Cognitive Rehabilitation and TBI

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Overview

- What is Cognition?
- Treatment of Cognitive dysfunction
 - Learning and memory
- Can treatment be Effective?



"It is much more important to know what sort of a patient has a disease than what sort of a disease a patient has."

> William Osler (1849- 1919)



What is Cognition?

Dictionary: "the act or process of knowing"



What is Cognition?

- <u>Receptive Functions</u>
 - Sensory input, paying attention, rapid processing incoming information
- Learning and Memory
 - Acquiring, storing, retrieving
- Thinking
 - Mental organization and manipulation
- Execution and expressive functions

– Acting upon and communicating intentions



Cognitive Deficits in TBI

- processing speed/understanding information
 - Slowness of thought
- Attention and Concentration
- Learning and memory of new Information
- Planning, organization, initiation of action
- Reasoning, problem solving, judgment
- Language and communication
- Spatial disorientation (problems with perception, direction)
- Difficulties recognizing own cognitive deficits/limits
- Inappropriate, embarrassing or impulsive behavior



Functional Outcomes from Inpatient Rehabilitation after Traumatic Brain Injury: How do Hispanics Fare?

> Juan Carlos Arango-Lasprilla, Ph.D. Mitchell Rosenthal, Ph.D. John DeLuca, Ph.D. David Cifu, M.D. Robin Hanks, Ph.D. Eugene Komaroff, Ph.D.

Archives of Physical Medicine and Rehabilitation, 2007, 88, 11-18



Functional Outcomes from Inpatient Rehabilitation after Traumatic Brain Injury: How do Hispanics Fare?

- Examine rehabilitation outcome in Hispanic vs Caucasian TBI survivors
- Longitudinal data of the TBI Model Systems database
- 3056 persons with TBI
 - 2745 Caucasian
 - 311 Hispanic
- Outcome from rehabilitation at
 - Admission
 - Discharge
 - 1-yr follow-up



Functional Independence Measure (FIM)

	Hispanics	whites	р
Admission	54.0 <u>+</u> 24.8	54.5 <u>+</u> 26.3	ns
Discharge	95.1 <u>+</u> 22.5	96.2 <u>+</u> 23.4	ns
1-yr follow-up	113.2 <u>+</u> 17.1	116.4 <u>+</u> 18.5	< .001



Arango et al, 2007

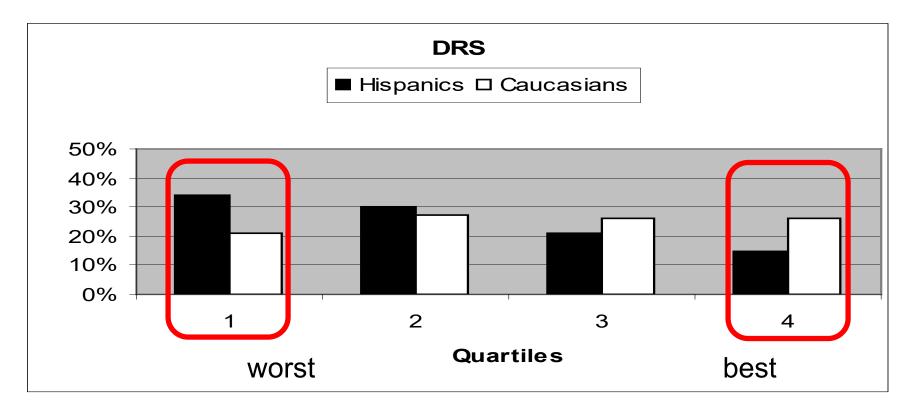
Disability Rating Scale

	Hispanics	whites	р
Admission	12.7 <u>+</u> 5.3	12.3 <u>+</u> 5.8	ns
Discharge	6.1 <u>+</u> 4.0	6.0 <u>+</u> 4.0	ns
1-yr follow-up	3.5 <u>+</u> 3.6	2.4 <u>+</u> 3.4	< .001



Arango et al, 2007

Disability Rating Scale (DRS) 1 year post Injury

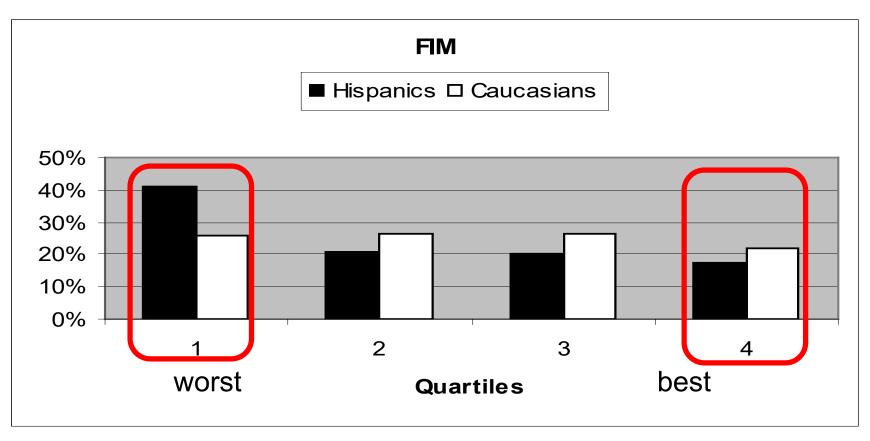


Odds ratio = 2.79



Arango et al., 2007

Functional Independence Measure (FIM) 1 year post Injury

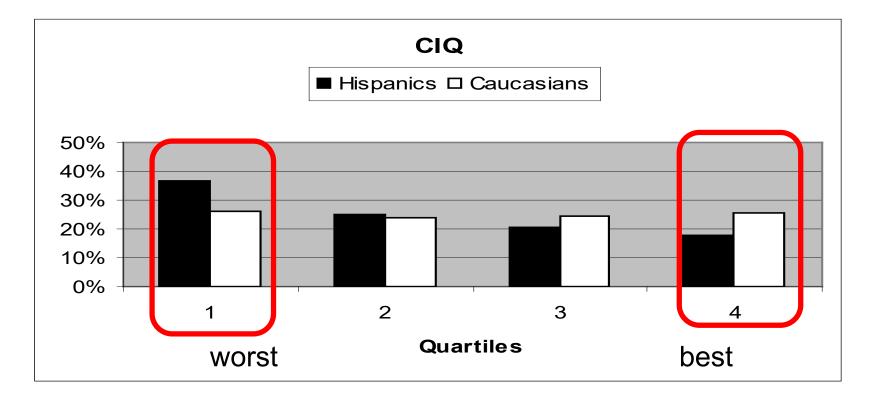


Odds ratio = 2.98



Arango et al., 2007

Community Integration Questionnaire (CIQ) 1 year post Injury



Odds ratio = 2.92



Arango et al., 2007

Functional Outcomes from Inpatient Rehabilitation after Traumatic Brain Injury: How do Hispanics Fare?

