

# Efficient Estimation of Causal Effects Under Two-Phase Sampling with Error-Prone Outcome and Treatment Measurements

ENAR 2025 Spring Meeting

---

**Keith Barnatchez**, Kevin Josey, Nima Hejazi, Giovanni Parmigiani, Bryan E. Shepherd, and Rachel Nethery

March 24<sup>th</sup>, 2025

Background

Methods

Data Application

Discussion

# Motivation

Explosion of the use of electronic health record (EHR) data for conducting **observational causal inference** studies

- Where, at a high level, one is interested in some measure of causal effect of a treatment  $A$  on an outcome of interest  $Y$

# Motivation

Explosion of the use of electronic health record (EHR) data for conducting **observational causal inference** studies

- Where, at a high level, one is interested in some measure of causal effect of a treatment  $A$  on an outcome of interest  $Y$

And for good reason!

- EHR data is typically cheaper to obtain, fairly representative of patient populations, rich in information on potential confounding factors  $X$ , and **big**

# Motivation

Explosion of the use of electronic health record (EHR) data for conducting **observational causal inference** studies

- Where, at a high level, one is interested in some measure of causal effect of a treatment  $A$  on an outcome of interest  $Y$

And for good reason!

- EHR data is typically cheaper to obtain, fairly representative of patient populations, rich in information on potential confounding factors  $X$ , and **big**

But EHR data tends to present numerous **challenges**

- Including **measurement error** in outcomes ( $Y$ ) and treatments ( $A$ ) of interest

## EHR data: in a perfect world

Suppose we aim to estimate the average treatment effect of a binary treatment  $A$  on  $Y$

$$\tau = \mathbb{E}[Y(1) - Y(0)]$$

## EHR data: in a perfect world

Suppose we aim to estimate the average treatment effect of a binary treatment  $A$  on  $Y$

$$\tau = \mathbb{E}[Y(1) - Y(0)]$$

$Y$	$A$	$\mathbf{X}$
$Y_1$	$A_1$	$\mathbf{X}_1$
$Y_2$	$A_2$	$\mathbf{X}_2$
$Y_3$	$A_3$	$\mathbf{X}_3$
$Y_4$	$A_4$	$\mathbf{X}_4$
$Y_5$	$A_5$	$\mathbf{X}_5$
$Y_6$	$A_6$	$\mathbf{X}_6$

In a perfect world, we'd have access to the true outcome + treatment values (+ covariates  $\mathbf{X}$ )

## EHR data: in a perfect world reality

$Y$	$Y^*$	$A$	$A^*$	$X$
NA	$Y_1^*$	NA	$A_1^*$	$X_1$
NA	$Y_2^*$	NA	$A_2^*$	$X_2$
NA	$Y_3^*$	NA	$A_3^*$	$X_3$
NA	$Y_4^*$	NA	$A_4^*$	$X_4$
NA	$Y_5^*$	NA	$A_5^*$	$X_5$
NA	$Y_6^*$	NA	$A_6^*$	$X_6$



## EHR data: in a perfect world reality

$Y$	$Y^*$	$A$	$A^*$	$X$
NA	$Y_1^*$	NA	$A_1^*$	$X_1$
NA	$Y_2^*$	NA	$A_2^*$	$X_2$
NA	$Y_3^*$	NA	$A_3^*$	$X_3$
NA	$Y_4^*$	NA	$A_4^*$	$X_4$
NA	$Y_5^*$	NA	$A_5^*$	$X_5$
NA	$Y_6^*$	NA	$A_6^*$	$X_6$

In practice, we often only have **error-prone measurements** of  $Y$  and  $A$ , denoted  $Y^*$  and  $A^*$

- Well-documented that using  $Y^*$  and  $A^*$  in place of  $Y$  and  $A$  can lead to severely **biased** causal effect estimates

## EHR data: a study design-based workaround

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

In practice, can sometimes spend time + money to obtain gold-standard measurements for a random (typically small) subset of this data

## EHR data: a study design-based workaround

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

In practice, [selection](#) into this subset can often be controlled, and may depend on the initial error-prone measurements:  $R \not\perp A^*, Y^*, X$

- Especially common sampling strategy if  $Y$  and/or  $A$  are rare

## EHR data: a study design-based workaround

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

In practice, [selection](#) into this subset can often be controlled, and may depend on the initial error-prone measurements:  $R \not\perp A^*, Y^*, X$

- **Intuition:** over-sample subjects who contribute more information to the target estimand

# Our Contributions

At a high-level, our work addresses the following question: How do we estimate causal effects nonparametrically when

At a high-level, our work addresses the following question: How do we estimate causal effects nonparametrically when

1. We have **error-prone** outcome + treatment measurements for all subjects

At a high-level, our work addresses the following question: How do we estimate causal effects nonparametrically when

1. We have **error-prone** outcome + treatment measurements for all subjects
2. We have **gold-standard** treatment + outcome measurements for a *subset* of our EHR data, where...

At a high-level, our work addresses the following question: How do we estimate causal effects nonparametrically when

1. We have **error-prone** outcome + treatment measurements for all subjects
2. We have **gold-standard** treatment + outcome measurements for a *subset* of our EHR data, where...
3. This subset was collected according to a sampling rule that is **dependent** on the initially observed data:  $X, A^*$  and  $Y^*$  all influence  $R$



At a high-level, our work addresses the following question: How do we estimate causal effects nonparametrically when

1. We have **error-prone** outcome + treatment measurements for all subjects
2. We have **gold-standard** treatment + outcome measurements for a *subset* of our EHR data, where...
3. This subset was collected according to a sampling rule that is **dependent** on the initially observed data:  $X, A^*$  and  $Y^*$  all influence  $R$

We present two **asymptotically equivalent** approaches to constructing efficient nonparametric causal effect estimators

