

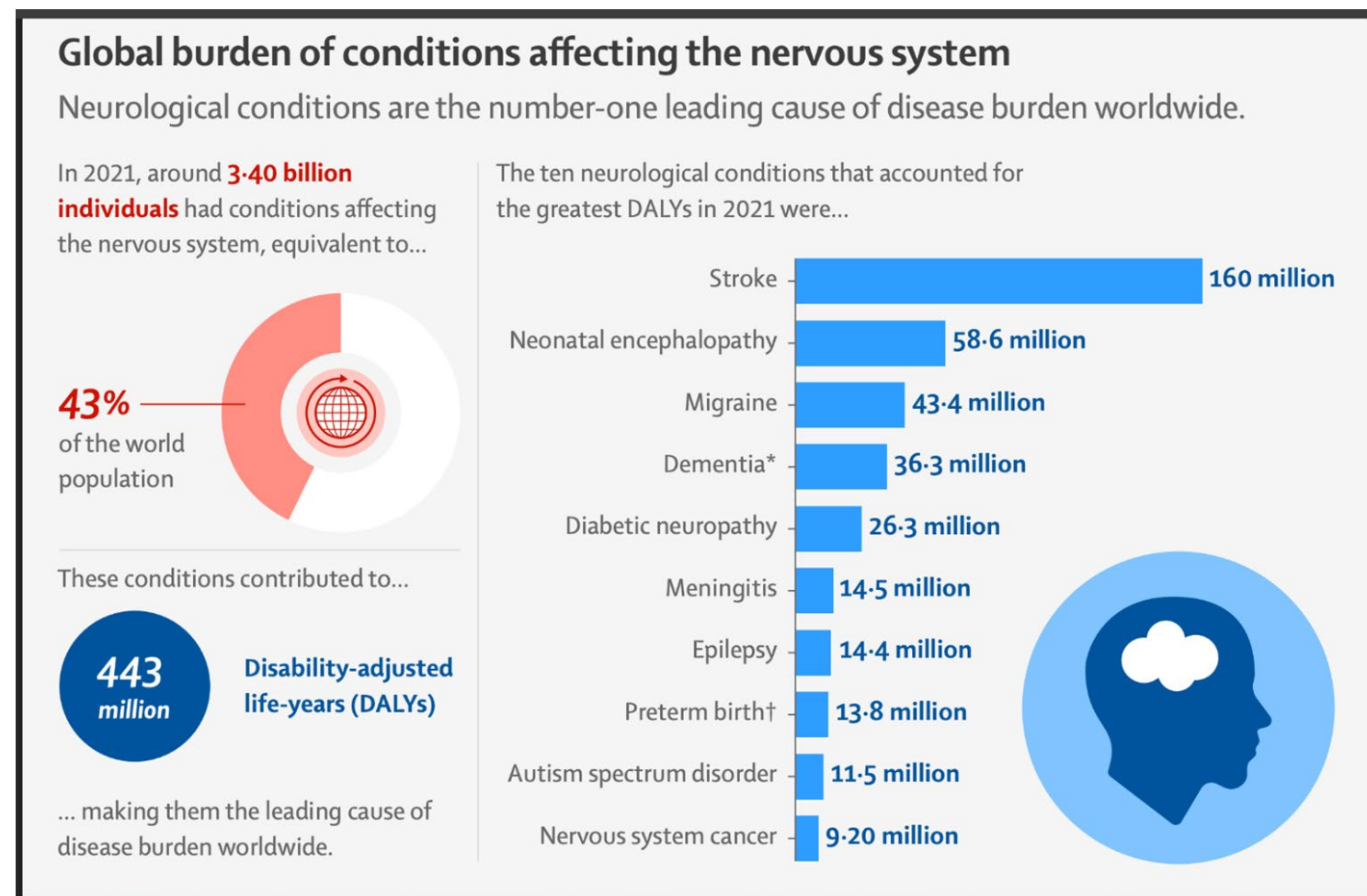
# Environmental Exposures and Neurodegeneration: From Risk to Resilience

*Burcin Ikiz, PhD*

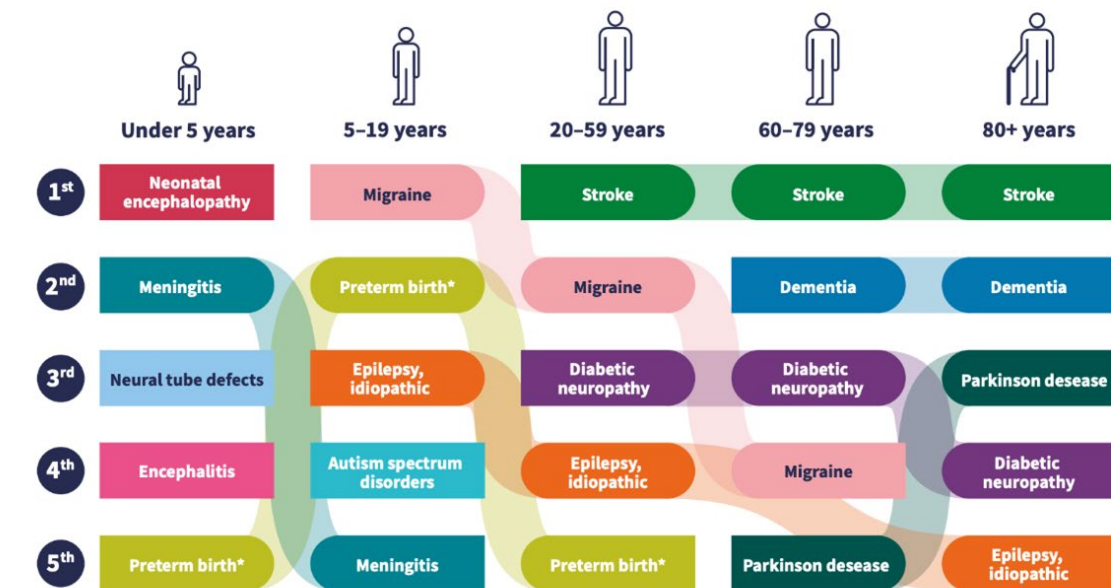
HEI Annual Conference  
April 27, 2026



# Why do we care? The global burden of neurodegenerative diseases (NDs)



Top 5 neurological conditions by age group (2021)



\*Rankings are isolated to disease DALYs due to neurological complications, as opposed to DALYs attributed to the entire condition.  
Source: Adapted from GBD2021 with adjustment using UN World Population Prospects.

