

# Lecture 6

## Regression for completely randomized experiment

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# Outline

- Using regression with no covariates
- Using regression with covariates adjustments
- Using regression with covariates adjustments and interactions
- The LRC-CPPT cholesterol data example
- Suggested readings: Imbens and Rubin Chapter 7

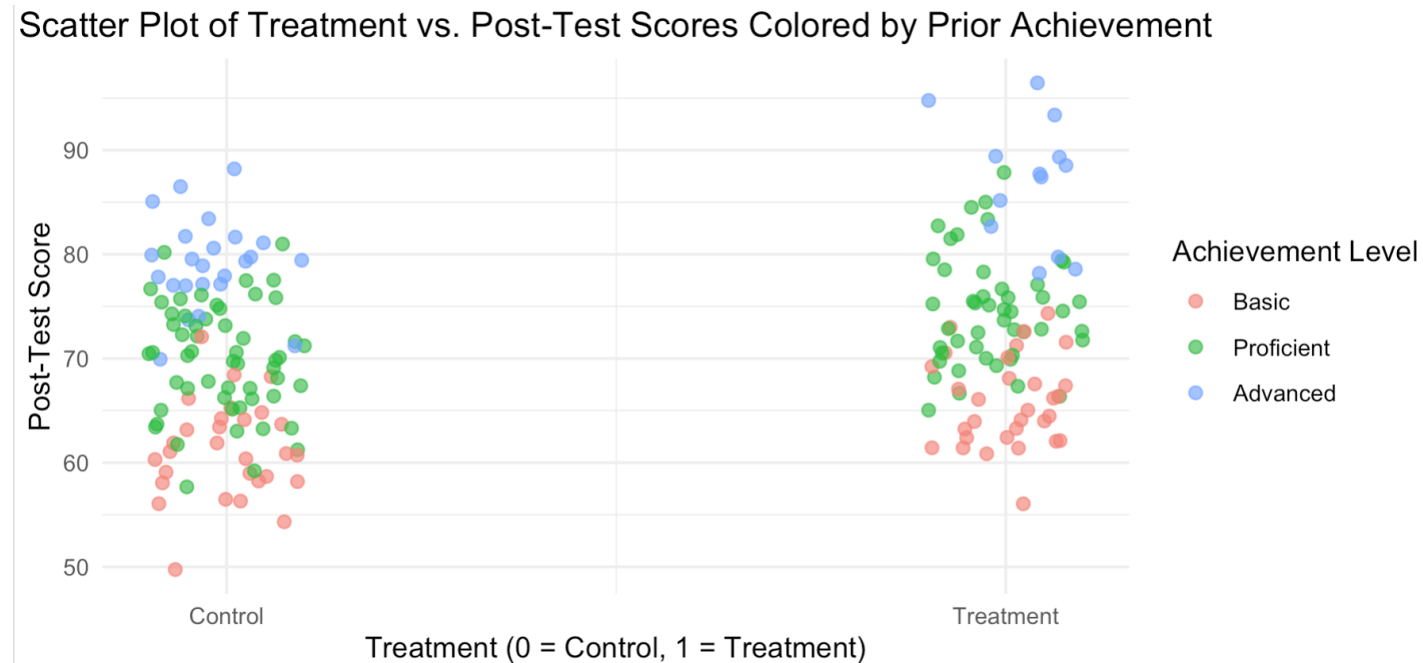
# Linear regression and causality

- Linear regression:

$$\mathbb{E}(Y_i | W_i, \mathbf{X}_i) = \alpha + \gamma W_i + \boldsymbol{\beta}^T \mathbf{X}_i$$

- Benefits of using linear regression:

- Adjust for confounding variables
  - Not need for completely randomized experiments as pre-treatment covariates are not confounded
- More accurate estimator if covariates explain part of the noise in the outcome



# Linear regression and causality

- Linear regression:

$$\mathbb{E}(Y_i|W_i, \mathbf{X}_i) = \alpha + \gamma W_i + \boldsymbol{\beta}^T \mathbf{X}_i$$

- Question:

- When can we interpret the coefficient(s) as causal effect?
- How can we do correct inference if we take into account the randomization procedure of treatment assignments?

- Some critiques

- In completely randomized experiments, covariates are not confounders
- Why do we want to assume a linear model if we don't need to?
  - Model  $\mathbb{E}(Y_i|W_i, \mathbf{X}_i) = \alpha + \gamma W_i + \boldsymbol{\beta}^T \mathbf{X}_i$  assumes same causal effect for all levels of  $\mathbf{X}_i$

“Experiments should be analyzed as experiments, not as observational studies”

---- David A. Freedman, 2006

# The LRC-CPPT cholesterol data

- An experiment to evaluate the effect of the drug cholestyramine on reducing cholesterol levels
- $N = 337$  patients are completely randomized
- **Pre-treatment covariates:** two cholesterol measurements before and after a suggestion of low-cholesterol diet, both measurements taken prior to the random assignment
  - $\text{cholp} = 0.25 \text{ chol1} + 0.75 \text{ chol2}$

**Table 7.1. Summary Statistics for PRC-CPPT Cholesterol Data**

	Variable	Control ( $N_c = 172$ )		Treatment ( $N_t = 165$ )		Min	Max
		Average	Sample (S.D.)	Average	Sample (S.D.)		
Pre-treatment	chol1	297.1	(23.1)	297.0	(20.4)	247.0	442.0
	chol2	289.2	(24.1)	287.4	(21.4)	224.0	435.0
	cholp	291.2	(23.2)	289.9	(20.4)	233.0	436.8
Post-treatment	cholf	282.7	(24.9)	256.5	(26.2)	167.0	427.0
	chold	-8.5	(10.8)	-33.4	(21.3)	-113.3	29.5
	comp	74.5	(21.0)	59.9	(24.4)	0	101.0

# The LRC-CPPT cholesterol data

- An experiment to evaluate the effect of the drug cholestyramine on reducing cholesterol levels
- $N = 337$  patients are completely randomized
- **Post-treatment outcomes:**
  - chol<sub>f</sub>: post-treatment average cholesterol level
  - chol<sub>d</sub> = chol<sub>f</sub> – chol<sub>p</sub>
  - comp: compliance rate, the percentage of individuals follow the treatment assignment

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	Variable	Control ( $N_c = 172$ )		Treatment ( $N_t = 165$ )		Min	Max
		Average	Sample (S.D.)	Average	Sample (S.D.)		
Pre-treatment	chol <sub>1</sub>	297.1	(23.1)	297.0	(20.4)	247.0	442.0
	chol <sub>2</sub>	289.2	(24.1)	287.4	(21.4)	224.0	435.0
	chol <sub>p</sub>	291.2	(23.2)	289.9	(20.4)	233.0	436.8
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# The LRC-CPPT cholesterol data

- Can we evaluate the drug effect by simply look at whether chold is positive or negative?
  - **No!** The before-after comparison is NOT necessarily causal
  - Even for the control group, chold is significantly negative
- The patient's post-treatment cholesterol should be highly correlated with his/her pre-treatment cholesterol level
- **How do we evaluate the causal effect after “adjusting for the pre-treatment cholesterol”?**
  - **Adjust for pre-treatment cholesterol by regression**

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# Linear regression with no covariates

- Neyman's approach

$$\hat{\tau}^{\text{dif}} = \bar{Y}_t^{\text{obs}} - \bar{Y}_c^{\text{obs}}$$

$$\bar{Y}_c^{\text{obs}} = \frac{1}{N_c} \sum_{i:W_i=0} Y_i^{\text{obs}} \quad \text{and} \quad \bar{Y}_t^{\text{obs}} = \frac{1}{N_t} \sum_{i:W_i=1} Y_i^{\text{obs}}$$

- An alternative way to get  $\hat{\tau}^{\text{dif}}$  is by linear regression

$$(\hat{a}, \hat{b}) = \arg \min_{(a,b)} \sum_{i=1}^N (Y_i - a - bW_i)^2$$

- It is easy to show that  $\hat{b} = \hat{\tau}^{\text{dif}}$

- $\arg \min_{(a,b)} \sum_{i=1}^n (Y_i - a - bW_i)^2 = \arg \min_{(a,b)} [\sum_{i:W_i=0} (Y_i - a)^2 + \sum_{i:W_i=1} (Y_i - a - b)^2]$