Background

Methods

Approach 1

Approach 2

Data Application

Discussion

Background

Methods

Approach 1

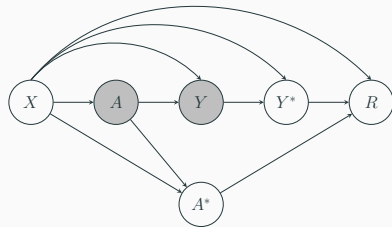
Approach 2

Data Application

Discussion

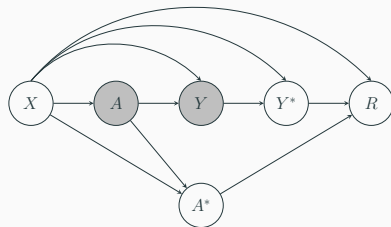
## Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



## Approach 1: Using the observed data distribution

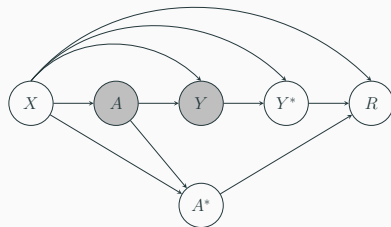
$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



**Main goal:** Estimate counterfactual means  $\mathbb{E}[Y(a)]$  ( $a \in \{0, 1\}$ ) *efficiently*, allowing  $R$  to depend on  $X$ ,  $Y^*$  and  $A^*$

# Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

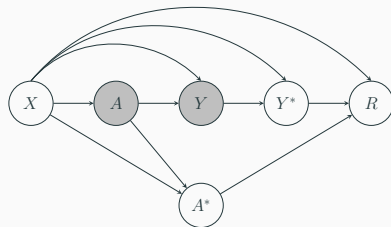


Construction of estimators follows standard **semiparametric estimation** pipeline:



# Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

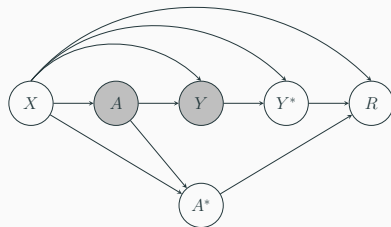


Construction of estimators follows standard **semiparametric estimation** pipeline:



# Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



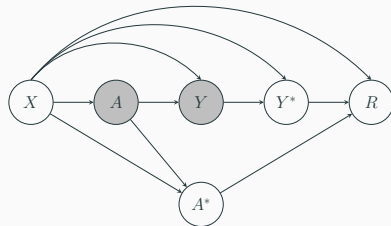
Construction of estimators follows standard **semiparametric estimation** pipeline:

$$\begin{array}{ccccccc}
 \text{causal estimand} & & & & \text{plug-in estimator} & & \text{debiased estimator} \\
 \mathbb{E}[Y(a)] & \xrightarrow{\text{Assumptions}} & \mathbb{E}[f(\text{data})] & \longrightarrow & \hat{\psi}_a^{\text{PI}} = \frac{1}{n} \sum_{i=1}^n \hat{f}(\text{data}_i) & \xrightarrow{\text{Additional work}} & \hat{\psi}_a^{\text{DB}}
 \end{array}$$



# Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

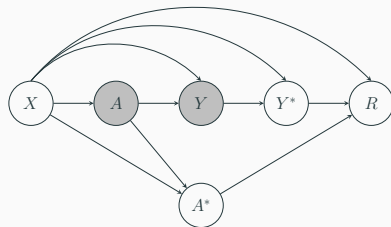


Construction of estimators follows standard **semiparametric estimation** pipeline:

$$\begin{array}{ccccccc}
 \text{causal estimand} & & \text{stat. estimand} & & \text{plug-in estimator} & & \text{debiased est.} \\
 \mathbb{E}[Y(a)] & \xrightarrow{\text{Assumptions}} & \mathbb{E}[f(\text{data})] & \longrightarrow & \hat{\psi}_a^{\text{PI}} = \frac{1}{n} \sum_{i=1}^n \hat{f}(\text{data}_i) & \xrightarrow{\text{(tedious) work}} & \hat{\psi}_a^{\text{OS}}
 \end{array}$$

## Approach 1: Using the observed data distribution

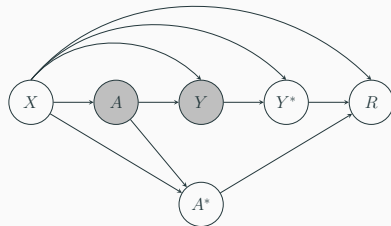
$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



Letting  $\mathbf{Z} = (\mathbf{X}, A^*, Y^*)$ , we derive a **plug-in** estimator

## Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



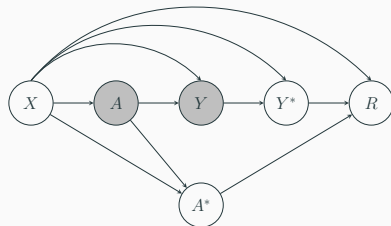
Letting  $\mathbf{Z} = (\mathbf{X}, A^*, Y^*)$ , we derive a **plug-in** estimator

$$\hat{\psi}_a^{\text{PI},1} = \frac{1}{n} \sum_{i=1}^n \frac{\hat{\mathbb{E}}[\hat{\lambda}_a(\mathbf{Z}) \cdot \hat{\mu}_a(\mathbf{Z}) | \mathbf{X}]}{\hat{\mathbb{E}}[\hat{\lambda}_a(\mathbf{Z}) | \mathbf{X}]}$$

where  $\hat{\lambda}_a$  and  $\hat{\mu}_a$  are imputation functions for  $A$  and  $Y$ , respectively

## Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



Letting  $\mathbf{Z} = (\mathbf{X}, A^*, Y^*)$ , we derive a **plug-in** estimator

