



<u>Long-term Impact of Military-relevant Brain</u>
<u>Injury Consortium/Chronic Effects of Neurotrauma Consortium:</u>

Lessons Learned on Collaborative Research

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Bottom Line Up Front: BLUF

- Mature, federal mTBI research consortium since 2013 with robust research infrastructure.
- 10 research studies have been completed with 197 publications and 12 additional grants funded.
- Currently, supporting 6 active research studies
 - 1,600+ (target 3,000) participant <u>Prospective Longitudinal Study</u>
 - Prospective <u>Biomarker Discovery</u> and <u>Novel Neuroimaging</u> studies
 - 2.2+ million-participant <u>Retrospective Database</u>, <u>Phenotypes</u> and <u>Health Economics big data studies using extant federal datasets
 </u>
- Key deliverable: A range of Knowledge Translation products have been developed and additional ones are underway for consumers, family, clinicians and researchers.
- How did we build an efficient, productive research program?

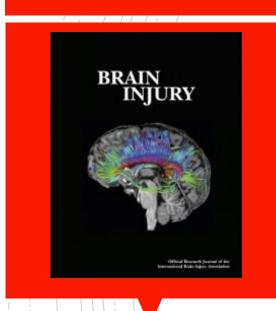
High-Level Overview

LIMBIC-CENC

www.limbic-cenc.org

- The <u>Chronic Effects</u> of <u>Neurotrauma Consortium</u> (CENC 2013-19) identified a range of differences between SM's and Veterans with and w/o mild TBI in the 1,700+ participant Longitudinal Study.
- CENC's unique combined dataset from the electronic medical, benefits, pharmacy and administrative records from VA and DoD of 2+ million unique Veterans revealed linkages between TBI and dementia, Parkinson's disease, chronic pain and suicide.
- The Long-term Impact of Military-relevant Brain Injury Consortium (LIMBIC 2019-24) commenced October 2019, is growing the Longitudinal Study to >3,000 participants, and continue to analyze large electronic dataset.
- LIMBIC-CENC's research teams have identified an association between TBI and dementia, biofluid markers of repetitive TBI and confirmation that pain and mental health disorders worsen dysfunction after mTBI.

Summaries of Protocols and Results



- CENC Special Issue: Brain Injury 2016; 30(12): 1397– 1514
 - Methodologies for 10 studies
 - Assessment Protocols and Tools

https://www.tandfonline.com/toc/ibij20/30/12?nav=tocList

- CENC Special Issue Brain Injury 2018; 32(9): 1149-1294
 - Findings for 10 studies through Spring 2018
 - Integration of findings across 7 clinical studies

https://www.tandfonline.com/toc/ibij20/32/10?nav=tocList

 LIMBIC Special Issue of Brain Injury scheduled for Fall 2021 on

"Practical Approaches to Assessing and Mitigating the Risk of Cognitive Decline after Concussion: Findings from the Long-term Impact of Military-relevant Brain Injury Consortium (LIMBIC)"



Long-Term Impact of Military-Relevant Brain Injury Consortium

Research Cores

Coordinating Center

Y Virginia Commonwealth University, Richmond, VA

Neuroimaging Core

 VA Salt Lake City Health Care System/University of Utah, Salt Lake City, UT

Biomarkers Core

Uniformed Services University of the Health Sciences/ National Institutes of Health, Bethesda, MD

Data and Biostatistics Core

- Hunter Holmes McGuire VA, Richmond, VA
- Virginia Commonwealth University, Richmond, VA
- VA Salt Lake City Health Care System/ University of Utah, Salt Lake City, UT
- University of Hawaii, Department of Speech, Honolulu, HI

Prospective Longitudinal Study

Enrollment Sites

- Hunter Holmes McGuire VA, Richmond, VA
- Michael E. DeBakey VA Medical Center, Houston, TX
- James A. Haley Veterans Hospital, Tampa Florida
- South Texas Veterans Hospital, San Antonio, TX Fort Belvoir Community Hospital, Alexandria, VA
- VA Portland Health Care System, Portland, OR
- Minneapolis VA Health Care System, Minneapolis, MN
- VA Boston Healthcare System, Boston, MA
- WG Hefner VA Medical Center, Salisbury, NC
- Eisenhower Army Medical Center, Fort Gordon, GA
- VA San Diego Health System/University of California-San Diego/Camp Pendleton, San Diego, CA

