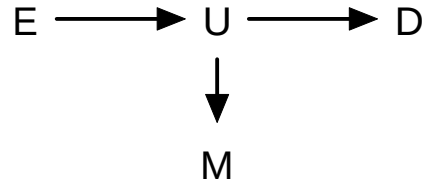


eAppendix

Detailed mathematical derivation:

Let E, U, M and D be four random variables with



Suppose the linear models among them are:

$$\begin{aligned}
 U &= \alpha_U + \beta_U E + \varepsilon_U \\
 M &= \alpha_M + \beta_M U + \varepsilon_M \\
 D &= \alpha_D + \beta_D U + \varepsilon_D
 \end{aligned}$$

Assume we have $E \sim N(\mu_E, \sigma_E^2)$, $\varepsilon_U \sim N(0, \sigma_U^2)$, $\varepsilon_M \sim N(0, \sigma_M^2)$ and $\varepsilon_D \sim N(0, \sigma_D^2)$, where

μ_E , σ_E^2 , σ_U^2 , σ_M^2 , and σ_D^2 are known.

$$\begin{aligned}
 \text{Var}(U) &= \beta_U^2 \sigma_E^2 + \sigma_U^2, \\
 \text{Var}(M) &= \beta_M^2 \sigma_U^2 + \sigma_M^2 = \beta_M^2 \beta_U^2 \sigma_E^2 + \beta_M^2 \sigma_U^2 + \sigma_M^2, \\
 \text{Var}(D) &= \beta_D^2 \sigma_U^2 + \sigma_D^2 = \beta_D^2 \beta_U^2 \sigma_E^2 + \beta_D^2 \sigma_U^2 + \sigma_D^2.
 \end{aligned}$$

Correlation between E and M

Therefore

$$\begin{aligned}
 M &= \alpha_M + \beta_M(\alpha_U + \beta_U E + \varepsilon_U) + \varepsilon_M \\
 &= \alpha_M + \beta_M \alpha_U + \beta_M \beta_U E + \beta_M \varepsilon_U + \varepsilon_M \\
 &= \alpha'_M + \beta'_M E + \varepsilon'_M,
 \end{aligned} \tag{1}$$

where

$$\begin{aligned}
 \alpha'_M &= \alpha_M + \beta_M \alpha_U, \\
 \beta'_M &= \beta_M \beta_U, \\
 \varepsilon'_M &= \beta_M \varepsilon_U + \varepsilon_M.
 \end{aligned}$$

Similarly, since

$$\begin{aligned}
U &= -\frac{\alpha_D}{\beta_D} + \frac{1}{\beta_D} D - \frac{\varepsilon_D}{\beta_D}, \\
M &= \alpha_M + \beta_M \left(-\frac{\alpha_D}{\beta_D} + \frac{1}{\beta_D} D - \frac{\varepsilon_D}{\beta_D} \right) + \varepsilon_M \\
&= \alpha_M + \beta_M \left(-\frac{\alpha_D}{\beta_D} \right) + \frac{\beta_M}{\beta_D} D - \frac{\beta_M}{\beta_D} \varepsilon_D + \varepsilon_M \\
&= \alpha''_M + \beta''_M D + \varepsilon''_M,
\end{aligned} \tag{2}$$

where

$$\begin{aligned}
\alpha''_M &= \alpha_M - \frac{\alpha_D \beta_M}{\beta_D}, \\
\beta''_M &= \frac{\beta_M}{\beta_D}, \\
\varepsilon''_M &= \varepsilon_M - \frac{\beta_M}{\beta_D} \varepsilon_D.
\end{aligned}$$

Therefore, we have

$$\begin{aligned}
\text{cov}(M, E) &= \beta_M \beta_U \sigma_E \sigma_M, \\
\text{cov}(M, U) &= \beta_M \sigma_U \sigma_M, \\
\text{cov}(M, D) &= \frac{\beta_M}{\beta_D} \sigma_D \sigma_M.
\end{aligned}$$

Correlation between E and D

Consider the correlation between E and D.

