

Least product relative error criterion based estimating  
equation approaches for the error-in-covariables  
multiplicative regression models

Qihua Wang

Academy of Mathematics and Systems Science, CAS

(Joint work with: Dahai Hu, University of Science and Technology of China)

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- 1 Introduction
- 2 Measurement Error Model
- 3 Conditional Mean Score Based Estimating Equation Approach
- 4 Corrected estimating equation method
- 5 Simulation Studies
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# Background

To handle the positive response, it is natural to consider the following multiplicative regression model,

$$Y_i = \exp(Z_i^T \beta_0) \varepsilon_i, \quad i = 1, \dots, n, \quad (1)$$

where  $Y_i$  is a scalar response variable,  $Z_i$  is a random covariate vector with the first component being 1 (intercept),  $\beta_0$  is the true regression parametric vector, and the error term  $\varepsilon$  is strictly positive.

# Least Absolute Relative Errors (LARE)

Chen et al. (2010, JASA) consider the following two types of relative errors:

- $|Y_i - \exp(Z_i^T \beta)| / Y_i$ ;
- $|Y_i - \exp(Z_i^T \beta)| / \exp(Z_i^T \beta)$ .

They proposed LARE criteria is to minimize

$$LARE_n(\beta) = \sum_{i=1}^n \left\{ \left| \frac{Y_i - \exp(Z_i^T \beta)}{Y_i} \right| + \left| \frac{Y_i - \exp(Z_i^T \beta)}{\exp(Z_i^T \beta)} \right| \right\}.$$

Denote the minimizer of  $LARE_n(\beta)$  as  $\hat{\beta}_{n,LARE}$ .

# Least Absolute Relative Errors (LARE)

- Advantage

- ▶ Scale free and Robust ;
- ▶ Relative error is concerned;
- ▶  $LARE_n(\beta)$  is strictly convex in  $\beta$  under some regular conditions;

- Disadvantage

- ▶ Nonsmooth;
- ▶ Computation is complicated;
- ▶ the limiting variance of  $\hat{\beta}_{n,LARE}$  involves the density of the error

## Least Product Relative Errors (LPRE)

To overcome the disadvantage of the LARE criteria, the product of the above two type relative errors are considered, namely,

$$\left| \frac{Y_i - \exp(Z_i^T \beta)}{Y_i} \right| \times \left| \frac{Y_i - \exp(Z_i^T \beta)}{\exp(Z_i^T \beta)} \right|.$$

- Wang et al. (2015, Test) developed a testing procedure to detect existence of the unknown change point and discussed a relative-based estimation of the change point.
- Chen et al.(2016, JMVA) study the least product relative error (LPRE) estimator.

# Least Product Relative Errors (LPRE)

## Advantage

- Smooth
- Convex



# Our Focus

Existing Question:

- The aforementioned LPRE methods commonly assume that covariates are observed precisely.
- We usually encounter corrupted data in practice, where the covariates are measured with error.

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# Measurement Error Model

- $Z_i = (V_i^T, X_i^T)^T$ 
  - ▶  $V_i$  :  $q \times 1$  vector of explanatory variables that are precisely measured with the first component being 1 (intercept),
  - ▶  $X_i$  :  $p \times 1$  vector of error-prone explanatory variable
- Classical additive measurement error model:

$$W_{i,j} = X_i + U_{i,j}, \quad j = 1, \dots, n_i, i = 1, \dots, n,$$

- ▶  $U$  and  $-U$  comes the same distribution
- ▶  $U$  is independent of  $(Z, \varepsilon)$

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## LPRE without Measurement Error

**When  $Z_i$ 's are precisely measured**, minimizing  $LPRE_n(\beta)$  is equivalent to **solving the following estimating equation**

$$U_n(\beta) = \sum_{i=1}^n \psi(Z_i, Y_i, \beta) = 0 \quad (2)$$

where  $\psi(Z_i, Y_i, \beta) = \{Y_i^{-1} \exp(Z_i^T \beta) - Y_i \exp(-Z_i^T \beta)\} Z_i$ .

With the assumption  $E[\varepsilon - 1/\varepsilon|Z] = 0$ , it's easy to see that  $E[\psi(Z_i, Y_i, \beta_0)] = 0$ .

## Conditional Mean Score Based Estimating Equation Approach

For simplification, denote the observed data  $\mathcal{O}_{i,r} = (Y_i, V_i, W_{i,r})$  and let  $\mathcal{U}_i = (Y_i, V_i, X_i)$  for  $i = 1, \dots, n$  and  $r = 1, \dots, n_i$ .

The key point is to find a function  $T^*(\mathcal{O}_{i,r}, \beta)$  such that

$$E[T^*(\mathcal{O}_{i,r}, \beta) | \mathcal{U}_i] = \psi(Z_i, Y_i, \beta).$$

If so, this leads to the following unbiased estimating equation,

$$\sum_{i=1}^n \left[ \frac{1}{n_i} \sum_{r=1}^{n_i} T^*(\mathcal{O}_{i,r}, \beta) \right] = \mathbf{0}. \quad (3)$$

This general idea has also been used in Hu and Lin (2004, JASA) and Wu et al.(2015, JASA).

