

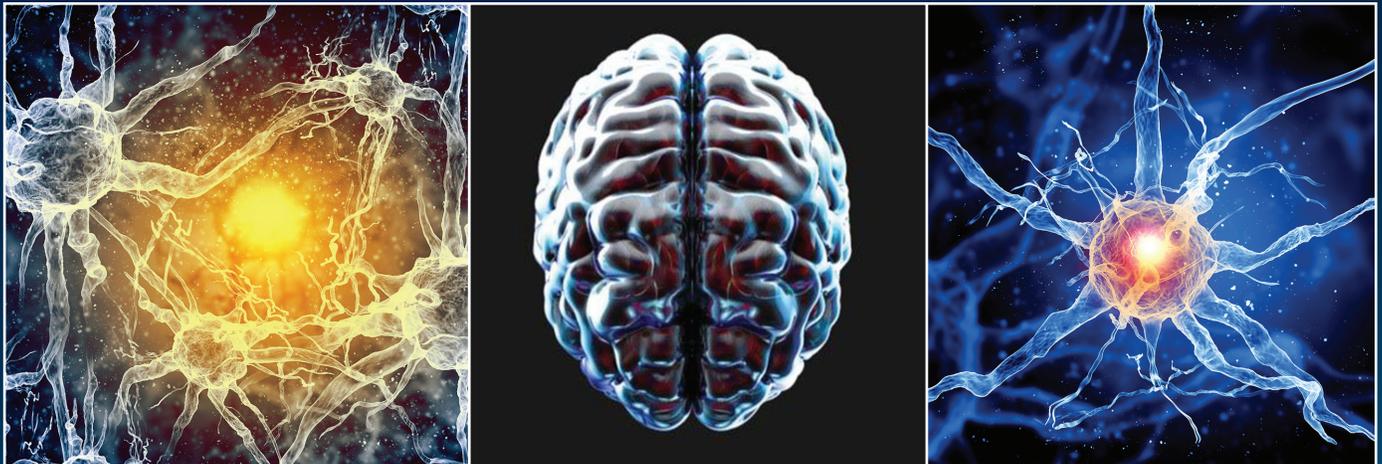
UNIFORMED SERVICES UNIVERSITY  
Center for the Study of Traumatic Stress

# PROGRAM

## 14th Annual Amygdala, Stress, and PTSD Conference: Risk, Resilience, and Recovery

APRIL 16, 2019

Sanford Auditorium & Lobby, Building B  
Uniformed Services University, Bethesda, MD



[www.AmygdalaPTSDconference.org](http://www.AmygdalaPTSDconference.org)

The Amygdala, Stress, and PTSD Conference at the Uniformed Services University brings together scientists and clinicians working towards solving the biological basis of stress, fear, and posttraumatic stress disorder.

SPONSORED BY:



The Center for the Study of Traumatic Stress (USU), Department of Psychiatry (USU),  
Neuroscience Program (USU), Department of Family Medicine (USU), and  
Department of Psychiatry (WRNMMC)



UNIFORMED SERVICES UNIVERSITY

Center for the Study of Traumatic Stress

---

# 14th Annual Amygdala, Stress, and PTSD Conference: Risk, Resilience, and Recovery

---

**APRIL 16, 2019**

**Sanford Auditorium & Lobby, Building B  
Uniformed Services University  
Bethesda, MD**



**[www.AmygdalaPTSDconference.org](http://www.AmygdalaPTSDconference.org)**

**SPONSORED BY:**

**The Center for the Study of Traumatic Stress (USU), Department of Psychiatry (USU),  
Neuroscience Program (USU), Department of Family Medicine (USU), and  
Department of Psychiatry (WRNMMC)**

**CSTS**



## Background

The Center for the Study of Traumatic Stress (CSTS) of the Uniformed Services University (USU), in collaboration with the USU Department of Psychiatry, USU Neuroscience Program, USU Department of Family Medicine, and the Walter Reed National Military Medical Center (WRNMMC), Department of Psychiatry, is pleased to present the *14th Annual Amygdala, Stress, and PTSD Conference: Risk, Resilience, and Recovery*.

The Amygdala, Stress, and PTSD Conference at the Uniformed Services University brings together scientists and clinicians working towards solving the biological basis of stress, fear, and posttraumatic stress disorder.

# Table of Contents

Agenda .....	3
Conference Speakers .....	4
Moderators .....	7
Conference Leadership .....	8
Conference Committee .....	10
Conference Posters .....	11
Continuing Education Credit .....	40



## AGENDA

---

0800-0900	Registration and Poster Review
0900-0905	Conference Announcements — <b>Derrick A. Hamaoka, MD</b>
0905-0915	Welcome and Introduction — <b>Robert J. Ursano, MD</b>
0915-1000	<i>Wake up to Sleep! A Translational Perspective of the Role of Sleep in Readiness and Resilience</i> — <b>Anne Germain, PhD</b>
1000-1045	<i>Resilience: The Science of Mastering Life's Greatest Challenges</i> — <b>Dennis S. Charney, MD</b>
1045-1115	Coffee Break and Poster Review in Lobby
1115-1145	Discussion Panel 1 — Moderator — <b>COL Christopher H. Warner</b>
1145-1245	Lunch
1245-1330	<i>Gene-activity and Proteins that Relate to Chronic PTSD Symptoms</i> — <b>Jessica M. Gill, PhD, RN</b>
1330-1415	<i>Mobilizing Hope in the Face of Despair: Applying Social Neuroscience Research in Brief Clinical Encounters</i> — <b>James L. Griffith, MD</b>
1415-1445	Coffee Break and Poster Review in Lobby
1445-1530	<i>Preclinical Development of Ketamine and the Metabolite 2R,6R-Hydroxynorketamine for Depression and Other Disorders</i> — <b>Irwin Lucki, PhD</b>
1530-1600	Discussion Panel 2 — Moderator — <b>COL Wendi Waits</b>
1600-1615	Closing Remarks and Presentation of Travel Award — <b>Robert J. Ursano, MD</b>

---

## Conference Speakers

### Dennis S. Charney, MD



Dennis S. Charney, MD, is Anne and Joel Ehrenkranz Dean of the Icahn School of Medicine at Mount Sinai and President for Academic Affairs for the Mount Sinai Health System. Dr. Charney is a world expert in the neurobiology and treatment of mood and anxiety disorders, making

fundamental contributions to the understanding of the causes of human anxiety, fear, and depression, and the discovery of new treatment for mood and anxiety disorders. His research on depression has led to discovery of new and novel therapies for treatment resistant depression including Ketamine and the first digital treatment for depression (EFMT). He has been honored with all of the major awards in his field for his scientific research, including World's Most Influential Scientific Minds 2014 and 2015, Ranked 48 out of 1,360 of Most Highly Cited Life Science Researchers in the World. His discovery of Ketamine for Treatment-Resistant Depression was named by Cleveland Clinic on its Top 10 list of 2017 Health Care Innovations. He holds 3 U.S. Patents, and 19 U.S. and Foreign Patent Applications, 10 of which are licensed to 2 companies. He has published 785 articles and book chapters, and 16 books, including *Resilience: The Science of Mastering Life's Greatest Challenges*, and *Charney & Nestler's Neurobiology of Mental Illness 5th Edition*. Charney was elected to the National Academy of Medicine in 2000, and the National Academy of Inventors in 2017.

### Anne Germain, PhD



Dr. Anne Germain is Professor of Psychiatry at the University of Pittsburgh School of Medicine at Western Psychiatric Institute and Clinic, and is Director of the University of Pittsburgh Sleep and Behavioral Neuroscience Center. She is Director of Military Sleep Tac-

tics of Resilience Research Team. At the University of Pittsburgh, Dr. Germain also holds secondary appointments in Psychology, and in Clinical and Translational Science.

Dr. Germain received a Bachelor of Science in psychology from McGill University in 1996, and completed her Ph.D. in Clinical Psychology from the Université de Montréal in 2001. She then pursued post-doctoral training in clinical sleep research and sleep neuroimaging at the University of Pittsburgh, and joined the Faculty in the Department of Psychiatry in 2005.

Dr. Germain's research program has two main areas of interest. A first area of interest focuses on the neural underpinnings and effects of acute sleep loss and chronic sleep disturbances occurring in the context of stress-related psychiatric disorders, with a special emphasis on posttraumatic stress disorder (PTSD) in military populations. To accomplish this, she uses multimodal sleep measurement methods including self-report measures, actigraphy and polysomnography, quantitative EEG, pharmacological probes, sleep neuroimaging techniques, as well as novel animal models. A second area of interest focuses on the development, adaptation, testing, and implementation of treatments targeting trauma-related sleep disturbances to enhance psychological resilience and to hasten recovery from trauma exposure.

Dr. Germain has published over 120 peer-reviewed articles, and recently co-edited a book entitled, *Sleep and Combat-Related PTSD* (Spinger, 2018). She is also the author of 25 book chapters and invited papers on insomnia, nightmares, treatments of sleep disorders, and sleep in the context of PTSD and other trauma-related disorders. Her h-index is 36. She has served on various committees of the Sleep Research Society and Ameri-

## Conference Speakers. Continued

can Academy of Sleep Medicine. She currently serves on the Editorial Board of the journal Behavioral Sleep Medicine, and is a regular peer reviewer for specialized journals on sleep, trauma, and psychiatry. She has served and continues to serve on various study sections for the Department of Defense, the NIH, the Canadian Institutes of Health Research, and the Department of Veterans Affairs.

### Jessica M. Gill, PhD, RN



Dr. Gill is a tenure track investigator at the National Institutes of Health (NIH) and co-director of the biomarkers core for the Center for Neurosciences and Regenerative Medicine. Dr. Gill has an established clinical and laboratory infrastructure to examine the biological mechanisms

of traumatic brain injury (TBI), and related comorbidities including posttraumatic stress disorder (PTSD), post-concussive disorder (PCD), depression and neurological deficits.

Current projects include a project coordinated with Madigan Army Medical Center. Findings include alterations in tau and amyloid beta in acute and chronic TBI patients using the SIMOA system and alterations in sleep regulatory proteins and the activity of genes that regulate sleep in patients with brain injuries. Other collaborations include analyses of epigenetic modifications in athletes with repeated TBIs. Dr. Gill plans to use both the insights and infrastructure from current projects to initiate this novel project to address the critical issue of the molecular mechanisms of TBI-related symptoms in patients with repeated injuries, and the role of sleep in these biomarkers and patient outcomes. The project is expected to initiate a program of research that will identify novel interventions to treat TBI related impairments to address this critical issue in patients with TBIs.

## Conference Speakers. Continued

### James L. Griffith, MD



James L. Griffith, M.D. is the Leon M. Yochelson Professor and Chair in the Department of Psychiatry and Behavioral Sciences at the George Washington University School of Medicine and Health Sciences. As a psychiatric educator, Dr. Griffith has developed treatment methods that use

cognitive and social science research to mobilize hope, strengthen resilience, and reduce stigmatization and social exclusion, reducing suffering and promoting mental health more effectively. Dr. Griffith has published extensively on family-centered treatment of psychosomatic disorders and chronic medical illnesses, including a book, *The Body Speaks: Therapeutic Dialogues for Mind-Body Problems*. His most recent book, *Religion that Heals, Religion that Harms*, addressed destructive uses of religion and ideology in clinical settings and received the Creative Scholarship Award from the Society for the Study of Psychiatry and Culture.

Dr. Griffith provides psychiatric care for immigrants, refugees, and survivors of political torture at Northern Virginia Family Services in Falls Church, VA. He has received the Human Rights Community Award from the United Nations Association of the National Capital Area and the Margaret B. and Cyril A. Schulman Distinguished Service Award from the George Washington University Medical Center, both for the training of mental health professionals and development of mental health services for survivors of political torture. He has received the Distinguished Teacher Award from the George Washington University School of Medicine and the 2003 Psychiatrist of the Year and 2014 Distinguished Service Award from the Washington Psychiatric Society. He was the 2017 recipient of the Oskar Pfister Award from the American Psychiatric Association for his contributions to the field of religion and psychiatry.

### Irwin Lucki, PhD



Irwin Lucki, Ph.D. is Chair of the Department of Pharmacology & Molecular Therapeutics at the Uniformed Services University of the Health Sciences (USUHS). Dr. Lucki received his Ph.D. (Biopsychology) from the University of Iowa in 1979. He conducted postdoctoral research in psychopharmacology at the

University of Pennsylvania in 1979-1982 before joining the faculty in 1984 as a researcher, an educator and Program Director of an NIMH predoctoral and postdoctoral training grant. He joined USUHS in 2016 as Professor of Pharmacology and Psychiatry.

The major focus of Dr. Lucki's research is the investigation of neural mechanisms underlying the behavioral effects of psychiatric medications and translation to clinical development. His laboratory develops new therapeutic approaches for treating depression and anxiety disorders based on studies involving relevant animal models and neurochemistry. Dr. Lucki started his career by studying the behavioral and physiological functions of different serotonin receptors and their relationship to the effects of SSRIs, eventually directing an NIMH Program Project Grant. Dr. Lucki also directed an NIMH Center grant on drug discovery in collaboration with industry partners. His research led to the development of vilazodone as an FDA-approved clinical antidepressant. He is currently investigating opioid receptor antagonists and ketamine and its metabolites for the rapid treatment of depression and other disorders. He has also conducted clinical pharmacology research studies with psychiatric patients and normal volunteers. Dr. Lucki is author of over 180 peer-reviewed publications and 40 reviews. Among awards for his research, he received the Young Psychopharmacologist Award from the American Psychological Association (1984) and the Distinguished Investigator Award from NARSAD (2005). Dr. Lucki is a Principal Editor for *Psychopharmacology*, and on the Editorial Boards for *Neuropsychopharmacology*, *Journal of Psychopharmacology* and *Neurobiology of Stress*. Dr. Lucki is a fellow of the American College of Neuropsychopharmacology (ACNP) and the American Psychological Association (APA).

## Moderators

### Colonel Wendi Waits



Colonel Wendi Waits was born in Portland, Oregon, and calls Oregon and California home. She received her bachelor's degree and commission from the United States Military Academy in 1994. She received her doctorate in medicine from the Uniformed Services University of the Health Sciences

(USUHS) in 1998 and completed a psychiatry residency at the National Capital Consortium (NCC) in 2002.

