

# BIOS762 - Notes

Mingwei Fei

December 4, 2022

## 1 Multinomial distribution

Get the covariance matrix for cross-sectional, prospective, retrospective sampling method.

### 1.1 Likelihood for one random variable

To calculate the covariance matrix, we will use the MGF and take derivatives. Or use the cumulant function KGF to get the covariance.

Use one random variable for the two way contingency table. While the Fisher information is the inverse of the covariance matrix, however we don't use Fisher information to calculate covariance matrix due to the math computation.

For one random variable  $Y$ :

$$\begin{aligned} p(\theta) &= \prod_{i=1}^n \prod_{j=1}^J \pi_j^{I(Y_i=j)}, \quad \theta = (\pi_1, \pi_2, \dots, \pi_J)' \\ \ln p(\theta) &= \sum_{i=1}^n \sum_{j=1}^J I(Y_i = j) \log(\pi_j) = \sum_{j=1}^J n_j \log(\pi_j) \\ M_X(t) &= E[\exp(t^T X)] = E[\exp(t^T (Y_1 + Y_2 + \dots + Y_n))] = E[\exp(t^T Y_1 + t^T Y_2 + \dots + t^T Y_n)] \\ &= E\left[\prod_{i=1}^n \exp(t^T Y_i)\right] \\ &= \prod_{i=1}^n E[\exp(t^T Y_i)] \quad (\text{by independence}) \\ &= \prod_{i=1}^n M_{Y_i}(t) = \prod_{i=1}^n P(Y_i = 1) e^{t y_i} \quad \text{by MGF of discrete variable } Y_i \\ &= \left( \sum_{j=1}^J \pi_j \exp(t_j) \right)^n \quad \text{by MGF of multinoulli} \end{aligned}$$

The MGF for bernoulli distribution

$$M_X(t) = 1 - p + p \exp(t), \quad K_X(t) = \log(1 - p + p \exp(t))$$

For multinomial distribution

$$M_X(t) = (1 - p + p \exp(t))^n, \quad K_X(t) = n \log(1 - p + p \exp(t))$$

$$E[n_j] = n\pi_j, \quad \text{Var}[n_j] = n\pi_j(1 - \pi_j), \quad \text{Cov}(n_j, n_k) = -n\pi_j\pi_k, (j \neq k)$$

Thus to compute covariance matrix

$$E(X_1 X_2) = \frac{\partial^2 M_X(t)}{\partial t_i \partial t_j} \Big|_{t_i=t_j=0}$$

$$= \frac{\partial \left( n(\pi_i e^{t_i}) (\sum_{k=1}^K \pi_k e^{t_k})^{n-1} \right)'}{\partial t_j}$$

$$= n(n-1) \left( \sum_{k=1}^K \pi_k e^{t_k} \right)^{n-2} \pi_i \pi_j \Big|_{t_i=t_j=0} = n(n-1) \pi_i \pi_j$$

$$E(X_i) = n\pi_i$$

$$\text{Cov}(X_i, X_j) = E(X_i X_j) - E(X_i)E(X_j) = n(n-1) \pi_i \pi_j - n^2 \pi_i \pi_j = -n\pi_i \pi_j$$

$$\text{Var}(X_i) = E(X_i^2) - E(X_i)^2$$

$$E(X_i^2) = \frac{\partial^2 M(t)}{\partial t \partial t} = \frac{\partial \left( n(\pi_i e^{t_i}) (\sum_{k=1}^K \pi_k e^{t_k})^{n-1} \right)'}{\partial t_i}$$

$$= n \left( \sum_{k=1}^K \pi_k e^{t_k} \right)^{n-1} \pi_i e^{t_i} + n(n-1) \left( \sum_{k=1}^K \pi_k e^{t_k} \right)^{n-2} \pi_i \pi_i e^{2t_i} \Big|_{t_i=0}$$

$$= n\pi_i + n(n-1) \pi_i^2 = n\pi_i(1 - \pi_i)$$

$$\text{Var}(X_i/n) = \frac{1}{n^2} \text{Var}(X_i) = \frac{1}{n} \pi_i(1 - \pi_i)$$

Thus the covariance matrix is

$$\Sigma = \begin{bmatrix} \pi_1(1 - \pi_1) & -\pi_1\pi_2 & \dots & -\pi_1\pi_j \\ -\pi_j\pi_i & \pi_i(1 - \pi_i) & \dots & \dots \\ \dots & \dots & \dots & \dots \\ \dots & \dots & \dots & \dots \end{bmatrix}$$

$$= \text{diag}(\pi_j) - \theta\theta^T$$

Here is the question, why do we think the covariance matrix of  $X$  is the covariance matrix of  $\pi$ ?

$$n^{-1}(n_1, n_2, \dots, n_I) = n^{-1} \sum_{i=1}^n [1(X_i = 1), 1(X_i = 2), \dots, 1(X_i = I)]$$

$$= E[1(X_i = 1), 1(X_i = 2), \dots, 1(X_i = I)] = [\pi_1, \pi_2, \dots, \pi_I]$$

## 1.2 Likelihood for multinomial sampling variable in contingency table

$$\begin{aligned}
p(\pi_{ij}) &= \prod_{i=1}^I \prod_{j=1}^J \pi_{ij}^{n_{ij}}, \quad \pi_{ij} > 0, \quad \sum_i \sum_j \pi_{ij} = 1 \\
\theta &= c(\pi_{11}, \pi_{12}, \pi_{21}) \\
\ln(\theta) &= \sum_i \sum_j n_{ij} \log \pi_{ij} = n_{11} \log \pi_{11} + n_{12} \log \pi_{12} + n_{21} \log \pi_{21} + n_{22} \log \pi_{22} \\
&= n_{11} \log \pi_{11} + n_{12} \log \pi_{12} + n_{21} \log \pi_{21} + n_{22} \log(1 - \pi_{11} - \pi_{12} - \pi_{21})
\end{aligned}$$

We can calculate the MLE estimate of  $\pi_{ij}$

$$\begin{aligned}
\frac{\partial \ln(\theta)}{\partial \pi} &= \frac{n_{11}}{\pi_{11}} - \frac{n_{22}}{(1 - \pi_{11} - \pi_{12} - \pi_{21})} = 0, \\
\pi_{11} &= \frac{n_{11}}{n_{22}} \pi_{22}, \quad \pi_{12} = \frac{n_{12}}{n_{22}} \pi_{22}, \quad \pi_{21} = \frac{n_{21}}{n_{22}} \pi_{22}, \quad \pi_{22} = \frac{n_{22}}{n} \\
\pi_{ij} &= \frac{n_{ij}}{n}
\end{aligned}$$

Similarly as above, we need to find the  $Cov(\theta)$ , start from finding  $Var(\pi_{11}, \pi_{12}), Cov(\pi_{11}, \pi_{12})$ .

## 1.3 Pearson Statistics

Question: why the Pearson Statistics use the square of difference between sample mean and expected mean, then divided by the expected mean?

We need to know what is the distribution of the Pearson Statistics. First, we start from the asymptotic distribution of the sample percentage  $\hat{\pi} = \frac{n_i}{n}$ .

$$\begin{aligned}
\sqrt{n} \left( \frac{n_1}{n} - \pi_1, \frac{n_2}{n} - \pi_2, \dots, \frac{n_I}{n} - \pi_I \right) &\xrightarrow{L} N(0, \Sigma^*) \\
\Sigma^* &= diag\{\pi\} - \pi \pi^T
\end{aligned}$$

We need to pay attention that, the  $\pi_1, \pi_2, \dots, \pi_I$  are joint distributed. The Pearson statistics comes from a function of  $(\frac{n_1}{n} - \pi_1, \frac{n_2}{n} - \pi_2, \dots, \frac{n_I}{n} - \pi_I)$ , which could use delta method. The normal distribution is always associated with chi-square distribution.

$$\begin{aligned}
\Gamma &= diag\{\pi_1, \pi_2, \dots, \pi_I\} \\
\sqrt{n} \Gamma^{-1/2} \left( \frac{n_1}{n} - \pi_1, \frac{n_2}{n} - \pi_2, \dots, \frac{n_I}{n} - \pi_I \right) &\xrightarrow{L} N(0, \Gamma^{-1/2} \Sigma^* \Gamma^{-1/2})
\end{aligned}$$

Because  $\Gamma$  is a diagonal matrix, so it could be multiplied directly to the left or right of a matrix, and it only works on the diagonal element.

$$\begin{aligned}
\Gamma^{-1/2}\Sigma^*\Gamma^{-1/2} &= \Gamma^{-1/2}\Gamma^{1/2}(I - \sqrt{\pi}^{\otimes 2})\left(\Gamma^{-1/2}\Gamma^{1/2}\right)^T \\
tr(I - \sqrt{\pi}^{\otimes 2}) &= I - 1 \\
tr(\Gamma^{-1/2}\Sigma^*\Gamma^{-1/2}) &= tr(\Sigma^*\Gamma^{-1/2}\Gamma^{-1/2}) = tr(\Sigma^*\Gamma^{-1}) \\
&= tr([\Gamma - \pi\pi^T]\Gamma^{-1}) = tr(\Gamma\Gamma^{-1}) - tr(\pi\pi^T\Gamma^{-1}) = I - 1
\end{aligned}$$

The Pearson Chi-square statistic is defined as

$$\chi^2 = n \sum_{j=1}^I \left( \frac{n_j}{n} - \pi_j \right)^2 / \pi_j = \left[ \sqrt{n} \Gamma^{-1/2} \left( \frac{n_1}{n} - \pi_1, \frac{n_2}{n} - \pi_2, \dots, \frac{n_I}{n} - \pi_I \right) \right]^{\otimes 2}$$

which converge to  $\chi^2(I - 1)$  as  $n \rightarrow \infty$ .

