \frac{\left(O - E\right)^2}{E}, df = k - 1$
Ca H	tegorical or ordinal outcome, two or more independent samples: I ₀ : Outcome and groups are independent	$\chi^2 = \Sigma \frac{(0-E)^2}{E}, df = (r-1)(c-1)$





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).





Associations

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9.3 Introduction to Correlation and Regression Analysis-Correlation

5 Sums

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



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CoVariance of X&Y

Not in book

9.3 Introduction to Correlation and Regression Analysis-Correlation

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

n 1	X	X ²	Y	γ ²	XY	5 Sums
2	36.0	1204.1	2030	4120900.0	73080.0	$\nabla \mathbf{w} \in \mathbf{r} \circ \mathbf{r}$
3	29.3	858.5	1440	2073600.0	42192.0	X = 652.1 $X = 2$
4	40.1	1608.0	2835	8037225.0	113683.5	
5	35.7	1274.5	3090	9548100.0	110313.0	$Y = 493340$ $Y^2 = 1$
6	42.4	1797.8	3827	14645929.0	162264.8	
7	40.3	1624.1	3260	10627600.0	131378.0	$\sum VV = 10211c$
8	37.3	1391.3	2690	7236100.0	100337.0	$\sum XY = 192110$
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	cov(x, y) = 1798.0
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	$s^2 = 9.9638$
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	

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

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25173.2 50934928.0

2.6

$\frac{1798.0}{\sqrt{(10.0)(485478.8)}}$ r = 0.82



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 $b_0 = \overline{Y} - b_1 \overline{X}$.

Line goes through $(\overline{X}, \overline{Y})$. Note b_1 has same sign as r.

And hence we have determined our regression line.

$$\hat{y} = b_0 + b_1 x$$





r = 0.82

slope formula



9.3 Introduction to Correlation and Regression Analysis-Regression

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





 $b_1 = r \frac{s_y}{r}$ $b_1 = 0.82 \frac{5}{696.8}$ $b_1 = 180.5$

 $b_0 = \overline{Y} - b_1 \overline{X}$ $b_0 = 2902 - (180.5)(38.4)$ $b_{0} = -4029.2$



9.4 Multiple Linear Regression Analysis

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

A multiple regression analysis is run and coefficients estimated.

SBP = 68.15 + 0.58BMI + 0.65AGE + 0.94MLS + 6.44TFH

Independent Variable	Regressio Coefficier	n nt <i>t</i>	<i>p</i> -value	You will often see this typ
Intercept	<i>b</i> ₀ =68.15	<i>t</i> ₀ =26.33	$0.0001 = p_0$	The <i>t</i> statistic is for $H_0: \beta_j$
BMI	$b_1 = 0.58$	$t_1 = 10.30$	$0.0001 = p_1$	The <i>p</i> -value is the probab
Age	$b_2 = 0.65$	$t_2 = 20.22$	$0.0001 = p_2$	this coefficient estimate c
Male sex	$b_3 = 0.94$	$t_3 = 1.58$	$0.1133 = p_3$	
Treatment for hypertension	$b_4 = 6.44$	$t_4 = 9.74$	$0.0001 = p_4$	larger in abs if it were tru

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be of output. $=0, H_1: \beta_i \neq 0.$ oility of getting or Jly 0. $t_j = \frac{b_j - 0}{\sqrt{\operatorname{var}(b_j)}}$



9.5 Multiple Logistic Regression Analysis



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 <- $glm(yy^{xx}, family=binomial(link="logit"))$ summary(logistic model) <- logistic_model\$coefficients[1] b0 <- logistic_model\$coefficients[2] b1 phat <- round($1/(1+\exp(-b0-b1*xx))$), digits = 4) <- round(phat/(1-phat) 0 <- data.frame(xx,yy,phat,O) df df

#scatter plot with curve xhat <- (1:4500)/100 vhat <- 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")



```
, digits = 4)
```