Her military assignments have included the following: 2ID Division Psychiatrist, Camp Casey, Korea; Assistant Chief and subsequently Chief of Inpatient Psychiatry, Tripler Army Medical Center (TAMC), Honolulu, HI; Chief of Behavioral Health Services, Schofield Barracks, Hawaii; Director for Behavioral Health, Fort Belvoir Community Hospital (FBCH), Fort Belvoir, VA; and Chief of Adolescent Inpatient Behavioral Health, FBCH. COL Waits has served as the Director for Behavioral Health, Walter Reed National Military Medical Center (WRNNMC), since July 2018.

Colonel Waits deployed to Iraq in 2006, where she served as Fitness Section Leader, 1972nd Combat Operational Stress Control (COSC) detachment, and to Afghanistan in 2010, where she served as Clinical

Operations (CLINOPS) OIC, 254th COSC detachment. She has supervised interns, residents, and other trainees as a member of the academic faculty at TAMC, Schofield Barracks, FBCH, and WRNNMC and has been an examiner for the American Board of Psychiatry and Neurology. COL Waits is a Clinical Associate Professor of Psychiatry at USUHS, a Distinguished Fellow of the American Psychiatric Association, a member of the Order of Military Medical Merit, and a recipient of the MEDCOM "A" Designator.

Colonel Waits' military education includes the AMEDD Officer Basic Course, the AMEDD Officer Advanced Course, and the Command and General Staff College Intermediate Level Education core course. Her awards and decorations include the Defense Meritorious Service Medal (1 award), Meritorious Service Medal (3 awards), Joint Service Commendation Medal (1 award), Army Commendation Medal (4 awards), Joint Service Achievement Medal (1 award), Army Achievement Medal (3 awards), National Defense Service Medal (1 service star), Global War on Terrorism Service Medal, Iraq Campaign Medal, Afghanistan Campaign Medal, Korean Defense Service Medal, Overseas Service Ribbon (5 awards), and the North Atlantic Treaty Organization Medal. Her badges include the Air Assault Badge and the Army Physical Fitness Badge.

### Colonel Christopher H. Warner



Christopher H. Warner, COL, MC, USA, FAPA, FAAFP. COL Warner is currently a Student and Class President at the National War College in Washington, DC. He is a graduate of the U.S. Military Academy and Uniformed Services University of the Health Sciences. He completed

his residency training in Family Practice and Psychiatry at Walter Reed Army Medical Center where he was named the General Graves B. Erskine Award winner. COL Warner has commanded Winn Army Community Hospital at Fort Stewart, Georgia and the 61st Multi-

functional Medical Battalion at Fort Hood, Texas. He served as the Consultant to The US Army Surgeon General for Psychiatry from 2012 to 2018. COL Warner has deployed twice in support of Operation Iraqi Freedom, where he was noted for his efforts in pushing behavioral health care forward to maneuver units and was named a recipient of the 2006 Surgeon General's Physician Recognition Award. COL Warner has published two books on military mental health and primary authored more than forty articles including recent entries in the *American Journal of Psychiatry*, *the Annals of Psychiatry*, and *Lancet*. His service and achievement have led to his selection to the Order of Military Medical Merit and the Army A Designator. Additionally, COL Warner is on the editorial board of *Academic Psychiatry* and the American College of Psychiatrists.

## Conference Leadership

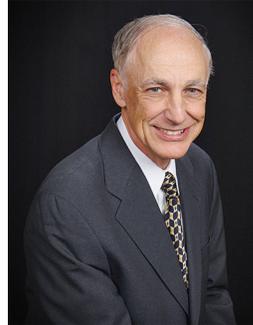
### Derrick A. Hamaoka, MD



Col (Dr.) Derrick A. Hamaoka serves as the Assistant Chair, Medical Education, for the Uniformed Services University of the Health Sciences Department of Psychiatry. In this capacity, he oversees the entire psychiatry education portfolio for the medical school to include the preclinical, clinical,

and advanced clinical rotations as well as the combat/operational psychiatry component for the medical school's field exercise, OPERATION BUSHMASTER. Col Hamaoka is a graduate of the Uniformed Services University of the Health Sciences School of Medicine (1999) and the University of Texas Health Science Center Psychiatry Residency Program (2003). Prior to serving in his current position, he was the Associate Program Director, University of Texas Health San Antonio Psychiatry Residency Program, leading one of the largest programs in the nation and responsible for the majority of the active duty Air Force psychiatry pipeline. He holds the Air Force Medical Corps Academic Grand Master (ME) Special Experience Identifier (SEI). He also serves as the Defense Institute for Medical Operations director and subject matter expert for the Mental Health Services After Disasters & Combat course, providing support/education for recent missions to Iraq, Sierra Leone, Tunisia, Colombia, Mexico, and Slovakia.

### Robert J. Ursano, MD



Dr. Ursano is Professor of Psychiatry and Neuroscience at the Uniformed Services University of the Health Sciences, Bethesda, Maryland and Director of the Center for the Study of Traumatic Stress. He served as Chairman of the Department of Psychiatry at USU for 25 years. In addition, Dr.

Ursano is Editor of *Psychiatry*, the distinguished journal of interpersonal and biological processes, founded by Harry Stack Sullivan. Dr. Ursano completed twenty years of service in USAF medical corps and retired as Colonel in 1991. He was educated at the University of Notre Dame and Yale University School of Medicine and did his psychiatric training at Wilford Hall USAF Medical Center and Yale University.

Dr. Ursano served as the Department of Defense representative to the National Advisory Mental Health Council of the National Institute of Mental Health and is a past member of the Veterans Affairs Mental Health Study Section and the National Institute of Mental Health Rapid Trauma and Disaster Grant Review Section. He is a Distinguished Life Fellow in the American Psychiatric Association and a Fellow of the American College of Psychiatrists. Dr. Ursano was the first Chairman of the American Psychiatric Association's Committee on Psychiatric Dimensions of Disaster. This work greatly aided the integration of psychiatry and public health in times of disaster and terrorism. Dr. Ursano was an invited participant to the White House Mental Health Conference in 1999. He has received the Department of Defense Humanitarian Service Award, the Lifetime Achievement Award of the International Society for Traumatic Stress Studies, and the American Psychiatric Association's Bruno Lima Award in Disaster Psychiatry. He is the recipient of the William C. Porter

*Continued on page 9*

*Robert J. Ursano, MD, Continued*

Award from the Association of Military Surgeons of the United States, the William Menninger Award of the American College of Physicians and the James Leonard Award of the Uniformed Services University. He is a frequent advisor on issues surrounding psychological response to trauma to the highest levels of the US Government and specifically to the Department of Defense leadership.

Dr. Ursano has served as a frequent member of the National Academies of Science, Institute of Medicine Committees and working groups including the Committee on Psychological Responses to Terrorism, Committee on PTSD, the Committee on Compensation for PTSD in Veterans and the Committee on Nuclear Preparedness; and the National Institute of Mental Health Task Force on Mental Health Surveillance After Terrorist Attack. In addition, he has served as a member of scientific advisory boards to the Secretary of Health and Human Services for disaster mental health and the Centers for Disease Control for preparedness and terrorism. Dr. Ursano is co-principal investigator of the largest NIMH grant ever given for the study of suicide in the U.S. Army. In collaboration with his co-principal investigators at Harvard University, the University of Michigan and the University of California, San Diego, the Army STARRS and STARRS-LS grants will be the Framingham Study of suicidal behavior, and address a national as well as DoD mental health need. In 2014, Dr. Ursano and Dr. Matthew Friedman of the VA National Center for PTSD co-founded the Friedman-Leahy Brain Bank supported through Senator Patrick Leahy (D-VT). It is the first human brain bank dedicated to PTSD. This joint effort of many people was a 12 year project developing concepts, pilot data and support. Dr. Ursano has over 300 publications and is co-author or editor of eight books.

## Conference Committee

**Derrick A. Hamaoka, MD, 2019 Chairman**

Col, USAF, MC, FS  
Assistant Chair and Associate Professor  
Department of Psychiatry  
Scientist  
Center for the Study of Traumatic Stress  
Uniformed Services University of the Health Sciences

**David Mears, PhD, 2019 Co-Chairman**

Associate Professor  
Department of Anatomy, Physiology, and Genetics  
Uniformed Services University of the Health Sciences

**Gary H. Wynn, MD, 2019 Scientific Co-Chairman**

LTC, MC, USA  
Assistant Chair and Professor  
Department of Psychiatry  
Scientist  
Center for the Study of Traumatic Stress  
Uniformed Services University of the Health Sciences

**Kwang Choi, PhD**

Assistant Professor  
Department of Psychiatry  
Scientist  
Center for the Study of Traumatic Stress  
Uniformed Services University of the Health Sciences

**Kelly L. Cozza, MD**

Associate Professor  
Department of Psychiatry  
Scientist  
Center for the Study of Traumatic Stress  
Uniformed Services University of the Health Sciences

**Eric Meyer, MD, 2019 CME Chair**

Maj, USAF, MC  
Assistant Professor  
Department of Psychiatry  
Scientist  
Center for the Study of Traumatic Stress  
Uniformed Services University of the Health Sciences

**James A. Naifeh, PhD**

Research Assistant Professor  
Department of Psychiatry  
Scientist  
Center for the Study of Traumatic Stress  
Uniformed Services University of the Health Sciences

**Holly H. Mash, PhD**

Research Assistant Professor  
Department of Psychiatry  
Scientist  
Center for the Study of Traumatic Stress  
Uniformed Services University of the Health Sciences

**Ioana M. Horotan-Enescu, MD**

Resident, PGY-3  
NCC Psychiatry Residency Program  
Walter Reed National Military Medical Center

**Amy E. Steward, TSgt, USAF**

Center for the Study of Traumatic Stress  
Uniformed Services University of the Health Sciences

**Katherine Pokorny, BA**

Research Assistant  
Center for the Study of Traumatic Stress  
Uniformed Services University of the Health Sciences

**We extend special thanks to:**

Robyn Hulvey, CMP, CGMP  
Assistant Director of Meetings  
Office of Education & Meetings

Henry M. Jackson Foundation for the Advancement of  
Military Medicine

*Thank you to the Henry M. Jackson Foundation for  
their kind support for the reception.*

## Conference Posters

### Presented in the Lobby

Impact of the Previous Night’s Sleep Disturbances on Post Traumatic Stress Symptoms among Individuals With and Without PTSD .....	13
Incidence Rates of Posttraumatic Stress Disorder in the US Military (2007-2015): Trends Across Military-Specific Variables.....	14
Risk and Resilience of Lifetime Stressful Experiences among U.S. Army Soldiers at Risk for Suicide.....	15
Inflammatory Cytokines and Cardiovascular Disease are Associated with Neuroimaging Findings after Acute Mild Traumatic Brain Injury.....	16
Elevated Tau Relates to Persistent PTSD, Depression and Post-Concussive Symptoms in Military Personnel with History of Traumatic Brain Injury.....	17
Prevalence of Medical Conditions and Healthcare Utilization in Treatment-Seeking, Bereaved Military Widows: A Prospective, Case-Controlled, Longitudinal Study .....	18
The Impact of Child Health on Parental PTSD .....	19
Whole-Genome Sequencing of 1,688 Veteran Twins to Detect Risk and Measure Rare Variation Associated with Major Depressive Disorder .....	20
Antidepressant Activity of the Selective Kappa Opioid Receptor Antagonist JNJ-67953964 in Mice .....	21
Safety and Confidence in Law Enforcement During Terrorist-Related Events: Association with Daily Life Activities.....	22
Effect of Prior Hurricane Experience on Preparedness in Florida Department of Health Workers .....	23
The 3 Predator Exposure Model as a Robust Animal Model of Traumatic Stress. ....	24
Does Damage to the Medial Frontal Gyrus in Chronic Mild TBI Patients Affect Post Traumatic Stress Symptoms? .....	25
Longitudinal Changes in Children’s Mental Health and Physical Injury Rates Following Father Death .....	26
Influence Of Attentional Bias on Scanpaths In Posttraumatic Stress Disorder: Assessment of Vigilance-Avoidance Scanning Patterns During Visual Search Paradigm .....	27

*Continued*

Mild Traumatic Brain Injury with Concurrent PTSD is Associated with Peripheral Tau Concentrations.....	28
The Relationship of PTSD to Anxiety and Depression: An Examination of the Moderating Effect of Sleep Quality and Nightmares.....	29
A Population Study of Incidence Rates of Posttraumatic Stress Disorder in the US Military (2007-2015): Analysis of Age, Sex, and Marital Status.....	30
Sex-related Differences of Intravenous Ketamine Infusion on Stress Hormone and Fear Memory in Rats.....	31
Gene Expression Differences in PTSD Are Uniquely Related to the Intrusion Symptom Cluster: A Transcriptome-Wide Analysis in Military Service Members .....	32
Nightmare Deconstruction and Reprocessing: Proof of Concept of a Novel Treatment for PTSD-Related Nightmares and Insomnia .....	33
Region- and Time-Dependent Gene Regulation in the Amygdala and Anterior Cingulate Cortex of a PTSD-Like Mouse Model.....	34
The Effects of Comorbid TBI and PTSD Symptoms among U.S. Active Duty Service Members on an Objective Cognitive Performance Measure.....	35
Fear Memory Extinction is Blocked In Rats Following Acute Glial Cell Inhibition: Implications for Targeting Neuroinflammation in Traumatic Stress Exposure.....	36
Sex Differences in Cued Fear Responses and Parvalbumin Cell Density in the Hippocampus Following Repetitive Concussive Brain Injuries in C57BL/6J Mice.....	37
Influence of Number of Awakenings during the Previous 1-3 Nights on Post Traumatic Stress Symptoms Among Individuals with PTSD .....	38
Sex-Related Differences of Mitochondrial DNA Copy Number in Active Military Service Members with PTSD .....	39

## Impact of the Previous Night's Sleep Disturbances on Post Traumatic Stress Symptoms among Individuals With and Without PTSD

### Authors

Quinn M. Biggs, Ph.D., M.P.H.,<sup>1</sup> Robert J. Ursano, M.D.,<sup>1</sup> Jing Wang, Ph.D.,<sup>1</sup> Gary H. Wynn, M.D.,<sup>1</sup> David S. Krantz, Ph.D.,<sup>1</sup> Russell B. Carr, M.D.,<sup>1,2</sup> Nicole M. Dacuyan, B.S.,<sup>1</sup> Katherine Pokorny, B.A.,<sup>1</sup> Ametisse Gover-Chamlou, B.A.,<sup>1</sup> Samantha Martinez, B.A.,<sup>1</sup> and Carol S. Fullerton, Ph.D.<sup>1</sup>

### Author Affiliations

1. Center for the Study of Traumatic Stress, Department of Psychiatry, Uniformed Services University of the Health Sciences, 4301 Jones Bridge Road, Bethesda, MD 20814
2. Walter Reed National Military Medical Center, 8901 Wisconsin Avenue, Bethesda, MD 20889

### ABSTRACT

**Background:** Sleep problems and Post Traumatic Stress Disorder (PTSD) are highly related, but it is not known whether sleep problems affect Post Traumatic Stress Symptoms (PTSS) on a daily basis. This study examined the relationship between the previous night's sleep disturbances and the following day's PTSS among individuals with and without PTSD.

