

**Web-based supporting materials for “Modeling Short- and  
Long-term Characteristics of Follicle Stimulating Hormone as  
Predictors of Severe Hot Flashes in Penn Ovarian Aging Study” by**

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## Details of posterior computations when assuming t model for longitudinal observations

**note:** in the case of assuming normal model for longitudinal observations, we are not updating  $m_{ij}$  but let  $m_{ij}$  be constant 1; and the missing longitudinal observation  $y_{ij}$  is drawn from  $N(\mu(t_{ij}), \sigma_i^2)$ .

### (1) update for longitudinal submodel

- **update** the mean profile class memberships  $D_i, i = 1, \dots, n$ : the full conditional posterior distribution  $[D_i | \cdot] \sim \text{Multinomial}(\tilde{\pi}_{i1}^D, \dots, \tilde{\pi}_{iK_D}^D)$ , where

$$\tilde{\pi}_{id}^D = \Pr(D_i = d | \cdot) = \frac{\pi_d^D |\Sigma_d|^{-\frac{1}{2}} \exp\left[-\frac{1}{2}(\mathbf{b}_i - \boldsymbol{\beta}_d)' \Sigma_d^{-1} (\mathbf{b}_i - \boldsymbol{\beta}_d)\right]}{\sum_{d=1}^{K_D} \pi_d^D |\Sigma_d|^{-\frac{1}{2}} \exp\left[-\frac{1}{2}(\mathbf{b}_i - \boldsymbol{\beta}_d)' \Sigma_d^{-1} (\mathbf{b}_i - \boldsymbol{\beta}_d)\right]}$$

- **update** the mean profile class parameters:

- **update**  $\boldsymbol{\beta}_d = (\beta_{d1}, \dots, \beta_{dL})^T$ :

assuming the prior for  $\beta_{d1} \sim N(0, v)$  and first order random walk prior  $\beta_{dl} \sim N(\beta_{d,l-1}, \tau_{\beta d}^2)$ ,

$l = 2, \dots, L$ , then the prior for  $\boldsymbol{\beta}_d$  can be written as:  $\pi(\boldsymbol{\beta}_d) = \left(\frac{1}{\sqrt{2\pi}\tau_{\beta d}}\right)^{L-1} \exp\{-\frac{1}{2}\boldsymbol{\beta}_d^T V_d \boldsymbol{\beta}_d\}$ ,

where  $V_d = \begin{pmatrix} v^{-1} & \mathbf{0} \\ \mathbf{0} & \mathbf{0} \end{pmatrix} + P^T P / \tau_{\beta d}^2$  and  $P = \begin{pmatrix} 1 & -1 & & & \\ & 1 & -1 & & \\ & & \ddots & \ddots & \\ & & & 1 & -1 \end{pmatrix}$  is the  $(L - 1) \times L$  penalty matrix. Then the full conditional posterior density for  $\boldsymbol{\beta}_d$  for  $d = 1, \dots, K_D$

is  $[\boldsymbol{\beta}_d | \cdot] \sim \text{MVN}(\tilde{\boldsymbol{\nu}}_d, \tilde{V}_d)$

$$\begin{aligned} \tilde{\boldsymbol{\nu}}_d &= \left[ V_d + \Sigma_d^{-1} \sum_{i=1}^n I(D_i = d) \right]^{-1} \left[ \Sigma_d^{-1} \sum_{i=1}^n I(D_i = d) \mathbf{b}_i \right] \\ \tilde{V}_d &= \left[ V_d + \Sigma_d^{-1} \sum_{i=1}^n I(D_i = d) \right]^{-1} \end{aligned}$$

- **update**  $\Sigma_d$ : assuming the prior for  $\Sigma_d \stackrel{ind}{\sim} \text{Inverse-Wishart}(m, \Lambda)$ , where  $m$  and  $\Lambda$  are the degrees of freedom and scale matrix, respectively, then the full conditional posterior density is,  $[\Sigma | \cdot] \sim \text{Inverse-Wishart}(\tilde{m}_d, \tilde{\Lambda}_d)$  where

$$\tilde{m}_d = m + n$$

$$\tilde{\Lambda}_d = \left[ \Lambda + \sum_{i=1}^n (\mathbf{b}_i - \boldsymbol{\beta}_d) (\mathbf{b}_i - \boldsymbol{\beta}_d)' \right]$$

- **update**  $\tau_{\beta d}^2$ : assuming  $\tau_{\beta d}^2 \sim \text{Inverse-Gamma}(v, e)$ , where  $v$  and  $e$  are the shape and rate parameters, then the full conditional posterior distribution is

$$[\tau_{\beta d}^2 | \cdot] \sim \text{Inverse-Gamma}\left(v + \frac{L-1}{2}, e + \frac{1}{2} \boldsymbol{\beta}_d^T P \boldsymbol{\beta}_d\right)$$

where  $L$  is the number of B spline basis functions.

