

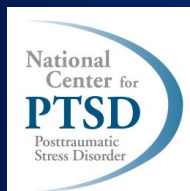
# Associations of PTSD, TBI, and Neurocognitive Performance Over Time

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# Longitudinal Associations among Posttraumatic Stress Disorder Symptoms, Traumatic Brain Injury, and Neurocognitive Functioning in Army Soldiers Deployed to the Iraq War

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Funding: VA Cooperative Studies Program #566

Prior data collection: DoD and VA CSR & D

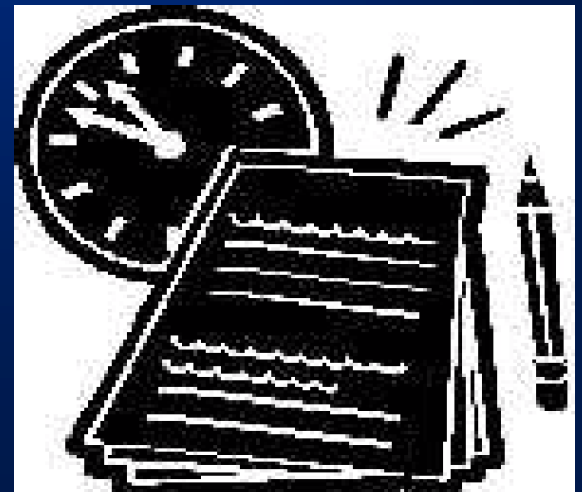
# Poll Question #1

- Do you work in any of the following clinical or clinical research setting(s) (select all that apply)?
  - Rehab (including polytrauma)
  - PTSD specialty care
  - Other mental health
  - Neurology/neurosurgery
  - Neuropsychology

# Background

# PTSD-related Neuropsychological Deficits: Meta-analytic findings

- ✓ Speed of information processing
- ✓ Attention/working memory
- ✓ Verbal learning  
Verbal memory  
Executive functioning



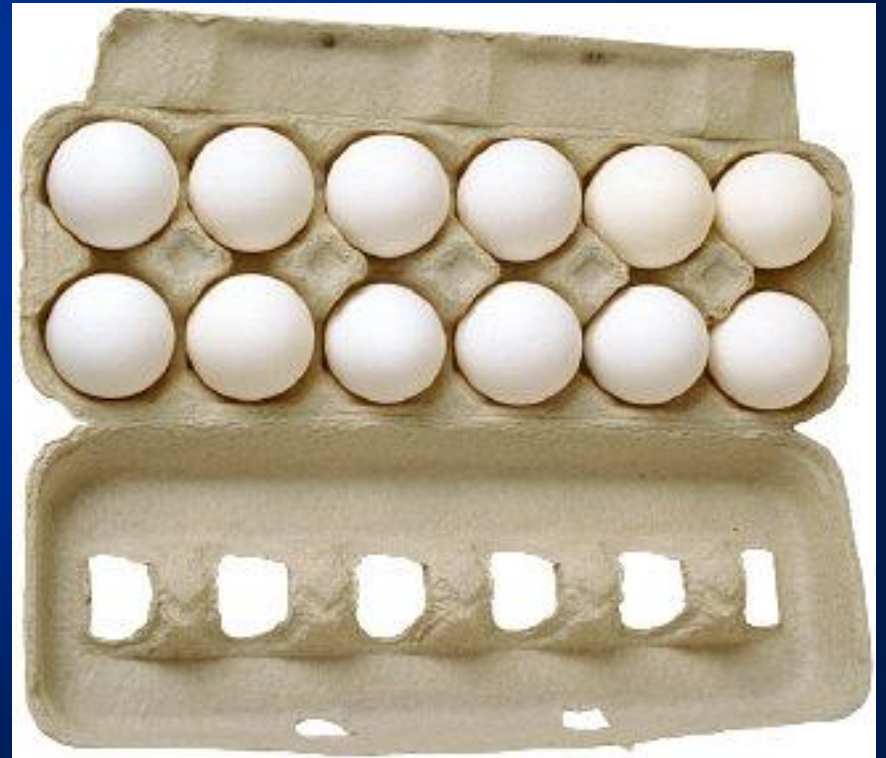
# PTSD-related Neuropsychological Deficits: Growing Evidence