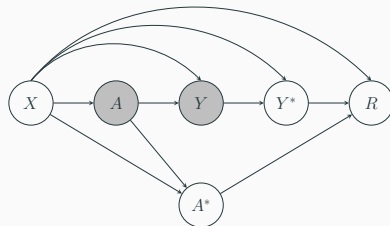
$$\hat{\psi}_a^{\text{PI},1} = \frac{1}{n} \sum_{i=1}^n \frac{\hat{\mathbb{E}}[\hat{\lambda}_a(\mathbf{Z}) \cdot \hat{\mu}_a(\mathbf{Z}) | \mathbf{X}]}{\hat{\mathbb{E}}[\hat{\lambda}_a(\mathbf{Z}) | \mathbf{X}]}$$

where  $\hat{\lambda}_a$  and  $\hat{\mu}_a$  are imputation functions for  $A$  and  $Y$ , respectively

- Interpretation: IPW on imputed values, after marginalizing out post-treatment variables

## Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



Letting  $\mathbf{Z} = (\mathbf{X}, A^*, Y^*)$ , we derive a **plug-in** estimator

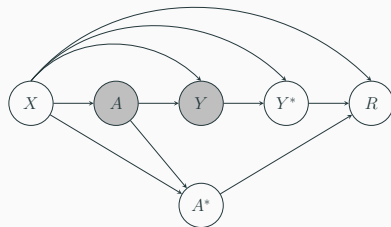
$$\hat{\psi}_a^{\text{PI},1} = \frac{1}{n} \sum_{i=1}^n \frac{\hat{\mathbb{E}}[\hat{\lambda}_a(\mathbf{Z}) \cdot \hat{\mu}_a(\mathbf{Z}) | \mathbf{X}]}{\hat{\mathbb{E}}[\hat{\lambda}_a(\mathbf{Z}) | \mathbf{X}]}$$

where  $\hat{\lambda}_a$  and  $\hat{\mu}_a$  are imputation functions for  $A$  and  $Y$ , respectively

- Drawback: Inference intractable when nuisance models are fit data-adaptively

## Approach 1: Using the observed data distribution

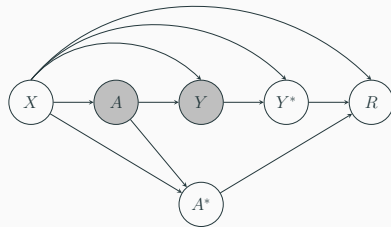
$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



To enable inference, we derive a one-step **debiased** estimator

## Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



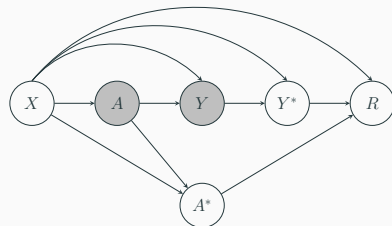
To enable inference, we derive a one-step **debiased** estimator

$$\hat{\psi}_a^{\text{OS},1} = \frac{1}{n} \sum_{i=1}^n \frac{\hat{\mathbb{E}}[\hat{\lambda}_a(\mathbf{Z}) \cdot \hat{\mu}_a(\mathbf{Z}) | \mathbf{X}]}{\hat{\mathbb{E}}[\hat{\lambda}_a(\mathbf{Z}) | \mathbf{X}]} + \widehat{\text{BC}}$$

where  $\widehat{\text{BC}}$  is a *bias correction* term based on the **efficient influence function** for  $\psi_a$

## Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

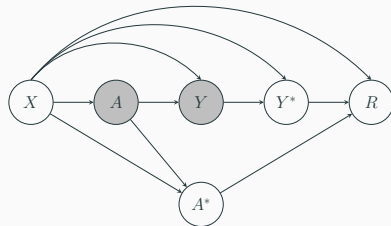


We document multiple properties of  $\hat{\psi}_a^{\text{OS},1}$



## Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

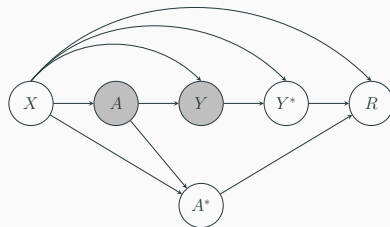


We document multiple properties of  $\hat{\psi}_a^{\text{OS},1}$

1. Bias correction enables **valid inference** when nuisance models are fit with flexible ML methods that converge at  $n^{1/4}$  rates, however...

## Approach 1: Using the observed data distribution

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



We document multiple properties of  $\hat{\psi}_a^{\text{OS},1}$

1. Bias correction enables **valid inference** when nuisance models are fit with flexible ML methods that converge at  $n^{1/4}$  rates, however...
2. This bias correction term introduces numerous **unstable** weighting terms that can harm finite sample performance
  - Particularly concerning, as validation samples tend to be small in practice

Background

Methods

Approach 1

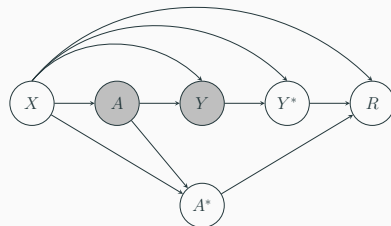
Approach 2

Data Application

Discussion

## Approach 2: Complete-data projection

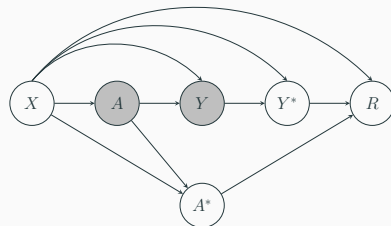
$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



Approach 2 is based on a well-developed, but relatively underutilized, framework for constructing **debiased estimators** under missing data (van der Laan and Robins 2003)

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

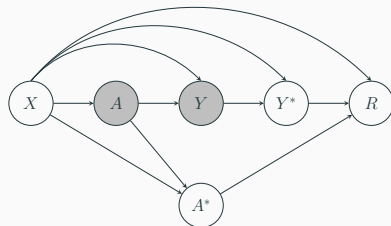


Approach 2 is based on a well-developed, but relatively underutilized, framework for constructing **debiased estimators** under missing data (van der Laan and Robins 2003)

- Links (i) the estimator we'd ideally construct under **complete data** to (ii) the observed data structure, where  $Y$  and  $A$  are partially missing