#### 1.4 Odds ratio

The covariance of odds ratio by delta method. We simplify  $2 \times 2$  table as  $\pi_{11} = \pi_1, \pi_{12} = \pi_2, \pi_{21} = \pi_3, \pi_{22} = \pi_4$ .

$$\begin{aligned}
g(\pi) &= \frac{\pi_{22}\pi_{11}}{\pi_{12}\pi_{21}} \quad \pi = (\pi_{11}, \pi_{12}, \pi_{21}, \pi_{22}) \\
\sqrt{n}(g(\hat{\pi}) - g(\pi)) &\xrightarrow{d} N\left(0, \left(\frac{\partial g(\pi)}{\partial \pi}\right) \Sigma \left(\frac{\partial g(\pi)}{\partial \pi}\right)^T\right) \\
\frac{\partial g(\pi)}{\partial \pi} &= \left( \frac{\partial g}{\partial \pi_{11}}, \frac{\partial g}{\partial \pi_{12}}, \frac{\partial g}{\partial \pi_{21}}, \frac{\partial g}{\partial \pi_{22}} \right)^T \\
&= \left( \frac{\pi_{22}}{\pi_{21}\pi_{12}}, \frac{-\pi_{11}\pi_{22}}{\pi_{21}\pi_{12}^2}, \frac{-\pi_{11}\pi_{22}}{\pi_{12}\pi_{21}^2}, \frac{\pi_{11}}{\pi_{21}\pi_{12}} \right)^T \\
\Sigma^* &= g(\pi)^2 \left( \frac{1}{\pi_{11}} + \frac{1}{\pi_{12}} + \frac{1}{\pi_{21}} + \frac{1}{\pi_{22}} \right)
\end{aligned}$$

So that,

$$Var(\hat{R}) = \frac{1}{n} \Sigma^*$$

We consider  $\log \hat{R}$  instead of  $\hat{R}$ , because  $\log \hat{R}$  converges rapidly to a normal distribution compared to  $\hat{R}$ .

$$\begin{aligned}
\log(\hat{R}) &= \log \pi_1 + \log \pi_2 - \log \pi_3 \log \pi_4 \\
\frac{\partial g(\pi)}{\partial \pi} &= \left( \frac{1}{\pi_{11}}, -\frac{1}{\pi_{12}}, -\frac{1}{\pi_{21}}, \frac{1}{\pi_{22}} \right)^T \\
\text{Var}(\log(\hat{R})) &= \frac{1}{n} \tilde{\Sigma} \\
\tilde{\Sigma} &= \left( \frac{\partial g(\pi)}{\partial \pi} \right)^T \Sigma \left( \frac{\partial g(\pi)}{\partial \pi} \right) \\
\log(\hat{R}) &= \frac{1}{n} \left( \frac{1}{\hat{\pi}_{11}} + \frac{1}{\hat{\pi}_{12}} + \frac{1}{\hat{\pi}_{21}} + \frac{1}{\hat{\pi}_{22}} \right) \\
s.e.\log(\hat{R}) &= \frac{1}{\sqrt{n}} \sqrt{\frac{1}{\hat{\pi}_{11}} + \frac{1}{\hat{\pi}_{12}} + \frac{1}{\hat{\pi}_{21}} + \frac{1}{\hat{\pi}_{22}}}
\end{aligned}$$

## 1.5 Retrospective vs. Prospective vs. Cross Sectional Study

### 1.5.1 Retrospective

For retrospective study, the Y is fixed

$$\begin{aligned}
\theta &= p(X = 1|Y = 1) = \frac{\pi_{11}}{\pi_{11} + \pi_{21}} \\
1 - \theta &= p(X = 0|Y = 1) = \frac{\pi_{21}}{\pi_{11} + \pi_{21}} \\
\gamma &= p(X = 1|Y = 0) = \frac{\pi_{12}}{\pi_{12} + \pi_{22}} \\
1 - \gamma &= p(X = 0|Y = 0) = \frac{\pi_{22}}{\pi_{12} + \pi_{22}}
\end{aligned}$$

$X|Y$  are binomial distribution, which is different from above multinomial distribution. And the  $X|Y = 0, X|Y = 1$  are independent.

$$\begin{aligned}
p(\theta, \gamma) &= \theta^{n_{11}} (1 - \theta)^{n_{21}} \gamma^{n_{12}} (1 - \gamma)^{n_{22}} \\
\ln p(\theta, \gamma) &= n_{11} \log \theta + n_{21} \log(1 - \theta) + n_{12} \log \gamma + n_{22} \log(1 - \gamma) \\
\frac{\partial \ln}{\partial \theta} &= \frac{n_{11}}{\theta} - \frac{n_{21}}{1 - \theta} = 0 \\
\hat{\theta} &= \frac{n_{11}}{n_{11} + n_{21}} \\
\frac{\partial \ln}{\partial \gamma} &= \frac{n_{12}}{\gamma} - \frac{n_{22}}{1 - \gamma} = 0 \\
\hat{\gamma} &= \frac{n_{12}}{n_{12} + n_{22}}
\end{aligned}$$

Then get covariance matrix by delta method, binomial distribution variance is  $np(1-p)$

$$\begin{aligned}
g(\theta) &= \frac{n_{11}n_{22}}{n_{21}n_{12}} = \frac{\theta/(1-\theta)}{\gamma/(1-\gamma)} \\
\sqrt{n}(\theta - \hat{\theta}) &\xrightarrow{d} N(0, \Sigma) \\
\Sigma &= \begin{bmatrix} \theta(1-\theta) & 0 \\ 0 & \gamma(1-\gamma) \end{bmatrix} \\
\sqrt{n}(g(\hat{\theta}) - g(\theta)) &\xrightarrow{d} N(0, g(\theta)' \Sigma^{New} g(\theta)^{rT}) \\
g(\theta)' &= \left( \frac{(1-\gamma)/\gamma}{1/(1-\theta)^2}, \frac{\theta/(1-\theta)}{-1/\gamma^2} \right)
\end{aligned}$$

The standard error for odds ratio in retrospective study

$$\begin{aligned}
se(\hat{R}) &= \hat{R} \sqrt{\frac{1}{n_{.1} \hat{\pi}_{X=2|Y=1} \hat{\pi}_{X=1|Y=1}} + \frac{1}{n_{.2} \hat{\pi}_{X=2|Y=2} \hat{\pi}_{X=1|Y=2}}} \\
\hat{\pi}_{X=2|Y=1} &= \frac{n_{21}}{n_{11} + n_{21}} \\
\hat{\pi}_{X=1|Y=1} &= \frac{n_{11}}{n_{11} + n_{21}} \\
\hat{\pi}_{X=2|Y=2} &= \frac{n_{12}}{n_{12} + n_{22}} \\
\hat{\pi}_{X=1|Y=2} &= \frac{n_{12}}{n_{12} + n_{22}} \\
n_{.1} &= n_{11} + n_{21}, \quad n_{.2} = n_{12} + n_{22} \\
se(\hat{R}) &= \frac{n_{22}n_{11}}{(n_{21}n_{12})} \sqrt{\frac{n_{11} + n_{21}}{n_{11}n_{21}} + \frac{n_{12} + n_{22}}{n_{12}n_{22}}} \\
&= \frac{n_{22}n_{11}}{(n_{21}n_{12})} \sqrt{\frac{1}{n_{11}} + \frac{1}{n_{12}} + \frac{1}{n_{21}} + \frac{1}{n_{22}}}
\end{aligned}$$

### 1.5.2 Prospective

The standard error for odds ratio in prospective study

$$se(\hat{R}) = \hat{R} \sqrt{\frac{1}{n_{1.} \hat{\pi}_{Y=2|X=1} \hat{\pi}_{Y=1|X=1}} + \frac{1}{n_{2.} \hat{\pi}_{Y=2|X=2} \hat{\pi}_{Y=1|X=2}}}$$

$$\hat{\pi}_{Y=2|X=1} = \frac{n_{12}}{n_{11} + n_{12}}$$

$$\hat{\pi}_{Y=1|X=1} = \frac{n_{11}}{n_{11} + n_{12}}$$

$$\hat{\pi}_{Y=2|X=2} = \frac{n_{22}}{n_{21} + n_{22}}$$

$$\hat{\pi}_{Y=1|X=2} = \frac{n_{21}}{n_{21} + n_{22}}$$

$$n_{1.} = n_{11} + n_{12}, \quad n_{2.} = n_{21} + n_{22}$$

$$se(\hat{R}) = \frac{n_{22}n_{11}}{(n_{21}n_{12})} \sqrt{\frac{n_{11} + n_{12}}{n_{11}n_{12}} + \frac{n_{21} + n_{22}}{n_{21}n_{22}}}$$

$$= \frac{n_{22}n_{11}}{(n_{21}n_{12})} \sqrt{\frac{1}{n_{11}} + \frac{1}{n_{12}} + \frac{1}{n_{21}} + \frac{1}{n_{22}}}$$

### 1.5.3 Cross-Sectional

For cross-sectional study, we only have the total n fixed. That is the difference for each scenario.

To calculate the covariance matrix, we will use the MGF and take derivatives. Or use the cumulant function KGF to get the covariance.

Use one random variable for the two way contingency table. While the Fisher information is the inverse of the covariance matrix, however we don't use Fisher information to calculate covariance matrix due to the math computation.

Show that the sample odds ratio  $\hat{R} = n_{22}n_{11}/(n_{21}n_{12})$  has the same standard error for cross-sectional, prospective and retrospective studies.

The standard error for odds ratio in cross sectional study

$$se(\hat{R}) = \frac{\hat{R}}{\sqrt{n}} \sqrt{\frac{1}{\hat{\pi}_{11}} + \frac{1}{\hat{\pi}_{12}} + \frac{1}{\hat{\pi}_{21}} + \frac{1}{\hat{\pi}_{22}}}$$

$$= \frac{n_{22}n_{11}}{(n_{21}n_{12})} \sqrt{\frac{1}{n_{11}} + \frac{1}{n_{12}} + \frac{1}{n_{21}} + \frac{1}{n_{22}}}$$

By comparing the above standard errors in three types of studies, we see that they have same standard errors. Odds ratio is invariant in terms of sampling method. Similarly the coefficient of a particular covariate is associated with the odds ratio of the covariate, which is invariant with prospective and retrospective studies. Check out p747.

## 1.6 Hypergeometric distribution

Derive the hypergeometric distribution

$$\begin{aligned}
 p(n_{11}|n_{1.}, n_{.1}, n, \Xi) &= \frac{p(n_{11}, n_{1.}, n_{.1}, |n)}{p(n_{1.}, n_{.1}, |n)} \\
 &= \frac{n!}{n_{11}!n_{12}!n_{21}!n_{22}!} \Xi^{n_{11}} \frac{n!}{n_{11}!n_{12}!n_{21}!n_{22}!} \\
 &= \frac{n!n_{1.}!(n - n_{1.})!}{n_{1.}!(n - n_{1.})!n_{11}!n_{12}!n_{21}!n_{22}!} \\
 &= \binom{n}{n_{1.}} \binom{n_{1.}}{n_{11}} \binom{n - n_{1.}}{n_{.1} - n_{11}}
 \end{aligned}$$

## 1.7 Contingency Table- Relationship between Poisson and Multinomial distribution

Consider a  $I \times J$  contingency table of cell counts, where each cell count is denoted by  $n_{ij}, i = 1, \dots, I, j = 1, \dots, J$ , and thus  $n_{ij}$  denotes the cell count of  $i$ th row and  $j$ th column, and  $n_{ij} \sim \text{Poisson}(\mu_{ij})$  and independent. Further, let  $n = \sum_{j=1}^J \sum_{i=1}^I n_{ij}$  denote the grand total.