**Methods:** Current and former U.S. service members ( $N = 80$ ) were assessed for probable PTSD at enrollment. PTSS were assessed four times daily and sleep disturbances were assessed once daily by self-report for 15 days using an ecological momentary assessment methodology. Eighteen items from the PTSD Checklist for DSM-5 were used to measure PTSS.

Nine items from the Pittsburgh Sleep Quality Index (PSQI) were adapted to measure sleep disturbances. Linear mixed models with control for demographic characteristics were applied to examine the association between sleep disturbances and PTSS. An interaction of sleep disturbances by PTSD group tested whether the association differed between individuals with and without PTSD.

**Results:** There was a significant interaction of previous night's sleep disturbances and PTSD group. In both groups, sleep disturbances were positively associated with PTSS. Among individuals with probable PTSD ( $n = 42$ ), one more sleep disturbance during the previous night was associated with a 3.75 increase in PTSS the following day ( $p < .001$ ). Among individuals without probable PTSD ( $n = 38$ ), one more sleep disturbance during previous night was associated with a 1.65 increase in PTSS the following day ( $p = .001$ ).

**Conclusions:** These first preliminary findings indicate that sleep disturbances affect next day PTSS and the influence is stronger for individuals with PTSD. Behavioral interventions to ameliorate sleep disturbances may reduce the daily burden of PTSD symptoms for individuals with PTSD.

**Keywords:** post traumatic stress disorder, sleep disturbances, symptom assessment, ecological momentary assessment, military personnel

# Incidence Rates of Posttraumatic Stress Disorder in the US Military (2007-2015): Trends Across Military-Specific Variables

## Authors

Jeffrey Cook, PhD <sup>(1,2)</sup>; Maegan M. Paxton, BS <sup>(1)</sup>; Jennifer Phillips, PhD <sup>(1,2)</sup>; Avni Patel, MPH <sup>(1)</sup>; Lisa French, PsyD <sup>(2)</sup>; David S. Riggs, PhD <sup>(1,2)</sup>; Tracey Koehlmoos, PhD <sup>(1)</sup>

## Author Affiliations

1. Uniformed Services University of the Health Sciences
2. Center for Deployment Psychology

## ABSTRACT

**Background:** Providing effective support and treatment for Posttraumatic Stress Disorder (PTSD) is a priority of the Department of Defense. Thus, it is critical to know how rates of PTSD differ across military-specific variables, including component (Active Duty [AD] or Reserve Component [RC]), branch of service, and rank. However, current estimates of PTSD in the US military vary widely dependent on the sample and study methodology.

**Methods:** This study examined the rates of initial diagnoses of PTSD across these variables in the US Military between 2007 and 2015. Data, consisting of direct care inpatient and outpatient records for AD and RC military were accessed via the Military Health System Data Repository. ICD-9 codes for the

initial diagnosis of PTSD were identified and rates calculated based on counts from the Defense Enrollment Eligibility Reporting System (DEERS).

**Results:** During the study period, rates of initial PTSD diagnosis were higher for AD members, with both AD and RC rates peaking in 2012 before declining through 2015. Rates were consistently highest for the Army, followed by the Marines, lowest for the Coast Guard, and intermediate for Navy and Air Force. Enlisted personnel had higher rates of PTSD than officers, and PTSD rates were inversely related to rank in both the Enlisted and Officer categories.

**Conclusions:** The present results provide a population analysis of the incidence rates of PTSD within the Military Health System. These findings highlight subgroups within the military with higher rates of PTSD diagnosis, which can enable DoD to tailor prevention and treatment interventions for these groups.

The opinions and assertions expressed herein are those of the author(s) and do not necessarily reflect the official policy or position of the Uniformed Services University or the Department of Defense. Additionally, the authors have no conflicts of interests to report.

## Risk and Resilience of Lifetime Stressful Experiences among U.S. Army Soldiers at Risk for Suicide

### Authors

Catherine L. Dempsey<sup>1</sup>, Ph.D., David M. Benedek<sup>1</sup>, MD., Kelly Zuromski<sup>2</sup>, Ph.D., Matthew K. Nock<sup>2</sup>, Ph.D., Tsz Hin Ng<sup>1</sup>, MPH., Charlotte Riggs<sup>1</sup>, MS., Catherine Broshek<sup>1</sup>, BA., Samantha Martinez<sup>1</sup>, BA., and Robert J. Ursano<sup>1</sup>, MD.

### Author Affiliations

1. Center for the Study of Traumatic Stress, Department of Psychiatry, Uniformed Services University of the Health Sciences.
2. Department of Psychology, Harvard University.

### ABSTRACT

**Background/ Objectives.** The purpose of this study is to identify the extent to which the presence of lifetime stressful events (military/deployment-related and family/social-related) are risk factors for suicide among US Army Soldiers as reported by third parties.

**Methods.** The data are from a psychological autopsy study conducted as part of the Army Study to Assess Risk and Resilience in Servicemembers (Army STARRS<sup>\*\*</sup>). Data were gathered from next of kin (NOK) and Army supervisors (SUP) using structured interviews. Propensity-matched controls were matched on known sociodemographic risk factors for suicide and Army history variables. Associations were examined using multivariate logistic regression.

**Results.** The odds of suicide were higher for those Soldiers who experienced lifetime interpersonal violence (sexual assault or rape) compared to propensity-matched controls as reported by SUP (OR =8.6

[95% CI= 1.1, 65.3]) and NOK (OR=4.2 [95% CI = 1.5, 11.5]). NOK reported exposure to the suicide of a close friend or relative NOK (OR=3.0 [95% CI = 1.5, 0.2]) as a risk factor for suicide. Interestingly, NOK and SUP reported service members who experienced a disaster were less likely to die by suicide compared to propensity-matched controls NOK (OR =0.2 [95% CI =0.1, 0.9]) and SUP (OR = 0.2 [95% CI = 0.0, 0.6]). SUP reports also demonstrated the protective effects of saving the life of a Soldier or civilian during deployment (SUP OR = 0.3 [95% CI = 0.1, 0.7]).

**Conclusions.** This study identified interpersonal violence and suicide of a close friend or relative were associated with increased odds of suicide death. Results suggest that exposure to disaster and saving the life of a Soldier or civilian, and the ability to handle stress may be protective of suicide. Our findings may assist both supervisors and family members in identifying those most at risk and inform preventive intervention efforts.

<sup>\*\*</sup> Army STARRS was sponsored by the Department of the Army and funded under cooperative agreement number U01MH087981 with the U.S. Department of Health and Human Services, National Institutes of Health, National Institute of Mental Health (NIH/NIMH). The contents are solely the responsibility of the authors and do not necessarily represent the views of the Department of Health and Human Services, NIMH, Department of the Army, Department of Defense, Uniformed Services University of the Health Sciences, or Center for the Study of Traumatic Stress.

## Inflammatory Cytokines and Cardiovascular Disease are Associated with Neuroimaging Findings after Acute Mild Traumatic Brain Injury

### Authors

Katie A Edwards, PhD, RN<sup>1,2</sup>; Cassandra L Pattinson, PhD<sup>1</sup>; Vivian Guedes, PhD,<sup>1</sup>; Jordan Peyer, BS<sup>1</sup>; Chen Lai, PhD<sup>1</sup>; Christina Devoto, MS<sup>1,2</sup>; Lawrence Latour, PhD<sup>1</sup>; Jessica M Gill, PhD, RN<sup>1</sup>

### Author Affiliations

1. National Institute of Nursing Research, National Institutes of Health, Bethesda, MD.
2. Henry M. Jackson Foundation for the Advancement of Military Medicine, Bethesda, MD.

### ABSTRACT

**Background/Objectives:** Identification of peripheral blood biomarkers related to mild traumatic brain injury (mTBI) that inform neuroimaging may be a safe, cost-effective method to identify individuals with intracranial injury. Previous data shows blood biomarkers discriminate mTBI subjects with/without magnetic resonance imaging (MRI) abnormalities (tau, NFL, GFAP), and with/without computed tomography (CT) findings (GFAP). In this study, we analyzed the relationship between peripheral blood levels of cytokines IL-6, IL-10, TNF-alpha, and VEGF and neuroimaging results following acute mTBI (<48 hours). Increased acute levels of circulating cytokines (IL-6, IL-10, VEGF, TNF-alpha) have been linked to worse clinical outcomes after moderate and severe TBIs. In addition, epidemiological, clinical, and preclinical studies have linked chronic inflammation and immune system activation to cardiovascular disease (CVD) pathogenesis. Associations between neuroimaging, CVD, and levels

of IL-6, IL-10, TNF-alpha, and VEGF in mTBI are unknown.

**Methods:** Participants presented to the emergency department with suspected mTBI (n=341) and underwent blood draw, CT, and MRI. Subjects were categorized into three neuroimaging groups: CT positive (n=94); MRI-only positive (n=112); control (negative neuroimaging, n=135). IL-6, IL-10, TNF-alpha, and VEGF plasma concentrations were measured using single-molecule array (SIMOA). We evaluated whether cytokine levels and/or CVD can predict specific neuroimaging groups.

**Results:** Neuroimaging outcomes were significantly associated to CVD histories of hyperlipidemia ( $p<0.001$ ) and hypertension ( $p=0.005$ ); neuroimaging outcomes were significantly associated to increased concentrations of IL-6 and VEGF ( $p's<0.001$ ). As such, these variables were put into a regression model. Results indicate hyperlipidemia and hypertension were not significant predictors, but IL-6 and VEGF differentiated patients with and without MRI abnormalities, with IL-6 being the strongest predictor ( $p's<0.001$ ).

**Conclusions:** Results suggest IL-6 and VEGF as promising markers of brain injury in patients with acute mTBI, and may differentiate those with CVD history. A multi-biomarker approach, including blood-based biomarkers and patient CVD history, may facilitate neuroimaging and clinical decision making.

## Elevated Tau Relates to Persistent PTSD, Depression and Post-Concussive Symptoms in Military Personnel with History of Traumatic Brain Injury

### Authors

Vivian A Guedes, Ph.D.<sup>1</sup>; Cassandra L Pattison, Ph.D.<sup>1</sup>; Pashtun Shahim, M.D., Ph.D.<sup>1,2</sup>; Patricia Taylor, MBA<sup>2</sup>; Kerri Dunbar, M.A.<sup>2</sup>; Vida Motamedi, BA<sup>1</sup>; Chen Lai, Ph.D.<sup>1</sup>; Christina Devoto, M.S.<sup>1</sup>; Jordan Peyer, BA<sup>1</sup>; Michael J. Roy, Ph.D.<sup>2</sup>; Jessica M Gill, R.N., Ph.D.<sup>1</sup>.

### Author Affiliations

- 1 National Institute of Nursing Research, National Institutes of Health.
- 2 Department of Medicine, Uniformed Services University of the Health Sciences.

### ABSTRACT

**Background/Objectives:** Posttraumatic stress disorder (PTSD), depression, and post-concussive syndrome (PCS) are highly prevalent in military populations and frequently occur comorbidly with traumatic brain injury (TBI), resulting in substantial health risks. Elucidating the biological variances that are associated with TBIs featuring comorbid chronic PTSD and PCS may ultimately enable the identification of the mechanisms that underlie these symptoms in military personnel to improve care and inform novel interventions. Tau, amyloid-beta (A $\beta$ ), and neurofilament light chain (NFL) have been associated with TBI-related pathological mechanisms. However, the relationship between biomarkers and behavioral alterations after TBIs remains to be un-

derstood. This study examined associations among TBI, blood biomarkers and symptoms of PTSD, depression, and PCS.

**Methods:** PTSD, depression and PCS were evaluated using PTSD checklist-civilian version (PCL), patient health questionnaire-9 (PHQ-9), and neurobehavioral symptom inventory (NSI), respectively, in military personnel (n = 109) with or without chronic TBI. Serum concentrations of tau, A $\beta$ 40, A $\beta$ 42, and NFL were measured using an ultra-sensitive assay.

**Results:** The TBI group reported significantly worse PTSD, depression, and post-concussive symptoms than the control group (no TBI). Within the TBI group, controlling for age, sex, time since last injury and anti-anxiety/depression medication use, Tau was positively correlated with PHQ (p = .01) and NSI (p = .008) total scores. Tau was also correlated with the subscales of NSI-somatosensory (p < .001), and PCL-negative mood (p = .047). NFL was correlated with PCL-hyperarousal (p = .017). However, A $\beta$ 40 and A $\beta$ 42A were not significantly correlated with the symptoms measured.

**Conclusions:** Our findings indicate that tau and NFL may play a role in maintenance of psychological symptoms in those who have experienced TBI. Furthermore, there is a critical need for studies of biomarkers longitudinally following TBI.

# Prevalence of Medical Conditions and Healthcare Utilization in Treatment-Seeking, Bereaved Military Widows: A Prospective, Case-Controlled, Longitudinal Study

## Authors

Kathryn Hefner, PhD<sup>1</sup>; Joscelyn Fisher, PhD<sup>1</sup>; Jing Zhou, MS<sup>1</sup>; Alexandra Burris BA<sup>1</sup>; Stephen Cozza, MD<sup>1</sup>

## Author Affiliations

1. Center for the Study of Traumatic Stress, Department of Psychiatry, Uniformed Services University of the Health Sciences

## ABSTRACT

**Background:** Spousal bereavement is associated with physical and mental health consequences, however, little is known about the specific impact of military bereavement. Due to their typically younger age at bereavement and likelihood of being impacted by sudden and violent deaths, military widows may be especially vulnerable to negative health outcomes.