- Largest multicenter longitudinal study on ethnicity and functional outcome in TBI
- No differences at admission and discharge to rehabilitation
- 1 year post discharge, Hispanics were roughly 2.5 times more likely than Caucasians to have worse outcome across all outcome measures
- Understanding variables which lead to worse long –term outcome in Hispanics is needed



Overview

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- Treatment of Cognitive dysfunction – Learning and memory
- Can treatment be Effective?



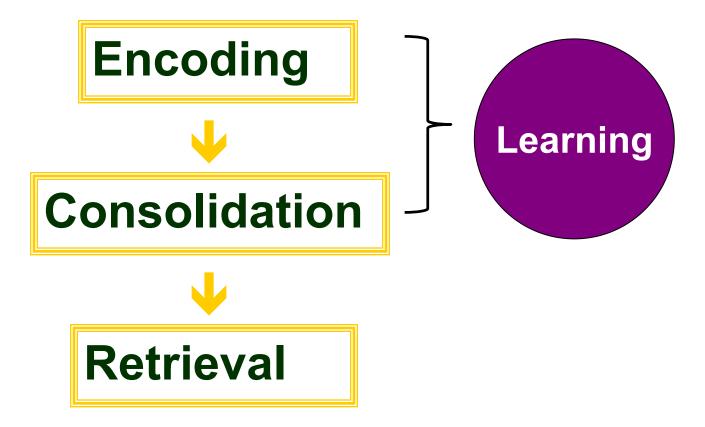
Defining Learning

- <u>Learning</u> "The *process* of acquiring new information"
- <u>Memory</u> "The *persistence* of learning in a state that can be revealed at a later time"

Squire, 1987



Learning and Memory Process





Identifying the Cause

- Retrieval failure hypothesis ?
- Acquisition deficits?



Acquisition Versus Retrieval Deficits in Traumatic Brain Injury: Implications for Memory Rehabilitation

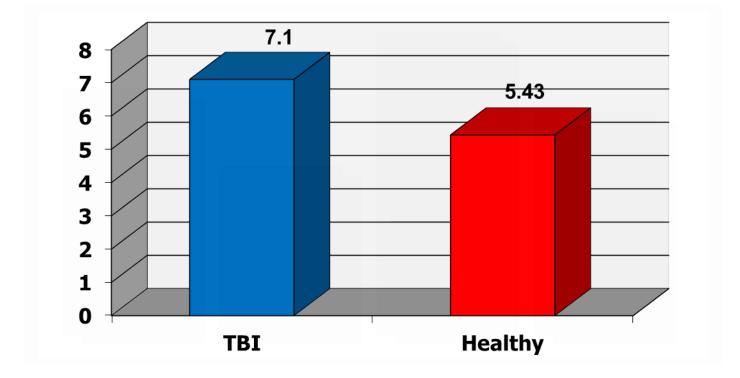
John DeLuca, PhD, Maria T. Schultheis, PhD, Nancy K. Madigan, PhD, Christopher Christodoulou, PhD, <u>Allison Averill, MD</u>

- 10 words Selective Reminding
- S's trained to learning criterion
 all words recalled on 2 consecutive trials
- S's examined on
 - Trials to reach criterion
 - Recall
 - Recognition