WHO 2025 Global Status Report on Neurology

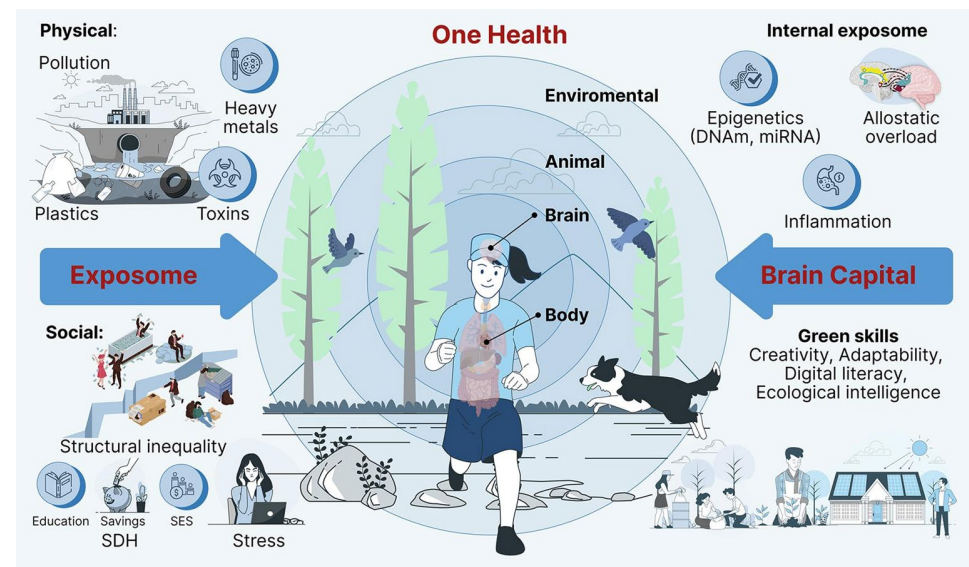
Steinmetz J et al., Global Burden of Neurological Diseases, The Lancet Neurology, 2024

# Beyond genetics: the modifiable risk factors of NDs



Up to ~45% of dementia risk is linked to modifiable factors

# Our brains' aging and ND risk are shaped by our environments

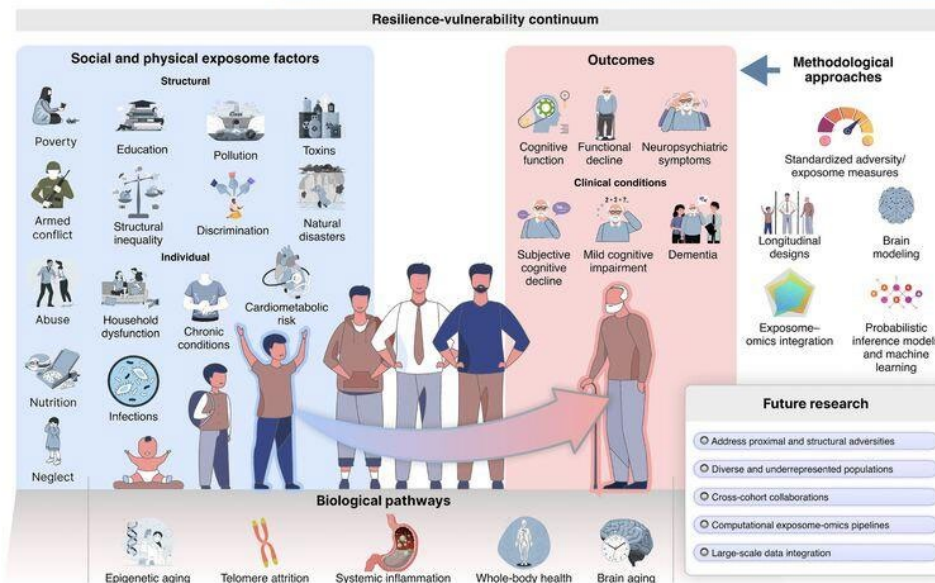


Ibanez et al., Neuron. 2024

## REVIEW

### Childhood adversities and the exposome in dementia risk and brain health

Sandra Baez<sup>a,b</sup>, Yehia Nabif<sup>c</sup> and Agustin Ibanez<sup>a,d,e,f,g</sup>

