

# Advanced Methods in Biostatistics I

## Lecture 11

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# Multivariate normality

- In order to perform inference on the linear model, we typically assume the response variable follows a multivariate normal distribution.
- Hence, it is important to understand the properties of this distribution.
- However, before discussing the multivariate normal distribution, let's briefly review the univariate normal distribution.

# The univariate normal distribution

- A random variable  $z$  follows a standard normal distribution if its density is

$$f_z(z) = \frac{1}{\sqrt{2\pi}} \exp(-z^2/2) \quad -\infty < z < \infty$$

- We say  $z \sim N(0, 1)$ ,
- A standard normal random variable has mean 0 and variance 1.

# The univariate normal distribution

- To obtain a normal random variable with arbitrary mean  $\mu$  and variance  $\sigma^2$  we use the transformation:

$$y = \mu + \sigma Z.$$

- Here  $E[y] = \mu$  and  $Var(y) = \sigma^2$ .

# The univariate normal distribution

- Conversely,

$$z = (y - \mu)/\sigma$$

is standard normal if  $y$  is a normal random variable with mean  $\mu$  and standard deviation  $\sigma$ .

# The univariate normal distribution

- The density for a normal random variable with mean  $\mu$  and standard deviation  $\sigma$  is given by

$$f_y(y) = f_z\left(\frac{y - \mu}{\sigma}\right) / \sigma.$$

- We write that  $y \sim N(\mu, \sigma^2)$ .

# Moment generating function

## Theorem

If  $y \sim N(\mu, \sigma^2)$  the moment generating function (m.g.f.) of  $y$  is given by

$$M_y(t) \equiv E[e^{ty}] = \exp\{\mu t + t^2 \sigma^2 / 2\}.$$

# The multivariate normal distribution

- Suppose  $z_1, z_2, \dots, z_p$  are independent identically distributed (i.i.d.) standard normal random variables.
- The joint density of  $\mathbf{z} = (z_1, z_2, \dots, z_p)'$  is then given by

$$\begin{aligned} f_{\mathbf{z}}(\mathbf{z}) &= \prod_{i=1}^p \frac{1}{\sqrt{2\pi}} \exp(-z_i^2/2) \\ &= (2\pi)^{-p/2} \exp(-\mathbf{z}'\mathbf{z}/2) \end{aligned}$$



# The multivariate normal distribution

- This is the multivariate standard normal distribution for a random vector  $\mathbf{z}$  with mean  $\mathbf{0}$  and variance  $\mathbf{I}$ .
- We write this as  $\mathbf{z} \sim N_p(\mathbf{0}, \mathbf{I})$ .
- Here  $p$  corresponds to the number of elements in  $y$ .

# The multivariate normal distribution

- To obtain a multivariate normal random variable with arbitrary mean  $\mu$  and variance-covariance matrix  $\Sigma$  we use the transformation:

$$\mathbf{y} = \mu + \Sigma^{1/2}\mathbf{z}.$$

where  $\Sigma^{1/2}$  is the symmetric square root of  $\Sigma$ .

- Here  $E[\mathbf{y}] = \mu$  and  $Var(\mathbf{y}) = \Sigma^{1/2}\Sigma^{1/2} = \Sigma$ , which is assumed to be positive definite.

# The multivariate normal distribution

- Conversely,

$$\mathbf{z} = \Sigma^{-1/2}(\mathbf{y} - \mu).$$

is a multivariate standard normal random variable if  $\mathbf{y}$  is a multivariate normal random variable with mean  $\mu$  and variance-covariance matrix  $\Sigma$ .

# The multivariate normal distribution

- The density of the non-standard multivariate normal distribution is given by

$$(2\pi)^{-n/2} |\Sigma|^{-1/2} \exp \left\{ -\frac{1}{2} (\mathbf{y} - \mu)' \Sigma^{-1} (\mathbf{y} - \mu) \right\}.$$

- In this setting, we say that  $\mathbf{y} \sim N_p(\mu, \Sigma)$ .

# The multivariate normal distribution

- Note that all full row rank linear transformations of the normal are also normal.
- That is,  $\mathbf{a} + \mathbf{A}\mathbf{y}$  is normal if  $\mathbf{A}$  has full row rank.
- We will also show that all conditional and submarginal distributions of the multivariate normal are also normal.

