

Statistical Methods for Analysis with Missing Data

Lecture 15: identifiability, nonignorability, pattern-mixture models

Mauricio Sadinle

Department of Biostatistics

W UNIVERSITY *of* WASHINGTON

So Far

The approaches that we have covered for handling missing data:

- ▶ Ad-hoc approaches (imputation, complete cases)
- ▶ Frequentist likelihood-based inference
- ▶ Bayesian inference
- ▶ Multiple imputation
- ▶ Inverse-probability weighting

Something they have in common:

- ▶ We have assumed MAR (or MCAR), sometimes avoiding to handle the response mechanism $p(r | z)$

Today's Lecture

- ▶ What if we want to move away from MAR?
- ▶ We will talk about some fundamental issues for handling missing data
 - ▶ Identifiability
 - ▶ Nonignorability
- ▶ This discussion naturally leads to *pattern-mixture models*
- ▶ Reading: Chapter 6 of the lecture notes of Davidian and Tsiatis

Back to the Basics: Lecture 1

- ▶ Y : study variable
- ▶ R : response indicator

$$\underbrace{p(y)}_{\text{what we want}} = p(y | R = 0)\underbrace{p(R = 0)}_{\text{what we can get}} + \underbrace{p(y | R = 1)p(R = 1)}_{\text{what we can get}}$$

We cannot recover $p(y | R = 0)$ nor $p(y)$ from observed data alone

The fundamental problem of inference with missing data: it is impossible without extra, usually untestable, assumptions on how missingness arises

Sample Data

- ▶ The *full-data sample* are independent and identically distributed (i.i.d.) draws from some distribution F

$$\{(Z_i, R_i)\}_{i=1}^n \stackrel{i.i.d.}{\sim} F$$

- ▶ R_i determines the part of Z_i that we get to observe: $Z_{i(R_i)}$
- ▶ We can think of the generative process, for each i :

$$Z_i \implies R_i \implies (Z_{i(R_i)}, R_i)$$

- ▶ In this lecture, we delete the subindex i to talk about
 - ▶ A generic draw from F
 - ▶ What we could recover provided an infinite sample size
 - ▶ Separate *identifiability* issues from *estimation* issues

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Types of Data

- ▶ Full data: (Z, R)
- ▶ Observed data: $(Z_{(R)}, R)$
- ▶ Missing data: $Z_{(\bar{R})}$

Relationship:

$$(Z, R) = (Z_{(\bar{R})}, Z_{(R)}, R)$$

Distributions of Interest

- ▶ Full-data distribution: joint distribution of (Z, R) with density

$$p(z, r) \equiv p(z_{(\bar{r})}, z_{(r)}, r), \text{ for all } r$$

- ▶ Observed-data distribution: joint distribution of $(Z_{(R)}, R)$ with density

$$p(z_{(r)}, r) = \int p(z_{(\bar{r})}, z_{(r)}, r) dz_{(\bar{r})}, \text{ for all } r$$

- ▶ Missing-data distribution, or *extrapolation* distribution: conditional distribution of $Z_{(\bar{R})}$ given $(Z_{(R)}, R)$

$$p(z_{(\bar{r})} \mid z_{(r)}, r) = \frac{p(z_{(\bar{r})}, z_{(r)}, r)}{p(z_{(r)}, r)}, \text{ for all } r$$

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The Full-Data Distribution

- ▶ Joint distribution of (Z, R) with density

$$p(z, r)$$

- ▶ Quantities of interest θ (parameters) depend on the full-data distribution

$$p(z, r) \longrightarrow p(z) = \sum_r p(z, r) \longrightarrow \theta = E[f(Z)] = \int f(z)p(z)dz$$

- ▶ For example, say $f(Z) = Z_j$, then

$$\theta_j = E(Z_j) = \int z_j p(z_j) dz_j = \int z_j p(z) dz$$

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The Observed-Data Distribution

- ▶ Given $R = r$, we observe $Z_{(r)}$
- ▶ We can estimate $p(z_{(r)} | R = r)$ and $p(R = r)$ from observed data
- ▶ The observed-data distribution is all we can hope to recover from data alone

$$p(z_{(r)}, r) = p(z_{(r)} | r)p(r)$$

- ▶ For example, say you can sample indefinitely from the joint distribution of

$$Z = (Z_1, Z_2) \text{ and } R = (R_1, R_2)$$

- ▶ If $R_j = 0$ you don't see the value of Z_j
- ▶ What we can estimate from such data:
 - ▶ $p(R = r), r \in \{0, 1\}^2$
 - ▶ $p(z_1 | R = 10)$
 - ▶ $p(z_2 | R = 01)$
 - ▶ $p(z_1, z_2 | R = 11)$

