# Modern Sampling Methods

Class 6: Covariate-Adaptive Randomization

January 11, 2022

### Outline

- Randomized Treatment Assignment
  - ► Simple Randomization
  - Efron's Biased Coin Design
  - Forced Balance
- Covariate-Adaptive Randomization / Stratification
  - ► Two-sample t-test
  - Regression with Strata Effects
  - Strata Saturated Regression

## Setup

Units  $i = 1, \ldots, n$ .

Treatment:  $T_i = 0, 1$ .

Potential Outcomes:  $Y_i(0), Y_i(1)$ 

Observed outcome:

$$Y_i = (1 - T_i)Y_i(0) + T_iY_i(1).$$

May also observe covariates  $X_i$  (invariant to treatment).

Average treatment effect:

$$ATE = E[Y_i(1) - Y_i(0)] = E[Y_i(1)] - E[Y_i(0)].$$

## Assignment

Recall: The assignment mechanism is the procedure that determines  $T_i$  given  $Y_i(0)$  and  $Y_i(1)$  (and given  $X_i$  if we also have covariates).

The assignment mechanism is key to understanding how observed outcomes (and treatments) are generated by potential outcomes.

## Assignment Mechanisms

Let 
$$W^{(n)} = \{(Y_i(0), Y_i(1), X_i)), i = 1, ..., n\}, T^{(i-1)} = (T_{i-1}, ..., T_1), \text{ and } T^{(0)} = \emptyset.$$

- 1. Simple/Complete/Pure Randomization ("Coin Flipping")
  - $ightharpoonup T_1, \ldots, T_n$  i.i.d.

• 
$$Pr(T_i = 1 | T^{(i-1)}, W^{(n)}) = Pr(T_i = 1) = p$$

- $Cov(T_i, T_j) = 0, \quad i \neq j$
- 2. Restricted Randomization
  - Allows for dependence across units in randomization assignment

$$Pr(T_i = 1 | T^{(i-1)}, W^{(n)}) = Pr(T_i = 1 | T^{(i-1)})$$

- $\quad \mathsf{Cov}(T_i,T_j) \neq 0, \quad i \neq j$
- 3. Covariate-Adaptive Randomization (Stratification)
  - Allows assignment to depend on baseline covariates

$$Pr(T_i = 1 | T^{(i-1)}, W^{(n)}) = Pr(T_i = 1 | X^{(i)}, T^{(i-1)})$$

- 4. Response-Adaptive Randomization
  - ▶ Allows assignment to depend on "earlier" outcomes

$$Pr(T_i = 1 | T^{(i-1)}, W^{(n)}) Pr(T_i = 1 | X^{(i)}, Y^{(i-1)}, T^{(i-1)})$$



# Simple Randomization

- Assignment Rule:  $Pr(T_i = 1) = \frac{1}{2}$
- ▶ Independence:  $Var(T_i) = \frac{1}{4}$ ,  $Cov(T_i, T_j) = 0$
- ▶ Balance: Large Sample vs. Small Sample
- ▶ Degree of Imbalance = (#Treated) (Target #Treated)

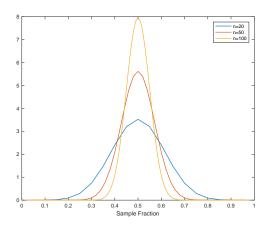
$$D_n = \left(\sum_{i=1}^n T_i\right) - p \cdot n$$

$$\left(\text{setting } p = \frac{1}{2}\right) = \sum_{i=1}^n \left(T_i - \frac{1}{2}\right)$$

$$= \frac{1}{2} \left(\#\text{Treated} - \#\text{Control}\right)$$

# Degree of Imbalance under Simple Randomization

$$\Pr(D_n = d) = rac{inom{n}{d + rac{n}{2}}}{2^n} \;,\;\; d = -rac{n}{2}, -rac{n}{2} + 1, \dots, rac{n}{2}$$



