Coursework 5: STAT 570

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1. Consider the data given in Table 1, which are a simplified version of those reported in Breslow and Day (1980). These data arose from a case-control study that was carried out to investigate the relationship between esophageal cancer and various risk factors. Disease status is denoted Y with Y=0 and Y=1 corresponding to without/with disease and alcohol consumption is represented by X with X=0 and X=1 denoting less than 80g and greater than or equal to 80g on average per day. Let the probabilities of high alcohol consumption in the cases and controls be denoted

$$p_1 = \mathbb{P}(X = 1 \mid Y = 1) \text{ and } p_2 = \mathbb{P}(X = 1 \mid Y = 0),$$
 (1)

respectively. Further, let X_1 be the number exposed from n_1 cases and X_2 be the number exposed from n_2 controls. Suppose $X_i \mid p_i \sim \text{Binomial}(n_i, p_i)$ in the case (i = 1) and control (i = 2) groups.

(a) Of particular interest in studies such as this is the odds ratio defined by

$$\theta = \frac{\mathbb{P}(Y = 1 \mid X = 1) / \mathbb{P}(Y = 0 \mid X = 1)}{\mathbb{P}(Y = 1 \mid X = 0) / \mathbb{P}(Y = 0 \mid X = 0)}.$$
 (2)

Show that the odds ratio is equal to

$$\theta = \frac{\mathbb{P}(X=1 \mid Y=1) / \mathbb{P}(X=0 \mid Y=1)}{\mathbb{P}(X=1 \mid Y=0) / \mathbb{P}(X=0 \mid Y=0)} = \frac{p_1/(1-p_1)}{p_2/(1-p_2)}.$$
 (3)

Solution: We have that

$$\mathbb{P}(Y = y \mid X = x) = \frac{\mathbb{P}(X = x \mid Y = y) \mathbb{P}(Y = y)}{\mathbb{P}(X = x)}$$
(4)

by Bayes' rule. Applying Equation 4 to Equation 2, we get

$$\theta = \frac{\left[\mathbb{P}(X=1 \mid Y=1)\,\mathbb{P}(Y=1)\right]/\left[\mathbb{P}(X=0 \mid Y=1)\,\mathbb{P}(Y=0)\right]}{\left[\mathbb{P}(X=0 \mid Y=1)\,\mathbb{P}(Y=1)\right]/\left[\mathbb{P}(X=0 \mid Y=0)\,\mathbb{P}(Y=0)\right]}.$$
 (5)

The $\mathbb{P}(Y=y)$ factors cancel and we obtain the first part of Equation 3. Using Equation 1, we substitute to obtain the second part of Equation 3.

$$\begin{array}{c|cccc} & X = 0 & X = 1 \\ \hline Y = 1 & 104 & 96 & 200 \\ Y = 0 & 666 & 109 & 775 \\ \hline \end{array}$$

Table 1: Case-control data: Y = 1 corresponds to the event of esophageal cancer, and X = 1 exposure to greater than 80g of alcohol per day. There are 200 cases and 775 controls.

(b) Obtain the MLE and a 90% confidence interval for θ , for the data of Table 1.

Solution: The likelihood and log-likelihood functions are

$$L(p_1, p_2) = \binom{n_1}{x_1} p_1^{x_1} (1 - p_1)^{n_1 - x_1} + \binom{n_2}{x_2} p_2^{x_2} (1 - p_2)^{n_2 - x_2}$$

$$l(p_1, p_2) = \log L(p_1, p_2)$$

$$= \sum_{i=1}^{2} \left[\log \binom{n_i}{x_i} + x_i \log p_i + (n_i - x_i) \log (1 - p_i) \right],$$
(6)

so the score function is

$$S(p_1, p_2) = \nabla \log L(p_1, p_2) = \begin{pmatrix} \frac{x_1 - n_1 p_1}{p_1 (1 - p_1)} \\ \frac{x_2 - n_2 p_2}{p_2 (1 - p_2)} \end{pmatrix}$$
(7)

Thus, the Fisher information is

$$I(p_1, p_2) = mathbb E[S(p_1, p_2) S(p_1, p_2)^{\mathsf{T}}] = \begin{pmatrix} \frac{n_1}{p_1(1-p_1)} & 0\\ 0 & \frac{n_2}{p_2(1-p_2)} \end{pmatrix}. \tag{8}$$

From Equation 7, we can solve $S(\hat{p}_1, \hat{p}_2) = \mathbf{0}$ to get the MLEs $\hat{p}_1 = x_1/n_1$ and $\hat{p}_2 = x_2/n_2$. Since the MLE is invariant to reparameterization, we have the MLE for θ :

$$\hat{\theta} = \frac{\hat{p}_1 / (1 - \hat{p}_1)}{\hat{p}_2 / (1 - \hat{p}_2)} = \frac{1992}{1417} \approx 5.640.$$
 (9)

We estimate the confidence interval for $\log \hat{\theta}$ which works since log is a monotonic transform. Using the delta method and Equation 8, we have that

$$\operatorname{Var}\left(\log \hat{\theta}\right) \approx \left(\nabla \log \hat{\theta}\right)^{\mathsf{T}} \left(I\left(\hat{p}_{1}, \hat{p}_{2}\right)\right)^{-1} \left(\nabla \log \hat{\theta}\right)$$

$$= \left(\frac{1}{\hat{p}_{1}(1-\hat{p}_{1})} \quad \frac{1}{\hat{p}_{2}(1-\hat{p}_{2})}\right) \begin{pmatrix} \frac{\hat{p}_{1}(1-\hat{p}_{1})}{n_{1}} & 0\\ 0 & \frac{\hat{p}_{2}(1-\hat{p}_{2})}{n_{2}} \end{pmatrix} \begin{pmatrix} \frac{\hat{p}_{1}(1-\hat{p}_{1})}{1}\\ \frac{1}{\hat{p}_{2}(1-\hat{p}_{2})} \end{pmatrix}$$

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

$$= \frac{1}{n_{1}\hat{p}_{1}} + \frac{1}{n_{1}\left(1-\hat{p}_{1}\right)} + \frac{1}{n_{2}\hat{p}_{2}} + \frac{1}{n_{2}\left(1-\hat{p}_{2}\right)}. \tag{10}$$

Numerically, this is $Var(\log \hat{\theta}) \approx 0.0307$.

The 90% confidence interval for $\log \hat{\theta}$ is approximately

$$\left(\log \hat{\theta} - \Phi^{-1}(0.95)\sqrt{\operatorname{Var}\left(\log \hat{\theta}\right)}, \log \hat{\theta} + \Phi^{-1}(0.95)\sqrt{\operatorname{Var}\left(\log \hat{\theta}\right)}\right), (11)$$

which is about (1.441, 2.018). Taking the exponent of both sides, we have a 90% confidence interval for $\hat{\theta}$ of (4.228, 7.524).

(c) We now consider a Bayesian analysis. Assume that the prior distribution for p_i is the beta distribution Beta (a, b) for i = 1, 2. Show that the posterior distribution $p_i \mid x_i$ is given by the beta distribution Beta $(a + x_i, b + n_i - x_i)$, i = 1, 2.

Solution: