

# Incomplete Data Analysis

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# Likelihood based inference with incomplete data

## Example: bivariate normal data with one variable subject to missingness

- ↪ Consider a medical study where measurements  $Y_1$  are taken at baseline (i.e., at the beginning of the study) and  $Y_2$  at follow-up.
- ↪ All individuals have their outcome recorded at baseline but some have their follow-up outcome missing.

	$Y_1$	$Y_2$	$R_2$
Units			
1			1
2			1
$\vdots$			$\vdots$
$m$			1
$\vdots$			0
$\vdots$			$\vdots$
$n$			0

# Likelihood based inference with incomplete data

Example: bivariate normal data with one variable subject to missingness

- ↪ We consider that  $\mathbf{Y} = (Y_1, Y_2)'$  follows a bivariate normal distribution, that is,  $\mathbf{Y} \sim N_2(\boldsymbol{\mu}, \boldsymbol{\Sigma})$ , with

$$\boldsymbol{\mu} = \begin{pmatrix} \mu_1 \\ \mu_2 \end{pmatrix}, \quad \text{and} \quad \boldsymbol{\Sigma} = \begin{pmatrix} \sigma_1^2 & \sigma_{12} \\ \sigma_{12} & \sigma_2^2 \end{pmatrix}.$$

- ↪ Remember that the density function of a bivariate normal distribution is given by

$$f_{\mathbf{Y}}(\mathbf{y}) = \frac{1}{2\pi} (\det \boldsymbol{\Sigma})^{-1/2} \exp \left\{ -\frac{1}{2} (\mathbf{y} - \boldsymbol{\mu})' \boldsymbol{\Sigma}^{-1} (\mathbf{y} - \boldsymbol{\mu}) \right\}.$$

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- ↪ The full data for the  $i$ th individual is  $(y_{1i}, y_{2i}, r_i)$ , where  $r_i = 1$  if  $y_{2i}$  is observed and  $r_i = 0$  if  $y_{2i}$  is missing, for  $i = 1 \dots, n$ .
- ↪ We assume, without loss of generality, that  $y_{2i}$  is observed for  $i = 1, \dots, m$  (i.e.,  $r_i = 1$ ,  $i = 1, \dots, m$ ) and  $y_{2i}$  is missing for  $i = m + 1, \dots, n$  (i.e.,  $r_i = 0$ ,  $i = m + 1, \dots, n$ ).
- ↪ The full data model is

$$f(\mathbf{y}, \mathbf{r} \mid \boldsymbol{\mu}, \boldsymbol{\Sigma}, \psi) = \prod_{i=1}^n f(y_{1i}, y_{2i}, r_i \mid \boldsymbol{\mu}, \boldsymbol{\Sigma}, \psi).$$

- ↪ Under ignorability, i.e., assuming that missing in  $Y_2$  is MAR (or MCAR) and that  $(\boldsymbol{\mu}, \boldsymbol{\Sigma})$  and  $\psi$  are distinct, we can ‘simply’ work with the likelihood of the observed data in order to estimate  $\boldsymbol{\mu}$  and  $\boldsymbol{\Sigma}$ .
- ↪ Remember that distinctness of the parameters formally implies that the parameter space of  $((\boldsymbol{\mu}, \boldsymbol{\Sigma}), \psi)$  is equal to the Cartesian product of their individual product spaces. Informally stated, this means that the model for the missing data mechanism does not contain information about the parameters of the complete data model.

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↪ The likelihood of the observed data, which is

$$\begin{aligned} L(\boldsymbol{\mu}, \boldsymbol{\Sigma} \mid \mathbf{y}_{\text{obs}}) &= \prod_{i=1}^m f(y_{1i}, y_{2i} \mid \boldsymbol{\mu}, \boldsymbol{\Sigma}) \prod_{i=m+1}^n f(y_{1i} \mid \mu_1, \sigma_1^2) \\ &= \prod_{i=1}^m \left\{ \frac{1}{2\pi} (\det \boldsymbol{\Sigma})^{-1/2} \exp \left[ -\frac{1}{2} (\mathbf{y}_i - \boldsymbol{\mu})^T \boldsymbol{\Sigma}^{-1} (\mathbf{y}_i - \boldsymbol{\mu}) \right] \right\} \\ &\quad \times \prod_{i=m+1}^n \left\{ \left( 2\pi\sigma_1^2 \right)^{-1/2} \exp \left[ -\frac{1}{2\sigma_1^2} (y_{1i} - \mu_1)^2 \right] \right\}. \end{aligned} \quad (1)$$