- (a) Derive the joint distribution of  $(n_{11}, n_{12}, \dots, n_{IJ})$  conditional on grand total  $n$ . By poisson distribution of each cell counts

$$\begin{aligned}
 n = \sum_{i=1}^I \sum_{j=1}^J n_{ij} &\sim \frac{\exp(-\mu) \mu^n}{n!}, \quad \mu = \sum_{i=1}^I \sum_{j=1}^J \mu_{ij} \\
 p(n_{11}, \dots, n_{IJ}|n) &= \frac{\prod_{i=1}^I \prod_{j=1}^J \frac{\exp(-\mu_{ij}) \mu_{ij}^{n_{ij}}}{n_{ij}!}}{\frac{\exp(-\mu) \mu^n}{n!}} \\
 &= \binom{n}{n_{11}n_{12}\dots n_{IJ}} \frac{\prod_{i=1}^I \prod_{j=1}^J \mu_{ij}^{n_{ij}}}{\mu^n} \\
 &= \binom{n}{n_{11}n_{12}\dots n_{IJ}} \prod_{i=1}^I \prod_{j=1}^J \left( \frac{\mu_{ij}}{\mu} \right)^{n_{ij}}
 \end{aligned}$$

The joint distribution is Multinomial  $(n; \pi_{11}, \pi_{12}, \dots, \pi_{IJ})$ , where  $\pi_{ij} = \frac{\mu_{ij}}{\sum_{i=1}^I \sum_{j=1}^J \mu_{ij}}$

- (b) Suppose all of the rows margins are assumed fixed. Derive the joint distribution



of  $(n_{11}, n_{12}, \dots, n_{ij})$ .

$$\begin{aligned}
n_{i+} &= \sum_{j=1}^J n_{ij} \\
n_{i+} &\sim \text{Poisson}\left(\sum_{j=1}^J \mu_{ij}\right) \\
p(n_{11}, \dots, n_{ij} | n_{i+}) &= \frac{\prod_{i=1}^I \prod_{j=1}^J \frac{\exp(-\mu_{ij}) \mu_{ij}^{n_{ij}}}{n_{ij}!}}{\prod_{i=1}^I \frac{\exp(-\mu_i) \mu_i^{n_{i+}}}{n_{i+}!}} \\
&= \prod_{i=1}^I \binom{n_{i+}}{n_{ij}} \prod_{i=1}^I \prod_{j=1}^J \left( \frac{\mu_{ij}}{\sum_{j=1}^J \mu_{ij}} \right)^{n_{ij}}
\end{aligned}$$

- (c) Suppose all of the columns margins are assumed fixed. Derive the joint distribution of  $(n_{11}, n_{12}, \dots, n_{ij})$ .

$$\begin{aligned}
n_{+j} &= \sum_{i=1}^I n_{ij} \\
n_{+j} &\sim \text{Poisson}\left(\sum_{i=1}^I \mu_{ij}\right) \\
p(n_{11}, \dots, n_{ij} | n_{+j}) &= \frac{\prod_{i=1}^I \prod_{j=1}^J \frac{\exp(-\mu_{ij}) \mu_{ij}^{n_{ij}}}{n_{ij}!}}{\prod_{j=1}^J \frac{\exp(-\mu_j) \mu_j^{n_{+j}}}{n_{+j}!}} \\
&= \prod_{j=1}^J \binom{n_{+j}}{n_{ij}} \prod_{i=1}^I \prod_{j=1}^J \left( \frac{\mu_{ij}}{\sum_{i=1}^I \mu_{ij}} \right)^{n_{ij}}
\end{aligned}$$

- (d) Suppose that  $I = 2$  and  $J = 2$ , and both the rows margins and column margins are fixed. Derive the joint distribution of  $(n_{11} | n_{1+}, n_{+1}n)$ , where  $n_{1+} = n_{11} + n_{12}$ ,  $n_{+1} = n_{11} + n_{21}$ .

$$\begin{aligned}
p(n_{11} | n_{1+}, n_{+1}n) &= \frac{p(n_{11}, n_{1+}, n_{+1}n)}{p(n_{1+}, n_{+1}n)} \\
p(n_{ij}) &= \prod_{i=1}^2 \prod_{j=1}^2 \frac{\exp(-\mu_{ij}) \mu_{ij}^{n_{ij}}}{n_{ij}!} \\
&= \frac{\exp(-\mu_{11}) \mu_{11}^{n_{11}}}{n_{11}!} \frac{\exp(-\mu_{12}) \mu_{12}^{n_{12}}}{n_{12}!} \frac{\exp(-\mu_{21}) \mu_{21}^{n_{21}}}{n_{21}!} \frac{\exp(-\mu_{22}) \mu_{22}^{n_{22}}}{n_{22}!} \\
n_{12} &= n_{1+} - n_{11}, \quad n_{21} = n_{+1} - n_{11}, \\
n_{22} &= n - n_{12} - n_{21} - n_{11} = n - n_{1+} - n_{+1} + n_{11} \\
p(n_{11}, n_{1+}, n_{+1}n) &= \frac{\exp(-\mu_{11}) \mu_{11}^{n_{11}}}{n_{11}!} \frac{\exp(-\mu_{12}) \mu_{12}^{n_{1+} - n_{11}}}{(n_{1+} - n_{11})!} \frac{\exp(-\mu_{21}) \mu_{21}^{n_{+1} - n_{11}}}{(n_{+1} - n_{11})!} \frac{\exp(-\mu_{22}) \mu_{22}^{n - n_{1+} - n_{+1} + n_{11}}}{(n - n_{1+} - n_{+1} + n_{11})!}
\end{aligned}$$

The Jacobian transformation matrix

$$J = \begin{pmatrix} \frac{\partial n_{11}}{\partial n_{11}} & \frac{\partial n_{11}}{\partial n_{12}} & \frac{\partial n_{11}}{\partial n_{21}} & \frac{\partial n_{11}}{\partial n_{22}} \\ \frac{\partial n_{12}}{\partial n_{11}} & \frac{\partial n_{12}}{\partial n_{12}} & \frac{\partial n_{12}}{\partial n_{21}} & \frac{\partial n_{12}}{\partial n_{22}} \\ \frac{\partial n_{21}}{\partial n_{11}} & \frac{\partial n_{21}}{\partial n_{12}} & \frac{\partial n_{21}}{\partial n_{21}} & \frac{\partial n_{21}}{\partial n_{22}} \\ \frac{\partial n_{22}}{\partial n_{11}} & \frac{\partial n_{22}}{\partial n_{12}} & \frac{\partial n_{22}}{\partial n_{21}} & \frac{\partial n_{22}}{\partial n_{22}} \end{pmatrix} = \begin{pmatrix} 1 & 0 & 0 & 0 \\ -1 & 1 & 0 & 0 \\ -1 & 0 & 1 & 0 \\ 1 & -1 & -1 & 1 \end{pmatrix}$$

$$\|J\| = 1$$

Then we can get the  $p(n_{1+}, n_{+1}, n)$  by summing over  $n_{11}$ . We have  $n_{11} \leq n_{1+}$ ,  $n_{11} \leq n_{+1}$ , and  $n_{11} \geq -n + n_{1+} + n_{+1}$ .

$$\begin{aligned} p(n_{11}, n_{1+}, n_{+1}n) &= \frac{\exp(-\mu_{11})\mu_{11}^{n_{11}}}{n_{11}!} \frac{\exp(-\mu_{12})\mu_{12}^{n_{1+}-n_{11}}}{(n_{1+}-n_{11})!} \frac{\exp(-\mu_{21})\mu_{21}^{n_{+1}-n_{11}}}{(n_{+1}-n_{11})!} \frac{\exp(-\mu_{22})\mu_{22}^{n-n_{1+}-n_{+1}+n_{11}}}{(n-n_{1+}-n_{+1}+n_{11})!} \\ &= \frac{\exp(-\sum_{i=1}^2 \sum_{j=1}^2 \mu_{ij}) \left(\frac{\mu_{11}\mu_{22}}{\mu_{12}\mu_{21}}\right)^{n_{11}} \left(\frac{\mu_{12}}{\mu_{22}}\right)^{n_{1+}} \left(\frac{\mu_{21}}{\mu_{22}}\right)^{n_{+1}} \mu_{22}^n}{n_{11}!(n_{1+}-n_{11})!(n_{+1}-n_{11})!(n-n_{1+}-n_{+1}+n_{11})!} \\ p(n_{1+}, n_{+1}n) &= \sum_{\max(0, -n+n_{1+}+n_{+1})}^{\min(n_{1+}, n_{+1})} \frac{\exp(-\sum_{i=1}^2 \sum_{j=1}^2 \mu_{ij}) \left(\frac{\mu_{11}\mu_{22}}{\mu_{12}\mu_{21}}\right)^{n_{11}} \left(\frac{\mu_{12}}{\mu_{22}}\right)^{n_{1+}} \left(\frac{\mu_{21}}{\mu_{22}}\right)^{n_{+1}} \mu_{22}^n}{n_{11}!(n_{1+}-n_{11})!(n_{+1}-n_{11})!(n-n_{1+}-n_{+1}+n_{11})!} \end{aligned}$$

So we can have

$$\begin{aligned} p(n_{11}|n_{1+}, n_{+1}n) &= \frac{p(n_{11}, n_{1+}, n_{+1}n)}{p(n_{1+}, n_{+1}n)} \\ &= \frac{\exp(-\sum_{i=1}^2 \sum_{j=1}^2 \mu_{ij}) \left(\frac{\mu_{11}\mu_{22}}{\mu_{12}\mu_{21}}\right)^{n_{11}} \left(\frac{\mu_{12}}{\mu_{22}}\right)^{n_{1+}} \left(\frac{\mu_{21}}{\mu_{22}}\right)^{n_{+1}} \mu_{22}^n}{n_{11}!(n_{1+}-n_{11})!(n_{+1}-n_{11})!(n-n_{1+}-n_{+1}+n_{11})!} \\ &\quad \Bigg/ \sum_{\max(0, -n+n_{1+}+n_{+1})}^{\min(n_{1+}, n_{+1})} \frac{\exp(-\sum_{i=1}^2 \sum_{j=1}^2 \mu_{ij}) \left(\frac{\mu_{11}\mu_{22}}{\mu_{12}\mu_{21}}\right)^{n_{11}} \left(\frac{\mu_{12}}{\mu_{22}}\right)^{n_{1+}} \left(\frac{\mu_{21}}{\mu_{22}}\right)^{n_{+1}} \mu_{22}^n}{n_{11}!(n_{1+}-n_{11})!(n_{+1}-n_{11})!(n-n_{1+}-n_{+1}+n_{11})!} \end{aligned}$$

Which we can rewrite

$$\begin{aligned} p(n_{11}|n_{1+}, n_{+1}n) &= \binom{n_{1+}}{n_{11}} \binom{n-n_{1+}}{n_{+1}-n_{11}} \left(\frac{\pi_{11}\pi_{22}}{\pi_{12}\pi_{21}}\right)^{n_{11}} \\ &\quad \Bigg/ \sum_{x=\max(0, -n+n_{1+}+n_{+1})}^{\min(n_{1+}, n_{+1})} \binom{n_{1+}}{x} \binom{n-n_{1+}}{n_{+1}-x} \left(\frac{\pi_{11}\pi_{22}}{\pi_{12}\pi_{21}}\right)^x \end{aligned}$$

- (e) Let  $\pi_{ij}$  denote the cell probability and assume  $n$  is fixed. Consider testing  $H_0$  :  $\pi_{ij} = \pi_{i+}\pi_{+j}$ ,  $i = 1, \dots, I$ ,  $j = 1, \dots, J$ . Derive the MLE of  $\pi_{ij}$  under  $H_0$ .