Recruiting Sites

- MacDill Air Force Base, FL
- Joint Base Lewis-McChord, WA
- ■Fort Steward, Columbia, SC
- ■Naval Amphibious Base, Coronado, CA



Additional Research Studies

Retrospective Database Study

∇ San Francisco VA Medical Center/University of California- San Francisco, San Francisco, CA

Phenotypes Study

VA Salt Lake City Health Care System/ University of Utah, Salt Lake City, UT

Health Economics Study

VA Palo Alto Health Care Systems/Stanford University Biomarkers Discovery Study

VUSUHS/NIH, Bethesda, MD

∇ North Florida/South Georgia Veterans Health System, Gainesville, FL

Novel Neuroimaging Study

VA Salt Lake City Health Care System/ University of Utah, Salt Lake City, UT

LIMBIC-CENC Legacy Study Sites

Epidemiology of mTBI and Neurosensory Outcomes

- ▲ San Francisco VA Medical Center, San Francisco, CA Tau Modification Study
- ▲ Roskamp Institute, Sarasota, FL/Barrows Institute, Phoenix AZ Otolith Dysfunction
- ▲ Mountain Home VA Medical Center, Mountain Home, TN Novel White Matter Imaging to Improve Diagnosis of Mild TBI
- ▲ VA San Diego Healthcare System, San Diego, CA ADAPT/EVOLVE
- ▲ University of Washington, Seattle, WA

Structural and Functional Neurobiology of Veterans Exposed to Primary Blast Forces

▲ WG Hefner VA Medical Center, Salisbury, NC DTI Phantom Study

▲ Baylor College of Medicine, Houston, TX

Clinical and Neuroimaging Correlates of Neurodegeneration in Military mTBI

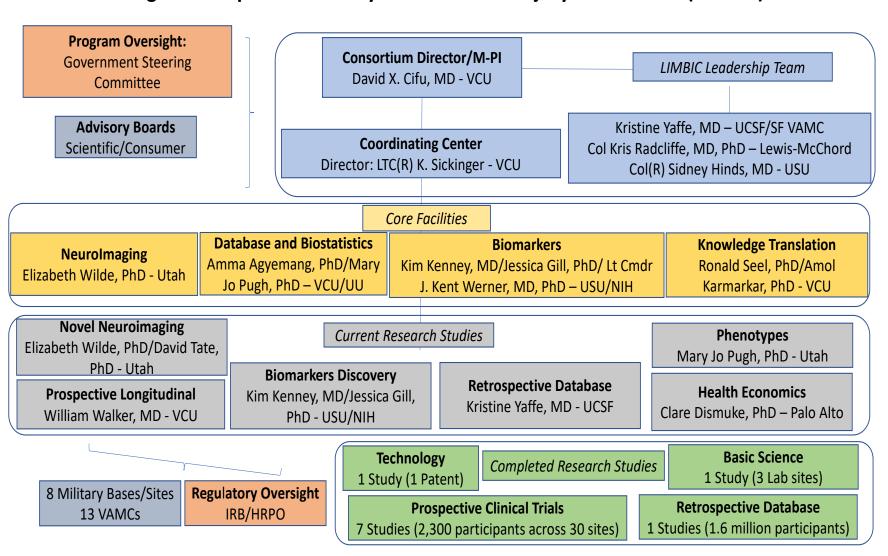
▲ Minneapolis VA Health Care System

Visual Sensory Impairments

▲ Iowa City VA Health Center, Iowa City, IA

Long-Term Impact of Military-relevant Brain Injury Consortium (LIMBIC) 2013-2024

Long-term Impact of Military-relevant Brain Injury Consortium (LIMBIC)



