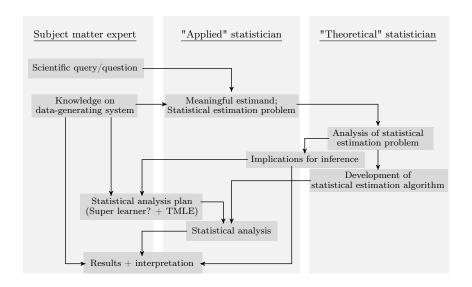
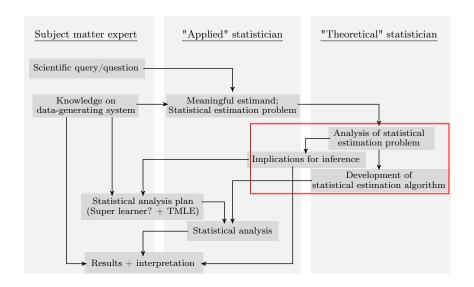
Day 1, Lecture 4

Targeted nonparametric inference

#### In this lecture, our goal is to:

- 1. Relate fundamental concepts of nonparametric efficiency theory to the construction of asymptotically linear estimators.
- Explain the particular role of the efficient influence curve in guiding the development of estimators, including the significance of second order remainders in achieving asymptotic linearity.





Key (technical) concepts we cannot avoid talking about:

- \* asympotically linear estimation
- \* efficient influence curve
- \* second-order remainders

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While TMLE can be applied without knowing about these concepts ...

- ... we cannot understand the purpose of TMLE without knowing a bit about the efficient influence curve.
- ... to understand what conditions are required for TMLE inference, we need to understand the second-order remainder.

#### Important notation:<sup>1</sup>

 $\triangleright$  For a function  $h: \mathcal{O} \to \mathbb{R}$  and distribution P

$$Ph = \mathbb{E}_{P}[h(O)] = \int hdP = \int_{\mathcal{O}} h(o)dP(o)$$

where  $\mathcal{O} = \mathbb{R}^d \times \{0,1\} \times \{0,1\}$  is the sample space of  $\mathcal{O} = (X,A,Y)$ .

 $\triangleright$  For the empirical measure  $\mathbb{P}_n$  of the sample  $O_1,\ldots,O_n$ :

$$\mathbb{P}_n h = \int h d\mathbb{P}_n = \frac{1}{n} \sum_{i=1}^n h(O_i);$$

note: the right-hand-side is really just the empirical average.

$$\triangleright X_n = o_P(1)$$
 means that  $X_n \stackrel{P}{\to} 0$ ;  $X_n = o_P(n^{-1/2})$  means that  $n^{1/2}X_n \stackrel{P}{\to} 0$ .

<sup>&</sup>lt;sup>1</sup>van der Vaart, A. W. (2000). Asymptotic statistics (Vol. 3). Cambridge university press.

A very desirable property —

<sup>&</sup>lt;sup>2</sup>recall:  $o_P(1)$  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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Then CLT + Slutsky implies:

$$\hat{\psi}_n \stackrel{\text{as}}{\sim} N(\psi_0, \text{Var}(\phi(P_0))/n).$$

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

<sup>&</sup>lt;sup>2</sup>recall:  $o_P(1)$  denotes a sequence which is converges to zero in probability.

**Simple example:** Estimator for the mean  $\psi_0 = \mathbb{E}_{P_0}[X]$ :

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

Then

$$\sqrt{n}(\hat{\psi}_{n,0} - \psi_0) = \sqrt{n} \frac{1}{n} \sum_{i=1}^{n} \underbrace{(X_i - \psi_0)}_{=\phi(P_0)(O_i)} = \sqrt{n} \mathbb{P}_n \phi(P_0)$$

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

**Simple example:** Estimator for the mean  $\psi_0 = \mathbb{E}_{P_0}[X]$ :

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

Then

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

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

**Simple example:** Estimator for the mean  $\psi_0 = \mathbb{E}_{P_0}[X]$ :

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

Then

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

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

**Simple example:** Estimator for the mean  $\psi_0 = \mathbb{E}_{P_0}[X]$ :

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

Then

$$\sqrt{n}(\hat{\psi}_{n,3} - \psi_0) = \sqrt{n} \frac{1}{n} \sum_{i=1}^{n} \underbrace{(X_i - \psi_0)}_{=\phi(P_0)(O_i)} + \frac{\sqrt{n}}{n^{1/2 - 0.1}} = \sqrt{n} \mathbb{P}_n \phi(P_0) + \underbrace{n^{0.1}}_{\to \infty}$$

 $\hat{\psi}_{n,3}$  is **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 than  $1/\sqrt{n}$ .

Data-adaptive machine learning estimators rarely achieve this rate.

 $<sup>^{3}</sup>$ recall:  $O_{P}(1)$  denotes a sequence which is bounded in probability.

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

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

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

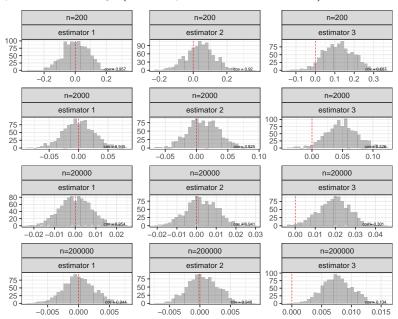
$$\sqrt{n}\hat{\psi}_{n,1} = \sqrt{n}\underbrace{\frac{1}{n}\sum_{i=1}^{n}X_{i}}_{P_{\rightarrow\psi_{0}}} + \underbrace{\frac{\sqrt{n}}{n}}_{N_{0}}, \quad \text{i.e.,} \quad \sqrt{n}(\hat{\psi}_{n,1} - \psi_{0}) = o_{P}(1).$$

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

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

# Asymptotic linearity (500 repeated simulations)



#### Repetition — our goal is $\sqrt{n}$ -consistency and asymptotic linearity.

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

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

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:

$$\hat{\psi}_n \stackrel{as}{\sim} N(\psi_0, \operatorname{Var}(\phi(P_0))/n).$$

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

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 curve.<sup>4</sup>

(Don't) be confused. We both talk about:

- the influence function for an estimator.
- ▶ the efficient influence function (curve) for the *estimation problem*.

<sup>&</sup>lt;sup>4</sup>also known as the efficient influence function, the pathwise derivative, the Neyman orthogonal score, the canonical gradient.

#### The von Mises expansion:

A sufficiently smooth  $\Psi:\mathcal{M}\to\mathbb{R}$  admits a certain distributional Taylor expansion:

$$\Psi(P) - \Psi(P') = \int \phi(P)(o)d(P - P')(o) + R_2(P, P'), \tag{1}$$

for distributions  $P, P' \in \mathcal{M}$  and a function  $\phi$  satisfying  $P\phi(P) = 0$  (mean zero) and  $P\phi(P)^2 < \infty$  (finite variance).

We will use (1) as a starting-point to start analyzing:

$$\hat{\psi}_n - \psi_0 = \Psi(\hat{P}_n) - \Psi(P_0).$$

when the model  $\mathcal{M}$  is assumed proper nonparametric, there exists \*one\* function  $\phi(P)$  fulfilling (1). This is called the efficient influence curve; we also denote it  $\phi^*(P)$ .

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  - for these situations we by the way have that  $P_0\phi(P_0)^2 \ge P_0\phi^*(P_0)^2$ . (which is why it is called the efficient influence curve).
- the efficient influence curve in nonparametric models indicates how to construct asymptotically linear (and efficient) estimators . . . the goal is to construct an asymptotically linear estimator which has influence function equal to the efficient influence curve.

