# Estimating the target

### Estimating the target



- one estimator is not more causal than another.
- different estimators are based on different nuisance parameters and have different statistical properties (bias/variance).

# G-formula versus IP-weighting

- G-formula 1. Estimate nuisance parameters  $f(a,x) = \mathbb{E}[Y \mid A = a, X = x]$  and the average over the marginal distribution  $\mu_X$  of X
  - 2. Plug in to estimate the ATE:

$$\hat{\psi}_n^{\text{g-formula}} = \tilde{\Psi}(\hat{f}_n, \hat{\mu}_X) = \int_{\mathbb{R}^d} (\hat{f}_n(1, x) - \hat{f}_n(0, x)) d\hat{\mu}_X(x)$$

- IP-weighting 1. Estimate nuisance parameters  $\pi(a \mid x) = P(A = a \mid X = x)$  and the average over the distribution P of O
  - 2. Plug in to estimate the ATE:

$$\hat{\psi}_{n}^{\mathsf{ipw}} = \tilde{\Psi}_{\mathsf{ipw}}(\hat{\pi}_{n}, \hat{P}_{n}) = \int_{\mathbb{R}^{d}} \left( \frac{\mathsf{a} \mathsf{y}}{\hat{\pi}_{n}(\mathsf{a} \mid \mathsf{x})} - \frac{(1-\mathsf{a})\mathsf{y}}{\hat{\pi}_{n}(\mathsf{a} \mid \mathsf{x})} \right) d\hat{P}_{n}(\mathsf{x})$$

### One-step estimation

- One-step 1. Estimate nuisance parameters  $f(a,x) = \mathbb{E}[Y \mid A = a, X = x], \ \pi(a \mid x) = \mathbb{E}[A \mid X = x]$ 
  - and the average over the distribution P of O
  - 2. Plug in to estimate the ATE:

$$\hat{\psi}_{n}^{\text{one}} = \tilde{\Psi}_{\text{one}}(\hat{f}_{n}, \hat{\pi}_{n}, \hat{P}_{n}) = \int_{\mathbb{R}^{d}} \sum_{a=0,1} \sum_{y=0,1} \left\{ \left( \frac{a}{\hat{\pi}_{n}(a \mid x)} - \frac{1-a}{\hat{\pi}_{n}(a \mid x)} \right) (y - \hat{f}_{n}(a, x)) + \hat{f}_{n}(1, x) - \hat{f}_{n}(0, x) \right\} d\hat{P}_{n}(o)$$

# G-formula versus IP-weighting

Estimation of the averages over  $\mu_X$  and P is straightforward using the empirical average over the observed data.

This yields:

Properties of the different estimators:

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- G-formula estimator requires estimator  $\hat{f}_n$  for conditional expectation f.
  - consistent if  $\hat{f}_n$  is consistent.

IP-weighted estimator requires estimator  $\hat{\pi}_n$  for the propensity score  $\pi$ .

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- the one-step estimator and the TMLE estimator share the same large-sample properties.
- one thing they have in common is an additional bias correction leads to consistency if either  $\hat{f}_n$  or  $\hat{\pi}_n$  is consistent (commonly known as double robustness).

### SMALL EXERCISE:

By the law of large numbers, the one-step estimator converges in probability to:

$$\mathbb{E}_{P_{\mathbf{0}}}\left[\left(\frac{A}{\pi(A\mid X)} - \frac{1-A}{\pi(A\mid X)}\right)\left(Y - f(A,X)\right) + f(1,X) - f(0,X)\right] \tag{1}$$

where  $(f,\pi)$  denotes the limit of  $(\hat{f}_n,\hat{\pi}_n)$ . Compute the right hand side of (1) when

- 1.  $f = f_0$  (i.e., the outcome regression is consistently estimated), and
- 2.  $\pi = \pi_0$  (i.e., the propensity score is consistently estimated).

TMLE estimator requires estimators  $\hat{f}_n$  and  $\hat{\pi}_n$  for conditional expectation f and propensity score  $\pi$ .

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Some other things we will see/explore:

• if  $\hat{\pi}_n$  is modeled correctly, the TMLE estimator will have lower variance than the IP-weighted estimator.