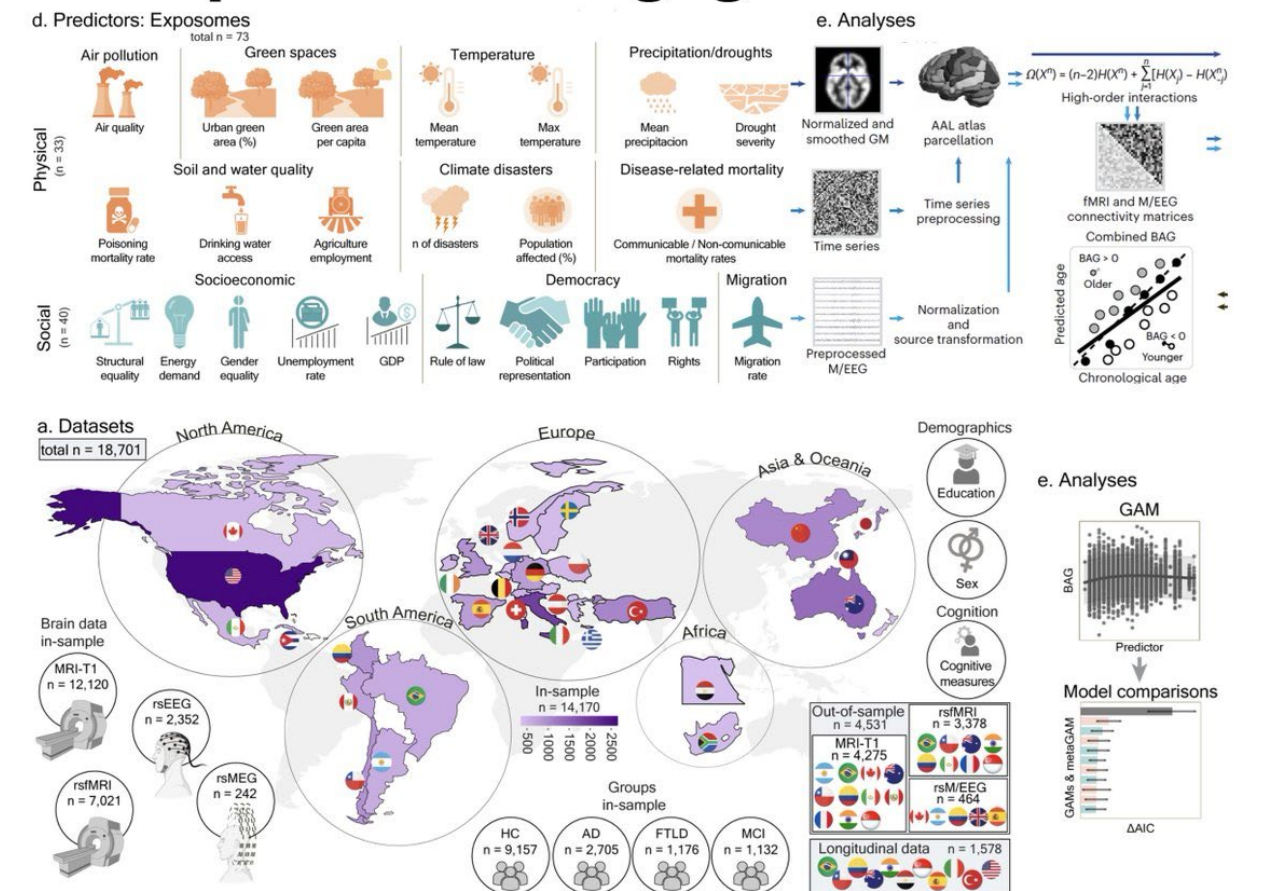


Baez et al., Curr Opin Psychi, 2026

## nature medicine

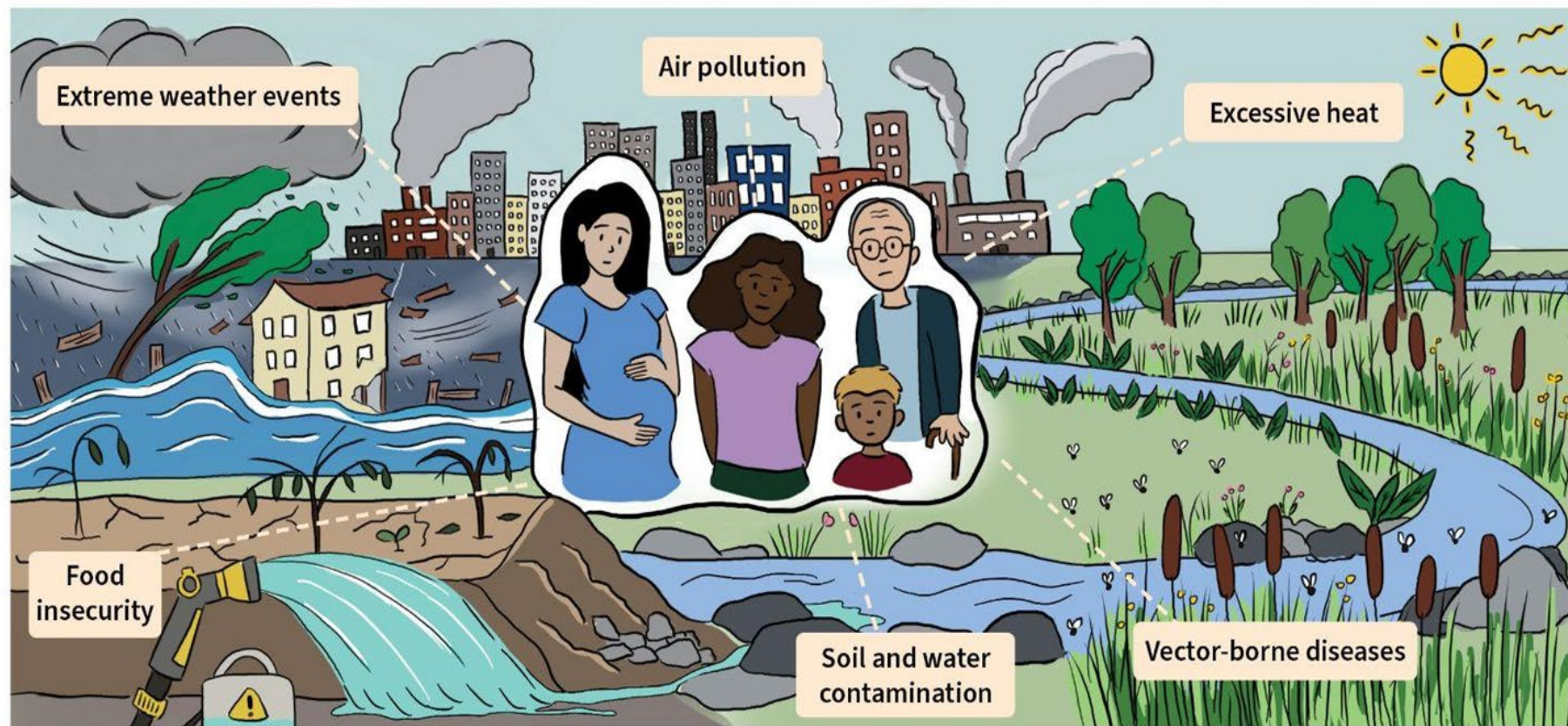
Article

### The exposome of brain aging across 34 countries



Legaz et al., Nature Medicine, 2026

# Environmental risk factors of ND




Associated brain diseases and effects			
<span style="color: blue;">●</span> Neurodevelopmental disorders <span style="color: magenta;">■</span> Neurodegenerative disorders <span style="color: yellow;">▲</span> Mental health disorder <span style="color: green;">◆</span> Intergenerational effects <span style="color: red;">◆</span> Other neurological conditions			
Factors		Direct effects on brain health	Effects on cognition and behavior
Excessive heat	<span style="color: blue;">●</span> <span style="color: red;">◆</span> <span style="color: magenta;">■</span> <span style="color: yellow;">▲</span>	Changes in blood-brain barrier (BBB) permeability, making the brain more vulnerable to inflammation and toxins.	Impairments in memory and attention, increased risk in mental health symptoms, aggression, and impulsivity.
Extreme weather events	<span style="color: yellow;">▲</span> <span style="color: green;">◆</span>	Brain volume alterations in the amygdala, hippocampus, and caudate.	Acute stress responses that can lead to depression, anxiety, and post-traumatic stress disorder.
Air pollution	<span style="color: blue;">●</span> <span style="color: magenta;">■</span>	Particulate matter from the air penetrates BBB leading to inflammation, oxidative stress, and neurodegeneration.	Symptoms of anxiety, depression and impaired cognitive functions.