An estimator 
$$\hat{\psi}_n = \Psi(\hat{P}_n)$$
 is asymptotically linear if, 
$$\sqrt{n} \Big( \Psi(\hat{P}_n) - \Psi(P_0) \Big) = \sqrt{n} \mathbb{P}_n \phi^*(P_0) + o_P(1). \quad (*)$$

$$\Psi(\hat{P}_n) - \Psi(P_0) = (\hat{P}_n - P_0)\phi^*(\hat{P}_n) + R_2(\hat{P}_n, P_0)$$

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+ (\mathbb{P}_{n} - P_{0})(\phi^{*}(\hat{P}_{n}) - \phi^{*}(P_{0}))$$

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Evaluating the von Mises expansion in an estimator  $\hat{P}_n$  and the true data-generating  $P_0$ :

$$\Psi(\hat{P}_{n}) - \Psi(P_{0}) = (\hat{P}_{n} - P_{0})\phi^{*}(\hat{P}_{n}) + R_{2}(\hat{P}_{n}, P_{0})$$

$$= -P_{0}\phi^{*}(\hat{P}_{n}) + R_{2}(P_{0}, \hat{P}_{n})$$

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$$+ R_{2}(\hat{P}_{n}, P_{0})$$

$$- \mathbb{P}_{n}\phi^{*}(\hat{P}_{n})$$
(\*3)

i.e., need (\*1)–(\*3) to be  $o_P(n^{-1/2})$ .

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+ R_2(\hat{P}_n, P_0) 
- \mathbb{P}_n \phi^*(\hat{P}_n)$$
(\*2)

- (\*1) is an empirical process term.
- (\*2) second-order bias term.
- (\*3) is called the efficient influence curve equation.

- ... about the empirical process term (\*1):
  - 1. can be handled by empirical process theory, if  $(\phi^*(P): P \in \mathcal{M})$  is assumed Donsker.<sup>5</sup>
  - 2. otherwise can handled by extra sample splitting.<sup>6, 7</sup>

<sup>&</sup>lt;sup>5</sup>Lemma 19.24 of van der Vaart, A. W. (2000): Asymptotic statistics yields then that  $(\mathbb{P}_n - P_0)(\phi^*(\hat{P}_n) - \phi^*(P_0)) = o_P(n^{-1/2})$ .

<sup>&</sup>lt;sup>6</sup>Zheng, W., & van der Laan, M. J. (2010). Asymptotic theory for cross-validated targeted maximum likelihood estimation.

<sup>&</sup>lt;sup>7</sup>Chernozhukov, V., Chetverikov, D., Demirer, M., Duflo, E., Hansen, C., Newey, W., & Robins, J. (2018). Double/debiased machine learning for treatment and structural parameters.

Side note: Usually, we will assume the Donsker class condition.

- this is a way of nonparametrically characterizing the complexity of nuisance parameters.
- classes of functions that are Donsker: Indicator functions, bounded monotone functions, Lipschitz parametric functions, smooth functions, . . .

Donsker classes also include traditional parametric functions.

We will not discuss this further. For a nice intro see Sections 4.2 and 4.3 of Kennedy, E. H. (2016): Semiparametric theory and empirical processes in causal inference.

### Estimator expansion

This is basically what is needed to understand the construction of TMLE.

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#### Conditions (asymptotic linearity and efficiency)

- (C1) Solve the efficient influence curve equation:  $\mathbb{P}_n \phi^*(\hat{P}_n) = o_P(n^{-1/2})$ .
- (C2) Remainder  $R(\hat{P}_n, P_0) = o_P(n^{-1/2})$ .
- (C3) Donsker class conditions for  $\{\phi^*(P): P \in \mathcal{M}\}$ .

Then:  $\Psi(\hat{P}_n) \stackrel{as}{\sim} N(\Psi(P_0), P_0 \phi^*(P_0)^2/n)$ .

#### Construction of estimators

$$\Psi(\hat{P}_{n}) - \Psi(P_{0}) = \mathbb{P}_{n}\phi^{*}(P_{0}) + o_{P}(n^{-1/2}) + R(\hat{P}_{n}, P_{0}) - \mathbb{P}_{n}\phi^{*}(\hat{P}_{n})$$

For a given target parameter  $\Psi : \mathcal{M} \to \mathbb{R}$ , we need to

- 1. Know the efficient influence curve, so that we can solve the efficient influence curve equation.
- 2. Analyze the remainder  $R(P, P_0) \coloneqq \Psi(P) \Psi(P_0) + P_0 \phi^*(P)$ .

Repetition: These are solely properties of the estimation problem, but also tell us how to construct estimators such as TMLE.

