National Institute of Neurological Disorders and Stroke

OPEN STAGE webinar series **The Neural Exposome: How NINDS is supporting Exposomic Research**



OCTOBER 23, 2023 1:00 PM - 2:00 PM ET

Register: https://go.nih.gov/4KXBlnZ

Office of Programs to Enhance Neuroscience Workforce Diversity

Meeting Reminders

Attendees are muted



Presentation is being recorded

Recording...

Submit questions in Q&A box

P Q&A

Live transcript available



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National Institute of Neurological Disorders and Stroke

National Institutes of Health National Institute of Neurological Disorders and Stroke

Mission is to seek fundamental knowledge about the brain and nervous system and to use that knowledge to reduce the burden of neurological disease for all people.



NIH NINDS Office of Neural Exposome and Toxicology (ONETOX)





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The Exposome



- Genetics has provided insight into the etiologies of inherited disease
- But most health risk factors cannot be explained by genetics alone
- Exposome: environmental exposures affecting the genome (Wild, 2005)
- Now <u>all</u> nonheritable factors affecting gene expression across the lifespan are considered
- New frontier of biomedical research to complement the genome
- Unlock a more holistic approach to disease prevention and more effective and personalized interventions



NeuroView A focus on the neural exposome

Amir P. Tamiz,¹ Walter J. Koroshetz,² Neel T. Dhruv,¹ and David A. Jett^{1,*} ¹Division of Translational Research, National Institute of Neurological Disorders and Stroke, NIH, 6001 Executive Blvd., Rockville, MD 20852, USA ²National Institute of Neurological Disorders and Stroke, NIH, 31 Center Drive, 8A31, Bethesda, MD 20892, USA ^{*}Correspondence: jettd@nih.gov https://doi.org/10.1016/j.neuron.2022.03.019

Many neurological disorders have complex etiologies that include **noninheritable factors, collectively called the neural exposome**. The National Institute of Neurological Disorders and Stroke is developing a new office with **goals to advance our understanding of the multiple causes of neurological illness and to enable the development of more effective interventions**.





Can exposome research help us have a better understanding of how to fix health disparities ?

https://www.alz.org/professionals/publichealth/public-health-topics/healthequity#:~:text=Black%20Americans%20are%20abo ut%20two,have%20Alzheimer's%20and%20other %20dementias.



Black Americans are about two times more likely than White Americans to have Alzheimer's and other dementias.



By 2050, Asian Americans are projected to comprise nearly 8% of those aged 65 and older.



Hispanic Americans are about one and one-half times more likely than White Americans to have Alzheimer's and other dementias.



Native Americans have high rates of chronic conditions, including conditions that are suspected risk factors for Alzheimer's, such as obesity, diabetes and hypertension.



Almost two-thirds of those living with Alzheimer's are women.

Funded 12 projects

Clinical Relevance of the Linkage between Environmental Toxicant Exposures and Alzheimer's Disease and Related Dementias	PAR-22-048	
ECHO Program NOFOs		
Climate Change and Health	NOT-ES-22-006	*
Research on Biopsychosocial Factors of Social Connectedness and Isolation on Health, Wellbeing, Illness, and Recovery	PAR-21-349 PAR-21-350	
Role of Environmental Stress in the Health Inequities of Alzheimer's Disease-Related Dementias	RFA-NS-24-024	*
Impact of the Microbiome Gut-Brain Axis on Alzheimer's Disease and Alzheimer's Disease Related Dementias	PAR-22-211	
NINDS Mission Relevant Pain Research	NOT-NS-22-050	
Administrative Supplements to Support research Infrastructure on Exposome Studies in Alzheimer's disease (AD) and AD-Related Dementias (ADRD)	NOT-AG-22-022	
Center for Exposome Research Coordination to Accelerate Precision Environmental Health	RFA-ES-23-010	*

Neural Exposome Funding Opportunities

* Active , visit website for upcoming due dates

*

Neural Exposome Outreach and Collaboration

- Society for Neuroscience Event
- SOT Symposium
- Funding Opportunity Webinars
- Request for Information
- Workshops



Wednesday, March 22, 8:00 AM to 10:45 AM, Room 205, Music City Center

Symposium Session: Expanding Our Knowledge of Neurological Disease Etiologies: Current Research on the Neural Exposome

Chair(s): David Jett, NIH/NINDS; and Cindy Lawler, NIEHS/NTP. Primary Endorser: Neurotoxicology Specialty Section

NIH Request for Information (RFI) on Interdisciplinary Research Opportunities that Bridge Neuroscience and Environmental Health Science, NOT-NS-22-101

NASHVILLE

2023





Summary

- Most disease and disorders cannot be explained by genes alone.
- A more holistic approach includes endogenous, exogenous, and behavioral factors collectively called the exposome.
- Many neurological diseases and disorders are now considering exposomic factors in intervention strategies.
- Will require new tools and data analyses.
- Complexity includes how to integrate the exposomic factors and how to interpret these factors across the lifespan.
- Most health inequities are probably driven by exposomic factors.