- **update** the mixing proportion  $\{\pi_d^D\}_d$ : assuming  $[\{\pi_d^D\}_d] \sim \text{Dirichlet}(e_1^D, \dots, e_{K_D}^D)$  then the full conditional posterior distribution is

$$[\{\pi_d^D\}_d | \cdot] \sim \text{Dirichlet}(\{e_d^D + \sum_{i=1}^n \text{I}(D_i = d)\}_d)$$

- **update** the variance class memberships  $C_i, i = 1, \dots, n$ : the full conditional posterior distribution  $[C_i | \cdot] \sim \text{Multinomial}(\tilde{\pi}_{i1}^C, \dots, \tilde{\pi}_{iK_C}^C)$  where

$$\tilde{\pi}_{ic}^C = \Pr(C_i = c | \cdot) = \frac{\pi_c^C \exp \left[ -\frac{1}{2} (\log \sigma_i^2 - \mu_c)^2 / \tau^2 \right]}{\sum_{c=1}^{K_C} \pi_c^C \exp \left[ -\frac{1}{2} (\log \sigma_i^2 - \mu_c)^2 / \tau^2 \right]}$$

- **update** the variance class parameters:

- **update**  $\mu_c$ : assuming the prior for  $\mu_c \stackrel{\text{ind}}{\sim} N(a, b)$ , then the full conditional posterior distribution is,  $[\mu_c | \cdot] \sim N(\tilde{a}, \tilde{b})$  where

$$\begin{aligned} \tilde{a} &= \frac{\sum_{i=1}^n \text{I}(C_i = c) \log \sigma_i^2 / \tau^2 + a/b}{1/b + \sum_{i=1}^n \text{I}(C_i = c) / \tau^2} \\ \tilde{b} &= \left( 1/b + \sum_{i=1}^n \text{I}(C_i = c) / \tau^2 \right)^{-1} \end{aligned}$$

- **update**  $\tau^2$ : assuming  $\tau^2 \sim \text{Inverse-Gamma}(v, e)$ , then the full conditional posterior distribution is  $[\tau^2 | \cdot] \sim \text{Inverse-Gamma} \left( v + \frac{n}{2}, e + \sum_{i=1}^n \sum_{c=1}^{K_C} \frac{1}{2} \text{I}(C_i = c) (\log \sigma_i^2 - \mu_c)^2 \right)$ .

- **update** the mixing proportions  $\{\pi_c^C\}_c$ : assuming  $[\{\pi_c^C\}_c] \sim \text{Dirichlet}(e_1^C, \dots, e_{K_C}^C)$  then the full conditional posterior distribution is

$$[\{\pi_c^C\}_c | \cdot] \sim \text{Dirichlet} \left( \{e_c^C + \sum_{i=1}^n \text{I}(C_i = c)\}_c \right)$$

- **update** the random effects  $\mathbf{b}_i, i = 1, \dots, n$  the full conditional posterior distribution is  $\mathbf{b}_i | \cdot \sim \text{MVN}(\tilde{\beta}_i, \tilde{\Sigma}_i)$ , where

$$\begin{aligned} \tilde{\Sigma}_i &= \left[ \Sigma^{-1} + \frac{1}{\sigma_i^2} \sum_{j=1}^{n_i} m_{ij} \phi_{ij} \phi'_{ij} + \mathbf{M}_{bi} \mathbf{M}_{bi}^T \right]^{-1} \\ \tilde{\beta}_i &= \tilde{\Sigma}_i \left[ \Sigma^{-1} \beta_{D_i} + \frac{1}{\sigma_i^2} \sum_{j=1}^{n_i} y_{ij} m_{ij} \phi_{ij} + (W_i - \alpha_0 - \mathbf{x}_i^T \lambda_0) \mathbf{M}_{bi} \right] \end{aligned}$$