# Example: ATE estimation

### EXAMPLE: Average treatment effect (ATE)

Observed data 
$$O = (X, A, Y) \in \mathbb{R}^d \times \{0, 1\} \times \{0, 1\} = \mathcal{O}$$

- \*  $X \in \mathbb{R}^d$  are covariates
- \*  $A \in \{0,1\}$  is a binary exposure variable (treatment decision)
- \*  $Y \in \{0,1\}$  is a binary outcome variable

 $O \sim P_0$  where  $P_0$  assumed to belong to nonparametric model  $\mathcal{M}$ .

We are interested in estimating the ATE:

$$\Psi(P) = \mathbb{E}_P \big[ \mathbb{E}_P \big[ Y \mid A = 1, X \big] - \mathbb{E}_P \big[ Y \mid A = 0, X \big] \big].$$

### EXAMPLE: Average treatment effect (ATE)

1. The efficient influence function:

$$\phi^{*}(P)(O) = \tilde{\phi}^{*}(f,\pi)(O)$$

$$= \left(\frac{A}{\pi(A \mid X)} - \frac{1 - A}{\pi(A \mid X)}\right) (Y - f(A,X)) + f(1,X) - f(0,X) - \Psi(P)$$

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2. The remainder:

$$\begin{split} R(P,P_0) &= \tilde{R}(f,\pi,f_0,\pi_0) \\ &= \int_{\mathbb{R}^d} \sum_{a=0,1} (2a-1) \frac{\pi_0(a\mid x) - \pi(a\mid x)}{\pi(a\mid x)} \big( f_0(a,x) - f(a,x) \big) d\mu_{0,X}(x) \end{split}$$

$$f(A, X) = \mathbb{E}_{P}[Y \mid A, X], \ \pi(A \mid X) = P(A = a \mid X)$$
  

$$f_{0}(A, X) = \mathbb{E}_{P_{0}}[Y \mid A, X], \ \pi_{0}(A \mid X) = P_{0}(A = a \mid X)$$

 $R(P, P_0) := \Psi(P) - \Psi(P_0) + P_0 \phi^*(P).$ 

2. Deriving the remainder for the ATE:

$$R(P, P_{0}) = \mathbb{E}_{P}[f(1, X) - f(0, X)] - \mathbb{E}_{P_{0}}[f_{0}(1, X) - f_{0}(0, X)]$$

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

$$+ \mathbb{E}_{P_{0}}[f(1, X) - f(0, X)] - \Psi(P)$$

$$\stackrel{*}{=} \int_{\mathbb{R}^{d}} \sum_{a=0,1} (2a - 1)\left(\frac{\pi_{0}(a \mid x)}{\pi(a \mid x)} - 1\right)(f_{0}(a, x) - f(a, x))d\mu_{0, X}(x)$$

$$= \int_{\mathbb{R}^{d}} \sum_{a=0,1} (2a - 1)\frac{\pi_{0}(a \mid x) - \pi(a \mid x)}{\pi(a \mid x)}(f_{0}(a, x) - f(a, x))d\mu_{0, X}(x)$$

the equality marked by \* is detailed on the next slide.

We used that:

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

$$= \mathbb{E}_{P_{0}}\left[\frac{2A - 1}{\pi(A\mid X)}(Y - f(A, X))\right]$$

$$= \mathbb{E}_{P_{0}}\left[\mathbb{E}_{P_{0}}\left[\frac{2A - 1}{\pi(A\mid X)}(Y - f(A, X))\middle|A, X\right]\right]$$

$$= \mathbb{E}_{P_{0}}\left[\frac{2A - 1}{\pi(A\mid X)}(f_{0}(A, X) - f(A, X))\right]$$

$$= \int_{\mathbb{R}^{d}} \sum_{a=0,1} \frac{2a - 1}{\pi(a\mid x)}(f_{0}(a, x) - f(a, x))\pi_{0}(a\mid x)d\mu_{0, X}(x)$$

$$= \int_{\mathbb{R}^{d}} \sum_{a=0,1} (2a - 1) \frac{\pi_{0}(a\mid x)}{\pi(a\mid x)}(f_{0}(a, x) - f(a, x))d\mu_{0, X}(x)$$

The remainder determines the asymptotic bias.