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Some other things we will see/explore:

- if  $\hat{\pi}_n$  is modeled correctly, the TMLE estimator will have lower variance than the IP-weighted estimator.
- the TMLE estimator may have larger variance than the g-formula estimator based on a correctly modeled  $\hat{f}_n$  (but it gives protection against the case that it is not).

Can't we just construct a good g-formula estimator???

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- ▶ a logistic regression great if correctly specified, but horrible if not.
- a random forest properly tuned?

Predictive performance of an estimator can be measured in terms of some distance<sup>1</sup> between:

1) the observed outcome:  $Y_i$ 

2) and the predicted conditional expectation:  $\hat{f}_n(A_i, X_i)$ 

<sup>&</sup>lt;sup>1</sup>Measured in terms of a *loss function*.

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- 1) the observed outcome: Y
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One example of a loss function  $\mathcal{L}(f)(O)$  is the negative log-likelihood loss:

$$\mathcal{L}(\hat{f}_n)(Y_i,A_i,X_i) = -\big(Y_i\log(\hat{f}_n(A_i,X_i)) + (1-Y_i)\log(1-\hat{f}_n(A_i,X_i))\big).$$

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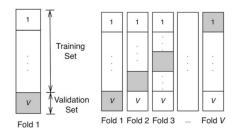
One example of a loss function  $\mathcal{L}(f)(O)$  is the negative log-likelihood loss:

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The estimator  $\hat{f}_n$  closest to the true  $f_0$  minimizes the risk:

$$\mathbb{E}_{P_0}[\mathcal{L}(\hat{f}_n)(Y_i,A_i,X_i)].$$

<sup>&</sup>lt;sup>1</sup>Measured in terms of a *loss function*.



The risk can be estimated in a cross-validation scheme.<sup>a</sup>

### I.e., for each sample split:

- 1. Each model is created and fitted on the training data:  $\hat{f}_n^{\text{train}}$ .
- 2. The quality of the model is checked on the validation data
  - Average of  $\mathcal{L}(\hat{f}_n^{\mathsf{train}})(O_i)$  in the validation sample.

<sup>&</sup>lt;sup>a</sup>To measure performance on independent data.

### Simulated example

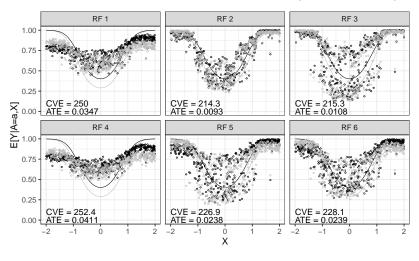
- $X \sim \text{Unif}(-2,2)$
- $X_1^{\text{noise}}, \dots, X_5^{\text{noise}} \sim N(0, 1)$
- ▶  $A \in \{0,1\}$  with distribution given X given by:

$$\mathsf{logit}\,\mathbb{E}[A\,|\,X] = \gamma_0 + \gamma_X^{\mathsf{T}}X$$

▶  $Y \in \{0,1\}$  with distribution given X and A given by:

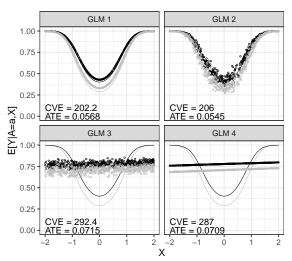
$$\operatorname{logit} \mathbb{E}[Y \mid A, X] = \beta_0 + \beta_A A + \beta_X^{\mathsf{T}} X^2$$

RF fitted with different values of tuning parameters (nodesize, mtry):



### Different GLM models

GLM models fitted with different covariates and functional form of covariates:



This is all about constructing a good estimator for the conditional expectation f.

This does not necessarily translate into a good estimator for the target  $\tilde{\Psi}(f, \mu_X)$ .

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This does not necessarily translate into a good estimator for the target  $\tilde{\Psi}(f, \mu_X)$ .

TMLE is all about constructing a g-formula estimator which is a good estimator for *the target*.

