

A Appendix

A.1 Posterior distributions in the MDA algorithm

A.1.1 Posterior distribution of $(d_i, \mathcal{W}_i, \mathbf{y}_{im})$

Let $\tilde{\mathbf{y}}_{im} = (\mathcal{W}_i, \mathbf{y}_{im})$ and $X_i = (x_{i1}, \dots, x_{iq}, \mathbf{y}_{io})'$. Let U_{ij_o} and U_{ij_m} be a partition of the $(Q+s_i) \times 1$ vector $(\theta_j, -1, \theta_{s_i-j})'$ according to the elements in X_i and $\tilde{\mathbf{y}}_{im}$. Let

$y_{ij}^* = -U_{ij_o}' X_i$ for model (10). For model (15), $y_{ij}^* = -U_{ij_o}' X_i - \sum_{k=1}^R \eta_k z_{ikj}$. The posterior distribution of $(d_i, \mathcal{W}_i, \mathbf{y}_{im})$ is given by

$$\begin{aligned} \pi(d_i, \mathcal{W}_i, \mathbf{y}_{im} | \mathbf{y}_{io}, \phi, v) &\propto f(d_i, \mathcal{W}_i) \left[\prod_{j=1}^{s_i} \sqrt{d_i \gamma_j} \right] \exp \left[-\sum_{j=1}^{s_i} \frac{d_i \gamma_j (y_{ij}^* - U_{ij_m}' \tilde{\mathbf{y}}_{im})^2}{2} \right] \\ &\propto d_i^{\frac{v+s_i+1}{2}-1} \exp \left[-d_i \frac{b_d + (\tilde{\mathbf{y}}_{im} - \mu_{w_i})' A_{w_i} (\tilde{\mathbf{y}}_{im} - \mu_{w_i})}{2} \right] \\ &\propto \left[1 + \frac{\frac{(\mathcal{W}_i - \mu_{w11})^2}{U_{w11}^2 b_d / b_a}}{b_a} \right]^{-\frac{b_a+1}{2}} \mathcal{G} \left(d_i \mid \frac{b_a+1}{2}, \frac{b_d + (\mathcal{W}_i - \mu_{w11})^2 L_{w11}^2}{2} \right) \\ &\quad N \left(\mathbf{y}_{im} \mid \mu_{im}, \frac{U_{w22} U_{w22}'}{d_i} \right) \end{aligned} \tag{19}$$

subject to $\mathcal{W}_i > 0$, where $\tilde{m} = m_i + 1$, A_0 is a $\tilde{m}_i \times \tilde{m}_i$ matrix with 1 at its (1, 1) entry

and 0 elsewhere, $A_{w_i} = A_0 + \sum_{j=1}^{s_i} \gamma_j U_{ij_m}' U_{ij_m}$, the lower triangle matrix

$L_{w_i} = \begin{bmatrix} L_{w11} & \boldsymbol{\theta} \\ L_{w21} & L_{w22} \end{bmatrix}$ satisfies $A_{w_i} = L_{w_i}' L_{w_i}$, L_{w11} is a scalar, $U_{w_i} = \begin{bmatrix} U_{w11} & \boldsymbol{\theta} \\ U_{w21} & U_{w22} \end{bmatrix} = L_{w_i}^{-1}$,

$B_{w_i} = \sum_{j=1}^{s_i} \gamma_j U_{ij_m}' y_{ij}^*$, $C_{w_i} = U_{w_i}' B_{w_i}$, and $\mu_{w_i} = A_{w_i}^{-1} B_{w_i} = U_{w_i} C_{w_i} = \begin{bmatrix} \mu_{w11} \\ \mu_{w12} \end{bmatrix}$,

$\mu_{im} = \mu_{w12} + U_{w21} L_{w11} (\mathcal{W}_i - \mu_{w11})$, $b_a = v + o_i$, $B_{w_i}' A_{w_i}^{-1} B_{w_i} = C_{w_i}' C_{w_i}$ and