9.5 Multiple Logistic Regression Analysis

Kours (x 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 $\hat{o}_i = \frac{\hat{p}_i}{1 - \hat{p}_i} = e^{\hat{\beta}_0 + \hat{\beta}_1 x_i}$ OR for a difference in x and for odds ratio $\hat{O}R = e^{\hat{\beta}_0 + \hat{\beta}_1 x_b} / e^{\hat{\beta}_0 + \hat{\beta}_1 x_a} = e^{\hat{\beta}_1 \Delta}$, $\Delta = x_b - x_a$. $\hat{\beta}_{0} = -5.66$ $\hat{\beta}_{1} = 0.21$ $\hat{OR} = e^{(0.21)(2)} = 1.5220$ OR for a difference of x=2Study 2 more hours and OR increases by 1.5.



c)	A (y)	\hat{p}	ô
	0	0.0120	0.0122
	0	0.0181	0.0184
	0	0.0272	0.0279
	0	0.0406	0.0423
	0	0.0603	0.0641
	1	0.0886	0.0972
	0	0.1284	0.1473
	0	0.1824	0.2232
	0	0.2527	0.3381
	0	0.3388	0.5124
	1	0.4371	0.7764
	0	0.5405	1.1764
	0	0.6406	1.7824
	1	0.7298	2.7008
	1	0.8036	4.0923
	1	0.8611	6.2008
	1	0.9038	9.3957
	1	0.9344	14.2365



9.6 Summary

Correlation

$$\operatorname{cov}(x, y) = \frac{1}{n-1} \left[\sum XY - \frac{1}{n} \left(\sum Y \right) \left(\sum X \right) \right]$$
$$s_x^2 = \frac{1}{n-1} \left[\sum X^2 - \frac{1}{n} \left(\sum X \right)^2 \right]$$
$$s_y^2 = \frac{1}{n-1} \left[\sum Y^2 - \frac{1}{n} \left(\sum Y \right)^2 \right]$$

$$r = \frac{\operatorname{cov}(x, y)}{\sqrt{s_x^2 s_y^2}}$$

Linear Regression

$$b_{1} = r \frac{S_{y}}{S_{x}} \qquad \hat{y} = b_{0} = \overline{Y} - b_{1}\overline{X}$$

Logistic Regression

$$\hat{p} = \frac{1}{1 + e^{-b_0 - b_1 x_1 - \dots - b_p x_p}}$$
$$ln\left(\frac{\hat{p}}{1 - \hat{p}}\right) = b_0 + b_1 x_1 + b_1 x_1 + b_1 x_2 + b_2 x_2 + b_1 x_2 + b_2 x_2 + b$$

 $\hat{O}R = e^{\hat{\beta}_1 \Delta_1 + \dots + \hat{\beta}_p \Delta_p}$

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$= b_0 + b_1 x$

logistic probability

 $\dots + b_p x_p$

log odds

odds ratio for difference Δ_j in x_j



10.1 Introduction to Nonparametric Testing – Sign Test

Example: Mark is training for 10K. *n*=20 daily runs.

Binomial Distribution, n=20, p=0.50.18 0.16 0.14 0.12 0.1 0.08 0.06 0.04 0.02 0 ~ 5 6 1 8 9 0 1 1 1 1 1 1 0 0 0 1 0 0 0