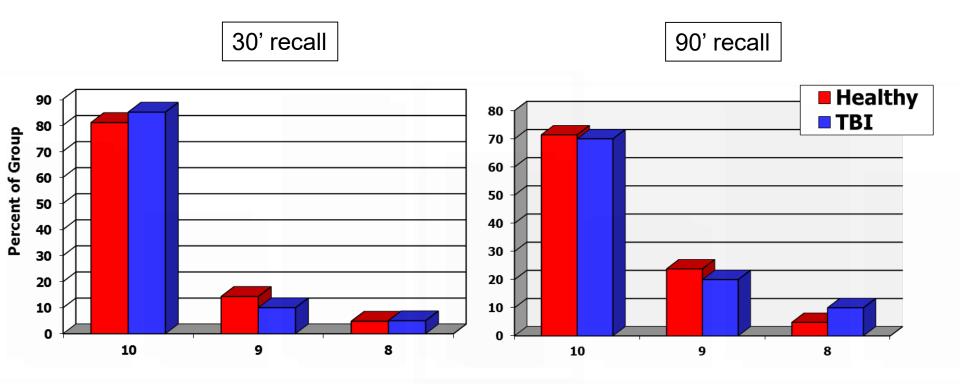
Archives of PM&R, 2000, 81, 1327-1333

Mean Trials to Criterion Open-Trial SRT

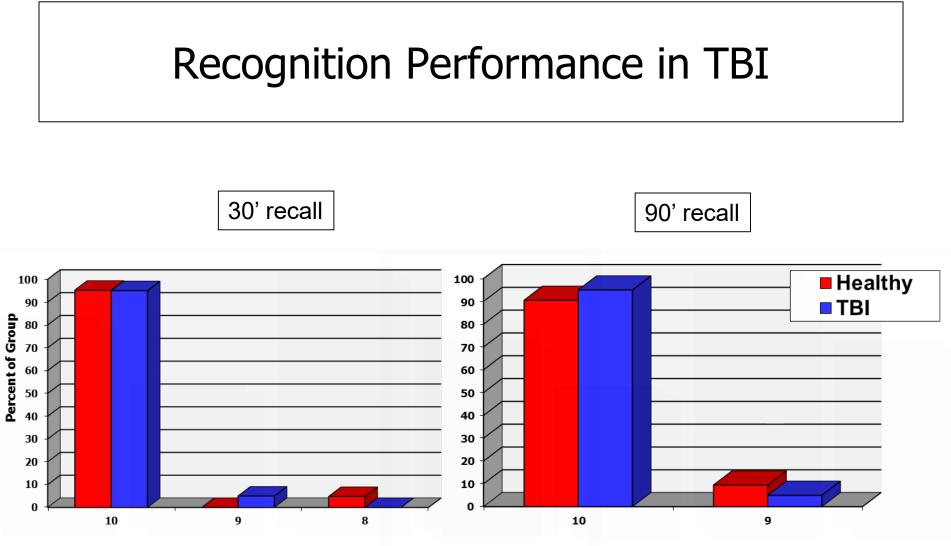




Recall Performance in TBI

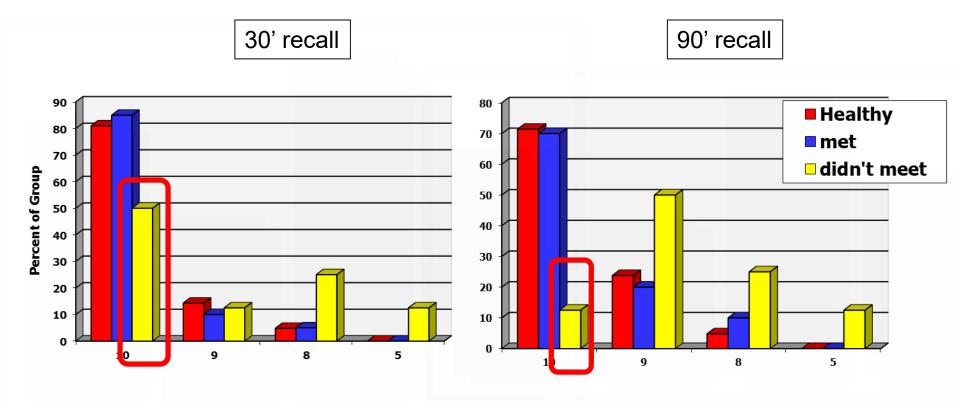








Recall in TBI who met and didn't meet criterion





Cognitive Performance in TBI Learners and non-Learners

	Learner	Non-learner	p-value	Cohen's d		
PS composite	7.1 (3.3)	4.5 (2.3)	.01	.79		
WM composite	9.7 (2.3)	7.7 (1.8)	.01	.97		
EF composite	8.5 (2.9)	5.8 (3.0)	.01	.92		
Verbal comp	10.0 (2.5)	7.8 (2.9)	.01	.81		
Percep comp	9.6 (3.0)	8.5 (3.1)	ns	.36		

No group differences in Anxiety or depression

Mean Standard score + SD



Chiou et al, 2015, *JCEN*

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Evidence-Based Cognitive Rehabilitation: Updated Review of the Literature From 2003 Through 2008

Keith D. Cicerone, PhD, Donna M. Langenbahn, PhD, Cynthia Braden, MA, CCC-SLP, James F. Malec, PhD, Kathleen Kalmar, PhD, Michael Fraas, PhD, Thomas Felicetti, PhD, Linda Laatsch, PhD, J. Preston Harley, PhD, Thomas Bergquist, PhD, Joanne Azulay, PhD, Joshua Cantor, PhD, Teresa Ashman, PhD

Substantial evidence to support interventions for:

- Attention
- Memory
- Social communication skills
- Executive functions
- Comprehensive-holistic neuropsychological rehabilitation program

"There is now sufficient information to support evidence-based protocols and implement empirically-supported treatments for cognitive disability after TBI and stroke"



Cognitive Rehabilitation

Sample RCT results



An RCT to Treat Learning Impairment in Traumatic Brain Injury: The TBI-MEM Trial

Nancy D. Chiaravalloti, PhD^{1,2}, Joshua Sandry, PhD^{1,2}, Nancy B. Moore, MA¹, and John DeLuca, PhD^{1,2} Neurorehabilitation and Neural Repair 2016, Vol. 30(6) 539–550 © The Author(s) 2015 Reprints and permissions: sagepub.com/journalsPermissions.nav DOI: 10.1177/1545968315604395 nnr.sagepub.com



69 Mod-severe TBI randomized to Tx vs Placebo

Context and imagery

Classification of Evidence: This study provides Class I evidence that the modified Story Memory Technique© behavioral intervention improves both objective memory and everyday memory in persons with TBI over 5 weeks



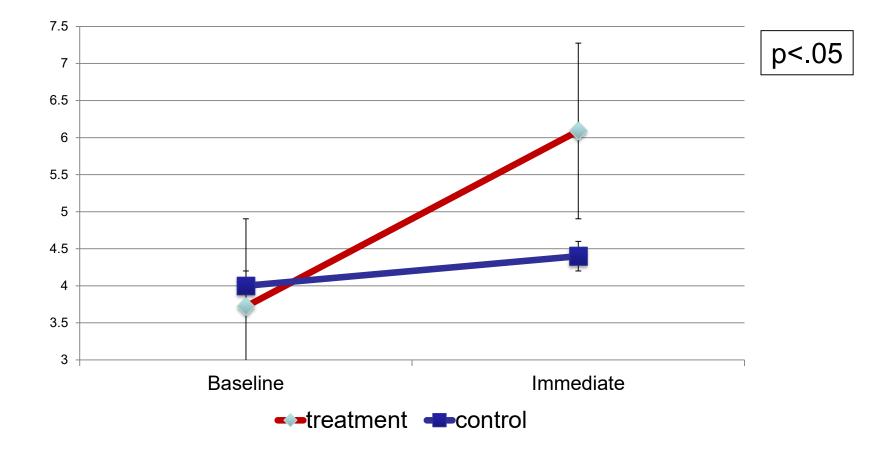
Modified Story Memory Technique

- Double-blind, placebo-controlled RCT
- 10 sessions
 - 2x per week for 5 weeks
 - 45-60 minutes in duration
- Two skills taught
 - Imagery (sessions 1-4)
 - Context (sessions 5-8)
- Generalization
 - How you use skills in daily life (sessions 9 and 10)