↪ Note that  $\mathbf{y}_{\text{obs}} = (y_{11}, \dots, y_{1n}, y_{21}, \dots, y_{2m})$  and  $\mathbf{y}_i = (y_{1i}, y_{2i})'$ .

# Likelihood based inference with incomplete data

Example: bivariate normal data with one variable subject to missingness

- ↪ Maximum likelihood estimates of  $\mu$  and  $\Sigma$  can be obtained by maximising the log likelihood function corresponding to the likelihood in (1) with respect to  $\mu$  and  $\Sigma$ .
- ↪ The likelihood equations, however, do not have an obvious solution.
- ↪ A more convenient parametrisation, for finding the ML estimates analytically in this example, uses a factored likelihood (Little and Rubin, 2002, Chapter 7), in which the several 'factors' or terms appearing in the likelihood have distinct parameters and can be maximised separately.

# Likelihood based inference with incomplete data

Example: bivariate normal data with one variable subject to missingness

## Known facts for a bivariate normal distribution

↪ If  $\mathbf{Y} = (Y_1, Y_2)' \sim N_2(\boldsymbol{\mu}, \boldsymbol{\Sigma})$ , then

$$Y_1 \sim N(\mu_1, \sigma_1^2),$$

$$Y_2 \mid Y_1 = y_1 \sim N(\beta_0 + \beta_1 y_1, \sigma_{2|1}^2),$$

$$\beta_0 = \mu_2 - \beta_1 \mu_1,$$

$$\beta_1 = \frac{\sigma_{12}}{\sigma_1^2},$$

$$\sigma_{2|1}^2 = \sigma_2^2 - \frac{\sigma_{12}^2}{\sigma_1^2}.$$

↪ Proofs of these results can be found in almost every book on multivariate analysis (or somewhere in the web!). I personally like the following book: Rencher, A.C. (2002) *Methods of Multivariate Analysis*, Wiley (see Chapter 4, pp. 82–111, about the multivariate normal distribution).

# Likelihood based inference with incomplete data

## Example: bivariate normal data with one variable subject to missingness

- ↪ Remember that we can write the joint density of  $Y_1$  and  $Y_2$  as the product of the conditional density of  $Y_2$  given  $Y_1 = y_1$  and the marginal density of  $Y_1$  (and from the results stated in the previous slide, we know what the parameters of these two distributions are)

$$f(y_1, y_2 \mid \phi) = f(y_2 \mid y_1, \beta_0, \beta_1, \sigma_{2|1})f(y_1 \mid \mu_1, \sigma_1).$$

- ↪ The crux here is that the parameter  $\phi \equiv \phi(\theta) = (\mu_1, \sigma_1, \beta_0, \beta_1, \sigma_{2|1})'$  is a one-to-one function of the original parameter  $\theta = (\mu_1, \sigma_1, \mu_2, \sigma_2, \sigma_{12})'$  and the parameters of the conditional and marginal densities are distinct since knowledge on  $(\mu_1, \sigma_1)$  does not imply any information about  $(\beta_0, \beta_1, \sigma_{2|1})$ .
- ↪ Saying that  $\phi$  is a one-to-one function of  $\theta$  means that all elements of  $\phi$  can be written as a function of elements of  $\theta$  and the other way around and that they are uniquely determined.
- ↪ In the previous slide we have already seen how to write the elements of  $\phi$  as a function of the elements of  $\theta$ .
- ↪ Similarly, the components of  $\theta$  other than  $\mu_1$  and  $\sigma_1$  can be expressed as the following functions of the components of  $\phi$