Fix randomness:

```
set.seed(5)
```

Fix a sample size:

Generate covariate  $X \in [-2, 2]$ :

Generate binary treatment decision A:

```
A <- rbinom(n, 1, prob=plogis(-0.25 + 1.2*X))
```

(corresponding to logit  $\mathbb{E}[A \mid X] = \gamma_0 + \gamma_X X$ )

Generate binary outcome Y according to

$$\mathsf{logit}\,\mathbb{E}[\,Y\mid A,X\,] = \beta_0 + \beta_A A + \beta_X X^2$$

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### First generate counterfactuals:

```
Y1 <- rbinom(n, 1, prob=plogis(-0.9 + 1.9*X^2 + 0.5*1))
Y0 <- rbinom(n, 1, prob=plogis(-0.9 + 1.9*X^2 + 0.5*0))
```

Generate binary outcome Y according to

$$\operatorname{logit} \mathbb{E}[Y \mid A, X] = \beta_0 + \beta_A A + \beta_X X^2$$

First generate counterfactuals:

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Y0 <- rbinom(n, 1, prob=plogis(-0.9 + 1.9*X^2 + 0.5*0))
```

We only observe the counterfactual outcome corresponding to the observed treatment level:

```
Y <- A*Y1 + (1-A)*Y0
```

### Observed data:

```
X A Y
1: -1.1991422 0 0
2: 0.7408744 1 0
3: 1.6675031 1 1
4: -0.8624022 0 1
5: -1.5813995 0 1
---
496: -0.3978523 1 0
497: -1.5069379 0 1
498: 1.8340120 1 1
499: 0.6349484 1 1
500: -0.5214807 0 1
```

### Observed data:

# X A Y 1: -1.1991422 0 0 2: 0.7408744 1 0 3: 1.6675031 1 1 4: -0.8624022 0 1 5: -1.5813995 0 1 -- 496: -0.3978523 1 0 497: -1.5069379 0 1 498: 1.8340120 1 1 499: 0.6349484 1 1 500: -0.5214807 0 1

### Counterfactual data:

	Х	Y1	YO
1:	-1.1991422	0	1
2:	0.7408744	1	0
3:	1.6675031	1	1
4:	-0.8624022	0	1
5:	-1.5813995	1	1
496:	-0.3978523	0	1
497:	-1.5069379	0	1
498:	1.8340120	1	1
499:	0.6349484	0	0
500:	-0.5214807	0	0

Simulating many observations of counterfactuals allows us to approximate the true ATE:

```
X <- runif(1e6, -2, 2)
Y1 <- rbinom(1e6, 1, prob=plogis(-0.9 + 1.9*X^2 + 0.5*1))
Y0 <- rbinom(1e6, 1, prob=plogis(-0.9 + 1.9*X^2 + 0.5*0))</pre>
```

The true ATE is then approximately:

```
(true.ate <- mean(Y1 - Y0))
```

since ATE = 
$$\mathbb{E}_{P_0}[Y^1] - \mathbb{E}_{P_0}[Y^0]$$
.

Fit correctly specified parametric model:

```
fit.glm <- glm(Y~A+X.squared, data=dt[, X.squared:=X^2],
    family=binomial)</pre>
```

Use model to estimate f(1, X) for all subjects:

And similarly f(0,X) for all subjects:

Then we can estimate the ATE by:

```
(fit.glm <- dt[, mean(pred.glm.A1-pred.glm.A0)])</pre>
```

Using a random forest (no tuning):

Using a misspecified parametric model:

```
fit.glm.mis <- glm(Y~A+X, data=dt, family=binomial)
dt[, pred.glm.mis.A1:=predict(fit.glm.mis, type="response",
    newdata=copy(dt)[, A:=1])]
dt[, pred.glm.mis.A0:=predict(fit.glm.mis, type="response",
    newdata=copy(dt)[, A:=0])]
(fit.glm.mis <- dt[, mean(pred.glm.mis.A1-pred.glm.mis.A0)])</pre>
```

We can investigate the properties of different estimators —

- We know the true value of ATE:  $\psi_0 \approx 0.0702$
- ▶ We have generated the outcome Y according to

$$\operatorname{logit} \mathbb{E}[Y \mid A, X] = \beta_0 + \beta_A A + \beta_X X^2$$

We have generated the treatment A according to

$$\operatorname{logit} \mathbb{E}[A \mid X] = \gamma_0 + \gamma_X X$$

If we repeat the experiment of drawing n observations we would every time end up with a different realization of the particular estimator.

#### Different estimators

G-formula estimator Using an estimator  $\hat{f}_n$  for  $f(a,X) = \mathbb{E}[Y \mid A = a, X]$ , estimate the ATE by:

$$\hat{\psi}_n^{\text{g-formula}} = \frac{1}{n} \sum_{i=1}^n \left\{ \hat{f}_n(1, X_i) - \hat{f}_n(0, X_i) \right\}$$

Inverse probability weighted estimator Using an estimator  $\hat{\pi}_n$  for  $\pi(a \mid X) = P(A = a \mid X)$ , estimate the ATE by:

$$\hat{\psi}_{n}^{\text{ipw}} = \frac{1}{n} \sum_{i=1}^{n} \left\{ \frac{A_{i} Y_{i}}{\hat{\pi}_{n}(A_{i} \mid X_{i})} - \frac{(1 - A_{i}) Y_{i}}{\hat{\pi}_{n}(A_{i} \mid X_{i})} \right\}$$

TMLE estimator Update the estimator  $\hat{f}_n \mapsto \hat{f}_n^*$  in a "targeted way" using the information from the estimator  $\hat{\pi}_n$ , then estimate the ATE by:

$$\hat{\psi}_{n}^{\text{tmle}} = \frac{1}{n} \sum_{i=1}^{n} \left\{ \hat{f}_{n}^{*}(1, X_{i}) - \hat{f}_{n}^{*}(0, X_{i}) \right\}$$

Today we will just (more or less blindly) use software to estimate TMLE.

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```
library(tmle)
```