- $\hat{a} = \bar{Y}_c^{\text{obs}}, \hat{a} + \hat{b} = \bar{Y}_t^{\text{obs}}$

- However, if we use R function `lm()`, the variance estimator is  $\frac{1}{N} \frac{(N_c-1)s_c^2 + (N_t-1)s_t^2}{N-2}$

- Different from  $\frac{s_c^2}{N_c} + \frac{s_t^2}{N_t}$  in Neyman's approach



# Causal interpretation of the linear model

- Linear model on the potential outcomes

$$Y_i(w) = \alpha + \tau_i w + \varepsilon_i^* = \alpha + \tau w + \varepsilon_i(w)$$

where  $\mathbb{E}(\varepsilon_i^*) = 0$  and  $\varepsilon_i(w) = \varepsilon_i^* + (\tau_i - \tau)w$

- Not really an assumption if  $w$  only has two values

$$Y_i(0) = \alpha + \varepsilon_i^*, Y_i(1) = Y_i(0) + \tau_i$$

- Assume that there is a super-population and the potential outcomes are i.i.d. samples
- The observed outcomes  $Y_i = W_i Y_i(1) + (1 - W_i) Y_i(0)$  are **not i.i.d. samples** under complete randomization
  
- Define PATE:  $\tau = \mathbb{E}(\tau_i) = \mathbb{E}(Y_i(1) - Y_i(0))$
- $\alpha = \mathbb{E}(Y_i(0))$  and  $\mathbb{E}(\varepsilon_i(w)) = 0$

# Linear regression with no covariates

- Causal model on the potential outcomes

$$Y_i(w) = \alpha + \tau_i w + \varepsilon_i^* = \alpha + \tau w + \varepsilon_i(w)$$

where  $\mathbb{E}(\varepsilon_i^*) = 0$  and  $\varepsilon_i(w) = \varepsilon_i^* + (\tau_i - \tau)w$

- If the treatment is binary ( $w = 0,1$ ), then the above model essentially has no assumption on  $Y_i(0)$  and  $Y_i(1)$
- If the treatment is continuous, the model assumes a linear but heterogenous causal effect on each individual
- How to estimate  $\tau$  from observed data?
- When does the above model imply the linear regression model on observed data?

$$Y_i^{\text{obs}} = \alpha + \tau W_i + \varepsilon_i$$

# Linear regression with no covariates

$$Y_i(w) = \alpha + \tau_i w + \varepsilon_i^* = \alpha + \tau w + \varepsilon_i(w)$$

We assume the following identification conditions

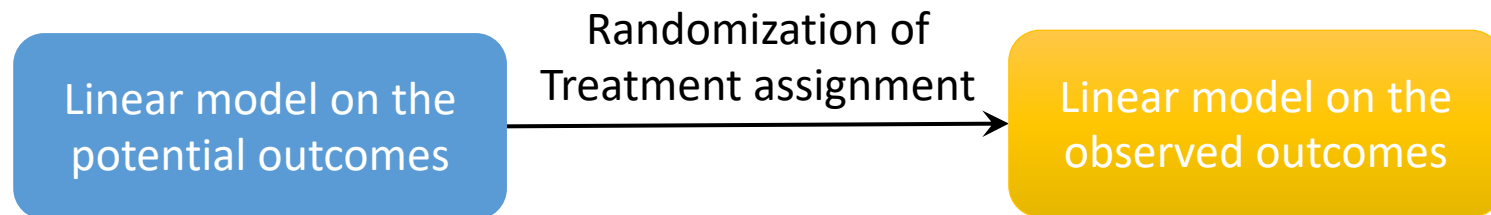
- **Randomization of the treatment:**

$$(Y(0), Y(1)) \perp W$$

- Satisfied in completely randomized experiments
- Then,  $\mathbb{E}(Y_i(w)) = \mathbb{E}(Y_i(w) | W_i = w) = \mathbb{E}(Y_i^{\text{obs}} | W_i = w) = \alpha + \tau w$
- So this implies a regression model

$$Y_i^{\text{obs}} = \alpha + \tau W_i + \varepsilon_i$$

where  $\varepsilon_i = \varepsilon_i(W_i) = \varepsilon_i^* + (\tau_i - \tau)W_i$



- What is the correct statistical inference?