Chiaravalloti et al, 2016, NNR

MAS Prose Memory Delayed Recall: pre to post treatment



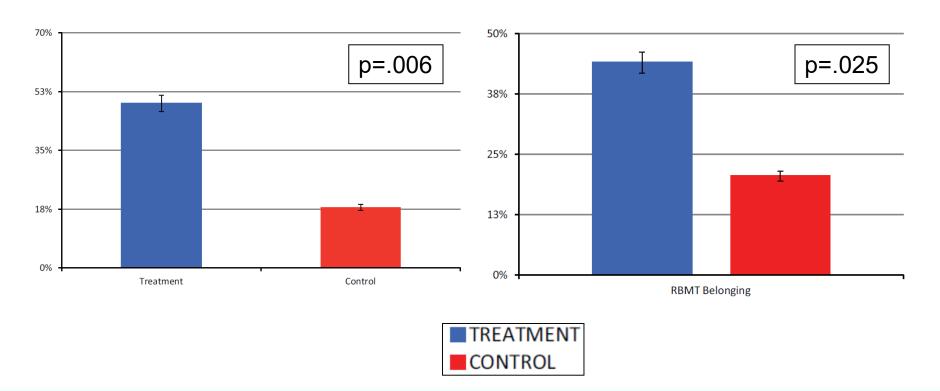


Chiaravalloti, et al., 2016, NNR, suppl data

Improvement on mSMT in TBI

% pre-post improvement in Prose Recall

% improving on RBMT – post tx

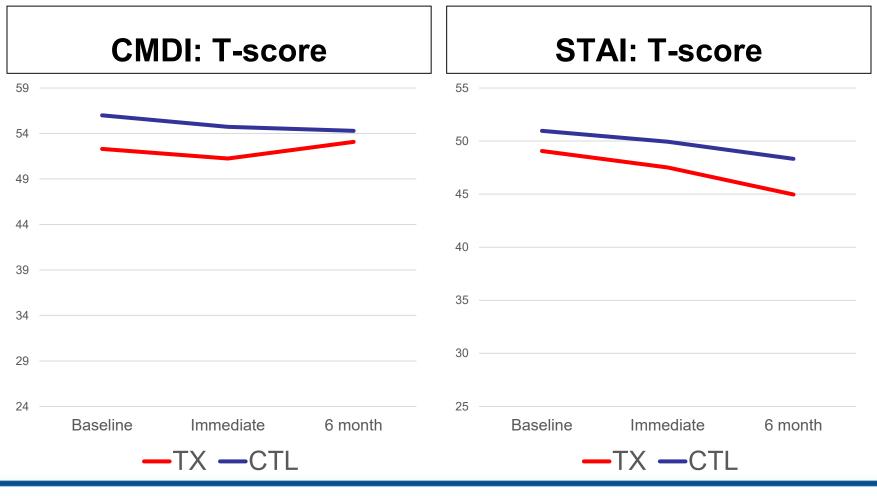




RBMT: Rivermead Behavior Memory test

Chiaravalloti et al, 2016, NNR

Depression and Anxiety





Chiaravalloti et al, 2016, NNR

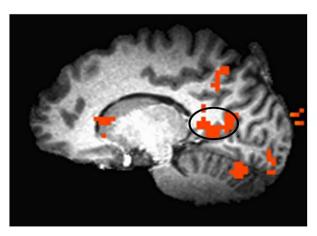
Examining the Efficacy of the Modified Story Memory Technique (mSMT) in Persons With TBI Using Functional Magnetic Resonance Imaging (fMRI): The TBI-MEM Trial

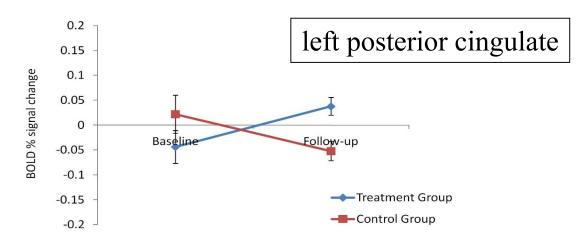
Nancy D. Chiaravalloti, PhD; Ekaterina Dobryakova, PhD; Glenn R. Wylie, DPhil; John DeLuca, PhD, ABPP

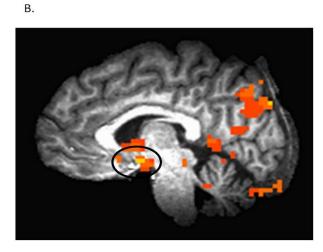


<u>Figure 1.</u> BOLD % signal change Posterior insula and cerebellum also showed increased activation in TX vs placebo

Α.







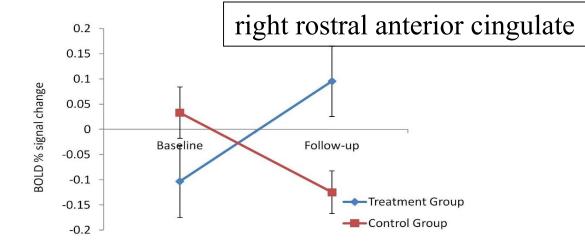
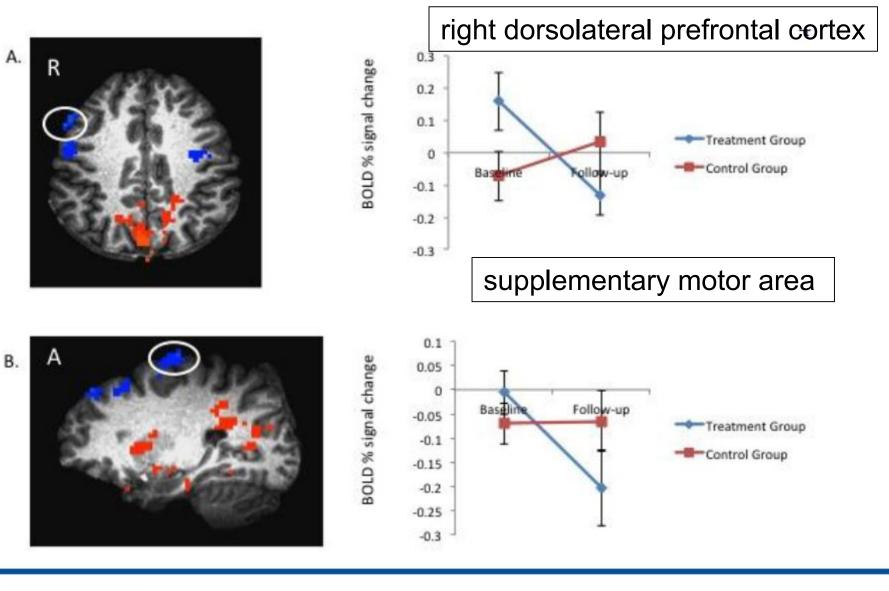




Figure 1. Areas of decreased BOLD % signal change





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- What is Cognition?
- Treatment of Cognitive dysfunction
 - Learning and memory

• Can treatment be Effective?