Food insecurity	<span style="color: blue;">●</span> <span style="color: green;">◆</span>	Lack of nutrients to the brain can impair growth and brain development.	Malnutrition can cause long-term cognitive effects during critical developmental periods and increase stress.
Soil and water contamination	<span style="color: magenta;">■</span>	Neurotoxins found in the environment accumulate to the top of the food chain (humans) and have neurodegenerative effects.	Lead to cognitive impairments, behavioural problems and developmental delays.
Vector-borne diseases	<span style="color: blue;">●</span> <span style="color: red;">◆</span> <span style="color: green;">◆</span> <span style="color: yellow;">▲</span> <span style="color: magenta;">■</span>	Inflammation, neurodegeneration, and birth defects, such as microcephaly.	Increased risk of mental health disorders (e.g., individuals diagnosed with Lyme disease).

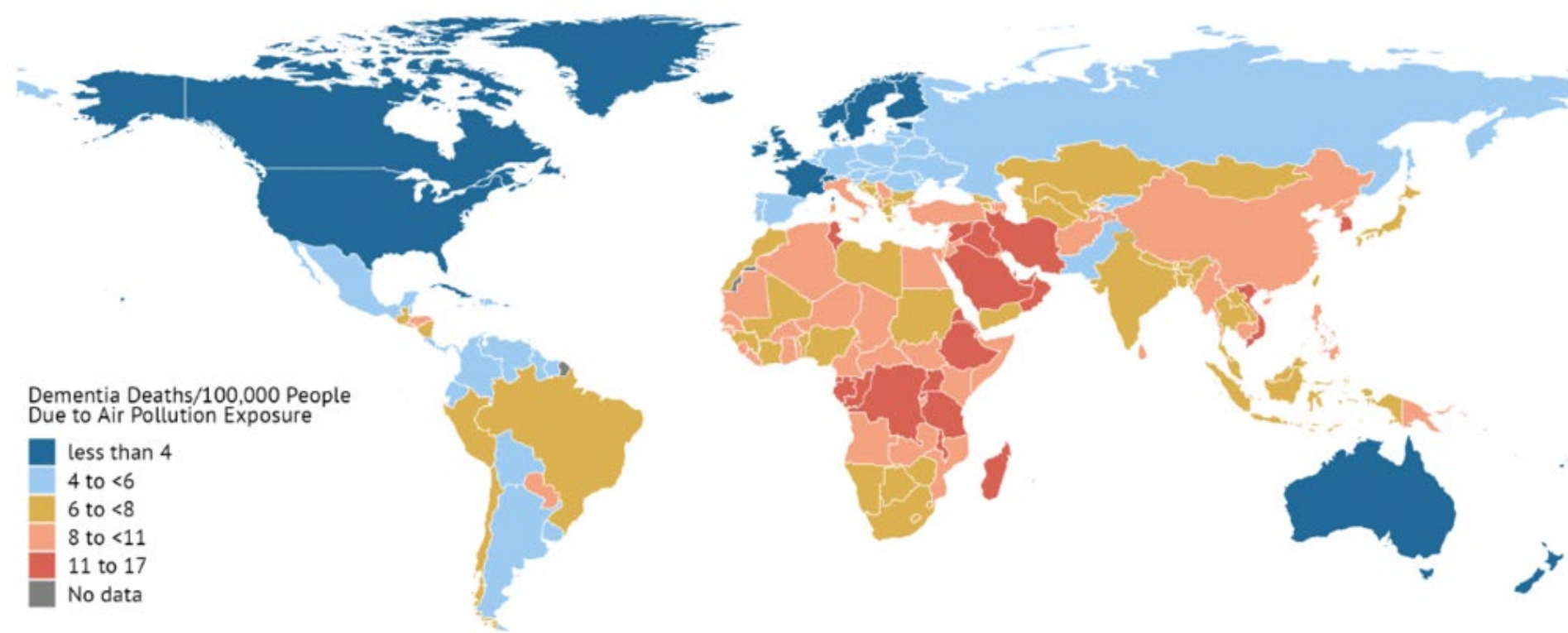
# Air Pollution: A major modifiable risk factor of NDs

- Air pollution is now recognized as a major contributor to neurodegeneration, including Alzheimer's disease, vascular dementia, and mild cognitive impairment.
- In 2023, approximately 626,000 dementia deaths globally (≈29%) were attributable to air pollution, with the highest burdens in Africa and Asia.
- Even modest pollution-related increases in dementia risk have large societal impacts, given population-wide exposure, aging demographics, and caregiving burdens, especially for women.


