

Hierarchical Bayesian model for certification of a country as “free” from an animal disease

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Outline:

- Background
- Disease Freedom
- Current Approaches
 - Qualitative
 - Quantitative
- Hierarchical Model
- Bayesian Approach: Gibbs Sampling
- Example: Newcastle Disease

Background:

- Risk analysis for trade in animal products.
- Countries are interested in new trading opportunities and maintaining current trade.
- Is the country disease free?
- Needed for risk analysis to make policy decisions.

Disease Freedom:

- “Disease freedom”
 - requires a *perfectly* sensitive test
 - *All* animals
 - *All* negative tests
- Level of Disease Freedom may not mean total freedom in all situations.
 - prevalence < threshold

Current Approaches:

- Qualitative
- Quantitative
 - Monte Carlo Simulation

Data:

- Two-stage cluster sample.
- Random selection of k herds and a random sample of n animals within each herd.

Bayesian Model:

Country level:

Z = “true” status of the country.

$$\begin{aligned} Z = 1 & \quad \alpha && \text{diseased} \\ = 0 & \quad 1 - \alpha \end{aligned}$$

Herd level: (latent data)

t_i = “true” status of herd i .

$$\begin{aligned} t_i &= 1 & \tau & \quad i^{\text{th}} \text{ herd is diseased} \\ &= 0 & 1 - \tau & \end{aligned}$$

Within herd level:

Assume two populations exist from which the i^{th} herd can be selected, either diseased or non-diseased.

λ_i = prevalence of the disease in the population from which the i^{th} herd was sampled.

π_i = prevalence of the disease in a diseased herd.

Prevalence Within Diseased Herds:

$$\pi_i \sim \text{beta}(a, b)$$

where (a, b) is random.

Prevalence Within Herds:

$$\begin{aligned} \lambda_i | \pi_i &= \begin{matrix} \pi_i & \tau \\ 0 & 1 - \tau \end{matrix} \\ &= \end{aligned}$$

So

$$\lambda_i | \pi_i, \tau \sim \pi_i \text{Bernoulli}(\tau)$$

Within herd level: (latent data)

v_{ij} = “true” status of animal j within herd i .

$$\begin{aligned} v_{ij} &= 1 && j^{th} \text{ animal in } i^{th} \text{ herd is diseased} \\ &= 0 && \text{not diseased} \end{aligned}$$

So

$$v_{ij} | Z = 1, t_i = 1 \sim \text{Bernoulli}(\pi_i)$$

Data:

$$\begin{aligned} X_{ij} &= 1 && \text{if the } j^{\text{th}} \text{ animal in the } i^{\text{th}} \text{ herd tests positive} \\ &= 0 && \text{otherwise test negative} \end{aligned}$$

Imperfect diagnostic test:

1. sensitivity $\eta = P(T^+|D) < 1$
2. specificity $\theta = P(T^-|\bar{D}) < 1$

$X_i = \sum_j X_{ij}$ = number of test positive animals in the i^{th} herd.

If $Z = 1$

$$X_i | Z = 1, \lambda_i, \eta, \theta \sim \text{Bin}[n_i, \lambda_i \eta + (1 - \lambda_i)(1 - \theta)]$$

If $Z = 0$

$$X_i | Z = 0 \sim \text{Bin}[n_i, (1 - \theta)]$$

Important: This model generates correlation in disease status between animals within each herd, but leaves the disease status independent between herds.

Bayesian Statistics:

\mathbf{X} is the data vector and Θ is the parameter set.

