## Appendix E: SAS PROC MIXED Syntax

We summarize the basic syntax of SAS proc mixed. The usual linear mixed effects model is

$$\mathbf{Y}_i = \mathbf{X}_i \boldsymbol{\beta} + \mathbf{Z}_i \mathbf{b}_i + \mathbf{e}_i$$

$$E(\boldsymbol{b}_i|\boldsymbol{x}_i) = \boldsymbol{0}$$
,  $var(\boldsymbol{b}_i|\boldsymbol{x}_i) = \boldsymbol{D}$ ,  $E(\boldsymbol{e}_i|\boldsymbol{x}_i,\boldsymbol{b}_i) = E(\boldsymbol{e}_i|\boldsymbol{x}_i) = \boldsymbol{0}$ ,  $var(\boldsymbol{e}_i|\boldsymbol{x}_i,\boldsymbol{b}_i) = var(\boldsymbol{e}_i|\boldsymbol{x}_i) = \boldsymbol{R}_i$ 

(proc mixed does not directly accommodate specifications of  $\mathbf{R}_i$  that depend on  $\mathbf{b}_i$ ). The model is usually written in software documentation in a streamlined form by "stacking" the contributions from each individual. Define

$$\mathbf{Y} = \begin{pmatrix} \mathbf{Y}_1 \\ \mathbf{Y}_2 \\ \vdots \\ \mathbf{Y}_m \end{pmatrix}, \quad \mathbf{e} = \begin{pmatrix} \mathbf{e}_1 \\ \mathbf{e}_2 \\ \vdots \\ \mathbf{e}_m \end{pmatrix}, \quad \mathbf{X} = \begin{pmatrix} \mathbf{X}_1 \\ \mathbf{X}_2 \\ \vdots \\ \mathbf{X}_m \end{pmatrix}, \quad \mathbf{R} = \begin{pmatrix} \mathbf{R}_1 & \mathbf{0} & \cdots & \mathbf{0} \\ \mathbf{0} & \mathbf{R}_2 & \cdots & \mathbf{0} \\ \vdots & \vdots & \ddots & \vdots \\ \mathbf{0} & \mathbf{0} & \cdots & \mathbf{R}_m \end{pmatrix},$$

$$\boldsymbol{b} = \begin{pmatrix} \boldsymbol{b}_1 \\ \boldsymbol{b}_2 \\ \vdots \\ \boldsymbol{b}_m \end{pmatrix}, \quad \boldsymbol{Z} = \begin{pmatrix} \boldsymbol{Z}_1 & \boldsymbol{0} & \cdots & \boldsymbol{0} \\ \boldsymbol{0} & \boldsymbol{Z}_2 & \cdots & \boldsymbol{0} \\ \vdots & \vdots & \ddots & \vdots \\ \boldsymbol{0} & \boldsymbol{0} & \cdots & \boldsymbol{Z}_m \end{pmatrix}, \quad \widetilde{\boldsymbol{D}} = \begin{pmatrix} \boldsymbol{D} & \boldsymbol{0} & \cdots & \boldsymbol{0} \\ \boldsymbol{0} & \boldsymbol{D} & \cdots & \boldsymbol{0} \\ \vdots & \vdots & \ddots & \vdots \\ \boldsymbol{0} & \boldsymbol{0} & \cdots & \boldsymbol{D} \end{pmatrix}.$$

Here,  $\widetilde{\boldsymbol{D}}$  has been displayed in the case where  $\operatorname{var}(\boldsymbol{b}_i|\boldsymbol{x}_i) = \boldsymbol{D}$  for all individuals (so is independent of  $\boldsymbol{x}_i$ ), but can be modified if this is relaxed, as in the dental study with the girls and boys having different matrices  $\boldsymbol{D}_G$  and  $\boldsymbol{D}_B$  (so depending on  $\boldsymbol{a}_i$ ). The model can be written consisely with these definitions as

$$\mathbf{Y} = \mathbf{X}\boldsymbol{\beta} + \mathbf{Z}\mathbf{b} + \mathbf{e}, \quad E(\mathbf{Y}|\tilde{\mathbf{x}}) = \mathbf{X}\boldsymbol{\beta}, \quad \text{var}(\mathbf{Y}|\tilde{\mathbf{x}}) = \mathbf{V} = \mathbf{Z}\tilde{\mathbf{D}}\mathbf{Z}^T + \mathbf{R}.$$
 (B.1)

The SAS documentation refers to  $\tilde{\boldsymbol{D}}$  as  $\boldsymbol{G}$ .