**Late adulthood**



Increased risk for cerebrovascular diseases and dementia



In 2023, **dementia** attributable to air pollution resulted in **626,000 deaths** and **11.6 million** healthy years of life lost.

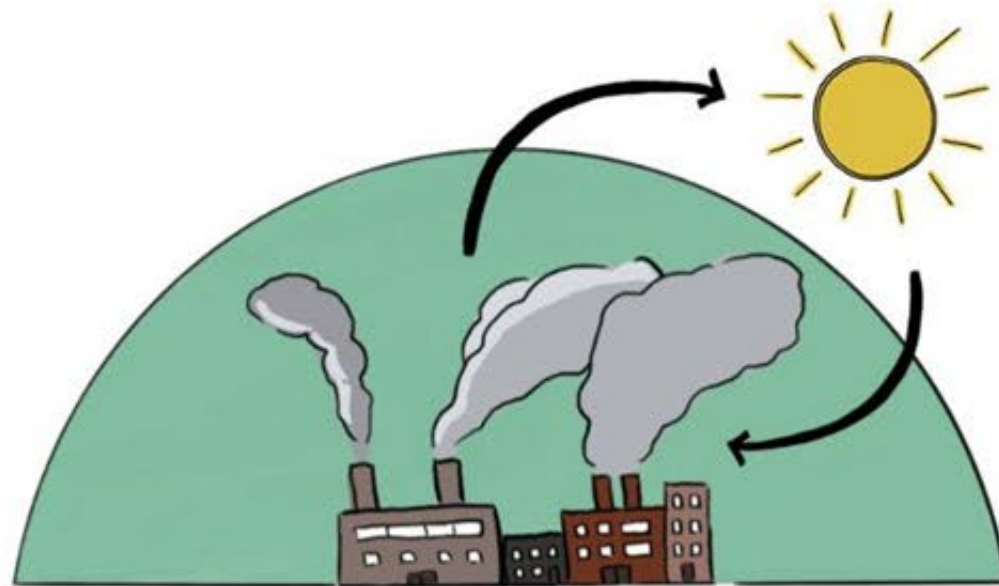


More than **1 in 4** dementia deaths

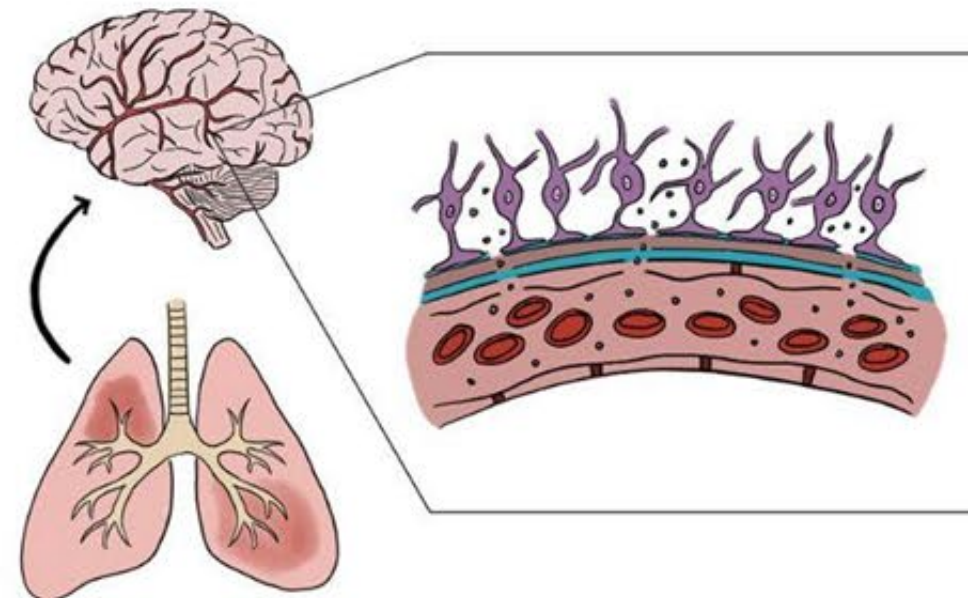
**FIGURE 17:** Age-standardized rates of death due to dementia attributable to air pollution in 2023.

# Mechanisms: how air pollution affects the ND risk

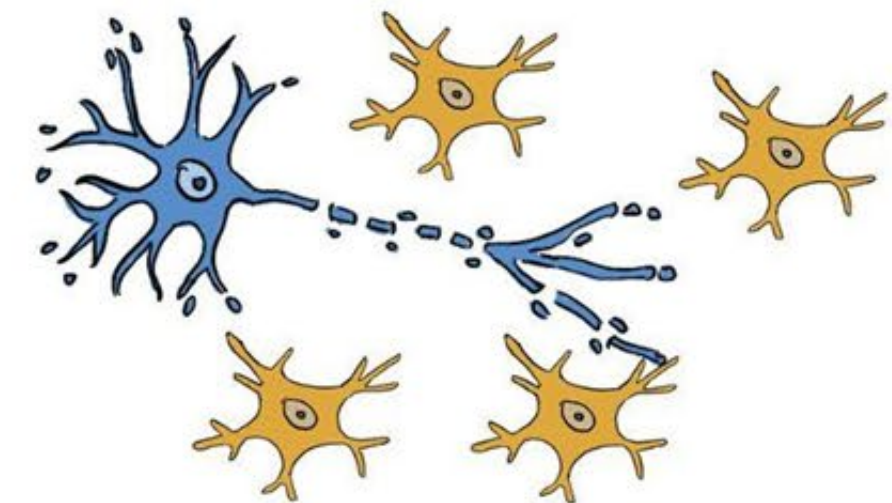
Some air pollution are greenhouse gases contributing to climate change, which in turn, can intensify air pollution.



