

# **9.S918: Statistical Inference in Brain and Cognitive Sciences**

**Week 2 Day 1: Causal inference, continued**

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# Potential-outcomes framework: brief recap

- In epidemiology and many other areas of statistics, causal inference was developed out of the idea of **potential outcomes** (Neyman 1923, Rubin 1974)
- Consider an outcome,  $Y$ , and a potential **treatment**  $A$
- **Example:**  
 $Y$ : an individual survives to the end of the year (0: no, 1: yes)  
 $A$ : an individual with heart disease receives a heart transplant (0: no, 1: yes)

# Potential-outcome random variables

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- Suppose that  $A$  is discrete; for this case,  $A \in \{0,1\}$
- The **potential outcomes**, or **counterfactual outcomes**, are random variables for  $Y$  for each potential value of  $A$

$Y^{a=0}$                       The value that  $Y$  would take if  $A$  were 0

$Y^{a=1}$                       The value that  $Y$  would take if  $A$  were 1

- **Counterfactual risk** is the **expected value** of each counterfactual-outcome random variable:

$$E[Y^{a=0}]$$

$$E[Y^{a=1}]$$

- Expected value, or expectation, is defined as follows:

$$E[X] = \sum_x xP(X = x)$$

- So we are interested in (and likewise for  $Y^{a=1}$ ):

$$E[Y^{a=0}] = \sum_y yP(Y^{a=0} = y) = 0 \times P(Y^{a=0} = 0) + 1 \times P(Y^{a=0} = 1) = \boxed{P(Y^{a=0} = 1)}$$

# Counterfactual data and causal effects

- Suppose we knew **what would happen** for each individual in the population under each value of the treatment
- Then we could compute the counterfactual risks:

$$E[Y^{a=0}] = 0.5 \qquad E[Y^{a=1}] = 0.5$$

- The **average causal effect** of treatment  $A$  is defined as the difference of counterfactual risks:

$$E[Y^{a=1}] - E[Y^{a=0}] = 0$$

- Here, treatment is **ineffective**

(Hernan & Robins, 2020, Table 1.1)

	$Y^{a=0}$	$Y^{a=1}$
Rheia	0	1
Kronos	1	0
Demeter	0	0
Hades	0	0
Hestia	0	0
Poseidon	1	0
Hera	0	0
Zeus	0	1
Artemis	1	1
Apollo	1	0
Leto	0	1
Ares	1	1
Athena	1	1
Hephaestus	0	1
Aphrodite	0	1
Cyclope	0	1
Persephone	1	1
Hermes	1	0
Hebe	1	0
Dionysus	1	0
$P(Y^{a=*}) = 1$	0.5	0.5

# Estimating causal effects

Remember,  $E[Y^{a=i}] = P(Y^{a=i} = 1)$

	$L$	$A$	$Y$	$Y^0$	$Y^1$
Rheia	0	0	0	0	?
Kronos	0	0	1	1	?
Demeter	0	0	0	0	?
Hades	0	0	0	0	?
Hestia	0	1	0	?	0
Poseidon	0	1	0	?	0
Hera	0	1	0	?	0
Zeus	0	1	1	?	1
Artemis	1	0	1	1	?
Apollo	1	0	1	1	?
Leto	1	0	0	0	?
Ares	1	1	1	?	1
Athena	1	1	1	?	1
Hephaestus	1	1	1	?	1
Aphrodite	1	1	1	?	1
Polyphemos	1	1	1	?	1
Persephone	1	1	1	?	1
Hermes	1	1	0	?	0
Hebe	1	1	0	?	0
Dionysus	1	1	0	?	0

- Naively, we might estimate the counterfactual risks  $P(Y^{a=i} = 1)$  directly from observed  $A$  and  $Y$ :

$$\hat{P}_{MLE}(Y = 1 | A = 0) = \frac{3}{7} \quad \hat{P}_{MLE}(Y = 1 | A = 1) = \frac{7}{13}$$

- But under what circumstances  $\hat{P}_{MLE}(Y | A = i) = \hat{P}_{MLE}(Y^{a=i} = 1)$ ?

- The following is certainly true:

$$\hat{P}_{MLE}(Y = 1 | A = i) = \frac{\text{Count}(Y = 1 \wedge A = i)}{\text{Count}(A = i)}$$

**CONSISTENCY:** when  
 $A = i, Y = Y^{a=i}$

$$= \frac{\text{Count}(Y^{a=1} = 1 \wedge A = i)}{\text{Count}(A = i)}$$

$$= \hat{P}_{MLE}(Y^{a=i} = 1 | A = i)$$

*Crucial step;  
make sure you  
understand it!*

- So, the following condition suffices:

$$P(Y^{a=i} = 1 | A = i) = P(Y^{a=i} = 1)$$

- This is called **EXCHANGEABILITY**:

$$Y^a \perp A | \{ \}$$

# Exchangeability and randomization

Goal:  $\hat{P}(Y^a = 1)$

- Why is a randomized experiment so powerful?
- Recap of exchangeability criterion:

$$Y^a \perp A \mid \{\}$$

- If we ourselves determine  $A$  in a way that is *truly blind* to  $Y^a$ , it **imposes** exchangeability!
- We can now go ahead and estimate

$$\hat{P}(Y^{a=i} = 1) = \hat{P}(Y = 1 \mid A = i)$$

- Hooray!!!