The  $H_0$  could be written as

$$H_0 : \pi_{ij} = \pi_{i+}\pi_{+j}$$

The multinomial distribution of  $\pi_{ij}$

$$p(\pi_{ij}) = \binom{n}{n_{11}n_{12}n_{21}n_{22}} \pi_{ij}^{n_{ij}}, \sum_{i=1}^I \sum_{j=1}^J \pi_{ij} = 1$$

The log-likelihood function

$$\log p(\pi_{ij}) = \log \binom{n}{n_{11}n_{12}n_{21}n_{22}} + n_{ij} \log \pi_{ij}, \sum_{i=1}^I \sum_{j=1}^J \pi_{ij} = 1$$

Under  $H_0$ , the log-likelihood

$$\log p(\pi_{ij}) = \log \binom{n}{n_{11}n_{12}n_{21}n_{22}} + n_{ij} \log \pi_{i+}\pi_{+j}, \sum_{i=1}^I \pi_{i+} = 1, \sum_{j=1}^J \pi_{+j} = 1$$

By Lagrangian multiplier theorem,

$$\begin{aligned} \ln(\pi_{ij}) &= n \log \binom{n}{n_{11}n_{12}n_{21}n_{22}} + \sum_{i=1}^I \sum_{j=1}^J n_{ij} \log \pi_{i+}\pi_{+j} + \lambda \left( \sum_{i=1}^I \sum_{j=1}^J \pi_{ij} - 1 \right), \\ &= n \log \binom{n}{n_{11}n_{12}n_{21}n_{22}} + \sum_{i=1}^I \sum_{j=1}^J n_{ij} \log \pi_{i+} + \sum_{j=1}^J \sum_{i=1}^I n_{ij} \log \pi_{+j} - \lambda \left( \sum_{i=1}^I \pi_{i+} - 1 \right) \end{aligned}$$

Take first derivative of log-likelihood

$$\begin{aligned} \frac{\partial \ln}{\partial \pi_{i+}} &= \frac{\sum_{j=1}^J n_{ij}}{\pi_{i+}} + \lambda = 0 \\ \hat{\pi}_{i+} &= \frac{\sum_{j=1}^J n_{ij}}{\lambda} \\ \sum_{i=1}^I \pi_{i+} &= 1, \quad \lambda = \sum_{j=1}^J \sum_{i=1}^I n_{ij} \\ \hat{\pi}_{i+} &= \frac{n_{i+}}{n} \end{aligned}$$

Similarly, we have  $\hat{\pi}_{+j} = \frac{n_{+j}}{n}$ , the MLE of  $\pi_{ij}$  under  $H_0$  is

$$\hat{\pi}_{ij} = \hat{\pi}_{i+}\hat{\pi}_{+j} = \frac{n_{i+}n_{+j}}{n^2}$$

- (f) Derive the likelihood ratio test for the hypothesis in part (e) and derive its asymptotic distribution under  $H_0$ . From part (e), we have the parameter estimates under  $H_0$ . While under alternative hypothesis, we have  $\mu_{ij} = n_{ij}$ .

$$\begin{aligned} LRT_n &= 2(LR(\pi_{H_1}) - LR(\pi_{H_0})) = 2 \left( \sum_{i=1}^I \sum_{j=1}^J n_{ij} \log \pi_{ij} - \sum_{i=1}^I \sum_{j=1}^J n_{ij} \log \pi_{i+} \pi_{+j} \right) \\ &= 2 \left( \sum_{i=1}^I \sum_{j=1}^J n_{ij} \log \frac{\pi_{ij}}{\pi_{i+} \pi_{+j}} \right) \\ &= 2 \left( \sum_{i=1}^I \sum_{j=1}^J n_{ij} \log \frac{n_{ij} n}{n_{i+} n_{+j}} \right) \sim \chi_{(I-1)(J-1)}^2 \end{aligned}$$

Note that the full model has  $(IJ - 1)$  parameters, and the null hypothesis has  $(I - 1) + (J - 1)$  parameters.

$$\begin{aligned} df &= I \times J - 1 - (I - 1) - (J - 1) \\ &= (I - 1)(J - 1) \end{aligned}$$

- (g) Suppose that  $\pi_{11}, \pi_{12}$  are parameters of interest and the rest of the parameters are treated as nuisance. Derive the conditional likelihood of  $(\pi_{11}, \pi_{12})$  and the conditional MLE's of  $(\pi_{11}, \pi_{12})$ . If not specified, we treat as general contingency table that total  $n$  is fixed. If only  $\pi_{11}, \pi_{12}$  are parameters of interest and the rest of the parameters are treated as nuisance, then we will set the rest of the parameters as one parameter, and get its distribution, which is to find the sufficient statistics for rest of the parameters. Write the Multinomial distribution in exponential family distribution.

We can find marginal distribution by summing over along all possible values of  $(n_{11}, n_{12})$ . Note that  $n_{11} \leq \min n_{1+} - n_{12}, n_{+1}$  for a given value of  $n_{12}$ . Similarly,  $n_{12} \leq \min n_{1+} - n_{11}, n_{+1}$  for a given value of  $n_{11}$ .

Additionally,

$$\begin{aligned} n &\geq n_{1+} + n_{+1} + n_{+2} - n_{11} - n_{12} \\ n_{11} + n_{12} &\geq \max 0, n_{+1} + n_{1+} + n_{+2} \end{aligned}$$

Let

$$\begin{aligned} S(n_{11}, n_{12}) &= \{(n_{11}, n_{12}) : n_{11} + n_{12} \geq \max 0, n_{+1} + n_{1+} + n_{+2}, \\ &\quad n_{11} \leq \min(n_{1+} - n_{12}, n_{+1}), n_{12} \leq \min(n_{1+} - n_{11}, n_{+1})\} \end{aligned}$$

The conditional distribution

$$\begin{aligned} p(n_{11}, n_{12} | n_{13}, \dots, n_{IJ}, n) &= \frac{p(n_{ij})}{p(S_n)} \\ &= \frac{\frac{1}{n_{11}! n_{12}!} \pi_{11}^{n_{11}} \pi_{12}^{n_{12}}}{\sum_{(x,y) \in S_n} \frac{1}{x! y!} \pi_{11}^x \pi_{12}^y} \end{aligned}$$

And  $\hat{\pi}_{11}, \hat{\pi}_{12}$  are the CMLE that maximize  $p(n_{11}, n_{12} | n_{13}, \dots, n_{IJ}, n)$ .

## 2 Practice

### 2.1 Contingency table parameters

- (a) Get MLE of  $\pi$  and prove CLT.

The multinomial distribution based on total  $n$ .

$$p(\theta) = n! \prod_{i=0}^1 \prod_{j=0}^1 \frac{\pi_{ij}^{n_{ij}}}{n_{ij}!}, \quad \theta = (\pi_{00}, \pi_{01}, \pi_{10}, \pi_{11})^T$$

$$\ln p(\theta) = \ln n! + \sum_{i=0}^1 \sum_{j=0}^1 n_{ij} \log(\pi_{ij}) - \log n_{ij}!$$

$$= \ln n! + n_{00} \log \pi_{00} + n_{01} \log \pi_{01} + n_{10} \log \pi_{10} + n_{11} \log(1 - \pi_{00} - \pi_{01} - \pi_{10})$$

The MLE of the  $\theta$  by taking derivative to the log-likelihood

$$\begin{aligned} \frac{\partial \ln(\theta)}{\partial \pi_{00}} &= \frac{n_{00}}{\pi_{00}} - \frac{n_{11}}{1 - \pi_{00} - \pi_{01} - \pi_{10}} = 0 \\ \frac{\partial \ln(\theta)}{\partial \pi_{01}} &= \frac{n_{01}}{\pi_{01}} - \frac{n_{11}}{1 - \pi_{00} - \pi_{01} - \pi_{10}} = 0 \\ \frac{\partial \ln(\theta)}{\partial \pi_{10}} &= \frac{n_{10}}{\pi_{10}} - \frac{n_{11}}{1 - \pi_{00} - \pi_{01} - \pi_{10}} = 0 \\ \hat{\pi}_{00} &= \frac{n_{00}}{n} \\ \hat{\pi}_{01} &= \frac{n_{01}}{n} \\ \hat{\pi}_{10} &= \frac{n_{10}}{n} \\ \hat{\pi}_{11} &= \frac{n_{11}}{n}, \quad n = n_{00} + n_{01} + n_{10} + n_{11} \end{aligned}$$

Let  $Z_i = I(X = x, Y = y) \sim \text{multi}(1, \pi_{00}, \pi_{01}, \pi_{10}, \pi_{11})$ .

$$Z_1 = I[(X, Y) = (0, 0)]$$

$$Z_2 = I[(X, Y) = (0, 1)]$$

$$Z_3 = I[(X, Y) = (1, 0)]$$

$$Z_4 = I[(X, Y) = (1, 1)]$$

$$p(\theta) = \prod_k \pi_k^{I(Z_k=1)}$$

$$M_Z(t) = E[\exp(t^T Z)] = E[\exp(t^T (Z_1 + Z_2 + \dots Z_n))] = E[\exp(t^T Z_1 + t^T Z_2 + \dots t^T Z_n)]$$

$$= E\left[\prod_{i=1}^n \exp(t^T Z_i)\right]$$

$$= \prod_{i=1}^n E[\exp(t^T Z_i)] \quad (\text{by independence})$$

$$= \prod_{i=1}^n M_{Z_i}(t) = \prod_{i=1}^n P(Z_i = 1) e^{t z_i} \quad \text{by MGF of discrete variable } Z_i$$

$$= \left( \sum_{j=1}^J \pi_j \exp(t_j) \right)^n \quad \text{by MGF of multinoulli}$$

Then the covariance matrix of  $\theta$  could be calculated by MGF.

