



Karolinska
Institutet

A Lifespan Perspective on Cognitive Reserve and Risk for Dementia

Matthew S. Panizzon¹, **Ida K. Karlsson**², Malin Ericsson², Marianne Nygaard³, Margaret Gatz⁴, Nancy L. Pedersen², & William S. Kremen¹

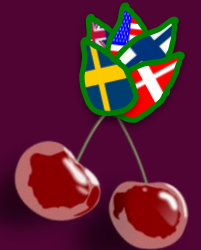
For the IGEMS Consortium

¹University of California, San Diego

²Karolinska Institutet (Department of Medical Epidemiology and Biostatistics)

³University of Southern Denmark

⁴University of Southern California



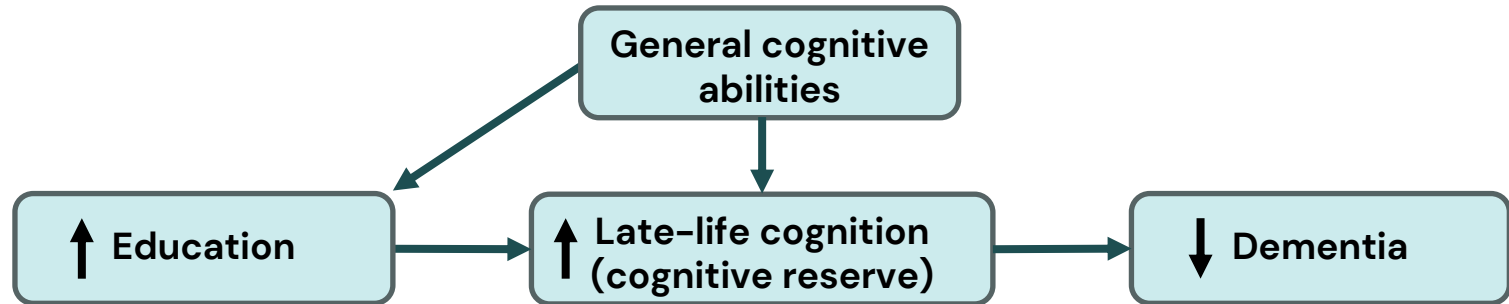
IGEMS consortium



- **The Interplay of Genes and Environment across Multiple Studies (IGEMS) consortium**
- Brings together twin studies of adult development and aging from Sweden, Denmark, Finland, Australia, the U.S.
- Pedersen et al. *IGEMS: The Consortium on Interplay of Genes and Environment Across Multiple Studies – An Update*. Twin Research and Human Genetics, 2019.
- **Current study:**
 - Data from the Swedish Twin Registry
 - Led by Matt Panizzon at UC San Diego

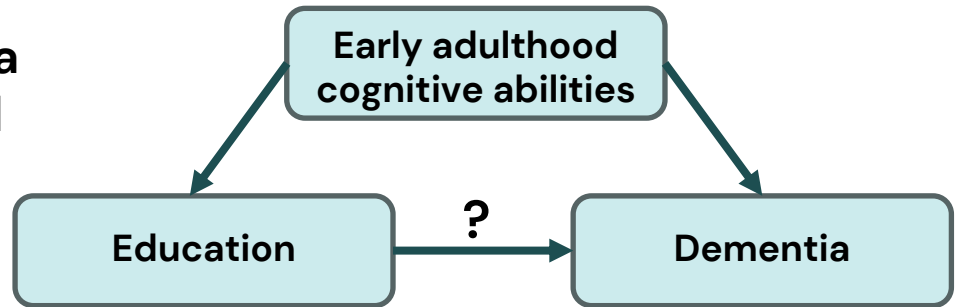
Background and aims

- Higher education = well-established protective factor for dementia
- Higher education → cognitive reserve
- Causal association?
- Explained by higher cognitive abilities → higher education?



