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*****
Analysis of the dental study data by fitting a general linear
regression model in time and gender structures using PROC MIXED.
- the repeated measurement factor is age (time)
- there is one "treatment" factor, gender

For each gender, the "full" mean model is a straight line in time.

We use the REPEATED statement of PROC MIXED with the
TYPE= options to fit the model assuming several different
covariance structures.

*****/
options ls=80 ps=59 nodate; run;
/*****
Read in the data set (See Example 1 of Chapter 4)

*****
data dent1; infile 'dental.dat';
input obsno child age distance gender;
ag = age*gender;
run;
/*****
Sort the data so we can do gender-by-gender fits.

*****/
proc sort data=dent1; by gender; run;
/*****
First the straight line model separately for each gender and
simultaneously for both genders assuming that the covariance
structure of a data vector is diagonal with constant variance; that
is, use ordinary least squares for each gender separately and
then together.

*****/
title "ORDINARY LEAST SQUARES FITS BY GENDER";
proc reg data=dent1; by gender;
model distance = age;

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run;
title "ORDINARY LEAST SQUARES FIT WITH BOTH GENDERS";
proc reg data=dent1;
model distance = gender age ag;
run;
/*****
Now use PROC MIXED to fit the more general regression model with
assumptions about the covariance matrix of a data vector. For all
of the fits, we use usual normal maximum likelihood (ML) rather
than restricted maximum likelihood (REML), which is the default.

We do this for each gender separately first using the unstructured
assumption. The main goal is to get insight into whether it might
be the case that the covariance matrix is different for each gender
(e.g. variation is different for each).

The SOLUTION option in the MODEL statement requests that the
estimates of the regression parameters be printed.

The R option in the REPEATED statement as used here requests that
the covariance matrix estimate be printed in matrix form. The
RCORR option requests that the corresponding correlation matrix
be printed.

*****/
* unstructured covariance matrix;
title "FIT WITH UNSTRUCTURED COVARIANCE FOR EACH GENDER";
proc mixed method=ml data=dent1; by gender;
class child;
model distance = age / solution;
repeated / type = un subject=child r rcorr;
run;
/*****
Now do the same analyses with both genders simultaneously.
Consider several models, allowing the covariance matrix to
be either the same or different for each gender using the
GROUP = option, which allows for different covariance
parameters for each GROUP (genders here).

For the fit using TYPE = CS (Compound symmetry) assumed the
same for each group, we illustrate how to fit the two
different parameterizations of the full model. For all other
fits, we just use the second parameterization.

*****/

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The CHISQ option in the MODEL statement requests that the Wald chi-square test statistics be printed for certain contrasts of the regression parameters (see the discussion of the OUTPUT). We only use this for the second parameterization -- the TESTS OF FIXED EFFECTS are tests of interest (different intercepts, slopes) in this case.

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*****
* compound symmetry with separate intercept and slope for:
* each gender;
title "COMMON COMPOUND SYMMETRY STRUCTURE";
proc mixed method=ml data=dent1;
class gender child;
model distance = gender age*gender / noint solution ;
repeated / type = cs subject = child r rcorr;
run;
* compound symmetry with the "difference" parameterization;
* same for each gender;
title "COMMON COMPOUND SYMMETRY STRUCTURE";
proc mixed method=ml data=dent1;
class gender child;
model distance = gender age*gender / solution chisq;
repeated / type = cs subject = child r rcorr;
run;
* ar(1) same for each gender;
title "COMMON AR(1) STRUCTURE";
proc mixed method=ml data=dent1;
class gender child;
model distance = gender age*gender / solution chisq;
repeated / type = ar(1) subject=child r rcorr;
run;
* one-dependant same for each gender;
title "COMMON ONE-DEPENDENT STRUCTURE";
proc mixed method=ml data=dent1;
class gender child;
model distance = gender age*gender / solution chisq;
repeated / type = toep(2) subject=child r rcorr;
run;
* compound symmetry, different for each gender;
title "SEPARATE COMPOUND SYMMETRY FOR EACH GENDER";
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proc mixed method=ml data=dent1;
class gender child;
model distance = gender age*gender / solution chisq;
repeated / type = cs subject=child r rcorr group=gender;
run;
```

```
* ar(1), different for each gender;
title "SEPARATE AR(1) FOR EACH GENDER";
proc mixed method=ml data=dent1;
class gender child;
model distance = gender age*gender / solution chisq;
repeated / type = ar(1) subject=child r rcorr group=gender;
run;
```

```
* one-dependent, different for each gender;
title "SEPARATE ONE-DEPENDENT FOR EACH GENDER";
proc mixed method=ml data=dent1;
class gender child;
model distance = gender age*gender / solution chisq;
repeated / type = toep(2) subject=child r rcorr group=gender;
run;
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*****
Examination of the AIC, BIC, and loglikelihood ratios from the
above fits indicates that
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- a model that allows a separate covariance matrix of the same type for each gender is preferred

- the compound symmetry structure for each gender is preferred

Thus, for this model, we fit

- the full model again, now asking for the covariance matrix of beta-hat to be printed using the COVB option;

- the reduced model (equal slopes)

- the full model using REML

This will allow a "full" vs. "reduced" likelihood ratio test of equal slopes to be performed (by hand from the output).

We fit the first parameterization this time, so that the estimates are interpreted as the gender-specific intercepts and slopes.

Thus, the TESTS OF FIXED EFFECTS in the output should be disregarded.

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*****
* full model again with covariance matrix of betahat printed:
title "FULL MODEL WITH COMPOUND SYMMETRY FOR EACH GENDER";
proc mixed method=ml data=dent1;
class gender child;
model distance = gender*age / noint solution covb;
repeated / type=cs subject=child r rcorr group=gender;
run;

* reduced model;
title "REDUCED MODEL WITH COMPOUND SYMMETRY FOR EACH GENDER";
proc mixed method=ml data=dent1;
class gender child;
model distance = gender age / noint solution covb;
repeated / type=cs subject=child r rcorr group=gender;
run;

* full model using REML (the default, so no METHOD= is specified);
* use ESTIMATE statement to estimate the mean for a boy of age 11;
title "FULL MODEL WITH COMPOUND SYMMETRY FOR EACH GENDER, REML";
proc mixed data=dent1;
class gender child;
model distance = gender*age / noint solution covb;
repeated / type=cs subject=child r rcorr group=gender;
estimate 'boy at 11' gender 0 1 gender*age 0 11;
run;

* also fit full model in first parameterization to get chi-square tests;
title "FULL MODEL, DIFFERENCE PARAMETERIZATION";
proc mixed method=ml data=dent1;
class gender child;
model distance = gender age gender*age / solution chisq covb;
repeated / type=cs subject=child r rcorr group=gender;
run;

```