

## Causal Inference (Part 2)

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<sup>1</sup>Adopted from the lecture notes of professor Marie Davidian

# Observational Studies

- We now consider observational studies.
- **Complication:** Individuals may receive treatment according to physician discretion or their own choice.
- Therefore, individuals who receive treatment 1 may have different characteristics than those who receive treatment 0.
- Thus, the Difference-in-Means estimator  $\hat{\tau}$  is no longer unbiased.
- We may want to utilize the auxiliary variable  $X$  to control the systematic difference.

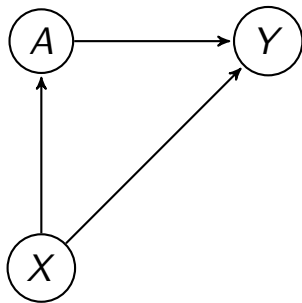
# Confounders

- If a variable is related to both  $A$  and outcome variable  $\{Y(1), Y(0)\}$ , then it is called a confounder.
- There could be several confounders (e.g. gender, race, age group, education level).
- Suppose that the confounding variables are known. Among individuals who share the same confounders, all factors associated with the selection and outcome are taken into account.
- That is, the assignment of treatment is effectively *at random* for individuals sharing the same confounders.
- Formally expressed as

$$\{Y(1), Y(0)\} \perp A \mid X^*$$

where  $X^*$  is the set of all confounders.

# Confounding factor example



- A: smoking
- Y: lung cancer
- X: stress

# No Unmeasured confounders assumption

- The auxiliary variable  $X$  contains the individual characteristics recorded in the observed data and available to the data analyst.
- In general, there is no guarantee that all the confounders are included in  $X$ .
- So, we assume that the confounders are all included in  $X$  so that

$$\{Y(1), Y(0)\} \perp A \mid X \quad (1)$$

holds. This is *No Unmeasured Confounders* (NUC) assumption. This is also called the *ignorability* assumption.

- **Fundamental difficulty:** It is impossible to verify from the observed data that there are no unmeasured confounders and that (1) is valid.

# Main Result

- Three assumptions

- ① SUTVA

$$Y = Y(1) \cdot A + Y(0) \cdot (1 - A)$$

- ② NUC assumption in (1).

- ③ Positivity assumption:

$$P(A = 1 \mid X) > 0$$

for all  $x \in \mathcal{X}$  with  $p_X(x) > 0$ .

- Under the three assumptions, we obtain the following.

$$E\{E(Y \mid X, A = a)\} = E\{Y(a)\} \quad (2)$$

for  $a = 0, 1$ .

# Justification

- SUTVA leads to

$$E\{Y \mid X, A = a\} = E\{Y(a) \mid X, A = a\}$$

- NUC assumption leads to

$$E\{Y(a) \mid X, A = a\} = E\{Y(a) \mid X\} \quad (3)$$

- Now, note that

$$E[E\{Y(a) \mid X\}] = \int_{\mathcal{X}} E\{Y(a) \mid X = x\} p_X(x) d\nu(x)$$

$$E[E\{Y(a) \mid X, A = a\}] = \int_{\mathcal{X}} E\{Y(a) \mid X = x, A = a\} p_X(x, a) d\nu(x)$$

Thus, given (3) holds, two terms are equal if  $p(x, a) > 0$  for all  $x \in \mathcal{X}$  with  $p_X(x) > 0$ .

- Implication of (2) :

$$\tau = E\{E(Y | X, A = 1)\} - E\{E(Y | X, A = 0)\}$$

depends on  $E(Y | X, A)$ , the regression function of  $Y$  on  $X$  and  $A$ .

- If

$$E(Y | X = x, A = a) = Q(x, a)$$

is known, we can obtain

$$\tau = E\{Q(X, 1)\} - E\{Q(X, 0)\}.$$

- In practice,  $Q(x, a)$  is unknown.



# Outcome regression model

- Idea: Let's use a statistical model for  $Q(x, a) = Q(x, a; \beta)$  with parameter  $\beta$ .
- For example, if  $Y$  is continuous, we can consider a linear regression model such as

$$Q(\mathbf{x}, a; \beta) = \beta_1 + \beta_2 a + \beta_3' \mathbf{x} + \beta_4' \mathbf{x} \cdot a$$

- Model parameter can be estimated from each sample.
- We can estimate  $\tau$  by

$$\hat{\tau} = \frac{1}{n} \sum_{i=1}^n \left\{ Q(x_i, 1; \hat{\beta}) - Q(x_i, 0; \hat{\beta}) \right\}$$

- Thus, the theory of imputation (= prediction) can be applied directly.