# Construct $T^*(\mathcal{O}_{i,r}, \beta)$

## Notation

- Take  $\hat{Z}_{i,r} = (V_i^T, W_{i,r}^T)^T$  and  $J = (0_{p \times q}, I_{p \times p})^T$ .  
Then  $\hat{Z}_{i,r} = Z_i + JU_{i,r}$ .
- Denote  $\varphi_0(\gamma) = E[\exp(U^T \gamma)]$  and  $\varphi_1(\gamma) = E[U \exp(U^T \gamma)]$ .
- Take

$$R_{i,r}^{(0)}(\beta) = \varphi_0^{-1}(\gamma) \exp(\hat{Z}_{i,r}^T \beta),$$

$$R_{i,r}^{(1)}(\beta) = \varphi_0^{-1}(\gamma) \exp(\hat{Z}_{i,r}^T \beta) \{ \hat{Z}_{i,r} - J \varphi_0^{-1}(\gamma) \varphi_1(\gamma) \}.$$

# Construct $T^*(\mathcal{O}_{i,r}, \beta)$

A simple algebraic manipulation yields

$$\exp(Z_i^T \beta) Z_i = E \left[ R_{i,r}^{(1)}(\beta) | \mathcal{U}_i \right], \quad (4)$$

$$\exp(Z_i^T \beta) = E \left[ R_{i,r}^{(0)}(\beta) | \mathcal{U}_i \right]. \quad (5)$$

Thus, the desired function  $T^*(\mathcal{O}_{i,r}, \beta)$  can be defined as

$$T^*(\mathcal{O}_{i,r}, \beta) = Y_i^{-1} R_{i,r}^{(1)}(\beta) - Y_i R_{i,r}^{(0)}(-\beta).$$

However,  $\varphi_0(\gamma)$  and  $\varphi_1(\gamma)$  in  $T^*(\mathcal{O}_{i,r}, \beta)$  are unknown.



## $\hat{\varphi}_0(\gamma)$ & $\hat{\varphi}_1(\gamma)$

Denote  $\xi_i = I(n_i > 1)$  and  $\tilde{n} = \sum_{i=1}^n \xi_i$ . Then,  $\varphi_k(\gamma)$ , ( $k = 0, 1$ ) can be estimated by

$$\hat{\varphi}_0(\gamma) = \left[ \frac{1}{\tilde{n}} \sum_{i=1}^n \frac{\xi_i}{n_i(n_i - 1)} \sum_{r \neq s} \exp(\gamma^T (W_{i,r} - W_{i,s})) \right]^{1/2},$$

$$\hat{\varphi}_1(\gamma) = \frac{1}{2\tilde{n}\hat{\varphi}_0(\gamma)} \sum_{i=1}^n \left\{ \frac{\xi_i}{n_i(n_i - 1)} \sum_{r \neq s} (W_{i,r} - W_{i,s}) \exp(\gamma^T (W_{i,r} - W_{i,s})) \right\}$$

## Construct $\hat{T}^*(\mathcal{O}_{i,r}, \beta)$

Substituting  $\varphi_0(\gamma)$  and  $\varphi_1(\gamma)$  in  $R_{i,r}^{(0)}(\beta)$  and  $R_{i,r}^{(1)}(\beta)$  with  $\hat{\varphi}_0(\gamma)$  and  $\hat{\varphi}_1(\gamma)$  yields  $\hat{R}_{i,r}^{(0)}(\beta)$  and  $\hat{R}_{i,r}^{(1)}(\beta)$ . Thereafter, the resulting estimating equation is given by

$$\sum_{i=1}^n \left[ \frac{1}{n_i} \sum_{r=1}^{n_i} \hat{T}^*(\mathcal{O}_{i,r}, \beta) \right] = \mathbf{0},$$

where  $\hat{T}^*(\mathcal{O}_{i,r}, \beta) = Y_i^{-1} \hat{R}_{i,r}^{(1)}(\beta) - Y_i \hat{R}_{i,r}^{(1)}(-\beta)$ . The solution of the above equation,  $\hat{\beta}_{CMS}$  say, can be defined as estimator of  $\beta$ .

## $\sqrt{n}$ -consistency

Define  $R_i^{(0)}(\beta) = n_i^{-1} \sum_{r=1}^{n_i} R_{i,r}^{(0)}(\beta)$ ,  $\hat{R}_i^{(0)}(\beta) = n_i^{-1} \sum_{r=1}^{n_i} \hat{R}_{i,r}^{(0)}(\beta)$ ,  
 $R_i^{(1)}(\beta) = n_i^{-1} \sum_{r=1}^{n_i} R_{i,r}^{(1)}(\beta)$  and  $\hat{R}_i^{(1)}(\beta) = n_i^{-1} \sum_{r=1}^{n_i} \hat{R}_{i,r}^{(1)}(\beta)$ . Define  
 $\mathcal{A}_k = \{i : n_i = k, i = 1, \dots, n\}$ ,  $k = 1, \dots, m$ .