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



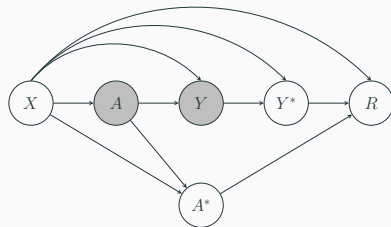
**Idea:** If we had complete data, could construct a plug-in estimator  $\hat{\psi}_a^{\text{PI},2} = \frac{1}{n} \sum_{i=1}^n \hat{m}_a(\mathbf{X}_i)$ ,

$$\hat{\psi}_a^{\text{PI},2} = \hat{\psi}_a^{\text{PI},1} + \frac{1}{n} \sum_{i=1}^n \left( m_a(\mathbf{X}_i) + \frac{I(A_i = a)}{p_a(\mathbf{X}_i)} (D_i - m_a(\mathbf{X}_i)) - \hat{m}_a(\mathbf{X}_i) \right)$$

where  $m_a(\mathbf{X}) = \mathbb{E}(Y|A = a, \mathbf{X})$  and  $p_a(\mathbf{X}) = \mathbb{P}(A = a|\mathbf{X})$

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



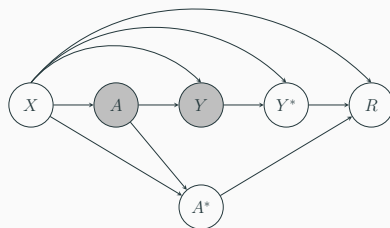
**Idea:** If we had complete data, could construct a plug-in estimator  $\hat{\psi}_a^{\text{PI},2} = \frac{1}{n} \sum_{i=1}^n \hat{m}_a(\mathbf{X}_i)$ , as well as an AIPW estimator

$$\hat{\psi}_a^{\text{OS},2} = \hat{\psi}_a^{\text{PI},2} + \frac{1}{n} \sum_{i=1}^n \left( \hat{m}_a(\mathbf{X}_i) + \frac{I(A_i = a)}{\hat{g}_a(\mathbf{X}_i)} \{Y_i - \hat{m}_a(\mathbf{X}_i)\} - \hat{\psi}_a^{\text{PI},2} \right)$$

where  $\hat{m}_a(\mathbf{X}) = \mathbb{E}(Y|A = a, \mathbf{X})$  and  $\hat{g}_a(\mathbf{X}) = \mathbb{P}(A = a|\mathbf{X})$

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



**Idea:** If we had complete data, could construct a plug-in estimator  $\hat{\psi}_a^{\text{PI},2} = \frac{1}{n} \sum_{i=1}^n \hat{m}_a(\mathbf{X}_i)$ , as well as an AIPW estimator

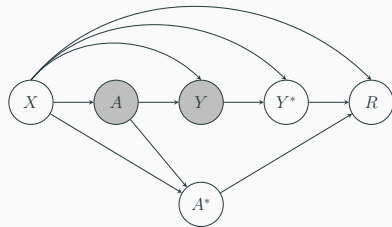
$$\hat{\psi}_a^{\text{OS},2} = \hat{\psi}_a^{\text{PI},2} + \frac{1}{n} \sum_{i=1}^n \left( \hat{m}_a(\mathbf{X}_i) + \frac{I(A_i = a)}{\hat{g}_a(\mathbf{X}_i)} \{Y_i - \hat{m}_a(\mathbf{X}_i)\} - \hat{\psi}_a^{\text{PI},2} \right)$$

Above,  $\hat{m}_a(\mathbf{X})$  and  $\hat{g}_a(\mathbf{X})$  can be estimated with weighted regressions that add weights  $R/\mathbb{P}(R=1|\mathbf{Z})$  to the underlying loss function



## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



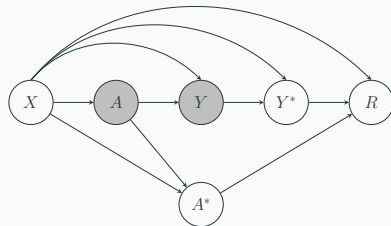
$$\hat{\psi}_a^{\text{OS},2} = \hat{\psi}_a^{\text{PI},2} + \frac{1}{n} \sum_{i=1}^n \left( \hat{m}_a(\mathbf{X}_i) + \frac{I(A_i = a)}{\hat{g}_a(\mathbf{X}_i)} \{Y_i - \hat{m}_a(\mathbf{X}_i)\} - \hat{\psi}_a^{\text{PI},2} \right)$$

**Issue:** The bias correction terms are only observed when  $R_i = 1$

- Above estimator is **infeasible**

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

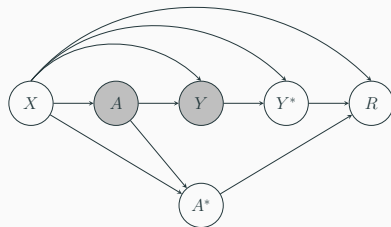


**Key idea:** Treat the bias correction terms as **pseudo-outcomes**:

$$\begin{aligned}\hat{\psi}_a^{\text{OS},2} &= \hat{\psi}_a^{\text{PI},2} + \frac{1}{n} \sum_{i=1}^n \left( \hat{m}_a(\mathbf{X}_i) + \frac{I(A_i = a)}{\hat{g}_a(\mathbf{X}_i)} \{Y_i - \hat{m}_a(\mathbf{X}_i)\} - \hat{\psi}_a^{\text{PI},2} \right) \\ &= \hat{\psi}_a^{\text{PI},2} + \frac{1}{n} \sum_{i=1}^n \underbrace{\chi_a(\mathbf{O}_i; \hat{m}_a, \hat{g}_a)}_{\text{pseudo outcome}}\end{aligned}$$