# Approximate Distribution for Degree of Imbalance under Simple Randomization

Since degree of imbalance is a sum over i.i.d. random variables,  $D_n = \sum_{i=1}^n \left(T_i - \frac{1}{2}\right)$ , we can approximate its behavior by a normal distribution,

$$\frac{1}{\sqrt{n}}D_n \ \stackrel{d}{\longrightarrow} \ N\left(0\ , \frac{1}{4}\right)$$

so, for large sample sizes,

$$\Pr(|D_n| \ge r) \approx 2\left[1 - \Phi\left(\frac{2r}{\sqrt{n}}\right)\right]$$

## Effect of Imbalance on Power

Recall  $H_0: ATE = 0$  vs.  $H_1: ATE \neq 0$ .

where

$$ATE = \alpha_1 - \alpha_0$$

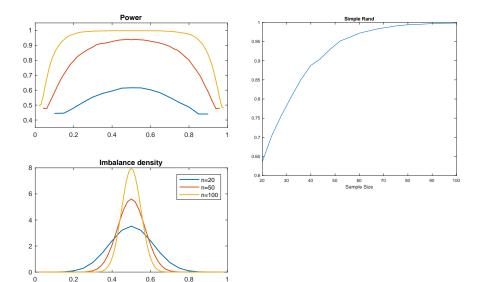
$$= E[Y_i \mid T_i = 1] - E[Y_i \mid T_i = 0]$$

$$\hat{\beta} = \overline{Y}_1 - \overline{Y}_0$$

Two Sample *t*-test: 
$$t = \frac{\beta}{\sqrt{\frac{\hat{\sigma}_1^2}{n_1} + \frac{\hat{\sigma}_0^2}{n_0}}}$$

 $\Rightarrow$  Reject if |t| > 1.96

## Simple Randomization - Power Simulation



Sample Fraction

# Efron's Biased Coin Design

Assignment Rule:

$$\Pr(T_i = 1 | T^{(i-1)}) = \begin{cases} \frac{1}{2} & \text{if } D_{i-1} = 0\\ q_e & \text{if } D_{i-1} < 0\\ 1 - q_e & \text{if } D_{i-1} > 0 \end{cases}$$

where 
$$q_e \in \left[\frac{1}{2},1\right]$$
, and  $D_{i-1} = \left(\sum_{j=1}^{i-1} T_j\right) - \frac{i-1}{2}$ 

- Special Cases:  $q_e=rac{1}{2}$ : Simple Randomization  $q_e=1$ : "Pairwise" Randomization"  $q_e=rac{2}{3}$ : Efron's suggested value
- $E(T_i) = \frac{1}{2}$ ,  $Var(T_i) = \frac{1}{4}$
- $ightharpoonup \Sigma = Var(T)$  and (Exact) distribution of  $D_n$  see Markaryan and Rosenberger (2010)



# Approximate Distribution for Degree of Imbalance under Efron's BC

•  $q_e = \frac{1}{2}$  Independent observations (Simple Randomization)

$$\frac{1}{\sqrt{n}}D_n \ \stackrel{d}{\longrightarrow} \ N\left(0\ , \frac{1}{4}\right)$$

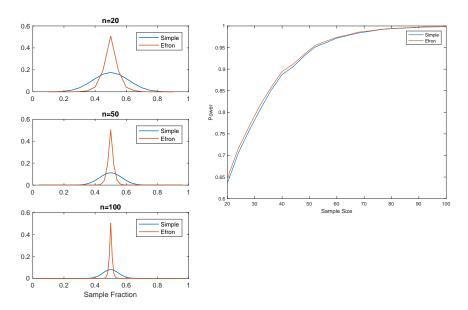
•  $q_e > \frac{1}{2}$  Dependent observations

$$\frac{1}{\sqrt{n}}D_n \stackrel{p}{\longrightarrow} 0$$

In fact,  $D_n = 0$  with positive probability for large (even) sample sizes.

$$\Pr(D_n = 0 | n \text{ even}) \longrightarrow \frac{2q_e - 1}{q_e} = \frac{1}{2} \left( \text{when } q_e = \frac{2}{3} \right)$$

## Efron's Biased Coin - Power Simulation



### Forced Balance Randomization

- Assignment Rule:  $\binom{n}{n/2}$  equally likely arrangements with n/2 assigned T=1 and n/2 assigned T=0. (n even)
- Example. Sequential allocation rule