$$\begin{aligned}
D &= \alpha_D + \beta_D(\alpha_U + \beta_U E + \varepsilon_U) + \varepsilon_D \\
&= \alpha_D + \beta_D \alpha_U + \beta_D \beta_U E + \beta_D \varepsilon_U + \varepsilon_D \\
&= \alpha'_D + \beta'_D E + \varepsilon'_D,
\end{aligned} \tag{3}$$

where

$$\begin{aligned}
\alpha'_D &= \alpha_D + \beta_D \alpha_U, \\
\beta'_D &= \beta_D \beta_U, \\
\varepsilon'_D &= \beta_D \varepsilon_U + \varepsilon_D.
\end{aligned}$$

$$\begin{aligned}
D &= \alpha_D + \beta_D \left(-\frac{\alpha_M}{\beta_M} + \frac{1}{\beta_M} M - \frac{\varepsilon_M}{\beta_M} \right) + \varepsilon_D \\
&= \alpha_D + \beta_D \left(-\frac{\alpha_M}{\beta_M} \right) + \frac{\beta_D}{\beta_M} M - \frac{\beta_D}{\beta_M} \varepsilon_M + \varepsilon_D \\
&= \alpha''_D + \beta''_D M + \varepsilon''_D,
\end{aligned} \tag{4}$$

where

$$\alpha''_D = \alpha_D - \frac{\alpha_M \beta_D}{\beta_M},$$

$$\beta''_D = \frac{\beta_D}{\beta_M},$$

$$\varepsilon''_D = \varepsilon_D - \frac{\beta_D}{\beta_M} \varepsilon_M.$$

Therefore, we have

$$\text{cov}(D, E) = \beta_M \beta_U \sigma_E \sigma_D,$$

$$\text{cov}(D, U) = \beta_M \sigma_U \sigma_D,$$

$$\text{cov}(D, M) = \frac{\beta_M}{\beta_D} \sigma_M \sigma_D.$$

Conditional covariance of E and D given M

$$\begin{aligned} \text{cor}(E, D) &= \beta'_D \\ &= \beta_D \beta_U \end{aligned}$$

$$\text{cor}(E, D | M) = \frac{\text{cov}(E, D | M)}{\sqrt{\text{var}(E | M)} \sqrt{\text{var}(D | M)}}$$

$$\text{cov}(E, D | M) = E[ED | M] - E[E | M]E[D | M]$$

According to section 1 and 2

$$M = \alpha'_M + \beta'_M E + \varepsilon'_M$$

$$E = -\frac{\alpha'_M}{\beta'_M} + \frac{M}{\beta'_M} + \frac{\varepsilon'_M}{\beta'_M}$$

$$D = \alpha''_D + \beta''_D M + \varepsilon''_D,$$

$$E[E | M] = -\frac{\alpha_M}{\beta_M} + \frac{1}{\beta'_M} M$$

$$E[D | M] = \alpha''_D + \beta''_D M$$

$$E[ED | M] = E[(-\frac{\alpha'_M}{\beta'_M} + \frac{M}{\beta'_M} + \frac{\varepsilon'_M}{\beta'_M}) (\alpha''_D + \beta''_D M + \varepsilon''_D) | M]$$

Since $E[\varepsilon'_M | M] = 0$ and $E[\varepsilon''_D | M] = 0$,

$$E[E | M] * E[D | M] = -\frac{\alpha'_M \alpha''_D}{\beta_M} - \frac{\alpha'_M \beta''_D M}{\beta_M} + \frac{M \alpha''_D}{\beta'_M} + \frac{M^2 \beta''_D}{\beta'_M} | M, \text{ we have}$$

$$E[ED | M] - E[E | M] E[D | M] = E[-\frac{\varepsilon'_M}{\beta'_M} \varepsilon''_D | M]$$

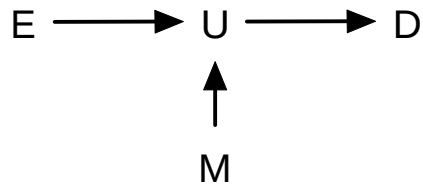
$$\begin{aligned}
&= E[(- \frac{\beta_M \varepsilon_U + \varepsilon_M}{\beta_M \beta_U})(\varepsilon_D - \frac{\beta_M}{\beta_D} \varepsilon_M) | M] \\
&= E[\frac{\varepsilon_M^2}{\beta_D \beta_U} | M] \\
&= \frac{\sigma_M^2}{\beta_D \beta_U} \\
\text{var}(E|M) &= \frac{1}{(\beta_M')^2} \text{var}(\varepsilon_M) \\
&= \frac{\beta_M^2 \sigma_U^2 + \sigma_M^2}{\beta_M^2 \beta_U^2} \\
\text{var}(D|M) &= \text{var}(\varepsilon_D) \\
&= \sigma_D^2 + \frac{\beta_D^2}{\beta_M^2} \sigma_M^2
\end{aligned}$$