-Techniques to improve learning



Pilot Study to Examine the Use of Self-Generation to Improve Learning and Memory in People With <u>Traumatic Brain Injury</u>

Yael Goverover, Nancy Chiaravalloti, John DeLuca

 The <u>generation effect</u> is: items generated by subjects are remembered better than items presented
 Robust effect in Healthy subjects

Little work in Clinical samples



American Journal of Occupational Therapy, 2010, 64, 540–546

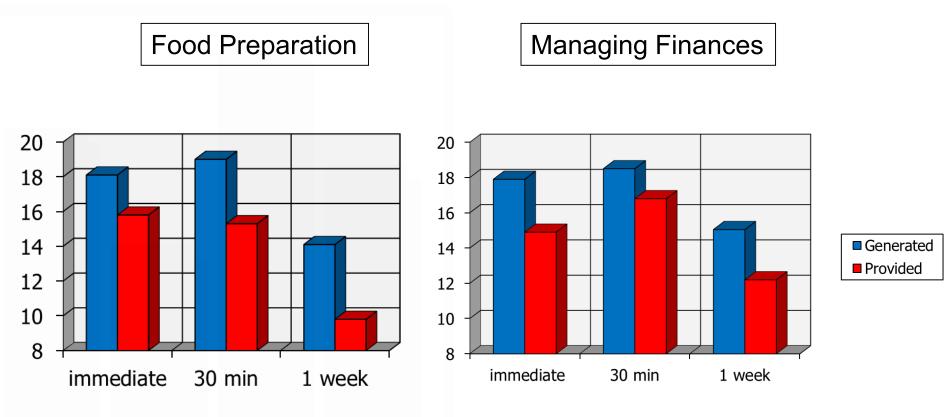
Everyday Life Tasks

- Participants were asked to complete two tasks:
- Food Preparation
- Financial
 - management
- Within-group design





Self-Generation and Everyday Life Activities in TBI





Goverover et al., 2010

Spaced Learning or "Spacing Effect"

- Is learning better if trials are spaced vs massed?
- Instructions on how to perform tasks were presented three times in two conditions:
 - Massed condition 1/2/3
 - Spaced condition 1____2___3
 - Within-group design

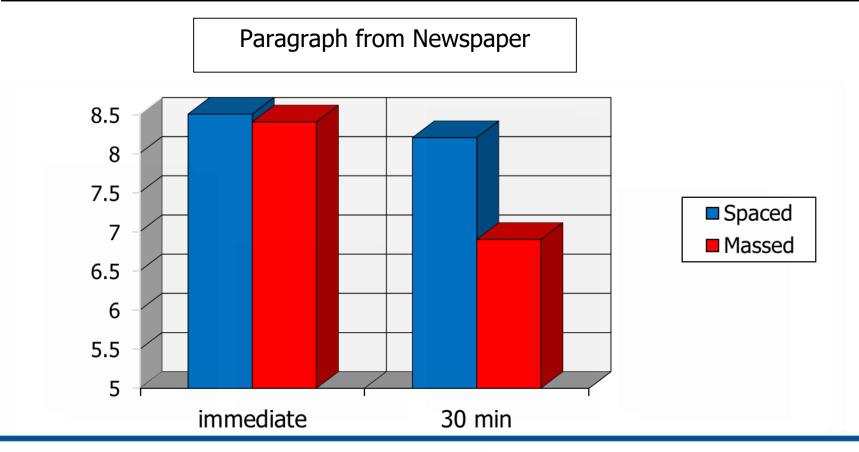


Goverover et al., J Exp Clin Neuro Psych, (2009)

Application of the Spacing Effect to Improve Learning and Memory for Functional Tasks in Traumatic Brain Injury: A Pilot Study

Yael Goverover, Juan Carlos Arango-Lasprilla, Frank G. Hillary,

Nancy Chiaravalloti, John DeLuca





The American Journal of Occupational Therapy September/October 2009, Volume 63, Number 5

Retrieval practice or "Testing Effect"

