

BIOS 755: Fixed versus Random effect models and Longitudinal versus Cross-sectional Effects

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Introduction

- ▶ Today we are going to be talking about time-varying and time-invariant covariates.
- ▶ Let \mathbf{X}_{ij} denote the time-varying covariates and $\mathbf{W}_{ij} = \mathbf{W}_i$ the time-invariant covariates.
- ▶ To analyze such data we could use

$$Y_{ij} = \mathbf{X}_{ij}\beta + \mathbf{W}_i\gamma + \alpha_i + e_{ij}$$

where $\mathbf{e}_i \sim N(0, \sigma^2)$.

Fixed effect model

- ▶ What if we didn't assume α_i was random, but rather estimated it from the data. What kind of model would we have?
- ▶ An issue with this model is that we couldn't estimate the time-invariant covariate effects:
 - ▶ **can't estimate both γ and the α_i 's.**

$$Y_{ij} = \mathbf{X}_{ij}\boldsymbol{\beta} + \mathbf{W}_i\boldsymbol{\gamma} + \alpha_i + e_{ij}$$

Fixed effect model

$$Y_{ij} = \mathbf{X}_{ij}\beta + \mathbf{W}_i\gamma + \alpha_i + e_{ij}$$

- ▶ We could estimate β
- ▶ For example, if we only had two observations:

$$(Y_{i2} - Y_{i1}) = (\mathbf{X}_{i2} - \mathbf{X}_{i1})'\beta + e_{i2}^*$$

which could be fitted using OLS of $(Y_{i2} - Y_{i1})$ on $(\mathbf{X}_{i2} - \mathbf{X}_{i1})$.

- ▶ Notice that this model removes the potential for bias due to confounding by all measured and unmeasured time-invariant characteristics of individuals (as long as the effect is constant over time).

Fixed effect model

- ▶ This approach can be expanded with the mean-centered model

$$Y_{ij}^* = \mathbf{X}_{ij}^* \boldsymbol{\beta} + e_{ij}$$

where $Y_{ij}^* = Y_{ij} - \bar{Y}_i$ and $\mathbf{X}_{ij}^* = \mathbf{X}_{ij} - \bar{\mathbf{X}}_i$.

- ▶ Or with the *first difference* model for $j = 2, \dots, n_i$

$$Y_{ij}^\dagger = \mathbf{X}_{ij}^\dagger \boldsymbol{\beta}^\dagger + e_{ij}^\dagger$$

where $Y_{ij}^\dagger = Y_{ij} - Y_{i1}$ and $\mathbf{X}_{ij}^\dagger = \mathbf{X}_{ij} - \mathbf{X}_{i1}$.

Fixed effect model

- ▶ This model removes all the variation due to time-invariant covariates.
- ▶ If the random intercept model is correct, the correlation e_{ij} can be ignored
 - ▶ assuming only time-invariant covariates are causing the dependence between measurements.
- ▶ To fit this model we'll use `proc glm` with an independent correlation matrix.

GO TO EXAMPLE

Random effect versus Fixed effect model

Some technical differences:

- ▶ The fixed effect model allows for correlation between α_i and \mathbf{X}_{ij} , and α_i and \mathbf{W}_i .
- ▶ The random effect model does not allow for this correlation and biases in β arise when violated.
- ▶ The random effect model allows for estimation of time-invariant fixed-effects, the fixed effect model does not.
- ▶ The random effect model is more efficient than the fixed effect model.
- ▶ Individuals must have more than 1 observation in the fixed effect model.

LONGITUDINAL VS CROSS-SECTIONAL EFFECTS

Introduction

- ▶ It is possible to allow for **longitudinal** and **cross-sectional** effects in longitudinal analyses.
- ▶ Such an approach acknowledges the two distinct sources of variation in a covariate:
 - ▶ one based on within-subject variation, and
 - ▶ one based on between-subject variation.
- ▶ Such a model recognized that longitudinal data provide information about
 - ▶ how individuals differ at any one occasion, and
 - ▶ how an individuals response varies over time.
- ▶ Commonly these effects are erroneously combined.

