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7.28 let  $p$  be the proportion of deaths from lung cancer in this plant.

$$H_0: p = 0.12 \quad H_1: p \neq 0.12$$

7.29 two-sided

7.30 let  $x$  be the number of deaths due to lung cancer.

$$P(x \geq 5 | H_0 \text{ is true}) = \sum_{i=5}^{20} C_{20}^i 0.12^i (1-0.12)^{20-i} \\ = 0.082719$$

binom.test(5, 20, 0.12, alt="two.sided")

7.31 .  $Z = \frac{\hat{p} - p_0}{\sqrt{p_0(1-p_0)/n}}$  where  $\hat{p} = 19/90$ ,  $p_0 = 0.12$ ,  $n = 90$ ,  $Z_0 = 2.6599$

when  $H_0$  is true,  $Z \sim N(0, 1)$

$$P(|Z| > Z_0 | Z \sim N(0, 1)) = 2 \times 0.00391 = 0.00782$$

7.32 the ~~not~~ proportion of deaths from lung cancer when deaths caused by IHD excluded is  $p_0$

$$p_0 = \frac{0.12}{1-0.4} = 0.2$$

$$\hat{p} = \frac{19}{90-18} = \frac{19}{72}, \quad n = 90 - 18 = 72$$

$$z_0 = \frac{\hat{p} - p_0}{\sqrt{p_0(1-p_0)/n}} = 1.355288$$

$$P(|Z| > z_0 | Z \sim N(0,1)) = 0.174576$$

7.51 one-sample t test

7.52 let  $\mu$  be the mean of stomach-cancer cases.

$$H_0: \mu = 2.88 \quad H_1: \mu \neq 2.88$$

$$T = \frac{\bar{X} - \mu}{s/\sqrt{n}} \stackrel{H_0}{\sim} t(19)$$

$$\text{when } \bar{X} = 2.65, s/\sqrt{n} = se = 0.11$$

$$T = -2.09091$$

$$\text{while } -t_{0.975}(19) = -2.093024 < T$$

thus ~~we~~ we reject  $H_0$  at 95% confidence level.

$$p = P(|T| > 2.09091 | T \sim t(19)) = 0.05021$$

7.53 `pwr.t.test(d = 0.2/0.11/sqrt(20), power = 0.8,`

`type = "one.sample", alt = "two.sided")`

$d = \frac{\text{difference between the means}}{\text{pooled standard deviation (here just } se/\sqrt{n} = s)}$

8.25  $\mu_1$ : mean concentration of drug A  
 $\mu_2$ : mean concentration of drug B

$$H_0: \mu_1 = \mu_2 \quad H_1: \mu_1 \neq \mu_2$$

8.26 use paired t test

8.27  $d_i$ : the difference between drug A and drug B

$$T = \frac{\bar{d}}{s_d/\sqrt{n}} \quad H_0: t(n-1)$$

$$\text{when } s_d = \sqrt{\frac{\sum (d_i - \bar{d})^2}{n-1}} = 3.0984$$

$$\bar{d} = 3.6$$

$$T = 3.6742$$

$$P(|T| > 3.6742 \mid T \sim t(n-1)) = 0.0051 \approx 0.05$$

thus we  reject  $H_0$  at 95% confidence level.

8.28.  $\bar{d} = 3.6$

8.29 
$$\frac{|d - \bar{d}|}{s_d/\sqrt{n}} < t_{0.975}(9)$$

$$-1.3835 < d < 5.8164$$

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### 9.7 wilcoxon rank-sum test

$F(x)$ : the p.d.f. of birthweight in treatment group

$G(x)$ : the p.d.f. of birthweight in control group

$$H_0: F(x) = G(x) \quad H_1: F(x) \neq G(x)$$

具体计算过程参照课件

$$W = 170.5, p = 0.01692$$

9.8 since the sample size is small, and the distribution of birthweight is unknown, parametric methods are not preferable.

9.15  $m_1$ : median duration of effusion in breast-fed baby.

