

# Segment 2: Randomized Studies

## Section 02: Randomization-Based Inference

# Randomization-Based Inference

For *population-level* inference we think of a sampling distribution dictated by:

- ▶ Randomly sampling units from a population
- ▶ Randomly assigning  $\mathbf{Z}$  in accordance with the assignment mechanism

For *sample-level* inference

- ▶ We are interested in inferring something about potential outcomes/causal effects in a given sample
  - ▶ (e.g,  $\tau_{SATE}$ )
- ▶ We are not focused on repeated samples from a population
  - ▶ So what is random? Where does variability in a given estimate come from?
- ▶ View  $\mathbf{Z}$  as the random quantity
- ▶ Envision re-randomization of the sample units in accordance with the assignment mechanism

# Randomization-Based Inference

When estimating a sample average causal effect ( $\tau_{SATE}$ ):

- ▶  $X, Y^c, Y^t$  in the sample are regarded as *fixed*
- ▶  $\mathbf{Z}$  is random
- ▶ “Sampling” different values of  $\mathbf{Z}$  for this sample generates variability
  - ▶ From the assignment mechanism
  - ▶ Every value of  $\mathbf{Z}$  gives different set of *observed* outcomes
- ▶ Will dictate a *randomization distribution* that plays a role analogous to a *sampling distribution*
- ▶ Will develop some intuition of how *study design* can dictate *uncertainty of (causal) estimates*

# Randomization-Based Inference

# Unbiasedness Under Completely Randomized Designs

Assignment mechanism:

$$Pr(\mathbf{Z}|\mathbf{X}, \mathbf{Y}^c, \mathbf{Y}^t) = \binom{n}{n_t}^{-1} \text{ if } \sum_{i=1}^n Z_i = n_t$$

and 0 otherwise, where  $n$  is the number of units in the sample and  $n_t$  is the desired number to receive the treatment

An example of an *ignorable* or *unconfounded* treatment design:

$$Z \perp\!\!\!\perp Y^t, Y^c$$

Observed-data comparison  $\hat{\tau} = \frac{1}{n_t} \sum Y_i - \frac{1}{n_c} \sum Y_i$ , is *unbiased* for  $\tau_{PATE}$  and  $\tau_{SATE}$

# How Design can Dictate Uncertainty

The rationale of randomization for unbiasedness is clear, but what about uncertainty?

- ▶ Unbiasedness tells us we should get the “right” answer *on average*
- ▶ Does not guarantee the estimate for any particular random assignment will be close to the truth
  - ▶ Could be “unlucky” and get poor *balance*
  - ▶ Particularly with small samples
- ▶ More “unlucky” randomizations  $\Rightarrow$  more variability in estimates across  $\mathbf{Z} \Rightarrow$  more uncertainty
- ▶ Reducing the chances of “unlucky” randomizations  $\rightarrow$  decrease the variance of effect estimates
- ▶ The comparison between Bernoulli randomization and the Completely Randomized Design is one example

# Fisher's Randomization Inference

- ▶ RA Fisher was the first to grasp the importance of randomization for assessing causal effects (1920s)
- ▶ **Basic idea** is to make inference based *solely on the assignment mechanism*
  - ▶ No modeling assumptions for the potential outcomes
  - ▶ Truly nonparametric
  - ▶ The assignment mechanism completely determines the randomization distribution of any specified test statistic
- ▶ Fisher focused on settings where
  - ▶ The assignment mechanism is **known**
  - ▶ Interest lies in a **sharp null hypothesis** of no causal effect whatsoever
  - ▶ Distinct from being interested in a “typical” or “average” causal effect

# Pieces of Fisher's Exact Method

- ▶ A *known* randomized assignment mechanism
- ▶ Observed data  $\mathbf{Z}$ ,  $\mathbf{X}$ ,  $\mathbf{Y}^{obs}$
- ▶ A *test statistic*,  $T$ 
  - ▶ A function of observed data  $(\mathbf{Y}^{obs}, \mathbf{X})$  and the (stochastic) assignment vector  $(\mathbf{Z})$
- ▶ A *sharp null hypothesis*
  - ▶ A null hypothesis that is sufficiently specific to “fill in” a hypothetical value for each unit's missing potential outcome
- ▶ The *randomization distribution*
  - ▶ Distribution of possible values of the test statistic under all possible assignments according to the assignment mechanism, assuming the null hypothesis is true
- ▶ A *p-value under the randomization distribution given the sharp null hypothesis*
  - ▶ Probability of observing a value of the test statistic at least as extreme under the null hypothesis