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

							TUDIC	0
Ctop 1. II . MD 4. vo II . M	Тм	vo-Sided Testα	.10	.05				
Step 1: H_0 . <i>MD</i> =4 <i>VS</i> . H_1 . <i>M</i>	D>2	ŀ, α=	0.05		Or	ne-Sided Test α	.05	.025
	х	P(X=x)	CumSum	CumSumR		n		
Step 2: Test Statistic	0	0.000	0.000	1.000		1		
	1	0.000	0.000	1.000		2		
r - the number of 1's	2	0.000	0.000	1.000		4		
x = the number of $+ 5$.	3	0.001	0.001	1.000		5	0	
	4	0.005	0.006	0.999		6	0	0
	5	0.015	0.021	0.994		7	0	0
Dinamial Distribution 20 05	6	0.037	0.058	0.979		8	1	0
Binomial Distribution, $n=20$, $p=0.5$	7	0.074	0.132	0.942		10	1	1
	8	0.120	0.252	0.868		11	2	1
0.14	9	0.160	0.412	0.748		12	2	2
0.12 -	10	0.176	0.588	0.588		13	3	2
0.1 -	11	0.160	0.748	0.412		14	3	2
0.08 -	12	0.120	0.868	0.252		15	3	3
0.06	13	0.074	0.942	0.132		17	4	4
	14	0.037	0.979	0.058		18	5	4
	15	0.015	0.994	0.021		19	5	4
O < ひ 3 & 5 6 1 8 9 0 < 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2 2	16	0.005	0.999	0.006		→ 20	5	5
	17	0.001	1.000	0.001		21	6	5
	18	0.000	1.000	0.000		23	7	6
$X = -15$ (or $n_{-}5-15$)	19	0.000	1.000	0.000		24	7	6
$\Lambda_{0.05} - 13$ (01 $n - 3 - 13$)	20	0,000	1,000	0.000		25	7	7
	20	0.000	1.000	0.000				

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.02	.01
.01	.005
0	
0	0
0	0
0	0
1	0
1	1
1	1
2	1
2	2
2	2
3	2
3	3
4	3
4	3
4	4
5	4
5	4
5	5
6	5

Tabla 6

data
5
3
5
3
4
4
6
6
6
4
6
5
5
5
4
5
5
5
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 RECALL

The hypothesis testing process consists of 5 Steps.

Step 3: Set-up the decision rule.







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Biostatistical Methods

10.1 Introduction to Nonparametric Testing – Sign Test

Two-Sided Test a

One-Sided Test α

19

20 21

If value > MD_{0} , +.

.10

.05

.05

.025

5

Table 6

Step 4: Compute the test statistic. x = 14 $x = (\text{the number of observations} > MD_0 = 4)$

Step 5: Because $x=14 < x_{\alpha}=15$, do not reject H_0 .

		-	0			
х	P(X=x)	CumSum	CumSumR			
5	0.015	0.021	0.994			
6	0.037	0.058	0.979			
14	0.037	0.979	0.058			
15	0.015	0.994	0.021			
See also Table 6						

N I	-1-	_
IN	ole	

If we used normal, we would reject H₀, $t=4.07 > t_{0.05,19}=2.093$. $t = \frac{\overline{X} - \mu_0}{s / \sqrt{n}} d_{f=n-1} \overline{X} = 4.8500 s = 0.9333$





If value $< MD_0$, –. If value = MD_0 , 0.

.02	.01
.01	.005
4	3
4	3
4	4

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 α . $H_0:MD_1=MD_2$ vs. $H_1:MD_1 \neq MD_2$, $\alpha=0.05$

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

						Rai	nks	
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	$R_{-}=37$
12	1	5		6		6		\mathbf{n}_{1} -37
		6	6	7.5	7.5	7.5	7.5	ח 10
		7		9		9		$K_{2}=18$
		12		10		10		2



Group 1	Group 2
Placebo	NewDrug
7	3
5	6
6	4
4	2
12	10
$n_1 = 5$	$n_2 = 5$



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.

$$U_{1} = n_{1}n_{2} + \frac{n_{1}(n_{1}+1)}{2} - R_{1} = (5)(5) + \frac{5(5+1)}{2} - 37 = 3$$

$$U_{2} = n_{1}n_{2} + \frac{n_{2}(n_{2}+1)}{2} - R_{2} = (5)(5) + \frac{5(5+1)}{2} - 18 = 22$$

$$U = \min(U_{1}, U_{2}) = \min(3, 22) = 3$$

$$U = \min(U_{1}, U_{2}) = \min(3, 22) = 3$$

Reject H_0 for small U.

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complete separation



Rankings

oup 1	Group 2	Group 1	Group 2
	1		1
	2	2	
	3		3
	4	4	
	5		5
6		6	
7			7
8		8	
9			9
10		10	
U :	= 0	U =	= 25

complete alternating

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 H₀ if $U \le U_{0.05,n_1,n_2}$.

Step 4: Compute test statistic. Already done, U=3.