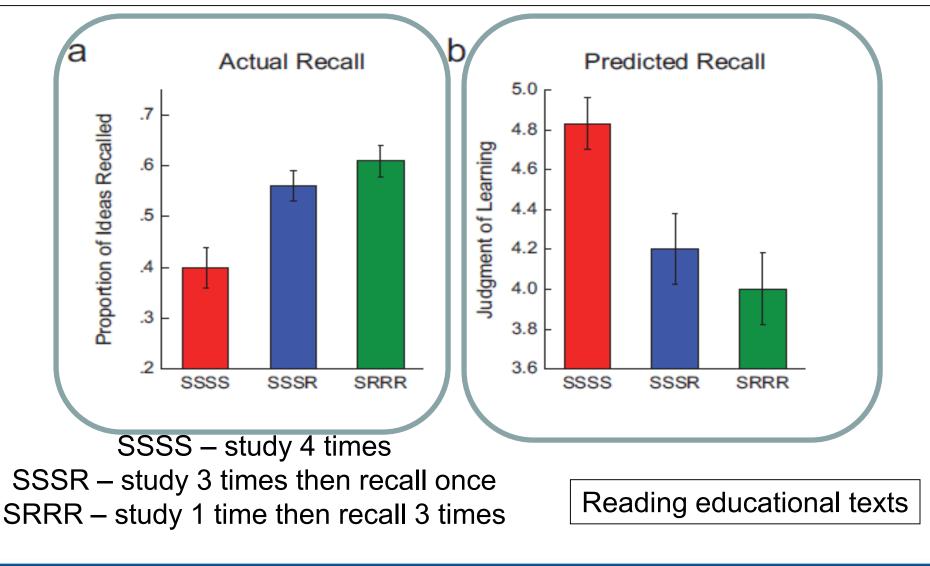
Which do you prefer for new learning

-4 opportunities to learn something

-1 opportunity then tested 3 times



Active Retrieval during Learning enhances deep and conceptual encoding

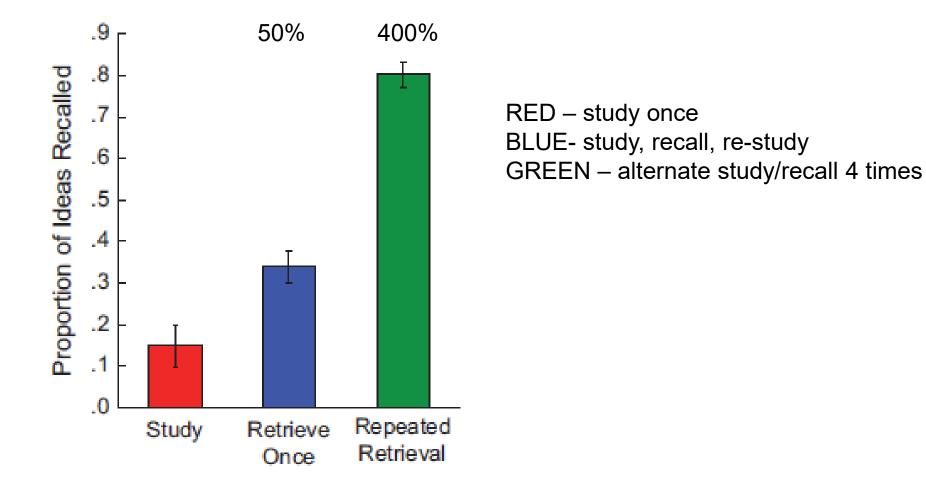


KESSLER FOUNDATION

Karpicke (2012) Psychological Science, 21(3) 157-163

Active Retrieval during Learning enhances deep and conceptual encoding

One week recall





Karpicke (2012) Psychological Science, 21(3) 157-163

Journal of the International Neuropsychological Society (2010), **16**, 1147–1150. Copyright © INS. Published by Cambridge University Press, 2010. doi:10.1017/S1355617710001128

BRIEF COMMUNICATION

Retrieval practice: A simple strategy for improving memory after traumatic brain injury

JAMES F. SUMOWSKI,^{1,2} HALI G. WOOD,¹ NANCY CHIARAVALLOTI,^{1,2} GLENN R. WYLIE,^{1,2} JEANNIE LENGENFELDER,^{1,2} and JOHN DELUCA^{1,2,3}

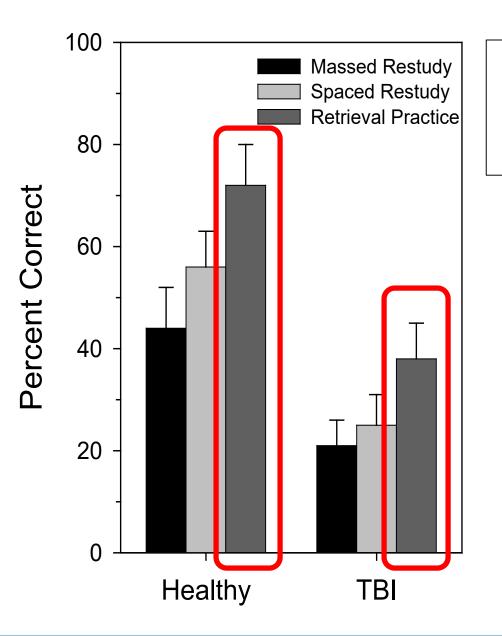


Testing Effect Demographics

	TBI	HC
	14	14
Age	38.4 <u>+</u> 12.6	44.9 <u>+</u> 12.1
Mdn age at TBI	26.4	
Mdn duration coma (days)	45.5	
Mdn verbal memory (CVLT)	1 st %ile	50 th %ile



Sumowski et al., JINS, (2010)



Testing Effect in TBI

Large effect size of Retrieval practice

Retrieval practice was best learning strategy In 93% of TBI S's

FOUNDATION

Sumowski et al, JINS (2010)





Archives of Physical Medicine and Rehabilitation

journal homepage: www.archives-pmr.org

Archives of Physical Medicine and Rehabilitation 2014;95:397-400



BRIEF REPORT

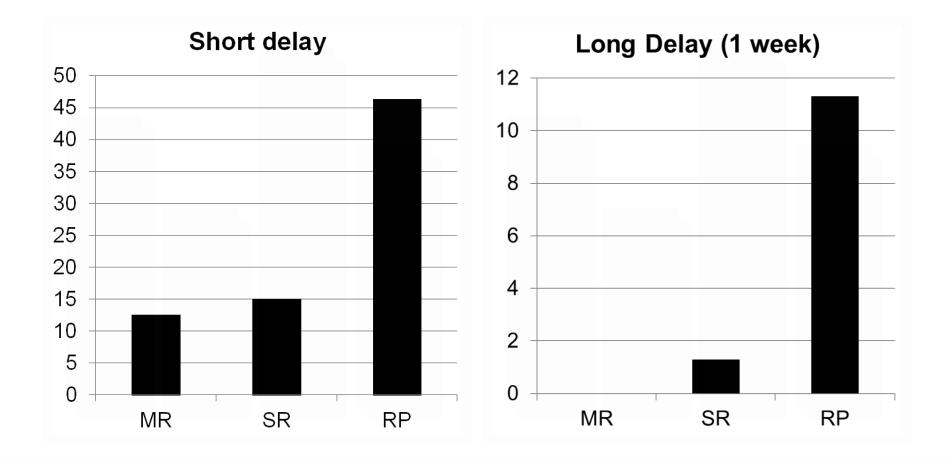
Retrieval Practice Improves Memory in Survivors of Severe Traumatic Brain Injury



James F. Sumowski, PhD,^{a,b} Julia Coyne, PhD,^{a,c} Amanda Cohen, BA,^a John DeLuca, PhD^{a,b}



Testing Effect in Severe TBI





Sumowski et al (2014), Arch PM&R

Overview

- What is Cognition?
- Treatment of Cognitive dysfunction
 - Learning and memory
- Can treatment be Effective?
 - Techniques to improve learning
- What else can be done?