For the ATE, the remainder has a really nice structure.

$$R(P, P_0) = \tilde{R}(f, \pi, f_0, \pi_0)$$

$$= \int_{\mathbb{R}^d} \sum_{a=0,1} (2a-1) \frac{\pi_0(a \mid x) - \pi(a \mid x)}{\pi(a \mid x)} (f_0(a, x) - f(a, x)) d\mu_{0,X}(x)$$

A "double robust" structure, which has some important implications.

$$|R(P, P_0)| = |\tilde{R}(f, \pi, f_0, \pi_0)|$$

$$\leq \sum_{a=0}^{\infty} \int_{\mathbb{R}^d} \frac{|\pi_0(a \mid x) - \pi(a \mid x)|}{\pi(a \mid x)} |f_0(a, x) - f(a, x)| d\mu_{0, X}(x)$$

$$|R(P, P_{0})| = |\tilde{R}(f, \pi, f_{0}, \pi_{0})|$$

$$\leq \sum_{a=0,1} \int_{\mathbb{R}^{d}} \frac{|\pi_{0}(a \mid x) - \pi(a \mid x)|}{\pi(a \mid x)} |f_{0}(a, x) - f(a, x)| d\mu_{0, X}(x)$$

$$\stackrel{*}{\leq} \sum_{a=0,1} \frac{1}{\pi(a \mid x)} \sqrt{\int_{\mathbb{R}^{d}} \left\{ \pi_{0}(a \mid x) - \pi(a \mid x) \right\}^{2} d\mu_{0, X}(x)}$$

$$\times \sqrt{\int_{\mathbb{R}^{d}} \left\{ f_{0}(a, x) - f(a, x) \right\}^{2} d\mu_{0, X}(x)}$$

$$|R(P, P_{0})| = |\tilde{R}(f, \pi, f_{0}, \pi_{0})|$$

$$\leq \sum_{a=0,1} \int_{\mathbb{R}^{d}} \frac{|\pi_{0}(a \mid x) - \pi(a \mid x)|}{\pi(a \mid x)} |f_{0}(a, x) - f(a, x)| d\mu_{0, X}(x)$$

$$\stackrel{*}{\leq} \sum_{a=0,1} \frac{1}{\pi(a \mid x)} \sqrt{\int_{\mathbb{R}^{d}} \left\{ \pi_{0}(a \mid x) - \pi(a \mid x) \right\}^{2} d\mu_{0, X}(x)}$$

$$\times \sqrt{\int_{\mathbb{R}^{d}} \left\{ f_{0}(a, x) - f(a, x) \right\}^{2} d\mu_{0, X}(x)}$$

Thus, if  $\pi(a \mid X) > \delta > 0$  a.s., then:

$$|\tilde{R}(\hat{f}_{n}^{*}, \hat{\pi}_{n}, f_{0}, \pi_{0})| \leq \sum_{a=0,1} \delta^{-1} \|\pi_{0}(a \mid \cdot) - \hat{\pi}_{n}(a \mid \cdot)\|_{\mu_{0}} \|f_{0}(a \mid \cdot) - \hat{f}_{n}(a \mid \cdot)\|_{\mu_{0}}$$

### What does this imply for estimation?

#### Double robustness in consistency

$$\begin{split} |\tilde{R}(\hat{f}_{n}^{*},\hat{\pi}_{n},f_{0},\pi_{0})| &\leq \sum_{a=0,1} \delta^{-1} \underbrace{\|\pi_{0}(a\mid\cdot) - \hat{\pi}_{n}(a\mid\cdot)\|_{\mu_{\mathbf{0}}}}_{o_{P}(1), \text{ or }} \underbrace{\|f_{0}(a\mid\cdot) - \hat{f}_{n}^{*}(a\mid\cdot)\|_{\mu_{\mathbf{0}}}}_{o_{P}(1)} \end{split}$$
 then  $\tilde{\Psi}(\hat{f}_{n}^{*}) - \tilde{\Psi}(f_{0}) = o_{P}(1)$ .