# Linear regression with no covariates

- regression model

$$Y_i^{\text{obs}} = \alpha + \tau W_i + \varepsilon_i$$

where  $\varepsilon_i = \varepsilon_i(W_i) = \varepsilon_i^* + (\tau_i - \tau)W_i$

- Follow the linear regression convention, we perform statistical inference conditional on  $(W_1, \dots, W_N)$ 
  - we treat assignment vectors as fixed
- Random sampling of the units
  - $(\varepsilon_i(0), \varepsilon_i(1))$  are independent across  $i$
  - This implies that  $\varepsilon_i$  in the linear regression model are **independent** as  $W_i$  are treated as fixed
  - But they may not follow the same distribution

# Homoscedastic error assumption

Homoscedastic error assumption:  $\mathbb{V}(\varepsilon_i(0)) = \mathbb{V}(\varepsilon_i(1)) = \sigma^2$

- Then  $\mathbb{V}(Y_i^{\text{obs}} | W_i) = \varepsilon_i = \varepsilon_i(W_i)$  always has variance  $\sigma^2$

- Under homoscedasticity, OLS estimates of the variance is

$$\hat{\sigma}_{Y|W}^2 = \frac{1}{N-2} \sum_{i=1}^N \hat{\varepsilon}_i^2 = \frac{1}{N-2} \sum_{i=1}^N (Y_i^{\text{obs}} - \hat{Y}_i^{\text{obs}})^2,$$

where the estimated residual is  $\hat{\varepsilon}_i = Y_i^{\text{obs}} - \hat{Y}_i^{\text{obs}}$ , and the predicted value  $\hat{Y}_i^{\text{obs}}$  is

$$\hat{Y}_i^{\text{obs}} = \begin{cases} \hat{\alpha}^{\text{ols}} & \text{if } W_i = 0, \\ \hat{\alpha}^{\text{ols}} + \hat{\tau}^{\text{ols}} & \text{if } W_i = 1. \end{cases}$$

- Same as the standard linear regression approach

# Heteroscedastic errors

- If we don't want to assume  $\mathbb{V}(\varepsilon_i(0)) = \mathbb{V}(\varepsilon_i(1))$ , then the homoscedastic error assumption fails
  - $\varepsilon_i$  has the same distribution for  $W_i = 0$ , and the same distribution for  $W_i = 1$
  - We should use same variance within the treated and control group
  - That leads to the variance estimator in Neyman's approach
- This is also called the Sandwich estimator that is robust to the violation of the homoscedastic noise assumption in linear regression
  - In R, it corresponds to Sandwich estimator with HC2 adjustment

# Linear regression with no covariates

## To summarize the logic

- We build a (linear) model on the potential outcomes
- This model implies a linear regression model on the observed outcome if  $(Y(0), Y(1)) \perp W$
- The coefficient on  $W_i$  in the linear regression model is the average causal effect (PATE)
- The linear regression model treats  $W$  as fixed so it works for any randomization assignment mechanism that satisfies  $(Y(0), Y(1)) \perp W$
- Noise in the linear regression model are independent as long as potential outcomes are independent across units
- **For statistical inference**
  - The OLS estimator is always unbiased
  - We can apply standard linear regression inference results if we assume  $V(\varepsilon_i(0)) = V(\varepsilon_i(1))$
  - If  $V(\varepsilon_i(0)) \neq V(\varepsilon_i(1))$ , we need to use the robust variance estimator