$$\begin{aligned} E(Z_1 Z_2) &= \frac{\partial^2 M_Z(t)}{\partial Z_i \partial Z_j} \Big|_{t_i=t_j=0} \\ &= \frac{\partial \left( n(\pi_i e^{t_i}) (\sum_{k=1}^K \pi_k e^{t_k})^{n-1} \right)'}{\partial t_j} \\ &= n(n-1) \left( \sum_{k=1}^K \pi_k e^{t_k} \right)^{n-2} \pi_i \pi_j \Big|_{t_i=t_j=0} = n(n-1) \pi_i \pi_j \end{aligned}$$

$$E(X_i) = n\pi_i$$

$$\text{Cov}(Z_i, Z_j) = E(Z_i Z_j) - E(Z_i)E(Z_j) = n(n-1)\pi_i \pi_j - n^2 \pi_i \pi_j = -n\pi_i \pi_j$$

$$\text{Var}(Z_i) = E(Z_i^2) - E(Z_i)^2$$

$$E(Z_i^2) = \frac{\partial \left( n(\pi_i e^{t_i}) (\sum_{k=1}^K \pi_k e^{t_k})^{n-1} \right)'}{\partial t_i}$$

$$= n \left( \sum_{k=1}^K \pi_k e^{t_k} \right)^{n-1} \pi_i e^{t_i} + n(n-1) \left( \sum_{k=1}^K \pi_k e^{t_k} \right)^{n-2} \pi_i \pi_i e^{2t_i} \Big|_{t_i=0}$$

$$= n\pi_i + n(n-1)\pi_i^2 = n\pi_i(1 - \pi_i)$$

$$\text{Var}(Z_i/n) = \frac{1}{n^2} \text{Var}(Z_i) = \frac{1}{n} \pi_i(1 - \pi_i)$$

Thus the covariance matrix is

$$\Sigma = \begin{bmatrix} \pi_{00}(1 - \pi_{00}) & -\pi_{00}\pi_{01} & -\pi_{00}\pi_{10} & -\pi_{00}\pi_{11} \\ -\pi_{01}\pi_{00} & \pi_{01}(1 - \pi_{01}) & -\pi_{01}\pi_{10} & -\pi_{01}\pi_{11} \\ -\pi_{10}\pi_{00} & -\pi_{10}\pi_{01} & \pi_{10}(1 - \pi_{10}) & -\pi_{10}\pi_{11} \\ -\pi_{11}\pi_{00} & -\pi_{11}\pi_{01} & -\pi_{11}\pi_{10} & \pi_{11}(1 - \pi_{11}) \end{bmatrix} = \text{diag}(\pi_{ij}) - \theta\theta^T$$

By Central limit theroem,

$$\sqrt{n}(\hat{\pi}_{00} - \pi_{00}, \hat{\pi}_{01} - \pi_{01}, \hat{\pi}_{10} - \pi_{10}, \hat{\pi}_{11} - \pi_{11})^T \xrightarrow{d} N(0, \Sigma)$$

- (b) Let R denote the odds ratio. Find the maximum likelihood estimate of log(R) and derive its asymptotic distribution.

By invariance of MLE:

$$\begin{aligned} R &= \frac{\pi_{00}\pi_{11}}{\pi_{01}\pi_{10}} \\ g(R) &= \log R = \log \pi_{00} + \log \pi_{11} - \log \pi_{01} - \log \pi_{10} \\ \log \hat{R} &= \log \hat{\pi}_{00} + \log \hat{\pi}_{11} - \log \hat{\pi}_{01} - \log \hat{\pi}_{10} \\ &= \log \frac{n_{00}n_{11}}{n_{01}n_{10}} \end{aligned}$$

By Central limit theorem, we have

$$\sqrt{n}(g(\hat{R}) - g(R)) \xrightarrow{d} N\left(0, \frac{\partial g(R)}{\partial \theta} \Sigma \frac{\partial g(R)}{\partial \theta}^T\right)$$

By delta method,

$$\begin{aligned} \frac{\partial g(R)}{\partial \theta} &= \left( \frac{1}{R} \frac{\partial R}{\partial \pi_{00}}, \frac{1}{R} \frac{\partial R}{\partial \pi_{01}}, \frac{1}{R} \frac{\partial R}{\partial \pi_{10}}, \frac{1}{R} \frac{\partial R}{\partial \pi_{11}} \right) \\ &= \left( \frac{1}{\pi_{00}}, -\frac{1}{\pi_{01}}, -\frac{1}{\pi_{10}}, \frac{1}{\pi_{11}} \right) \\ \Sigma^R &= \frac{\partial g(R)}{\partial \theta} \Sigma \frac{\partial g(R)}{\partial \theta}' \\ &= \left( \frac{1}{\pi_{00}}, -\frac{1}{\pi_{01}}, -\frac{1}{\pi_{10}}, \frac{1}{\pi_{11}} \right) \begin{bmatrix} \pi_{00}(1 - \pi_{00}) & -\pi_{00}\pi_{01} & -\pi_{00}\pi_{10} & -\pi_{00}\pi_{11} \\ -\pi_{01}\pi_{00} & \pi_{01}(1 - \pi_{01}) & -\pi_{01}\pi_{10} & -\pi_{01}\pi_{11} \\ -\pi_{10}\pi_{00} & -\pi_{10}\pi_{01} & \pi_{10}(1 - \pi_{10}) & -\pi_{10}\pi_{11} \\ -\pi_{11}\pi_{00} & -\pi_{11}\pi_{01} & -\pi_{11}\pi_{10} & \pi_{11}(1 - \pi_{11}) \end{bmatrix} \begin{bmatrix} \frac{1}{\pi_{00}} \\ -\frac{1}{\pi_{01}} \\ -\frac{1}{\pi_{10}} \\ \frac{1}{\pi_{11}} \end{bmatrix} \\ &= \left( \frac{1}{\pi_{00}} + \frac{1}{\pi_{01}} + \frac{1}{\pi_{10}} + \frac{1}{\pi_{11}} \right) \end{aligned}$$

We have the asymptotic distribution of  $\log(R)$

$$\sqrt{n}(\log \hat{R} - \log R) \xrightarrow{d} N\left(0, \left(\frac{1}{\pi_{11}} + \frac{1}{\pi_{12}} + \frac{1}{\pi_{21}} + \frac{1}{\pi_{22}}\right)\right)$$

- (c) Construct an approximate 95% confidence interval for the odds ratio  $R$ .

From part (b), we have the asymptotic normal distribution of  $\log R$ . We have the asymptotic distribution of  $R$ .

$$\begin{aligned} f &= \exp(g) = R, & f(g)' &= R \\ \sqrt{n}(f(\hat{g}) - f(g)) &\xrightarrow{d} N\left(0, f(g)' \left(\frac{1}{\pi_{11}} + \frac{1}{\pi_{12}} + \frac{1}{\pi_{21}} + \frac{1}{\pi_{22}}\right) f(g)^{T'}\right) \\ \sqrt{n}(\hat{R} - R) &\xrightarrow{d} N\left(0, R^2 \left(\frac{1}{\pi_{11}} + \frac{1}{\pi_{12}} + \frac{1}{\pi_{21}} + \frac{1}{\pi_{22}}\right)\right) \\ (\hat{R} - R) &\xrightarrow{d} N\left(0, \frac{1}{n} R^2 \left(\frac{1}{\pi_{11}} + \frac{1}{\pi_{12}} + \frac{1}{\pi_{21}} + \frac{1}{\pi_{22}}\right)\right) \end{aligned}$$

The 95% confidence interval for the odds ratio  $R$

$$\{R : \hat{R} - 1.96\hat{R}\sqrt{\frac{1}{\pi_{11}} + \frac{1}{\pi_{12}} + \frac{1}{\pi_{21}} + \frac{1}{\pi_{22}}} \leq R \leq \hat{R} + 1.96\hat{R}\sqrt{\frac{1}{\pi_{11}} + \frac{1}{\pi_{12}} + \frac{1}{\pi_{21}} + \frac{1}{\pi_{22}}}\}$$

- (d) Under the assumptions of part (a), further assume that  $\pi_{1+} = \pi_{11} + \pi_{10} = \frac{\exp(\alpha)}{1+\exp(\alpha)}$  and  $\pi_{+1} = \pi_{11} + \pi_{01} = \frac{\exp(\alpha+\beta)}{1+\exp(\alpha+\beta)}$ . Derive the maximum likelihood estimates of  $(\alpha, \beta)$ , denoted by  $(\hat{\alpha}; \hat{\beta})$ .

$$\begin{aligned} \pi_{01} + \pi_{11} &= \frac{\exp(\alpha)}{1 + \exp(\alpha)} \\ \exp(\alpha) &= \frac{\pi_{10} + \pi_{11}}{\pi_{01} + \pi_{00}}, & \alpha &= \log\left(\frac{\pi_{10} + \pi_{11}}{\pi_{01} + \pi_{00}}\right) \\ \pi_{10} + \pi_{11} &= \frac{\exp(\alpha + \beta)}{1 + \exp(\alpha + \beta)} \\ \alpha + \beta &= \log\left(\frac{\pi_{01} + \pi_{11}}{\pi_{10} + \pi_{00}}\right) \\ \beta &= \log\left(\frac{\pi_{01} + \pi_{11}}{\pi_{10} + \pi_{00}}\right) - \log\frac{\pi_{10} + \pi_{11}}{\pi_{01} + \pi_{00}}, & \beta &= \log\left(\frac{(\pi_{01} + \pi_{11})(\pi_{01} + \pi_{00})}{(\pi_{10} + \pi_{00})(\pi_{10} + \pi_{11})}\right) \end{aligned}$$

By invariance of MLE,

$$\begin{aligned} \hat{\alpha} &= \log\left(\frac{\hat{\pi}_{10} + \hat{\pi}_{11}}{\hat{\pi}_{01} + \hat{\pi}_{00}}\right) = \log\left(\frac{n_{10} + n_{11}}{n_{01} + n_{00}}\right) \\ \hat{\beta} &= \log\left(\frac{(\hat{\pi}_{01} + \hat{\pi}_{11})(\hat{\pi}_{01} + \hat{\pi}_{00})}{(\hat{\pi}_{10} + \hat{\pi}_{00})(\hat{\pi}_{10} + \hat{\pi}_{11})}\right) = \log\left(\frac{(n_{01} + n_{11})(n_{01} + n_{00})}{(n_{10} + n_{00})(n_{10} + n_{11})}\right) \end{aligned}$$



- (e) Using the assumptions of part (d), derive the asymptotic distribution of  $(\alpha, \beta)$  (properly normalized).

