

# Example for large sample credible intervals

## Example 1: Prevalence of Recessive Gene

If gene frequencies are in equilibrium, the genotypes  $AA$ ,  $Aa$ , and  $aa$  occur with probabilities  $(1-\theta)^2$ ,  $2\theta(1-\theta)$ , and  $\theta^2$  respectively, where  $\theta$  represents the overall prevalence of the recessive  $a$  gene in the population. Plato et al. (1964) published the following data on a haptoglobin type in a sample of 190 people:

Haptoglobin Type	AA	Aa	aa
Count	112	68	10

Let's regard the vector  $\mathbf{x} = (x_1, x_2, x_3) = (112, 68, 10)$  as a realization of the random variable  $\mathbf{X} \sim \text{Multinomial}((1-\theta)^2, 2\theta(1-\theta), \theta^2)$ .

To save some time and allow us to focus on the results of interest here, I'll give you the likelihood function, its first and second derivatives with respect to  $\theta$ , and the form of the posterior:

### Preliminary Results

#### General form of Multinomial pmf

$$f(\mathbf{x}|\mathbf{p}) = \frac{n!}{x_1!x_2!\dots x_k!} p_1^{x_1} p_2^{x_2} \dots p_k^{x_k}$$

#### Likelihood function

In our example,  $p_1 = (1-\theta)^2$ ,  $p_2 = 2\theta(1-\theta)$ , and  $p_3 = \theta^2$ .

$$\begin{aligned}\mathcal{L}(\theta|\mathbf{x}) &= f(\mathbf{x}|\theta) \\ &= \{(1-\theta)^2\}^{x_1} \{2\theta(1-\theta)\}^{x_2} \{\theta^2\}^{x_3}\end{aligned}$$

I'm going to leave this in terms of  $x_1$ ,  $x_2$ , and  $x_3$  for now.

#### Log-likelihood function

$$\begin{aligned}\ell(\theta|\mathbf{x}) &= \log[\mathcal{L}(\theta|\mathbf{x})] \\ &= \log [\{(1-\theta)^2\}^{x_1} \{2\theta(1-\theta)\}^{x_2} \{\theta^2\}^{x_3}] \\ &= x_1 \log\{(1-\theta)^2\} + x_2 \log\{2\theta(1-\theta)\} + x_3 \log\{\theta^2\}\end{aligned}$$

### First and second derivatives of log-likelihood function

The first derivative of the log-likelihood is:

$$\frac{d}{d\theta} \ell(\theta|\mathbf{x}) = \dots = \frac{-2x_1\theta}{\theta(1-\theta)} + \frac{x_2(1-2\theta)}{\theta(1-\theta)} + \frac{2x_3(1-\theta)}{\theta(1-\theta)}$$

The second derivative of the log-likelihood is:

$$\frac{d^2}{d\theta^2} \ell(\theta|\mathbf{x}) = \dots = -\frac{2x_1 + x_2}{(1-\theta)^2} - \frac{2x_3 + x_2}{\theta^2}$$

### Maximum likelihood estimator

Setting the first derivative equal to 0, we obtain a maximum likelihood estimator of

$$\hat{\theta}^{MLE} = \frac{X_2 + 2X_3}{2n}.$$

It can be verified that this gives a global maximum of the likelihood function.

### Posterior Distribution

Suppose we adopt a prior of  $\Theta \sim \text{Uniform}(0, 1)$

The prior distribution for  $\Theta$  has density  $f_{\Theta}(\theta) = \begin{cases} 1 & \text{if } \theta \in [0, 1] \\ 0 & \text{otherwise} \end{cases}$ .

Additionally, in part (a) we showed that  $f_{\mathbf{X}|\Theta}(\mathbf{x}|\theta) = \{(1-\theta)^2\}^{x_1} \{2\theta(1-\theta)\}^{x_2} \{\theta^2\}^{x_3}$ .

Applying Bayes' Rule, we find that

$$f_{\Theta|\mathbf{X}}(\theta|\mathbf{x}) = \dots = \begin{cases} c\{(1-\theta)^2\}^{x_1} \{2\theta(1-\theta)\}^{x_2} \{\theta^2\}^{x_3} & \text{if } \theta \in [0, 1] \\ 0 & \text{otherwise} \end{cases}$$

The integral is kind of annoying, but can be done.

### Problems for you

There is just one written problem here. Do this problem before continuing to lab 11.

#### 1. Find a large-sample normal approximation to the posterior distribution for $\theta$ .

Use the approximation centered at the maximum likelihood estimate  $\hat{\theta}^{MLE}$ . Your answer will be in terms of  $x_1$ ,  $x_2$ ,  $x_3$ , and  $n$ .

**Solution:**

The observed Fisher information at the maximum likelihood estimate is:

$$\begin{aligned} J(\hat{\theta}^{MLE}) &= -\frac{d^2}{d\theta^2} \ell(\theta|\mathbf{x})|_{\theta=\hat{\theta}^{MLE}} \\ &= \frac{2x_1 + x_2}{(1 - \hat{\theta}^{MLE})^2} + \frac{2x_3 + x_2}{(\hat{\theta}^{MLE})^2} \\ &= \frac{2x_1 + x_2}{(1 - \frac{x_2+2x_3}{2n})^2} + \frac{2x_3 + x_2}{(\frac{x_2+2x_3}{2n})^2} \end{aligned}$$

The approximate posterior distribution for  $\Theta$  is therefore

$$\Theta|X_1 = x_1, X_2 = x_2, X_3 = x_3 \sim \text{Normal} \left[ \frac{x_2+2x_3}{2n}, \left\{ \frac{2x_1+x_2}{(1-\frac{x_2+2x_3}{2n})^2} + \frac{2x_3+x_2}{(\frac{x_2+2x_3}{2n})^2} \right\}^{-1} \right]$$