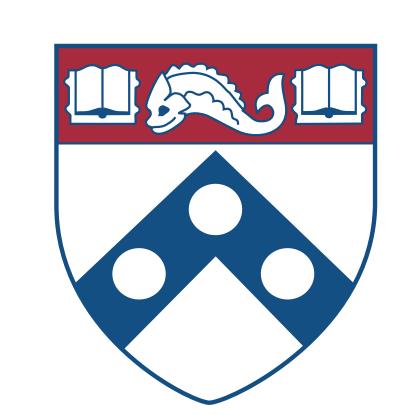
Extended Sensitivity Analysis for Heterogeneous Unmeasured Confounding with an Application to Sibling Studies of Returns to Education



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Heterogeneous Patterns of Confounding

- A sensitivity analysis assesses how robust causal conclusions from an observational study are to departures from random assignment due to unmeasured confounding.
- In a stratified study, the **sensitivity value** is the smallest bound on the maximal bias present in any stratum, while allowing for *arbitrarily unfavorable patterns of confounding*, such that the qualitative conclusions of the study change.
- If the investigator suspects hidden confounding is **present but heterogeneous** across strata *the least favorable patterns of confounding may be very unlikely,* leading the sensitivity analysis to be **overly pessimistic**.

Goal of the Paper: Extend sensitivity analysis to place bounds on both maximal and typical biases present in strata, to account for the presence of heterogeneous confounding.

Example: Returns to Education

Ashenfelter and Rouse (1998):

- Collected survey data on 340 monozygotic twin pairs from the Twinsburg Twins Festival in Twinsburg, Ohio over three summers (1991-1993).
- I=40 pairs where one twin had at least two years of college education ($Z_{ij}=1$) and the other no more than a HS degree ($Z_{ij'}=0$).
- Included twin pairs where both twins held a job at the time of survey collection; log wage data reflecting the 1989-1993 period $(R_{ij} = Z_{ij}r_{Tij} + (1 Z_{ij})r_{Cij})$.
- Retrospective survey \implies no measures of baseline ability, such as IQ recorded earlier in life (a potential unobserved confounder u_{ij}).



Naive analysis assuming no unmeasured confounding:

- p-value ≈ 0.0001 (Fisher's Sharp Null $H_0: \mathbf{r}_T = \mathbf{r}_C$)
- 95% CI: [16%, 43%]

A Model for Unmeasured Confounding

• Assuming **no unmeasured confounding** between twins, treatment assignment in a twin study looks like a randomized experiment:

$$\mathbb{P}(\mathbf{Z} = \mathbf{z} | \mathbf{z} \in \Omega) = 1/2^{I}, \ \Omega = \{\mathbf{z} : z_{i1} + z_{i2} = 1, \forall i = 1, ..., I\}.$$

• In the presence of unmeasured confounding ($u_{i1} \neq u_{i2}$), distribution of treatment allocations is unknown:

$$\mathbb{P}_{\boldsymbol{\pi}}(\mathbf{Z} = \mathbf{z} | \mathbf{z} \in \Omega) = \prod_{i=1}^{I} \pi_{i1}^{Z_{i1}} (1 - \pi_{i1})^{1 - Z_{i1}}$$

because $\pi_{ij} = \mathbb{P}(Z_{ij} = 1 | \mathbf{Z} \in \Omega)$ are **unobserved**.

• Departures from randomization parameterized by $\Gamma > 1$, a **bound on the maximal odds of treatment assignment** in any twin pair:

$$\pi_i^*/(1-\pi_i^*) \le \Gamma, \ \pi_i^* = \max\{\pi_{i1}, \pi_{i2}\}, \ \forall i = 1, \dots, I.$$

• Conventional Sensitivity Analysis under this model: For a test statistic $t = \mathbf{Z}^T \mathbf{q}(\mathbf{R})$ and given maximal bias bound Γ , find the pattern of π that maximizes p-value $P(\pi)$ under reference distribution \mathbb{P}_{π} ,

$$P_{\Gamma}^* = \max_{\boldsymbol{\pi} \in \mathcal{P}_{\Gamma}} P(\boldsymbol{\pi})$$

where $\mathcal{P}_{\Gamma} = \{ \boldsymbol{\pi} : 1/2 \leq \pi_i^* \leq \Gamma/(1+\Gamma), \forall i = 1, \dots, I \}$

Conventional Sensitivity Analysis of Example

- Worst case pattern of confounding: Amounts to assigning twin with higher hourly wage the largest allowable probability of attending college \implies average and maximal π_i^* coincide.
- Sensitivity value: $\Gamma^* \approx 2.36$ where $\Gamma^* = \inf\{\Gamma: P_{\Gamma}^* \geq 0.05/2\}$.

