



Andrea L.C. Schneider MD PhD

Assistant Professor Department of Neurology, Division of Neurocritical Care Department of Biostatistics, Epidemiology, and Informatics University of Pennsylvania Perelman School of Medicine

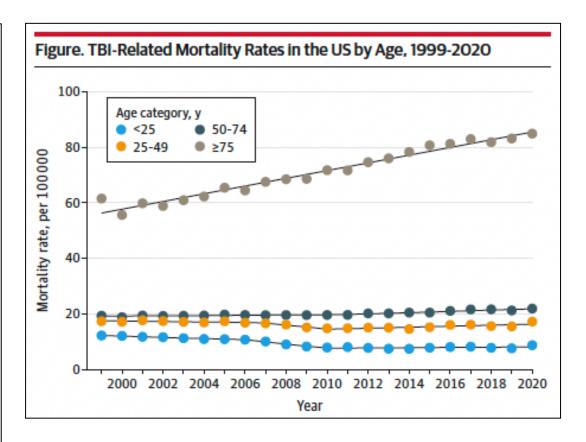
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TBI-Related Mortality in the United States

	Age-standardized mortality rate per 100 000 (95% CI)		1999-2020 AAPC	P value for
	1999	2020	(95% CI) ^a	trenda
Overall	18.64 (18.31 to 18.96)	19.34 (19.05 to 19.63)	0.14 (-0.60 to 0.88)	.72
Sex				
Male	28.90 (28.31 to 29.49)	30.17 (29.64 to 30.70)	0.12 (-0.66 to 0.91)	.76
Female	9.49 (9.18 to 9.81)	9.31 (9.04 to 9.59)	-0.05 (-0.35 to 0.25)	.73
Race and ethnicity				
Non-Hispanic American Indian or Alaska Native	24.19 (20.10 to 28.62)	30.89 (26.81 to 35.11)	1.04 (-0.96 to 3.08)	.31
Non-Hispanic Asian or Pacific Islander	10.42 (9.05 to 11.80)	8.49 (7.71 to 9.26)	-0.87 (-1.36 to -0.37)	.001
Non-Hispanic Black	20.40 (19.39 to 21.41)	21.29 (20.43 to 22.16)	-0.11 (-0.84 to 0.62)	.76
Hispanic	14.67 (13.66 to 15.67)	13.04 (12.41 to 13.67)	-0.87 (-1.67 to -0.07)	.03
Non-Hispanic White	19.03 (18.65 to 19.41)	20.85 (20.47 to 21.22)	0.46 (0.22 to 0.70)	<.001
Age, y ^b				
<25	12.19 (11.97 to 12.41)	8.66 (8.48 to 8.84)	-1.92 (-2.90 to -0.93)	<.001
25-49	17.33 (17.07 to 17.58)	17.08 (16.84 to 17.33)	-0.32 (-1.47 to 0.84)	.59
50-74	19.32 (18.97 to 19.68)	21.81 (21.62 to 22.11)	0.64 (0.48 to 0.80)	<.001
≥75	61.43 (60.23 to 62.63)	84.78 (83.60 to 85.97)	1.97 (1.80 to 2.15)	<.001





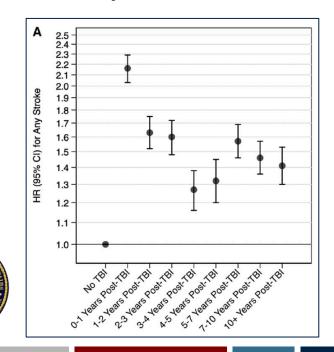


Long-term Outcomes after TBI in Older Adults

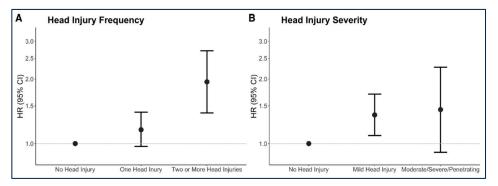
- Stroke
- Cognitive Decline and Dementia
- Epilepsy
- ► How do we study long-term outcomes after TBI when very few studies designed to study TBI from the time of injury have long-term follow-up (i.e., 30+ years)?
 - Leverage data from ongoing, deeply phenotyped epidemiologic studies such as the Atherosclerosis Risk in Communities (ARIC) Study
- ► The study of long-term outcomes after TBI in older adults has methodological challenges that must be addressed
 - Repeated measures
 - Attrition and missing data

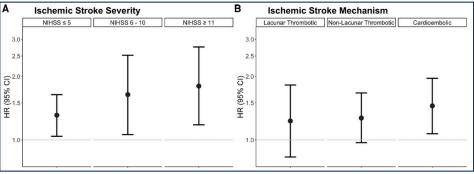
Long-term Outcomes after TBI – Stroke

- ► 306,796 Veterans with TBI matched 1:1 to 306,796 Veterans without TBI
- ► TBI associated with 1.7 times the risk of incident (ischemic or hemorrhagic) stroke over a median of 5 years



► TBI associated with 1.3 times the risk of ischemic stroke among 12,813 community-dwelling participants followed for a median of 27 years







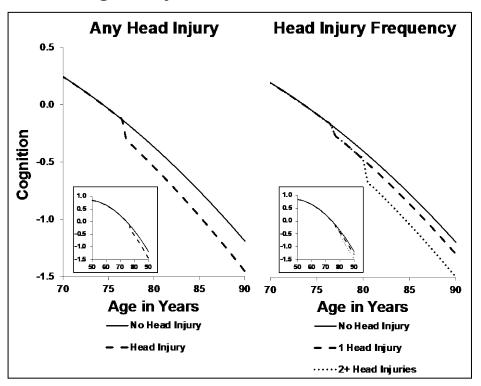
⁻ Elser H, Pappalardo LW, Gottesman RF, Coresh J, Diaz-Arrastia R, Mosley TH, Kasner SE, Koton S, **Schneider ALC**. Head Injury and Risk of Incident Ischemic Stroke in Community-Dwelling Adults. *Stroke*. 2024 Jun:55(6):1562-1571.