Fine particles from air pollution can irritate the lungs causing systemic effects on the brain and penetrate the blood-brain barrier.



This can lead to inflammation and neurodegeneration.



Source: Neuro Climate Working Group

**Systemic inflammation:** Fine particles trigger inflammation and oxidative stress that damage brain cells

**Direct brain exposure:** Fine particles enter the bloodstream and reach the brain

**Blood-brain barrier disruption:** Pollution weakens the brain's protective barrier

**Vascular effects:** Increased heart diseases and stroke raise dementia risk

# Populations most at -risk

- **Older adults:** Higher risk of cognitive decline, dementia, and stroke.
- **People with pre-existing conditions:** Neurological, cardiovascular, or mental health disorders increase susceptibility.
- **Low-income and marginalized communities:** Higher exposure and fewer protective resources.

**Women:** Greater caregiving burden and higher dementia prevalence in later life.



# Challenges & knowledge gaps

- Synergistic impacts are poorly understood
- Incomplete mechanistic understanding
- Limited causal inference
- Confounding factors
- Inadequate or imprecise air pollution exposure assessment
- Underdiagnosis and surveillance gaps
- Equity gaps
- Low policy integration
- Fragmented systems



# Restoring environments, restoring minds



# Better air quality, better brain health

## PLOS MEDICINE

RESEARCH ARTICLE

Air quality improvement and cognitive decline in community-dwelling older women in the United States: A longitudinal cohort study

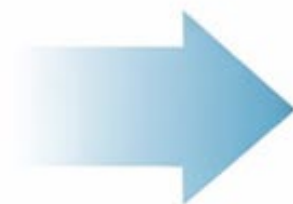
Diana Younan<sup>1,2,\*</sup>, Xinhui Wang<sup>2,3</sup>, Joshua Millstein<sup>1</sup>, Andrew J. Petkus<sup>2</sup>, Daniel P. Beavers<sup>3</sup>, Mark A. Espeland<sup>3</sup>, Helena C. Chui<sup>2</sup>, Susan M. Resnick<sup>4</sup>, Margaret Gatz<sup>5</sup>, Joel D. Kaufman<sup>6</sup>, Gregory A. Wellenius<sup>7</sup>, Eric A. Whitset<sup>8</sup>, JoAnn E. Manson<sup>9</sup>, Stephen R. Rapp<sup>10</sup>, Jiu-Chuan Chen<sup>1,2</sup>

<sup>1</sup> Department of Population and Public Health Sciences, University of Southern California, Los Angeles.

## Association of improved air quality with lower dementia risk in older women

Xinhui Wang<sup>a,1</sup>, Diana Younan<sup>b,2,1</sup>, Joshua Millstein<sup>b</sup>, Andrew J. Petkus<sup>a</sup>, Erika Garcia<sup>b</sup>, Daniel P. Beavers<sup>c</sup>, Mark A. Espeland<sup>c</sup>, Helena C. Chui<sup>a</sup>, Susan M. Resnick<sup>d</sup>, Margaret Gatz<sup>e</sup>, Joel D. Kaufman<sup>f,g,h</sup>, Gregory A. Wellenius<sup>i</sup>, Eric A. Whitset<sup>j,k</sup>, JoAnn E. Manson<sup>l</sup>, Stephen R. Rapp<sup>m,n</sup>, and Jiu-Chuan Chen<sup>a,b,2</sup>