  Urn: n/2 balls labeled T (or 1) and n/2 balls labeled C (or 0)

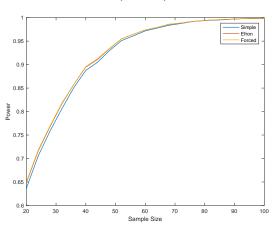
$$\Pr(T_i = 1 | T^{(i-1)}) = \frac{\frac{n}{2} - \sum_{j=1}^{i-1} T_j}{n - (i-1)}$$

- ►  $E(T_i) = \frac{1}{2}$ ,  $Var(T_i) = \frac{1}{4}$ , Dependence:  $Cov(T_i, T_j) = -\frac{1}{4(n-1)}$
- $\triangleright D_n \dots$

Simple: 
$$\frac{1}{\sqrt{n}}D_n \stackrel{d}{\longrightarrow} N\left(0, \frac{1}{4}\right)$$

Efron's BC:  $D_n \stackrel{p}{\longrightarrow} 0$ 

Forced:  $D_n = 0$  (*n* even)



#### Baseline Covariate Stratification

- Construct strata
- 2. Apply randomization within strata

Stratification: Partition covariate space into a finite number of strata.

e.g. 
$$X_{1,i} = \text{region} \in \{1,2,3,4\}$$
 (N, S, E, W)  $X_{2,i} = \text{income category} \in \{0,1\}$  (low, high) Strata  $\mathscr{S} = \{(1,0),(1,1),(2,0),(2,1),(3,0),(3,1),(4,0),(4,1)\}$   $|\mathscr{S}| = 4 \times 2 = 8$ 

Let S denote the map from covariate values into strata values,  $S(X_i) = S_i \in \mathscr{S}$ 

For each strata  $s \in \mathcal{S}$ , apply randomization ((e.g. Simple, Efron's BC, Forced Balance) with target fraction of treated p(s).

Assignment Rule:

$$Pr(T_i = 1 | S^{(i)}, T^{(i-1)})$$

Degree of Imbalance for each strata s:

$$D_n(s) = \sum_{i=1}^n (T_i - p(s)) \mathbf{1} \{S_i = s\}$$

Suppose p = p(s).

$$\left\{\frac{1}{\sqrt{n}}D_n(s):\ s\in\mathscr{S}\right\}\stackrel{d}{\longrightarrow} N\big(0,\ \tau\ \Omega_D\big)$$

where  $\Omega_D = \mathsf{diag}\left(\mathsf{Pr}(S_i = 1), \dots, \mathsf{Pr}(S_i = |\mathscr{S}|)\right)$ 

- ▶ Simple Randomization:  $\tau = p(1-p)$
- Efron's Biased Coin Design:  $\tau = 0$
- ▶ Forced Balance based on  $\binom{n(s)}{[pn(s)]}$  re-arrangments:  $\tau = 0$

au= 0: less imbalance, but more dependence in observations

Behavior of tests for ATE=0 will depend on the value of au

### Inference on ATE with Covariates

Suppose  $(Y_i(0), Y_i(1), X_i)$  i.i.d. (across i). Then stratified randomization implies unconfoundedness w.r.t. strata:

$$(Y_i(0), Y_i(1)) \perp T_i | S_i$$

$$H_0: ATE = 0 \text{ vs. } H_1: ATE \neq 0.$$

- ► Two Sample *t*-test
- Regression with Strata Effects
- Saturated Regression

see Bugni, Canay, and Shaikh (2018, 2019)

## Two Sample *t*-test

$$t=rac{\overline{Y}_1-\overline{Y}_0}{\sqrt{rac{\hat{\sigma}_1^2}{n_1}+rac{\hat{\sigma}_0^2}{n_0}}}\;, \qquad ext{Reject if } |t|>1.96$$

Regression equivalent:

$$Y_i = \gamma + \beta T_i + \varepsilon_i$$

t-test for  $\beta = 0$  with heteroskedasticity-robust standard errors

Result:

(i) Under  $H_0$ ,

$$t \stackrel{d}{\longrightarrow} N(0, \sigma_{t-test}^2)$$

where  $\sigma_{t-test}^2 \leq 1$ ;