# Propensity score approach

- The treatment indicator function  $A$  takes values in  $\{0, 1\}$ .
- The propensity score function  $\pi(X) = P(A = 1 \mid X)$  reflects the “propensity” for choosing  $A = 1$  for individuals with characteristics  $X$ .
- Originally proposed by Rosenbaum and Rubin (1983).

# Inverse propensity score weighting

- For now, suppose that  $\pi(X)$  is known.
- Estimation of  $\mu_1 = E\{Y(1)\}$ : Use

$$\hat{\mu}_1 = \frac{1}{n} \sum_{i=1}^n \frac{A_i Y_i}{\pi(X_i)}$$

- Estimation of  $\mu_0 = E\{Y(0)\}$ : Use

$$\hat{\mu}_0 = \frac{1}{n} \sum_{i=1}^n \frac{(1 - A_i) Y_i}{1 - \pi(X_i)}$$

- The unbiasedness requires the same three assumptions (SUTVA, NUC, and Positivity) in addition to  $\pi(x)$  known.

# Justification

- If  $\pi(x)$  known

$$\hat{\tau} = \frac{1}{n} \sum_{i=1}^n \left\{ \frac{A_i Y_i}{\pi(X_i)} - \frac{(1 - A_i) Y_i}{1 - \pi(X_i)} \right\}$$

- Observational studies: Posit and fit propensity model  $\pi(x; \gamma)$

$$\hat{\tau} = \frac{1}{n} \sum_{i=1}^n \left\{ \frac{A_i Y_i}{\pi(X_i; \hat{\gamma})} - \frac{(1 - A_i) Y_i}{1 - \pi(X_i; \hat{\gamma})} \right\}$$

- It is based on the assumption that the propensity model  $\pi(x; \gamma)$  is correctly specified.

# Outcome regression vs inverse propensity weighted estimator

- Both require SUTVA, NUC, and the positivity assumption.
- Outcome regression estimator: requires a model assumption on  $E(Y \mid x, a) = Q(x, a; \beta)$ .
- IPW estimator: requires a model assumption on  $P(A = 1 \mid x) = \pi(x; \gamma)$
- Maybe a doubly robust estimator can be developed to protect against model misspecification.

# Double robust estimation of $\tau$

- Given fitted models  $\pi(x; \hat{\gamma})$  and  $Q(x, a; \hat{\beta})$ , we can consider

$$\hat{\tau}_{\text{DR}} = \hat{\mu}_{1,\text{DR}} - \hat{\mu}_{0,\text{DR}} \quad (4)$$

where

$$\hat{\mu}_{1,\text{DR}} = \frac{1}{n} \sum_{i=1}^n \left[ Q(x_i, 1; \hat{\beta}) + \frac{A_i}{\pi(x_i; \hat{\gamma})} \left\{ Y_i - Q(x_i, 1; \hat{\beta}) \right\} \right]$$

and

$$\hat{\mu}_{0,\text{DR}} = \frac{1}{n} \sum_{i=1}^n \left[ Q(x_i, 0; \hat{\beta}) + \frac{1 - A_i}{1 - \pi(x_i; \hat{\gamma})} \left\{ Y_i - Q(x_i, 0; \hat{\beta}) \right\} \right]$$

# Justification



- $\hat{\tau}_{\text{DR}}$  is doubly robust.
- It is efficient (i.e. optimal) when both the propensity and the outcome regression models are correctly specified.
  - It achieves the smallest variance among the class of estimators of the form

$$\hat{\tau}_h = \frac{1}{n} \sum_{i=1}^n \left[ \frac{A_i Y_i}{\pi(X_i; \hat{\gamma})} - \frac{(1 - A_i) Y_i}{1 - \pi(X_i; \hat{\gamma})} - \{A_i - \pi(X_i; \hat{\gamma})\} h(X_i) \right]$$

- That is, the optimal  $h(x)$  is

$$h^*(X) = \frac{E(Y | X, A = 1)}{\pi(X)} + \frac{E(Y | X, A = 0)}{1 - \pi(X)}$$

- Based on the semiparametric theory (Tsiatis, 2006).

## REFERENCES

- Rosenbaum, P. R. and D. B. Rubin (1983), 'The central role of the propensity score in observational studies for causal effects', *Biometrika* **70**, 41–55.
- Tsiatis, A. A. (2006), *Semiparametric Theory and Missing Data*, Springer.