$$\mu_2 = \beta_0 + \beta_1 \mu_1$$

$$\sigma_{12} = \beta_1 \sigma_1^2$$

$$\sigma_2^2 = \sigma_{2|1}^2 + \beta_1^2 \sigma_1^2$$



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↪ Finally, the likelihood for the observed data simplifies to

$$\begin{aligned} L(\boldsymbol{\theta} \mid \mathbf{y}_{\text{obs}}) &= \prod_{i=1}^m f(y_{1i}, y_{2i} \mid \boldsymbol{\theta}) \prod_{i=m+1}^n f(y_{1i} \mid \boldsymbol{\theta}) \\ &= \left\{ \prod_{i=1}^m f(y_{1i} \mid \mu_1, \sigma_1) f(y_{2i} \mid y_{1i}, \beta_0, \beta_1, \sigma_{2|1}) \right\} \prod_{i=m+1}^n f(y_{1i} \mid \mu_1, \sigma_1) \\ &= \prod_{i=1}^m f(y_{2i} \mid y_{1i}, \beta_0, \beta_1, \sigma_{2|1}) \prod_{i=1}^n f(y_{1i} \mid \mu_1, \sigma_1) \\ &= \prod_{i=1}^m \left\{ (2\pi\sigma_{2|1}^2)^{-1/2} \exp \left[ -\frac{1}{2\sigma_{2|1}^2} (y_{2i} - (\beta_0 + \beta_1 y_{1i}))^2 \right] \right\} \\ &\quad \times \prod_{i=1}^n \left\{ (2\pi\sigma_1^2)^{-1/2} \exp \left[ -\frac{1}{2\sigma_1^2} (y_{1i} - \mu_1)^2 \right] \right\} \end{aligned} \quad (2)$$

# Likelihood based inference with incomplete data

Example: bivariate normal data with one variable subject to missingness

- ↪ The likelihood in (2) is much more tractable, from an analytical point of view, than the one in (1).
- ↪ The first term is the density for  $m$  observations from a conditional normal distribution with mean  $\beta_0 + \beta_1 y_1$  and variance  $\sigma_{2|1}^2$ .
- ↪ The second factor is the density of an independent sample of size  $n$  from a normal distribution with mean  $\mu_1$  and variance  $\sigma_1^2$ .
- ↪ Because we know that the parameters of the two terms are distinct, ML estimates of  $\phi$  can be obtained by independently maximising the likelihoods corresponding to these two components.

# Likelihood based inference with incomplete data

Example: bivariate normal data with one variable subject to missingness

→ After some calculations, we obtain

$$\hat{\mu}_1 = \frac{1}{n} \sum_{i=1}^n y_{1i}, \quad \hat{\sigma}_1^2 = \frac{1}{n} \sum_{i=1}^n (y_{1i} - \hat{\mu}_1)^2$$

$$\hat{\beta}_0 = \bar{y}_2 - \hat{\beta}_1 \bar{y}_1, \quad \bar{y}_j = \frac{1}{m} \sum_{i=1}^m y_{ji}, \quad j = 1, 2,$$

$$\hat{\beta}_1 = \frac{s_{12}}{s_1^2}, \quad s_{12} = \frac{1}{m} \sum_{i=1}^m (y_{1i} - \bar{y}_1)(y_{2i} - \bar{y}_2), \quad s_1^2 = \frac{1}{m} \sum_{i=1}^m (y_{1i} - \bar{y}_1)^2,$$

$$\hat{\sigma}_{2|1}^2 = s_2^2 \frac{s_{12}^2}{s_1^2}, \quad s_2^2 = \frac{1}{m} \sum_{i=1}^m (y_{2i} - \bar{y}_2)^2.$$

→ Note that under a complete case analysis, the likelihood would be

$$L(\theta \mid y_{11}, \dots, y_{1m}, y_{21}, \dots, y_{2m}) = \prod_{i=1}^m f(y_{1i}, y_{2i} \mid \mu, \Sigma).$$