Take

$$v_i = Y_i^{-1} R_i^{(1)}(\beta_0) - Y_i R_i^{(1)}(-\beta_0),$$
$$r_i = \frac{E(1/\varepsilon + \varepsilon)}{2(1 - \rho_1)\varphi_0^2(\gamma_0)} \{h_i^{(1)}(\gamma_0) - 2\varphi_0^{-1}(\gamma_0)\varphi_1(\gamma_0)h_i^{(0)}(\gamma_0)\},$$

where  $\rho_1 = \lim |\mathcal{A}_1|/n$ ,  $h_i^{(0)}(\gamma) = \frac{1}{n_i(n_i-1)} \sum_{r \neq s} \exp\{\gamma^T(W_{i,r} - W_{i,s})\}$   
and  $h_i^{(1)}(\gamma) = \frac{1}{n_i(n_i-1)} \sum_{r \neq s} (W_{i,r} - W_{i,s}) \exp\{\gamma^T(W_{i,r} - W_{i,s})\}$ .

Furthermore, define  $V_0 = E[(1/\varepsilon + \varepsilon)ZZ^T]$ .

## Theorem 1

*Under Regularity Conditions,  $\hat{\beta}_{CMS}$  exists and is unique in a neighbourhood of  $\beta_0$  with probability converging to 1 as  $n \rightarrow \infty$ , and  $\hat{\beta}_{CMS} \xrightarrow{P} \beta_0$ . In addition,*

$$\sqrt{n}(\hat{\beta}_{CMS} - \beta_0) \xrightarrow{D} N(0, \Gamma_{CMS}),$$

*where  $\Gamma_{CMS} = V_0^{-1} \Sigma_{CMS} V_0^{-1}$  and*

*$\Sigma_{CMS} = \lim_{n \rightarrow \infty} \frac{1}{n} \sum_{i=1}^n E(v_i - \xi_i J r_i)^{\otimes 2}$ .  $\Gamma_{CMS}$  can be estimated by plug-in method.*

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## Naive Method and Bias

Define  $\bar{W}_{i,\cdot} = \frac{1}{n_i} \sum_{r=1}^{n_i} W_{i,r}$ , and  $\hat{Z}_i = (V_i^T, \bar{W}_{i,\cdot}^T)^T = Z_i + J\bar{U}_{i,\cdot}$ , where  $\bar{U}_{i,\cdot} = \frac{1}{n_i} \sum_{r=1}^{n_i} U_{i,r}$ . A naive computable estimating function  $U_{nv}(\beta)$  can be obtained as follow

$$U_{nv}(\beta) = \sum_{i=1}^n \left\{ Y_i^{-1} \exp(\hat{Z}_i^T \beta) - Y_i \exp(-\hat{Z}_i^T \beta) \right\} \hat{Z}_i = \sum_{i=1}^n \psi(\hat{Z}_i, Y_i, \beta) \quad (6)$$

by replacing  $Z_i$  in (2) with  $\hat{Z}_i$ . Let  $\beta_{Naive}$  be the solution of  $U_{nv}(\beta) = 0_{(p+q) \times 1}$ .  $\beta_{Naive}$  is then the naive-LPRE estimator.

# Naive Method and Bias

A simple algebraic manipulation leads to

$$\begin{aligned} & E[\psi(\hat{Z}_i, Y_i, \beta) | Y_i, Z_i] \\ &= \varphi_0^{n_i} \left( \frac{\gamma}{n_i} \right) \psi(Z_i, Y_i, \beta) + J \left\{ Y_i^{-1} \exp(Z_i^T \beta) + Y_i \exp(-Z_i^T \beta) \right\} \varphi_0^{n_i-1} \left( \frac{\gamma}{n_i} \right) \varphi_1 \left( \frac{\gamma}{n_i} \right) \quad (7) \\ &:= I_{1n}(\beta) + I_{2n}(\beta). \end{aligned}$$

- Bias:  $E[\psi(\hat{Z}_i, Y_i, \beta_0)] = E[I_{2n}(\beta_0)]$ , which might not  $\mathbf{0}$ , resulting in an biased estimating function;
- loss of efficiency: The factor  $\varphi_0^{n_i} \left( \frac{\gamma}{n_i} \right)$  in  $I_{1n}(\beta)$ .

## Corrected estimating equation

In view of the bias and the loss of efficiency, we can construct an unbiased estimating function as

$$U^*(\beta) = \sum_{i=1}^n \tilde{\psi}_i$$

where

$$\tilde{\psi}_i = \left\{ \varphi_0^{n_i} \left( \frac{\gamma}{n_i} \right) \right\}^{-1} \left\{ \psi(\hat{Z}_i, Y_i, \beta) - I_{2n}(\beta) \right\}.$$

However,  $X_i$  in  $I_{2n}(\beta)$  can not be observed.