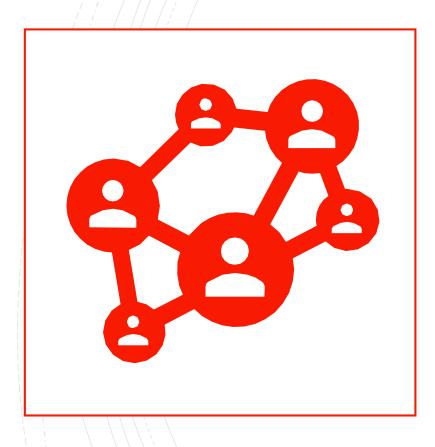


Knowledge Translation Center Building research capacity and output from the research infrastructure Dementia Risk and Risk Reduction Tools Patient Education Videos Clinical Pearls Postcards LIMBIC 2020-21 Gold Standard Research Assessments Searchable Scientific Reference Catalog Secured funding to expand into LIMBIC. Expand PLS to 17 recruitment and 11-testing sites centers to Publish Initiate Publish initial Prospective additional prospective research Longitudinal Study. 2014-15 research Biomarker and articles. articles. Neuroimaging Initiate one single Initiate three Studies. Establish mega-Present research center and one 3database from 9 single center findings at center prospective Expand extant datasets prospective **CENC 2013** national studies. Database studies meetings. studies to 3 Implement Round Develop research Present unique areas of Initiate 8-site l of research RFPs. Obtain and administrative approach and focus: Technology additional grant infrastructure. project. findings at **Epidemiology** funding Implement 3-site national - Phenotypes. **Basic Science** meetings. - Health Initiate one 4-Enroll 2.000+ Sponsor National project. **Economics** center prospective participants Meeting. Obtain study. additional grant Add Knowledge funding. Translation Center.

LIMBIC-CENC: Progressively building upon the experience and infrastructure of an established consortium to study long-term effects

Add Health Economist.

Keys to Building a Collaborative Program



- The PI had established research and professional relationships (and a positive reputation) with all the key leaders of the team.
- The initial team of researchers had been connected academically for 5+ years before the RFP.
- The leadership team began having weekly 1-hour calls >12 months before the RFP was released to vet ideas, establish organization matrices and build workable collaborative programs.
- Individuals who didn't attend calls, couldn't compromise, or weren't able to adapt to the collaborative model were not included.

Keys to Building a Collaborative Program

- The initial aspirational goals of the researchers were high; however, expediency dictated that the "goals" of the funding agencies needed to be met.
- Research projects and new expertise were introduced progressively over the initial 5-year cycle
- All funding and organizational decisions were done transparently and in a democratic fashion.
- Excellence and success were praised, while failures were ameliorated and optimized.

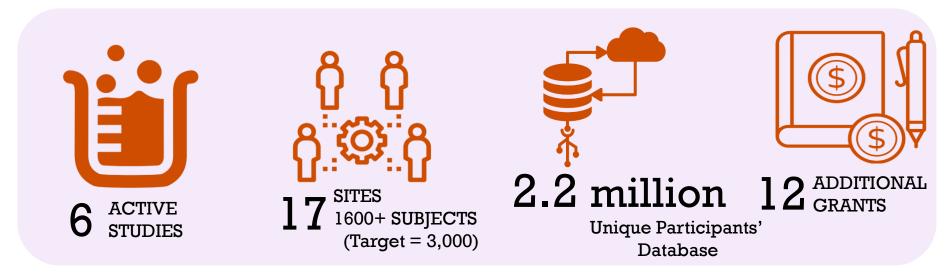


Keys to Building a Collaborative Program

- Everyone/No one is a superstar.
- Young/New investigators are given seats at the table, mentored and given increasing responsibility.
- Change and innovation are welcomed at all levels.
- All work is interdisciplinary.
- "Persons-served" are hired, used to guide ideas, vet products, and provide critical insights.
- KISS Stick to the core missions, ignore shiny objects and keep it real.

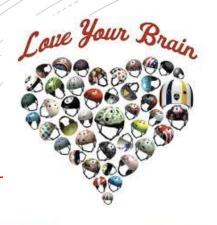
LIMBIC-CENC: Getting It Done Together



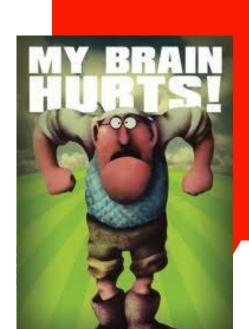


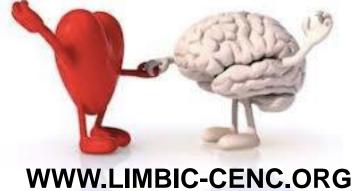
LIMBIC 2013 - 2024: 50+ researchers; 19 Universities, 14 VAMCs and 8 DoD facilities



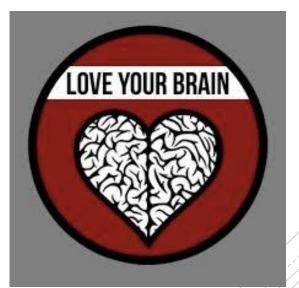












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