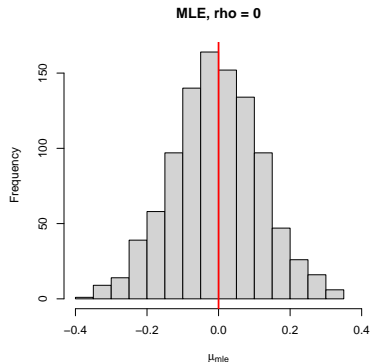
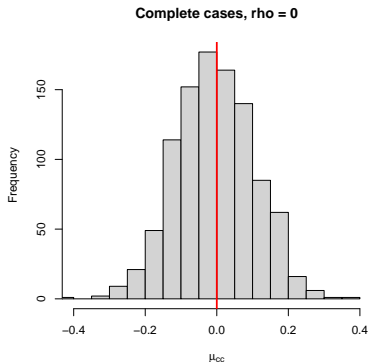
# Likelihood based inference with incomplete data

Example: bivariate normal data with one variable subject to missingness

- ↪ We have conducted a simulation study to ascertain the performance of both the mle and complete cases approach in estimating  $\mu_2$ . Note that  $\hat{\mu}_2 = \hat{\beta}_0 + \hat{\beta}_1 \hat{\mu}_1$ .
- ↪ We have that  $\hat{\mu}_2^{\text{CCA}} = \frac{1}{m} \sum_{i=1}^m y_{2i}$ .
- ↪ We have considered to simulate data that  $\mu_1 = \mu_2 = 0$ ,  $\sigma_1^2 = \sigma_2^2 = 1$ , and different correlations, namely  $\rho \in \{0, 0.5, 0.9\}$  (thus implying  $\sigma_{12} \in \{0, 0.5, 0.9\}$ ).
- ↪ We simulated 1000 datasets, each of size  $n = 100$ .
- ↪ Induced missingness mechanism is MAR. Specifically, we have considered that  $r_i = 0$  (i.e.,  $y_{2i}$  is missing) if  $y_{1i} > z_p$ , where  $z_p = \Phi^{-1}(a)$ , for  $a = 0.8$  (distinctness of parameters assumption is also satisfied).

# Likelihood based inference with incomplete data

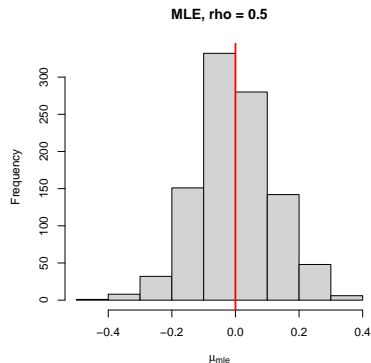
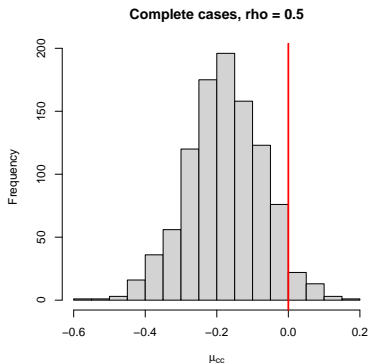
Example: bivariate normal data with one variable subject to missingness



→ Note that when the correlation between  $Y_1$  and  $Y_2$  is zero, i.e.,  $Y_1$  and  $Y_2$  are independent, the mle estimator of  $\mu_2$  reduces to the complete cases estimator.

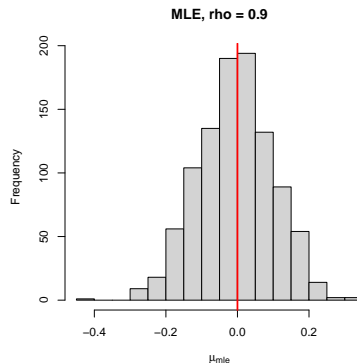
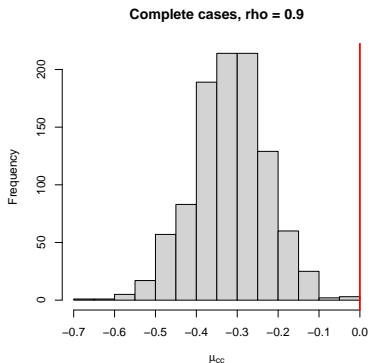
# Likelihood based inference with incomplete data