- $Y \in \mathbb{R} \text{ or } Y \in \{0,1\}$
- ▶  $A \in \{0, 1\}$
- X a vector, matrix or a data frame

- gform
  - ightharpoonup optional regression formula for the propensity score  $\pi$
  - ▶ on the form A~X1+X2
  - (overrides call to SuperLearner)
- Qform
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- gbound
  - truncation of predicted probabilities of treatment

#### On a sidenote — tomorrow

- ▶ Q.SL.library
  - optional vector of prediction algorithms to use for SuperLearner in initial estimation of f
- ▶ g.SL.library
  - optional vector of prediction algorithms to use for SuperLearner in initial estimation of  $\pi$
- Q.discreteSL
  - if TRUE, a discrete super learner is used (rather than ensemble)
  - default is FALSE
- g.discreteSL
  - if TRUE, a discrete super learner is used (rather than ensemble)
  - default is FALSE

**Note:** The discrete super learner simply picks an algorithm from its library by minimizing the cross-validated empirical risk with respect a loss function.

What were the estimated IP weights?

```
summary(fit.tmle$g$g1W)
```

```
Min. 1st Qu. Median Mean 3rd Qu. Max. 0.04751 0.19441 0.49405 0.49400 0.79710 0.94109
```

Note that weights close to 0 or to 1 would indicate positivity issues.

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Note that weights close to 0 or to 1 would indicate positivity issues.

What truncation level was used?

```
fit.tmle$gbound
```

[1] 0.03598084 1.00000000

I.e., no weights were truncated.

### Explorations based on simulated data

As part of the exercise we want to explore —

- 1. Comparing g-formula estimators for different estimators for f; either different logistic regressions or different machine learning algorithms.
- 2. Properties of the g-formula estimator and the IP-weighted estimator, compared to the TMLE estimator.
- 3. Double robustness: Misspecification of the outcome regression (f).

The exercise is described in detail in: day1-practical1.pdf.

Some relevant concepts linking what we have seen by now to the rest of today + tomorrow.

A very desirable property —

 $<sup>^{2}</sup>o_{P}(1)$  denotes a sequence which is converges to zero in probability.

The empirical measure  $\mathbb{P}_n$  of the sample  $O_1, \ldots, O_n$ :  $\mathbb{P}_n h = \int h d\mathbb{P}_n = \frac{1}{n} \sum_{i=1}^n h(O_i).$ 

#### A very desirable property —

An estimator  $\hat{\psi}_n$  is  $\sqrt{n}$ -consistent and asymptotically linear with influence function  $\phi(P_0)(O)$  if <sup>2</sup>

$$\sqrt{n}(\hat{\psi}_n - \psi_0) = \sqrt{n} \mathbb{P}_n \phi(P_0) + o_P(1),$$

where  $\mathbb{E}_{P_0}[\phi(P_0)(O)] = 0$  and  $\mathbb{E}_{P_0}[\{\phi(P_0)(O)\}^2] < \infty$ .