# Moment generating function

## Theorem

If  $\mathbf{y} \sim N_p(\boldsymbol{\mu}, \Sigma)$  the moment generating function (m.g.f.) of  $\mathbf{y}$  is

$$M_{\mathbf{y}}(\mathbf{t}) \equiv E[e^{\mathbf{t}'\mathbf{y}}] = \exp\{\boldsymbol{\mu}'\mathbf{t} + \frac{1}{2}\mathbf{t}'\Sigma\mathbf{t}\}.$$

# Moment generating function

Two important properties of moment generating functions:

- If two random vectors have the same moment generating function, they have the same density.
- Two random vectors are independent if and only if their joint moment generating function factors into the product of their two separate moment generating functions;

## Properties

Let  $\mathbf{y} \sim N_p(\boldsymbol{\mu}, \Sigma)$ , and let  $\mathbf{a}$  be a  $p \times 1$  vector,  $\mathbf{b}$  a  $k \times 1$  vector, and  $\mathbf{C}$  a  $k \times p$  matrix with rank  $= k \leq p$ , then

- $x = \mathbf{a}'\mathbf{y} \sim N(\mathbf{a}'\boldsymbol{\mu}, \mathbf{a}'\Sigma\mathbf{a})$ .
- $\mathbf{x} = \mathbf{C}\mathbf{y} + \mathbf{b} \sim N_p(\mathbf{C}\boldsymbol{\mu} + \mathbf{b}, \mathbf{C}\Sigma\mathbf{C}')$ .



# Example

- Let  $\mathbf{Z} = (Z_1, Z_2)' \sim N_2(\mathbf{0}, \mathbf{I})$ , and let  $\mathbf{A}$  be the linear transformation matrix

$$\mathbf{A} = \begin{pmatrix} 1/2 & -1/2 \\ -1/2 & 1/2 \end{pmatrix}.$$

- Let  $\mathbf{Y} = (Y_1, Y_2)'$  be the linear transformation

$$\mathbf{Y} = \mathbf{AZ} = \begin{pmatrix} (Z_1 - Z_2)/2 \\ (Z_2 - Z_1)/2 \end{pmatrix}.$$

- Now  $\mathbf{Y} \sim N(\mathbf{0}, \Sigma)$  where  $\Sigma = \mathbf{AA}'$ .

## Theorem

If  $\mathbf{y} \sim N_p(\boldsymbol{\mu}, \Sigma)$ , then any  $r \times 1$  subvector of  $\mathbf{y}$  has a  $r$ -variate normal distribution.

- It follows directly from this result that if  $\mathbf{y} \sim N_p(\boldsymbol{\mu}, \Sigma)$  then  $y_i \sim N(\mu_i, \sigma_i^2)$  for  $i = 1, \dots, p$ .
- Thus, joint normality implies marginal normality.
- The converse is not necessarily true.

- Let  $\mathbf{y} \sim N_n(\boldsymbol{\mu}, \Sigma)$  be partitioned as follows:

$$\mathbf{y} = \begin{pmatrix} \mathbf{y}_1 \\ \mathbf{y}_2 \end{pmatrix},$$

where  $\mathbf{y}_1$  is  $p \times 1$  and  $\mathbf{y}_2$  is  $q \times 1$  with  $p + q = n$ .

# Partitioning

- Then, the mean and covariance matrix are correspondingly partitioned as

$$\boldsymbol{\mu} = \begin{pmatrix} \boldsymbol{\mu}_1 \\ \boldsymbol{\mu}_2 \end{pmatrix}$$

and

$$\begin{aligned} \boldsymbol{\Sigma} &= \begin{pmatrix} \boldsymbol{\Sigma}_{11} & \boldsymbol{\Sigma}_{12} \\ \boldsymbol{\Sigma}_{21} & \boldsymbol{\Sigma}_{22} \end{pmatrix} \\ &= \begin{pmatrix} \text{var}(\mathbf{y}_1) & \text{cov}(\mathbf{y}_1, \mathbf{y}_2) \\ \text{cov}(\mathbf{y}_2, \mathbf{y}_1) & \text{var}(\mathbf{y}_2) \end{pmatrix}. \end{aligned}$$

# Marginal distribution

- The marginal distributions are  $\mathbf{y}_1 \sim N_p(\boldsymbol{\mu}_1, \Sigma_{11})$  and  $\mathbf{y}_2 \sim N_q(\boldsymbol{\mu}_2, \Sigma_{22})$ .

## Theorem

If

$$\mathbf{y} = \begin{pmatrix} \mathbf{y}_1 \\ \mathbf{y}_2 \end{pmatrix}$$

is  $N_{p+q}(\boldsymbol{\mu}, \Sigma)$ , then  $\mathbf{y}_1$  and  $\mathbf{y}_2$  are independent if  $\Sigma_{12} = \mathbf{0}$ .

# Independence

- However, if  $\mathbf{y}_1 \sim N_p(\boldsymbol{\mu}_1, \Sigma_{11})$  and  $\mathbf{y}_2 \sim N_q(\boldsymbol{\mu}_2, \Sigma_{22})$ , and  $\Sigma_{12} = \Sigma'_{21} = \mathbf{0}$ , this does not necessarily mean that  $\mathbf{y}_1$  and  $\mathbf{y}_2$  are independent.
- We also need  $\mathbf{y}$  to be jointly normal.



## Corollary

If  $\mathbf{y} \sim N_p(\boldsymbol{\mu}, \Sigma)$ , then any two individual variables  $y_i$  and  $y_j$  are independent if  $\sigma_{ij} = 0$ .