**Method:** Using outpatient medical records from wives of active duty military service members (SMs), we compared prevalences of medical diagnoses and frequency of healthcare visits among treatment-seeking military widow cases (n=1375) and matched (on age, baseline healthcare utilization, SM deployment and rank), time-yoked control military wives (n=1375), from one year prior (Yr-1) to two years following (Yr+1 and Yr+2) SM death. Preva-

lence risk ratios and confidence intervals were used to compare medical condition prevalence rates and outpatient healthcare visits between cases and controls across each time period.

**Results:** Compared to controls, prevalences of ill-defined conditions and mental health conditions (MHCs) increased among cases from Yr-1 to Yr+1 as well as from Yr-1 to Yr+2. Physical and mental healthcare visits generally increased for both cases and controls throughout the study period; however, in Yr+2, widow cases accrued greater numbers of physical healthcare visits as compared to controls. Mental health visits also increased following SM death in Yr+1 and Yr+2 among widows.

**Conclusions:** The increase in ill-defined physical health conditions among widows in the early years following SM death suggests bereavement may be manifested via somatic symptoms, corroborating other recent research indicating military spouses often manifest somatic indicators of stress (Steenkamp et al., 2018). The combination of increases in physical health conditions, MHCs and healthcare utilization among widows highlights the need for improved access to healthcare services that are prepared to adequately treat grief-related conditions.

## The Impact of Child Health on Parental PTSD

### Authors

Elizabeth Hisle-Gorman, MSW, PhD<sup>1</sup>, Apryl Susi, MS<sup>1</sup>, Gregory Gorman, MD, MPH<sup>1,2</sup>

### Author Affiliations

1. Department of Pediatrics, Uniformed Services University, Bethesda, MD 20814
2. Walter Reed National Military Medical Center, Bethesda MD 20814

### ABSTRACT

**Background/Objective:** Research indicates parenthood increased veterans’ likelihood of PTSD care. Results cannot determine if parenthood increases risk or increases motivation for treatment. We sought to examine the impact a child’s mental health condition on parental treatment for PTSD after controlling for deployment related variables.

**Methods:** Utilizing the Military Healthcare System (MHS) database we identified parents injured 2002-16, with a child/children aged 2-16 at time of injury. Parental inclusion in the Ill, Injured, and Wounded Warrior dataset identified service members diagnosed with PTSD and other injuries. Parental records included total pre-injury deployment, battle injuries, and number of injuries. ICD-9 codes identified outpatient child mental healthcare for two years preced-

ing injury. Family data identified number of children, and age of youngest child. Logistic regression determined odds of PTSD diagnosis and treatment by child mental health diagnosis, controlling for family and military factors.

**Results:** There were 234,911 injured parent with 1 or more injury; 1,312 (0.56%) were battle injured, parents were deployed a man of 1.38 years, and had a median of 1.1 injures not including PTSD. There were a median of 1.8 children per family and the median youngest age was 6.2 years. In unadjusted and adjusted analysis parents of a child with a mental health diagnosis were 7% (OR 1.07 [95% CI 1.04-1.10]) more likely to have diagnosed PTSD. Odds of parental PTSD were also increased with each additional child (2%), total pre-injury deployment years (37%), number of injuries (351%), and sustaining a battle injury (196%), odds of PTSD decreased with age of the youngest child (2%).

**Conclusion:** Odds of PTSD diagnosis was increased in injured parents of children with mental health diagnoses. Results may relate to underlying family risk, or suggest family stress exacerbates PTSD risk. Results suggest providing preventative and family focused care may ameliorate some PTSD risk.

---

### Odds of PTSD in Injured Parents

	Unadjusted Odds of PTSD OR [95% CI]	Adjusted Odds of PTSD OR [95% CI]
Child with Mental Health Diagnosis	1.07 [1.04-1.10]	1.07 [1.04-1.10]
Pre-Injury Deployment Length in Years	1.39 [1.38-1.41]	1.37 [1.36-1.39]
Each Additional Child in Family	1.05 [1.04-1.07]	1.02 [1.01-1.03]
Age in Years of Youngest Child in Family	0.97 [0.97-0.98]	0.99 [0.99-0.99]
Each Additional Parental Injury	3.71 [3.62-3.81]	3.51 [3.42-3.60]
Sustaining Injury in Battle	8.02 [7.18-8.96]	1.96 [1.72-2.24]

---

# Whole-Genome Sequencing of 1,688 Veteran Twins to Detect Risk and Measure Rare Variation Associated with Major Depressive Disorder

## Authors

Daniel Hupalo PhD<sup>1</sup>, Christopher Forsberg MS<sup>2</sup>, Jack Goldberg PhD<sup>2,3</sup>, William S. Kremen PhD<sup>4,5</sup>, Michael J. Lyons PhD<sup>6</sup>, Anthony Soltis PhD<sup>1</sup>, Coralie Viollet PhD<sup>1</sup>, Robert J. Ursano MD<sup>7</sup>, Murray B. Stein PhD<sup>4</sup>, Carol E. Franz PhD<sup>4</sup>, Yan V. Sun PhD<sup>8</sup>, Viola Vaccarino PhD<sup>8</sup>, Nicholas L. Smith PhD<sup>2,3</sup>, Clifton L. Dalgard PhD<sup>1</sup>, Matthew D. Wilkerson PhD<sup>1</sup>, Harvey B. Pollard PhD<sup>1</sup>

## Author Affiliations

1. The American Genome Center, Collaborative Health Initiative Research Program, Uniformed Services University, Bethesda, MD
2. Seattle Epidemiologic Research and Information Center, Office of Research and Development, U.S. Department of Veteran Affairs, Seattle, WA
3. Department of Epidemiology, University of Washington, Seattle, WA
4. Department of Psychiatry, University of California, San Diego, San Diego, CA
5. VA San Diego Center of Excellence for Stress and Mental Health, San Diego, CA
6. Department of Psychological & Brain Sciences, Boston University, Boston, MA
7. Department of Psychiatry, Uniformed Services University, Bethesda, MD
8. Department of Epidemiology, Emory University, Atlanta, GA

## ABSTRACT

**Background:** Major Depressive Disorder (MDD) is a complex neuropsychiatric disease that can have large impacts on military readiness. However, new insights from genomic associations allow for prediction of an individual's lifetime genetic risk, and provide for the possibility of proactive therapy. Initial studies have identified 47 regions associated with MDD among individuals of European ancestry.

**Methods:** Here, we report the sequencing of 1,688 whole genomes at 30X mean depth, including 958 monozygotic twins, 626 dizygotic twins, and 73 singletons sequenced in an effort to identify rare genetic variation associated with MDD. Sequencing results were hierarchically clustered using 54 quality metrics to identify and remove bias, resulting in 70 samples being excluded. Veterans were classified as having MDD based on psychiatric interview diagnostic criteria. Using previously published genome-wide association summary statistics from one independent study, we calculated a polygenic risk score (PRS) using 44 loci.

**Results:** No significant difference was found in PRS among Veterans with a binary history of MDD compared to those without a history as determined by logistic regression. When considering the variable describing the lifetime accrual of symptoms of MDD, there was a trend of increasing PRS among those with symptoms compared to without symptoms by increasing symptom count as determined by negative-binomial regression. Additionally, when considering the severity of illness, there was a significant difference in PRS when comparing Veterans with severe depression to those without severe disease as determined by logistic regression.

**Conclusions:** A deeper exploration of the 44 loci is underway. These preliminary results may represent a replication of the MDD associations in a new independent cohort and provide support for undertaking a deeper investigation of MDD rare variation. Precise quantification of genetic risk can potentially provide an early warning allowing for the avoidance of exposures that may cause this complex and chronic disease.

## Antidepressant Activity of the Selective Kappa Opioid Receptor Antagonist JNJ-67953964 in Mice

### Authors

Moriah L. Jacobson, PhD<sup>1</sup>, Hildegard A. Wulf, MA<sup>1</sup>,  
Caroline A. Browne, PhD<sup>1</sup>, Irwin Lucki, PhD<sup>1,2</sup>

### Author Affiliations

1. Department of Pharmacology and Molecular Therapeutics, Uniformed Services University, Bethesda, MD
2. Department of Psychiatry, Uniformed Services University, Bethesda, MD

### ABSTRACT

**Background:** Major depressive disorder (MDD) is a leading cause of disability worldwide that is precipitated and/or exacerbated by stress exposure. Dysregulation of the endogenous opioid system is implicated in the emergence of MDD. Stress increases dynorphin-induced activation of kappa opioid receptor (KOR) signaling, which is known to produce negative affect, dysphoria, and aversion. KOR blockade produces behavioral effects in rodent tests used to screen novel antidepressant compounds that are indicative of potential antidepressant-like activity. JNJ-67953964 (previously LY2456302 and CERC-501) is the only selective KOR antagonist currently in clinical trials for MDD, yet there are no preclinical studies systematically evaluating this compound in animals exposed to stress. In these experiments

we investigated the ability of JNJ-67953964 to ameliorate the behavioral deficits produced by exposure to a chronic mild stress paradigm.

**Method:** Adult male C57BL/6J mice were exposed to four weeks of unpredictable chronic mild stress (UCMS). After three weeks of stress, JNJ-67953964 (10 mg/kg) was administered for 12 days. Behavioral assessments included nest building, sucrose preference test, forced swim test (FST), and the hot plate.

**Result:** Exposure to UCMS reduced nesting, increased immobility on the FST, induced deficits in sucrose preference, and produced thermal hyperalgesia. Exposure to JNJ-67953964 reversed all of these deficits. JNJ-67953964 reversal of stress-induced behavioral deficits persisted for 3 weeks post treatment cessation.

**Conclusions:** Our data show that the KOR antagonist JNJ-67953964 is capable of counteracting a stress-induced phenotype on several behavioral tests. In addition, this is the first study to evaluate the effects of chronic mild stress on nesting, and we demonstrated that nesting is a useful assay to test the effects of stress. In sum, these results encourage further clinical development of JNJ-67953964 as a therapeutic for stress-related disorders.

## Safety and Confidence in Law Enforcement During Terrorist-Related Events: Association with Daily Life Activities

### Authors

Holly B. Herberman Mash, PhD<sup>1</sup>, Carol S. Fullerton, PhD<sup>1</sup>, Joshua C. Morganstein, MD<sup>1</sup>, Brian W. Flynn, EdD<sup>1</sup>, & Robert J. Ursano, MD<sup>1</sup>

### Author Affiliations

1. Center for the Study of Traumatic Stress  
Department of Psychiatry  
Uniformed Services University of the Health Sciences  
Bethesda, Maryland

### ABSTRACT

**Objective:** This study examined the relationship of perceived safety and confidence in law enforcement to changes in daily life activities during the Washington, D.C. sniper attacks, which occurred over a 3-week period in October, 2002.

**Methods:** Participants were 1238 Washington, D.C. area residents assessed using an internet survey that included items related to safety (at work, at home, and in general), confidence in law enforcement, and changes in routine activities of daily life. These changes were defined as either increases or decreases in the following activities: 1)being in large public places (e.g., shopping malls); 2)getting gas; 3) sending one's child(ren) to school and activities; 4) attending large public gatherings (e.g., concerts or sporting events); 5)travelling by public transpor-

tation; 6)travelling by automobile; and 7)attending faith-based activities. Univariate and multivariate logistic regression analyses investigated the relationship of confidence in law enforcement and perceived safety to any changes in routine life activities.

**Results:** A majority of participants (52%, n=640) reported changes in daily life activities, with approximately one-third identifying changes related to being in large places (37%, n=461) and getting gas (36%, n=445). Perceived safety was associated with confidence in law enforcement ( $r = .32, p < .001$ ). After adjusting for demographics, lower feelings of safety and less confidence in law enforcement, both independently and together in separate models, were related to a higher likelihood of changes in daily activities.

**Conclusions:** Terrorist events affect feelings of safety and disrupt routine activities. Safety is both an individual perception and community experience. Focus on strengthening community relationships with law enforcement and community resources that provide support, may enhance perceived safety, decrease feelings of threat, and maintain community members' involvement in daily life activities. Additional research is needed to further articulate how perceived safety and confidence in law enforcement interact to influence community behaviors following terrorist and other disaster events.

## Effect of Prior Hurricane Experience on Preparedness in Florida Department of Health Workers

### Authors

Holly B. Herberman Mash, PhD<sup>1</sup>, Joshua C. Morganstein, MD<sup>1</sup>, Carol S. Fullerton, PhD<sup>1</sup>  
Leming Wang, MS<sup>1</sup>, Alexander G. Liu, MPH<sup>1</sup>, Sumr Farooq, BA<sup>1</sup>, & Robert J. Ursano, MD<sup>1</sup>

### Author Affiliation

1. Center for the Study of Traumatic Stress  
Department of Psychiatry  
Uniformed Services University of the Health Sciences  
Bethesda, Maryland

### ABSTRACT

**Objective:** The extent to which one feels prepared may influence initial responses to disaster exposure, such as fear, anxiety, and helplessness, and affect the time it takes to recover. This study examined preparedness behaviors in Florida Department of Health (FDOH) workers following the 2004 and 2005 hurricane seasons, and the effect of hurricane experience on feelings of preparedness.

**Methods:** Participants were FDOH workers assessed 9 months following four hurricanes and one tropical storm during a seven-week period in August and September of 2004 (n=4676) and one year later, after the 2005 hurricane season (n=3460). During the first assessment, participants' ages ranged from 18-79 years (M=47.81). Most were female (79.8%), married (65.5%), White (71.6%). Nearly half had less

than a BA/BS degree (49.6%). Participants completed a questionnaire that included items examining hurricane preparedness, defined by having the following resources: 1) home emergency preparedness plan; 2) 2+ days of food and water; 3) flashlight; 4) portable radio; 5) spare batteries; 6) emergency phone numbers; and 7) plan to communicate with family/friends. Additional items assessed the extent to which participants felt prepared before and after the 2004 and 2005 hurricanes, indicating whether experience affected preparedness over time.

**Results:** A majority of participants at Time 1 (72%) and Time 2 (77%) reported having at least six preparedness items, demonstrating a high level of hurricane preparedness. Perceived levels of preparedness increased from before the 2004 hurricane season (23%) to after the hurricanes (41%), decreased slightly in the period before the 2005 hurricane season (34%), increasing again following hurricane exposure (47%), suggesting an effect of experience on feelings of preparedness.