Example

- ▶ Suppose there is a study on the impact of talkativeness and well-being (subjectively measured).
- ▶ There could be a positive association between talkativeness and well-being between-people:
 - ▶ people who are (on average) more talkative than others are (on average) happier than others.
- ▶ There could be a positive association between talkativeness and well-being within-people:
 - ▶ people who are more talkative today (than they usually are) are happier today (than they usually are).

Example

- ▶ **These effects may be equal,**
 - ▶ if a person is 20% more talkative (either between or within), we predict they will score 15% higher on their well-being (on average).

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- ▶ **These effects may be different,**
 - ▶ if person A is 20% more talkative than person B (on average), we predict person A will score 5% higher than person B on their well-being (on average), while
 - ▶ if person A is 20% more talkative today (than they usually are), we predict they will score 20% higher than usual on their well-being (on average).

Example

- ▶ **These effects may be equal,**
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 - ▶ if person A is 20% more talkative today (than they usually are), we predict they will score 20% higher than usual on their well-being (on average).
- ▶ **These effects may be the opposite,**
 - ▶ if person A is 20% more talkative than person B (on average), we predict person A will score 5% higher than person B on their well-being (on average), while
 - ▶ if person A is 20% more talkative today (than they usually are), we predict they will score 10% lower than usual on their well-being (on average).

Example

- ▶ **Between person effects**
 - ▶ **if person A is 20% more talkative than person B (on average), we predict person A will score 5% higher than person B on their well-being (on average), while**
 - ▶ if person A is 20% more talkative today (than they usually are), we predict they will score 20% higher than usual on their well-being (on average).

- ▶ **if person A is 20% more talkative than person B (on average), we predict person A will score 5% higher than person B on their well-being (on average), while**
- ▶ if person A is 20% more talkative today (than they usually are), we predict they will score 10% lower than usual on their well-being (on average).

Example

► Within person effects

- if person A is 20% more talkative than person B (on average), we predict person A will score 5% higher than person B on their well-being (on average), while
- **if person A is 20% more talkative today (than they usually are), we predict they will score 20% higher than usual on their well-being (on average).**
- if person A is 20% more talkative than person B (on average), we predict person A will score 5% higher than person B on their well-being (on average), while
- **if person A is 20% more talkative today (than they usually are), we predict they will score 10% lower than usual on their well-being (on average).**

Longitudinal and cross-sectional information

- ▶ Assessment of within-subject changes in the response due to aging (for example) can only be achieved within a longitudinal study design.
- ▶ What would happen with a cross-sectional design?
- ▶ Recall the Muscatine Coronary Risk Factor (MCRF) study which had five cohorts of children, initially aged 5–7, 7–9, 9–11, 11–13, and 13–15.
- ▶ Goal: determine whether the risk for obesity increases with age.
- ▶ Measurements were taken in 1977, 1979, and 1981.
- ▶ Suppose we have BMI as an outcome of interest.

Model

- ▶ To combine longitudinal and cross-sectional effects we will include both effects in the model.
- ▶ For example, we can use the linear mixed effects model:

$$Y_{ij} = \mathbf{X}_{ij}^* \boldsymbol{\beta}^{(L)} + \mathbf{X}_{i1}' \boldsymbol{\beta}^{(C)} + \mathbf{W}_i' \boldsymbol{\gamma} + \mathbf{Z}_{ij}' \mathbf{b}_i + e_{ij}$$

where $\mathbf{X}_{ij}^* = \mathbf{X}_{ij} - \mathbf{X}_{i1}$.

- ▶ Here, $\boldsymbol{\beta}^{(C)}$ represent the cross-section effects while $\boldsymbol{\beta}^{(L)}$ are the longitudinal effects.
- ▶ **This is one example, another example is using $\mathbf{X}_{ij}^* = \mathbf{X}_{ij} - \bar{\mathbf{X}}_i$ and $\bar{\mathbf{X}}_i$.**

Interpretation

- For the baseline measurement, it's straightforward to show that

$$E(Y_{i1}) = \mathbf{X}'_{i1}\boldsymbol{\beta}^{(C)} + \mathbf{W}'_i\boldsymbol{\gamma}$$

so $\boldsymbol{\beta}^{(C)}$ is the expected difference in the baseline outcome for a 1 unit change in \mathbf{X}_{i1} .