## Corrected estimating equation

Note that

$$\begin{aligned} & E \left[ Y_i^{-1} \exp(\hat{Z}_i^T \beta) + Y_i \exp(-\hat{Z}_i^T \beta) | \mathcal{U}_i \right] \\ &= \left\{ Y_i^{-1} \exp(Z_i^T \beta) + Y_i \exp(-Z_i^T \beta) \right\} \varphi_0^{n_i} \left( \frac{\gamma}{n_i} \right). \end{aligned} \quad (8)$$

From (8), we have

$$\begin{aligned} & Y_i^{-1} \exp(Z_i^T \beta) + Y_i \exp(-Z_i^T \beta) \\ &= E \left[ \varphi_0^{-n_i} \left( \frac{\gamma}{n_i} \right) \left\{ Y_i^{-1} \exp(\hat{Z}_i^T \beta) + Y_i \exp(-\hat{Z}_i^T \beta) \right\} | \mathcal{U}_i \right]. \end{aligned} \quad (9)$$

## Corrected estimating equation

Therefore, we can define  $\psi_i^*$  as follow,

$$\begin{aligned} \psi_i^* = & \left\{ \varphi_0^{n_i} \left( \frac{\gamma}{n_i} \right) \right\}^{-1} \left\{ \psi(\hat{Z}_i, Y_i, \beta) \right. \\ & \left. - J \left[ Y_i^{-1} \exp(\hat{Z}_i^T \beta) + Y_i \exp(-\hat{Z}_i^T \beta) \right] \frac{\varphi_1(\gamma/n_i)}{\varphi_0(\gamma/n_i)} \right\} \end{aligned}$$

by replacing  $Y_i^{-1} \exp(Z_i^T \beta) + Y_i \exp(-Z_i^T \beta)$  in  $\tilde{\psi}_i$  with  $\varphi_0^{-n_i} \left( \frac{\gamma}{n_i} \right) \left\{ Y_i^{-1} \exp(\hat{Z}_i^T \beta) + Y_i \exp(-\hat{Z}_i^T \beta) \right\}$ .

## Corrected estimating equation

However,  $\varphi_0(\gamma)$  and  $\varphi_1(\gamma)$  in  $\psi_i^*$  are unknown. Define

$$\hat{\psi}_i^* = \{\hat{\varphi}_0^{n_i}(\frac{\gamma}{n_i})\}^{-1} \left\{ \psi(\hat{Z}_i, Y_i, \beta) - J[Y_i^{-1} \exp(\hat{Z}_i^T \beta) + Y_i \exp(-\hat{Z}_i^T \beta)] \frac{\hat{\varphi}_1(\gamma/n_i)}{\hat{\varphi}_0(\gamma/n_i)} \right\},$$

by replacing  $\varphi_0(\gamma/n_i)$  and  $\varphi_1(\gamma/n_i)$  in  $\psi^*$  with  $\hat{\varphi}_0(\gamma/n_i)$  and  $\hat{\varphi}_1(\gamma/n_i)$  given in the previous section, and we obtain an resultant estimating equation for  $\beta_0$  as follow

$$\sum_{i=1}^n \hat{\psi}_i^* = \mathbf{0}.$$

Let  $\hat{\beta}_{CEE}$  be the solution to the above estimating equation.

## Corrected estimating equation

Denote  $\tilde{R}_i^{(1)}(\beta) = \varphi_0^{-n_i}(\gamma/n_i) \exp(\hat{Z}_i^T \beta) \{ \hat{Z}_i - J \frac{\varphi_1(\gamma/n_i)}{\varphi_0(\gamma/n_i)} \}$ .

Let

$$\tilde{v}_i = Y_i^{-1} \tilde{R}_i^{(1)}(\beta_0) - Y_i \tilde{R}_i^{(1)}(-\beta_0),$$

$$\tilde{r}_{i,k} = \frac{E(1/\varepsilon + \varepsilon)}{2(1 - \rho_1)\varphi_0^2(\gamma_0/k)} \{ h_i^{(1)}(\gamma_0/k) - 2\varphi_0^{-1}(\gamma_0/k)\varphi_1(\gamma_0/k)h_i^{(0)}(\gamma_0/k) \},$$

where  $h_i^{(k)}(\gamma)$  ( $k = 0, 1$ ) are defined as the above section. Let

$$\rho_k = \lim_{n \rightarrow \infty} |\mathcal{A}_k|/n.$$

## Theorem 2

*Under regularity Conditions,  $\hat{\beta}_{CEE}$  exists and is unique in a neighbourhood of  $\beta_0$  with probability converging to 1 as  $n \rightarrow \infty$ , and  $\hat{\beta}_{CEE} \xrightarrow{p} \beta_0$ . In addition,*

$$\sqrt{n}(\hat{\beta}_{CEE} - \beta_0) \xrightarrow{D} N(0, \Gamma_{CEE}),$$

*where  $\Gamma_{CEE} = V_0^{-1} \Sigma_{CEE} V_0^{-1}$  and*

*$\Sigma_{CEE} = \lim_{n \rightarrow \infty} \frac{1}{n} \sum_{i=1}^n E\{\tilde{v}_i - \xi_i J \sum_{k=1}^m \rho_k \tilde{r}_{i,k}\}^{\otimes 2}$ . Furthermore,*