$m_2$ : median duration of effusion in bottle-fed baby.

$$H_0: m_1 = m_2 \quad H_1: m_1 < m_2$$

### 9.17 wilcoxon signed-rank test

$$9.18 \quad H_0: m_1 = m_2 \quad H_1: m_1 \neq m_2$$

具体计算过程参照课件

$$W^+ = 61, E(W^+) = 138, \text{Var}(W^+) = 1079$$

$$p = 0.01986$$

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$$H_0: p_1 = p_2 \quad H_1: p_1 \neq p_2$$

$$Z = \frac{\hat{p}_1 - \hat{p}_2}{\sqrt{\bar{p}(1-\bar{p})(\frac{1}{n_1} + \frac{1}{n_2})}} \stackrel{H_0}{\sim} N(0,1) \quad \bar{p} = \frac{p_1 + kp_2}{1+k}$$

$$\text{Power} = P(Z > Z_{1-\frac{\alpha}{2}} \mid |p_1 - p_2| = \Delta \neq 0)$$

$$= P\left(\frac{\hat{p}_1 - \hat{p}_2}{\sqrt{\bar{p}\bar{q}(\frac{1}{n_1} + \frac{1}{n_2})}} > Z_{1-\frac{\alpha}{2}} \mid |p_1 - p_2| = \Delta \neq 0\right)$$

$$= P\left(\frac{\hat{p}_1 - \hat{p}_2 - \Delta}{\sqrt{p_1q_1/n_1 + p_2q_2/n_2}} > Z_{1-\frac{\alpha}{2}} \frac{\sqrt{\bar{p}\bar{q}(\frac{1}{n_1} + \frac{1}{n_2})}}{\sqrt{p_1q_1/n_1 + p_2q_2/n_2}} - \frac{\Delta}{\sqrt{p_1q_1/n_1 + p_2q_2/n_2}}\right)$$

$$\stackrel{H_1}{\sim} \frac{\hat{p}_1 - \hat{p}_2 - \Delta}{\sqrt{p_1q_1/n_1 + p_2q_2/n_2}} \stackrel{H_1}{\sim} N(0,1)$$

$$\Rightarrow 1-\beta = \Phi\left(\frac{\Delta}{\sqrt{p_1q_1/n_1 + p_2q_2/n_2}} - Z_{1-\frac{\alpha}{2}} \frac{\sqrt{\bar{p}\bar{q}(\frac{1}{n_1} + \frac{1}{n_2})}}{\sqrt{p_1q_1/n_1 + p_2q_2/n_2}}\right)$$

$$Z_{1-\beta} = \frac{\Delta}{\sqrt{p_1q_1/n_1 + p_2q_2/n_2}} - Z_{1-\frac{\alpha}{2}} \frac{\sqrt{\bar{p}\bar{q}(\frac{1}{n_1} + \frac{1}{n_2})}}{\sqrt{p_1q_1/n_1 + p_2q_2/n_2}}$$

$$n_2 = kn_1 \text{ 代入}$$

$$\text{得 } n_1 = \left[ \sqrt{\bar{p}\bar{q}(1+\frac{1}{k})} Z_{1-\frac{\alpha}{2}} + \sqrt{p_1q_1 + \frac{p_2q_2}{k}} Z_{1-\beta} \right]^2 / \Delta^2$$

$$n_2 = kn_1$$



11.12 推导 E(MSW) 和 E(MSB) 两个表达式

$$y_{ij} = \mu + \alpha_i + e_{ij}, \quad \alpha_i \sim N(0, \sigma_A^2), \quad e_{ij} \sim N(0, \sigma^2) \quad i=1, \dots, k, j=1, \dots, n_i$$

$$MSW = \frac{1}{n-k} \sum_i^k \sum_j^{n_i} (y_{ij} - \bar{y}_i)^2$$

$$\bar{y}_i = \frac{1}{n_i} \sum_j^{n_i} y_{ij} = \mu + \alpha_i + \frac{1}{n_i} \sum_j^{n_i} e_{ij}$$

$$\therefore E(y_{ij} - \bar{y}_i) = E(e_{ij} - \frac{1}{n_i} \sum_j^{n_i} e_{ij}) = 0$$