#### Result: (cont'd)

- (ii) Moreover, the test is conservative,  $\sigma_{t-test}^2 < 1$ , unless:
  - No stratification
     So, for all previous simple and restricted randomization cases (ignoring covariates), two sample t-test is not conservative
     or -
  - ▶  $E[Y_i(t)|S_i] = E[Y_i(t)]$  for t = 0, 1So, two sample t-test is *not* conservative when, there's no (mean) heterogeneity by strata
  - Assignment by simple randomization
    So, stratified block randomization (forced balance by strata), two sample t-test will be too conservative

# Regression with Strata Effects

$$Y_i = \beta T_i + \sum_{s \in \mathscr{S}} \gamma_s \mathbf{1} \{ S_i = s \} + \varepsilon_i^{se}$$

- $\beta = ATE$
- ▶ t-test for  $\beta = 0$  with heteroskedasticity-robust standard errors

$$t^{\mathsf{se}} = rac{\hat{eta}}{\mathsf{std} \; \mathsf{err}(\hat{eta})}$$

- "Within" estimation (deviations from strata means)
  - ▶ Same  $\hat{\beta}$
  - ▶ Different std  $err(\hat{\beta})$ , in general

#### Result:

(i) Under  $H_0$ ,

$$t^{se} \stackrel{d}{\longrightarrow} N(0, \sigma_{t^{se}}^2)$$

where  $\sigma_{t^{se}}^2 \leq 1$ ;

- (ii) Moreover, the test is conservative,  $\sigma_{t^{\rm se}}^2 < 1$  unless
  - As before, no stratification, no heterogeneity by strata, simple randomization
    - or -
  - ▶  $p = \frac{1}{2}$ Strata effects has an additional condition leading to non-conservativeness. This condition  $p = \frac{1}{2}$  is common.

#### Notes:

- ▶ If  $\tau = 0$ ,  $\sigma_{t^{se}}^2 = \sigma_{t-test}^2$ . So, two-sample *t*-test and strata effects regression have same power (asymptotically).
- ▶ Imbens-Rubin (2015) consider other stratified block randomization designs and show  $\hat{\beta}$  not consistent

## Strata Saturated Regression

- ▶ p(s) = p: Two Sample *t*-test and Regression with Strata Effects
- $\triangleright$  p(s) possibly nonconstant: Saturated Regression

$$Y_i = \sum_{s \in \mathscr{S}} \gamma_s \mathbf{1}\{S_i = s\} + \sum_{s \in \mathscr{S}} \theta_s \mathbf{1}\{T_i = 1, S_i = s\} + \varepsilon_i^{sat}$$

- $\theta_s = ATE(s)$  and  $\beta = ATE = \sum_{s \in \mathscr{S}} \theta_s \Pr(S_i = s)$
- $\blacktriangleright$  *t*-test for  $\beta$  with heteroskedasticity-robust standard errors

$$\hat{\beta} = \sum_{s \in \mathscr{S}} \hat{\theta}_s \left( \frac{\sum_{i=1}^n \mathbf{1}\{S_i = s\}}{n} \right) \quad \text{and} \quad t^{se} = \frac{\hat{\beta}}{\textit{std err}(\hat{\beta})}$$



#### Result:

(i) Under  $H_0$ ,

$$t^{sat} \stackrel{d}{\longrightarrow} N(0, \sigma_{t^{sat}}^2)$$

where  $\sigma_{t^{sat}}^2 \geq 1$ ;

- (ii) Moreover, the test is invalid (does not control size),  $\sigma_{t^{\rm sat}}^2 > 1$  unless
  - No stratification (ignoring covariates)- or -
  - No heterogeneity by strata:  $E[Y_i(1) - Y_i(0)|S_i] = E[Y_i(1) - Y_i(0)]$

#### Notes:

- ► For two sample *t*-test and strata effects regression, the usual heteroskedasticity-robust variance estimator is too *large*, in general.
  - For saturated regression, the usual heteroskedasticity-robust variance estimator is too *small*, in general.
- ▶ When  $\tau=0$  and p(s)=p, all three tests have the same asymptotic power. But, when  $\tau>0$ , the saturated regression approach is most powerful.

## Simple Randomization - Local Power Simulation