- Further, the model for the within-subject changes is

$$\begin{aligned} E(Y_{ij} - Y_{i1}) &= \mathbf{X}^*_{ij}\boldsymbol{\beta}^{(L)} + \mathbf{X}'_{i1}\boldsymbol{\beta}^{(C)} + \mathbf{W}'_i\boldsymbol{\gamma} - \left(\mathbf{X}^*_{i1}\boldsymbol{\beta}^{(L)} + \mathbf{X}'_{i1}\boldsymbol{\beta}^{(C)} + \mathbf{W}'_i\boldsymbol{\gamma} \right) \\ &= (\mathbf{X}_{ij} - \mathbf{X}_{i1})'\boldsymbol{\beta}^{(L)} \end{aligned}$$

so $\boldsymbol{\beta}^{(L)}$ is the expected within-person change from baseline in the outcome for a 1 unit increase in \mathbf{X}_{ij} from baseline.

Example

- ▶ Let A_{ij} be the age of person i at measurement j and Y_{ij} their BMI. One option is to fit the model

$$Y_{ij} = \beta_0 + \beta_1 A_{ij} + b_{0i} + \epsilon_{ij}$$

- ▶ Separating out the cross-sectional and longitudinal effects of age we have

$$Y_{ij} = \beta_0 + \beta_1^{(L)} A_{ij}^* + \beta_1^{(C)} A_{i1} + b_{0i} + \epsilon_{ij}$$

where $A_{ij}^* = A_{ij} - A_{i1}$ is the change in age from baseline.

- ▶ When there is not a well-defined baseline measurement, I prefer the average model.

Example

- Separating out the cross-sectional and longitudinal effects of age we have

$$Y_{ij} = \beta_0^* + \beta_1^{(L)} A_{ij}^* + \beta_1^{(C)} A_{i1} + b_{0i}^* + \varepsilon_{ij}$$

where $A_{ij}^* = A_{ij} - \bar{A}_i$ and \bar{A}_i is the persons average age in the study.

- For subjects with a baseline age that was 1 unit higher, we expect their baseline outcome to be $\beta_1^{(C)}$ units larger.
- During our study, if an individual ages 1 year from baseline, the expected change in the outcome is $\beta_1^{(L)}$ units from baseline.

Example

- ▶ We can also do this with the average, where our model is

$$Y_{ij} = \beta_0 + \beta_1^{(L)} A_{ij}^* + \beta_1^{(C)} \bar{A}_i + b_{0i} + \varepsilon_{ij}$$

$$\text{with } \bar{A}_i = \frac{\sum_{j=1}^{n_i} A_{ij}}{n_i}.$$

- ▶ For subjects with an average age that was 1 unit higher, we expect their average outcome to be $\beta_1^{(C)}$ units larger.
- ▶ During our study, if an individual ages 1 year, the expected change in the outcome is $\beta_1^{(L)}$ units from their average outcome.

$\beta^{(L)}$ versus $\beta^{(C)}$

- Notice that when $\beta^{(L)} = \beta^{(C)} = \beta$

$$\begin{aligned} Y_{ij} &= \mathbf{X}_{ij}^* \beta^{(L)} + \mathbf{X}_{i1}' \beta^{(C)} + \mathbf{W}_i' \gamma + \mathbf{Z}_{ij}' b_i + e_{ij} \\ &= \mathbf{X}_{ij}' \beta + \mathbf{W}_i' \gamma + \mathbf{Z}_{ij}' b_i + e_{ij} \end{aligned}$$

which is the “standard” mixed effects model

- So the hypothesis test $H_0 : \beta^{(L)} = \beta^{(C)}$ tests whether longitudinal and cross section effects are equal.

Longitudinal and cross-sectional information

- ▶ Could we measure the effect of age using only data from 1977?
- ▶ Differences in $\beta^{(L)}$ versus $\beta^{(C)}$ can arise when there are cohort or period effects.
- ▶ When we assume $\beta^{(L)} = \beta^{(C)} = \beta$, i.e., use the “standard” linear mixed effects model, we get an estimate for β that is a combination of $\beta^{(L)}$ and $\beta^{(C)}$.