# Orthogonal transformations

## Theorem

Let  $\mathbf{y} \sim N_p(\boldsymbol{\mu}, \sigma^2 \mathbf{I})$ , and let  $\mathbf{T}$  be an orthogonal constant matrix. Then  $\mathbf{T}\mathbf{y} \sim N_p(\mathbf{T}\boldsymbol{\mu}, \sigma^2 \mathbf{I})$ .

# Orthogonal transformations

- The theorem states that mutually independent normal random variables with common variance remain mutually independent with common variance under orthogonal transformations.
- Orthogonal matrices correspond to rotations and reflections about the origin, i.e., they preserve the vector length:

$$\|\mathbf{T}\mathbf{y}\|^2 = (\mathbf{T}\mathbf{y})'(\mathbf{T}\mathbf{y}) = \mathbf{y}'\mathbf{T}'\mathbf{T}\mathbf{y} = \mathbf{y}'\mathbf{y} = \|\mathbf{y}\|^2.$$

# Conditional distributions

- Suppose that  $\mathbf{y}_1$  and  $\mathbf{y}_2$  are jointly multivariate normal with  $\Sigma_{12} \neq 0$ , i.e.

$$\begin{pmatrix} \mathbf{y}_1 \\ \mathbf{y}_2 \end{pmatrix} \sim N \left( \begin{pmatrix} \boldsymbol{\mu}_1 \\ \boldsymbol{\mu}_2 \end{pmatrix}, \begin{pmatrix} \Sigma_{11} & \Sigma_{12} \\ \Sigma_{21} & \Sigma_{22} \end{pmatrix} \right).$$

- Further assume that  $\Sigma_{11}$  is nonsingular.

# Conditional distributions

- The conditional distribution of  $\mathbf{y}_2 \mid \mathbf{y}_1$  is  $N(\mu_{\mathbf{y}_2|\mathbf{y}_1}, \Sigma_{\mathbf{y}_2|\mathbf{y}_1})$ , where

$$\mu_{\mathbf{y}_2|\mathbf{y}_1} = \mu_2 + \Sigma_{21}\Sigma_{11}^{-1}(\mathbf{y}_1 - \mu_1)$$

$$\Sigma_{\mathbf{y}_2|\mathbf{y}_1} = \Sigma_{22} - \Sigma_{21}\Sigma_{11}^{-1}\Sigma'_{12}.$$

# Example

- Suppose that  $y$  and  $\mathbf{x}$  are jointly multivariate normal with

$$\begin{pmatrix} y \\ \mathbf{X} \end{pmatrix} \sim N \left( \begin{pmatrix} \mu_y \\ \boldsymbol{\mu}_x \end{pmatrix}, \begin{pmatrix} \sigma_y^2 & \Sigma'_{xy} \\ \Sigma_{xy} & \Sigma_x \end{pmatrix} \right).$$

where  $y$  is a scalar and  $\mathbf{X}$  is a  $p \times 1$  vector.

- Consider now predicting  $y$  given  $\mathbf{X} = \mathbf{x}$ .
- A good estimate for this would be  $E[y \mid \mathbf{X} = \mathbf{x}]$ .

# Example

- Our results suggest that  $\mathbf{y} \mid \mathbf{X} = \mathbf{x}$  is normal with mean:

$$\begin{aligned}\mu_{\mathbf{y}|\mathbf{x}} &= \mu_y + \Sigma'_{xy}\Sigma_x^{-1}(\mathbf{x} - \mu_x) \\ &= \mu_y - \boldsymbol{\mu}'_x\Sigma_x^{-1}\Sigma_{xy} + \mathbf{x}'\Sigma_x^{-1}\Sigma_{xy} \\ &= \beta_0 + \mathbf{x}'\boldsymbol{\beta}\end{aligned}$$

where

$$\beta_0 = \mu_y - \boldsymbol{\mu}'_x\Sigma_x^{-1}\Sigma_{xy}$$

and

$$\boldsymbol{\beta} = \Sigma_x^{-1}\Sigma_{xy}.$$

# Example

- Consider the case of simple linear regression.
- Here,

$$\beta_1 = \frac{\text{Cov}(x, y)}{\text{Var}(x)} = \rho(x, y) \frac{\sqrt{\text{Var}(y)}}{\sqrt{\text{Var}(x)}}.$$

and

$$\beta_0 = \bar{y} - \bar{x}\beta_1.$$



# Example

- Hence, the conditional mean in this case mirrors the linear model.
- The slope is the inverse of the variance-covariance matrix of the predictors times the cross correlations between the predictors and the response.

# Example

- This example provides motivation for the linear model in cases when the joint normality of the predictor and response is conceptually reasonable.
- Though we note that such joint normality is not always reasonable, such as when the predictors are binary, even though the linear model remains well justified.