By Central limit theorem and delta method,

$$\begin{aligned}\xi &= (\alpha, \beta)^T \\ g(\xi) &= \left\{ \log \left( \frac{\pi_{10} + \pi_{11}}{\pi_{01} + \pi_{00}} \right), \log \left( \frac{(\pi_{01} + \pi_{11})(\pi_{01} + \pi_{00})}{(\pi_{10} + \pi_{00})(\pi_{10} + \pi_{11})} \right) \right\}^T \\ \sqrt{n}(g(\hat{\xi}) - g(\xi)) &\xrightarrow{d} N(0, \Sigma^N) \\ \Sigma^N &= \frac{\partial g(\xi)}{\partial \pi} \Sigma \frac{\partial g(\xi)}{\partial \pi}^T\end{aligned}$$

$\Sigma^N$  is calculated by delta method,

$$\begin{aligned}\frac{\partial g(\alpha)}{\partial \pi_{00}} &= -\frac{1}{(\pi_{01} + \pi_{00})} = -\frac{1}{\pi_{0+}} \\ \frac{\partial g(\alpha)}{\partial \pi_{01}} &= -\frac{1}{(\pi_{01} + \pi_{00})} = -\frac{1}{\pi_{0+}} \\ \frac{\partial g(\alpha)}{\partial \pi_{10}} &= \frac{1}{(\pi_{10} + \pi_{11})} = \frac{1}{\pi_{1+}} \\ \frac{\partial g(\alpha)}{\partial \pi_{11}} &= \frac{1}{(\pi_{10} + \pi_{11})} = \frac{1}{\pi_{1+}} \\ \frac{\partial g(\beta)}{\partial \pi_{00}} &= \frac{(\pi_{10} - \pi_{01})}{(\pi_{01} + \pi_{00})(\pi_{00} + \pi_{10})} = -\frac{1}{(\pi_{10} + \pi_{00})} + \frac{1}{(\pi_{01} + \pi_{00})} = -\frac{1}{\pi_{+0}} + \frac{1}{\pi_{0+}} \\ \frac{\partial g(\beta)}{\partial \pi_{01}} &= \frac{1}{(\pi_{01} + \pi_{11})} + \frac{1}{(\pi_{01} + \pi_{00})} \\ \frac{\partial g(\beta)}{\partial \pi_{10}} &= -\frac{1}{(\pi_{10} + \pi_{00})} - \frac{1}{(\pi_{10} + \pi_{11})} \\ \frac{\partial g(\beta)}{\partial \pi_{11}} &= \frac{(\pi_{10} - \pi_{01})}{(\pi_{10} + \pi_{11})(\pi_{01} + \pi_{11})} = -\frac{1}{(\pi_{10} + \pi_{11})} + \frac{1}{(\pi_{01} + \pi_{11})} \\ \frac{\partial g(\xi)}{\partial \pi} &= \begin{bmatrix} -\frac{1}{\pi_{0+}} & -\frac{1}{\pi_{0+}} & \frac{1}{\pi_{1+}} & \frac{1}{\pi_{1+}} \\ \frac{1}{\pi_{0+}} - \frac{1}{\pi_{+0}} & \frac{1}{\pi_{0+}} + \frac{1}{\pi_{+1}} & -\frac{1}{\pi_{+0}} - \frac{1}{\pi_{1+}} & \frac{1}{\pi_{+1}} - \frac{1}{\pi_{1+}} \end{bmatrix} \\ \Sigma^N &= \frac{\partial g(\xi)}{\partial \pi} \Sigma \frac{\partial g(\xi)}{\partial \pi}^T \\ &= \begin{pmatrix} \frac{1}{\pi_{11}} + \frac{1}{\pi_{12}} + \frac{1}{\pi_{21}} + \frac{1}{\pi_{22}} \end{pmatrix}\end{aligned}$$

- (f) Under the model of part (d), show that  $(\pi_{1+}\pi_{0+})^{-1} + (\pi_{+1}\pi_{+0})^{-1} \leq (\pi_{1+}\pi_{+0})^{-1} + (\pi_{+1}\pi_{0+})^{-1}$ .

$$\begin{aligned}
& (\pi_{1+}\pi_{0+})^{-1} + (\pi_{+1}\pi_{0+})^{-1} - (\pi_{1+}\pi_{0+})^{-1} - (\pi_{+1}\pi_{0+})^{-1} \\
&= \frac{\pi_{0+} - \pi_{+0}}{\pi_{1+}\pi_{0+}\pi_{0+}} + \frac{\pi_{+0} - \pi_{0+}}{\pi_{+1}\pi_{0+}\pi_{0+}} \\
&= \frac{(\pi_{0+} - \pi_{+0})(\pi_{+1} - \pi_{1+})}{\pi_{1+}\pi_{0+}\pi_{0+}\pi_{+1}} \\
&= \frac{(\pi_{01} - \pi_{10})^2}{\pi_{1+}\pi_{0+}\pi_{0+}\pi_{+1}} \geq 0
\end{aligned}$$

From above, we have  $(\pi_{1+}\pi_{0+})^{-1} + (\pi_{+1}\pi_{0+})^{-1} \leq (\pi_{1+}\pi_{0+})^{-1} + (\pi_{+1}\pi_{0+})^{-1}$ .

## 2.2 Logistic Regression

Consider independent observations  $(X_1, Y_1), \dots, (X_n, Y_n)$  where  $Y_i$  takes values 0 and 1. Suppose that  $X_i|Y_i = m \sim N(\mu_m, \sigma^2)$  and  $P(Y_i = m) = \pi_m$  for  $m = 0, 1$ , where  $\pi_0 + \pi_1 = 1$ , and  $0 < \pi_0 < 1$ .

(a) Show that  $P(Y_i = m|X_i), m = 0, 1$ , satisfies the logistic model, that is

$$\text{logit}(P(Y_i = 1|X_i, \alpha)) = \alpha_0 + \alpha_1 X_i$$

We have distribution of  $P(Y_i = m|X_i), m = 0, 1$

$$\begin{aligned}
P(Y_i = m|X_i, \alpha) &= \frac{P(Y_i, X_i)}{P(X_i)} = \frac{P(X_i|Y_i)P(Y_i)}{P(X_i)} \\
P(Y_i = 1|X_i, \alpha) &= \frac{P(X_i|Y_i = 1)P(Y_i = 1)}{P(X_i)} \\
&= \frac{\exp(-1/2\sigma^2(x_i - \mu_1)^2)\pi_1}{\exp(-1/2\sigma^2(x_i - \mu_i)^2)\pi_1 + \exp(-1/2\sigma^2(x_i - \mu_1)^2)\pi_0} \\
P(Y_i = 0|X_i, \alpha) &= \frac{P(X_i|Y_i = 0)P(Y_i = 0)}{P(X_i)} \\
&= \frac{\exp(-1/2\sigma^2(x_i - \mu_1)^2)\pi_0}{\exp(-1/2\sigma^2(x_i - \mu_i)^2)\pi_1 + \exp(-1/2\sigma^2(x_i - \mu_1)^2)\pi_0} \\
\text{logit}(P(Y_i = 1|X_i, \alpha)) &= \log \frac{P(Y_i = 1|X_i, \alpha)}{P(Y_i = 0|X_i, \alpha)} \\
&= \log(\pi_1/\pi_0) - \frac{(x_i - \mu_1)^2}{2\sigma^2} + \frac{(x_i - \mu_0)^2}{2\sigma^2} \\
&= \log(\pi_1/\pi_0) + \frac{\mu_0^2 - \mu_1^2}{2\sigma^2} + \frac{(\mu_1 - \mu_0)}{\sigma^2} x_i \\
\text{In which, } \alpha &= (\alpha_0, \alpha_1) = \left( \log(\pi_1/\pi_0) + \frac{\mu_0^2 - \mu_1^2}{2\sigma^2}, \frac{(\mu_1 - \mu_0)}{\sigma^2} \right)^T
\end{aligned}$$

- (b) Based on the logistic model in part (a), give the explicit form of the Newton-Raphson algorithm for calculating the maximum likelihood estimate of  $\alpha$ , denoted by  $\hat{\alpha} = (\hat{\alpha}_0, \hat{\alpha}_1)$ , and derive the asymptotic covariance matrix of  $\alpha$ .  
 $Y_i|X_i$  follows a binomial distribution

$$\begin{aligned}
p(Y_i|\alpha) &= P(Y_i = 1|X_i, \alpha)^{I(y_i=1)} P(Y_i = 0|X_i, \alpha)^{I(y_i=0)} \\
\log p(Y_i|\alpha) &= I(y_i = 1)\log P(Y_i = 1|X_i, \alpha) + I(y_i = 0)\log P(Y_i = 0|X_i, \alpha) \\
\ln p(Y_i|\alpha) &= \sum_{i=1}^n I(y_i = 1)\log P(Y_i = 1|X_i, \alpha) + I(y_i = 0)\log P(Y_i = 0|X_i, \alpha) \\
&= \sum_{i=1}^n I(y_i = 1)\log P(Y_i = 1) + (1 - I(y_i = 1))\log(1 - P(Y_i = 1)) \\
&= \sum_{i=1}^n I(y_i = 1)\log P(Y_i = 1)/(1 - P(Y_i = 1)) + \log(1 - P(Y_i = 1))
\end{aligned}$$

Let  $\theta = \log P(Y_i = 1)/(1 - P(Y_i = 1))$

$$\begin{aligned}
\ln p(Y_i|\theta) &= \sum_{i=1}^n I(y_i = 1)\theta - \log(1 + \exp(\theta)) \\
\ln p(Y_i|\alpha) &= \sum_{i=1}^n y_i(\alpha_0 + \alpha_1 x_i) - \log(1 + \exp(\alpha_0 + \alpha_1 x_i))
\end{aligned}$$

Find MLE for  $\alpha$

$$\begin{aligned}
\frac{\partial \ln p(Y_i|\alpha)}{\partial \alpha_0} &= \sum_{i=1}^n y_i - (1 + \exp(\alpha_0 + \alpha_1 x_i))^{-1} \exp(\alpha_0 + \alpha_1 x_i) \\
\frac{\partial \ln p(Y_i|\alpha)}{\partial \alpha_1} &= \sum_{i=1}^n y_i x_i - (1 + \exp(\alpha_0 + \alpha_1 x_i))^{-1} \exp(\alpha_0 + \alpha_1 x_i) x_i \\
\frac{\partial \ln^2 p(Y_i|\alpha)}{\partial \alpha_0^2} &= - \sum_{i=1}^n \frac{\exp(\alpha_0 + \alpha_1 x_i)}{[1 + \exp(\alpha_0 + \alpha_1 x_i)]^2}, \quad E\left[-\frac{\partial \ln^2 p(Y_i|\alpha)}{\partial \alpha_0^2}\right] = n\pi_1(1 - \pi_1) \\
\frac{\partial \ln^2 p(Y_i|\alpha)}{\partial \alpha_1^2} &= - \sum_{i=1}^n \frac{\exp(\alpha_0 + \alpha_1 x_i)}{[1 + \exp(\alpha_0 + \alpha_1 x_i)]^2} x_i x_i^T \\
\frac{\partial \ln^2 p(Y_i|\alpha)}{\partial \alpha_0 \alpha_1} &= - \sum_{i=1}^n \frac{\exp(\alpha_0 + \alpha_1 x_i)}{[1 + \exp(\alpha_0 + \alpha_1 x_i)]^2} x_i \\
I_n(\alpha) &= -E\left[\frac{\partial \ln^2 p(Y_i|\alpha)}{\partial \alpha^2}\right] \\
&= \begin{bmatrix} n\pi_1(1 - \pi_1) & \sum_{i=1}^n \pi_1(1 - \pi_1)x_i \\ \sum_{i=1}^n \pi_1(1 - \pi_1)x_i & \sum_{i=1}^n \pi_1(1 - \pi_1)x_i x_i^T \end{bmatrix}
\end{aligned}$$

So the N-R algorithm is

$$\alpha_{k+1} = \alpha_k - I_n(\alpha_k)^{-1} \frac{\partial \ln p(Y_i | \alpha_k)}{\partial \alpha_k}$$

The asymptotic distribution of  $\alpha$  by CLT and covariance matrix

$$\begin{aligned} \sqrt{n}(\hat{\alpha} - \alpha) &\xrightarrow{d} N(0, \Sigma) \\ \Sigma &= \left\{ \frac{1}{n} I_n(\alpha) \right\}^{-1} \end{aligned}$$

- (c) Write down the joint distribution of  $\{(X_i Y_i) : i = 1, 2, \dots, n\}$  and calculate the maximum likelihood estimate of  $\theta$ , denoted by  $\theta_F$ , and its asymptotic covariance matrix.