**Conclusions:** Findings illustrate the high level of preparedness among FDOH workers, who are trained in disaster response. Disaster-related experiences may further influence feelings of preparedness and have emergency public health implications for enhancing preparedness behaviors that optimize disaster response and improve community safety.

## The 3 Predator Exposure Model as a Robust Animal Model of Traumatic Stress

### Authors

Matthew May, M.S.<sup>1</sup>, Isaac Jeong, B.S.<sup>1</sup>, Rachel Taylor, Ph.D.,<sup>1</sup> Nicole Moore, Ph.D.<sup>1</sup>, Emily Lowery-Gionta, Ph.D.,<sup>1\*</sup> Liana Matson, Ph.D.<sup>1\*</sup>

### Author Affiliations

1. Performance Assessment and Chemical Evaluation Laboratory, Behavioral Biology Branch, Center for Military Psychiatry and Neuroscience, Walter Reed Army Institute of Research, 503 Robert Grant Avenue, Silver Spring, MD 20910

\* Asterisk indicates co-last authorship

### ABSTRACT

**Background:** An estimated 12-20% of soldiers exposed to psychological trauma during the course of a deployment in present-day conflicts go on to develop adverse symptoms of post-traumatic stress disorder (PTSD). The Performance Assessment and Chemical Evaluation (PACE) Laboratory identifies and tests novel compounds for efficacy using a preclinical rat model of traumatic stress to develop treatments for combat-related stress.

**Methods:** The PACE laboratory developed a three predator stress model, combining the live exposure of multiple predators: (snake, cat, and ferret). Exploratory and anxiety-like behaviors are measured in male rats under basal conditions, as well as 24

hours, 48 hours, and 192 hours (7 days) following a single-day sequential exposure to three predator species (snake, ferret and cat) to model a threat to life exposure. Rats spend 5-10 minutes with each of the predators, in a protective enclosure that maximizes sensory exposure to each predator.

**Results:** Predator-exposed (n = 33) and control sham (n = 35) group data was compiled from multiple studies to analyze model effectiveness and to assess the trajectory of traumatic-stress related behaviors. Preliminary analyses indicate that predator exposure results in increased anxiety-like behavior in the elevated plus maze compared to control animals at 24H using the anxiety index formula by Cohen et al. 2008 ( $t = 2.363$ ,  $p = 0.02$ ) but not at 48H or 192H.

**Conclusions:** Elevated plus maze data from multiple studies support the hypothesis that the 3 predator exposure increases anxiety 24 hours after exposure. Additionally, rats showing increased anxiety-like behaviors at baseline also demonstrate elevated anxiety-like behaviors across time compared to animals showing low anxiety-like behaviors at baseline. There are individual differences in anxiety-like response to predator exposure that can be used to model the trajectory of human response to traumatic stress. Additional predictive factors are being evaluated.

## Does Damage to the Medial Frontal Gyrus in Chronic Mild TBI Patients Affect Post Traumatic Stress Symptoms?

### Authors

Dominic E. Nathan, PhD<sup>1,2,3</sup>, Louis M. French, PsyD<sup>1,2,4</sup>, Wei Liu, PhD<sup>4,5</sup>, John Ollinger, PhD<sup>4</sup>, Grant H. Bonavia, MD, PhD<sup>2,4</sup>, Gerard Riedy, PhD, MD<sup>2,4</sup>

### Author Affiliations

1. Center for Neuroscience and Regenerative Medicine, Bethesda, MD, USA
2. Uniformed Services University, Bethesda, MD, USA
3. Henry M. Jackson Foundation, Bethesda, MD, USA
4. National Intrepid Center of Excellence, Bethesda, MD, USA
5. North Tide LLC, Sterling, VA, USA

### ABSTRACT

**Background:** Differentiating between post traumatic stress disorder (PTSD) and chronic mild Traumatic Brain Injury (mTBI) is challenging due to significant symptom overlap, and the absence of objective indicators. The focus of this work is to examine how post-traumatic stress influences task-free resting state brain connectivity of the default mode network in a sample of military chronic mTBI subjects.

**Methods:** Control subjects (N=44, age=36±9.7 years, M=28) without a history of TBI were compared with chronic mTBI subjects with low (N=58, PCL-C total < 30), medium (N= 124, PCL-C total = 31–49), and high (N= 105, PCL-C total ≥60) post-traumatic stress symptoms. Functional MRI data were acquired on a 3T scanner for a six-minute task-free scan. Voxel-wise t-tests were performed on

the default mode network to compare the control group, and each of the mTBI groups, using age as a nuisance regressor.

**Results:** A pattern of disrupted connectivity was observed in the right medial frontal gyrus (Brodmann area 10) across all chronic mTBI subjects. However, connectivity within a network of brain regions comprising the anterior cingulate, temporal lobes and parahippocampal gyrus, varied with intensity of PCL-C scores. Damage to Brodmann area 10 could disrupt executive function and attention processes involved with emotional regulation and memory encoding, rather than a fear related response.

**Conclusions:** The results suggest a potential contextual dependent construct of fear, in which the brain attempts to regulate memories and emotions relating to prior traumatic experiences. Such consolidation of memories or emotions may occur during the absence of explicit external stressful stimuli, and may differ from a fear related response that involves the amygdala. Future work will test amygdala connectivity in the task free context. Results from this study have the potential for the development of in-vivo indicators for future research pertaining to PTSD and chronic mTBI.

**Disclaimer:** Views expressed in this abstract are those of the author(s) and do not necessarily reflect the official policy or position of the Department of the Navy, Army, Department of Defense, nor the U.S. Government.

## Longitudinal Changes in Children's Mental Health and Physical Injury Rates Following Father Death

### Authors

Christin M. Ogle, Ph.D.<sup>1</sup>, Joscelyn E. Fisher, Ph.D.<sup>1</sup>,  
Jing Zhou, M.S.<sup>1</sup>, Ametisse N. Gover-Chamlou,  
B.A.<sup>1</sup>, Stephen J. Cozza, M.D.<sup>1</sup>

### Author Affiliation

1. Center for the Study of Traumatic Stress,  
Department of Psychiatry, Uniformed Services  
University of the Health Sciences

### ABSTRACT

**Objective:** In a prospective, case-controlled study, we examined longitudinal changes in prevalence rates of mental health conditions, physical injuries, and outpatient healthcare visits among children of deceased active-duty male service members compared to their non-bereaved peers.

**Methods:** International Classification of Diseases codes drawn from outpatient medical records were used to determine the rates of mental health conditions, physical injuries, and outpatient healthcare visits among bereaved children ( $n = 1142$ ;  $M$  age = 8.8,  $SD = 5.0$ ) and time-yoked controls ( $n = 1142$ ;  $M$  age = 9.2,  $SD = 4.7$ ) matched on age, baseline physical and mental health utilization, service member parent deployment history, and service member parent rank. Logistic regressions were used to examine prevalence risk ratios one year prior (Yr-1)

and two years following (Yr+1, Yr+2) father death in bereaved compared to non-bereaved children.

**Results:** Comparisons of pre- and post-loss prevalence rates of mental health conditions among bereaved children indicated that rates of depression, anxiety, stress disorder/PTSD, and adjustment disorder were two- to nine-times higher 1- to 2-years following father death. Prevalence of behavioral disturbances and fractures also increased in the second year post-loss. With the exception of depression, no pre- to post-loss increases in prevalence of mental health conditions and physical injuries were observed among non-bereaved youth. Comparisons of longitudinal changes in prevalence of mental health conditions and physical injuries between bereaved and non-bereaved children indicated that only adjustment disorder increased at a significantly higher rate from Yr-1 to Yr+1 in bereaved compared to non-bereaved youth. In addition, the average number of mental healthcare visits was significantly higher for bereaved compared to non-bereaved youth both years post-loss.

**Conclusions:** Father death is associated with increased prevalence of psychiatric conditions, physical injuries, and mental health care utilization 1- to 2-years post-loss in children from treatment-seeking military families.

# Influence Of Attentional Bias on Scanpaths In Posttraumatic Stress Disorder: Assessment of Vigilance-Avoidance Scanning Patterns During Visual Search Paradigm

## Authors

Hyuk Oh, Ph.D.<sup>1</sup>, Matthew Reinhard, Psy.D.<sup>2</sup>, Nathaniel Allen, B.S.<sup>2</sup>, Arghavan Hamedi, M.A.<sup>2</sup>, Kyle Jaquess, Ph.D.<sup>2</sup>, Michelle Costanzo, Ph.D.<sup>2</sup>

## Author Affiliations

- 1 Department of Kinesiology, University of Maryland, College Park, MD 20742, USA.
- 2 War Related Illness and Injury Study Center, Veterans Affairs Medical Center, Washington, DC, 20422, USA.

## ABSTRACT

**Objective:** Eye-tracking tasks allow investigation of sustained attention to threat, which has been implicated in posttraumatic stress disorder (PTSD). Antisaccade performance in the context of affective cues has revealed inhibitory control deficits in a Veteran PTSD population. This study aims to examine visual scanning patterns in Veterans with PTSD by quantifying the spatial arrangements of visual fixations during eye-tracking tasks with and without cognitive-emotional interference.

**Method:** Thirty-six veterans were recruited from the Washington DC Veterans Affairs Medical Center, and classified into two groups (17 PTSD diagnosed vs. 19 controls) based on the Clinician Administered PTSD Scale (CAPS). Participants completed pro-saccade (PS) and antisaccade (AS) eye movement

direction tasks under two conditions; (1) standard condition (STD) that utilized square and circle images, and (2) face condition (FACE) that utilized neutral and negative emotional face images. The nearest neighbor index (NNI) algorithm was employed to measure the spatial pattern of gaze points. Two Gaussian generalized linear models were used to estimate the linear relationships between the NNI and three predictors (Group, Condition, and Direction), and between the NNI and CAPS.

**Results:** The linear model indicated that: (a) there is a significant linear relationship between the NNI and three predictors; (b) eye tracking patterns in veterans with PTSD were less clustered than controls regardless of Condition or Direction; (c) all participants' clusters were tighter during STD and PS; (d) NNI and CAPS were positively correlated.

**Conclusions:** This study revealed that the Veterans with PTSD exhibited face avoidance and more scattered scanning in the antisaccade eye-tracking tasks, and such avoidance symptom of PTSD can be successfully assessed through visual scanning pattern analysis. Thus, our results demonstrated that this task and analytical approach can be used as a novel cost-effective tool for assessing PTSD in a veteran population with the future possibility of informing diagnosis.

## Mild Traumatic Brain Injury with Concurrent PTSD is Associated with Peripheral Tau Concentrations

### Authors

Cassandra L. Pattinson, Ph.D.<sup>1</sup>, Jessica M. Gill, R.N. Ph.D.<sup>1</sup>, Tracey Brickell, Ph.D.<sup>2</sup>, Louis M. French, Ph.D.<sup>3</sup>, Sara M. Lippa, Ph.D.<sup>2</sup>, Rael Lange, Ph.D.<sup>2</sup>

### Author Affiliations

1. National Institutes of Health, National Institute of Nursing Research
2. Defense and Veterans Brain Injury Center, Walter Reed National Military Medical Center
3. National Intrepid Center of Excellence (NICoE), Walter Reed National Military Medical Center

### ABSTRACT

**Objective:** Mild traumatic brain injuries (mTBIs) with comorbid post-traumatic stress disorder (PTSD) is common in military personnel following recent deployments. Given that TBI and PTSD are associated with an increased risk of Alzheimer's disease and related dementias, we aimed to examine if mTBI and comorbid PTSD had cumulative effects on peripheral tau concentrations in two cohorts of military veterans and service members.

**Method:** Cohort 1 included military personnel who were treated at Walter Reed National Military Medical Center following mTBI. Participants were divided into three groups: mTBI-Neg/PTSD-Neg (n=18), mTBI-Pos/PTSD-Neg (n=64), and mTBI-Pos/PTSD-Pos (n=21); Age M=34.1years (SD=10.1). Cohort 2

included personnel who were referred to the Madigan Sleep Disorders Clinic. Participants were split into four groups: mTBI-Neg/PTSD-Neg (n=59), mTBI-Neg/PTSD-Pos (n=11), mTBI-Pos/PTSD-Neg (n=17), and mTBI-Pos/PTSD-Pos (n=25); Age M=33.3 years (SD=7.9). The cohorts were analyzed separately using ANOVA models, with Bonferroni adjustment.

**Results:** For Cohort 1, ANOVA showed significant group differences on tau ( $p < .01$ ). Post-hoc analyses revealed that the mTBI-Pos/PTSD-Pos group had significantly higher tau than the mTBI-Neg/PTSD-Neg group ( $p = .02$ ) and trended towards higher than the mTBI-Pos/PTSD-Neg group ( $p = .06$ ). The mTBI-Pos/PTSD-Pos and mTBI-Neg/PTSD-Pos groups did not significantly differ ( $p > .05$ ). These results were validated with Cohort 2; ANOVA showed group differences on tau ( $p < .05$ ). Post-hoc analyses revealed that the mTBI-Pos/PTSD-Pos group had significantly higher tau than both the mTBI-Neg/PTSD-Neg and mTBI-Pos/PTSD-Neg groups ( $p < .05$ ).

**Conclusion:** These results are the first to identify that a history of mTBI and comorbid PTSD is associated with increased peripheral tau. This provides insight into potential pathways for intervention and monitoring to ameliorate negative outcomes following mTBI in military personnel.

## The Relationship of PTSD to Anxiety and Depression: An Examination of the Moderating Effect of Sleep Quality and Nightmares

### Authors

Maegan M. Paxton, BS<sup>1</sup>, Larissa L. Tate, MPS<sup>1</sup>,  
Patricia T. Spangler, PhD<sup>1,2</sup>, Eva Asabre, MPH<sup>2</sup>,  
and David S. Riggs, PhD<sup>1,3</sup>

### Author Affiliations

1. Uniformed Services University of the Health Sciences
2. Center for the Study of Traumatic Stress
3. Center for Deployment Psychology

### ABSTRACT

**Background and Objectives:** PTSD is a significant health concern for the United States military population. It causes significant distress in a service member's life and is often accompanied by other mental health symptoms. As many as 70% of service members with PTSD experience significant sleep disturbance and upwards of 90% report trauma-related nightmares. These sleep problems are known to exacerbate PTSD symptom development as well as negatively impact patient functioning. Patients with PTSD also often report symptoms of anxiety and depression. The numerous co-occurring conditions and varied presentations make PTSD challenging to treat. Interventions may be improved by better understanding the relationship of these comorbid conditions. Because sleep disturbances are hallmark features of PTSD, anxiety, and depression, we examined the role of sleep disturbances in the relations among these disorders.

