

**MATH-449 - Biostatistics**  
**EPFL, Spring 2022**  
**Problem Set 1**

1. Determine whether each of these questions are phrased as causal questions or not (yes or no).
  - a) Does the Moderna vaccine reduce the risk of severe COVID-19 infection?
  - b) Do women with breast cancer survive longer than men with prostate cancer?
  - c) Is the life expectancy in Switzerland longer than the life expectancy in Italy?
  - d) Does drinking 0.5 L beer compared to 0.5 L Coca Cola at 19h00 affect the quality of sleep?
  - e) Would drinking a cup of coffee 2 hours before your exam improve your performance?
2. Based on the definition of a causal effects in the lecture slides, argue whether the following statements about a covariate  $L \in \mathbb{R}$ , a treatment  $A = 0, 1$  and an outcome  $Y \in \mathbb{R}$  are right or wrong (there is no guarantee that  $A$  is randomly assigned).
  - a)  $\mathbb{E}(Y^{a=1} | L = l) - \mathbb{E}(Y^{a=0} | L = l)$  is a causal effect .
  - b)  $\mathbb{E}(Y | A = 1, L = l) - \mathbb{E}(Y | A = a, L = l)$  is a causal effect .
  - c)  $\mathbb{E}(Y^{a=1} | A = 1, L = l) - \mathbb{E}(Y^{a=0} | A = 1, L = l)$  is a causal effect .
  - d)  $\frac{\mathbb{E}(Y^{a=1})}{\mathbb{E}(Y^{a=0})}$  is an average over individual level (additive) causal effects.
3. Translate these English sentences to mathematical (counterfactual) statements.
  - a) The average causal effect of receiving a COVID-19 vaccine ( $A = 1$ ) vs placebo ( $A = 0$ ) on mortality after one year ( $Y = 1$  is death,  $Y = 0$  is alive) in the entire population of interest.
  - b) The average causal effect of receiving a COVID-19 vaccine ( $A = 1$ ) vs placebo ( $A = 0$ ) on mortality after one year ( $Y = 1$  is death,  $Y = 0$  is alive) among those who received placebo in the observed (factual) data.
  - c) The average causal effect of receiving a COVID-19 vaccine ( $A = 1$ ) vs placebo ( $A = 0$ ) on mortality after one year ( $Y = 1$  is death,  $Y = 0$  is alive) among those who received treatment in the observed (factual) data.
  - d) The average causal effect of receiving a COVID-19 vaccine ( $A = 1$ ) vs placebo ( $A = 0$ ) on mortality after one year ( $Y = 1$  is death,  $Y = 0$  is alive) in men ( $X = 1$ ).
  - e) Are your answers in a)-d) estimands, estimators or estimates?
4. Suppose investigators had access to data from a study in which they observed for each patient a binary outcome  $Y$ , a binary treatment  $A$  and a 4-level baseline covariate  $L$ . The parameters of the joint density of  $(L, A, Y)$  were computed from the data and summarized in Table 1 (where we suppose that the sample size was so large, that sampling variability is not a concern).
  - a) From the parameters in Table 1, compute  $\mathbb{E}[Y]$ .
  - b) Suppose now that the data did not in fact arise from a regular observational study, but had instead come from a special trial. Upon recruitment into the study, each patient's covariate  $L$  is measured and then they are sorted into groups based on that covariate's value. In each group, the investigators conduct a separate experiment, which are identical except they use a special coin to randomize patients to either treatment ( $a = 1$ ) or control ( $a = 0$ ), with "heads" corresponding to treatment and "tails" corresponding to control. The probabilities for heads for each of these sub-trials is given by the column labeled  $P(A = 1 | L = l)$ . Assume consistency holds ( $Y^A = Y$ ), and that patients perfectly complied with their assignments. With the information in the table, compute the effect of treatment  $\mathbb{E}[Y^{a=1} - Y^{a=0} | L = l]$  for each subgroup  $L = l$  that was targeted in each of the sub-trials. What additional assumptions did you use along the way, that was justified given the source of the data?

- c) From the quantities computed in part a), use laws of probability to compute the average treatment effect, among the whole population,  $\mathbb{E}[Y^{a=1} - Y^{a=0}]$ .
- d) Draw a directed acyclic graph (DAG) that could depict the mechanism that generated the observed data.
- e) The data analyst for the study approaches you and said they made a terrible mistake: when preparing the column  $P(A = 1 | L = l)$  in Table 1, they reverse coded the treatment variable, so in fact the true values of the treatment propensities are 1 minus those listed in the table. What will be the values of the previously computed parameters, and explain in words why these changes did (or did not occur).

	$P(Y = 1   A = a, L = l)$	$P(A = 1   L = l)$	$P(L = l)$
	$a = 1$	$a = 0$	
$l = 1$	.1	.8	.2
$l = 2$	.2	.7	.4
$l = 3$	.3	.6	.6
$l = 4$	.4	.5	.1

Table 1: Parameters of  $P_{L,A,Y}$  observed in the conditionally randomized trial.

5. Consider a covariate  $L \in \mathbb{R}$ , a treatment  $A = 0, 1$  and an outcome  $Y \in \mathbb{R}$ .

- a) Investigator 1 claims that  $A \perp\!\!\!\perp Y \implies A \perp\!\!\!\perp Y | L$ . Show that the statement is wrong.
- b) Investigator 2 claims that  $A \perp\!\!\!\perp Y | L \implies A \perp\!\!\!\perp Y$ . Show that the statement is wrong.