The Transmissibility of Highly Pathogenic Avian Influenza in Commercial Poultry in Industrialized Countries

Tini Garske, Paul Clarke, Azra C. Ghani

Technical Appendix

1 The information matrix for discrete times

In the main paper, we assumed that distribution times followed a Weibull distribution which is a continuous distribution. However, our infection time data is discretized into whole days, and the likelihood must be discretized to reflect this.

To calculate the maximum likelihood (ML) estimate, substitute $W(t_j - t_i; \kappa, \eta)$ for $w(t_j - t_i; \kappa, \eta)$ in equation (6) of the main paper. The Weibull cumulative distribution function (CDF) is given by

$$W(T;\kappa,\eta) = 1 - \exp\left[-(\eta T)^{\kappa}\right], \qquad (A.1)$$

and for every occurrence of $w(t_j - t_i; \kappa, \eta)$ we substituted

$$w_{ij} = W(T + 1/2; \kappa, \eta) - W(T - 1/2; \kappa, \eta)$$
(A.2)

$$= \exp\left[-\left(\eta(T-1/2)\right)^{\kappa}\right] - \exp\left[-\left(\eta(T+1/2)\right)^{\kappa}\right]$$
(A.3)

$$= E^{-} - E^{+}$$
 (A.4)

To calculate variance-covariance matrix in equation (8) of the main paper, we use the standard relationship

$$V(\theta) = J^{-1}(\theta) \tag{A.5}$$

where $J(\theta)$ is the observed information matrix given by

$$J(\hat{\theta}) = -\begin{pmatrix} \frac{\partial^2 \ln L}{\partial \kappa^2} & \frac{\partial^2 \ln L}{\partial \kappa \partial \eta} \\ \frac{\partial^2 \ln L}{\partial \eta \partial \kappa} & \frac{\partial^2 \ln L}{\partial \eta^2} \end{pmatrix}$$
(A.6)

with $\hat{\theta} = (\kappa, \eta)$ and $\ln L$ the log-likelihood with the cumulative density for the Weibull substituted.

To calculate J it is first necessary to evaluate the first and second derivatives with respect to κ and $\eta,$ given by

$$\frac{\partial}{\partial\kappa}E^{\pm} = \beta_{\kappa\pm}E^{\pm} \tag{A.7}$$

$$\frac{\partial}{\partial \eta} E^{\pm} = \beta_{\eta\pm} E^{\pm} \tag{A.8}$$

$$\frac{\partial^2}{\partial \kappa^2} E^{\pm} = \beta_{\kappa\kappa\pm} E^{\pm} = \left(\beta_{\kappa\pm}^2 + \frac{\partial}{\partial \kappa} \beta_{\kappa\pm}\right) E^{\pm} \tag{A.9}$$

$$\frac{\partial}{\partial\eta}E^{\pm} = \beta_{\eta\pm}E^{\pm}$$
(A.8)
$$\frac{\partial^{2}}{\partial\kappa^{2}}E^{\pm} = \beta_{\kappa\kappa\pm}E^{\pm} = \left(\beta_{\kappa\pm}^{2} + \frac{\partial}{\partial\kappa}\beta_{\kappa\pm}\right)E^{\pm}$$
(A.9)
$$\frac{\partial^{2}}{\partial\eta\partial\kappa}E^{\pm} = \beta_{\kappa\eta\pm}E^{\pm} = \left(\beta_{\kappa\pm}\beta_{\eta\pm} + \frac{\partial}{\partial\eta}\beta_{\kappa\pm}\right)E^{\pm}$$
(A.10)

$$\frac{\partial^2}{\partial \eta^2} E^{\pm} = \beta_{\eta\eta\pm} E^{\pm} = \left(\beta_{\eta\pm}^2 + \frac{\partial}{\partial \eta}\beta_{\eta\pm}\right) E^{\pm} \,. \tag{A.11}$$

With $T^{\pm} = T \pm 1/2$, we have

$$\beta_{\kappa\pm} = -\left(\eta T^{\pm}\right)^{\kappa} \ln\left(\eta T^{\pm}\right) \tag{A.12}$$

$$\beta_{\eta\pm} = -\frac{\kappa}{\eta} \left(\eta T^{\pm}\right)^{\kappa} \tag{A.13}$$

$$\beta_{\kappa\kappa\pm} = \beta_{\kappa\pm}^2 - \left(\eta T^{\pm}\right)^{\kappa} \ln^2\left(\eta T^{\pm}\right) \tag{A.14}$$

$$\beta_{\kappa\eta\pm} = \beta_{\kappa\pm}\beta_{\eta\pm} - \frac{(\eta T^{\pm})}{\eta} \left(1 + \kappa \ln\left(\eta T^{\pm}\right)\right) \tag{A.15}$$

$$\beta_{\eta\eta\pm} = \beta_{\eta\pm}^2 - \frac{\kappa}{\eta^2} \left(\eta T^{\pm}\right)^{\kappa} \left(\kappa - 1\right). \tag{A.16}$$