Model: (or likelihood)

$$p(\mathbf{X}|\Theta)$$

Prior distribution:

$$p(\Theta)$$

Joint distribution:

$$p(\mathbf{X}, \Theta) = p(\mathbf{X}|\Theta)p(\Theta)$$

Posterior distribution: (Bayes Theorem)

$$p(\Theta|\mathbf{X}) = \frac{p(\mathbf{X}|\Theta)p(\Theta)}{\int p(\mathbf{X}|\Theta)p(\Theta)d\Theta}$$

$$\propto p(\mathbf{X}|\Theta)p(\Theta)$$

Gibbs Sampler: (ref. Gelfand and Smith 1990)

Two parameter case: $\Theta = (\theta_1, \theta_2)$ $p(\Theta|\mathbf{X}) = p(\theta_1, \theta_2|\mathbf{X})$

Want the marginals, $p(\theta_1|\mathbf{X})$ and $p(\theta_2|\mathbf{X})$

Can get the conditional marginals, $p(\theta_1|\mathbf{X}, \theta_2)$ and $p(\theta_2|\mathbf{X}, \theta_1)$

Initial value $(\theta_1^{(0)}, \theta_2^{(0)})$ for $h = 1, \dots, Reps$

1. Sample $\theta_1^{(h)}$ from $p(\theta_1|\mathbf{X}, \theta_2^{(h-1)})$.
2. Sample $\theta_2^{(h)}$ from $p(\theta_2|\mathbf{X}, \theta_1^{(h)})$.
3. Set $h = h + 1$ and go to 1.

Inferences are calculated using the simulated data after the chain has stabilized.

$$(\theta_1^{(BurnIn+1)}, \theta_2^{(BurnIn+1)}), \dots, (\theta_1^{(Reps)}, \theta_2^{(Reps)})$$

The theory behind the Gibbs sampler says, $\Theta^{(1)}, \dots, \Theta^{(Reps)}$ are realizations of a stationary Markov Chain, with transition probability from $\Theta^{(h-1)}$ to $\Theta^{(h)}$,

$$T(\Theta^{(h-1)}, \Theta^{(h)}) = p(\theta_1 | \mathbf{X}, \theta_2^{(h-1)}) p(\theta_2 | \mathbf{X}, \theta_1^{(h)})$$

Justified by Ergodic theory, we can calculate estimates of the parameter using means. For example,

$$\hat{\theta}_1 = \frac{1}{Reps} \sum_h \theta_1^{(i)} \xrightarrow{a.s.} E[\theta_1 | \mathbf{X}],$$

$$Reps \rightarrow \infty.$$

Priors:

- Country level, $\alpha \sim \text{beta}(a_\alpha, b_\alpha)$.
- Herd level, $\tau \sim \text{beta}(a_\tau, b_\tau)$.
- Within herd level, $\pi_i \sim \text{beta}(a, b)$.
- The distribution from which the prevalence a diseased herd results from is assumed to be a beta distribution with unknown parameters (a, b) . Adaptive Rejection Sampling.
- Sensitivity, $\eta \sim \text{beta}(a_\eta, b_\eta)$.
- Specificity, $\theta \sim \text{beta}(a_\theta, b_\theta)$.

The Likelihood:

$$X_{ij} | Z = 1, \{v_{ij}\}, \{t_i\}, \{\pi_i\}, \eta, \theta \sim \text{Bernoulli}[\eta^{v_{ij}} (1 - \theta)^{1-v_{ij}}].$$

$$X_{ij} | Z = 0 \sim \text{Bernoulli}[(1 - \theta)].$$

Parameter Set: $\Theta = \{Z, \{v_{ij}\}, \{t_i\}, \{\pi_i\}, (a, b), \alpha, \tau, \eta, \theta\}$

Joint Posterior:

$$p(\Theta|\mathbf{X}) \propto p(\mathbf{X}|\Theta)p(\Theta)$$

Conditional Marginal Posterior Distributions:

- “true” status of each animal.