#### Asymptotic linearity (easier to establish due to double robust structure)

$$|\tilde{R}(\hat{f}_{n}^{*},\hat{\pi}_{n},f_{0},\pi_{0})| \leq \sum_{a=0,1} \delta^{-1} \underbrace{\|\pi_{0}(a\mid\cdot) - \hat{\pi}_{n}(a\mid\cdot)\|_{\mu_{0}}}_{=o_{P}(n^{-1/4})} \underbrace{\|f_{0}(a\mid\cdot) - \hat{f}_{n}^{*}(a\mid\cdot)\|_{\mu_{0}}}_{=o_{P}(n^{-1/4})}$$
i.e., 
$$\tilde{R}(\hat{f}_{n}^{*},\hat{\pi}_{n},f_{0},\pi_{0}) = o_{P}(n^{-1/2}).$$

I.e., bias is converging at fast enough rate for reliable confidence intervals.

Side note: Showing the double robustness in consistency . . .

Say we have estimators  $(\hat{f}_n^*, \hat{\pi}_n)$ ;

- converging to  $(f, \pi)$
- solving the efficient influence curve equation.

Per definition,  $\tilde{R}(\hat{f}_n^*,\hat{\pi}_n,f_0,\pi_0) = \tilde{\Psi}(\hat{f}_n^*) - \tilde{\Psi}(f_0) + P_0\tilde{\phi}^*(\hat{f}_n^*,\hat{\pi}_n).$ 

l.e., 
$$\tilde{\Psi}(\hat{f}_n^*) - \tilde{\Psi}(f_0) = -P_0\tilde{\phi}^*(\hat{f}_n^*, \hat{\pi}_n) + \tilde{R}(\hat{f}_n^*, \hat{\pi}_n, f_0, \pi_0)$$
  
=  $(\mathbb{P}_n - P_0)\phi^*(\hat{f}_n^*, \hat{\pi}_n) + \tilde{R}(\hat{f}_n^*, \hat{\pi}_n, f_0, \pi_0).$ 

The first term is an empirical process term which equals  $(\mathbb{P}_n - P_0)\tilde{\phi}^*(f,\pi)$  plus an  $o_P(n^{-1/2})$ -term.

This then gives

$$\tilde{\Psi}(\hat{f}_n^*) - \tilde{\Psi}(f_0) = \underbrace{(\mathbb{P}_n - P_0)\tilde{\phi}^*(f,\pi)}_{\text{LLN applies}} + \tilde{R}(\hat{f}_n^*,\hat{\pi}_n,f_0,\pi_0) + o_P(n^{-1/2})$$

which yields that  $\tilde{\Psi}(\hat{f}_n^*) - \tilde{\Psi}(f_0) = o_P(1)$  if  $\tilde{R}(\hat{f}_n^*, \hat{\pi}_n, f_0, \pi_0) = o_P(1)$ .

#### Double robustness in consistency

• consistent estimation of target parameter requires only that either f or  $\pi$  is consistently estimated.

can be important, but even more important is:

#### "Rate double robustness"

- convergence rate requirements for estimators f and  $\pi$  are seriously weakened;
- asymptotic linearity is achieved when these are estimated (consistently) at rate at least n<sup>-1/4</sup>.8

<sup>&</sup>lt;sup>8</sup> and this is actually possible with certain machine learning algorithms; and if these are combined in super learner there is something called the oracle property, which says that the super learner achieves the rate of convergence of the *best* estimator in its library.

#### EXAMPLE: Average treatment effect (ATE)

1. The efficient influence function:

$$\phi^*(P)(O) = \tilde{\phi}^*(f,\pi)(O) = \left(\frac{A}{\pi(A|X)} - \frac{1-A}{\pi(A|X)}\right) (Y - f(A,X)) + f(1,X) - f(0,X) - \Psi(P)$$

2. The remainder:

$$R(P, P_0) = \tilde{R}(f, \pi, f_0, \pi_0)$$

$$= \int_{\mathbb{R}^d} \sum_{a=0,1} (2a-1) \frac{\pi_0(a \mid x) - \pi(a \mid x)}{\pi(a \mid x)} (f_0(a, x) - f(a, x)) d\mu_{0,X}(x)$$

Deriving these is done once for a given target parameter  $\Psi: \mathcal{M} \to \mathbb{R}$ .