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

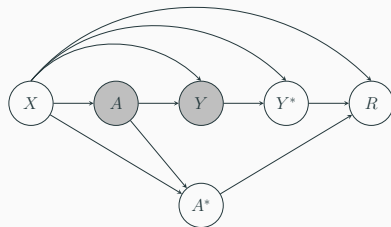


**Key idea:** Perform AIPW on the **pseudo-outcome**:

$$\hat{\psi}_a^{\text{OS},2} = \hat{\psi}_a^{\text{PI},2} + \frac{1}{n} \sum_{i=1}^n \chi_a(\mathbf{O}_i; \hat{m}_a, \hat{g}_a)$$

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1

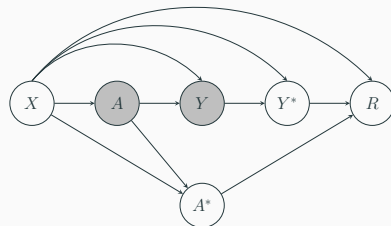


**Key idea:** Perform AIPW on the **pseudo-outcome**:

$$\hat{\psi}_a^{\text{OS},2} = \hat{\psi}_a^{\text{PI},2} + \frac{1}{n} \sum_{i=1}^n \left( \hat{\varphi}_a(\mathbf{Z}_i) + \frac{R_i}{\mathbb{P}(R_i = 1 | \mathbf{Z}_i)} \{ \chi_a(\mathbf{O}_i; \hat{m}_a, \hat{g}_a) - \hat{\varphi}_a(\mathbf{Z}_i) \} \right)$$

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



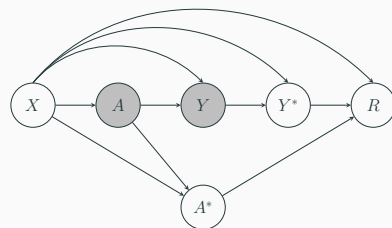
**Key idea:** Perform AIPW on the **pseudo-outcome**:

$$\hat{\psi}_a^{\text{OS},2} = \hat{\psi}_a^{\text{PI},2} + \frac{1}{n} \sum_{i=1}^n \left( \hat{\varphi}_a(\mathbf{Z}_i) + \frac{R_i}{\mathbb{P}(R_i = 1 | \mathbf{Z}_i)} \{ \chi_a(\mathbf{O}_i; \hat{m}_a, \hat{g}_a) - \hat{\varphi}_a(\mathbf{Z}_i) \} \right)$$

where  $\hat{\varphi}_a(\mathbf{Z}) = \hat{\mathbb{E}}[\chi_a(\mathbf{O}; \hat{m}_a, \hat{g}_a) | \mathbf{Z}, R = 1]$

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



**Key idea:** Perform AIPW on the **pseudo-outcome**:

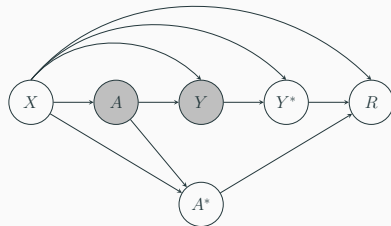
$$\hat{\psi}_a^{\text{OS},2} = \hat{\psi}_a^{\text{PI},2} + \frac{1}{n} \sum_{i=1}^n \left( \hat{\varphi}_a(\mathbf{Z}_i) + \frac{R_i}{\mathbb{P}(R_i = 1 | \mathbf{Z}_i)} \{ \chi_a(\mathbf{O}_i; \hat{m}_a, \hat{g}_a) - \hat{\varphi}_a(\mathbf{Z}_i) \} \right)$$

where  $\hat{\varphi}_a(\mathbf{Z}) = \mathbb{E}[\chi_a(\mathbf{O}_i; \hat{m}_a, \hat{g}_a) | \mathbf{Z}, R = 1]$

- Feasible estimator!
- When sampling probabilities are known,  $\hat{\psi}_a^{\text{OS},2}$  is doubly-robust in the traditional sense

## Approach 2: Complete-data projection

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	0
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	0
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
$Y_5$	$Y_5^*$	$A_5$	$A_5^*$	$X_5$	0
$Y_6$	$Y_6^*$	$A_6$	$A_6^*$	$X_6$	1



Connections between  $\hat{\psi}_a^{\text{OS},1}$  and  $\hat{\psi}_a^{\text{OS},2}$ :

- Can be shown that asymptotically, Approach 1 and Approach 2 estimators are *equivalent*
  - Approach 2 can be viewed as a re-parameterization of Approach 1
- Both estimators have unique sources of finite sample **instability**

Background

Methods

Data Application

Discussion



# The Vanderbilt Comprehensive Care Clinic (VCCC)

EHR database with information on  $\approx 1900$  patients living with HIV between 1998-2011 that began receiving care at the VCCC

- Numerous variables recorded with error
  - Date of antiretroviral therapy (ART) initiation
  - Occurrence of AIDS-defining events (ADEs)

# The Vanderbilt Comprehensive Care Clinic (VCCC)

EHR database with information on  $\approx 1900$  patients living with HIV between 1998-2011 that began receiving care at the VCCC

- Numerous variables recorded with error
  - Date of antiretroviral therapy (ART) initiation
  - Occurrence of AIDS-defining events (ADEs)
- Reliable measurements on demographic information, risk factors and baseline lab measurements (baseline covariates,  $\mathbf{X}$ )

# The Vanderbilt Comprehensive Care Clinic (VCCC)

EHR database with information on  $\approx 1900$  patients living with HIV between 1998-2011 that began receiving care at the VCCC

- Numerous variables recorded with error
  - Date of antiretroviral therapy (ART) initiation
  - Occurrence of AIDS-defining events (ADEs)
- Reliable measurements on demographic information, risk factors and baseline lab measurements (baseline covariates,  $\mathbf{X}$ )
- Team at Vanderbilt University Medical Center **manually validated** every patient's observation in the database
  - Effectively yielding a **fully validated** dataset alongside an **error-prone** dataset

# The Vanderbilt Comprehensive Care Clinic (VCCC)

EHR database with information on  $\approx 1900$  patients living with HIV between 1998-2011 that began receiving care at the VCCC