One of Our First Grantees: Erica Glasper, PhD



Associate Professor Department of Neuroscience

Postdoctoal training: Princeton Neuroscience Institute

Doctoral training: The Ohio State University

From Social Disruption to Neural Compromise: Establishing Markers and Mediators

- PIs: Erica R. Glasper, PhD and Gretchen Neigh, PhD
- Research on Biopsychosocial Factors of Social Connectedness and Isolation on Health, Wellbeing, Illness, and Recovery
- Investigation into mechanistic link between social bond disruption and subsequent increases in neuroinflammation and compromised neural mitochondrial function



Researching Aspects of the Neural Exposome: One Stage at a Time

Dr. Erica R. Glasper The Ohio State University Wexner Medical Center

ERICA R. GLASPER, PH.D

- ASSOCIATE PROFESSOR
 - DEPARTMENT OF NEUROSCIENCE
 - INSTITUTE FOR BEHAVIORAL MEDICINE RESEARCH
- CO-DIRECTOR, DISCOVERY PREP, NIGMS FUNDED POST-BACCALAUREATE RESEARCH PROGRAM
- FACULTY MENTOR
 - NEUROSCIENCE GRADUATE PROGRAM
 - BIOLOGICAL SCIENCES GRADUATE PROGRAM



The Ohio State University



Hormones



0

Psychoneuroimmunology

WHAT ARE MY RESEARCH INTERESTS?

Social Neuroscience, Behavior, & Health Lab



Goal: To understand how neuroendocrine & neuroimmune processes alter social behavior and health.

MY RESEARCH TRAINING



1998-2002, Psychology and Biology, RANDOLPH-MACON COLLEGE

Advisor: Kelly G. Lambert



2002-2006: Psychobiology and Behavioral Neuroscience, THE OHIO STATE UNIVERSITY

PhD Advisor: A. Courtney Devries



2006-2011: Behavioral Neuroscience, PRINCETON UNIVERSITY

Postdoc Advisor: Elizabeth Gould



NIH FUNDING MECHANISMS PAVED THE WAY TOWARD MY SUCCESS

3 Summer Undergraduate Research Fellowships

NINDS Diversity Supplement

F31 (NINDS)

Merck-United Negro College Fund Postdoctoral Fellowship / F32 (NIA)

RO1 (NINDS)

HAD I BEEN STUDYING THE NEURAL EXPOSOME MY ENTIRE CAREER?















Been et al., 2022, Neuroscience and Biobehavioral Reviews

BASIC SCIENCE APPROACH TO SOCIAL BONDING & LONELINESS

California mouse (Peromyscus californicus)

Genetically monogamous

Strong mate bonds and bonds with offspring

Negative behavioral and physiological response to pair bond dissolution



Image © Mark A. Chappell

Manipulation of social bonds in the laboratory can facilitate examination of the impact of bonding and bond dissolution on behavior and physiology

1R01NS125589-01

From Social Disruption to Neural Compromise: Establishing Markers and Mediators

Hypothesis: The neural-glial response engaged by social bond disruption increases neuroinflammation and compromises neural mitochondrial function through disruptions in OT signaling

Pls:



Erica R. Glasper The Ohio State University



Gretchen N. Neigh Virginia Commonwealth University











National Institute of Neurological Disorders and Stroke

THE OHIO STATE UNIVERSITY COLLEGE OF MEDICINE

THE OHIO STATE UNIVERSITY CENTER FOR CLINICAL AND TRANSLATIONAL SCIENCE





Acknowledgement

University of Maryland

Molly Hyer, PhD Priyanka Agarwal, MS Nicole Palin, MS Allison Whitaker **Terrence Hunter** Luke Hallgarth Neilesh Sud Robyn Harper Jhansi Katakam Cyrus Ameri Collin Kaufman Amanda Holmes Allison Whitaker Samuel Doty, MD

The Ohio State University

Shakeera Walker, MS Amber Valentino, MA Janet Chen, MD Zachary Weisenseel JaNiya Ulysse Emily Oakley Noah Holmes Rita Beyene Jessie Bontatibus

University of Wisconsin-Madison Farrah Madison, PhD

Virginia Commonwealth University Gretchen Neigh, PhD

Opportunities for Exposome Research







<u>Data</u>

- > Human studies
- > Biobanks
- > Geospatial data
- > Wearables
- > Biomarkers

<u>Tools</u>

- > Data Bases (CTD)
- > Training
- > Analysis (HHEAR)
- > Screening
- > Omics technology

Team Science

- > Blueprint ICs
- > NIH programs
- > Consortia
- > CDC, EPA, DoD
- > Non-profits



