

CEP Group Meeting
Update on *paper 3* of my PhD

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Overview

- ▶ **Aim:** to research the *short-term* effects of childhood exposures to non-persistent EDCs on neurodevelopment in childhood, and how the metabolome **might** *mediate* these effects.
- ▶ **How:** by making use of the principles and criteria of triangulation.
- ▶ **Population:** the HELIX sub-cohort, consisting of $N = 1200$ mother-child pairs.
- ▶ **Exposures:** non-persistent EDCs (phenols, phthalates, and organophosphate compounds), measured in childhood in a pool of two urine samples.
- ▶ **Outcome:** Attention Network Test, to provide a measure of the efficiency of three different functions of attention.
- ▶ **Mediators:** urine metabolites (focus on corticosteroids).

The Research Questions



Figure 1: Simplified *DAG*.

Issues and limitations:

- ▶ The effect of exposure to these chemicals on neurodevelopment has been studied already, although there are clear issues with that study (e.g., wrong model, DAG **not** tailored to the exposures of interest).
- ▶ The effect of exposure to these chemicals on these **newly** measured metabolites could be identifiable, although they were measured in the same samples. This question poses some challenges since they were all measured in urine (inter-individual variability in urine dilution).
- ▶ The effect of these **newly** measured metabolites on neurodevelopment could be identifiable, since they were measured in the urine samples (**which aliquot?**) collected before the visit.

Triangulation

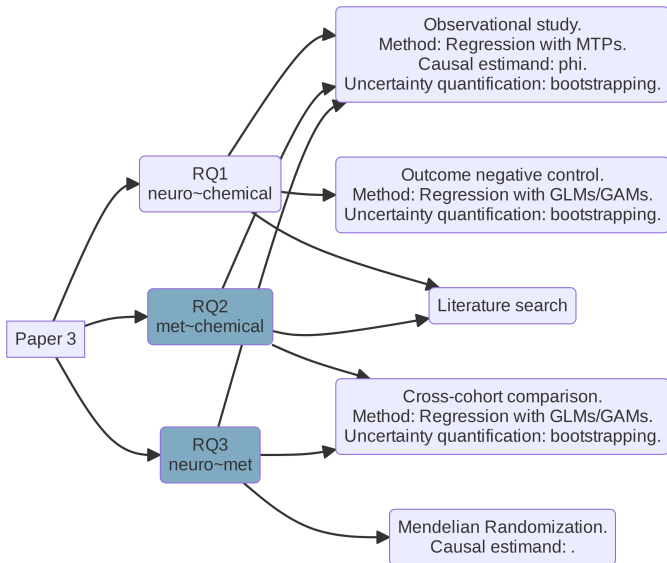


Figure 2: Diagram summarizing research questions and methods.

myphd R package

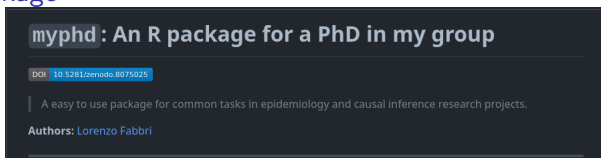


Figure 3: myphd

- ▶ URL manual: <https://isglobal-cep.github.io/myphd/>
- ▶ To install it: `remotes::install_github("isglobal-cep/myphd")`

```
myphd::create_formula(  
  dat,  
  outcome,  
  exposure,  
  covariates,  
  method,  
  add_inter_exposure,  
  add_splines_exposure,  
  df_splines,  
  threshold_smooth,  
  threshold_k  
)
```

```
myphd::describe_data(  
  dat,  
  id_var,  
  grouping_var  
)
```

```
myphd::preproc_data(  
  dat,  
  outcome = NULL,  
  dic_steps,  
  id_var,  
  by_var  
)
```

Current setup

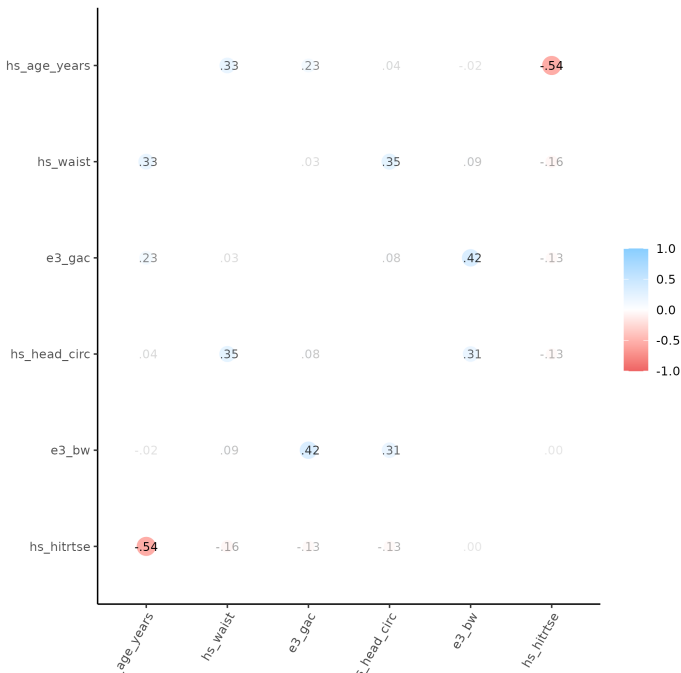
Variables:

- ▶ Clinical outcome: Attention Network Test (`hs_hitrtse`, Hit Reaction Time Standard Error).

Methods:

- ▶ Weights estimation (`chemical ~ covariates`): energy balancing.
- ▶ Effect estimation (`outcome ~ chemical + covariates`): glm with natural splines for exposure.

Population description: correlations



Preliminary results

Next steps