# Linear regression with covariates adjustment

- What are model assumptions on the potential outcomes that lead to

$$Y_i^{\text{obs}} = \alpha + \tau W_i + \boldsymbol{\beta}^T \mathbf{X}_i + \varepsilon_i$$

a linear model on the observed outcome

- Assumption 1:  $\mathbb{E}(Y_i(0) | \mathbf{X}_i) = \alpha + \boldsymbol{\beta}^T \mathbf{X}_i$
- Assumption 2: CATE  $\tau(\mathbf{x}) = \mathbb{E}(\tau_i | \mathbf{X}_i = \mathbf{x}) \equiv \tau = \text{PATE}$  constant across levels of  $\mathbf{X}_i$ 
  - We can allow for heterogeneous causal effect but need  $\mathbb{E}(\tau_i - \tau | \mathbf{X}_i) = 0$   
(individual causal effects are independent from the pre-treatment covariates)
- Then  $\mathbb{E}(Y_i(w) | \mathbf{X}_i) = \mathbb{E}(Y_i(0) + \tau_i w | \mathbf{X}_i) = \alpha + \tau w + \boldsymbol{\beta}^T \mathbf{X}_i$
- Unconfoundedness property:  
 $(Y(0), Y(1)) \perp W | X$
- $\mathbb{E}(Y_i^{\text{obs}} | W_i = w, \mathbf{X}_i = \mathbf{x}) = \mathbb{E}(Y_i(w) | \mathbf{X}_i = \mathbf{x}) = \alpha + \tau w + \boldsymbol{\beta}^T \mathbf{X}_i$ 
  - Statistical inference is conditional on both  $\mathbf{X}_i$  and  $W_i$



# OLS with covariates adjustment

$$(\hat{\alpha}^{\text{ols}}, \hat{\tau}^{\text{ols}}, \hat{\beta}^{\text{ols}}) = \arg \min_{\alpha, \tau, \beta} \sum_{i=1}^N \left( Y_i^{\text{obs}} - \alpha - \tau \cdot W_i - X_i \beta \right)^2$$

- The estimator  $\hat{\tau}^{\text{ols}}$  is unbiased for the causal estimand  $\tau$
- Even if the model is **incorrect** (either the violation of  $\mathbb{E}(Y_i(0) | \mathbf{X}_i) = \alpha + \boldsymbol{\beta}^T \mathbf{X}_i$  or  $\tau \equiv \mathbb{E}(\tau_i | \mathbf{X}_i = \mathbf{x})$ ),  $\hat{\tau}^{\text{ols}}$  still converges to the PATE  $\mathbb{E}(\tau_i)$  under complete randomization

## Efficiency gain from regression

- If the model is correct, we have

$$\mathbb{V}(\hat{\tau}^{\text{ols}}) \approx \frac{\mathbb{E}\{\mathbb{V}(Y_i(1) | \mathbf{X}_i)\}}{N_t} + \frac{\mathbb{E}\{\mathbb{V}(Y_i(0) | \mathbf{X}_i)\}}{N_c} \leq \frac{\sigma_c^2}{N_c} + \frac{\sigma_t^2}{N_t}$$

- If  $\mathbf{X}_i$  is predictive of the (potential) outcomes, we have a more accurate estimator
- If the linear model is **incorrect**, the efficiency might be lost (Freedman 2008, *Adv. Appl. Math.*)

# Estimate of the variance of $\hat{\tau}^{\text{ols}}$ with covariates adjustment

- Assume homoscedastic error assumption:

$$\mathbb{V}(\varepsilon_i(0)) = \mathbb{V}(\varepsilon_i(1)) = \sigma^2 = \mathbb{V}(Y_i^{\text{obs}} | W_i, \mathbf{X}_i)$$

We can follow standard linear regression inference and estimate variance of  $\hat{\tau}^{\text{ols}}$  as

$$\hat{\mathbb{V}}_{\text{sp}}^{\text{homo}} = \frac{1}{N(N-1-\dim(\mathbf{X}_i))} \cdot \frac{\sum_{i=1}^N \left( Y_i^{\text{obs}} - \hat{\alpha}^{\text{ols}} - \hat{\tau}^{\text{ols}} - \mathbf{X}_i \hat{\beta}^{\text{ols}} \right)^2}{\bar{W} \cdot (1 - \bar{W})}$$