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Rheia  
Kronos  
Demeter  
Hades  
Hestia  
Poseidon  
Hera  
Zeus  
Artemis  
Apollo  
Leto  
Ares  
Athena  
Hephaestus  
Aphrodite  
Polyphemus  
Persephone  
Hermes  
Hebe  
Dionysus

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# Does loss of randomization make things hopeless?

- In the real world, many datasets are **not** randomized this way
- **Example:** let's imagine some other variable that might affect whether treatment  $A$  is applied; e.g.,  $L$  = whether the patient was in critical condition (1=yes, 0=no)
- In general,  $L$  will be related to  $Y^a$ 
  - E.g., in this example, patients in critical condition are surely more likely to die overall!

$A \perp Y^a \mid \{ \}$



	$L$
Rheia	0
Kronos	0
Demeter	0
Hades	0
Hestia	0
Poseidon	0
Hera	0
Zeus	0
Artemis	1
Apollo	1
Leto	1
Ares	1
Athena	1
Hephaestus	1
Aphrodite	1
Polyphemus	1
Persephone	1
Hermes	1
Hebe	1
Dionysus	1



# Conditional exchangeability

- But now suppose we have observed (i.e., it's in our dataset) the factor  $L$  that affected whether the treatment  $A$  was applied
- If the following condition holds, it can help us estimate the counterfactual risks  $P(Y^a)$ :

$$A \perp Y^a \mid L$$

- That is,  $L$  captures all the information available in  $A$  that is relevant to all  $Y^a$
- This is called **CONDITIONAL EXCHANGEABILITY**

	$L$	$A$	$Y^{a=0}$	$Y^{a=1}$	$Y$
Rheia	0	0	0	1	0
Kronos	0	0	1	0	1
Demeter	0	0	0	0	0
Hades	0	0	0	0	0
Hestia	0	1	0	0	0
Poseidon	0	1	1	0	0
Hera	0	1	0	0	0
Zeus	0	1	0	1	1
Artemis	1	0	1	1	1
Apollo	1	0	1	0	1
Leto	1	0	0	1	0
Ares	1	1	1	1	1
Athena	1	1	1	1	1
Hephaestus	1	1	0	1	1
Aphrodite	1	1	0	1	1
Polyphemus	1	1	0	1	1
Persephone	1	1	1	1	1
Hermes	1	1	1	0	0
Hebe	1	1	1	0	0
Dionysus	1	1	1	0	0



# Using conditional exchangeability

- Can we estimate  $P(Y^{a=i} = 1 | L)$ ?
- It turns out we can!

$$P(Y^{a=i} = 1 | L) = P(Y^{a=i} = 1 | L, A) \quad \text{CONDITIONAL EXCHANGEABILITY}$$

$$\hat{P}_{\text{MLE}}(Y^{a=i} = 1 | L = j, A = k) = \frac{\text{Count}(Y^{a=i} = 1, L = j, A = k)}{\text{Count}(L = j, A = k)}$$

- In estimating this condprob:
  - when  $i = k$  we use CONSISTENCY
  - WHEN  $i \neq k$  we have "missing data", so ignore those instances

CONSISTENCY: when  $A = i, Y = Y^{a=i}$

	$L$	$Y$
Rheia	0	0
Kronos	0	1
Demeter	0	0
Hades	0	0
Hestia	0	0
Poseidon	0	0
Hera	0	0
Zeus	0	1
Artemis	1	1
Apollo	1	1
Leto	1	0
Ares	1	1
Athena	1	1
Hephaestus	1	1
Aphrodite	1	1
Polyphemus	1	1
Persephone	1	1
Hermes	1	0
Hebe	1	0
Dionysus	1	0

$L$	$A$	$\text{Count}(L, A)$	$\text{Count}(Y^{a=0} = 1, L, A)$	$\text{Count}(Y^{a=1} = 1, L, A)$	$\hat{P}_{\text{MLE}}(Y^{a=i} = 1   L)$
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0	0	4	1	?	1/4
0	1	4	?	1	1/4
1	0	3	2	?	2/3
1	1	9	?	6	2/3

*This is just like estimating  $P(Y | L, A)$ !*

# Using conditional exchangeability

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- We originally characterized our goal as estimating the **counterfactual risks**  $E[Y^{a=i}] = P(Y^{a=i} = 1)$
- With conditional exchangeability, we estimated  $P(Y^{a=i} = 1 | L)$  – these are called **stratum-specific risks** (where each **stratum** is a value of  $L$ )
- Often, this may be all you need or want
  - If the causal effect of  $A$  depends on  $L$ , then "summarizing out"  $L$  discards information!
- But there are situations where the basic counterfactual risk  $E[Y^{a=i}]$  may be of interest
  - e.g., "how many lives would it save if everyone who came to the hospital with heart disease received a heart transplant?"
- But we can recover the basic counterfactual risks through **standardization** (or the mathematically equivalent **inverse probability weighting**)