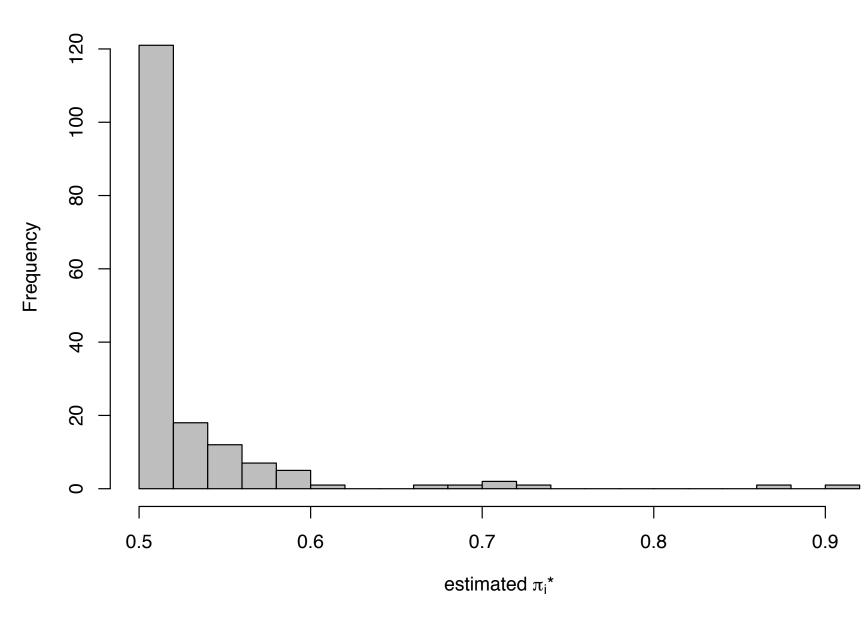
Is 2.36 a plausible bound on the maximal bias among twins due to unmeasured confounding?

• **Example: Ability Bias** – Could disparities in ability between twins in our study increase the odds that one of the twins went to college by more than a factor of 2.36?

Cross-Study Calibration

A partial solution to the above open question: Calibrate Γ^* to an estimate of maximal bias due to disparities in ability from a calibration study that has comparable design and information on baseline IQ [using calibration methods from Hsu and Small (2013)]. For example,

Estimated π_i^* from 171 same-sex sibling pairs in the Wisconsin Longitudinal Study (WLS) where $Z_{i1}+Z_{i2}=1$:



- $\max \pi_i^*/(1 \max \pi_i^*) = 9.3$ and $\bar{\pi}^*/(1 \bar{\pi}^*) = 1.1 \implies$ mostly modest biases, some moderate and few very large biases.
- $\Gamma^* < 9.3 \implies$ Ashenfelter study **likely sensitive** to plausible levels of ability bias but also suggests *maximal and typical biases due to IQ disparities are quite different*.

Extended Sensitivity Analysis

Our Contribution: Adapt the conventional sensitivity analysis to provide a less pessimistic view of the study's robustness to hidden bias when heterogeneous confounding leads to differing bounds on maximal and typical bias.

Extended Sensitivity Analysis (ESA): A two-parameter sensitivity analysis that simultaneously bounds the maximal and typical biases by Γ and $\bar{\Gamma}$, respectively:

$$P_{\Gamma,\bar{\Gamma}}^* = \max_{oldsymbol{\pi} \in \mathcal{P}_{\Gamma,\bar{\Gamma}}} P(oldsymbol{\pi})$$

where $\mathcal{P}_{\Gamma,\bar{\Gamma}}$ is the set of π that respect both bounds on maximal and typical biases.

Properties:

- Returns a curve of $(\Gamma, \overline{\Gamma})$ changepoints, analogous to Γ^* .
- If desired, bias parameters can be calibrated to population-level bias information (important in cross-study calibration).
- Can recover conventional sensitivity analysis when $\bar{\Gamma} = \Gamma$.
- Computationally feasible can be expressed as a standard QP [Fogarty and Small (2016)].

Superpopulation Model for Paired Studies

- Sample a treated subject from an infinite population of treated subjects and record $X_{i1} = x_i$ then sample a control subject from an infinite population of controls conditional on $X_{i2} = x_i$.
- Under this model, π_{ij} is a realization of random variable Π_{ij} $\Longrightarrow \pi_i^*$ a realization of random variable $\Pi_i^* = \max\{\Pi_{i1}, \Pi_{i2}\}.$

Why assume a population when inferential framework is conditional on sample?

Why Population-Level Bias Calibration?

Bias estimates from calibration sample are relevant to study sample only insofar as the two samples believably came from some common superpopulation!

A difficulty with constructing a set $\mathcal{P}_{\Gamma,\bar{\Gamma}}$ in terms of bounds on population-level bias:

• Although $1/2 \le \Pi_i^* \le \Gamma/(1+\Gamma) \implies 1/2 \le \max \pi_i^* \le \Gamma/(1+\Gamma)$, the same correspondence does not hold for typical biases,

$$1/2 \le \mathbb{E}[\Pi_i^*] \le \bar{\Gamma}/(1+\bar{\Gamma}) \implies 1/2 \le \bar{\pi}_i^* \le \bar{\Gamma}/(1+\bar{\Gamma}).$$

Implication for constructing $\mathcal{P}_{\Gamma,\bar{\Gamma}}$: A deterministic set $\mathcal{P}_{\Gamma,\bar{\Gamma}}$ respecting population-level bias bounds would allow for an allocation π such that $\bar{\pi}^*$ and $\max \pi_i^*$ coincide, however unlikely \Longrightarrow back in the conventional sensitivity analysis.