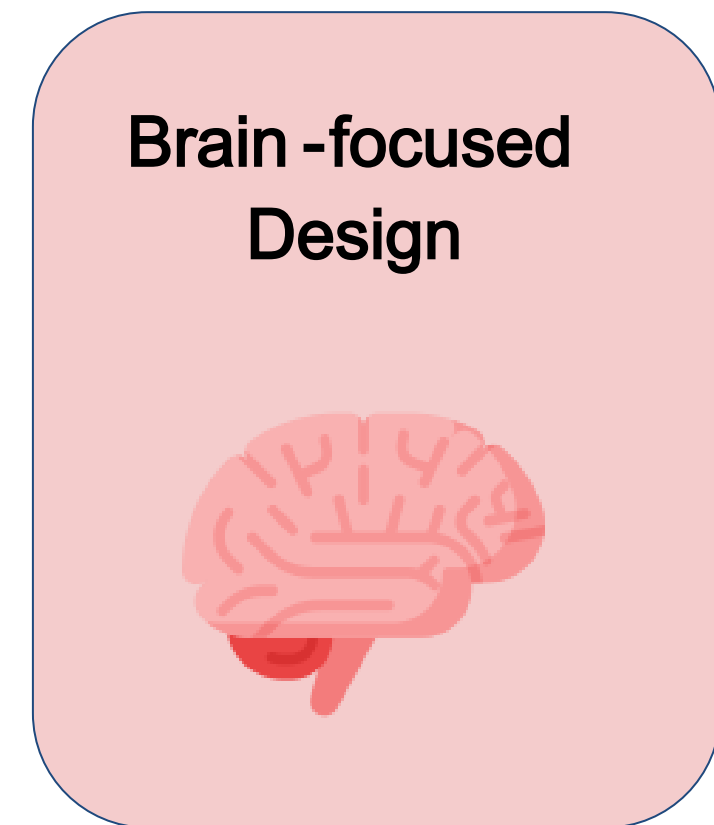
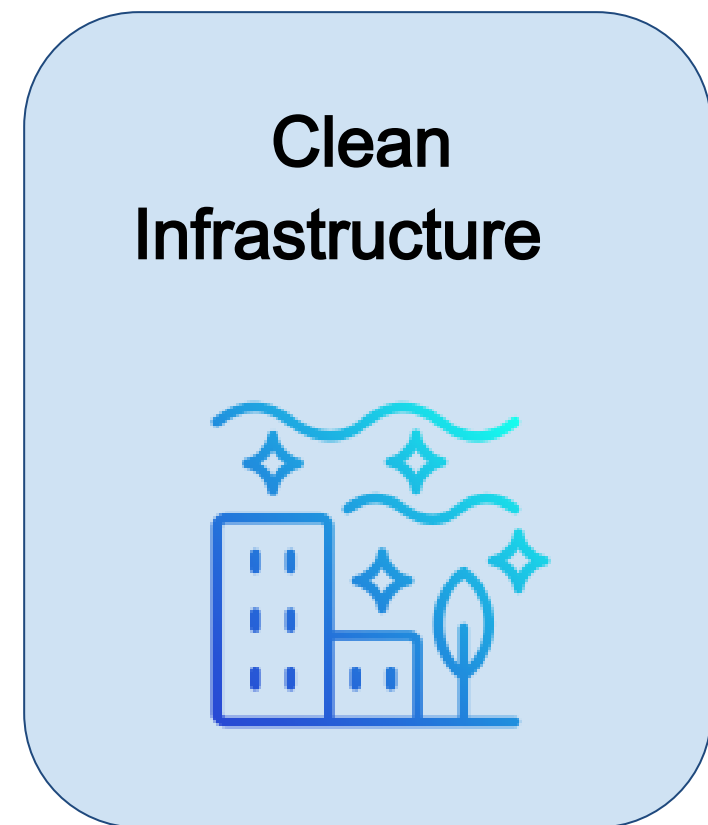
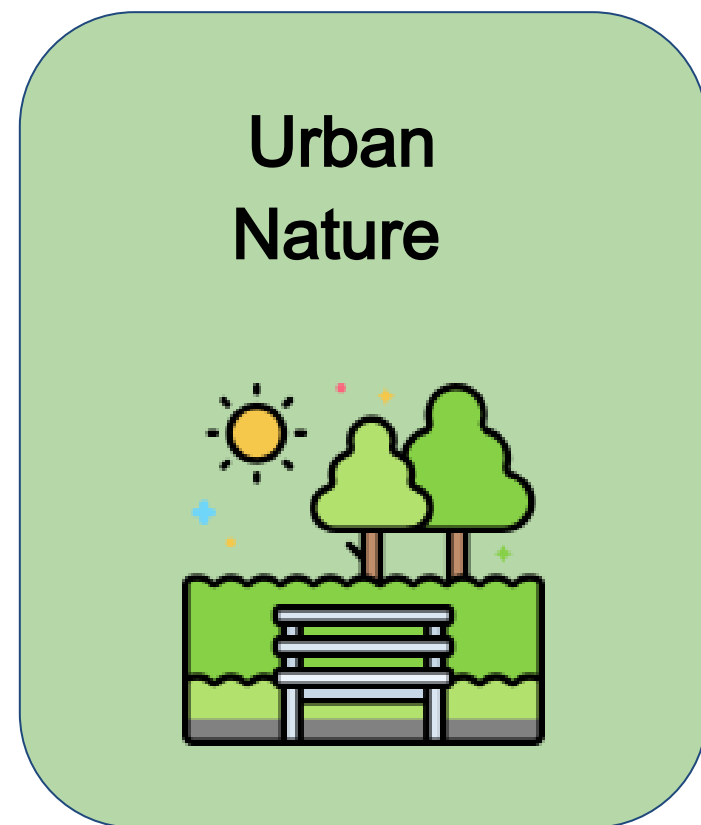
↓ Air Pollution  
(PM<sub>2.5</sub>, NO<sub>2</sub>)



## Brain Health Benefits

- ~20% lower dementia risk
- Slower cognitive decline
- ≈ 2~3 years younger cognitive age

# Brain health co-benefits of climate and clean air policies



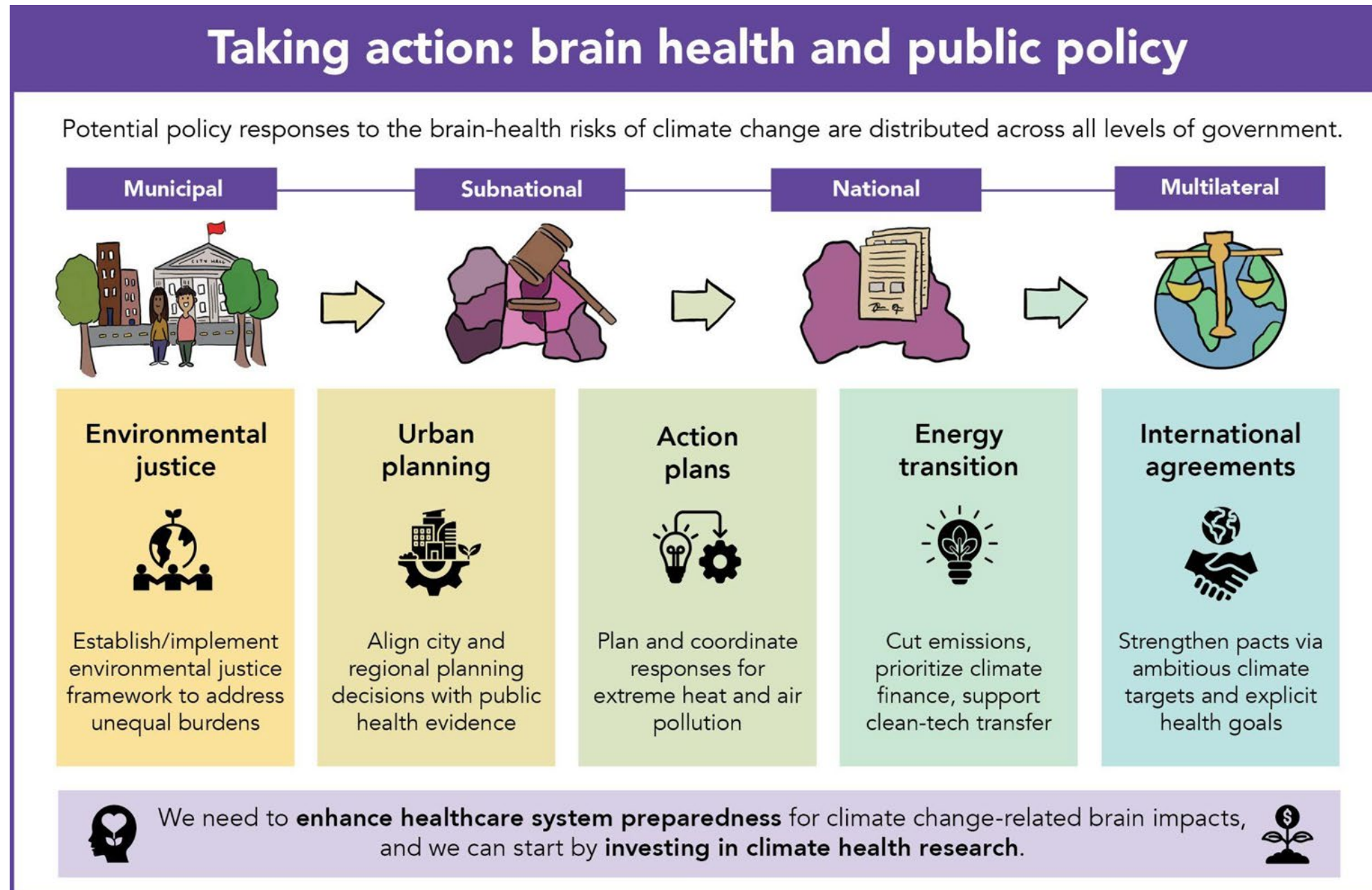
Cleaner energy → cleaner air → healthier brains

