

Discussion on Stringhini & Chadeau-Hyam – mechanisms in LIFEPATH

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## Key lessons (Silvia): Inequalities in mortality and functioning

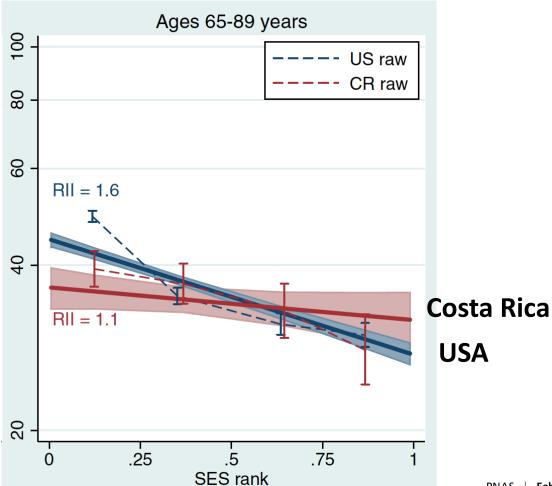


- Pooling cohorts not harmonised ex-ante is possible: socioeconomic inequalities in health everywhere
  - Can we understand better heterogeneity, across countries and time? replicability vs understanding 'stage' of 'inequalities transition'
- Theory model of 'inequalities transition': Explain differences across countries in staging of health inequalities, e.g., Costa Rica
- SEP affects both biology and function
  - What are the dynamics of these relationships? E.g., early childhood disadvantage, BMI, adult SEP
- SEP has independent association with health
  - Residual confounding, 'statistical controlling' enough?



### Exploring why Costa Rica outperforms the United States in life expectancy: A tale of two inequality gradients

Luis Rosero-Bixby<sup>a,1</sup> and William H. Dow<sup>a,b</sup>



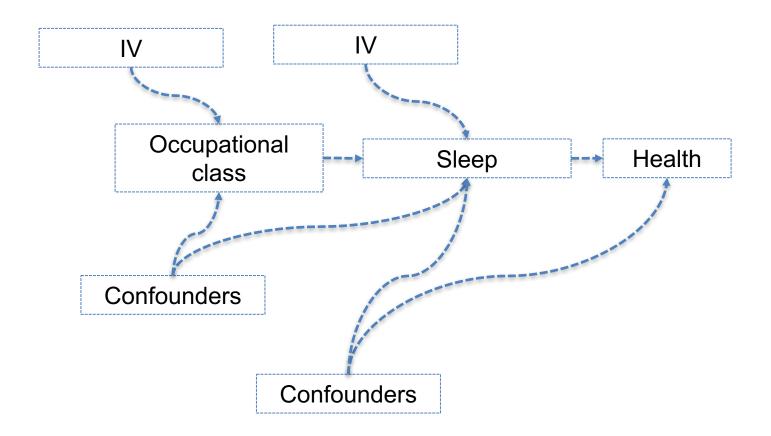






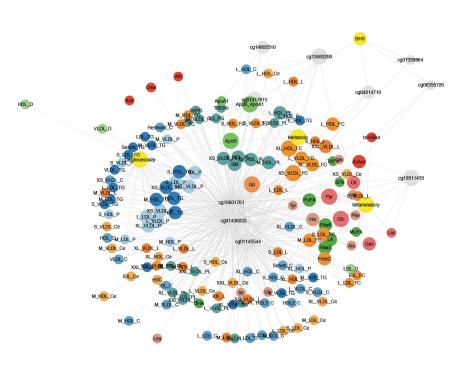
- Relationship between SEP and health partly mediated by behavioural risk factors, but not fully explained
- Baron & Kenny (1986): strong assumptions, counterfactual approach might reveal difference results (e.g., sleep)
- Relative importance different risk factors open question
- Why do lower SEP groups engage into poor behaviours? Stress? knowledge? preferences? parenting? environment?
- What are the causal dynamics? e.g., does income leads to poorer behaviour, or is poorer behaviour the cause of lower income, or do both share common cause
- Differences between cohorts (countries) in pathways
- If SEP independent factor (Lancet, BMJ), why mechanisms?