*$\Gamma_{CEE}$  can be estimated by plug-in method.*

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# Several Methods

- **CMS**:(Our proposal) Conditional Mean Score based estimating equation approach;
- **CEE**:(Our proposal) Corrected Estimating Equation approach;
- Full: LPRE with the true value of  $X$ ;
- Naive: LPRE with the  $X$  replaced by the average of all the surrogate values

# Simulation Models

- Model:  $Y = \exp(c_0 + \alpha_0 V^* + \gamma_0 X)\varepsilon$ .
  - ▶  $(c_0, \alpha_0, \gamma_0) = (1, 1, 2)$
  - ▶  $(V^*, X)$ : bivariate normal distribution with  $Var(X) = Var(V^*) = 1$  and  $\rho(X, V^*) = 0.5$
  - ▶  $\varepsilon$ : the log-standard normal distribution
- Replication: 1000 times.
- Each of sample size:  $n=200, 300$  or  $500$ .



# Measurement Error Model

- $V^*$  : Measured precisely
- $X$  : Classical additive model:

$$W_{i,j} = X_i + U_{i,j}, \quad i = 1, \dots, n; j = 1, \dots, n_i$$

- The distribution of  $U$ 
  - ▶  $U \sim N(0, \sigma_u^2)$
  - ▶ the standardized  $N(0, 1)$  distribution truncated between  $-c$  and  $c$  and scaled to have standard deviation of  $\sigma_u$
  - ▶  $\sigma_u = 0.5$  or  $0.75$ , inducing a signal-to-noise ratio of 0.8 or 0.64

# Measurement Error Model

- $n_i$

- ▶ **Case 1:**  $n_i = 3$  for all subjects

- ▶ **Case 2:** 1/3 of the population:  $n_i = 1$ ;

- 1/3 of the population:  $n_i = 2$ ;

- 1/3 of the population:  $n_i = 3$ ;

# Scenario 1

- $n_i = 3$  for all the subjects
- $U \sim N(0, \sigma_u^2)$ ,  $\sigma_u = 0.5$  or  $0.75$

# Scenario 1

$n$	$\sigma_u$	method	$\hat{c}$			$\hat{\alpha}$			$\hat{\gamma}$			
			Bias	SE	MSE	Bias	SE	MSE	Bias	SE	MSE	
200	0.50	Full	-0.0031	0.0752	0.0057	-0.0011	0.0872	0.0076	-0.0010	0.0899	0.0081	
		Naive	-0.0034	0.0874	0.0076	0.1003	0.1025	0.0206	-0.1988	0.0997	0.0495	
		CMS	-0.0042	0.0948	0.0090	-0.0048	0.1178	0.0139	0.0148	0.1397	0.0197	
		CEE	-0.0030	0.0894	0.0080	-0.0036	0.1091	0.0119	0.0119	0.1168	0.0138	
	0.75	Full	0.0036	0.0759	0.0058	0.0031	0.0870	0.0076	0.0001	0.0879	0.0077	
		Naive	0.0041	0.1039	0.0108	0.2034	0.1182	0.0553	-0.3967	0.1095	0.1694	
		CMS	0.0034	0.1452	0.0211	-0.0261	0.2074	0.0437	0.0686	0.3243	0.1099	
		CEE	0.0040	0.1127	0.0127	-0.0173	0.1405	0.0200	0.0424	0.1646	0.0289	
	300	0.50	Full	-0.0017	0.0643	0.0041	-0.0016	0.0718	0.0052	-0.0006	0.0711	0.0051
			Naive	-0.0008	0.0755	0.0057	0.0995	0.0837	0.0169	-0.2034	0.0812	0.0479
			CMS	-0.0016	0.0800	0.0064	-0.0042	0.0965	0.0093	0.0069	0.1126	0.0127
			CEE	-0.0011	0.0767	0.0059	-0.0047	0.0885	0.0079	0.0053	0.0966	0.0094
0.75		Full	-0.0009	0.0607	0.0037	0.0010	0.0695	0.0048	0.0008	0.0702	0.0049	
		Naive	-0.0044	0.0896	0.0080	0.1996	0.0940	0.0487	-0.3979	0.0883	0.1661	
		CMS	-0.0059	0.1143	0.0131	-0.0166	0.1752	0.0310	0.0308	0.2581	0.0676	
		CEE	-0.0040	0.0950	0.0090	-0.0142	0.1106	0.0124	0.0252	0.1327	0.0182	
500		0.50	Full	-0.0001	0.0482	0.0023	-0.0019	0.0554	0.0031	0.0008	0.0568	0.0032
			Naive	0.0016	0.0571	0.0033	0.0997	0.0664	0.0143	-0.1986	0.0642	0.0436
			CMS	0.0025	0.0624	0.0039	-0.0024	0.0782	0.0061	0.0068	0.0939	0.0089
			CEE	0.0021	0.0592	0.0035	-0.0021	0.0698	0.0049	0.0046	0.0745	0.0056
	0.75	Full	-0.0013	0.0505	0.0026	-0.0022	0.0537	0.0029	0.0006	0.0551	0.0030	
		Naive	-0.0015	0.0674	0.0045	0.1956	0.0738	0.0437	-0.3992	0.0705	0.1643	
		CMS	-0.0028	0.0932	0.0087	-0.0250	0.1577	0.0255	0.0331	0.2451	0.0612	
		CEE	-0.0024	0.0703	0.0049	-0.0119	0.0865	0.0076	0.0121	0.1041	0.0110	