$b_d = \nu + \sum_{j=1}^{s_i} \gamma_j y_{ij}^{*2} - C_{w_i}' C_{w_i}$. In SAS IML, L_{w_i} can be computed can use the following syntax

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index=mtilde:1;
Lwi = (root(Awi[index,index]))[index,index];
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Equation (19) implies that we can draw $(\mathcal{W}_i, d_i, y_{im})$ sequentially from

$$\begin{aligned} d_i^* &\sim \mathcal{G}\left(\frac{b_a}{2}, \frac{b_d}{2}\right), \quad \mathcal{W}_i | d_i^* \sim N^+\left(\mu_{wi1}, \frac{U_{wi1}^2}{d_i}\right), \\ d_i | \mathcal{W}_i, d_i^* &\sim \mathcal{G}\left(\frac{b_a+1}{2}, \frac{b_d + (\mathcal{W}_i - \mu_{wi1})^2 L_{wi1}^2}{2}\right), \\ y_{im} | d_i, \mathcal{W}_i, d_i^* &\sim N\left(\mu_{im}, \frac{U_{w22} U_{w22}'}{d_i}\right). \end{aligned} \quad (20)$$

The marginal distribution of \mathcal{W}_i is $t^+\left(\mu_{wi1}, U_{wi1}^2 \frac{b_d}{b_a}, b_a\right)$, but the marginal distribution of d_i is not gamma due to the restriction $\mathcal{W}_i > 0$.

The posterior distribution of y_{im} can be equivalently written as

$$y_{im} | d_i, \mathcal{W}_i, y_{io}, \phi, V \sim N\left(\mu_{im}, \frac{V_{y_m}}{d_i}\right), \quad (21)$$

where h is the index of the first missing observation for subject i in pattern s , \tilde{U}_{ij_m} is a sub-vector of $(\theta_j, -1, \theta_{s_i-j})'$ corresponding to the elements in $y_{im}, y_{ij}^{**} = y_{ij}^* - \psi_j \mathcal{W}_i$, $V_{y_m} = (\sum_{j=h}^s \gamma_j \tilde{U}_{ij_m} \tilde{U}_{ij_m}')$ and $\mu_{im} = \hat{V}_{y_m} \sum_{j=h}^s \gamma_j \tilde{U}_{ij_m} y_{ij}^{**}$.

A.1.2 Posterior distribution of g in step PX1

The posterior distribution of g with a Harr prior g^{-1} and Jacobian g^{n-p} is

$$\begin{aligned}
\text{pos}(g) &\propto g^{n-p} g^{-1} \pi \left(Y_m, gd_1, \dots, gd_n, W, \boldsymbol{\theta}_1, \frac{\gamma_1}{g}, d_{1_\psi}, \dots, \boldsymbol{\theta}_p, \frac{\gamma_p}{g}, d_{p_\psi}, \nu | Y_o \right) \\
&\propto g^{n-p} g^{-1} \prod_{i=1}^n \left\{ (gd_i)^{\frac{\nu+1}{2}-1} \exp \left[-g \frac{d_i(\nu + \mathcal{W}_i^2)}{2} \right] \right\} \\
&\quad \prod_{j=1}^p \left\{ \left(\frac{\gamma_j}{g} \right)^{\frac{n_w+2j+r-p-3}{2}} \exp \left[-\frac{\gamma_j \tilde{\boldsymbol{\theta}}_j' E_j \tilde{\boldsymbol{\theta}}_j}{2g} \right] \right\} \\
&\propto g^{\frac{n(1+\nu)-p(n_w+r)}{2}-1} \exp \left[-g \frac{\sum_{i=1}^n d_i(\nu + \mathcal{W}_i^2)}{2} \right] \exp \left[-\frac{\sum_{j=1}^p \gamma_j \tilde{\boldsymbol{\theta}}_j' E_j \tilde{\boldsymbol{\theta}}_j}{2g} \right].
\end{aligned}$$