Background and aims

- Higher education = well-established protective factor for dementia
- Higher education → cognitive reserve
- Causal association?
- Explained by higher cognitive abilities → higher education?
- **Does education predict dementia if we account for early adulthood cognitive abilities?**



Methods

- Swedish Twin Registry: Screening Across the Lifespan Twin (SALT) study, conducted 1998–2002
- Male twins born 1936–1958 → N = 13,771
- Mean age: 52.7 (standard deviation: 5.6)

- Education information → International Standard Classification of Education (ISCED) system

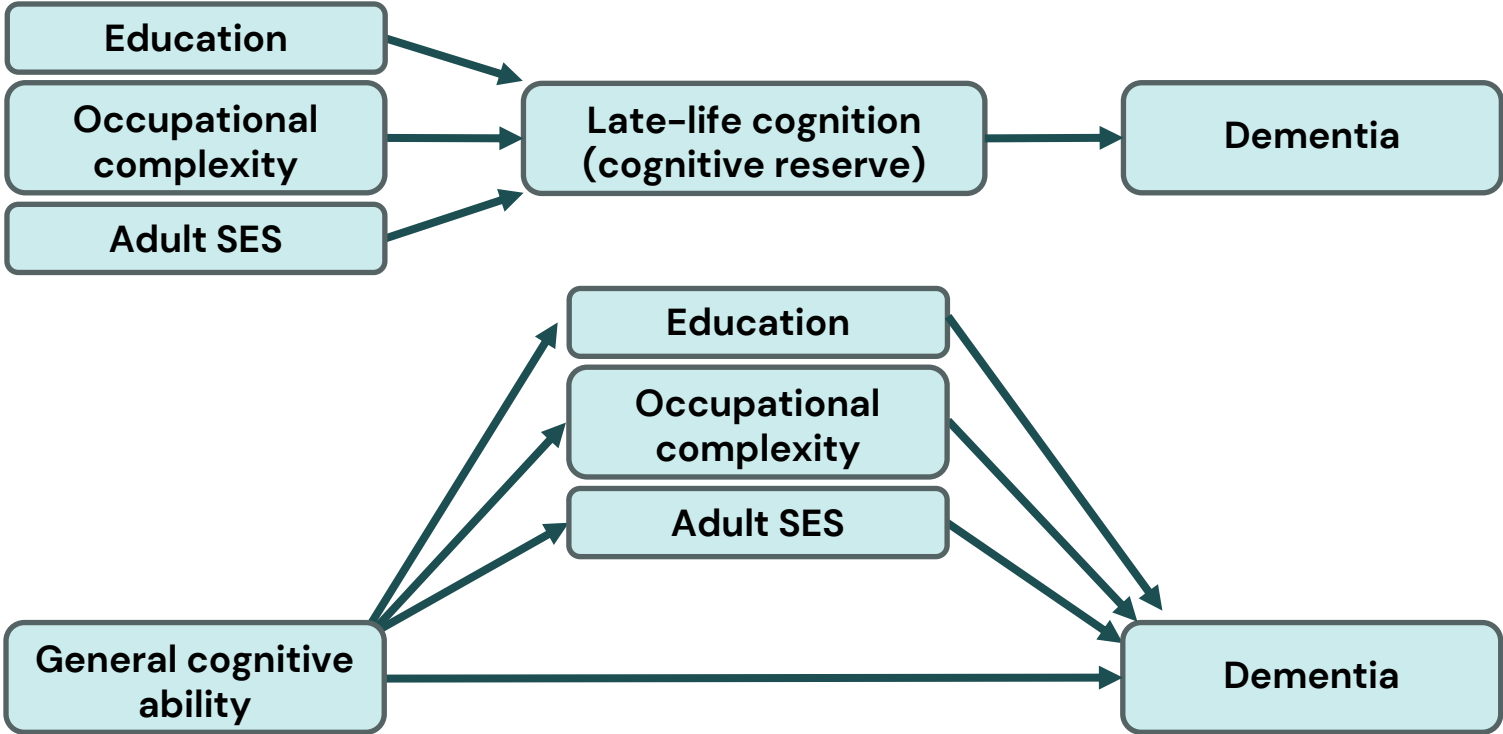
- In addition: occupational complexity, adult socioeconomic status