# Simulation Results for Scenario 1

- The naive estimators for  $\gamma_0$  and  $\alpha_0$  are always seriously biased. Furthermore, the bias of the naive estimator is larger as  $\sigma_u$  becomes larger.
- The two proposed estimators  $\hat{\beta}_{CMS}$  and  $\hat{\beta}_{CEE}$  can effectively correct the biases caused by measurement error.
- The SE of  $\hat{\beta}_{CMS}$  and  $\hat{\beta}_{CEE}$  become smaller as the sample size  $n$  increases.
- The SE of  $\hat{\beta}_{CMS}$  is larger than that of  $\hat{\beta}_{CEE}$ .

## Scenario 2

- $n_i = 3$  for all the subjects
- $U$ : the standardized  $N(0, 1)$  distribution truncated between  $-c$  and  $c$  and scaled to have standard deviation of  $\sigma_u$ , where  $c = 2$ ,  $\sigma_u = 0.5$  or  $0.75$

# Scenario 2

$n$	$\sigma_u$	method	$\hat{c}$			$\hat{\alpha}$			$\hat{\gamma}$			
			Bias	SE	MSE	Bias	SE	MSE	Bias	SE	MSE	
200	0.50	Full	0.0019	0.0764	0.0058	-0.0007	0.0862	0.0074	0.0032	0.0866	0.0075	
		Naive	0.0036	0.0898	0.0081	<b>0.1006</b>	0.0992	0.0199	<b>-0.1984</b>	0.0943	0.0483	
		CMS	0.0021	0.0910	0.0083	-0.0015	0.1054	0.0111	0.0058	0.1101	0.0122	
		CEE	0.0028	0.0902	0.0081	-0.0026	0.1053	0.0111	0.0082	0.1097	0.0121	
	0.75	Full	0.0025	0.0760	0.0058	0.0016	0.0894	0.0080	-0.0019	0.0879	0.0077	
		Naive	0.0047	0.0979	0.0096	<b>0.2001</b>	0.1178	0.0539	<b>-0.4018</b>	0.1068	0.1728	
		CMS	0.0004	0.1031	0.0106	-0.0005	0.1298	0.0168	0.0087	0.1460	0.0214	
		CEE	0.0036	0.1015	0.0103	-0.0096	0.1355	0.0184	0.0199	0.1561	0.0248	
	300	0.50	Full	-0.0011	0.0627	0.0039	-0.0004	0.0716	0.0051	-0.0011	0.0696	0.0048
			Naive	-0.0014	0.0713	0.0051	<b>0.0972</b>	0.0837	0.0165	<b>-0.1988</b>	0.0789	0.0457
			CMS	-0.0013	0.0729	0.0053	-0.0038	0.0861	0.0074	0.0045	0.0898	0.0081
			CEE	-0.0013	0.0726	0.0053	-0.0048	0.0861	0.0074	0.0063	0.0910	0.0083
0.75		Full	0.0003	0.0634	0.0040	0.0035	0.0734	0.0054	0.0021	0.0732	0.0054	
		Naive	0.0008	0.0859	0.0074	<b>0.2038</b>	0.0959	0.0507	<b>-0.3941</b>	0.0856	0.1627	
		CMS	0.0009	0.0903	0.0082	-0.0009	0.1072	0.0115	0.0119	0.1199	0.0145	
		CEE	0.0023	0.0890	0.0079	-0.0037	0.1099	0.0121	0.0206	0.1216	0.0152	
500		0.50	Full	0.0005	0.0476	0.0023	0.0020	0.0540	0.0029	-0.0008	0.0553	0.0031
			Naive	0.0007	0.0555	0.0031	<b>0.0999</b>	0.0644	0.0141	<b>-0.1978</b>	0.0622	0.0430
			CMS	0.0003	0.0557	0.0031	-0.0024	0.0669	0.0045	0.0047	0.0702	0.0050
			CEE	0.0005	0.0559	0.0031	-0.0020	0.0680	0.0046	0.0048	0.0709	0.0050
	0.75	Full	-0.0001	0.0487	0.0024	-0.0034	0.0560	0.0032	0.0032	0.0562	0.0032	
		Naive	-0.0030	0.0658	0.0043	<b>0.1961</b>	0.0700	0.0434	<b>-0.3942</b>	0.0653	0.1596	
		CMS	-0.0024	0.0670	0.0045	-0.0065	0.0804	0.0065	0.0141	0.0928	0.0088	
		CEE	-0.0023	0.0682	0.0047	-0.0083	0.0811	0.0066	0.0148	0.0954	0.0093	

## Simulation Results for Scenario 2

- Results for Scenario 2 shows similar patterns as results for Scenario 1 except that the SE of  $\hat{\beta}_{CEE}$  is a little larger than that of  $\hat{\beta}_{CMS}$ .