Visual learning/memory

Inhibition/gating

(e.g., Aase et al., 2017; Psychiatry Res; DeGutis et al., 2015: J Int Neuropsych Soc)

# Most research cross-sectional



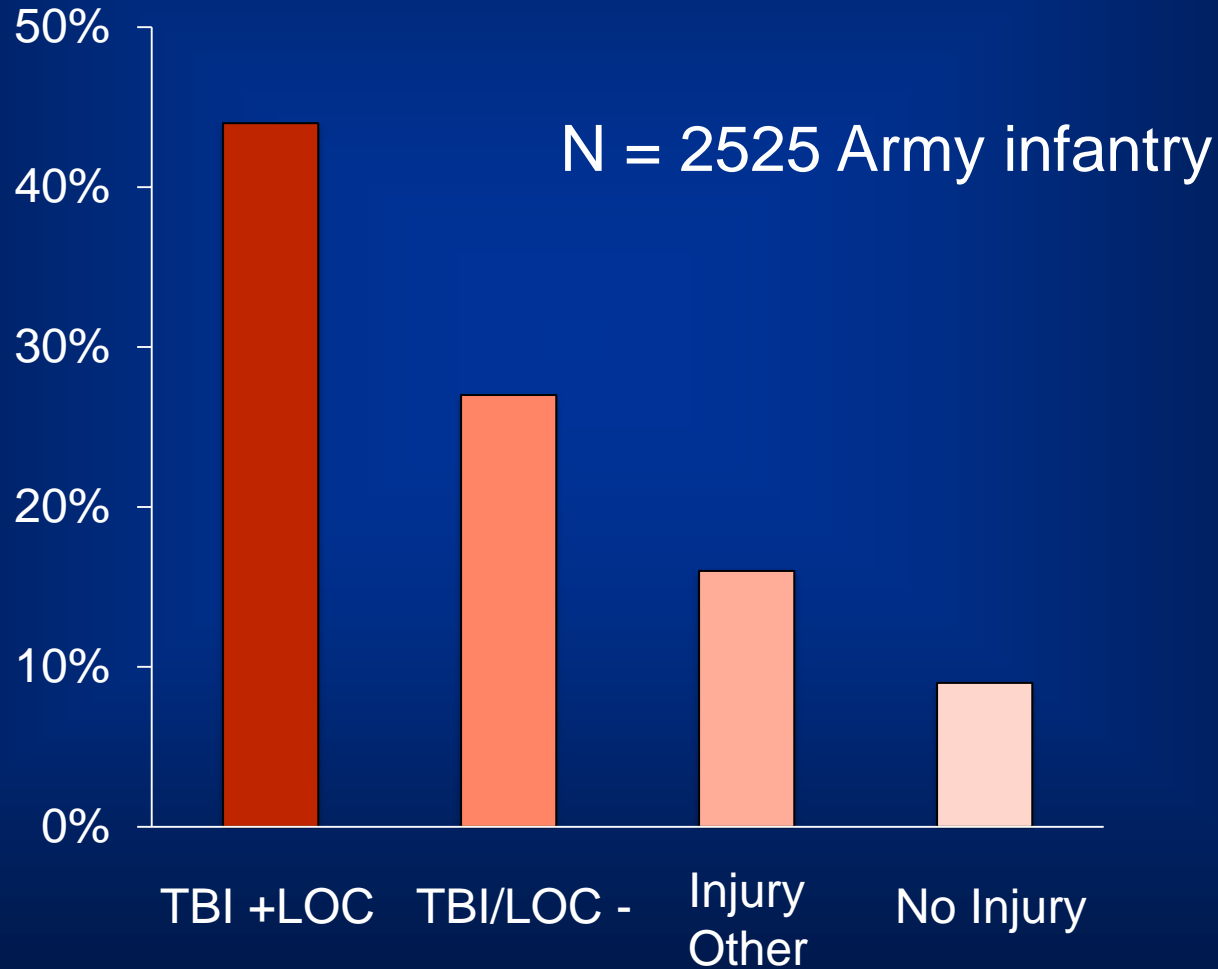
## Objective 1:

To examine longitudinal associations between neuropsychological performance and PTSD symptoms



# Hoge et al. (2008; *N Eng J Med*)

PTSD



# Mild TBI Increases Risk of PTSD

- Bryant et al. (2010; Am J Psych)

n = 1084 civilians with traumatic injuries

At 12 mos., mild TBI patients ~2x more likely to develop new:

	Adj OR	CI
PTSD	1.92	1.08, 3.40
Panic	2.10	1.03, 4.14
Social Phobia	2.07	1.03, 4.16
Agoraphobia	1.94	1.13, 3.39

Functional impairment related to psychiatric status.

# 6-year Follow-up

- O'Donnell et al. (2016; J Clin Psychiatry)

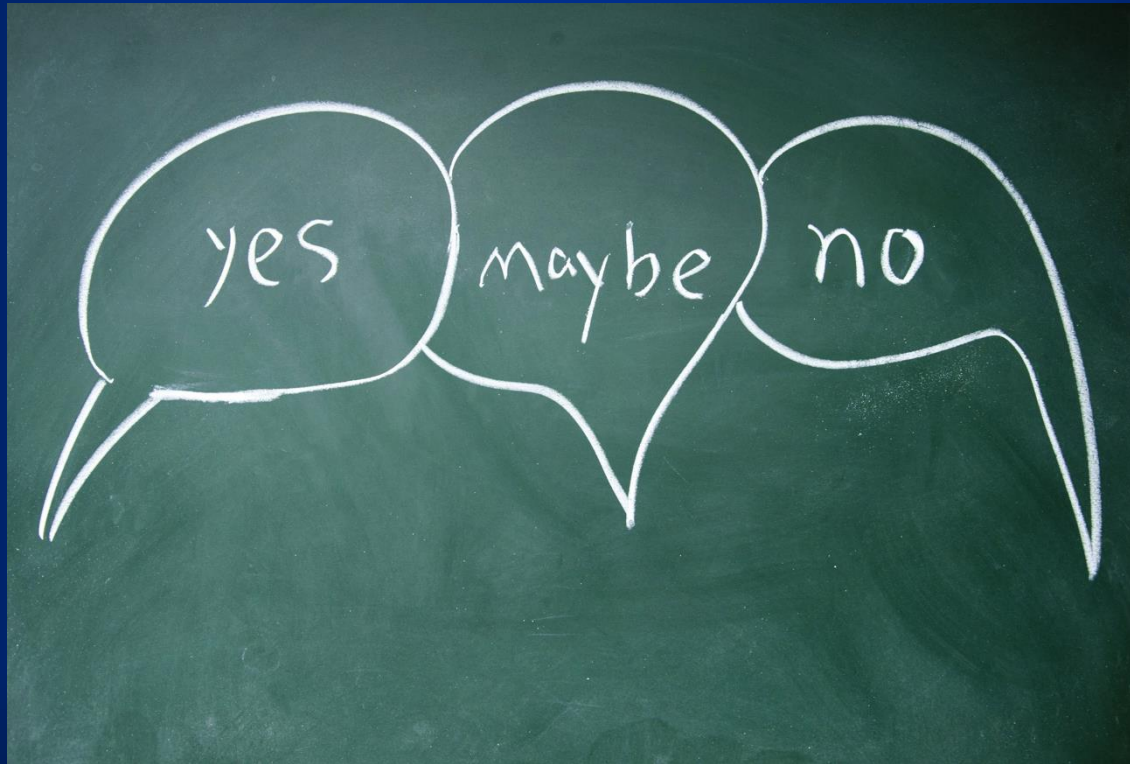
At 72 mos., mild TBI patients still ~2x more likely to have PTSD.

	Adj OR	CI
PTSD (DSM-IV)	2.32	1.11, 4.84
PTSD (DSM-5)	1.92	0.89, 4.14

PTSD dx, >12x more likely to report high disability.

mTBI → disability only with psychiatric co-morbidity.