$$\begin{aligned} \text{Var}(y_{ij} - \bar{y}_i) &= \text{Var}(e_{ij} - \frac{1}{n_i} \sum_j^{n_i} e_{ij}) \\ &= \text{Var}(e_{ij}) + \frac{1}{n_i^2} \text{Var}(\sum_j^{n_i} e_{ij}) - 2\text{cov}(e_{ij}, \frac{1}{n_i} \sum_j^{n_i} e_{ij}) \\ &= \sigma^2 + \frac{\sigma^2}{n_i} - \frac{2}{n_i} \sigma^2 \\ &= (1 - \frac{1}{n_i}) \sigma^2 \end{aligned}$$

$$\begin{aligned} E(MSW) &= \frac{1}{n-k} \sum_i^k \sum_j^{n_i} E(y_{ij} - \bar{y}_i)^2 \\ &= \frac{1}{n-k} \sum_i^k \sum_j^{n_i} [E^2(y_{ij} - \bar{y}_i) + \text{Var}(y_{ij} - \bar{y}_i)] \\ &= \frac{1}{n-k} \sum_i^k \sum_j^{n_i} (1 - \frac{1}{n_i}) \sigma^2 \\ &= \sigma^2 \end{aligned}$$

$$MSB = \frac{1}{k-1} \sum_i^k (\bar{y}_i - \bar{y})^2$$

$$\begin{aligned} \bar{y} &= \frac{1}{n} \sum_i^k \sum_j^{n_i} y_{ij} = \frac{1}{n} \sum_i^k \sum_j^{n_i} (\mu + \alpha_i + e_{ij}) \\ &= \mu + \frac{1}{n} \sum_i^k n_i \alpha_i + \frac{1}{n} \sum_i^k \sum_j^{n_i} e_{ij} \end{aligned}$$

$$\therefore \bar{y}_i - \bar{y} = \alpha_i - \frac{1}{n} \sum_i^k n_i \alpha_i + \frac{1}{n_i} \sum_j^{n_i} e_{ij} - \frac{1}{n} \sum_i^k \sum_j^{n_i} e_{ij}$$

$$E(\bar{y}_i - \bar{y}) = 0$$

$$\text{Var}(\bar{y}_i - \bar{y}) = \text{Var}(\alpha_i - \frac{1}{n} \sum_i^k n_i \alpha_i) + \text{Var}(\frac{1}{n_i} \sum_j^{n_i} e_{ij} - \frac{1}{n} \sum_i^k \sum_j^{n_i} e_{ij})$$

$$\begin{aligned} \text{Var}(\alpha_i - \frac{1}{n} \sum_i^k n_i \alpha_i) &= \text{Var}(\alpha_i) + \text{Var}(\frac{1}{n} \sum_i^k n_i \alpha_i) - 2\text{cov}(\alpha_i, \frac{1}{n} \sum_i^k n_i \alpha_i) \\ &= \sigma_A^2 + \frac{\sum_i^k n_i^2 \sigma_A^2}{n^2} - \frac{2n_i \sigma_A^2}{n} \end{aligned}$$

$$\text{Var}(\frac{1}{n_i} \sum_j^{n_i} e_{ij} - \frac{1}{n} \sum_i^k \sum_j^{n_i} e_{ij}) = \frac{\sigma^2}{n_i} + \frac{\sigma^2}{n} - \frac{2}{n} \sigma^2 = (\frac{1}{n_i} - \frac{1}{n}) \sigma^2$$