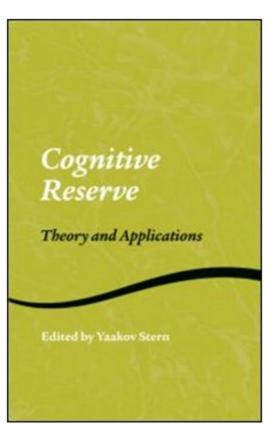
-Cognitive Reserve



Cognitive Reserve Hypothesis

Persons with higher lifetime intellectual enrichment can better withstand diseaserelated neuropathology without suffering cognitive impairment or dementia, likely due to more efficient neurocognitive processing.

> Stern et al., *JINS* 2002;8:448-460. Stern et al., *Cereb Cortex* 2005;15:394-402.









Archives of Physical Medicine and Rehabilitation

journal homepage: www.archives-pmr.org

Archives of Physical Medicine and Rehabilitation 2013;94:2562-4



BRIEF REPORT

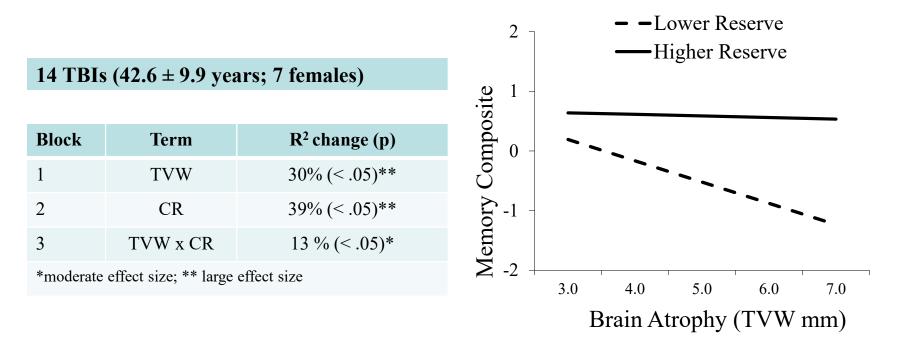
Education Attenuates the Negative Impact of Traumatic Brain Injury on Cognitive Status



James F. Sumowski, PhD,^{a,b} Nancy Chiaravalloti, PhD,^{a,b,c} Denise Krch, PhD,^{b,c} Jessica Paxton, PhD,^{a,b} John DeLuca, PhD^{a,b,d}



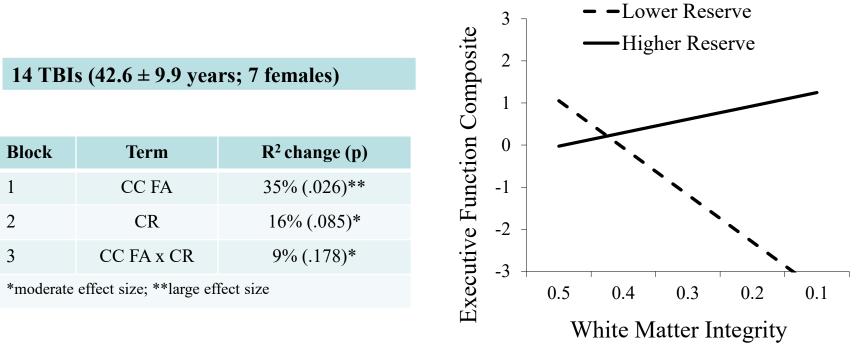
Cognitive Reserve Moderates the Effect of TBI Brain Pathology on Memory





Sumowski et al (2013), Archives PM&R

Cognitive Reserve Moderates the Effect of TBI Brain Pathology on Executive Function



(Corpus Callosum FA)



Sumowski et al (2013), Archives PM&R

Cognitive Reserve and Rehabilitation

- Higher cognitive reserve protects TBI subjects
 from TBI-related cognitive decline
- Can we identifying "at risk" patients for cognitive impairment?
- Can one build up a "cognitive reserve"?
 - "neuroprotective" against developing cognitive impairment?



What is the Potential Impact of Cognitive Rehabilitation?

ACTIVE study



Published in final edited form as: JAMA. 2002 November 13; 288(18): 2271–2281.

Effects of Cognitive Training Interventions With Older Adults:

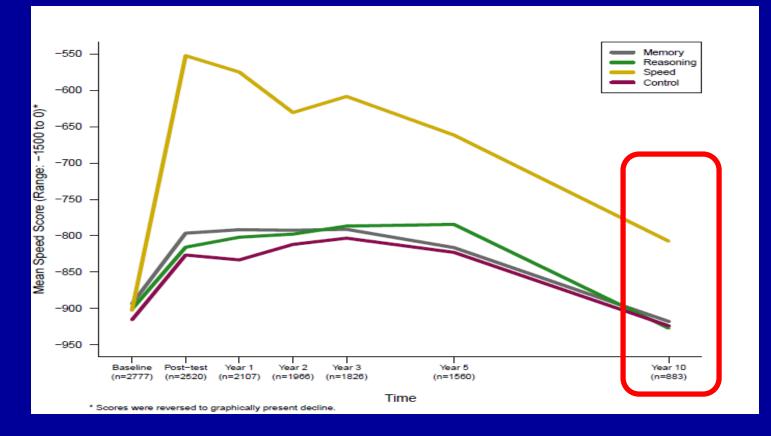
A Randomized Controlled Trial

Karlene Ball, PhD, Daniel B. Berch, PhD, Karin F. Helmers, PhD, Jared B. Jobe, PhD, Mary D. Leveck, PhD, Michael Marsiske, PhD, John N. Morris, PhD, George W. Rebok, PhD, David M. Smith, MD, Sharon L. Tennstedt, PhD, Frederick W. Unverzagt, PhD, and Sherry L. Willis, PhD for the ACTIVE Study Group

2832 Participants 65-94 yo Randomized into 1 of 4 groups (10 sessions of group training) verbal episodic memory (n=711) reasoning (n=705) Processing speed (n=712) no contact control (n=704) Outcome: immediate, 1yr, 2yr, 3 yr, 5 yr 10yr post training