where,  $\phi_{ij} = (\phi_1(t_{ij}), \dots, \phi_L(t_{ij}))^T$  a vector of B spline basis functions evaluated at time  $t_{ij}$  such that  $\mu_i(t_{ij}) = \mathbf{b}_i^T \phi_{ij}$  where  $y_{ij} \sim N(\mu_i(t_{ij}), \sigma_i^2/m_{ij})$  and  $\mathbf{M}_{bi}$  is defined such that  $\int_T \mu_i(t) \theta_0(t) dt = \int_T \mathbf{b}_i^T \phi(t) \psi^0(t)^T \tilde{\boldsymbol{\theta}}_0 dt = \mathbf{b}_i^T G_T^0 \tilde{\boldsymbol{\theta}}_0 = \mathbf{b}_i^T \mathbf{M}_{bi}$ .

- **update** the variances  $\sigma_i^2, i = 1, \dots, n$

$$\begin{aligned}\pi(\sigma_i^2 | \cdot) &\propto (\sigma_i^2)^{-\frac{n_i}{2}-1} \exp \left[ -\sum_{c=1}^{K_C} I(C_i = c) \frac{(\log \sigma_i^2 - \mu_c)^2}{2\tau^2} - \frac{1}{2\sigma_i^2} \sum_{j=1}^{n_i} m_{ij} (y_{ij} - \mathbf{b}_i^T \phi_{ij})^2 \right] \\ &\times \exp \left[ -\frac{1}{2} \left( W_i - \alpha_0 - \mathbf{x}_i^T \boldsymbol{\lambda}_0 - \int_T \mu_i(t) \theta_0(t) dt \right)^2 \right]\end{aligned}$$

where,  $\mathbf{x}_i$  is a vector of baseline covariate including subject specific residual variance  $\frac{v_i}{v_i-2} \sigma_i^2$ . Since there is no closed form of the full conditional posterior density, the draws for  $\sigma_i^2, i = 1, \dots, n$  at each iteration of the Gibbs sampling are obtained using the inverse cumulative distribution sampling method.

- **update**  $m_{ij}, j = 1, \dots, n_i$ , given that  $m_{ij} \sim \text{gamma}(v/2, v/2)$  where  $v/2$  and  $v/2$  are the shape and rate parameter in gamma distribution, then the full conditional posterior distribution for  $m_{ij}$  is  $m_{ij} \sim \text{gamma} \left( \frac{v+1}{2}, \frac{1}{2} \left( \frac{(y_{ij} - \mu(t_{ij}))^2}{\sigma_i^2} + v \right) \right)$ .
- **update** missing longitudinal data: missing  $y_{ij}$  at time  $t_{ij}$  is drawn from  $t(\mu(t_{ij}), \sigma_i^2, v)$ .

## (2) update for outcome probit submodel:

- **update**  $W_i, i = 1, \dots, m$

$$\begin{aligned}[W_i | o_i = 0, \cdot] &\sim N(\eta_i^W, 1) I_{(-\infty, 0)}(\cdot) \\ [W_i | o_i = 1, \cdot] &\sim N(\eta_i^W, 1) I_{(0, \gamma_2)}(\cdot) \\ [W_i | o_i = 2, \cdot] &\sim N(\eta_i^W, 1) I_{(\gamma_2, \infty)}(\cdot)\end{aligned}$$

where,  $\eta_i^W = \alpha_0 - \mathbf{x}_i^T \boldsymbol{\lambda}_0 - \int_T \mu_i(t) \theta_0(t) dt$  and  $\gamma_2$  is the cutoff.

- **update** cutoff  $\gamma_2$ : assuming flat prior on  $\gamma_2$ , then the full conditional posterior density for  $\gamma_2$  is  $\text{Unif}(\text{Max}_{o_i=1} W_i, \text{Min}_{o_i=2} W_i)$ .
- **update**  $(\alpha, \boldsymbol{\lambda})'$ : Assuming independent prior for  $(\alpha, \boldsymbol{\lambda})' \sim \text{MVN}(\boldsymbol{\nu}_{\alpha\lambda}, V_{\alpha\lambda})$ , then the full conditional posterior density  $\sim \text{MVN}(\tilde{\boldsymbol{\nu}}_{\alpha\lambda}, \tilde{V}_{\alpha\lambda})$  where