Methods

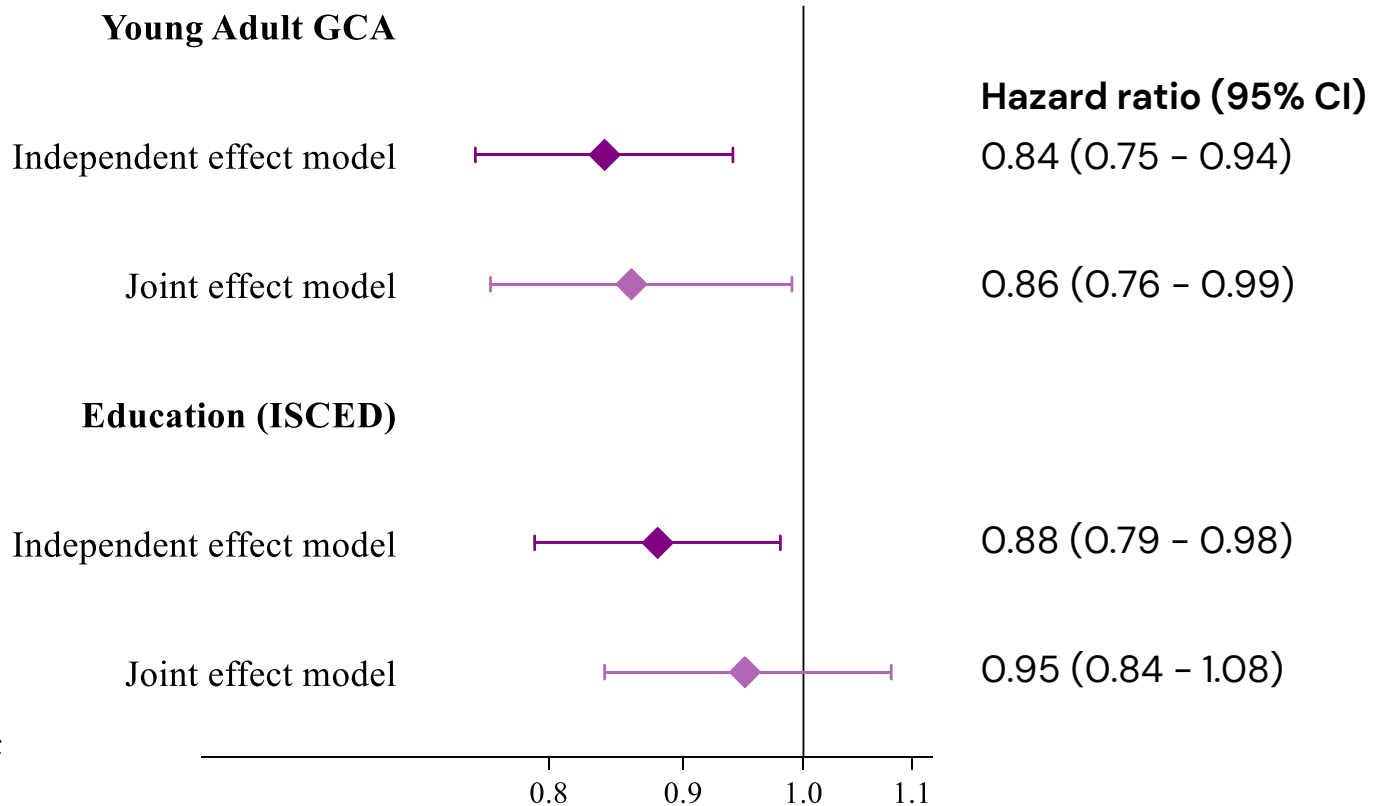


- Linkage to the Conscription Register → cognitive abilities in early adulthood
- Dementia diagnoses: National healthcare registers, linkage through 2016
 - National Patient Register (inpatient + outpatient specialist care)
 - Causes of Death Register
 - Prescribed Drug Register (dementia medication, used as proxy for diagnosis)
- Cox proportional hazard model
 - Followed from age ≥ 50 to dementia diagnosis, death, or end of follow-up

Methods



Results



Young adult general cognitive abilities: Continuous measure; mean 19.0 (SD 6.3)

Education (ISCED): Continuous measure, 5-point scale

Results

Occupational Complexity

Independent effect model



Hazard ratio (95% CI)

0.90 (0.80 - 1.09)

Joint effect model



0.96 (0.85 - 1.08)

Adult SES

Independent effect model



0.86 (0.77 - 0.97)