**Methods:** Participants were 51 treatment-resistant (i.e., not fully responsive to SSRI and/or SNRI treat-

ment) active duty service members and veterans of OIF, OEF, and OND. Participants were recruited for participation in a randomized control trial examining the effectiveness of Riluzole as an augmentation medication for PTSD. Riluzole is a glutamate modulator that may improve outcomes compared to treatment with SSRIs or SNRIs alone.

**Results:** Notably, sleep disturbance was found to be a moderating factor in the relationship of PTSD to anxiety and depression. Also, nightmares moderated the relationship of PTSD to depression. Additional analyses indicate that the relationship between PTSD and anxiety and depression is strengthened as symptoms of sleep disturbance and nightmares increase.

**Conclusions:** These results demonstrate the potentially important role sleep plays in understanding the comorbidity of PTSD and anxiety or depression. By targeting sleep disturbances in treatment for PTSD, patients may see greater reductions in symptoms of comorbid anxiety and depression than treatment for PTSD alone, perhaps leading to increased functioning and quality of life.

*The opinions and assertions expressed herein are those of the author(s) and do not necessarily reflect the official policy or position of the Uniformed Services University or the Department of Defense. Additionally, the authors have no conflicts of interests to report.*

## A Population Study of Incidence Rates of Posttraumatic Stress Disorder in the US Military (2007-2015): Analysis of Age, Sex, and Marital Status

### Authors

Maegan M. Paxton, BS<sup>1</sup>; Jeffrey Cook, PhD<sup>1,2</sup>;  
Jennifer Phillips, PhD<sup>1,2</sup>; Avni Patel, MPH<sup>1</sup>; Lisa French, PsyD<sup>2</sup>; David S. Riggs, PhD<sup>1,2</sup>; Tracey Koe-hlmoos, PhD<sup>1</sup>

### Author Affiliations

1. Uniformed Services University of the Health Sciences
2. Center for Deployment Psychology

### ABSTRACT

**Background:** The rate of Posttraumatic Stress Disorder (PTSD) in the military is a frequently-studied and often-contested area in trauma research. Prevalence rates vary widely from 1.4% to 41.3%. Further, systematic reviews suggest a rate between 8% and 15%. Additionally, the literature differs in terms of rates across demographic factors. The wide range of rates reported across the literature may be due to factors such as the size and composition of the sample as well as the method of assessment.

**Methods:** By accessing the Military Health System Data Repository (MDR), this study aimed to determine the overall rate and demographic trends of initial diagnosis of PTSD in the US Military population between 2007 and 2015. PTSD diagnosis was determined by ICD-9 code in direct care inpatient

and outpatient records for Active Duty and Reserve Component military. Rates were calculated based on counts from the Defense Enrollment Eligibility Reporting System (DEERS).

**Results:** During the study period, the rate of PTSD diagnosis peaked in 2012 (9.69 per 1000-person years) before declining through 2015. Notably, rates of PTSD diagnosis in females steadily increased after 2012, while male rates decreased. Rates were consistently highest among those over age 40 and lowest in those under 20. Single Service members had the lowest PTSD diagnosis rates, followed by married, and then divorced personnel.

**Conclusions:** These findings provide the most comprehensive view of rates of PTSD diagnoses in the military to date. Additionally, they highlight important differences across basic demographic characteristics within the US military. Therefore, these findings may help to identify service members at greater risk of developing PTSD as well as to inform targeted interventions.

*The opinions and assertions expressed herein are those of the author(s) and do not necessarily reflect the official policy or position of the Uniformed Services University or the Department of Defense. Additionally, the authors have no conflicts of interests to report.*

## Sex-related Differences of Intravenous Ketamine Infusion on Stress Hormone and Fear Memory in Rats

### Authors

Kennett D Radford, PhD<sup>1</sup>, Michael Zhang, BS<sup>2,3</sup>,  
Rina Y Berman, BS<sup>2</sup>, T John Wu, PhD<sup>4</sup>, and Kwang  
H Choi, PhD<sup>1,2,3</sup>

### Author Affiliations

1. Daniel K. Inouye Graduate School of Nursing, Uniformed Services University, Bethesda, MD, 20814
2. Department of Psychiatry, Uniformed Services University, Bethesda, MD, 20814
3. Center for the Study of Traumatic Stress, Uniformed Services University, Bethesda, MD, 20814
4. Department of Obstetrics and Gynecology, Uniformed Services University, Bethesda, MD, 20814

### ABSTRACT

**Background:** The U.S. Department of Defense has recently opened combat roles to women that were previously restricted to men. As a result of this policy change, military doctors and nurses can anticipate an increased frequency of combat-related injuries in female service members in future military conflicts. Ketamine, an NMDA receptor antagonist, is a preferred battlefield analgesic due to its hemodynamic stability and a lack of respiratory suppression in wounded service members. However, ketamine administration in the peri-trauma period can produce dissociation and hallucination which may strengthen the traumatic memory formation. We previously reported that subanesthetic intrave-

nous (IV) ketamine infusion (0–10 mg/kg) dose-dependently increased stress hormone corticosterone (CORT) levels and fear memory in male rats.

**Method:** We investigated the effects of IV ketamine infusion on those measures in intact female rats. Adult female Sprague-Dawley rats received a 2-hour IV ketamine infusion (0 or 10 mg/kg) immediately after the fear conditioning (3 times of auditory tone and mild footshock pairing). Spontaneous locomotor activity was monitored during the infusion and plasma CORT levels were measured using the ELISA method. Fear memory retrieval, fear extinction, and fear recall were tested between 2 and 4 days after the fear conditioning/ketamine infusion.

**Results:** The IV ketamine infusion reduced locomotor activity and elevated plasma CORT levels in female rats. The elevated CORT levels following ketamine infusion were greater in female rats than in male rats. The ketamine infusion following fear conditioning enhanced fear memory and this effect appeared to be greater in female rats than in male rats.

**Conclusions:** The study is in progress to confirm the current findings and to further investigate potential contribution of sex hormones (estrogen and progesterone) to the effects of IV ketamine infusion on fear-related disorders.

**Funding:** TriService Nursing Research Program and Center for the Study of Traumatic Stress

# Gene Expression Differences in PTSD Are Uniquely Related to the Intrusion Symptom Cluster: A Transcriptome-Wide Analysis in Military Service Members

## Authors

Heather L. Rusch, MS<sup>1</sup>, Jeffrey Robinson, PhD<sup>1</sup>, Sijung Yun, PhD<sup>2</sup>, Nicole D. Osier, PhD<sup>2</sup>, Christina Martin, BS<sup>1</sup>, Chris R. Brewin, PhD<sup>3</sup>, & Jessica M. Gill, PhD<sup>1</sup>

## Author Affiliations:

1. National Institute of Nursing Research, National Institutes of Health, Bethesda, MD 20892, United States
2. Yotta Biomed, LLC, Bethesda, MD 20817, United States
3. University College London, London WC1E 6BT, United Kingdom

## Corresponding Author:

Heather L. Rusch  
3 Center Drive, Building 3, Room 5E/26; Bethesda, MD 20892  
Tel: 301-496-5198; E-mail: heather.rusch@nih.gov

## ABSTRACT

**Background:** Posttraumatic stress disorder (PTSD) is associated with wide-spread immune dysregulation; however, little is known about the gene expression differences attributed to each PTSD symptom cluster: intrusions (flashbacks), avoidance, negative alterations in cognition and mood, and arousal symptoms. This is an important consideration when identifying diagnostic and treatment response markers in highly comorbid populations with overlapping symptoms.

**Methods:** We utilized a transcriptome-wide analysis of differential gene expression in peripheral blood

by comparing military service members: (1) with vs. without PTSD, (2) with high vs. low PTSD cluster symptom severity, and (3) with improved vs. not improved PTSD symptoms following 4 to 8 weeks of sleep-focused treatment. Data were analyzed at a  $\pm 2.0$ -fold change magnitude,  $FDR \leq 0.05$  with subsequent pathway analysis.

**Results:** In participants with PTSD ( $n=39$ ), 89 differentially expressed genes were identified. In participants with high intrusion symptoms ( $n=22$ ), 1040 differentially expressed genes were identified. No differentially expressed genes were identified for the remaining PTSD symptom clusters, which overlap with other conditions. Ten discrete genes (*C5orf24*, *RBAK*, *CREBZF*, *CD69*, *PMAIP1*, *AGL*, *ZNF644*, *ANKRD13C*, *ESCO1*, and *ZCCHC10*) were upregulated in participants with high intrusion symptoms at baseline and downregulated in participants with improved symptoms following treatment. Pathway analysis identified upregulated immune response systems and metabolic networks with a NF- $\kappa$ B hub linked to high intrusion symptoms, which were downregulated with symptom reduction.

**Conclusion:** Between group (PTSD vs. control) gene expression differences were almost entirely attributed to the intrusion symptom cluster (98%). We previously reported that brain volume alterations were unrelated to overall PTSD severity, but inversely related to intrusion symptom scores. Taken together, our findings provide a novel reference dataset, which can inform the development of precise PTSD diagnostic biomarkers and therapeutic targets.

## Nightmare Deconstruction and Reprocessing: Proof of Concept of a Novel Treatment for PTSD-Related Nightmares and Insomnia

### Authors

Patricia Spangler, PhD<sup>1</sup>, Alvi Azad, DO<sup>1,2</sup>, James West, MD<sup>1,2</sup>, David Benedek, MD<sup>1,2</sup>, Catherine Dempsey, PhD<sup>1</sup>, Keke Schuler, PhD<sup>1</sup>, Megan Paxton, BA<sup>3</sup>, Eva Asabre, MPH<sup>1</sup>, Ashley Phares, BA<sup>1</sup>, Jessica Canales, BA<sup>1</sup>

### Author Affiliations

1. Center for the Study of Traumatic Stress, Uniformed Services University, Bethesda, MD, 20814
2. Department of Psychiatry, Uniformed Services University, Bethesda, MD, 20814
3. Department of Medical and Clinical Psychology, Uniformed Services University, Bethesda, MD, 20814

### ABSTRACT

**Background/Objectives:** Nightmares and insomnia are signature symptoms of posttraumatic stress disorder (PTSD) and are commonly refractory following evidence-based treatment (EBT). EBTs such as Prolonged Exposure and Cognitive Processing Therapy utilize fear memory extinction and reconsolidation but do not target sleep symptoms. Given that nightmares may play a role in trauma memory maintenance, developing a psychotherapy that activates trauma memory through exposure to nightmare images may facilitate trauma memory reconsolidation. This proof-of-concept study is investigating Nightmare Deconstruction and Reprocessing (NDR), a three-stage treatment that integrates exposure to nightmare images, meaning-making and reprocessing, and nightmare reconstruction and

rehearsal. Our aims are to test NDR's plausibility as a treatment for PTSD-related nightmares and to test the feasibility of methods intended for use in a future large-scale randomized controlled trial. **Methods:** Treatment plausibility will be tested by analyzing pre-to-post-treatment changes in nightmare and insomnia. Methods being tested include collection of blood samples within a prescribed circadian window (0800-1200) and participant compliance with daily download of physiologic data from a wristband device and completion of psychometric measures. Participants are up to 30 combat veterans with trauma-related nightmares and insomnia who are being recruited at Walter Reed National Military Medical Center. Primary outcomes are nightmare severity (Disturbing Dreams and Nightmare Severity Index) and insomnia severity (Pittsburgh Sleep Quality Index). The Empatica E4 wristband enables collection of actigraphic data to measure sleep disturbance and heart rate variability and electrodermal activity to monitor in-session stress during nightmare exposure. Finally, brain-derived neurotrophic factor and inflammatory markers (IL-2, IL-6, and CRP) will be assayed from blood samples taken at baseline, first exposure to nightmare images, and post-treatment.

**Results:** Results for the current study are pending.

**Discussion:** Results on NDR's plausibility as a treatment and feasibility of study methods will provide evidence for testing NDR in a large-scale randomized clinical trial.

## Region- and Time-Dependent Gene Regulation in the Amygdala and Anterior Cingulate Cortex of a PTSD-Like Mouse Model

### Authors

Mikiei Tanaka Ph.D.<sup>1</sup>, Hongyun Li Ph.D.<sup>1</sup>, Xijun Zhang Ph.D.<sup>2</sup>, Jatinder Singh Ph.D.<sup>2</sup>, Clifton Dalgard Ph.D.<sup>1,2</sup>, Matthew Wilkerson Ph.D.<sup>†‡</sup>, and Yumin Zhang Ph.D.<sup>1,2</sup>

### Author Affiliations

1. Department of Anatomy, Physiology and Genetics
2. Collaborative Health Initiative Research Program (CHIRP), Uniformed Services University of Health Sciences, 4301 Jones Bridge Rd, Bethesda, MD 20814, USA

### ABSTRACT

**Background/Objectives:** PTSD is developed by exposure to a threatening and/or a horrifying event and characterized by the presence of re-experiencing, hyperarousal, and avoidance for a prolonged period of time. However, the stage-dependent alteration of gene regulation in emotional regulatory regions is not well-elucidated.

**Method:** PTSD-like mouse model was constructed by electric foot shock (1.5 mA for 2 sec ×8 times) followed by situational reminders. Fear memory sustainability was assessed by measuring avoidance latency and freezing behavior in the same context up to 5 weeks post stress (PS). Anxiety-like behavior was assessed by open field test, elevated plus maze

test, and light/dark box test at 2 and 4 weeks PS. Acoustic startle response test was performed at 2 and 4 weeks PS. The amygdala (AMY) and anterior cingulate cortex (ACC) were dissected from the mice at 2 and 5 weeks PS and employed RNA-sequencing analysis.

**Results:** Contextual fear memory was sustained up to 5 weeks PS. Anxiety-like behavior was shown after 2 weeks and continued at 4 weeks PS, while exaggerated startle response emerged after 4 weeks PS. RNA sequencing revealed more than 1,000 differentially expressed genes were identified at 2 weeks PS in both regions. The number of the regulated genes remained constant in AMY at 5 weeks PS, whereas those in ACC were plummeted. Although synaptic remodeling and endocrine system were the most enriched signaling pathways in both ACC and AMY, the individual gene expression profile was regulated differentially in a region- and time-dependent manner. In addition, several genes associated with PTSD involved in Hypothalamic-Pituitary-Adrenal axis were regulated.