# Mild TBI and persistent neuropsychological deficits?



# Poll Question #2

- Are most Veterans that you care for with both PTSD and history of TBI (pick one answer):
  - More comfortable with a mental health etiology for cognitive deficits
  - More comfortable with TBI as an etiology for cognitive deficits
  - Equally comfortable/uncomfortable with either etiology

## Objective 2:

To examine longitudinal associations of TBI with both PTSD symptoms and neuropsychological performance

Gaps in knowledge regarding long-term OEF/OIF deployment mental health outcomes.



## Objective 3:

To examine longitudinal associations both in the short-term and long-term



# Methods

**Iraq  
Deployment**



**12 mo**

**M = 7.9 years**

**Pre  
2003-05**

**Post  
2004-06**

**Follow-up  
2009-14**

**VA CSP 566**



# Eligibility for CSP 566

Iraq deployment

Prior permission for contact for future research

Pre and post neuropsych assessment

Valid neuropsychological performances

No physical condition precluding testing

Living in US

# Measures

Demographics

Written survey

Military history

Written survey/DMDC records

Combat exposure

DRRI Combat Exp Scale

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PTSD severity

PTSD Checklist (PCL-C)\*

TBI

Structured interview

\*Highly correlated with CAPS (PTSD gold standard interview) summary score at long-term follow-up

# Neuropsychological Measures

Effort/Engagement

TOMM, Trial 1

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Verbal learning

WMS-III Verbal Paired Assoc I

Verbal memory

WMS-III Verbal Paired Assoc II

Visual learning

WMS Visual Reproductions IR

Visual memory

WMS Visual Reproductions DR

Simple reaction time

ANAM Simple Reaction Time



Sustained attention

NES Continuous Perf Test



# VA CSP 566 (Long-term Follow-Up) Procedures

- Phone interview (CAPS, TBI interview)



- Mailed questionnaires

- In-person  
neurocognitive tests



	Pre-deployment	Long-term follow-up
Age, mean (SD), years	26.1 (6.1)	35.3 (6.1)
% women	5.9	5.9
% racial/ethnic minority	28.8	28.8
Education		
% high school/GED	66.9	21.3
% > high school	33.1	78.7
Duty status		
% regular active duty	84.6	31.3
% reservist	15.4	17.7
% military veteran	0	51.1
Cumulative OEF/OIF deployments		
0	98.5	0
1	1.5	39.3
>1	0	61.7
% PTSD cases	4.8	19.9

# TBI Characteristics

<b>Pre- to post-deployment (n = 73 reporting TBI)</b>	<b>%</b>
> 1 TBI	37.0
Mild TBI (most serious event)	89.0
Time, most recent event to post-deployment	
<1 month	0
1 – 3 months	1.4
>3 months – 1 year	79.4
> 1 year	19.2
<b>Post-depl. to long-term follow-up (n = 60 reporting TBI)</b>	<b>%</b>
> 1 TBI	43.3
Mild TBI (most serious event)	83.3
Time, most recent event to long-term follow-up	
<1 month	0
1 – 3 months	1.7
>3 months – 1 year	3.3
> 1 year	95.0



# Predicting PTSD Severity from Earlier Neuropsych and TBI

Variable	Post-deployment PCL	Long-term follow-up PCL
Earlier PCL-C score	(+) $p < .001$	(+) $p < .001$
Visual Reproductions, immed recall	$p = .05$	(-) $p = .02$
Visual Reproductions delayed recall	ns	(-) $p = .01$
Verbal Paired Assoc, I	ns	ns
Verbal Paired Assoc II	ns	ns
Simple Reaction Time throughput	ns	ns
CPT log-transformed omissions	ns	ns
TBI present	(+) $p < .001$	(+) $p = .01$
Combat severity (DRRI)	ns	(+) $p < .001, < .002$

Adjusted for age, race, education, marital status, duty status

# Relationships of PTSD Symptom Change and TBI to Post-deployment Neuropsychological Performance

Variable	Neuropsychological outcomes					
	Visual Reprod		Verbal Pairs		Simple RT	CPT
	Immed	Delay	I	II	Throughput	Omissions*
Pre-deployment neuropsych value	(+) P<.001	(+) P<.001	(+) P<.001	(+) P<.001	(+) P<.001	(+) P<.001
PTSD severity increase (PCL-C)	ns	ns	ns	(-) P=.03	(-) P=.05	ns
TBI present during interval	ns	ns	ns	ns	ns	ns
Combat severity (DRRI)	ns	ns	ns	ns	ns	ns