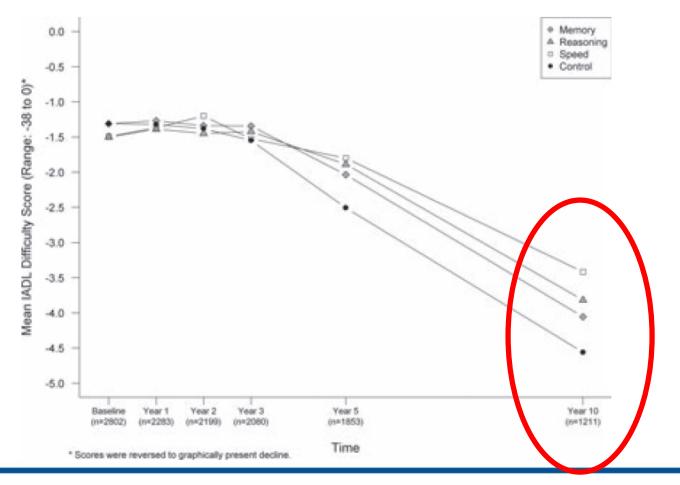
ACTIVE Results at 10 years

Mean PS Score



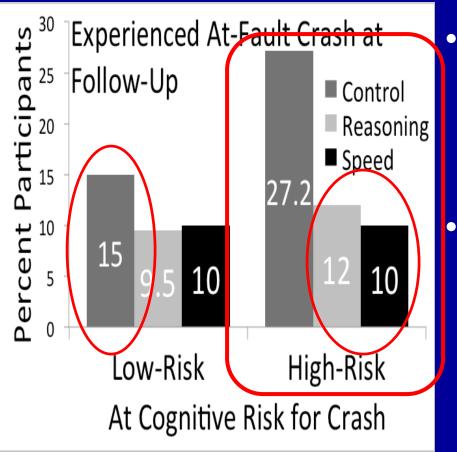
(JAGS 2014)

ACTIVE Results at 10 years Mean IADL





Reduction in auto crashes: Comparison of speed & reasoning training & control group: 5- and 10-yr findings



Reduced Crashes for participants in <u>Speed</u> <u>& Reasoning Training</u> at high risk for crash
For training groups, % high risk participants who crashed was reduced to level of low

risk participants

Courtesy of Dr. George Rebok

What is the Potential Impact of Cognitive Rehabilitation?

- ACTIVE trial in healthy aging:
- Cog Rehab and Dementia Risk:
 - 78.8% relative risk reduction of dementia 10 years later
 - magnitude of this effect is greater than:
 - the of 20-40% relative risk reduction antihypertensive medications provide over 3-5 years against *:
 - stroke
 - coronary heart disease
 - heart failure

* Lancet 2003;362:1527-35



Edwards et al (2017) Alzheimer's & Dementia

Time Magazine 1/20/2003

I love my life way too much to just hand it over to Alzheimer's.

Fighting Back with Aricept'

ARICEPT" is well tolerated but may not be for everyone

Some people may experience nausea, diarrhea, not sleeping well, vomiting, muscle cramps, feeling very tired, or hot wanting to eat. In studies, these side effects were usually mild and went away over time. Some people taking ARICEPTE may

experience fainting. People at risk for ulcers should tell their doctors because their condition may get worse.

Please see additional important product information on accompanying page

When my memory started failing, I knew I had to see my doctor. He put me on ARICEPT." Now I'm doing better."

> If a person forgets names, places or facts – and has trouble with everyday things like reading or shopping – it may not be normal aging. It could be Alzheimer's disease. So it's important to see a doctor as soon as you can.

There is no cure for Alzheimer's. But a prescription drug called ARICEPT[®] has been used by millions of people to help their symptoms.

In studies, ARICEPT[®] has been proven to work for mild to moderate Alzheimer's. It has helped people improve their memory over time. It has also helped them to keep doing everyday things on their own, longer.

Ask your doctor if ARICEPT[®] is right for you or your loved one. It is the #1 prescribed drug for Alzheimer's in the world. The sooner you know it's Alzheimer's, the sooner ARICEPT[®] can help.



Strength in the face of Alzheimer's

To learn more and to receive a memory checklist, call 800-760-6029 ext.91 or visit www.aricept.com

 Individual responses to ARICEPT® can be different – people may get better, stay the same or not get better.

"When my memory started Failing, I knew I had to see my Doctor. He put me **On ARICEPT.** Now I'm doing better"



Video Games and Cognitive Rehabilitation

– Can I tell my client to use "brain games" or "video games" for cognitive rehabilitation?

A Consensus on the Brain Training Industry from the Scientific Community

Max-Planck-Institut für Bildungsforschung Max Planck Institute for Human Development

October 20, 2014



75 Leading Cognitive Psychologists & Cognitive Neuroscientists Representing 48 Universities

"We object to the claim that brain games offer consumers a scientifically grounded avenue to reduce or reverse cognitive decline when there is no compelling scientific evidence to date that they do."



"A Consensus on the Brain Training Industry...", accessed (April 29, 2015), http://longevity3.stanford.edu/blog/2014/10/15/the-consensus-on-the-braintraining-industry-from-the-scientific-community/

Lumosity to Pay \$2 Million to Settle FTC Deceptive Advertising Charges for Its "Brain Training" Program

- "Lumosity preyed on consumers' fears about age-related cognitive decline, suggesting their games could stave off memory loss, dementia, and even Alzheimer's disease, But Lumosity simply did not have the science to back up its ads."
- Lumosity claimed that training would:
 - 1) improve performance on everyday tasks, in school, at work, and in athletics
 - 2) delay age-related cognitive decline and protect against mild cognitive impairment, dementia, and Alzheimer's disease
 - 3) reduce cognitive impairment associated with health conditions, including stroke, traumatic brain injury, PTSD, ADHD, the side effects of chemotherapy, and Turner syndrome, and that scientific studies proved these benefits.

https://www.ftc.gov/news-events/press-releases/2016/01/lumosity-pay-2-million-settle-ftc-deceptive-advertising-charges



Federal Trade Commission Press release, 1/6/2015

Conclusions

- Cognitive rehabilitation works in TBI
- Treatment is available and effective
- Cognitive rehabilitation works through neuroplasticity in the brain
- Can we build a "cognitive reserve" after TBI?
- Future research needed to look at factors which predict response to treatment



Acknowledgments

Nancy Chiaravalloti, Ph.D. Yael Goverover, Ph.D. Jean Lengenfelder, Ph.D. Glenn Wylie, D.Phil Nancy Moore, M.S. James Sumowski, Ph.D. Kathy Chiou, Ph.D. Denise Krch, Ph.D. Joshua Sandry, Ph.D.







Thank You