A.1.3 Posterior distribution of h in step PX2

The posterior distribution of h with a Harr prior h^{-1} and Jacobian h^{n-p} is

$$\text{pos}(h) \propto h^{n-p} h^{-1} \exp \left(-h^2 \frac{\sum_{i=1}^n d_i \mathcal{W}_i^2}{2} \right) \exp \left(-\frac{\sum_{j=1}^p \gamma_j \underline{\psi}_j^2 \frac{4d_{j_\psi}}{\pi^2}}{2h^2} \right).$$

The posterior distribution of $H = h^2$ is

$$\text{pos}(H) \propto \text{pos}(h) \left| \frac{\partial h}{\partial H} \right| \propto H^{\frac{n-p-1}{2}} \exp \left(-H \frac{\sum_{i=1}^n d_i \mathcal{W}_i^2}{2} \right) \exp \left(-\frac{\sum_{j=1}^p \gamma_j \underline{\psi}_j^2 \frac{4d_{j_\psi}}{\pi^2}}{2H} \right).$$

A.1.4 Prior and posterior distributions of ν

We firstly derive the PC prior for ν . The Kullback-Leibler (KL) distance between the multivariate t distribution $t(\boldsymbol{\mu}, \frac{\nu-2}{\nu} \Sigma, \nu)$ and the normal distribution $N(\boldsymbol{\mu}, \Sigma)$ is

$$\begin{aligned}
\text{KL}(\nu) &= \int t\left(\mathbf{x} | \boldsymbol{\mu}, \frac{\nu-2}{\nu}\Sigma, \nu\right) \log t\left(\mathbf{x} | \boldsymbol{\mu}, \frac{\nu-2}{\nu}\Sigma, \nu\right) d\mathbf{x} \\
&\quad - \int t\left(\mathbf{x} | \boldsymbol{\mu}, \frac{\nu-2}{\nu}\Sigma, \nu\right) \log \phi(\mathbf{x} | \boldsymbol{\mu}, \Sigma) d\mathbf{x} \\
&= \frac{p}{2} \left[1 + \log\left(\frac{2}{\nu-2}\right) \right] + \log \Gamma\left(\frac{\nu+p}{2}\right) - \log \Gamma\left(\frac{\nu}{2}\right) \\
&\quad - \frac{\nu+p}{2} \left[\Psi\left(\frac{\nu+p}{2}\right) - \Psi\left(\frac{\nu}{2}\right) \right].
\end{aligned}$$

since the first integration equals

$$\log \Gamma\left(\frac{\nu+p}{2}\right) - \log \Gamma\left(\frac{\nu}{2}\right) - \frac{\nu+p}{2} \left[\Psi\left(\frac{\nu+p}{2}\right) - \Psi\left(\frac{\nu}{2}\right) \right] - \frac{1}{2} \log |\Sigma| - \frac{p}{2} \log(\nu-2) - \frac{p}{2} \log(\pi)$$

by Kotz and Nadarajah [76], and the second integration equals

$$-\frac{p}{2} \log(2\pi) - \frac{1}{2} \log |\Sigma| - \frac{p}{2}.$$

By the definition of the PC prior [52], the density is $\lambda \exp(-\lambda d(\nu)) \left| \frac{\partial d(\nu)}{\partial \nu} \right|$, where $d(\nu) = \sqrt{2\text{KL}(\nu)}$.