The syntax for proc mixed is geared to the *subject-specific* linear mixed effects model; however, the procedure can also be used to fit *population-averaged* linear models of the form

$$\mathbf{Y}_i = \mathbf{X}_i \boldsymbol{\beta} + \boldsymbol{\epsilon}_i$$

or in "stacked" form

$$\mathbf{Y} = \mathbf{X}\boldsymbol{\beta} + \boldsymbol{\epsilon}$$
,  $E(\mathbf{Y}|\tilde{\mathbf{X}}) = \mathbf{X}\boldsymbol{\beta}$ ,  $var(\boldsymbol{\epsilon}|\tilde{\mathbf{X}}) = var(\mathbf{Y}|\tilde{\mathbf{X}}) = \mathbf{V}$ ,

where V does not have a specific structure; the structure of V is specified fully by the analyst and is not induced by the model.

For either type of model, the form of the **population mean** is either induced (SS model) or is specified explicitly (PA model). The model statement is of course the mechanism by which the analyst specifies the form the population mean, which is  $X_i\beta$  for i or  $X\beta$  for all individuals, stacked, in the usual SAS way.

In the context of a PA model, we have used the repeated statement to specify the overall covariance matrix V. For the SS linear mixed effects model, the repeated statement is used to specify the form of the **within-individual** covariance model  $R_i$  or, equivalently, R above. For this model the random statement is used to specify the assumption on  $var(b_i|x_i)$   $(\widetilde{D})$ .

Here is a summary of the basic form of a call to proc mixed.

```
proc mixed data=dataset method= (ML,REML);
class classification variables;
model response = columns of X / solution;
random columns of Z / type= subject= group= ;
repeated classification variable for time / type= subject= group=
run;
```

proc mixed statement

method=REML is the default; no method= required in this case

model statement

- columns of X are variables (class or continuous) corresponding to variables associated with fixed effects  $\beta$
- Intercept is assumed unless noint option after slash
- ullet solution is an option and must be invoked to get the estimates of eta

## random statement

- Describes the matrix  $\widetilde{\boldsymbol{D}} = \text{var}(\boldsymbol{b}|\widetilde{\boldsymbol{x}})$  (i.e. the matrices  $\text{var}(\boldsymbol{b}_i|\boldsymbol{x}_i)$  making up the blocks of  $\widetilde{\boldsymbol{D}}$
- columns of Z are variables (class or continuous), i.e. variables associated with random effects
- subject= tells mixed what class variable denotes the grouping determining the individuals
- type= allows choice of matrix (e.g. un, unstructured)
- group= allows **D** to be different according to this class variable (e.g. dental study, boys, girls)

## repeated statement

- Describes the matrix  $\mathbf{R} = \text{var}(\mathbf{e}|\tilde{\mathbf{x}})$  (i.e. the matrices  $\mathbf{R}_i = \text{var}(\mathbf{e}_i|\mathbf{x}_i)$ )
- If  $var(\mathbf{e}_i \mathbf{x}_i) = \sigma^2 \mathbf{I}_{n_i}$  is the same for all *i* repeated statement is **NOT** needed
- subject= tells mixed what class variable denotes the grouping determining the individuals
- type= allows choice other than diagonal (e.g. ar(1), cs, etc.)
- The optional classification variable before the slash is for situations with unbalanced data and nondiagonal type so that observations can be correctly attributed to the times at which they were taken
- group= allows  $\mathbf{R}_i$  to be different depending on group membership (e.g. dental study,  $\text{var}(\mathbf{e}_i|\mathbf{x}_i) = \sigma_G^2 \mathbf{I}_{n_i}$  girls,  $\text{var}(\mathbf{e}_i|\mathbf{x}_i) = \sigma_B^2 \mathbf{I}_{n_i}$  boys)

The foregoing syntax makes clear that, to implement a linear PA model using proc mixed with the repeated statement, we simply make a correspondence between this model and the model (B.1) with **no** random effects **b**, From purely **operational** point of view (but **not** an **interpretation** point of view), the models have the same structure – a mean plus a deviation with components of length  $n_i$ , each of which has a covariance matrix. Thus, purely to specify these covariance matrices for the PA model, the repeated statement can be used.

See the SAS documentation for proc mixed for much more detail on the use of these statements and available options.