Therefore the entries of the information matrix are

$$\frac{\partial^{2} \ln L}{\partial \kappa^{2}} = \sum_{j=k}^{N} \left[\frac{\sum_{i \in S_{j}} \left(\beta_{\kappa\kappa-} E^{-} - \beta_{\kappa\kappa+} E^{+}\right)}{\sum_{i \in S_{j}} \left(E^{-} - E^{+}\right)} - \frac{\left(\sum_{i \in S_{j}} \left(\beta_{\kappa-} E^{-} - \beta_{\kappa+} E^{+}\right)\right)^{2}}{\left(\sum_{i \in S_{j}} \left(E^{-} - E^{+}\right)\right)^{2}} \right]$$
(A.17)
$$\frac{\partial^{2} \ln L}{\partial \kappa \eta} = \sum_{j=k}^{N} \left[\frac{\sum_{i \in S_{j}} \left(\beta_{\kappa\eta-} E^{-} - \beta_{\kappa\eta+} E^{+}\right)}{\sum_{i \in S_{j}} \left(E^{-} - E^{+}\right)} - \frac{\left(\sum_{i \in S_{j}} \left(\beta_{\kappa-} E^{-} - \beta_{\kappa+} E^{+}\right)\right) \left(\sum_{i \in S_{j}} \left(\beta_{\eta-} E^{-} - \beta_{\eta+} E^{+}\right)\right)}{\left(\sum_{i \in S_{j}} \left(E^{-} - E^{+}\right)\right)^{2}} \right]$$
(A.18)
$$\partial^{2} \ln L = \sum_{i \in S_{j}}^{N} \left[\sum_{i \in S_{j}} \left(\beta_{\eta\eta-} E^{-} - \beta_{\eta\eta+} E^{+}\right)\right) \left(\sum_{i \in S_{j}} \left(\beta_{\eta-} E^{-} - \beta_{\eta+} E^{+}\right)\right) \right]$$

$$\frac{\partial^2 \ln L}{\partial \eta^2} = \sum_{j=k}^{N} \left[\frac{\sum_{i \in S_j} (\beta_{\eta\eta-}E^- - \beta_{\eta\eta+}E^+)}{\sum_{i \in S_j} (E^- - E^+)} - \frac{\left(\sum_{i \in S_j} (\beta_{\eta-}E^- - \beta_{\eta+}E^+)\right)^2}{\left(\sum_{i \in S_j} (E^- - E^+)\right)^2} \right]$$
(A.19)

Once this is done standard formulae for a) the inverse of a two-by-two matrix and b) the conditional normal distribution, can be used to calculate $V = J^{-1}$ and generate draws from the bivariate normal distribution. For completeness, we give each of these results below.

2 Inversion of the information matrix

The variance-covariance matrix is obtained by inverting the information matrix. The general inversion formula for a 2×2 matrix

$$M = \begin{pmatrix} A & B \\ C & D \end{pmatrix} \tag{A.20}$$

is given by

$$M^{-1} = \begin{pmatrix} \frac{D}{\det} & -\frac{B}{\det} \\ -\frac{C}{\det} & \frac{A}{\det} \end{pmatrix} \quad \text{with} \quad \det = AD - BC \,. \tag{A.21}$$

3 Marginal and conditional distributions

The distribution function of a bivariate normal distribution is given by

$$P(x_1, x_2) = \frac{1}{2\pi\sigma_1\sigma_2\sqrt{1-\rho^2}} \exp\left[-\frac{z}{2(1-\rho^2)}\right]$$
(A.22)

with

$$z = \frac{(x_1 - \mu_1)^2}{\sigma_1^2} - \frac{2\rho(x_1 - \mu_1)(x_2 - \mu_2)}{\sigma_1\sigma_2} + \frac{(x_2 - \mu_2)^2}{\sigma_2^2}.$$
 (A.23)

The variance-covariance matrix V determines the variances and correlations via $(\Sigma - \Sigma) = (-2)^2$

$$V = \begin{pmatrix} \Sigma_{11} & \Sigma_{12} \\ \Sigma_{21} & \Sigma_{22} \end{pmatrix} = \begin{pmatrix} \sigma_1^2 & \rho \sigma_1 \sigma_2 \\ \rho \sigma_1 \sigma_2 & \sigma_2^2 \end{pmatrix}.$$
 (A.24)

The marginal probability for any x_i is given by the univariate normal distribution

$$P(x_i) = \frac{1}{\sigma_i \sqrt{2\pi}} \exp\left[-\frac{(x_i - \mu_i)^2}{2\sigma_i^2}\right].$$
 (A.25)

The conditional probability of x_2 given that $x_1 = a$ is also a normal distribution, but with mean $\bar{\mu_2}$ and variance $\bar{\sigma_2^2}$, which are given as

$$\bar{\mu}_2 = \mu_2 + \frac{\Sigma_{21}}{\Sigma_{11}} (a - \mu_1) \tag{A.26}$$

$$\bar{\sigma}_2^2 = \Sigma_{22} - \frac{\Sigma_{21} \Sigma_{12}}{\Sigma_{11}} \tag{A.27}$$