The joint distribution of  $\{(X_i Y_i) : i = 1, 2, \dots, n\}$

$$\begin{aligned} p(X_i, Y_i) &= P(X_i | Y_i) P(Y_i) \\ p(Y_i = 1, X_i) &= \frac{1}{\sqrt{2\pi}\sigma} \exp(-1/2\sigma^2(x_i - \mu_1)^2) \pi_1 \\ p(Y_i = 0, X_i) &= \frac{1}{\sqrt{2\pi}\sigma} \exp(-1/2\sigma^2(x_i - \mu_0)^2) \pi_0 \\ p(X_i, Y_i) &= P(Y_i = 1, X_i)^{I(y_i=1)} P(Y_i = 0, X_i)^{I(y_i=0)} \\ &= \left\{ \frac{1}{\sqrt{2\pi}\sigma} \exp(-1/2\sigma^2(x_i - \mu_1)^2) \pi_1 \right\}^{y_i} \left\{ \frac{1}{\sqrt{2\pi}\sigma} \exp(-1/2\sigma^2(x_i - \mu_0)^2) \pi_0 \right\}^{1-y_i} \\ \log p(X_i, Y_i) &= \log \frac{1}{\sqrt{2\pi}\sigma} + y_i \log \pi_1 + (1 - y_i) \log(1 - \pi_1) - \frac{(x_i - \mu_1)^2}{2\sigma^2} y_i - \frac{(x_i - \mu_0)^2}{2\sigma^2} (1 - y_i) \end{aligned}$$

The log-likelihood function of  $\{(X_i Y_i) : i = 1, 2, \dots, n\}$

$$\log p(X, Y) = n \log \frac{1}{\sqrt{2\pi}\sigma} + \sum_{i=1}^n y_i \log \pi_1 + (1 - y_i) \log(1 - \pi_1) - \frac{(x_i - \mu_1)^2}{2\sigma^2} y_i - \frac{(x_i - \mu_0)^2}{2\sigma^2} (1 - y_i)$$

The MLE of  $\theta$  could get by taking derivatives to log-likelihood function

$$\begin{aligned}
\frac{\partial \ln p(X, Y | \theta)}{\partial \pi_1} &= \sum_{i=1}^n y_i / \pi_1 - (1 - y_i) / (1 - \pi_1) = 0 \\
\frac{\partial \ln p(X, Y | \theta)}{\partial \mu_1} &= \sum_{i=1}^n \frac{y_i (x_i - \mu_1)}{\sigma^2} = 0 \\
\frac{\partial \ln p(X, Y | \theta)}{\partial \mu_0} &= \sum_{i=1}^n \frac{(1 - y_i) (x_i - \mu_0)}{\sigma^2} = 0 \\
\frac{\partial \ln p(X, Y | \theta)}{\partial \sigma^2} &= -\frac{n}{2} 1 / \sigma^2 + \sum_{i=1}^n \frac{(x_i - \mu_1)^2 y_i}{2\sigma^4} + \sum_{i=1}^n \frac{(x_i - \mu_0)^2 (1 - y_i)}{2\sigma^4} = 0 \\
\hat{\sigma}^2 &= \frac{\sum_{i=1}^n [(x_i - \mu_1)^2 y_i + (x_i - \mu_0)^2 (1 - y_i)]}{n} \\
\hat{\pi}_1 &= \frac{\sum_{i=1}^n y_i}{n}, \quad \hat{\mu}_1 = \frac{\sum_{i=1}^n x_i y_i}{\sum_{i=1}^n y_i}, \quad \hat{\mu}_0 = \frac{\sum_{i=1}^n x_i (1 - y_i)}{\sum_{i=1}^n (1 - y_i)}
\end{aligned}$$

The Fisher information matrix

$$\begin{aligned}
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \pi_1^2} &= \sum_{i=1}^n -\frac{y_i}{\pi_1^2} - \frac{(1-y_i)}{(1-\pi_1)^2}, & E\left[-\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \pi_1^2}\right] &= \frac{1}{\pi_1(1-\pi_1)} \\
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \mu_1^2} &= \sum_{i=1}^n -\frac{y_i}{\sigma^2}, & E\left[-\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \mu_1^2}\right] &= \frac{\pi_1}{\sigma^2} \\
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \mu_0^2} &= \sum_{i=1}^n -\frac{(1-y_i)}{\sigma^2}, & E\left[-\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \mu_0^2}\right] &= \frac{1-\pi_1}{\sigma^2} \\
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial (\sigma^2)^2} &= \frac{n}{2(\sigma^2)^2} - \sum_{i=1}^n \frac{(x_i - \mu_1)^2 y_i}{(\sigma^2)^3} - \sum_{i=1}^n \frac{(x_i - \mu_0)^2 (1-y_i)}{(\sigma^2)^3} \\
E\left[-\frac{\partial \ln^2 p(X, Y|\theta)}{\partial (\sigma^2)^2}\right] &= \frac{1}{2\sigma^4} \\
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \pi_1 \mu_1} &= 0 \\
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \pi_1 \mu_0} &= 0 \\
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \pi_1 \sigma} &= 0 \\
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \mu_1 \mu_0} &= 0 \\
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \mu_1 \sigma} &= \sum_{i=1}^n -\frac{y_i(x_i - \mu_1)}{(\sigma^2)^2}, & E\left[-\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \mu_1 \sigma}\right] &= 0 \\
\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \mu_0 \sigma} &= \sum_{i=1}^n -\frac{(1-y_i)(x_i - \mu_0)}{(\sigma^2)^2}, & E\left[-\frac{\partial \ln^2 p(X, Y|\theta)}{\partial \mu_0 \sigma}\right] &= 0
\end{aligned}$$

So we have covariance matrix, by CLT

$$\begin{aligned}
I(\theta) &= E\left[-\frac{1}{n} \frac{\partial \ln^2 p(X, Y|\theta)}{\partial \theta^2}\right], & &= \begin{bmatrix} \frac{1}{\pi_1(1-\pi_1)} & 0 & 0 & 0 \\ 0 & \frac{\pi_1}{\sigma^2} & 0 & 0 \\ 0 & 0 & \frac{1-\pi_1}{\sigma^2} & 0 \\ 0 & 0 & 0 & \frac{1}{2\sigma^4} \end{bmatrix} \\
\sqrt{n}(\hat{\theta} - \theta) &\xrightarrow{d} N(0, \Sigma), & \Sigma(\theta) &= I(\theta)^{-1} = \begin{bmatrix} \pi_1(1-\pi_1) & 0 & 0 & 0 \\ 0 & \frac{\sigma^2}{\pi_1} & 0 & 0 \\ 0 & 0 & \frac{\sigma^2}{1-\pi_1} & 0 \\ 0 & 0 & 0 & 2\sigma^4 \end{bmatrix}
\end{aligned}$$

(d) Calculate the asymptotic covariance matrix of  $h(\hat{\theta}^F)$ .

$$\begin{aligned}
h(\theta^F) &= (\alpha_0, \alpha_1) = \left( \log\left(\frac{\pi_1}{1-\pi_1}\right) + \frac{\mu_0^2 - \mu_1^2}{2\sigma^2}, \frac{(\mu_1 - \mu_0)}{\sigma^2} \right)^T \\
\frac{\partial h(\theta^F)}{\partial \pi_1} &= \left( \frac{1}{\pi_1} + \frac{1}{1-\pi_1}, 0 \right)^T \\
\frac{\partial h(\theta^F)}{\partial \mu_1} &= \left( -\frac{\mu_1}{\sigma^2}, \frac{1}{\sigma^2} \right)^T \\
\frac{\partial h(\theta^F)}{\partial \mu_0} &= \left( \frac{\mu_0}{\sigma^2}, -\frac{1}{\sigma^2} \right)^T \\
\frac{\partial h(\theta^F)}{\partial \sigma^2} &= \left( -\frac{(\mu_0^2 - \mu_1^2)}{2\sigma^4}, -\frac{(\mu_1 - \mu_0)}{\sigma^4} \right)^T \\
\sqrt{n}(h(\hat{\theta}^F) - h(\theta^F)) &\xrightarrow{d} N(0, \Sigma_h)
\end{aligned}$$

By delta method,

$$\begin{aligned}
\Sigma^h &= h(\theta^F)' \Sigma(\theta) (\theta^F)^T \\
&= \begin{bmatrix} \frac{1}{\pi_1} + \frac{1}{1-\pi_1} & -\frac{\mu_1}{\sigma^2} & \frac{\mu_0}{\sigma^2} & -\frac{(\mu_0^2 - \mu_1^2)}{2\sigma^4} \\ 0 & \frac{1}{\sigma^2} & -\frac{1}{\sigma^2} & -\frac{(\mu_1 - \mu_0)}{\sigma^4} \end{bmatrix} \begin{bmatrix} \pi_1(1-\pi_1) & 0 & 0 & 0 \\ 0 & \frac{\sigma^2}{\pi_1} & 0 & 0 \\ 0 & 0 & \frac{\sigma^2}{1-\pi_1} & 0 \\ 0 & 0 & 0 & 2\sigma^4 \end{bmatrix} \begin{bmatrix} \frac{1}{\pi_1} + \frac{1}{1-\pi_1} & 0 \\ -\frac{\mu_1}{\sigma^2} & \frac{1}{\sigma^2} \\ \frac{\mu_0}{\sigma^2} & -\frac{1}{\sigma^2} \\ -\frac{(\mu_0^2 - \mu_1^2)}{2\sigma^4} & -\frac{(\mu_1 - \mu_0)}{\sigma^4} \end{bmatrix} \\
&= \begin{bmatrix} \frac{1}{\pi_1(1-\pi_1)} + \frac{\mu_0}{(1-\pi_1)\sigma^2} + \frac{\mu_1}{\pi_1\sigma^2} + \frac{(\mu_0^2 - \mu_1^2)^2}{2\sigma^4} & -\frac{1}{\sigma^2} \left( \frac{\mu_0}{1-\pi_1} + \frac{\mu_1}{\pi_1} \right) + \frac{(\mu_1 - \mu_0)(\mu_0^2 - \mu_1^2)}{\sigma^4} \\ -\frac{1}{\sigma^2} \left( \frac{\mu_0}{1-\pi_1} + \frac{\mu_1}{\pi_1} \right) + \frac{(\mu_1 - \mu_0)(\mu_0^2 - \mu_1^2)}{\sigma^4} & \frac{1}{\sigma^2 \pi_1 (1-\pi_1)} + \frac{2(\mu_1 - \mu_0)^2}{\sigma^4} \end{bmatrix}
\end{aligned}$$

(e) In this part, suppose that  $\mu_0 = \mu_1$ . Show that  $Cov(\hat{\alpha})^{-1}Cov(h(\hat{\theta}^F))$  converges to a matrix which does not depend on  $\theta$ . Interpret this result.