- Numerous variables recorded with error
  - Date of antiretroviral therapy (ART) initiation
  - Occurrence of AIDS-defining events (ADEs)
- Reliable measurements on demographic information, risk factors and baseline lab measurements (baseline covariates,  $\mathbf{X}$ )
- Team at Vanderbilt University Medical Center **manually validated** every patient's observation in the database
  - Effectively yielding a **fully validated** dataset alongside an **error-prone** dataset

**Causal estimand:** Average causal effect of starting ART within 1 month of first visit ( $A$ ) on 3-year (post initial visit) risk of suffering an ADE ( $Y$ )

## VCCC application

“Reveal” increasingly larger shares of the validated data  $\mathbb{P}(R = 1)$ , over-sampling those with  $A^* = 1$  and  $Y^* = 1$

- E.g. 5%, 10%, 15%, ...
- Implement proposed estimators at each share

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
	$Y_2^*$		$A_2^*$	$X_2$	0
	$Y_3^*$		$A_3^*$	$X_3$	0
	$Y_4^*$		$A_4^*$	$X_4$	0
	$Y_5^*$		$A_5^*$	$X_5$	0
	$Y_6^*$		$A_6^*$	$X_6$	0
	$Y_7^*$		$A_7^*$	$X_7$	0
	$Y_8^*$		$A_8^*$	$X_8$	0
	$Y_9^*$		$A_9^*$	$X_9$	0
	$Y_{10}^*$		$A_{10}^*$	$X_{10}$	0

# VCCC application

“Reveal” increasingly larger shares of the validated data  $\mathbb{P}(R = 1)$ , over-sampling those with  $A^* = 1$  and  $Y^* = 1$

- E.g. 5%, 10%, 15%, ...
- Implement proposed estimators at each share

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	1
	$Y_3^*$		$A_3^*$	$X_3$	0
	$Y_4^*$		$A_4^*$	$X_4$	0
	$Y_5^*$		$A_5^*$	$X_5$	0
	$Y_6^*$		$A_6^*$	$X_6$	0
	$Y_7^*$		$A_7^*$	$X_7$	0
	$Y_8^*$		$A_8^*$	$X_8$	0
	$Y_9^*$		$A_9^*$	$X_9$	0
	$Y_{10}^*$		$A_{10}^*$	$X_{10}$	0

# VCCC application

“Reveal” increasingly larger shares of the validated data  $\mathbb{P}(R = 1)$ , over-sampling those with  $A^* = 1$  and  $Y^* = 1$

- E.g. 5%, 10%, 15%, ...
- Implement proposed estimators at each share

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	1
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	1
	$Y_4^*$		$A_4^*$	$X_4$	0
	$Y_5^*$		$A_5^*$	$X_5$	0
	$Y_6^*$		$A_6^*$	$X_6$	0
	$Y_7^*$		$A_7^*$	$X_7$	0
	$Y_8^*$		$A_8^*$	$X_8$	0
	$Y_9^*$		$A_9^*$	$X_9$	0
	$Y_{10}^*$		$A_{10}^*$	$X_{10}$	0

# VCCC application

“Reveal” increasingly larger shares of the validated data  $\mathbb{P}(R = 1)$ , over-sampling those with  $A^* = 1$  and  $Y^* = 1$

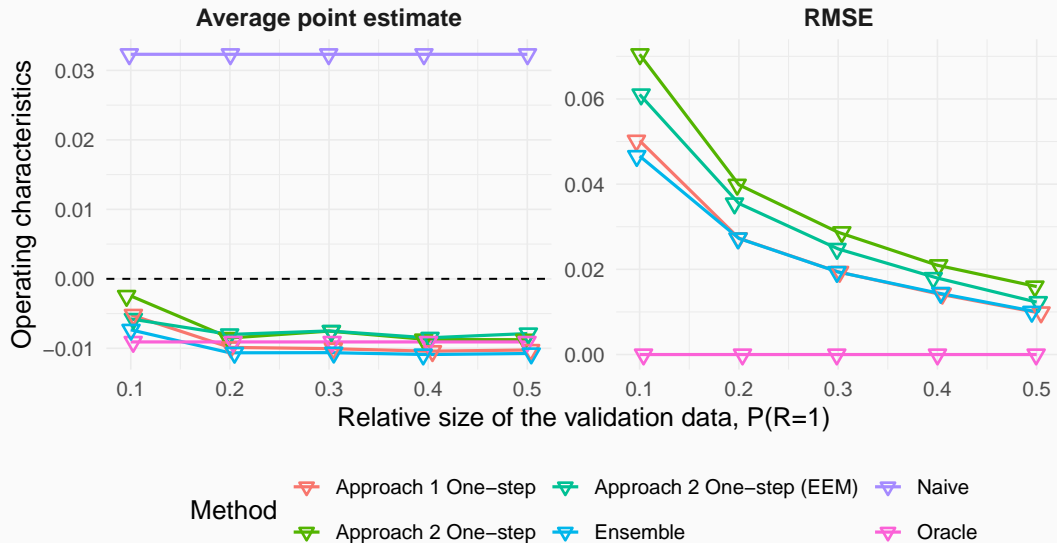
- E.g. 5%, 10%, 15%, ...
- Implement proposed estimators at each share

$Y$	$Y^*$	$A$	$A^*$	$X$	$R$
$Y_1$	$Y_1^*$	$A_1$	$A_1^*$	$X_1$	1
$Y_2$	$Y_2^*$	$A_2$	$A_2^*$	$X_2$	1
$Y_3$	$Y_3^*$	$A_3$	$A_3^*$	$X_3$	1
$Y_4$	$Y_4^*$	$A_4$	$A_4^*$	$X_4$	1
	$Y_5^*$		$A_5^*$	$X_5$	0
	$Y_6^*$		$A_6^*$	$X_6$	0
	$Y_7^*$		$A_7^*$	$X_7$	0
	$Y_8^*$		$A_8^*$	$X_8$	0
	$Y_9^*$		$A_9^*$	$X_9$	0
	$Y_{10}^*$		$A_{10}^*$	$X_{10}$	0