### Can we assess mediation in a crosssectional dataset in a population cohort?



## Key Lesson (Marc): Methylation and (abstract)modern art





**OMICs Integration** 



Sirodot Régine

### Key lessons (Marc)



- Lower SEP associated with accelerated (epigenetic) ageing as measured by methylation
  - How important is this targeted finding in the broader picture of health, functioning and mortality? Independent?
  - Can we intervene on methylation to change disease, functioning or survival?
- The fact that we find correlations between early life SEP and late-life health does not mean that early life is the only (or 'best') period on which to intervene
  - Inequalities start early, but are interventions in early life more 'effective' than those in mid or later life?
  - Normative question, e.g., inequalities in early life 'more unfair' than inequalities in later life?



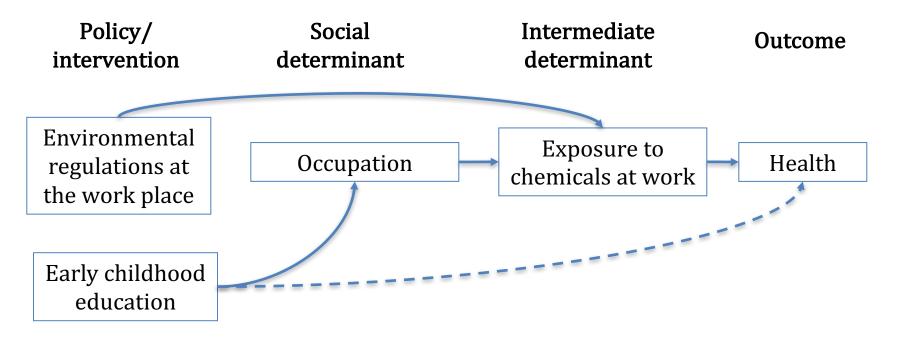
#### Two policy questions

1. Can we intervene on (some component of) socioeconomic position to influence health?

- 2. Can we intervene on the pathways that link socioeconomic status and health?
  - Are there clear biological pathways on which we can intervene
  - Should we prioritise some pathways over others?
  - Can we intervene on methylation pathways

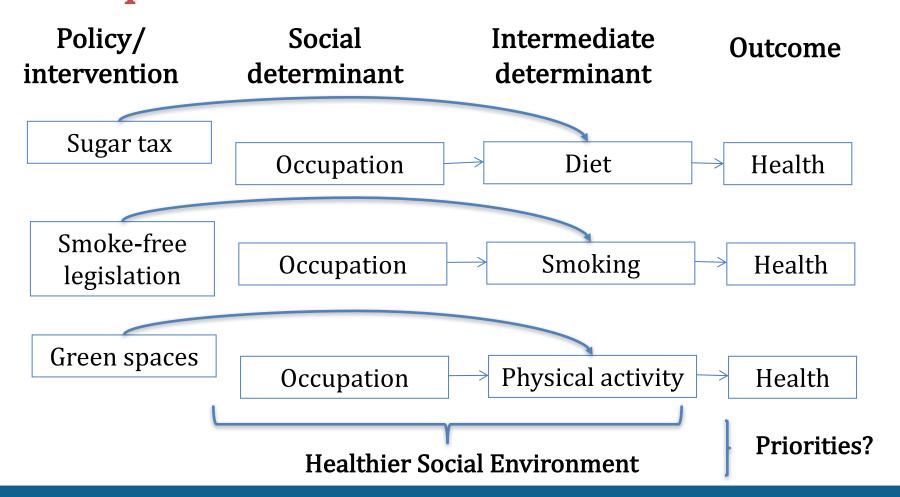
# Policy and socioeconomic inequalities in health





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# Methylation: Implications for policy



- Helpful to 'capture' ageing-related change associated with SEP,
  but unclear how independent from all unobservables
- Evidence of changes in methylation come from behaviour changes, e.g., smoking cessation; to what extent a change in SEP would change methylation pathways?
  - Evidence of direct interventions on methylation processes limited (I think)
- Not necessarily a problem, as we advance science: it might just take time before we can derive policy implications



#### In conclusion

- Lifepath has revealed important evidence on (a) the consistency of health inequalities; (b) the potential role of several mediating pathways from behaviour to methylation
- However, gaps remained for drawing policy implications:
  - Causal processes
  - 2. **Reversibility:** How interventions would alter pathways & outcomes
  - 3. **Cross-cohort variation:** We need more theory & hypothesis testing
  - 4. Implications for social environment, but priorities unclear
  - 5. **Importance** of methylation still unclear, and implications for policy likely only in the **distant future**