$$\begin{aligned}
\therefore E(MSB) &= \frac{1}{k-1} \sum_{i=1}^k \sum_{j=1}^{n_i} E(\bar{y}_i - \bar{y})^2 \\
&= \frac{1}{k-1} \sum_{i=1}^k \sum_{j=1}^{n_i} [E^2(\bar{y}_i - \bar{y}) + \text{Var}(\bar{y}_i - \bar{y})] \\
&= \frac{1}{k-1} \sum_{i=1}^k \sum_{j=1}^{n_i} \left( \sigma_A^2 + \frac{\sum_{i=1}^k n_i^2 \sigma_A^2}{n^2} - \frac{2 \sum_{i=1}^k n_i \sigma_A^2}{n} + \frac{\sigma^2}{n_i} - \frac{\sigma^2}{n} \right) \\
&= (n \sigma_A^2 + \frac{\sum_{i=1}^k n_i^2 \sigma_A^2}{n} - \frac{2 \sum_{i=1}^k n_i \sigma_A^2}{n} + k \sigma^2 - \sigma^2) / (k-1) \\
&= [n \sigma_A^2 - \frac{\sum_{i=1}^k n_i^2 \sigma_A^2}{n} + (k-1) \sigma^2] / (k-1) \\
&= \frac{n - \frac{\sum_{i=1}^k n_i^2}{n}}{k-1} \sigma_A^2 + \sigma^2
\end{aligned}$$

when  $n_1 = n_2 = \dots = n_k$ ,  $\frac{n - \frac{\sum_{i=1}^k n_i^2}{n}}{k-1} = n_1$

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12.37 计算前可对数据进行对数变换

random effect model:  $y_{ij} = \mu + \alpha_i + e_{ij}$   $k=30$ where  $\alpha_i \sim N(0, \sigma_A^2)$ ,  $e_{ij} \sim N(0, \sigma^2)$  $y_{ij}$ : temperature at  $j$ th location on  $i$ th day

$$MSB = \frac{SSB}{k-1} = \frac{\sum_{i=1}^k \sum_{j=1}^{n_i} (\bar{y}_i - \bar{y})^2}{k-1} = 0.02934 / 131.906$$

$$MSW = \frac{SSW}{n-k} = \frac{\sum_{i=1}^k \sum_{j=1}^{n_i} (y_{ij} - \bar{y}_i)^2}{n-k} = 0.0006735 / 2.861$$

$$\text{where } \bar{y}_i = \frac{\sum_{j=1}^{n_i} y_{ij}}{n_i}, \bar{y} = \frac{\sum_{i=1}^k \sum_{j=1}^{n_i} y_{ij}}{\sum_{i=1}^k n_i} \quad n_1 = n_2 = \dots = n_{30} = 21$$

$$\hat{\sigma}_A^2 = \frac{MSB - MSW}{21} = \dots$$

12.38. here,  $k=21$ ,  $n_1 = n_2 = \dots = n_{21} = 30$ 

$$y_{ij} = \mu + \alpha_j + e_{ij} \quad \alpha_j \sim N(0, \sigma_B^2), e_{ij} \sim N(0, \sigma^2)$$

$$H_0: \sigma_B^2 = 0 \quad H_1: \sigma_B^2 \neq 0$$

$$F = \frac{MSB}{MSW} \stackrel{H_0}{\sim} F(k-1, n-k)$$

$$MSB = \frac{\sum_{i=1}^k \sum_{j=1}^{n_i} (\bar{y}_i - \bar{y})^2}{k-1} = 0.0083 / 35.65$$

$$MSW = \frac{\sum_{i=1}^k \sum_{j=1}^{n_i} (y_{ij} - \bar{y}_i)^2}{n-k} = 0.00179 / 7.929$$

$$\Rightarrow \text{where } \bar{y}_i = \frac{\sum_{j=1}^k y_{ij}}{k}$$

$$F^* > F_{0.95}(20, 609), \text{ reject } H_0.$$



12.3.9

fixed effect model:  $y_{ij} = \mu + \alpha_i + e_{ij}$ ,  $e_{ij} \sim N(0, \sigma^2)$

$H_0: \alpha_i = \alpha_j$ ,  $H_1: \alpha_i \neq \alpha_j$  ( $\forall i \neq j, i < j$ )

t test (unpaired)

then use Bonferroni method to get correct assessments.

12.68 & 12.69 按照课件计算公式即可

1. OR 适用于 case-control study

RR, RD, OR 都可用于 prospective study

因为在 RR 和 RD 的计算中涉及 exposed 和 not exposed 中的患病率, 而在回顾性研究中患病率是无法估计的, OR 的计算中涉及的是人群暴露率, 这在两种研究中都可获得估计。

2.