Step 5: Conclusion. Do not reject H_0 because $U=3>U_{0.05,5,5}=2$. Interpret.

Two-	Sided	Test	α = 1	0.05																
										n	1									
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:

H₀: The median difference is zero (H₀: $\delta = 0$)

H₁: The median difference is positive (H₁: $\delta > 0$)

We will calculate a test statistic W the smaller of W+ and W_- .

- W+ = sum of positive ranks
- $W_{-} =$ sum of negative ranks

$$\longrightarrow$$
 W=min(W+, W_-)

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. **D.B.** Rowe



 $H_0: \delta \leq 0$ vs. $H_1: \delta > 0$ δ is population version of *d*.



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:

D.B. Rowe

$$H_0: \delta \leq 0 \text{ vs. } H_1: \delta > 0$$

 δ is population version of d.

Step 2: Select the test statistic.

- $W_{\perp} = \text{sum of positive ranks} = 32$
- W_{-} = sum of negative ranks = 4
- $W = \min(W_1, W_2) = \min(4, 32) = 4$

Reject H_0 for small W.

b	а	d	sorted	sign	rank	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
	n=8					

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				
W = 0	W = 10	V = 16	W - 2				



IF

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 H_0 if $W \le W_{\alpha,n}$.



Step 4: Compute test statistic. Already done, *W*=4.

Step 5: Conclusion. Reject H_0 because $W=4 \le W_{0.05,8}=6$. Interpret.

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

The hypothesis testing process consists of 5 Steps.

Step 1: Set up the hypotheses and determine the level of significance α .

 $H_0: MD_1 = MD_2 \dots = MD_k$ vs. $H_1:$ at least two MD's different reject for "large" disparities H.

We will assume the medians are equal and see how different from equal.



Test



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 α .

vs. H₁: at least two μ 's different $H_0: \mu_1 = \mu_2 \dots = \mu_k$ reject for "large" disparities F = MSB/MSE.

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.

- $H_0: MD_1 = MD_2 = MD_3$ vs. $H_1:$ at least two different
- 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 H_0 if $H \ge 5.656$.

Step 4: Compute test statistic.

H = 7.52

Step 5: Conclusion

Reject H_0 because 7.52 > 5.656, and conclude difference in median albumin.

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Tł	nree grou
n ₁	n ₂
5	4







Sample size order doesn't matter.



10.5 Summary

Sign Test (one sample) x = number of observations > MD_0

Mann-Whitney U Test

$$U_{1} = n_{1}n_{2} + \frac{n_{1}(n_{1}+1)}{2} - R_{1}$$
$$U_{2} = n_{1}n_{2} + \frac{n_{2}(n_{2}+1)}{2} - R_{2}$$
$$U = \min(U_{1}, U_{2})$$

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

$$H = \left(\frac{12}{N(N+1)}\sum_{j=1}^{k}\frac{R_j^2}{n_j}\right)$$

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-3(N+1)



11.1 Introduction to Survival Data

Survival analysis measures two pieces of information

- 1) Whether the event occurred, 1 = yes, 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.



t=0.0 : survival probability=1.00 t=2.0 : survival probability=0.90 t=9.0 : survival probability=0.50 (Median) t=10.0: survival probability=0.45

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curves.



11.6 Summary

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

Actuarial Life Table

- $N_{t} = \# \text{ event free during interval } t$ $D_{t} = \# \text{ who die in interval } t$ $C_{t} = \# \text{ censored in interval } t$
- N_{t*} = avg. # at risk in interval t, N_{t*} = $N_t C_t/2$
- $q_t = \text{prop. die in interval } t, q_t = D_t / N_t$ $p_t = \text{prop. survive in interval } t, p_t = 1 - q_t$

 S_t = prop. survive past interval tCan plot S_t vs. t.

Kaplan-Meier Life Table $SE(S_t) = S_t$ Chi-Square Test Cox Proportional Hazards Model $h(t) = h_0(t) \exp(b_1 x_1 + b_2 x_2 + \dots + b_n x_n)$





Questions?

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