The posterior distribution of ν is given by

$$\begin{aligned}
\pi(\nu | \Sigma, \boldsymbol{\psi}, \boldsymbol{\theta}_j, \gamma_j) &\propto \pi(\nu) \prod_{i=1}^n t_p(\mathbf{y}_{io} | \boldsymbol{\mu}_{io}, \Omega_{io}, \nu) \\
&\quad T_{\nu+o_i} \left[\lambda_{io}^*(\mathbf{y}_{io} - \boldsymbol{\mu}_{io}) \sqrt{\frac{\nu+o_i}{\nu + (\mathbf{y}_{io} - \boldsymbol{\mu}_{io})' \Omega_{io}^{-1} (\mathbf{y}_{io} - \boldsymbol{\mu}_{io})}} \right] I(\nu > \nu_l),
\end{aligned}$$

where $t_p(\mathbf{y}_{io} | \boldsymbol{\mu}_{io}, \Omega_{io}, \nu) \propto \frac{\Gamma(\frac{\nu+o_i}{2})}{\Gamma(\frac{\nu}{2}) \nu^{o_i/2}} [1 + (\mathbf{y}_{io} - \boldsymbol{\mu}_{io})' \Omega_{io}^{-1} (\mathbf{y}_{io} - \boldsymbol{\mu}_{io}) / \nu]^{-\frac{\nu+o_i}{2}}$, and

$$\lambda_{io}^* = \Sigma_{io}^{-1} \boldsymbol{\psi}_{io} / \sqrt{1 + \boldsymbol{\psi}_{io}' \Sigma_{io}^{-1} \boldsymbol{\psi}_{io}}.$$

For subjects with no intermittent missing data, the skew-t density function can be computed without matrix inversion using the following relationship

$$\lambda_{io}^*(\mathbf{y}_{io} - \boldsymbol{\mu}_{io}) = \sum_{t=1}^s \lambda_t R_t \underline{\boldsymbol{\psi}}_t \text{ and}$$

$(\mathbf{y}_{io} - \boldsymbol{\mu}_{io})' \Omega_{io}^{-1} (\mathbf{y}_{io} - \boldsymbol{\mu}_{io}) = \sum_{t=1}^s \lambda_t r_t^2 - [\sum_{t=1}^s \lambda_t r_t \underline{\psi}_t]^2 / [1 + \sum_{t=1}^s \lambda_t \underline{\psi}_t^2]$, where
 $r_j = y_{ij} - \sum_{t=1}^{j-1} \beta_{jt} y_{it} - \sum_{k=1}^Q \alpha_{kj} x_{ik}$ for model (10) and $r_j = y_{ij} - \sum_{t=1}^{j-1} \beta_{jt} y_{it} - \sum_{k=1}^R \eta_k z_{ik} - \sum_{k=1}^Q \alpha_{kj} x_{ik}$
for model (15).

The candidate ν^* is generated from $\log(\nu^* - \nu_l) \sim N[\log(\nu - \nu_l), c^2]$. It will be

accepted with probability $\alpha_\nu = \min \left\{ 1, \frac{(\nu^* - \nu_l) \pi(\nu^*) \prod_{i=1}^n f(\mathbf{y}_{io} | \boldsymbol{\theta}_i, \gamma_i, \nu^*)}{(\nu - \nu_l) \pi(\nu) \prod_{i=1}^n f(\mathbf{y}_{io} | \boldsymbol{\theta}_i, \gamma_i, \nu)} \right\}$.

A.2 Adaption of the MDA algorithm for MMRM-sn

For MMRM-sn (i.e. $\nu \equiv \infty$), the MDA algorithm \mathcal{A} can easily adapted by ignoring steps P2 and PX1, and setting $d_i \equiv 1$ and $d_i^* \equiv 1$ in drawing $(\boldsymbol{\theta}_j, \gamma_j, d_{\psi_j}, \rho_j)$'s and $(\mathcal{W}_i, \mathbf{y}_{im})$'s. For example, $(\mathcal{W}_i, \mathbf{y}_{im})$ in step I can be imputed by modifying the posterior distribution (20) as

$$\mathcal{W}_i \sim N^+(\mu_{wi1}, U_{w11}^2) \text{ and } \mathbf{y}_{im} | \mathcal{W}_i \sim N(\mu_{im}, U_{w22} U_{w22}').$$