Example: bivariate normal data with one variable subject to missingness



# Likelihood based inference with incomplete data

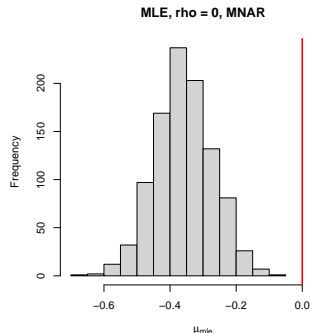
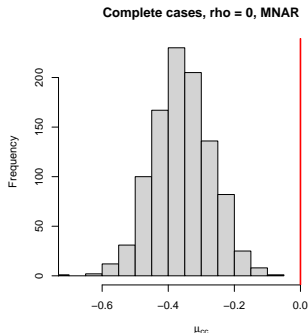
Example: bivariate normal data with one variable subject to missingness



# Likelihood based inference with incomplete data

## Example: bivariate normal data with one variable subject to missingness

- We have repeated the same simulation exercise but we now break the assumption of MAR data and simulate MNAR data.
- Specifically, we have considered that  $r_i = 0$  (i.e.,  $y_{2i}$  is missing) if  $y_{2i} > z_p$ , where  $z_p = \Phi^{-1}(a)$ , for  $a = 0.8$ .

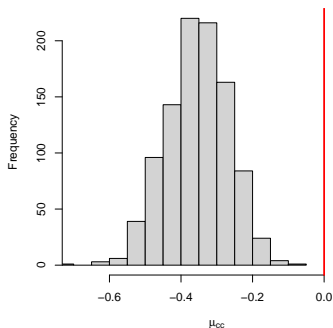




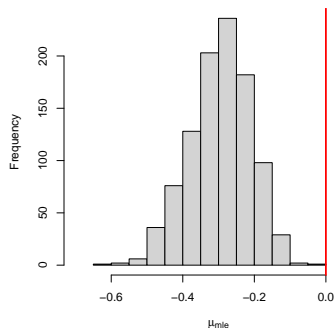
# Likelihood based inference with incomplete data

Example: bivariate normal data with one variable subject to missingness

Complete cases,  $\rho = 0.5$ , MNAR



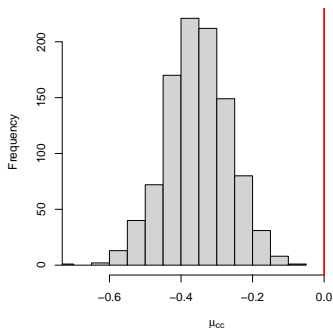
MLE,  $\rho = 0.5$ , MNAR



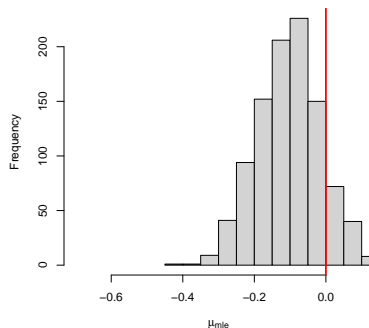
# Likelihood based inference with incomplete data

Example: bivariate normal data with one variable subject to missingness

Complete cases,  $\rho = 0.9$ , MNAR



MLE,  $\rho = 0.9$ , MNAR



# Likelihood based inference with incomplete data

## Final remarks

- ↪ Under the ignorability assumption, maximum likelihood estimates can be validly obtained through the observed data likelihood.
- ↪ In this specific example covered here we were able to factorise the observed data likelihood and thus easily obtained the ML estimates.
- ↪ However, general missing data patterns often do not have the particular forms that allow explicit maximum likelihood estimates to be calculated by exploiting factorisations of the likelihood.
- ↪ Furthermore, for some models a factorisation exists, but the parameters in the factorisation are not distinct, and thus maximising the factors separately does not maximise the likelihood.
- ↪ We will now learn about the EM algorithm, an iterative procedure especially designed to deal with incomplete data scenarios.
- ↪ It goes without saying that our ML estimates are only as good as the model we posit for the sampling distribution.