Heat mitigation → better sleep, learning, productivity

Resilient infrastructure → reduced trauma and stress

Safer environment → more outdoor activity and community interaction

# Policy-relevant solutions



- Clean energy transitions
- Heat standards for housing and workplaces
- Urban cooling and green infrastructure
- Health-inclusive climate and energy policy
- Data and metrics that include brain health outcomes

NCWG State-of-Science Report, Climate Change and Brain Health: From Evidence to Action  
(coming soon)

# Wanna learn more?



[neuroclimate.org](https://neuroclimate.org)

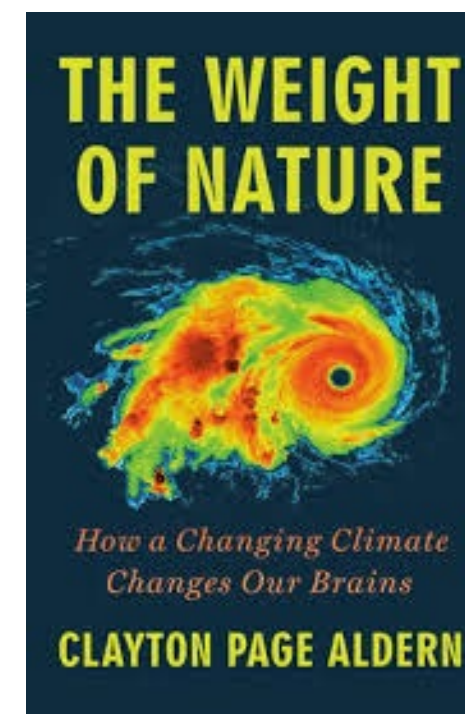
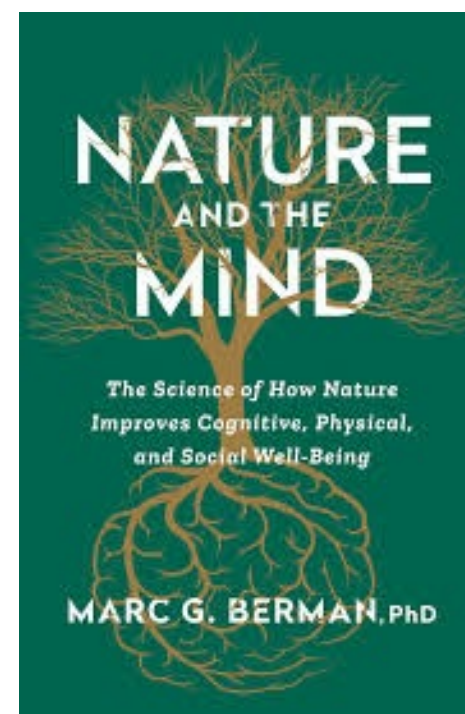
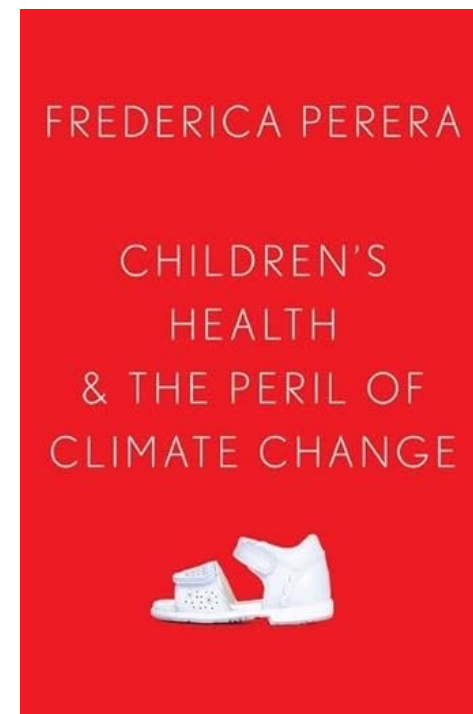
OPINION **FIRST OPINION**

## The hidden neurological toll of wildfires

It can take months, years, even decades to understand the damage done, especially to children



**FIRST  
OPINION**  
NEWSLETTER  
The smartest thinkers in life  
sciences on what's happening —  
and what's to come



# Thank you!

Interested in the intersection of climate change and brain health? Join our Neuro Climate Working Group.



Wanna connect?

Please email me at [burcin@stanford.edu](mailto:burcin@stanford.edu)