Causal inference in health disparities research and the "No-Multiple-Versions-of-Treatment" assumption

"When I use a word, it means just what I choose it to mean—neither more nor less.—"

Humpty Dumpty, Through the Looking Glass by Lewis Carroll

Health disparities research investigates causes of health differences across populations, often relying on observational data for causal inference. The Neyman-Rubin potential outcomes framework defines causal effects as the difference between two potential outcomes (what-if scenarios). It relies on a set of identification assumptions (Figure 1).

To avoid a violation of this assumption, it is important that regardless of which component of SES is changed, the same effect would occur

Exchangeability (no confounding)

Positivity (non-zero probability of receiving treatment) Stable Unit Treatment Value Assumption (SUTVA)

- No interference
- No-multiple-versionsof-the-treatment

Figure 1. Identification assumptions under the potential outcomes framework.

The no-multiple-versions-of-the-treatment assumption implies that:

"... the exposure (treatment) specified in the analysis must have enough precision that any variation within the exposure specification would not result in a different outcome." [1]

Less of a concern in randomized controlled trials because the intervention is pre and welldefined, i.e., the intervention means just what the researchers choose it to mean neither more nor less.

In health disparities research, the studied exposures often lead to violations of this assumption. However, this is seldom discussed in applications.

In the example, we show multiple versions of the exposure *Medium* 5 SES. The issue arises when each version could lead to a different outcome, i.e., when $Y_{v1} \neq Y_{v2}$.

From previous research, it is known that different components of SES have a different impact on health outcomes. Thus, each version is likely to lead to a different outcome, i.e., SES as exposure is not well-defined.

Main consequences:

- We would not know which causal contrast is being estimated from the data, i.e., which causal effect, if any, is being estimated. Difficult to justify the use of estimates for decision-making.^[2]
- Different versions might require different sets of covariates for adjustment, opening back door paths we are unaware of.
- Violations of this assumption could explain (in part) the heterogeneous results obtained in health inequalities research and replicability issues.

EXAMPLE: Socioeconomic status (SES)

Looking into the effect of SES on a health outcome Y, specifically the contrast between a scenario where the SES level is set to *Medium* vs the *Low* SES scenario.

SES is often measured as a composite index containing information on Education,



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Income and Occupation. SES is a "compound treatment".

A Medium SES score could be achieved in various ways, which leads to various "hidden" versions of the Medium SES exposure level as seen in Figure 2.

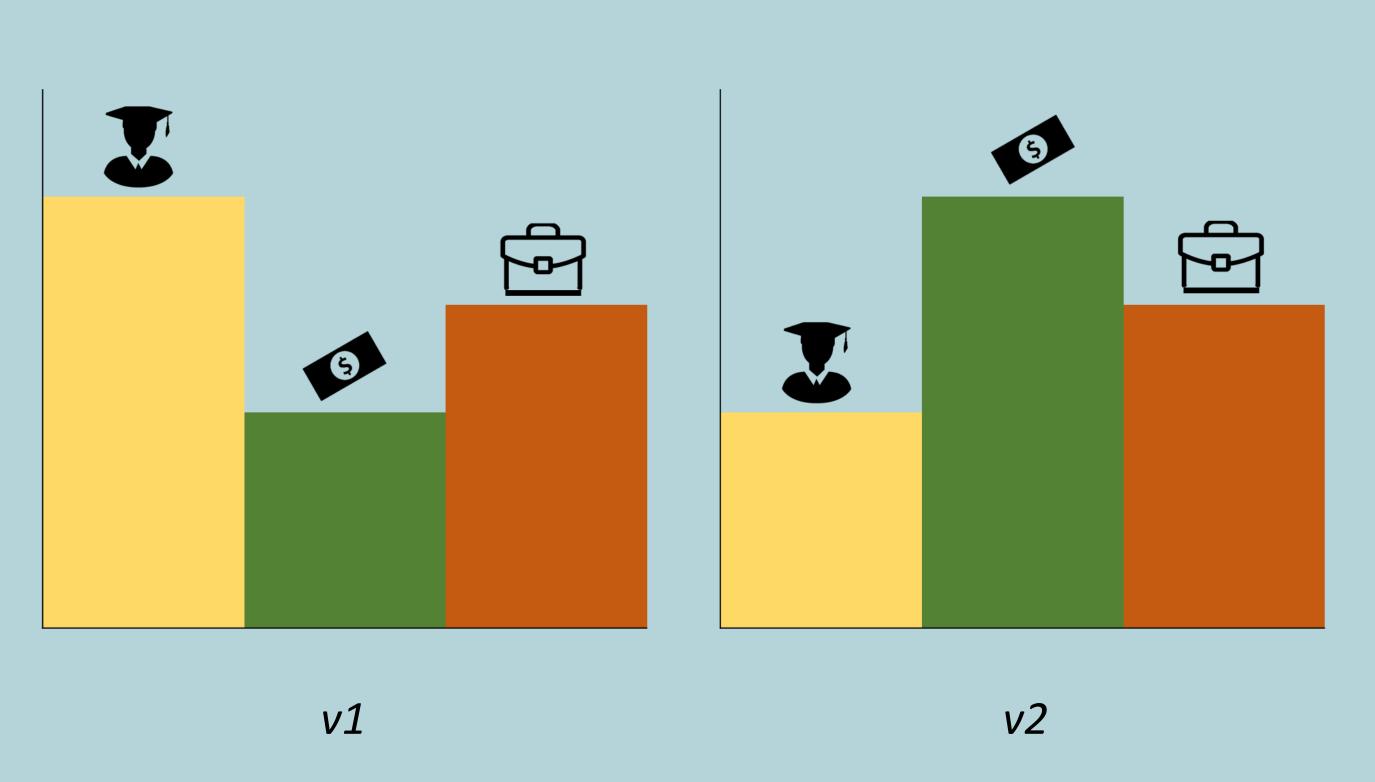


Figure 2. Examples of "hidden" version of the exposure *Medium* SES"

The No-Multiple-Versions-of-Treatment assumption implies a well-defined research question.

Considerations

- . If data collection is planned, predefine the causal contrast (research question) to be estimated as precisely as possible.
- For composite measures (e.g. SES), consider the analysis of the components separately.
- When using registry or pre-collected data neither option above might be available to the researchers.
- . In certain scenarios, it is plausible to identify an "overall treatment effect" even under violations of this assumption.[2-3]

References

- 1. Rehkopf, D.H., Glymour, M.M., Osypuk, T.L. The Consistency Assumption for Causal Inference in Social Epidemiology: When a Rose Is Not a Rose. Curr Epidemiol Rep, (2016).
- 2. Hernán M.A, VanderWeele T.J. Compound Treatments and Transportability of Causal Inference. Epidemiology, (2011). 3. VanderWeele T.J, Hernán M.A. Causal Inference Under Multiple Versions of Treatment. J Causal Inference, (2013).