Therefore

$$\text{bias} = \text{cor}(E, D | M) - \text{cor}(E, D) = \frac{\frac{\sigma_M^2}{\beta_U \beta_D}}{\sqrt{\frac{\beta_M^2 \sigma_U^2 + \sigma_M^2}{\beta_M^2 \beta_U^2}} \sqrt{\sigma_D^2 + \frac{\alpha_M^2 \beta_D^2}{\beta_M^2} \sigma_M^2}} - \beta_D \beta_U \quad (5)$$

The second model

Let E, U, M and D be four random variables with.



Suppose the linear models among them are:

$$U = \alpha_U + \beta_{UE}E + \beta_{UM}M + \varepsilon_U,$$

$$D = \alpha_D + \beta_D U + \varepsilon_D,$$

Assume we have $E \sim N(\mu_E, \sigma_E^2)$, $M \sim N(\mu_M, \sigma_M^2)$, $\varepsilon_U \sim N(0, \sigma_U^2)$, $\varepsilon_M \sim N(0, \sigma_M^2)$ and $\varepsilon_D \sim N$

$(0, \sigma_D^2)$, where μ_E , σ_E^2 , μ_M , σ_M^2 , σ_U^2 , and σ_D^2 are known.

$$\mu_U = \alpha_U + \beta_{UE}\mu_E + \beta_{UM}\mu_M,$$

$$\mu_D = \alpha_D + \beta_D \mu_U$$

$$\text{Var}(U) = \beta_{UE}^2 \sigma_E^2 + \beta_{UM}^2 \sigma_M^2 + \sigma_U^2,$$

$$\text{Var}(D) = \beta_D^2 \sigma_U^2 + \sigma_D^2.$$

Therefore

$$\begin{aligned} D &= \alpha_D + \beta_D(\alpha_U + \beta_{UE}E + \beta_{UM}M + \varepsilon_U) \\ &= \alpha'_D + \beta'_{DE}E + \beta'_{DM}M + \varepsilon'_D, \end{aligned}$$

where

$$\alpha'_D = \alpha_D + \beta_D\alpha_U,$$

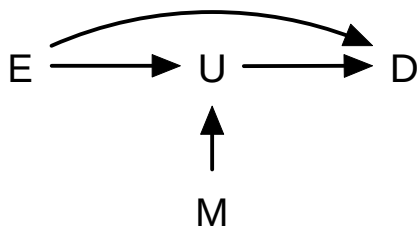
$$\beta'_{DE} = \beta_D\beta_{UE},$$

$$\beta'_{DM} = \beta_D\beta_{UM},$$

$$\varepsilon'_D = \beta_D\varepsilon_U + \varepsilon_D.$$

The third model

Let E, U, M and D be four random variables with



Suppose the linear models among them are:

$$U = \alpha_U + \beta_{UE}E + \beta_{UM}M + \varepsilon_U,$$

$$D = \alpha_D + \beta_{DE}E + \beta_{DU}U + \varepsilon_D.$$

Therefore

$$\begin{aligned} D &= \alpha_D + \beta_{DU}(\alpha_U + \beta_{UE}E + \beta_{UM}M + \varepsilon_U) + \beta_{DE}E + \varepsilon_D \\ &= \alpha_D + \beta_{DU}\alpha_U + (\beta_{DE} + \beta_{UE}\beta_{DU})E + \beta_{UM}\beta_{DU}M + \beta_{DU}\varepsilon_U + \varepsilon_D \\ &= \alpha'_D + \beta'_{DE}E + \beta'_{DM}M + \varepsilon'_D, \end{aligned}$$

where

$$\alpha'_D = \alpha_D + \beta_D\alpha_U,$$

$$\beta'_{DE} = \beta_{DE} + \beta_{UE}\beta_{DU},$$

$$\beta'_{DM} = \beta_{UM}\beta_{DU},$$

$$\varepsilon'_D = \beta_{DM}\varepsilon_U + \varepsilon_D.$$