A.3 Adaption of the MDA algorithm for MMRM-t

For MMRM-t (i.e. $\underline{\psi}_j \equiv 0$ for $j = 1, \dots, p$, $\mathcal{W}_i \equiv 0$), the MDA algorithm \mathcal{A} needs the following modifications:

1. Step PX2 is no longer needed.
2. Step P1: Remove \mathcal{W}_i from the model. Set $d_{\psi_j} \equiv 0$ and $\boldsymbol{\theta}_j = (\alpha_{1j}, \dots, \alpha_{qj}, \beta_{j1}, \dots, \beta_{j,j-1})'$. Sample $(\boldsymbol{\theta}_j, \gamma_j)$'s from

$$\pi(\boldsymbol{\theta}_j, \gamma_j | d_i)'s, \nu, Y_o, Y_m) \propto \gamma_j^{\frac{n_w + 2j + r^* - p - 1}{2}} \exp \left[-\frac{\gamma_j}{2} \tilde{\boldsymbol{\theta}}_j' (E_j + \sum_{i \leq n_j} d_i \tilde{\mathbf{x}}_{ij} \tilde{\mathbf{x}}_{ij}') \tilde{\boldsymbol{\theta}}_j \right]$$

for $j = 1, \dots, p$, where E_j is the $(q+j) \times (q+j)$ leading principle submatrix of the $(q+p) \times (q+p)$ matrix $E = \begin{bmatrix} M & M\alpha_0' \\ \alpha_0 M & \alpha_0 M\alpha_0' + A_w \end{bmatrix}$ and r^* is the rank of M . If the

inverse Wishart or Jeffrey's prior (with fixed A_w and n_w) instead of the hierarchical prior of Huang and Wand [49] is used, step P0 shall be ignored.

3. Step I: draw $d_i \sim \mathcal{G}\left(\frac{b_a}{2}, \frac{b_d}{2}\right)$ and $\mathbf{y}_{im} \sim N(\mu_{w_i}, (d_i A_{w_i})^{-1})$ since

$$\begin{aligned} \pi(d_i, \mathbf{y}_{im} | \mathbf{y}_{io}, \nu, \phi) &\propto f(d_i) d_i^{\frac{s_i}{2}} \exp\left[-\frac{\sum_{j=1}^{s_i} d_i \gamma_j (y_{ij}^* - \tilde{U}_{ij_m}' \mathbf{y}_{im})^2}{2}\right] \\ &\propto \left\{ d_i^{\frac{\nu+o_i}{2}-1} \exp\left[-\frac{d_i b_d}{2}\right] \right\} \left\{ d_i^{\frac{m_i}{2}} \exp\left[-\frac{d_i (\mathbf{y}_{im} - \mu_{w_i})' A_{w_i} (\mathbf{y}_{im} - \mu_{w_i})}{2}\right] \right\}, \end{aligned}$$

where $f(d_i) \propto d_i^{\nu/2-1} \exp(-d_i \nu)$, $A_{w_i} = \sum_{j=1}^{s_i} \gamma_j \tilde{U}_{ij_m}' \tilde{U}_{ij_m}$, $B_{w_i} = \sum_{j=1}^{s_i} \gamma_j \tilde{U}_{ij_m}' y_{ij}^*$, $\mu_{w_i} = A_{w_i}^{-1} B_{w_i}$,

$$b_a = \nu + o_i \text{ and } b_d = \nu + \sum_{j=1}^{s_i} \gamma_j y_{ij}^{*2} - B_{w_i}' A_{w_i}^{-1} B_{w_i}.$$

4. Step PX1: g is randomly drawn from

$$g^{\frac{n\nu-p(n_w+r^*)}{2}-1} \exp\left[-g \frac{\nu \sum_{i=1}^n d_i}{2}\right] \exp\left[-\frac{\sum_{j=1}^p \gamma_j \tilde{\boldsymbol{\theta}}_j' E_j \tilde{\boldsymbol{\theta}}_j}{2g}\right].$$