# Standardization

$L$  = whether the patient was in critical condition (1=yes, 0=no)

$L$	$A$	$\hat{P}_{MLE}(Y^{a=i}   L)$
0	0	1/4
0	1	1/4
1	0	2/3
1	1	2/3

- By the law of total probability,

*We have just estimated this*

$$P(Y^{a=i}) = \sum_j P(Y^{a=i} | L) P(L)$$

*We can estimate this from the data, too*

- Expanding the sum and plugging in our estimates we get:

	$L$	$A$	$Y$	$Y^0$	$Y^1$
Rheia	0	0	0	0	?
Kronos	0	0	1	1	?
Demeter	0	0	0	0	?
Hades	0	0	0	0	?
Hestia	0	1	0	?	0
Poseidon	0	1	0	?	0
Hera	0	1	0	?	0
Zeus	0	1	1	?	1
Artemis	1	0	1	1	?
Apollo	1	0	1	1	?
Leto	1	0	0	0	?
Ares	1	1	1	?	1
Athena	1	1	1	?	1
Hephaestus	1	1	1	?	1
Aphrodite	1	1	1	?	1
Polyphemus	1	1	1	?	1
Persephone	1	1	1	?	1
Hermes	1	1	0	?	0
Hebe	1	1	0	?	0
Dionysus	1	1	0	?	0

$$P(Y^{a=i} = 1) = P(Y^{a=i} = 1 | L = 0)P(L = 0) + P(Y^{a=i} = 1 | L = 1)P(L = 1)$$

$$= \frac{1}{4} \times \frac{2}{5} + \frac{2}{3} \times \frac{3}{5}$$

$$= \frac{1}{10} + \frac{2}{5}$$

$$= \frac{5}{10} = \frac{1}{2}$$

(Because  $\hat{P}_{MLE}(Y^{a=i} | L)$  are the same for  $a = 0$  and  $a = 1$ , this work gives us the result for both counterfactual treatments, and the risk ratio is 1)

$$\hat{P}_{MLE}(L = 1) = \frac{12}{20} = \frac{3}{5}$$

# IDENTIFIABILITY of causal effects

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- IDENTIFIABILITY means, our assumptions allow the causal effect we are interested in to be ***uniquely estimated*** from the available data (set of observed/measured variables)
  - "Uniquely estimate": if we had an arbitrary large quantity of data, we could estimate the causal effect with arbitrarily high accuracy and precision
- Simple case of unidentifiability: Hernan & Robins's heart transplant example, if  $L$  (severity of disease) affects probability of a heart transplant and we don't measure it
  - Suppose that people with transplants have lower survival rates:  $\hat{P}_{MLE}(Y|A = 1) < \hat{P}_{MLE}(Y|A = 0)$ 
    - Could be because heart transplants are dangerous
    - Or: sicker people are more likely to get transplants!

*Not an issue of sample size—no amount of data would help!*

# The three criteria for identifiability

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- **Consistency:**  $Y = Y^{a=i}$  whenever  $A = i$ 
  - Consequence: different individuals' outcomes don't affect each other
  - Consequence: there can be no "multiple versions" of the same treatment  $A$  in terms of their influence on  $Y$
- **Conditional Exchangeability:** for all  $i$ ,  $Y^{a=i} \perp A \mid Z$  for some set of observed variables  $Z$ 
  - Consequence: there can be no "hidden common causes" or "hidden mediators" of  $A$  and  $Y^{a=i}$
- **Positivity:** for all  $i$  and all values of  $Z$ ,  $P(A = i \mid Z) > 0$ 
  - e.g., in our example, it can't be the case that individuals with heart disease are *always* given transplants
- If all three criteria hold, we can estimate causal effects

# Summary of intro to potential outcomes

- The potential outcomes framework formalizes causal effects (or risks) through **counterfactual outcome** (also called **potential outcome**) **variables**
- At most one counterfactual outcome is observable in each datum, so causal effects cannot in general be naively estimated from data

	$L$	$A$	$Y$	$Y^0$	$Y^1$
Rheia	0	0	0	0	?
Kronos	0	0	1	1	?
Demeter	0	0	0	0	?
Hades	0	0	0	0	?
Hestia	0	1	0	?	0
Poseidon	0	1	0	?	0

- However, if the three following conditions hold, the data can be viewed as a **conditionally randomized experiment** and causal effects can be estimated

**Consistency**

$$Y = Y^{a=i} \text{ whenever } A = i$$

**Conditional Exchangeability**

$$\exists Z. \forall i. Z \text{ is observed} \\ \wedge Y^{a=i} \perp A | Z$$

**Positivity**

$$\forall i. \forall Z. P(A = i | Z) > 0$$

- This analysis also sheds light on the power of randomized experiments: they offer **unconditional exchangeability**