**Conclusion:** In our PTSD-like mouse model, global gene expression profile showed synaptic remodeling and endocrine systems are most critical for disease development. The dynamic gene regulation could provide a clue to the molecular mechanism for disease development.

## The Effects of Comorbid TBI and PTSD Symptoms among U.S. Active Duty Service Members on an Objective Cognitive Performance Measure

### Authors

Larissa L. Tate, MPS<sup>1</sup>, Maegan M. Paxton, BS<sup>1</sup>, Louis M. French, PsyD<sup>1,2,3</sup>, Wendy A. Law, PhD<sup>1,2</sup>, Katherine W. Sullivan, SLP<sup>2,3</sup>, & David S. Riggs, PhD<sup>1</sup>

### Author Affiliations

1. Uniformed Services University of the Health Sciences,
2. Walter Reed National Military Medical Center, National Intrepid Center of Excellence
3. Defense and Veterans Brain Injury Center

### ABSTRACT

**Background and Objectives:** More than 2.8 million US military personnel have deployed to Iraq and Afghanistan since September 11, 2001. Both traumatic brain injury (TBI) and posttraumatic stress disorder (PTSD) have been labeled “signature wounds” of these conflicts, and often co-occur as brain injuries are frequently sustained during traumatic events. Nearly 380,000 service members were diagnosed with a TBI between 2000 and 2017, and upwards of 440,000 service members returned home from these conflicts with PTSD. Both of these debilitating conditions can result in considerable cognitive concerns for a significant number of service members. Despite their frequent comorbidity and overlapping symptoms, there is a dearth of knowledge regarding the combined effects of TBI and PTSD on cognitive performance.

**Methods:** The present study examined the potentially synergistic effects of comorbid TBI and PTSD symptoms (PTS) on cognitive functioning using data from 211 active duty service members previ-

ously collected as part of a larger clinical database at the National Intrepid Center of Excellence’s Brain Fitness Center (BFC) at Walter Reed National Military Medical Center. In accordance with the BFC’s normal standard of care procedures, service members completed a variety of self-reports, including the PTSD Checklist (PCL-C), as well as an objective cognitive assessment, the Automated Neuropsychological Assessment Metrics version 4 Traumatic Brain Injury (ANAM-4 TBI).

**Results:** Notably, service members with TBI did not significantly differ from those with PTS-only on cognitive performance; however, individuals with both TBI and PTS performed worse than those with either condition alone.

**Conclusions:** Our findings suggest there may be something unique about the combination of TBI and PTSD and their subsequent influence on cognitive performance. Results illustrate the complexity of the relationship between PTSD and TBI and highlight the need for further research in this area.

*The views expressed in this abstract are those of the author(s) and do not reflect the official policy of the Uniformed Services University, Department of Army, Navy, Air Force, Department of Defense, or U.S. Government. The identification of specific products, scientific instrumentation, or organizations is considered an integral part of the scientific endeavor and does not constitute endorsement or implied endorsement on the part of the author, DoD, or any component agency. Additionally, the authors have no conflicts of interest to report.*

## Fear Memory Extinction is Blocked In Rats Following Acute Glial Cell Inhibition: Implications for Targeting Neuroinflammation in Traumatic Stress Exposure

### Authors

Rachel Taylor, PhD<sup>1</sup>, Matthew May, MS<sup>1</sup>, Nicole Moore, PhD<sup>1</sup>, Liana Matson, PhD<sup>1</sup>, Emily Lowery-Gionta, PhD<sup>1</sup>

### Author Affiliations

1. Walter Reed Army Institute of Research, Silver Spring, MD

### ABSTRACT

**Background:** Emerging evidence suggests a role for both central and peripheral inflammation in post-traumatic stress disorder (PTSD). Moreover, inflammation has been reported to disrupt fear memory extinction as observed in preclinical fear conditioning paradigms. Ibudilast, a PDE4 inhibitor and putative glial cell inhibitor, has been shown to reduce neuroinflammation as well as inflammation-induced behavioral impairments. Here, we characterized Ibudilast for its efficacy in facilitating fear extinction in rats to better understand the role of inflammation in fear learning.

**Methods:** Adult male rats underwent a single fear conditioning session in which delivery of light and tone cues were associated with the delivery of a mild footshock. The following week, Ibudilast (3 or 10 mg/kg, i.p.) or its vehicle was administered

once-daily 1-h prior to a 10-min extinction test for a total of four days, in which freezing behavior was measured. Additionally, behavioral performance in acoustic startle response and elevated plus maze tests was measured under drug-free conditions at various timepoints before and after fear conditioning occurred. Terminal blood and various brain regions were collected for later evaluation of inflammatory cytokine levels.

**Results:** Ibudilast administration to shock-exposed rats resulted in a sustained level of freezing behavior, suggesting fear extinction was delayed or blocked. Freezing behavior was also observed in the sham-exposed, Ibudilast-treated control group, however the levels did not reach those of the shock-exposed groups and only appeared on the 2<sup>nd</sup>-4<sup>th</sup> test day. Moreover, freezing behavior decreased over test days for the sham-exposed, Ibudilast-treated group, but did not reach sham-exposed, vehicle-treated levels.

**Conclusions:** Although experiments are ongoing, the present results suggests that while a mild Ibudilast-specific effect on motor ability may be involved, Ibudilast modulates aspects of fear learning relative to extinction. These results may have clinically-relevant implications for current pharmacological targets of traumatic stress exposure.

## Sex Differences in Cued Fear Responses and Parvalbumin Cell Density in the Hippocampus Following Repetitive Concussive Brain Injuries in C57BL/6J Mice

### Authors

Laura B. Tucker, M.S.<sup>1,2</sup>, Brian S. Winston<sup>2</sup>, Jiong Liu, M.D.<sup>2</sup>, Alexander G. Velosky, B.A.<sup>2</sup>, Amanda H. Fu, M.D.<sup>1,2</sup>, Antigone A. Grillakis, B.S.<sup>3</sup>, and Joseph T. McCabe, Ph.D.<sup>1,2,3</sup>

### Author Affiliations

1. Pre-Clinical Studies Core, Center for Neuroscience and Regenerative Medicine, Uniformed Services University of the Health Sciences, Bethesda, MD
2. Department of Anatomy, Physiology & Genetics, Uniformed Services University of the Health Sciences, Bethesda, MD
3. Graduate Program in Neuroscience, Uniformed Services University of the Health Sciences, Bethesda, MD

### ABSTRACT

**Background/Objectives:** Repetitive concussive brain injuries (rCBI) may be a risk factor for depression and anxiety disorders, including post-traumatic stress disorder. Animal models of brain injury afford the opportunity for controlled study of the effects of injury on functional outcomes. Despite the participation of female athletes in contact sports, few pre-clinical studies include both sexes in TBI research.

**Methods:** Male and cycling female C57BL/6J mice sustained rCBI (3x) at 24-hr intervals and were tested in a context and cued fear conditioning para-

digm to assess amygdala and hippocampal function. Mice were also tested in the elevated zero maze and tail suspension test for anxiety- and depressive-like symptoms, respectively.

**Results:** All mice with rCBI showed less freezing behavior than sham control mice during the fear conditioning context test. Injured male, but not female, mice also froze less in response to the auditory cue (tone). Injured mice spent more time in the open quadrants of the elevated zero maze, suggesting decreased anxiety, but there were no differences between injured mice and sham-controls in depressive-like activity on the tail suspension test. Pathologically, injured mice showed increased astrogliosis in the injured cortex and white matter tracts (optic tracts and corpus callosum). There were no changes in the number of parvalbumin-positive interneurons in the cortex or amygdala, but injured male mice had fewer parvalbumin-positive neurons in the hippocampus.

**Conclusions:** Parvalbumin-reactive interneurons of the hippocampus have been previously demonstrated to be involved in hippocampal-cortical interactions required for memory consolidation, and it is possible memory changes in the fear-conditioning paradigm following rCBI are the result of more subtle imbalances in excitation and inhibition both within the amygdala and hippocampus, and between more widespread brain regions that are injured following a diffuse brain injury.

## Influence of Number of Awakenings during the Previous 1-3 Nights on Post Traumatic Stress Symptoms Among Individuals with PTSD

### Authors

Jing Wang, Ph.D.,<sup>1</sup> Robert J. Ursano, M.D.,<sup>1</sup> Quinn M. Biggs, Ph.D., M.P.H.,<sup>1</sup> David S. Krantz, Ph.D.,<sup>1</sup> Gary H. Wynn, M.D.,<sup>1</sup> Russell B. Carr, M.D.,<sup>1,2</sup> Nicole M. Dacuyan, B.S.,<sup>1</sup> and Carol S. Fullerton, Ph.D.<sup>1</sup>

### Author Affiliations

1. Center for the Study of Traumatic Stress, Department of Psychiatry, Uniformed Services University of the Health Sciences, 4301 Jones Bridge Road, Bethesda, MD 20814
2. Walter Reed National Military Medical Center, 8901 Wisconsin Avenue, Bethesda, MD 20889

### ABSTRACT

**Background:** Little is known how sleep affects Post Traumatic Stress Symptoms (PTSS) from day to day. This study examined the relationship between the number of awakenings from sleep during the previous 1-3 nights and daily level of PTSS among individuals with Post Traumatic Stress Disorder (PTSD).

**Methods:** Subjects ( $N = 80$ ) were assessed for probable PTSD at enrollment and individuals with probable PTSD ( $n = 42$ ) were included in the analyses. Using an ecological momentary assessment methodology, PTSS were assessed four times daily and number of awakenings was assessed once daily by self-report for 15 days. An overall awakenings variable (i.e., person mean) and a previous night awakenings variable (i.e., the difference between the previous night and the individual's mean) were created for number of awakenings from the sleep

during the previous night. Similar variables were created for number of awakening from the sleep during the previous two or three nights. Linear mixed models were applied, with controlling for demographic characteristics.

**Results:** Number of awakenings during the previous night was not associated with PTSS ( $b = 0.86, p = .064$ ). However, number of awakenings during the previous two ( $b = 1.76, p = .035$ ) or three nights ( $b = 1.56, p = .029$ ) was positively associated with PTSS. The overall variables of number of awakenings were not associated with PTSS in all three models.

**Conclusions:** These preliminary findings suggest that number of awakenings did not have an immediate influence on the following day's PTSS. However, two or three days of disruptive sleep awakenings may have an accumulative effect with more awakenings associated with higher PTSS. Assessing and monitoring changes in sleep and implementing interventions to improve sleep quality may reduce PTSS for individuals with PTSD.

**Keywords:** post traumatic stress disorder, sleep, symptom assessment, ecological momentary assessment

**Disclaimer:** The opinions and assertions expressed herein are those of the author(s) and do not necessarily reflect the official policy or position of the Uniformed Services University, Department of Army/Navy/Air Force, Department of Defense, or U.S. Government.

## Sex-Related Differences of Mitochondrial DNA Copy Number in Active Military Service Members with PTSD

### Authors

Lei Zhang MD<sup>1</sup>, Xian-Zhang Hu MD & PhD<sup>1</sup>, Xiaoxia Li BS<sup>1</sup>, David M. Benedek MD<sup>1</sup>, Carol S. Fullerton PhD<sup>1</sup>, Gary Wynn MD<sup>1</sup>, Biomarker Study Group, Robert J. Ursano MD<sup>1</sup>

### Author Affiliations

1. Center for the Study of Traumatic Stress, Department of Psychiatry, USUHS, Bethesda, MD 20814, USA

### ABSTRACT

**Background:** The lifetime prevalence of PTSD has been observed to be about 10–12% in women and 5–6% in men suggesting a sex-related difference in PTSD prevalence. PTSD is associated with mitochondrial DNA copy number (mtDNAcn), which is an emerging systemic index of mitochondrial biogenesis and function. Together, these results lead to a hypothesis that there is a sex-related difference between mtDNAcn levels, which may allow mtDNAcn levels to be used as a potential biomarker to identify PTSD or monitor PTSD symptoms.

**Methods:** mtDNAcn was assessed with a TaqMan assay using white blood cells from service members with (n=137; male: 123, female: 14) or without (n=673; male: 606, female: 67) PTSD, who served during combat operations in the Afghanistan and/ or Iraq wars between 2008 and 2016. The limitation of this study is a relatively small female sample size, and so a comprehensive with a larger female

sample size study might be needed in the future. The Institutional Review Board at the Uniformed Service University of Human Sciences approved all study procedures and all participants were given written informed consent. PTSD symptoms were assessed using the PTSD Checklist (PCL), a 17-item, DSM-based, self-report questionnaire with well-established validity and reliability. PTSD diagnosis was determined based on endorsement of DSM-IV criteria and a PCL total score  $\geq 44$ . The severity of PTSD symptoms was determined using the PCL total score. Subjects with depression, substance abuse, or any medication use were excluded. Demographic data, such as age, gender, and race, were collected.

**Results:** mtDNAcn was significantly higher in female service members with PTSD compared with either male/female non-PTSD controls or male service members with PTSD ( $p < 0.05$ ). However, there was no significant difference of mtDNAcn between male service members with PTSD and male/female non-PTSD controls.

**Discussion:** This study provides the first evidence showing significantly increased mtDNAcn levels in female service members with PTSD, but not in male service members with PTSD. This suggests that altered mtDNAcn in PTSD may be sex-dependent, at least relating to active service members, and reflect impaired energy metabolism, which might represent a novel aspect of PTSD pathophysiology and serve as a biomarker for PTSD in females.

## Continuing Education Credit

The Amygdala, Stress, and PTSD Conference, in conjunction with the Center for Deployment Psychology at the Uniformed Services University is pleased to offer continuing education credits for Physicians, Psychologists, Social Workers, and Nurs-

es. The credits are available through live in person attendance.

*Continuing Education made possible in part by the generous support of the Center for Deployment Psychology.*

---

## American Psychiatric Association (APA)

### Accreditation

This activity has been planned and implemented in accordance with the accreditation requirements and policies of the Accreditation Council for Continuing Medical Education through the joint providership of the American Psychiatric Association (APA) and Society of Uniformed Services Psychiatrists. The APA is accredited by the ACCME to provide continuing medical education for physicians.

### Designation

The APA designates this live activity for a maximum of 5 *AMA PRA Category 1 Credit*.™ Physicians should claim only the credit commensurate with the extent of their participation in the activity.