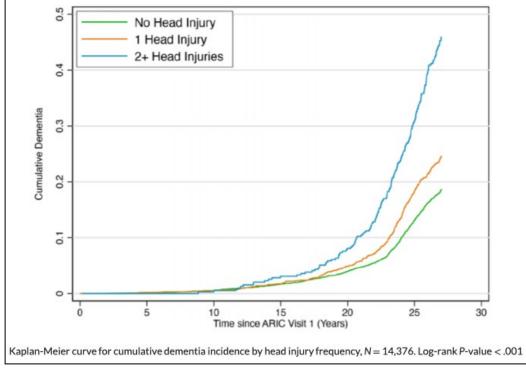
⁻ Schneider ALC, Peltz CB, Li Y, Bahorik A, Gardner RC, Yaffe K. Traumatic Brain Injury and Long-Term Risk of Stroke Among US Military Veterans. Stroke. 2023 Aug;54(8):2059-2068.

Long-term Outcomes after TBI – Cognitive Decline and Dementia

Over 30-years, the difference in cognitive decline between individuals with versus without TBI is equivalent to individuals with TBI being 7.4 years older at baseline



► TBI was associated with 1.4 times the risk of dementia over a median of 25 years





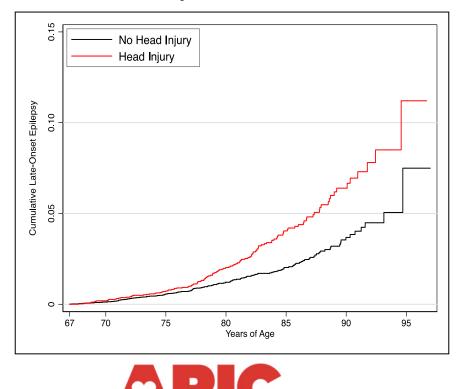
⁻ Schneider ALC, Selvin E, Latour L, Turtzo LC, Coresh J, Mosley T, Ling G, Gottesman RF. Head Injury and 25-Year Risk of Dementia. Alzheimer's and Dementia. 2021 Sep;17(9):1432-1441.

⁻ Schneider ALC, Pike JR, Elser H, Coresh J, Mosley TH, Diaz-Arrastia R, Gottesman RF. Traumatic brain injury and cognitive change over 30 years among community-dwelling older adults. Alzheimer's and Dementia. 2024 Jul 5. doi: 10.1002/alz.14104. Online ahead of print.

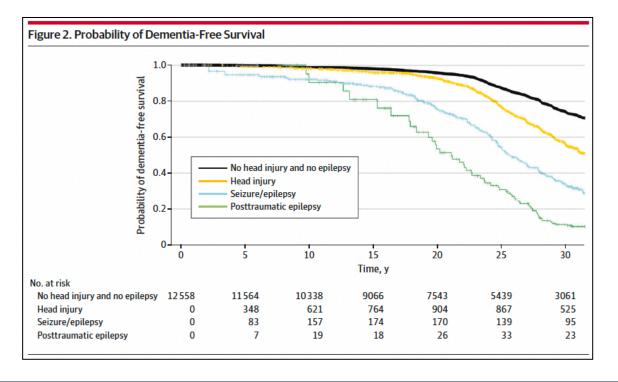


Long-term Outcomes after TBI – Epilepsy and Dementia

► Among 8,878 participants, TBI was associated ► Over a median of 25-years of follow-up of with 1.9 times the risk of epilepsy over a median of 11 years



12,558 individuals, post-traumatic epilepsy was associated with greater dementia risk than TBI or epilepsy alone



⁻ Schneider ALC, Gottesman RF, Krauss GL, Gugger J, Diaz-Arrastia R, Kucharska-Newton A, Huang J, Johnson EL. Association of Head Injury with Late-Onset Epilepsy: Results from the Atherosclerosis Risk in Communities (ARIC) Cohort. Neurology. 2022 Feb 22;98(8):e808-e817.

⁻ Schneider ALC. Law CA, Gottesman RF, Krauss G, Huang J, Kucharska-Newton A, Jensen FE, Gugger JJ, Diaz-Arrastia R, Johnson EL. Post-traumatic Epilepsy and Dementia Risk. JAMA Neurology. 2024 Feb 26;81(4):346-53.



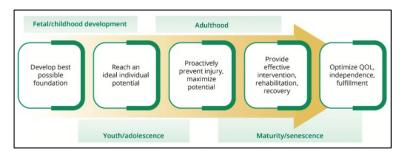
Research Gaps and Recommendations for Research Priorities

- Research to date has focused on TBI as an independent risk factor for later-life outcomes
- ► <u>Recommendation</u>: Focus on contextualizing associations of TBI with long-term outcomes through the lens of "brain health"

 | Brain health is a continuous state of attaining and maintaining the optimal |

Brain health is a continuous state of attaining and maintaining the optimal neurologic function that best supports one's physical, mental, and social well-being through every stage of life.

- Recommendation: Consider how pre- and post-injury risk and resilience factors across the life course may influence associations of TBI with long-term outcomes
 - Medical comorbidities
 - Social and environmental determinants of health



 Recommendation: Prioritize leveraging existing longitudinal datasets to gain new insights (with appropriate consideration of methodological challenges)