### TMLE as an estimation procedure

#### TMLE is a two-step procedure:

- Step 1 Construct initial estimator  $\hat{P}_n$  for P such that  $R(\hat{P}_n, P_0) = o_P(n^{-1/2})$ .
- Step 2 Update the estimator  $\hat{P}_n \mapsto \hat{P}_n^*$  such that  $\hat{P}_n^*$  solves the efficient influence curve equation.

TMLE and the estimating equation (EE) estimator both solve the efficient influence curve equation ... that is why they share the same asymptotic (large-sample) properties.

We will talk more later about deriving efficient influence curves, but just note for now that it is very much tied to the notion of derivatives along smooth parametric submodels.

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The von Mises expansion (1) implies a related notion of smoothness called pathwise differentiability, i.e.,

$$\frac{d}{d\varepsilon}\bigg|_{\varepsilon=0}\Psi(P_{\varepsilon}) = \int \phi(P)(o)b(o)dP(o), \tag{2}$$

for every smooth submodel  $\{P_{\varepsilon}: \varepsilon \in \mathbb{R}\} \subset \mathcal{M}$ , where  $P_{\varepsilon}$  has density density  $p_{\varepsilon}$ , and for which  $P_{\varepsilon=0}=P_0$ .

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for every smooth submodel  $\{P_{\varepsilon} : \varepsilon \in \mathbb{R}\} \subset \mathcal{M}$ , where  $P_{\varepsilon}$  has density density  $p_{\varepsilon}$ , and for which  $P_{\varepsilon=0} = P_0$ .

Equation (2) can be used to derive the efficient influence curve.

For example one evaluates the pathwise derivative along submodels defined for a mean-zero function  $h: \mathcal{O} \to \mathbb{R}$  as:

$$p_{\varepsilon}(o) = p(o)(1 + \varepsilon h(o)).$$

and solves the integral equation on the right hand side of (2).

Note that this submodel has score function  $\frac{d}{d\varepsilon}\Big|_{\varepsilon=0}\log p_{\varepsilon}(o)=h(o)$ .

(And there are many other useful tricks in this process).

Another common strategy is to assume the data are discrete and then compute the Gateaux derivative:

$$\frac{d}{d\varepsilon}\Big|_{\varepsilon=0} \Psi((1-\varepsilon)p(o)+\varepsilon\delta_{o'}),$$

which equals the influence curve  $\phi(P)(o')$  directly.

Note that this really corresponds to computing the pathwise derivative along the particular submodel of the form  $(1-\varepsilon)p(o)+\varepsilon\delta_{o'}$  for which the right hand side of (2) is  $\phi(P)(o')$ .

In this exercise we continue the simulation setting of Practical 1, now to explore —

- 1. Inference for estimators based on the efficient influence curve;
- 2. Variance estimation and coverage of confidence intervals;
- 3. Small-sample properties, particularly under positivity violations.

This is described in detail in: day1-practical2.pdf.

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- these exercises emphasize the asymptotic equivalence of TMLE and estimating equation (EE) estimation;
- there may be small-sample differences in performance (often argued in favor of TMLE, but this is not necessarily a general result);
- otherwise, the differences are not so important for the ATE estimation problem. BUT, for other problems (e.g., in survival analysis), the substitution property of TMLE may be crucial.

### Summary

#### "Classical" causal inference estimators:

- consistency of g-formula estimators rely on correct specification of the outcome regression  $f(a,x) = \mathbb{E}[Y \mid A = a, X = x]$
- consistency of ipw estimators rely on correct specification of the propensity score π(a | x) = P(A = a | X = x)
- both types of estimators work poorly with machine learning (cross-validation does not help)

### TMLE (and other estimators based on the efficient influence curve):

- built-in bias correction
- double robustness in consistency
- inference based on the efficient influence curve
- even when incorporating machine learning (under conditions!!)