## Scenario 3

- 1/3 of the population:  $n_i = 1$ ;  
1/3 of the population:  $n_i = 2$ ;  
1/3 of the population:  $n_i = 3$ ;
- $U \sim N(0, \sigma_u^2)$ ,  $\sigma_u = 0.5$  or  $0.75$

# Scenario 3

$n$	$\sigma_u$	method	$\hat{c}$			$\hat{\alpha}$			$\hat{\gamma}$			
			Bias	SE	MSE	Bias	SE	MSE	Bias	SE	MSE	
200	0.50	Full	-0.0031	0.0752	0.0057	-0.0011	0.0872	0.0076	-0.0010	0.0899	0.0081	
		Naive	-0.0062	0.0993	0.0099	<b>0.1702</b>	0.1165	0.0426	<b>-0.3440</b>	0.1091	0.1302	
		CMS	-0.0058	0.1084	0.0118	-0.0140	0.1461	0.0215	0.0287	0.1855	0.0352	
		CEE	-0.0050	0.1036	0.0108	-0.0112	0.1341	0.0181	0.0227	0.1519	0.0236	
	0.75	Full	0.0036	0.0759	0.0058	0.0031	0.0870	0.0076	0.0001	0.0879	0.0077	
		Naive	0.0041	0.1271	0.0162	<b>0.3300</b>	0.1396	0.1283	<b>-0.6454</b>	0.1212	0.4312	
		CMS	0.0064	0.1656	0.0275	-0.0119	0.2363	0.0560	0.0415	0.3422	0.1188	
		CEE	0.0081	0.1500	0.0226	-0.0230	0.2000	0.0405	0.0640	0.2849	0.0853	
	300	0.50	Full	-0.0017	0.0643	0.0041	-0.0016	0.0718	0.0052	-0.0006	0.0711	0.0051
			Naive	-0.0001	0.0851	0.0072	<b>0.1747</b>	0.0932	0.0392	<b>-0.3543</b>	0.0885	0.1334
			CMS	-0.0029	0.0902	0.0082	-0.0136	0.1305	0.0172	0.0232	0.1803	0.0331
			CEE	-0.0011	0.0872	0.0076	-0.0092	0.1091	0.0120	0.0136	0.1286	0.0167
0.75		Full	-0.0009	0.0607	0.0037	0.0010	0.0695	0.0048	0.0008	0.0702	0.0049	
		Naive	-0.0036	0.1073	0.0115	<b>0.3258</b>	0.1088	0.1180	<b>-0.6485</b>	0.0969	0.4299	
		CMS	-0.0068	0.1340	0.0180	-0.0155	0.2016	0.0409	0.0334	0.3147	0.1001	
		CEE	-0.0031	0.1196	0.0143	-0.0253	0.1551	0.0247	0.0485	0.2265	0.0536	
500		0.50	Full	-0.0001	0.0482	0.0023	-0.0019	0.0554	0.0031	0.0008	0.0568	0.0032
			Naive	0.0010	0.0651	0.0042	<b>0.1756</b>	0.0756	0.0366	<b>-0.3479</b>	0.0724	0.1263
			CMS	0.0016	0.0718	0.0052	-0.0046	0.1004	0.0101	0.0133	0.1443	0.0210
			CEE	0.0018	0.0692	0.0048	-0.0025	0.0847	0.0072	0.0087	0.1024	0.0106
	0.75	Full	-0.0013	0.0505	0.0026	-0.0022	0.0537	0.0029	0.0006	0.0551	0.0030	
		Naive	-0.0041	0.0837	0.0070	<b>0.3225</b>	0.0876	0.1117	<b>-0.6490</b>	0.0804	0.4277	
		CMS	-0.0022	0.1092	0.0119	-0.0245	0.1826	0.0340	0.0389	0.3002	0.0916	
		CEE	-0.0048	0.0945	0.0090	-0.0205	0.1293	0.0171	0.0294	0.1875	0.0360	

## Simulation Results for Scenario 3

- Results for Scenario 3 shows similar patterns as results for Scenario 1.
- The SE of Scenario 3 is a litter larger than that of Scenario 1. The reason is that  $1/3$  of the population has only one surrogate.