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$$\sqrt{n}(\hat{\psi}_n - \psi_0) = \sqrt{n} \mathbb{P}_n \phi(P_0) + o_P(1),$$

where 
$$\mathbb{E}_{P_0}[\phi(P_0)(O)] = 0$$
 and  $\mathbb{E}_{P_0}[\{\phi(P_0)(O)\}^2] < \infty$ .

Then CLT + Slutsky implies:

$$\hat{\psi}_n \stackrel{as}{\sim} N(\Psi(P_0), \text{Var}(\phi(P_0))/n).$$

The estimator behaves asymptotically as an average of the influence function.

 $<sup>^{2}</sup>o_{P}(1)$  denotes a sequence which is converges to zero in probability.

**Simple example:** Estimator for the mean  $\psi_0 = \mathbb{E}[X]$ :

$$\hat{\psi}_{n,0} = \frac{1}{n} \sum_{i=1}^{n} X_i$$

Then

$$\sqrt{n}(\hat{\psi}_n - \psi_0) = \sqrt{n} \frac{1}{n} \sum_{i=1}^n \underbrace{(X_i - \psi_0)}_{=\phi(P_0)(O_i)} = \sqrt{n} \mathbb{P}_n \phi(P_0)$$

 $\hat{\psi}_{n,0}$  is linear and thus asymptotically linear.

**Simple example:** Estimator for the mean  $\psi_0 = \mathbb{E}[X]$ :

$$\hat{\psi}_{n,1} = \frac{1}{n} \sum_{i=1}^{n} X_i + \frac{1}{n}$$

Then

$$\sqrt{n}(\hat{\psi}_{n} - \psi_{0}) = \sqrt{n} \frac{1}{n} \sum_{i=1}^{n} \underbrace{(X_{i} - \psi_{0})}_{=\phi(P_{0})(O_{i})} + \frac{\sqrt{n}}{n} = \sqrt{n} \mathbb{P}_{n} \phi(P_{0}) + \underbrace{\frac{1}{\sqrt{n}}}_{=o_{P}(1)}$$

 $\hat{\psi}_{\mathit{n},1}$  is asymptotically linear.

**Simple example:** Estimator for the mean  $\psi_0 = \mathbb{E}[X]$ :

$$\hat{\psi}_{n,2} = \frac{1}{n} \sum_{i=1}^{n} X_i + \frac{1}{n^{1/2 + 0.1}}$$

Then

$$\sqrt{n}(\hat{\psi}_n - \psi_0) = \sqrt{n} \frac{1}{n} \sum_{i=1}^n \underbrace{(X_i - \psi_0)}_{=\phi(P_0)(O_i)} + \frac{\sqrt{n}}{n^{1/2 + 0.1}} = \sqrt{n} \mathbb{P}_n \phi(P_0) + \underbrace{\frac{1}{n^{0.1}}}_{=o_P(1)}$$

 $\hat{\psi}_{\it n,2}$  is asymptotically linear.

**Simple example:** Estimator for the mean  $\psi_0 = \mathbb{E}[X]$ :

$$\hat{\psi}_{n,3} = \frac{1}{n} \sum_{i=1}^{n} X_i + \frac{1}{n^{1/2 - 0.1}}$$

Then

$$\sqrt{n}(\hat{\psi}_n - \psi_0) = \sqrt{n} \frac{1}{n} \sum_{i=1}^n \underbrace{(X_i - \psi_0)}_{=\phi(P_0)(O_i)} + \frac{\sqrt{n}}{n^{1/2 - 0.1}} = \sqrt{n} \mathbb{P}_n \phi(P_0) + \underbrace{n^{0.1}}_{\to \infty}$$

 $\hat{\psi}_{\textit{n},3}$  is  $\mathbf{not}$  asymptotically linear.

An estimator  $\hat{\psi}_n$  has rate of convergence  $r_n \to \infty$  if <sup>3</sup>

$$r_n(\hat{\psi}_n - \psi_0) = O_P(1)$$
, i.e.,  $\hat{\psi}_n - \psi_0 = O_P(1/r_n)$ .

The convergence rate  $r_n$  tells us how fast  $\hat{\psi}_n$  centers around  $\psi_0$ , with the difference  $\hat{\psi}_n - \psi_0$  behaving like  $1/r_n$ .

- One wants negligible bias such as to obtain reliable confidence intervals for  $\psi_0$ .
- ▶ The bias of an asymptotically linear estimator converges to zero at a rate faster the  $1/\sqrt{n}$ .