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The Extrapolation Distribution

- ▶ Given $R = r$, we observe $Z_{(r)}$, but we don't observe $Z_{(\bar{r})}$
- ▶ There is no way of estimating $p(z_{(\bar{r})} | z_{(r)}, r)$ without assumptions

$$\underbrace{p(z_{(\bar{r})}, z_{(r)}, r)}_{\text{what we want}} = \overbrace{p(z_{(\bar{r})} | z_{(r)}, r)}^{\text{how to extrapolate}} \underbrace{p(z_{(r)}, r)}_{\text{what we can get}}$$

- ▶ We say that $p(z_{(\bar{r})} | z_{(r)}, r)$, and therefore $p(z, r)$, are not *identifiable*
- ▶ *Identifying assumptions* explicitly or implicitly amount to constructing $p(z_{(\bar{r})} | z_{(r)}, r)$ from $p(z_{(r)}, r)$

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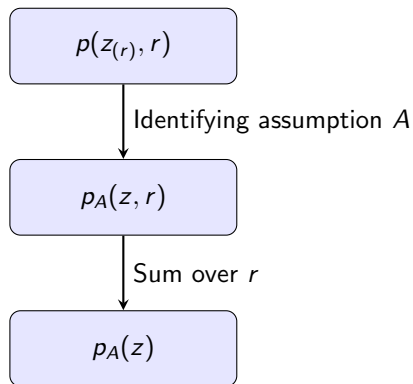
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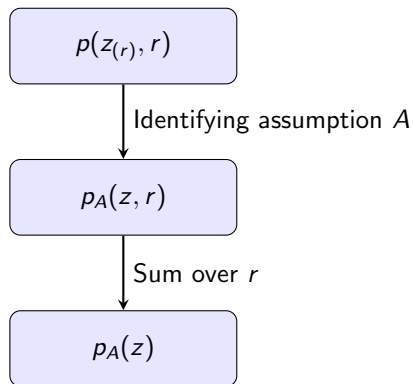
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General Identification Strategy



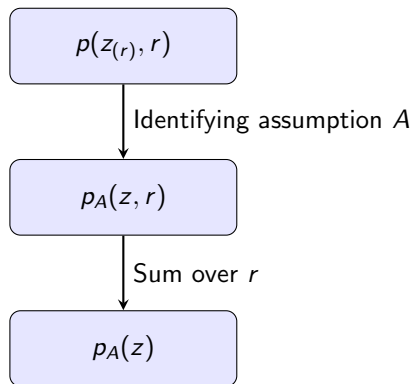
- ▶ Note that MAR (ignorability) gives you a shortcut to go from $p(z(r), r)$ to $p_{MAR}(z)$
- ▶ Otherwise, how do people specify identifying assumptions?

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Factorizations of the Full-Data Distribution

Selection model factorization:

$$p(z, r) = p(r | z)p(z)$$

- ▶ The response mechanism $p(r | z)$ represents the way in which values of study variables get *selected* into the sample
- ▶ Natural factorization when we initially had a model $\{p(z | \theta)\}_\theta$ in mind, say had we not had missing data
- ▶ Allows us to continue using model $\{p(z | \theta)\}_\theta$
- ▶ Identifying assumptions are expressed as restriction on response mechanism $p(r | z)$
- ▶ We have focused on this approach so far under MAR:

$$p(r | z) = p(r | z_{(r)})$$

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Pattern-mixture model factorization:

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- ▶ Requires models for distribution of Z given each value $R = r$
- ▶ Distribution of study variables is obtained as a *mixture* of *pattern-specific* models

$$p(z) = \sum_r p(z | r)p(r)$$

- ▶ This gives an alternative approach for handling missing data

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Pattern-Mixture Models

- ▶ The pattern-mixture model factorization explicitly reveals:

$$\begin{aligned} p(z) &= \sum_r p(z | r) p(r) \\ &= \sum_r p(z_{(\bar{r})} | z_{(r)}, r) p(z_{(r)} | r) p(r) \\ &= \sum_r \underbrace{p(z_{(\bar{r})} | z_{(r)}, r)}_{\text{needs identifying assumption}} \underbrace{p(z_{(r)} | r) p(r)}_{\text{can be estimated from data}} \end{aligned}$$