Joint effect model



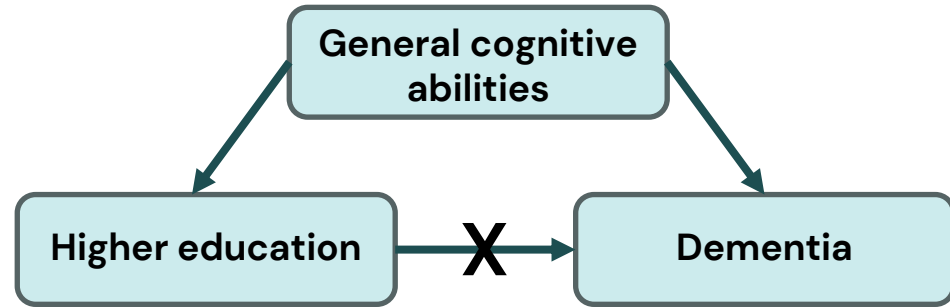
0.93 (0.82 - 1.05)

Occupational complexity:
Continuous measure; mean 3.3
(SD 0.4)

Adult SES: Continuous measure ;
mean 46.2 (SD 21.3)

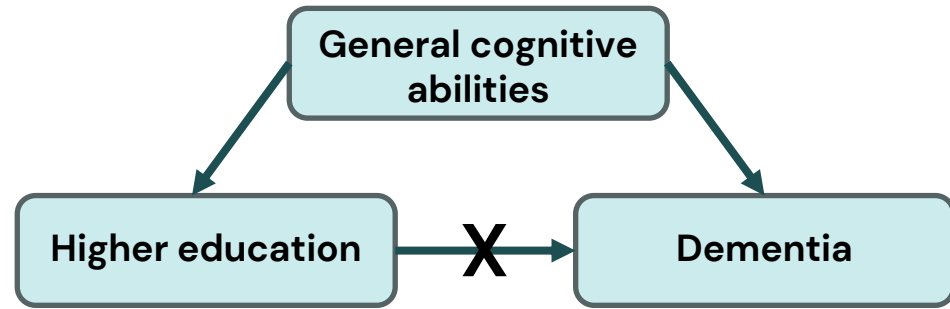
0.8 0.9 1.0 1.1

Discussion



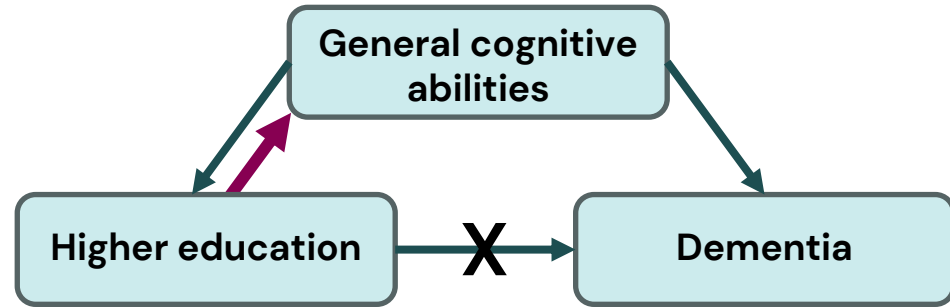
- Lower education is one of the most well-established modifiable risk factors for ADRD
 - Current study: Higher education → lower risk of ADRD
 - However: No association when accounting for early adulthood general cognitive abilities
- Supports confounding through general cognitive abilities

Discussion



- Further supported by similar findings for:
 - Occupational complexity
 - Adult socioeconomic status
- Findings align with prior work
 - E.g. Kremen et al. *Influence of young adult cognitive ability and additional education on later-life cognition*. Proc Natl Acad Sci U S A. 2019.
 - Nygaard et al. *Influence of young adult cognitive ability on the association between lifetime education and later-life cognitive function – a study in Danish twins*. NKG, Thursday Ber2:PO2; Poster tour: 12.45, Ber2:1.

Conclusion



- NOTE! Bidirectional association: Higher education → increased general cognitive abilities
- Early adulthood cognitive abilities robustly associated with lower risk of dementia, even when controlling for education, occupational complexity, or adult SES
- Education → dementia = downstream of general cognitive abilities (same for occupational complexity and adult SES)



**Karolinska
Institutet**