$$\begin{aligned}\tilde{\boldsymbol{\nu}}_{\alpha\lambda} &= \left[ V_{\alpha\lambda}^{-1} + \sum_{i=1}^n \mathbf{z}_i \mathbf{z}_i' \right]^{-1} \left[ V_{\alpha\lambda}^{-1} \boldsymbol{\nu}_{\alpha\lambda} + \sum_{i=1}^n \left( W_i - \int_T \mu_i(t) \theta_0(t) dt \right) \mathbf{z}_i \right] \\ \tilde{V}_{\alpha\lambda} &= \left[ V_{\alpha\lambda}^{-1} + \sum_{i=1}^n \mathbf{z}_i \mathbf{z}_i' \right]^{-1}\end{aligned}$$

where  $\mathbf{z}_i = (1, \mathbf{x}_i)'$  is a vector of constant 1, baseline covariates and residual variance  $\frac{v}{v-2} \sigma_i^2$

- **update**  $\tilde{\boldsymbol{\theta}}_0 = (\tilde{\theta}_{01}, \dots, \tilde{\theta}_{0K})'$ : assuming the prior for  $\tilde{\theta}_{01} \sim N(0, v_{\theta_0})$  and first order random walk prior  $\tilde{\theta}_{0k} \sim N(\tilde{\theta}_{k-1}, \tau_{\theta_0}^2)$ ,  $k = 2, \dots, K$ , then the prior for  $\tilde{\boldsymbol{\theta}}_0$  can be written as:  $\pi(\tilde{\boldsymbol{\theta}}) = \left( \frac{1}{\sqrt{2\pi}\tau_{\theta_0}} \right)^{L-1} \exp\{-\frac{1}{2}\tilde{\boldsymbol{\theta}}_0^T V_{\theta_0} \tilde{\boldsymbol{\theta}}_0\}$ , where  $V_{\theta_0} = \begin{pmatrix} v_{\theta_0}^{-1} & \mathbf{0} \\ \mathbf{0} & \mathbf{0} \end{pmatrix} + P_{\theta_0}^T P_{\theta_0} / \tau_{\theta_0}^2$  with  $P_{\theta_0} =$

$$\begin{pmatrix} 1 & -1 & & & \\ & 1 & -1 & & \\ & & \ddots & \ddots & \\ & & & 1 & -1 \end{pmatrix}$$

is the  $(K-1) \times K$  penalty matrix. Then the full conditional posterior density for  $\tilde{\boldsymbol{\theta}}_0$  is  $[\tilde{\boldsymbol{\theta}}_0 | \cdot] \sim MVN(\tilde{\boldsymbol{\nu}}_{\theta_0}, \tilde{V}_{\theta_0})$  where

$$\begin{aligned} \tilde{\boldsymbol{\nu}}_{\theta_0} &= \left[ V_{\theta_0}^{-1} + \sum_{i=1}^n \mathbf{z}_{\theta i} \mathbf{z}_{\theta i}' \right]^{-1} \left[ V_{\theta_0}^{-1} \boldsymbol{\nu}_{\theta_0} + \sum_{i=1}^n (W_i - \alpha_0 - \mathbf{x}_i^T \boldsymbol{\lambda}_0) \mathbf{z}_{\theta i} \right] \\ \tilde{V}_{\theta_0} &= \left[ V_{\theta_0}^{-1} + \sum_{i=1}^n \mathbf{z}_{\theta i} \mathbf{z}_{\theta i}' \right]^{-1} \end{aligned}$$

where  $\mathbf{z}_{\theta i}$  is a vector of  $K$  elements defined such that  $\int_T \mu_i(t) \theta_0(t) dt = \int_T \mathbf{b}_i^T \boldsymbol{\phi}(t) \boldsymbol{\psi}^0(t)^T \tilde{\boldsymbol{\theta}}_0 dt = \mathbf{b}_i^T G_T^0 \tilde{\boldsymbol{\theta}}_0 = \tilde{\boldsymbol{\theta}}_0^T \mathbf{z}_{\theta i}$ .

- **update**  $\tau_{\theta_0}^2$ : assuming  $\tau_{\theta_0}^2 \sim IG(f, g)$ , where  $f$  and  $g$  are the shape and rate parameters, then the full conditional posterior distribution is  $[\tau_{\theta_0}^2 | \cdot] \sim IG\left(f + \frac{K-1}{2}, g + \frac{1}{2}\tilde{\boldsymbol{\theta}}_0^T P_{\theta_0} \tilde{\boldsymbol{\theta}}_0\right)$ , where  $K$  is the number of B spline basis functions to express  $\theta_0(t)$  in the ordinal probit submodel.