Adjusted for age, race, education, marital status, duty status

\*log-transformed

# Relationships of PTSD Symptom Change and TBI to Long-term Neuropsychological Outcomes

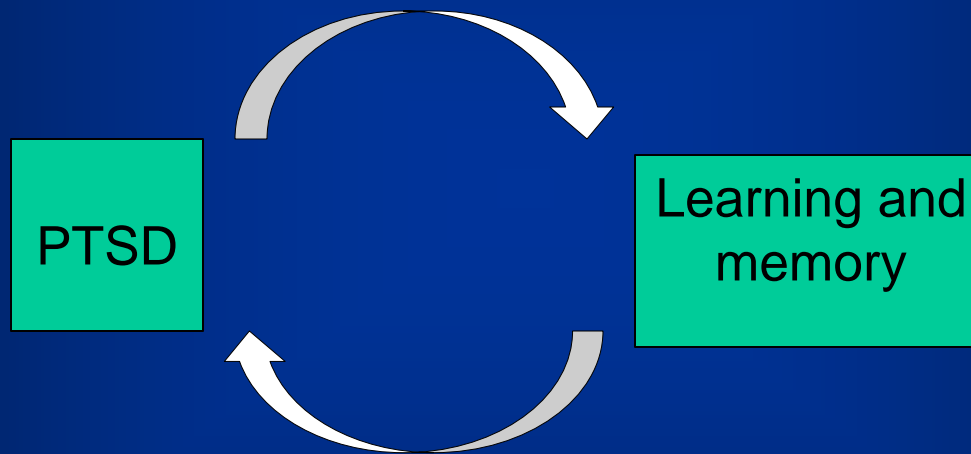
Variable	Neuropsychological outcomes					
	Visual Reprod		Verbal Pairs		Simple RT	CPT
	Immed	Delay	I	II	Throughput	Omissions*
Post-deployment neuropsych value	(+) P<.001	(+) P<.001	(+) P<.001	(+) P<.001	(+) P<.001	(+) P =.02
PTSD severity increase (PCL-C)	(-) P = .03	(-) P =.001	ns	(-) P =.03	ns	ns
TBI present during interval	ns	ns	ns	ns	ns	ns
Combat severity (DRRI)	ns	ns	ns	ns	ns	ns

Adjusted for age, race, education, marital status, duty status

\*log-transformed

# Summary

- Better visual learning and memory predictive of less severe PTSD symptoms on subsequent assessments.



- Although cognitive performances were WNL, PTSD symptom increases associated with poorer learning and memory (both assessments).

# Summary

- TBI not associated with neuropsychological deficits.
- In contrast, TBI associated with more severe PTSD symptoms at both post-deployment and long-term follow-up.
- Relationship of TBI with poorer PTSD outcomes not explained by combat severity

# Limitations

- TBI screened retrospectively; most mild
- Limited range of neuropsychological functions assessed
- Common co-morbidities not assessed (e.g., chronic pain, depression)
- Too few women to examine gender

# Conclusions

- Relationships among PTSD, TBI, and neuropsychological functioning may over time lead to sustained emotional and neurocognitive symptoms.
- Assessing for, and addressing, PTSD in Veterans presenting with TBI is important over both the short and long term.
- Augmentative strategies to enhance cognitive skills, and especially memory processes, may be helpful.

# NDHS Phase I Scientific Team

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**Co-PI:** Susan P. Proctor, DSc

**Co-Is:** Robert Kane, PhD  
COL (ret) Paul Amoroso, MD, MPH

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Roberta F. White, PhD



## NDHS Phase II: VA CSP #566

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Biostatistician:	M. Aslan
Boston Site PI:	B. Marx
Seattle Site PIs:	M. McFall/ M. Jakupcak
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S. Proctor, P. Schnurr, G. Huang, T. Gleason	

# QUESTIONS/COMMENTS?

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For more information about PTSD:

<http://www.ptsd.va.gov>

THANK YOU!