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Identifying Assumptions for Pattern-Mixture Models

- ▶ Identifying assumptions in the framework of pattern mixture models amount to specifying how to construct

$$\{p(z_{(\bar{r})} | z_{(r)}, r)\}_r$$

from

$$\{p(z_{(r)}, r)\}_r$$

- ▶ Once $p_A(z_{(\bar{r})} | z_{(r)}, r)$ is specified, according to an assumption A , this defines a full-data density

$$p_A(z_{(\bar{r})}, z_{(r)}, r) = p_A(z_{(\bar{r})} | z_{(r)}, r)p(z_{(r)}, r)$$

- ▶ Note that this in turn implies a response mechanism

$$p_A(r | z_{(\bar{r})}, z_{(r)}) = \frac{p_A(z_{(\bar{r})}, z_{(r)}, r)}{\sum_{r'} p_A(z_{(\bar{r})}, z_{(r')}, r')}$$

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$$p_A(z_{(\bar{r})}, z_{(r)}, r) = p_A(z_{(\bar{r})} \mid z_{(r)}, r)p(z_{(r)}, r)$$

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- ▶ Assumptions that lead to response mechanisms that are not particular cases of MAR are *nonignorable*

Identifying Assumptions for Pattern-Mixture Models

- ▶ Identifying assumptions in the framework of pattern mixture models amount to specifying how to construct

$$\{p(z_{(\bar{r})} \mid z_{(r)}, r)\}_r$$

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Comments on Pattern-Mixture Models

Advantages:

- ▶ Identifiability requirements are more explicit than with selection models: easier to understand what is it that you are assuming
- ▶ Provides a natural framework for sensitivity analyses

Limitations:

- ▶ We cannot continue using model $\{p(z | \theta)\}_{\theta}$
- ▶ Parameters of scientific interest do not explicitly appear in the model
- ▶ Requires per-pattern model, say $\{p(z_{(r)} | r, \theta_r)\}_{\theta_r}$
- ▶ For general pattern of nonresponse we would need $2^K - 1$ models, one for each pattern in $\{0, 1\}^K$ (minus $\mathbf{0}_K$)
- ▶ Most developments under this approach assume monotone nonresponse (e.g., dropout)

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Dropout in Longitudinal Study

If missingness comes only from subjects dropping out

- ▶ Missingness patterns are uniquely summarized by the dropout time

$$D = 1 + \sum_{j=1}^T R_j$$

- ▶ The *observed data* are obtained as realizations of

$$(Z_{(D)}, D)$$

where, if $D = d$, $Z_{(d)} = (Z_1, \dots, Z_{d-1})$ and $Z_{(\bar{d})} = (Z_d, \dots, Z_T)$

- ▶ Pattern-mixture model requires modeling
 - ▶ $p(D = d)$: simply take empirical frequency
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A Simple Pattern-Mixture Model Under Dropout

In some situations, the following simple implementation of pattern-mixture models (PMMs) might be reasonable

- ▶ Idea: for each dropout group, model observed data and extrapolate to missing data
- ▶ Example:

- ▶ For each d , fit

$$E(Y_j | D = d) = \beta_{0d} + \beta_{1d}t_j,$$

using data from $j < d$, and predict for $j \geq d$

- ▶ This implies

$$E(Y_j) = E[E(Y_j | D = d)] = \sum_d p(D = d)\beta_{0d} + t_j \sum_d p(D = d)\beta_{1d}$$

- ▶ All parameters $p(d)$, β_{0d} , β_{1d} , $d = 1, \dots, T$, can be directly estimated from the observed data (provided dropout starts at time $D = 3$)
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Identifying Assumptions for PMMs Under Dropout

- ▶ In general, how to obtain $p(z_{(\bar{d})} | z_{(d)}, d)$ from $p(z_{(d)}, d)$?
- ▶ Note that

$$\begin{aligned} p(z_{(\bar{d})} | z_{(d)}, d) &= p(z_d, \dots, z_T | z_1, \dots, z_{d-1}, d) \\ &= \prod_{\ell=d}^T p(z_\ell | z_1, \dots, z_{d-1}, \dots, z_{\ell-1}, d) \\ &= \prod_{\ell=d}^T p(z_\ell | z_{(\ell)}, d) \end{aligned}$$

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The Complete-Case Identifying Assumption

Little (JASA 1993) proposed to tie the extrapolation distributions to the distribution of complete cases:

$$p_{CC}(z_\ell \mid z_{(\ell)}, D = d) \equiv p(z_\ell \mid z_{(\ell)}, D = T + 1),$$

for all $\ell \geq d$, $d = 1, \dots, T$.