	Y=0	Y=1	
X=0	A	B	Y: 疾病发生情况
X=1	C	D	X: 暴露情况

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

$$\hat{p}_1 = \frac{D}{C+D}, \quad \hat{p}_2 = \frac{B}{A+B}$$

$$= \frac{AD}{BC}$$

$$n_1 = C+D, \quad n_2 = A+B$$

$$\text{Var}(\ln \hat{OR}) = \text{Var}\left(\ln \frac{\hat{p}_1}{1 - \hat{p}_1}\right) + \text{Var}\left(\ln \frac{\hat{p}_2}{1 - \hat{p}_2}\right)$$

由 delta-method, 令  $g(x) = \ln \frac{x}{1-x}$ ,  $g'(x) = \frac{1}{x(1-x)}$

$$\text{则 } \text{Var}\left(\ln \frac{\hat{p}_1}{1 - \hat{p}_1}\right) \approx (g'(\hat{p}_1))^2 \frac{\hat{p}_1(1 - \hat{p}_1)}{n_1}$$

$$= \frac{1}{\hat{p}_1^2(1 - \hat{p}_1)^2} \cdot \frac{\hat{p}_1(1 - \hat{p}_1)}{n_1} = \frac{1}{n_1 \hat{p}_1(1 - \hat{p}_1)} = \frac{1}{C} + \frac{1}{D}$$

$$\text{同理可得 } \text{Var}\left(\ln \frac{\hat{p}_2}{1 - \hat{p}_2}\right) \approx \frac{1}{n_2 \hat{p}_2(1 - \hat{p}_2)} = \frac{1}{A} + \frac{1}{B}$$

$$\text{故 } \text{Var}(\ln \hat{OR}) \approx \frac{1}{A} + \frac{1}{B} + \frac{1}{C} + \frac{1}{D}$$

假设 Mc Nemar's test 中列联表如下:

		Case		
		Exposed	Not exposed	Total
Control	Exposed	$n_{11}$	$n_{12}$	$n_{11}+n_{12}$
	Not exposed	$n_{21}$	$n_{22}$	$n_{21}+n_{22}$
	Total	$n_{11}+n_{21}$	$n_{12}+n_{22}$	$n$

共有几个对, 在 Mantel-Haenszel test 中可对应分成  $n$  层 ( $a_i+b_i=1, c_i+d_i=1$ )

有  $n_{11}$  层:

	exposed	Not exposed
case	1	0
control	1	0

$$O_i = a_i = 1$$

$$E_i = \frac{(a_i+b_i)(a_i+c_i)}{n_i} = 1$$

$$V_i = \frac{(a_i+b_i)(a_i+c_i)(b_i+d_i)(c_i+d_i)}{n_i^2(n_i-1)}$$

有  $n_{12}$  层

	exposed	Not exposed
case	0	1
control	1	0

$$= 0$$

$$O_i = 0$$

$$E_i = \frac{1}{2}$$

$$V_i = \frac{1}{4}$$

有  $n_{21}$  层

	exposed	Not exposed
case	1	0
control	0	1

$$O_i = 1$$

$$E_i = \frac{1}{2}$$

$$V_i = \frac{1}{4}$$

有  $n_{22}$  层

	exposed	Not exposed
case	0	1
control	0	1

$$O_i = 0$$

$$E_i = 0$$

$$V_i = 0$$

$$\text{则 } O = \sum O_i = n_{11} + n_{21}$$

$$E = \sum E_i = n_{11} + \frac{1}{2}n_{12} + \frac{1}{2}n_{21}$$

$$V = \sum V_i = \frac{1}{4}n_{12} + \frac{1}{4}n_{21}$$

$$\chi_{MH}^2 = \frac{(O-E)^2}{V} = \frac{(n_{12}-n_{21})^2}{n_{12}+n_{21}} \quad H_0: \chi^2(u)$$

其中  $n_{12}+n_{21}=n_D$  (不一致对数), 故  $\chi_{MH}^2$  即为 Mc Nemar's test 中的统计量.