## Operating characteristics: 3-year ADE risk

Treatment: ART initiation within one month of first visit



# Roadmap

---

Background

Methods

Data Application

Discussion

Our work addresses the following high-level question:

How do we estimate causal effects nonparametrically when **(1)** we have **error-prone** outcome + treatment measurements, **(2)** we have **gold-standard** treatment + outcome measurements for a *subset* of our data, where **(3)** that subset was collected according to a sampling rule that depends on the initially observed data

We've presented plug-in + **debiased** one-step estimators that accommodate these sampling schemes (and general two-phase sampling schemes). Currently working on

- Further studying approaches that improve the finite-sample behavior of the one-step estimators
- Development of R package implementing Approach 2 for general missing data patterns

Thank you!

Working paper coming soon!

## References

van der Laan, M. J. and Robins, J. M. (2003). *Unified methods for censored longitudinal data and causality*. Springer.

## Appendix

# Identification

First, let  $\mathbf{W} = (Y^*, A^*)$ . Then, we have

$$\begin{aligned}\mathbb{E}(Y(a)) &= \mathbb{E}_{\mathbf{X}} \mathbb{E}(Y(a) | \mathbf{X}) \\ &= \mathbb{E}_{\mathbf{W}(a) | \mathbf{X}} [\mathbb{E}_{\mathbf{X}} \mathbb{E}(Y(a) | \mathbf{X}, \mathbf{W}(a))] \\ &= \mathbb{E}_{\mathbf{W}(a) | \mathbf{X}} [\mathbb{E}_{\mathbf{X}} \mathbb{E}(Y(a) | \mathbf{X}, \mathbf{W}(a), A = a, R = 1)] \\ &= \mathbb{E}_{\mathbf{W} | \mathbf{X}, A} [\mathbb{E}_{\mathbf{X}} \mathbb{E}(Y | \mathbf{X}, \mathbf{W}, A = a, R = 1)]\end{aligned}$$

The problem: the density  $p(w|x, a)$  is unidentified

- We only see  $A$  in the validation data, which is **not** independent of  $\mathbf{W}$  so we can't just condition on  $R = 1$

Notice

$$\mathbb{E}(Y(a)) = \sum_w \sum_x \sum_y y \cdot p(y|w, x, a, r = 1) \cdot p(w|x, a) \cdot p(x)$$



# Identification

Notice

$$\begin{aligned}\mathbb{E}(Y(a)) &= \sum_w \sum_x \sum_y y \cdot p(y|w, x, a, r = 1) \cdot p(w|x, a) \cdot p(x) \\ &= \sum_w \sum_x \sum_y y \cdot p(y|w, x, a, r = 1) \cdot \frac{p(a|x, w)p(w|x)p(x)}{p(a|x)p(x)} \cdot p(x)\end{aligned}$$

# Identification

Notice

$$\begin{aligned}\mathbb{E}(Y(a)) &= \sum_w \sum_x \sum_y y \cdot p(y|w, x, a, r = 1) \cdot p(w|x, a) \cdot p(x) \\ &= \sum_w \sum_x \sum_y y \cdot p(y|w, x, a, r = 1) \cdot \frac{p(a|x, w)p(w|x)p(x)}{p(a|x)p(x)} \cdot p(x) \\ &= \sum_w \sum_x \sum_y y \cdot p(y|w, x, a, r = 1) \cdot \frac{p(a|x, w, r = 1)p(w|x)}{\sum_{w'} p(a|w', x)p(w'|x)} \cdot p(x)\end{aligned}$$

# Identification

Notice

$$\begin{aligned}\mathbb{E}(Y(a)) &= \sum_w \sum_x \sum_y y \cdot p(y|w, x, a, r = 1) \cdot p(w|x, a) \cdot p(x) \\ &= \sum_w \sum_x \sum_y y \cdot p(y|w, x, a, r = 1) \cdot \frac{p(a|x, w)p(w|x)p(x)}{p(a|x)p(x)} \cdot p(x) \\ &= \sum_w \sum_x \sum_y y \cdot p(y|w, x, a, r = 1) \cdot \frac{p(a|x, w, r = 1)p(w|x)}{\sum_{w'} p(a|w', x)p(w'|x)} \cdot p(x)\end{aligned}$$

Key idea: Use Bayes rule to re-express un-identified density in terms of identifiable ones

- Then, overall expression is identified – just need to define terms and “collapse” the above expectations

## Plug-in implementation

$Y_i$	$Y_i^*$	$A_i$	$A_i^*$	$X_i$	$R_i$	$\hat{\lambda}_a$	$\hat{\mu}_a$	$\hat{\pi}_a$	$\hat{\eta}_a$
1	0	0	1	3.2	1	NA	NA	NA	NA
0	1	1	1	2.1	1	NA	NA	NA	NA
1	1	0	0	0.3	1	NA	NA	NA	NA
1	0	0	1	1.6	0	NA	NA	NA	NA
0	1	0	1	4.8	0	NA	NA	NA	NA
1	1	0	0	0.9	0	NA	NA	NA	NA

Step 1: use validation data to fit  $\hat{\lambda}_a$  and  $\hat{\mu}_a$

## Plug-in implementation

$Y_i$	$Y_i^*$	$A_i$	$A_i^*$	$\mathbf{X}_i$	$R_i$	$\hat{\lambda}_a$	$\hat{\mu}_a$	$\hat{\pi}_a$	$\hat{\eta}_a$
1	0	0	1	3.2	1	0.2	0.72	NA	NA
0	1	1	1	2.1	1	0.43	0.18	NA	NA
1	1	0	0	0.3	1	0.64	0.77	NA	NA
1	0	0	1	1.6	0	0.03	0.55	NA	NA
0	1	0	1	4.8	0	0.81	0.11	NA	NA
1	1	0	0	0.9	0	0.39	0.83	NA	NA