If  $\mu_0 = \mu_1$ , then  $\alpha = (\alpha_0, \alpha_1)^T = (\log(\pi_1/\pi_0), 0)^T$  The covariance matrix of  $\alpha$

$$\begin{aligned}
\alpha_0 &= \log(\pi_1/\pi_0) \\
\ln p(Y_i|\alpha) &= \sum_{i=1}^n y_i(\alpha_0) - \log(1 + \exp(\alpha_0)) \\
\frac{\partial \ln p(Y_i|\alpha)}{\partial \alpha_0} &= \sum_{i=1}^n y_i - \frac{\exp \alpha_0}{1 + \exp \alpha_0} \\
\frac{\partial \ln^2 p(Y_i|\alpha)}{\partial \alpha_0^2} &= \sum_{i=1}^n -\frac{\exp \alpha_0}{(1 + \exp \alpha_0)^2} \\
I_n(\alpha) &= E\left[-\frac{\partial \ln^2 p(Y_i|\alpha)}{\partial \alpha_0^2}\right] = \sum_{i=1}^n \frac{\exp \alpha_0}{(1 + \exp \alpha_0)^2} \\
\log p(\theta) &= n \log \frac{1}{\sqrt{2\pi}\sigma} + \sum_{i=1}^n y_i \log \pi_1 + (1 - y_i) \log(1 - \pi_1) - \frac{(x_i - \mu)^2}{2\sigma^2} \\
\frac{\partial \ln p(\theta)}{\partial \pi_1} &= \sum_{i=1}^n \frac{y_i}{\pi_1} - \frac{1 - y_i}{1 - \pi_1} \\
\frac{\partial \ln^2 p(\theta)}{\partial \pi_1^2} &= \sum_{i=1}^n -\frac{y_i}{\pi_1^2} - \frac{1 - y_i}{(1 - \pi_1)^2}, \quad E\left[-\frac{\partial \ln^2 p(\theta)}{\partial \pi_1^2}\right] = n \frac{\pi_1}{(1 - \pi_1)} \\
\frac{\partial \ln p(\theta)}{\partial \mu} &= \sum_{i=1}^n \frac{x_i - \mu}{\sigma^2} \\
\frac{\partial \ln^2 p(\theta)}{\partial \mu^2} &= \sum_{i=1}^n -\frac{1}{\sigma^2} \\
\frac{\partial \ln p(\theta)}{\partial \sigma^2} &= -\frac{n}{2} \frac{1}{\sigma^2} + \sum_{i=1}^n \frac{(x_i - \mu)^2}{2\sigma^4} \\
\frac{\partial \ln^2 p(\theta)}{\partial (\sigma^2)^2} &= \frac{n}{2(\sigma^2)^2} - \sum_{i=1}^n \frac{(x_i - \mu)^2}{\sigma^6}, \quad E\left[-\frac{\partial \ln^2 p(\theta)}{\partial (\sigma^2)^2}\right] = \frac{n}{2\sigma^4} \\
\frac{\partial \ln^2 p(\theta)}{\partial \mu \sigma^2} &= \sum_{i=1}^n -\frac{x_i - \mu}{\sigma^4}, \quad E\left[-\frac{\partial \ln^2 p(\theta)}{\partial \mu \sigma^2}\right] = 0
\end{aligned}$$



Then we have Fisher information  $I_n(\theta)$

$$\begin{aligned}
I_n(\theta) &= E\left[-\frac{\partial^2 \ln^2 p(\theta)}{\partial \theta^2}\right] \\
&= \begin{bmatrix} n \frac{\pi_1}{(1-\pi_1)} & 0 & 0 \\ 0 & \frac{n}{\sigma^2} & 0 \\ 0 & 0 & \frac{n}{2\sigma^4} \end{bmatrix} \\
Cov(\hat{\alpha})^{-1} &= I_n(\alpha) = n\pi_1(1-\pi_1) \\
\frac{\partial h}{\partial \theta} &= \left(\frac{1}{\pi_1(1-\pi_1)}, 0, 0\right)
\end{aligned}$$

Then we have

$$\begin{aligned}
Cov(\hat{\alpha})^{-1} \Sigma^h &= I_n(\alpha) \frac{\partial h}{\partial \theta} I_n(\theta)^{-1} \frac{\partial h^T}{\partial \theta} \\
&= n\pi_1(1-\pi_1) \left(\frac{1}{\pi_1(1-\pi_1)}, 0, 0\right) \begin{bmatrix} \pi_1(1-\pi_1)/n & 0 & 0 \\ 0 & \sigma^2/n & 0 \\ 0 & 0 & 2\sigma^4/n \end{bmatrix} \left(\frac{1}{\pi_1(1-\pi_1)}, 0, 0\right)^T \\
&= 1
\end{aligned}$$

So we have  $Cov(\hat{\alpha})^{-1} Cov(h(\hat{\theta}^F))$  converges to a matrix which does not depend on  $\theta$ .

- (f) Now suppose that  $\pi_1$  is known. Will the results of (b) - (e) be changed? Please explain. If so, then derive the corresponding results and compare with those obtained above.

If  $\pi_1$  is known,

- (i) For (b), does not change as the parameters are  $\alpha = (\alpha_0, \alpha_1)^T$  which does not involve  $\pi_1$ .

$$\begin{aligned}
I_n(\alpha) &= -E\left[\frac{\partial^2 \ln^2 p(Y_i|\alpha)}{\partial \alpha^2}\right] \\
&= \begin{bmatrix} n\pi_1(1-\pi_1) & \sum_{i=1}^n \pi_1(1-\pi_1)x_i \\ \sum_{i=1}^n \pi_1(1-\pi_1)x_i & \sum_{i=1}^n \pi_1(1-\pi_1)x_i x_i^T \end{bmatrix} \\
Cov(\alpha) &= I_n(\alpha)^{-1} = \frac{1}{[\sum_{i=1}^n nx_i^2 - (\sum_{i=1}^n x_i)^2]\pi_1(1-\pi_1)} \begin{bmatrix} \sum_{i=1}^n nx_i^2 & -\sum_{i=1}^n x_i \\ -\sum_{i=1}^n x_i & n \end{bmatrix}
\end{aligned}$$

- (ii) For (c), it involves  $\pi_1$ , so the result will change. We have covariance matrix

for  $\theta = (\mu_1, \mu_0, \sigma^2)^T$ ,

$$I(\theta) = E\left[-\frac{1}{n} \frac{\partial \ln^2 p(X, Y|\theta)}{\partial \theta^2}\right], \quad = \begin{bmatrix} \frac{\pi_1}{\sigma^2} & 0 & 0 \\ 0 & \frac{1-\pi_1}{\sigma^2} & 0 \\ 0 & 0 & \frac{1}{2\sigma^4} \end{bmatrix}$$

$$\sqrt{n}(\hat{\theta} - \theta) \xrightarrow{d} N(0, \Sigma), \quad \Sigma = I(\theta)^{-1} = \begin{bmatrix} \frac{\sigma^2}{\pi_1} & 0 & 0 \\ 0 & \frac{\sigma^2}{1-\pi_1} & 0 \\ 0 & 0 & 2\sigma^4 \end{bmatrix}$$

(iii) For (d), the  $h(\theta)$  does not involve  $\pi_1$ , but the Jacobian matrix and  $I(\theta)$  will change when  $\pi_1$  is known. We have covariance matrix for  $h(\theta) = c(\mu, \sigma^2)$ .

$$h(\theta^F) = (\alpha_0, \alpha_1) = \left( \log\left(\frac{\pi_1}{1-\pi_1}\right) + \frac{\mu_0^2 - \mu_1^2}{2\sigma^2}, \frac{(\mu_1 - \mu_0)}{\sigma^2} \right)^T$$

$$\sqrt{n}(h(\hat{\theta}^F) - h(\theta^F)) \xrightarrow{d} N(0, \Sigma_h)$$

$$h(\theta^F)' = \begin{bmatrix} -\frac{\mu_1}{\sigma^2} & \frac{\mu_0}{\sigma^2} & -\frac{(\mu_0^2 - \mu_1^2)}{2\sigma^4} \\ \frac{1}{\sigma^2} & -\frac{1}{\sigma^2} & -\frac{(\mu_1 - \mu_0)}{\sigma^4} \end{bmatrix}$$

$$\Sigma(\theta) = \begin{bmatrix} \frac{\sigma^2}{\pi_1} & 0 & 0 \\ 0 & \frac{\sigma^2}{1-\pi_1} & 0 \\ 0 & 0 & 2\sigma^4 \end{bmatrix}$$

$$\Sigma^h = h(\theta^F)' \Sigma(\theta) (\theta^F)^T$$

$$= \begin{bmatrix} \frac{\mu_0}{(1-\pi_1)\sigma^2} + \frac{\mu_1}{\pi_1\sigma^2} + \frac{(\mu_0^2 - \mu_1^2)^2}{2\sigma^4} & -\frac{1}{\sigma^2} \left( \frac{\mu_0}{1-\pi_1} + \frac{\mu_1}{\pi_1} \right) + \frac{(\mu_1 - \mu_0)(\mu_0^2 - \mu_1^2)}{\sigma^4} \\ -\frac{1}{\sigma^2} \left( \frac{\mu_0}{1-\pi_1} + \frac{\mu_1}{\pi_1} \right) + \frac{(\mu_1 - \mu_0)(\mu_0^2 - \mu_1^2)}{\sigma^4} & \frac{2(\mu_1 - \mu_0)^2}{\sigma^4} \end{bmatrix}$$

(iv) For (e), the only parameter that need to estimate is  $\alpha_0 = \log(\pi_1/(1 - \pi_1))$ , which is now known. The question is meaningless.