How can we remedy this?

Making $\mathcal{P}_{\Gamma,\bar{\Gamma}}$ Stochastic

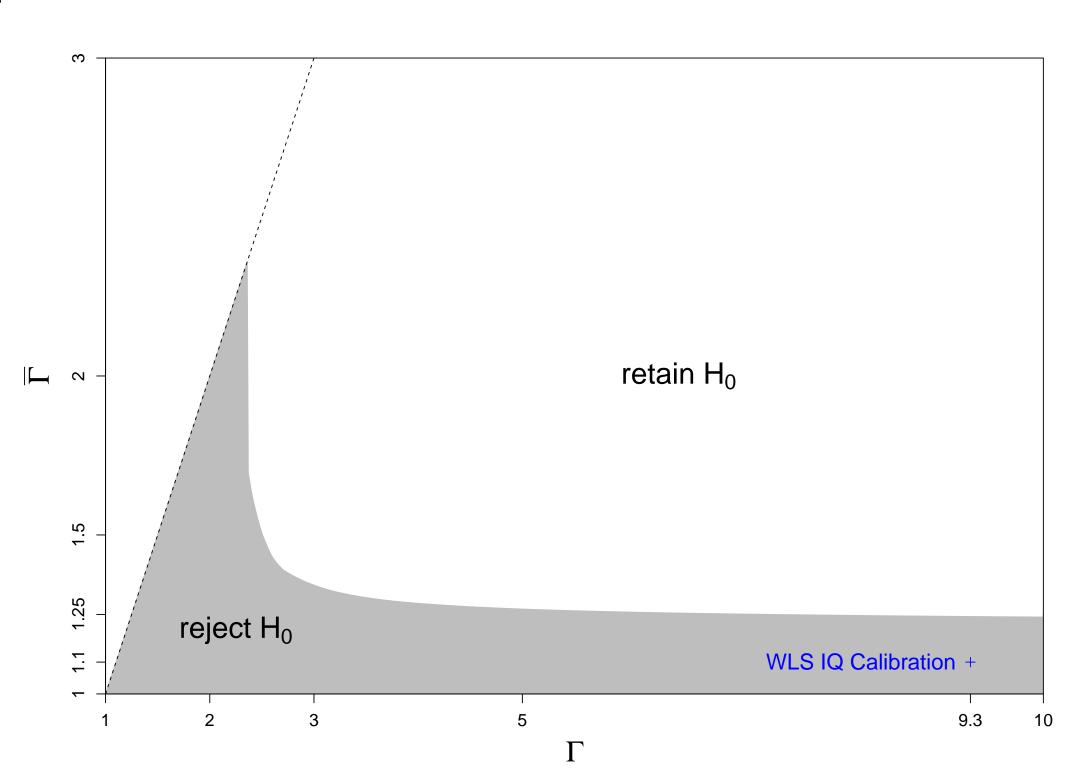
A strategy from Berger and Boos (1994): Maximize p-value over $1-\beta$ confidence set for a nuisance parameter and add β to get a valid p-value for tests based on non-pivotal quantities.

Modification for ESA: π takes the place of the nuisance parameter, need $\Pi \in \mathcal{P}_{\Gamma,\bar{\Gamma}}$ with probability at least $1-\beta$, and β is added to the maximal p-value to account for stochastic nature of set.

Ensuring $\mathbb{P}(\Pi \in \mathcal{P}_{\Gamma,\bar{\Gamma}}) \geq 1 - \beta$: Can use fact that $\bar{\Pi}^*$ is a sample average, conservative variance estimate for Π_i^* that depends only on $\bar{\Gamma}$ and Γ (Bhatia-Davis Inequality), and apply CLT.

Calibrated Sensitivity Curve

Rather than a single sensitivity parameter, ESA returns a **sensitivity curve** of $(\Gamma, \overline{\Gamma})$ below which we cannot reject H_0 , calibrated to estimates of **both** typical and maximal biases due to disparities in IQ:



Takeaway: Unlike the conventional analysis, the calibrated ESA suggests that the Ashenfelter study **is not sensitive** to plausible patterns of ability bias!

Special Cases:

- At (Γ, Γ) we recover the conventional sensitivity analysis $\Longrightarrow \Gamma^* \approx 2.36$.
- If no plausible bound on maximal bias, taking $\lim_{\Gamma_n \to \infty} (\Gamma_n, \bar{\Gamma})$ recovers single-parameter sensitivity analysis bounding only the typical bias $\Longrightarrow \bar{\Gamma}^* \approx 1.22$.

Concluding Remarks

- In summary, the extended sensitivity analysis generalizes the conventional sensitivity analysis by allowing the researcher to place bounds on both the maximal and typical bias.
- As illustrated with the Ashenfelter study, this will generally result in less conservative conclusions when there is reason to believe the bounds on maximal and typical bias do not coincide.

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