Step 1: use validation data to fit  $\hat{\lambda}_a$  and  $\hat{\mu}_a$

## Plug-in implementation

$Y_i$	$Y_i^*$	$A_i$	$A_i^*$	$\mathbf{X}_i$	$R_i$	$\hat{\lambda}_a$	$\hat{\mu}_a$	$\hat{\pi}_a$	$\hat{\eta}_a$
1	0	0	1	3.2	1	0.2	0.72	NA	NA
0	1	1	1	2.1	1	0.43	0.18	NA	NA
1	1	0	0	0.3	1	0.64	0.77	NA	NA
1	0	0	1	1.6	0	0.03	0.55	NA	NA
0	1	0	1	4.8	0	0.81	0.11	NA	NA
1	1	0	0	0.9	0	0.39	0.83	NA	NA

Step 2: regress  $\hat{\lambda}_a$  and  $\hat{\mu}_a$  on  $\mathbf{X}$  and  $A^*$  to yield  $\hat{\pi}_a$  and  $\hat{\eta}_a$

## Plug-in implementation

$Y_i$	$Y_i^*$	$A_i$	$A_i^*$	$\mathbf{X}_i$	$R_i$	$\hat{\lambda}_a$	$\hat{\mu}_a$	$\hat{\pi}_a$	$\hat{\eta}_a$
1	0	0	1	3.2	1	0.2	0.72	0.77	0.23
0	1	1	1	2.1	1	0.43	0.18	0.41	0.26
1	1	0	0	0.3	1	0.64	0.77	0.59	0.80
1	0	0	1	1.6	0	0.03	0.55	0.08	0.53
0	1	0	1	4.8	0	0.81	0.11	0.72	0.18
1	1	0	0	0.9	0	0.39	0.83	0.38	0.79

Step 2: regress  $\hat{\lambda}_a$  and  $\hat{\mu}_a$  on  $\mathbf{X}$  and  $A^*$  to yield  $\hat{\pi}_a$  and  $\hat{\eta}_a$

## Plug-in implementation

$Y_i$	$Y_i^*$	$A_i$	$A_i^*$	$\mathbf{X}_i$	$R_i$	$\hat{\lambda}_a$	$\hat{\mu}_a$	$\hat{\pi}_a$	$\hat{\eta}_a$
1	0	0	1	3.2	1	0.2	0.72	0.77	0.23
0	1	1	1	2.1	1	0.43	0.18	0.41	0.26
1	1	0	0	0.3	1	0.64	0.77	0.59	0.80
1	0	0	1	1.6	0	0.03	0.55	0.08	0.53
0	1	0	1	4.8	0	0.81	0.11	0.72	0.18
1	1	0	0	0.9	0	0.39	0.83	0.38	0.79

Step 3: construct  $\hat{\psi}_a^{\text{PI}} = \frac{1}{n} \sum_{i=1}^n \frac{\hat{\eta}_a(\mathbf{X}_i, A_i^*)}{\hat{\pi}_a(\mathbf{X}_i, A_i^*)}$



## Equivalence of Approach 1 and Approach 2 EICs

$$\begin{aligned}\phi_a^{\text{obs}}(\mathbf{O}) &= \frac{R}{\mathbb{P}(R = 1|\mathbf{Z})} \chi_a^{\text{full}}(\mathbf{O}) \\ &\quad - \left( \frac{R}{\mathbb{P}(R = 1|\mathbf{Z})} - 1 \right) \varphi_a(\mathbf{Z})\end{aligned}$$

## Equivalence of Approach 1 and Approach 2 EICs

$$\begin{aligned}\phi_a^{\text{obs}}(\mathbf{O}) &= \frac{R}{\mathbb{P}(R=1|\mathbf{Z})} \left[ \frac{I(A=a)}{e_a(\mathbf{X})} (Y - m_a(\mathbf{X})) + m_a(\mathbf{X}) - \psi_a \right] \\ &\quad - \left( \frac{R}{\mathbb{P}(R=1|\mathbf{Z})} - 1 \right) \varphi_a(\mathbf{Z})\end{aligned}$$

## Equivalence of Approach 1 and Approach 2 EICs

$$\begin{aligned}\phi_a^{\text{obs}}(\mathbf{O}) &= \frac{R}{\mathbb{P}(R=1|\mathbf{Z})} \left[ \frac{I(A=a)}{\pi_a(\mathbf{X})} \left( Y - \frac{\eta_a(\mathbf{X})}{\pi_a(\mathbf{X})} \right) + \frac{\eta_a(\mathbf{X})}{\pi_a(\mathbf{X})} - \psi_a \right] \\ &\quad - \left( \frac{R}{\mathbb{P}(R=1|\mathbf{Z})} - 1 \right) \varphi_a(\mathbf{Z})\end{aligned}$$

## Equivalence of Approach 1 and Approach 2 EICs

$$\begin{aligned}\phi_a^{\text{obs}}(\mathbf{O}) &= \frac{R}{\mathbb{P}(R=1|\mathbf{Z})} \left[ \frac{I(A=a)}{\pi_a(\mathbf{X})} \left( Y - \frac{\eta_a(\mathbf{X})}{\pi_a(\mathbf{X})} \right) + \frac{\eta_a(\mathbf{X})}{\pi_a(\mathbf{X})} - \psi_a \right] \\ &\quad - \left( \frac{R}{\mathbb{P}(R=1|\mathbf{Z})} - 1 \right) \left( \frac{\lambda_a(\mathbf{Z})\mu_a(\mathbf{Z})}{\pi_a(\mathbf{X})} - \frac{\lambda_a(\mathbf{Z})\eta_a(\mathbf{X})}{\pi_a(\mathbf{X})^2} + \frac{\eta_a(\mathbf{X})}{\pi_a(\mathbf{X})} - \psi_a \right)\end{aligned}$$