- The robust variance estimator (Sandwich estimator) without assuming homoscedasticity

$$\hat{\mathbb{V}}_{\text{sp}}^{\text{hetero}} = \frac{1}{N(N-1-\dim(\mathbf{X}_i))} \cdot \frac{\sum_{i=1}^N (W_i - \bar{W})^2 \cdot \left( Y_i^{\text{obs}} - \hat{\alpha}^{\text{ols}} - \hat{\tau}^{\text{ols}} - \mathbf{X}_i \hat{\beta}^{\text{ols}} \right)^2}{(\bar{W} \cdot (1 - \bar{W}))^2}$$

# Linear regression with covariates adjustment and interactions

What if the assumption  $\tau \equiv \tau(\mathbf{x}) = \mathbb{E}(\tau_i | \mathbf{X}_i = \mathbf{x})$  constant across levels of  $\mathbf{X}_i$  is incorrect?

- Assume CATE  $\tau(\mathbf{x}) = \mathbb{E}(\tau_i | \mathbf{X}_i = \mathbf{x}) = \tau + \boldsymbol{\gamma}^T (\mathbf{x} - \bar{\mathbf{X}})$ 
  - $\tau$  is still the population average treatment effect
- Still assume  $\mathbb{E}(Y_i(0) | \mathbf{X}_i) = \alpha + \boldsymbol{\beta}^T \mathbf{X}_i$
- Then  $\mathbb{E}(Y_i(w) | \mathbf{X}_i) = \mathbb{E}(Y_i(0) + \tau_i w | \mathbf{X}_i) = \alpha + \tau w + \boldsymbol{\beta}^T \mathbf{X}_i + \boldsymbol{\gamma}^T (\mathbf{X}_i - \bar{\mathbf{X}}) w$
- When does the above model imply the linear regression model with interactions on observed data?
$$Y_i^{\text{obs}} = \alpha + \tau W_i + \boldsymbol{\beta}^T \mathbf{X}_i + \boldsymbol{\gamma}^T (\mathbf{X}_i - \bar{\mathbf{X}}) W_i + \varepsilon_i$$
  - Unconfoundedness property  $\rightarrow$  check by yourself
  - In completely randomized experiments, with the interaction terms, we can always guarantee no efficiency loss even when the linear model is wrong (Peng's book section 6.2.2)

# Results on the LRC-CPPT cholesterol data

- We estimate the PATE for both the post-treatment cholesterol level `cholp` and compliance
  - A considerable reduction of the variance of  $\hat{\tau}^{ols}$  for `cholp` when we add the pre-treatment cholesterol levels in the regression
  - Our goal is always estimating PATE even after “covariates adjustment”
  - In randomized experiments satisfying  $(Y(0), Y(1)) \perp W$ , adjusting for covariates or not, our estimate of PATE is always valid, we only change the efficiency of our estimate

Covariates	Effect of Assignment to Treatment on			
	Post-Cholesterol Level		Compliance	
	$\hat{\tau}$	$\widehat{\text{s. e.}}$	$\hat{\tau}$	$\widehat{\text{s. e.}}$
No covariates	-26.22	(3.93)	-14.64	(3.51)
<code>cholp</code>	-25.01	(2.60)	-14.68	(3.51)
<code>chol1</code> , <code>chol2</code>	-25.02	(2.59)	-14.95	(3.50)
<code>chol1</code> , <code>chol2</code> , interacted with $W$	-25.04	(2.56)	-14.94	(3.49)

# The LRC-CPPT cholesterol data

## A bit explanation about compliance

- If we compare between control and treatment group, we are evaluating the causal effect of “being assigned”, not the causal effect of actually taking the drug
- Compliance lower in the treatment group possibly due to the side effect of the drug
- Can we just throw away individuals who do not follow the treatment and estimate the causal effect of taking the drug based on the rest individuals? **No**
- Will discuss more about compliance in later lectures

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# Why do we use linear regression in randomized experiments?

- Covariate adjustment can be used to improve efficiency in randomized experiments
  - Always add interaction terms (between each covariate and treatment) to guarantee power improvement
- In completely randomized experiments
  - No need to worry about model misspecification
  - Treatment and covariates are independent