# Implementing a Fisher's Exact Test

of the sharp null hypothesis

- ▶ Decide on a test statistic,  $T$
- ▶ Specify the *sharp null hypothesis* of no causal effect whatsoever
  - ▶  $H_0 : Y_i^0 = Y_i^1 \quad \forall i$
  - ▶ Stronger than a hypothesis about the *average* causal effect
  - ▶ Unobserved potential outcome is *implied* for every unit under the sharp null:  $Y_i^{miss} = Y_i^{obs}$
- ▶ Calculate the test statistic in the observed data,  $T^{obs}$
- ▶ “Re-randomize” units from the assignment mechanism
  - ▶ Calculate  $T^{sim}$  for each simulation from the assignment mechanism,  $\mathbf{Z}^{sim}$
- ▶ Compare  $T^{obs}$  to the distribution of  $T^{sim}$ 
  - ▶ Calculate the exact p-value:  $Pr(|T| > T^{obs})$
  - ▶ If  $T^{obs}$  looks very different than most of  $T^{sim}$ ,  $\Rightarrow$  reject  $H_0$

# Fisher's Exact Test: Toy Example

Observed Data, Completely Randomized Experiment

Unit	Z	$Y^0$	$Y^1$
1	0	66	?
2	0	0	?
3	0	0	?
4	1	?	0
5	1	?	607
6	1	?	436

# Fisher's Exact Test: Toy Example

Observed Data, Completely Randomized Experiment

Unit	Z	$Y^0$	$Y^1$
1	0	66	?
2	0	0	?
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4	1	?	0
5	1	?	607
6	1	?	436

► Test statistic,  $T = T(\mathbf{Z}, Y^{obs})$ :

$$\sum_i Z_i Y_i^{obs} / n_t - \sum_i (1 - Z_i) Y_i^{obs} / n_c$$

►  $T^{obs} = 325.6$

# Fisher's Exact Test: Toy Example

One “Re-Randomization”, Completely Randomized Experiment

Unit	$Z^{sim}$	$Y^0$	$Y^1$
1	0	66	?
2	0	0	?
3	1	0	?
4	1	?	0
5	0	?	607
6	1	?	436

- ▶  $H_0 : Y_i^0 = Y_i^1$  for  $i = 1, 2, \dots, 6$
- ▶ Simulate  $\mathbf{Z}$  from the randomization distribution
- ▶ “Fill in” potential outcomes *under the null hypothesis*
- ▶ Recalculate  $T = T(\mathbf{Z}, Y^{obs})$
- ▶  $T^{sim} = -79.0$

Basis of test: Where does  $T^{obs}$  fall in the distribution of would-be values of  $T^{sim}$  under every possible assignment to  $\mathbf{Z}$ , assuming that  $H_0$  is true?

# The Randomization Distribution

$Z_1$	$Z_2$	$Z_3$	$Z_4$	$Z_5$	$Z_6$	$T$
0	0	0	1	1	1	325.6
0	0	1	0	1	1	325.6
0	0	1	1	0	1	-79.0
0	0	1	1	1	0	35.0
0	1	0	0	1	1	325.6
0	1	0	1	0	1	-79.0
0	1	0	1	1	0	35.0
0	1	1	0	0	1	-79.0
0	1	1	0	1	0	35.0
0	1	1	1	0	0	-369.6
1	0	0	0	1	1	369.6
1	0	0	1	0	1	-35.0
1	0	0	1	1	0	79.0
1	0	1	0	0	1	-35.0
1	0	1	0	1	0	79.0
1	0	1	1	0	0	-325.6
1	1	0	0	0	1	-35.0
1	1	0	0	1	0	79.0
1	1	0	1	0	0	-325.6
1	1	1	0	0	0	-325.6

# Test Result

- ▶ How likely (under the randomization distribution) are we to have observed a value of the test statistic at least as large as  $T^{obs}$ ?
- ▶ **Exact p-value:**  $Pr(|T| \geq T^{obs}) = 8/20 = 0.40$  for the sharp null
- ▶ Under the sharp null of absolutely no effect, the observed difference could be due to chance

# Comments and Extensions

- ▶ With large  $n$ , impossible to enumerate all possible  $\mathbf{Z}^{sim}$ 
  - ▶ Simulate a (large) random sample of  $\mathbf{Z}^{sim}$  instead
- ▶ Extensions to other randomization schemes straightforward
  - ▶ If we know the **assignment mechanism** we can always simulate  $\mathbf{Z}^{sim}$  from it
- ▶ Nonzero null:  $H_0 : Y_i^1 - Y_i^0 = \tau$ 
  - ▶ Equivalently,  $H_0 : Y^1 = Y^0 + \tau$
- ▶ Fisher interval estimates
  - ▶ Calculate Fisher p-value under a range of possible  $\tau$
  - ▶ The uncertainty interval for  $\tau$  will be all values for which we fail to reject  $H_0 : Y^0 = Y^1 + \tau$
- ▶ Choice of test statistics
  - ▶ Many options, context/problem dependent