## Scenario 4

- 1/3 of the population:  $n_i = 1$ ;
- 1/3 of the population:  $n_i = 2$ ;
- 1/3 of the population:  $n_i = 3$ ;
- $U$ : the standardized  $N(0, 1)$  distribution truncated between  $-c$  and  $c$  and scaled to have standard deviation of  $\sigma_u$ , where  $c = 2$ ,  $\sigma_u = 0.5$  or  $0.75$

# Scenario 4

$n$	$\sigma_u$	method	$\hat{c}$			$\hat{\alpha}$			$\hat{\gamma}$			
			Bias	SE	MSE	Bias	SE	MSE	Bias	SE	MSE	
200	0.50	Full	0.0019	0.0764	0.0058	-0.0007	0.0862	0.0074	0.0032	0.0866	0.0075	
		Naive	0.0045	0.1002	0.0101	0.1688	0.1100	0.0406	-0.3356	0.1026	0.1231	
		CMS	0.0026	0.1009	0.0102	-0.0053	0.1180	0.0140	0.0128	0.1283	0.0166	
		CEE	0.0027	0.1006	0.0101	-0.0063	0.1188	0.0142	0.0148	0.1287	0.0168	
	0.75	Full	0.0025	0.0760	0.0058	0.0016	0.0894	0.0080	-0.0019	0.0879	0.0077	
		Naive	0.0021	0.1185	0.0141	0.3150	0.1334	0.1170	-0.6320	0.1130	0.4122	
		CMS	0.0002	0.1236	0.0153	-0.0053	0.1604	0.0257	0.0166	0.1919	0.0371	
		CEE	0.0007	0.1250	0.0156	-0.0123	0.1596	0.0256	0.0272	0.1910	0.0372	
	300	0.50	Full	-0.0011	0.0627	0.0039	-0.0004	0.0716	0.0051	-0.0011	0.0696	0.0048
			Naive	-0.0015	0.0810	0.0066	0.1674	0.0931	0.0367	-0.3378	0.0871	0.1217
			CMS	-0.0010	0.0831	0.0069	-0.0063	0.0966	0.0094	0.0102	0.1091	0.0120
			CEE	-0.0010	0.0829	0.0069	-0.0071	0.0979	0.0096	0.0123	0.1105	0.0124
0.75		Full	0.0003	0.0634	0.0040	0.0035	0.0734	0.0054	0.0021	0.0732	0.0054	
		Naive	-0.0050	0.1041	0.0109	0.3191	0.1065	0.1131	-0.6278	0.0913	0.4024	
		CMS	-0.0026	0.1072	0.0115	-0.0051	0.1292	0.0167	0.0170	0.1581	0.0253	
		CEE	-0.0030	0.1100	0.0121	-0.0094	0.1300	0.0170	0.0269	0.1575	0.0255	
500		0.50	Full	0.0005	0.0476	0.0023	0.0020	0.0540	0.0029	-0.0008	0.0553	0.0031
			Naive	0.0026	0.0631	0.0040	0.1706	0.0731	0.0345	-0.3383	0.0670	0.1190
			CMS	0.0020	0.0631	0.0040	-0.0013	0.0793	0.0063	0.0032	0.0848	0.0072
			CEE	0.0026	0.0636	0.0040	-0.0021	0.0798	0.0064	0.0045	0.0837	0.0070
	0.75	Full	-0.0001	0.0487	0.0024	-0.0034	0.0560	0.0032	0.0032	0.0562	0.0032	
		Naive	-0.0035	0.0799	0.0064	0.3123	0.0835	0.1045	-0.6264	0.0720	0.3975	
		CMS	-0.0004	0.0790	0.0062	-0.0065	0.1013	0.0103	0.0147	0.1225	0.0152	
		CEE	-0.0003	0.0808	0.0065	-0.0099	0.1013	0.0104	0.0203	0.1203	0.0149	

## Simulation Results for Scenario 4

- Results for Scenario 4 shows similar patterns as results for Scenario 2.
- The SE of Scenario 4 is a litter larger than that of Scenario 2. The reason is that  $1/3$  of the population has only one surrogate.

- 1 Introduction
- 2 Measurement Error Model
- 3 Conditional Mean Score Based Estimating Equation Approach
- 4 Corrected estimating equation method
- 5 Simulation Studies
- 6 Real Data Analysis

# Data Description

- **Data:** ACTG315 data, which is available at <https://www.urmc.rochester.edu/biostat/people/faculty/wusite/datasets/ACTG315LongitudinalDataViralLoad.cfm>
- **Aim:** The relationship between viral load and CD4+ cell counts of the first two days of treatment.
- **Response Y:** The average load of viral load of the first two days of treatment
- **Predictor X:** The average counts of CD4+ cell counts of the first two days of treatment.



- **Measurement Error Model:**

$$W_{i,r} = X_i + U_{i,r}, \quad r = 1, \dots, n_i, \quad i = 1, \dots, 45$$

- **Model:**

$$Y_i = \exp(c_0 + \gamma_0 X_i) \varepsilon_i$$

# Data Analysis

Table: Analysis of the ACTG315 data with LS(Least Square), CMS, and CEE

	LS		CMS		CEE	
	Est	p-value	Est	p-value	Est	p-value
$c_0$	12.217(0.436)	0	12.383(0.401)	0	12.424(0.416)	0
$\gamma_0$	-0.377(0.216)	0.040	-0.491(0.212)	0.010	-0.514(0.220)	0.010

*Thank you!*