### Physician CME Certificates

At the conclusion of the conference, physician participants will be provided with an opportunity to claim hours of participation and receive an official CME certificate by completing the online CME conference evaluation.

1. Go to <http://apapsy.ch/amygdala>
2. Click ACCESS ACTIVITY and log in with your APA username and password or create a new account.
3. Enter the Group ID: SUSP191 on the Purchase page and click SUBMIT to

proceed to the evaluation and certificate. Select “AMA PRA Category 1 Physician” as your certificate type.

You must claim your CME credit within 60 days of the conference. Certificates will not be issued after June 16, 2019. For customer support, please email [apalms@psych.org](mailto:apalms@psych.org).

### Disclosure Information

Derrick A. Hamaoka, MD – Nothing to disclose

Eric Meyer II, MD – Nothing to disclose

Katy Pokorny – Nothing to disclose

Dennis S. Charney, MD – *(See next page)*

Anne Germain, PhD – CEO and Equity – Rehat, LLC, Consultant – Jazz Pharmaceuticals, Inc.

Jessica M. Gill, PhD, RN. – Nothing to disclose

James L. Griffith, MD – Nothing to disclose

Irwin Lucki, PhD – Consultant – Alkermes

COL Wendi Waits, MD – Capital Equity – Reimbursify

COL Christopher H. Warner, MD – Nothing to disclose

Kwang Choi, PhD – Nothing to disclose

Kelly L. Cozza, MD – Nothing to disclose

Iona M. Horotan-Enescu, MD – Nothing to disclose

Holly H. Mash, PhD – Nothing to disclose

David Mears, PhD – Nothing to disclose

James A. Naifeh, PhD – Nothing to disclose  
 Amy E. Steward, TSgt, USAF – Nothing to disclose  
 Robert J. Ursano, MD – Nothing to disclose  
 Gary H. Wynn, MD – Nothing to disclose

Dennis S. Charney, MD  
 Ongoing Research Support P01HL131478 (PI: Fayad) 03/17/17-02/28/22 National Heart, Lung, and Blood Institute/NIH/DHHS Stress and atherosclerotic plaque macrophages – a systems biology approach Role: Co-Investigator N/A (PI: Bevilacqua) 07/15/18-07/15/20 Brain and Behavior Research Foundation Intranasal NPY

Role: Key Personnel, Co-Mentor 23629 (PI: Balchandani) 01/15/16-01/14/19 Brain and Behavior Research Foundation Development of 7T MRI protocol to establish imaging biomarkers for depression Role: Key Personnel, Mentor Completed Research Support (Past 3 Years K23MH099223 (PI: Iacoviello) 08/19/13 – 07/31/18 (Transferred) NIMH A Novel Cognitive Training Intervention for Depression Role: Key Personnel, Mentor 2013098 (PI: Murrugh) 07/01/13-06/30/17 (Ended Early) Doris Duke Charitable Foundation Ketamine Plus Lithium as a Novel Pharmacotherapeutic Strategy for Treatment-Resistant Depression

Role: Key Personnel, Mentor K23MH094707 (PI: Murrugh) 07/15/11-05/31/16 NIH Functional MRI Studies of Emotion in Depression and Rapid Anti-depressant Response

Named co-inventor US Patent No. 9,592,207 - Intranasal Administration of Ketamine to Treat Depression (Issued March 14, 2017); US Patent No. 9,539,220 - Methods for Treating Suicidal Ideation (Issued January 10, 2017) ; US Patent No. 8,785,500 - Intranasal Administration of Ketamine to Treat Depression (Issued July 22, 2014) ; US Patent No. 10,123,737 - Systems and Methods for Treating a Psychiatric Disorder (Issued November 13, 2018) ; US CON Patent Appl No. 16/189,059 – and related foreign patent applications - Systems and Methods for Treating a Psychiatric Disorder US Provisional Patent Appl No. 62/649,469 – Brain Plasticity Following Cognitive-Emotional Training ; US CON Patent Appl No. 14/974,576 and related foreign patent

applications - Method for Treating Post Traumatic Stress Disorder (PTSD) ; US Serial No. 14/889,746 and a related foreign patent application - Treatment of Mood and Anxiety Disorders ; US CON Patent Appl Nos. 15/379,013 and 15/417,689 - Intranasal Administration of Ketamine to Treat Depression.

### Meeting Objectives

At the end of this educational activity, the learner will be able to:

- Understand how psychological stress alters brain function.
- Understand psychobiological mechanisms of human resilience to stress.
- You can train to be more resilient.
- Implications for your own life.
- Describe the importance of sleep as a biological force multiplier in psychological health, resilience, and readiness.
- Explain the interaction between sleep and neural circuits underlying threat- and goal-oriented behaviors.
- Detail the nature and impact of sleep-focused interventions on psychological health.
- Describe the proteins that relate to PTSD as well as traumatic brain injury (TBI) symptoms.
- Describe gene-activity profiles that relate to chronic PTSD symptoms.
- Determine the long-term biological changes that relate to TBI and PTSD, which may have morbidity risks associated with them.
- Conceptualize hope as a practice, i.e. “something you do” rather than “something you feel.”
- Demonstrate rapid assessment of patients’ competencies for mobilizing hope in stressful circumstances.
- Formulate and design interventions to mobilize hope when a patient is demoralized.
- Show how empirical social psychology and social neuroscience can expand the scope and potency of psychotherapeutic interventions to mobilize hope and counter demoralization.
- The state of research on developing rapid acting antidepressants for treatment-resistant depression.
- The potential benefits of using ketamine in the treatment of mental health disorders.
- The potential negatives of using ketamine in the treatment of mental health disorders.

## Brooke Army Medical Center (BAMC)

Free Continuing Education (CE) Credits are available for this workshop. This event is sponsored through the Department of Behavioral Health, Brooke Army Medical Center. Department of Behavioral Health, Brooke Army Medical Center is approved by the American Psychological Association to sponsor continuing education for psychologists.

Department of Behavioral Health, Brooke Army Medical Center maintains responsibility for this program and its content. 3.75 credit hours will be given for full attendance at the workshop. No partial credit will be provided. A sign-in /sign-out sheet as well as an evaluation form must be completed in order to receive CE credits and a certificate.

---

## PESI, Inc.

**Program Content provided by Center for Deployment Psychology Continuing Education hours provided by PESI, Inc.**

**Program Title:** 14th Annual Amygdala, Stress, and PTSD Conference: Risk, Resilience, and Recovery

**Date:** April 16, 2019

**Location:** Uniformed Services University, Bethesda, MD

**PESI Program Code:** 72883BET (Please reference this code when contacting PESI regarding this program)

**Duration of Instructional Content:** 225 minutes

*Must attend entire seminar to receive credit. No partial credit will be awarded.*

**ADDICTION COUNSELORS:** This activity consists of 3.5 clock hours of continuing education instruction. Credit requirements and approvals vary per state board regulations. Please save the course outline, the certificate of completion you receive from the activity and contact your state board or organization to determine specific filing requirements.

**COUNSELORS:** This intermediate activity consists of 3.75 clock hours of continuing education instruc-

tion. Credit requirements and approvals vary per state board regulations. Please save the course outline, the certificate of completion you receive from the activity and contact your state board or organization to determine specific filing requirements.

**MARRIAGE & FAMILY THERAPISTS:** This activity consists of 225 minutes of continuing education instruction. Credit requirements and approvals vary per state board regulations. You should save this course outline, the certificate of completion you receive from the activity and contact your state board or organization to determine specific filing requirements.

**NURSES/NURSE PRACTITIONERS/CLINICAL NURSE SPECIALISTS:** This activity consists of 3.75 clock hours of continuing education instruction. Credit requirements and approvals vary per state board regulations. Please save the course outline, the certificate of completion you receive from this activity and contact your state board or organization to determine specific filing requirements.

**SOCIAL WORKERS:** PESI, Inc., #1062, is approved to offer social work continuing education by the Association of Social Work Boards (ASWB) Approved Continuing Education (ACE) program. Organizations, not individual courses, are approved as ACE providers. State and provincial regulatory boards have the final authority to determine whether

an individual course may be accepted for continuing education credit. PESI, Inc. maintains responsibility for the course. ACE provider approval period: January 27, 2017 - January 27, 2020. Social Workers completing this course receive 3.75 Clinical Practice continuing education credits for completing this course. Course Level: Intermediate. A certificate of attendance will be awarded at the end of the program to social workers who complete the program evaluation. Full attendance is required; no partial credits will be offered for partial attendance.

**PENNSYLVANIA SOCIAL WORKERS, MARRIAGE & FAMILY THERAPISTS AND PROFESSIONAL COUNSELORS:** This intermediate activity consists of 3.75 clock hours of continuing education instruction. Credit requirements and approvals vary per state board regulations. Please contact your licensing board to determine if they accept programs or providers approved by other national or state licensing boards. A certificate of attendance will be awarded at the end of the program to participants who are in full attendance and who complete the program evaluation. Full attendance is required; no partial credits will be offered for partial attendance.

**OTHER PROFESSIONS:** This activity qualifies for 225 minutes of instructional content as required by many national, state and local licensing boards and professional organizations. Save your course outline

and certificate of completion, and contact your own board or organization for specific requirements.

Credits listed are for full attendance only. Certificates of Completion are emailed 3-4 weeks after the event and after attendance has been verified to those who attend the full seminar. Please see “live seminar schedule” for full attendance start and end times. NOTE: Boards do not allow credit for breaks or lunch. If your profession is not listed, please contact your licensing board to determine your continuing education requirements and check for reciprocal approval. Materials that are included in this course may include interventions and modalities that are beyond the authorized practice of mental health professionals. As a licensed professional, you are responsible for reviewing the scope of practice, including activities that are defined in law as beyond the boundaries of practice in accordance with and in compliance with your professions standards. PESI, Inc. offers continuing education programs and products under the brand names PESI, PESI Healthcare, PESI Rehab and Psychotherapy Networker.

\*\*CE hours for this activity are being provided by PESI, Inc. Have an inquiry on continuing education credit that is not listed here? Please contact Bridget Schaub at PESI, Inc. before the event. You may reach her via email at [bschaub@pesi.com](mailto:bschaub@pesi.com).

## Amygdala Conference Training Titles and Learning Objectives

1. **Dennis S. Charney, MD — “Resilience: The Science of Mastering Life’s Greatest Challenges”**
  - a. Learning Objectives
    - i. Articulate how psychological stress alters brain function
    - ii. Appraise the psychobiological mechanisms of human resilience to stress
    - iii. Point out training methods that increase resilience
    - iv. Communicate the implications of improved resilience in their own lives
2. **Anne Germain, PhD — “Wake up to Sleep! A Translational Perspective of the Role of Sleep in Readiness and Resilience”**
  - a. Learning Objectives
    - i. Deduce the importance of sleep as a biological force multiplier in psychological health, resilience, and readiness
    - ii. Assess the interaction between sleep and neural circuits underlying threat- and goal-oriented behaviors
    - iii. Communicate the nature and impact of sleep-focused interventions on psychological health
3. **Jessica M. Gill, PhD, RN — “Gene-activity and Proteins that Relate to Chronic PTSD Symptoms”**
  - a. Learning Objectives
    - i. Point out the proteins that relate to PTSD as well as traumatic brain injury (TBI) symptoms
    - ii. Diagnose gene-activity profiles that relate to chronic PTSD symptoms
    - iii. Determine the long-term biological changes that relate to TBI and PTSD, which may have morbidity risks associated with them
4. **James L. Griffith, MD— “Mobilizing Hope in the Face of Despair: Applying Social Neuroscience Research in Brief Clinical Encounters”**
  - a. Learning Objectives
    - i. Appraise how hope is a practice, i.e. “something you do” rather than “something you feel”

- ii. Consider methods on assessing patients' competencies for mobilizing hope in stressful circumstances
- iii. Provide an overview of interventions to mobilize hope when a patient is demoralized
- iv. Analyze how empirical social psychology and social neuroscience can expand the scope and potency of psychotherapeutic interventions to mobilize hope and counter demoralization.

**5. Irwin Lucki, PhD — “Preclinical Development of Ketamine and the Metabolite 2R,6R-Hydroxynorketamine For Depression and Other Disorders”**

a. Learning Objectives

- i. Characterize the state of research on developing rapid-acting antidepressants for treatment-resistant depression.
- ii. Critique the potential benefits of using ketamine in the treatment of mental health disorders.
- iii. Assess the potential negatives of using ketamine in the treatment of mental health disorders.



Notes

---



---



---



---



---



---



---



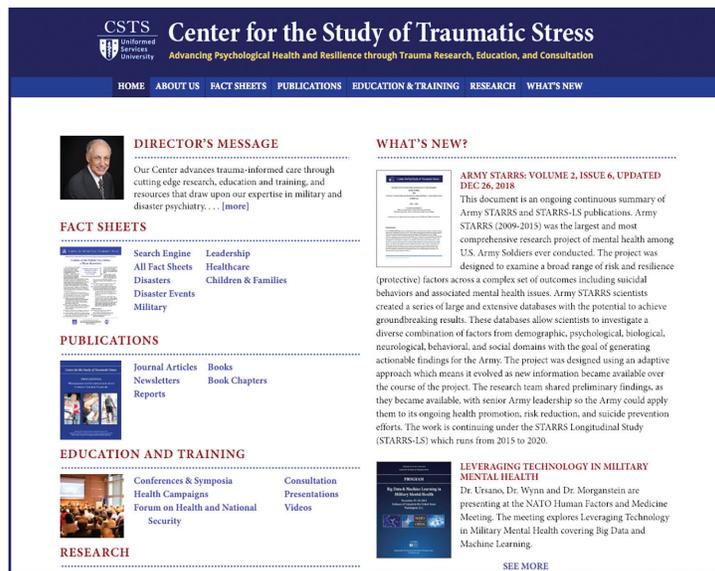
---



---



---



Please visit us at [www.CSTSONline.org](http://www.CSTSONline.org)

Follow us on Twitter and like us on Facebook and receive notices when we add content.



[www.twitter.com/CSTS\\_USU](https://www.twitter.com/CSTS_USU) | [www.facebook.com/USU.CSTS](https://www.facebook.com/USU.CSTS)





Center for the Study of Traumatic Stress  
Department of Psychiatry  
Uniformed Services University  
4301 Jones Bridge Road, Bethesda, MD 20814-4799  
[www.CSTSONline.org](http://www.CSTSONline.org)