Data-adaptive machine learning estimators rarely achieve this rate.

 $<sup>^3</sup>O_P(1)$  denotes a sequence which is bounded in probability.

$$\sqrt{n}\hat{\psi}_{n,1} = \sqrt{n}\underbrace{\frac{1}{n}\sum_{i=1}^{n}X_{i}}_{\stackrel{P}{\rightarrow}\psi_{0}} + \underbrace{\frac{\sqrt{n}}{n}}_{\stackrel{N}{\rightarrow}0}, \quad \text{i.e.,} \quad \sqrt{n}(\hat{\psi}_{n,1} - \psi_{0}) = o_{P}(1).$$

$$\sqrt{n}\hat{\psi}_{n,2} = \sqrt{n}\underbrace{\frac{1}{n}\sum_{i=1}^{n}X_{i}}_{\stackrel{P}{\to}\psi_{0}} + \underbrace{\frac{\sqrt{n}}{n^{1/2+0.1}}}_{\stackrel{N}{\to}0}, \text{ i.e., } \sqrt{n}(\hat{\psi}_{n,3} - \psi_{0}) = o_{P}(1).$$

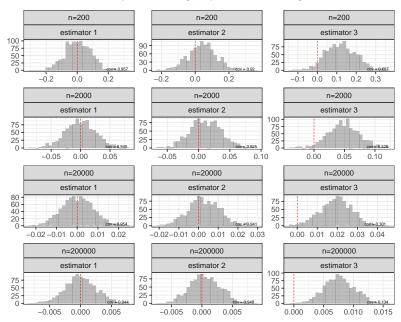
$$\sqrt{n}\hat{\psi}_{n,3} = \sqrt{n}\underbrace{\frac{1}{n}\sum_{i=1}^{n}X_{i}}_{\stackrel{P}{\rightarrow}\psi_{0}} + \underbrace{\frac{\sqrt{n}}{n^{1/2-0.1}}}_{\stackrel{\rightarrow}{\rightarrow}\infty}, \quad \text{i.e.,} \quad \sqrt{n}(\hat{\psi}_{n,3} - \psi_{0}) \stackrel{P}{\rightarrow} \infty.$$

$$\sqrt{n}\hat{\psi}_{n,1} = \sqrt{n}\underbrace{\frac{1}{n}\sum_{i=1}^{n}X_{i}}_{P_{\psi_{0}}} + \underbrace{\frac{\sqrt{n}}{n}}_{N}, \quad \text{i.e.,} \quad \sqrt{n}(\hat{\psi}_{n,1} - \psi_{0}) = o_{P}(1).$$

$$\sqrt{n}\hat{\psi}_{n,2} = \sqrt{n}\underbrace{\frac{1}{n}\sum_{i=1}^{n}X_{i}}_{P_{\rightarrow \psi_{0}}} + \underbrace{\frac{\sqrt{n}}{n^{1/2+0.1}}}_{0}, \text{ i.e., } \sqrt{n}(\hat{\psi}_{n,3} - \psi_{0}) = o_{P}(1).$$

$$\sqrt{n}\hat{\psi}_{n,3} = \sqrt{n}\underbrace{\frac{1}{n}\sum_{i=1}^{n}X_{i}}_{P\rightarrow 0} + \underbrace{\frac{\sqrt{n}}{n^{1/2-0.1}}}_{\rightarrow \infty}, \quad \text{i.e.,} \quad \sqrt{n}(\hat{\psi}_{n,3} - \psi_{0}) \stackrel{P}{\rightarrow} \infty.$$

[The remainder term that determines the asymptotic bias the estimator].



A key component in constructing a  $\sqrt{n}$ -consistent and asymptotically linear estimator, even when using machine learning estimation, is the so-called the efficient influence function (also known as the canonical gradient).

A key component in constructing a  $\sqrt{n}$ -consistent and asymptotically linear estimator, even when using machine learning estimation, is the so-called the efficient influence function (also known as the canonical gradient).

- The efficient influence function provides a nonparametric lower bound for the estimation problem.
- Tells us how to do bias-correction.
- With the bias-correction, the remainder term that we need to control to have  $\sqrt{n}$ -consistency and asymptotic linearity admits a nice structure that we *can* control.

Rest of today + tomorrow (TMLE).