$$v_{ij}|Z = 1, X_{ij} = 1, t_i = 1, \text{“rest”} \sim \text{Bernoulli}(P(D|X_{ij} = 1))$$

$$P(D|X_{ij} = 1) = \frac{\eta\pi_i}{\eta\pi_i + (1 - \theta)(1 - \pi_i)}$$

$$v_{ij}|Z = 1, X_{ij} = 0, t_i = 1, \text{“rest”} \sim \text{Bernoulli}(P(D|X_{ij} = 0))$$

$$P(D|X_{ij} = 0) = \frac{(1 - \eta)\pi_i}{(1 - \eta)\pi_i + \theta(1 - \pi_i)}$$

$$v_{ij}|Z = 1, t_i = 0, \text{“rest”} \sim \text{Bernoulli}(0)$$

- “true” status of each herd

If $v_{ij} = 1$ for any j within herd i , then

$$t_i | \{X_{ij}\}, \text{“rest”} \sim \text{Bernoulli}(1).$$

If $v_{ij} = 0$ for all j within herd i , then

$$t_i | \{X_{ij}\}, \text{“rest”} \sim \text{Bernoulli} \left(\frac{(1 - \pi)^n \tau}{(1 - \pi_i)^n + 1(1 - \tau)} \right).$$

If $Z = 0$, then

$$t_i | Z = 0, \text{“rest”} \sim \text{Bernoulli}(0).$$

- prevalence of a diseased herd.

$$\pi_i | \{X_{ij}\}, \text{“rest”} \sim \text{beta} \left(a + \sum_j v_{ij}, b + n - \sum_j v_{ij} \right)$$

- hyperparameters for the distribution on the π_i .

$(a, b) | \{X_{ij}\}, \text{“rest”}$ sampled using Adaptive Rejection.

- probability the country is diseased.

$$\alpha | \{X_{ij}\}, \text{“rest”} \sim \text{beta} (a_\alpha + Z, b_\alpha + (1 - Z))$$

- probability a herd is diseased.

$$\tau | \{X_{ij}\}, \text{“rest”} \sim \text{beta} \left(a_\tau + \sum_i t_i, b_\tau + k - \sum_i t_i \right)$$

- sensitivity.

$$\eta|\{X_{ij}\}, \text{“rest”} \sim \text{beta} \left(a_\eta + \sum x_{ij}v_{ij}, b_\eta + \sum (1 - x_{ij})v_{ij} \right)$$

- specificity.

$$\theta|\{X_{ij}\}, \text{“rest”} \sim \text{beta}(a_\theta + \sum (1 - x_{ij})(1 - v_{ij}), b_\theta + \sum x_{ij}(1 - v_{ij}))$$

- country status.

$$Z|\{X_{ij}\}, \text{“rest”} \sim \text{Bernoulli} \left(\frac{\alpha(1 - \tau)^k}{\alpha(1 - \tau)^k + (1 - \alpha)} \right)$$

Adaptive Rejection Sampling: (ref. Gilks and Wild 1992)

$$\prod_{i=1}^k \left[\frac{\Gamma(a+b)}{\Gamma(a)\Gamma(b)} \pi_i^{a-1} (1-\pi_i)^{b-1} p(a) \right]^{t_i}$$

log-concavity in a given b .

Steps To Perform The Gibbs Sampler:

Given the initial values:

1. $v_{ij}^{(h)} \sim \text{Bernoulli}, 1, \dots, k, j = 1, \dots, n.$
2. $t_i^{(h)} \sim \text{Bernoulli}, i = 1, \dots, k.$
3. sample (a, b) using Adaptive Rejection.
4. $\pi_i^{(h)} \sim \text{beta}, i = 1, \dots, k.$
5. $\alpha^{(h)} \sim \text{beta}.$
6. $\tau^{(h)} \sim \text{beta}.$
7. $\eta^{(h)} \sim \text{beta}.$
8. $\theta^{(h)} \sim \text{beta}.$
9. $Z^{(h)} \sim \text{Bernoulli}.$

Conclusions:

- Bayesian approach.
 - Uses prior knowledge.
 - Latent data.
 - Imperfect test.
- Variable within herd-level prevalence.
- Country/Herd/Animal inference.
- Priors for other risk analysis.

Further Work:

- Variable sample size within herds.
- Application to continuous surveillance.

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