- ▶ The distributions for $D = T + 1$ are identifiable from the complete cases
- ▶ This strategy could also be used with nonmonotone missingness
- ▶ **HW4**: say $T = 3$, write down this restriction for $\ell \geq d$, $d = 1, 2, 3$.

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The extrapolation distributions could also be obtained from the closest dropout pattern where ℓ is available:

$$p_{NC}(z_\ell | z_{(\ell)}, D = d) \equiv p(z_\ell | z_{(\ell)}, D = \ell + 1),$$

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- ▶ Among observations with $D = \ell + 1$ we get to observe z_ℓ and $z_{(\ell)}$
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Here, the extrapolation distributions are obtained from all available cases where ℓ is available:

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- ▶ **HW4:** under monotone nonresponse, the AC assumption is equivalent to MAR

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Observational Equivalence

- ▶ Two full-data distributions are said to be *observationally equivalent* if their implied observed-data distributions are the same
- ▶ This is, say I have two full-data distributions

$$p_A(z_{(\bar{r})}, z_{(r)}, r)$$

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$$\int p_A(z_{(\bar{r})}, z_{(r)}, r) dz_{(\bar{r})} = \int p_B(z_{(\bar{r})}, z_{(r)}, r) dz_{(\bar{r})}$$

for all $(z_{(r)}, r)$, then they are *observationally equivalent*

- ▶ **HW4:** the full-data distributions obtained under the CC, NC, and AC restrictions are observationally equivalent
- ▶ This is an important feature in *sensitivity analysis!* (next class)

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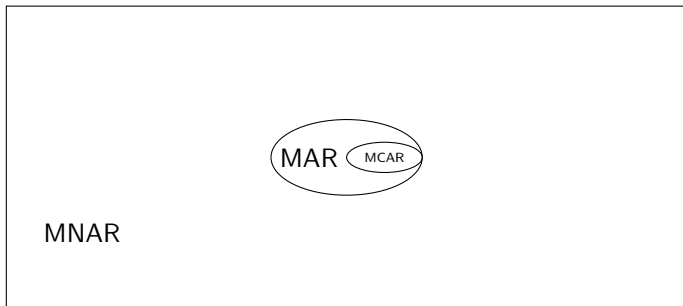
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Summary

Main take-aways from today's lecture:

- ▶ The fundamental problem of inference with missing data: *it is impossible without extra, usually untestable, assumptions on how missingness arises*
- ▶ Pattern-mixture models provide an alternative way of thinking about missing data
- ▶ Remember the universe of missing-data assumptions:



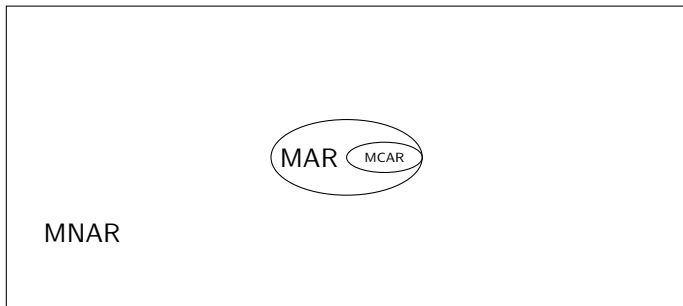
Next lecture:

- ▶ More on nonignorable missing data (MNAR), and sensitivity analysis

Summary

Main take-aways from today's lecture:

- ▶ The fundamental problem of inference with missing data: *it is impossible without extra, usually untestable, assumptions on how missingness arises*
- ▶ Pattern-mixture models provide an alternative way of thinking about missing data
- ▶ Remember the universe of missing-data assumptions:



Next lecture:

- ▶ More on nonignorable missing data (MNAR), and sensitivity analysis