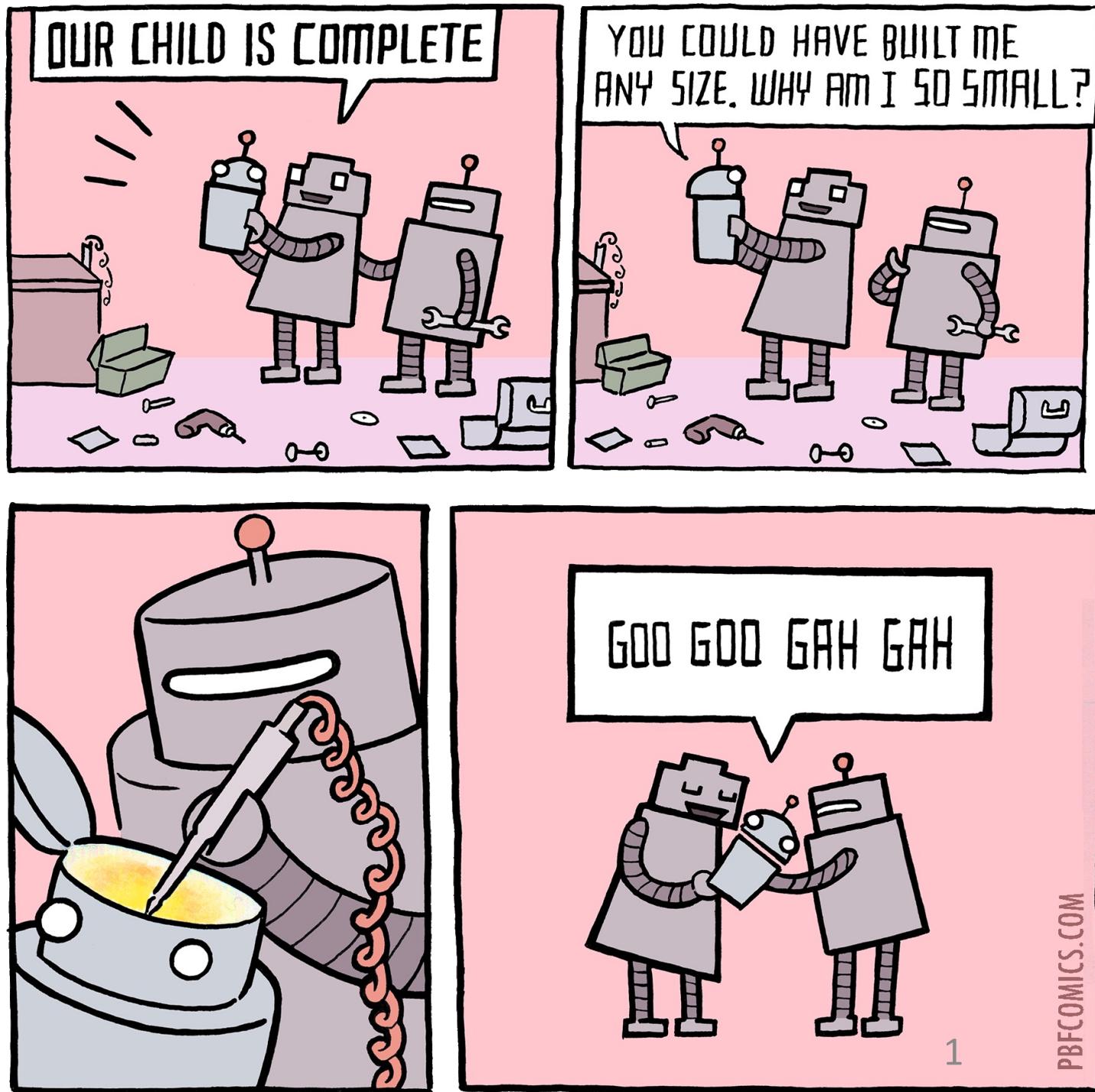
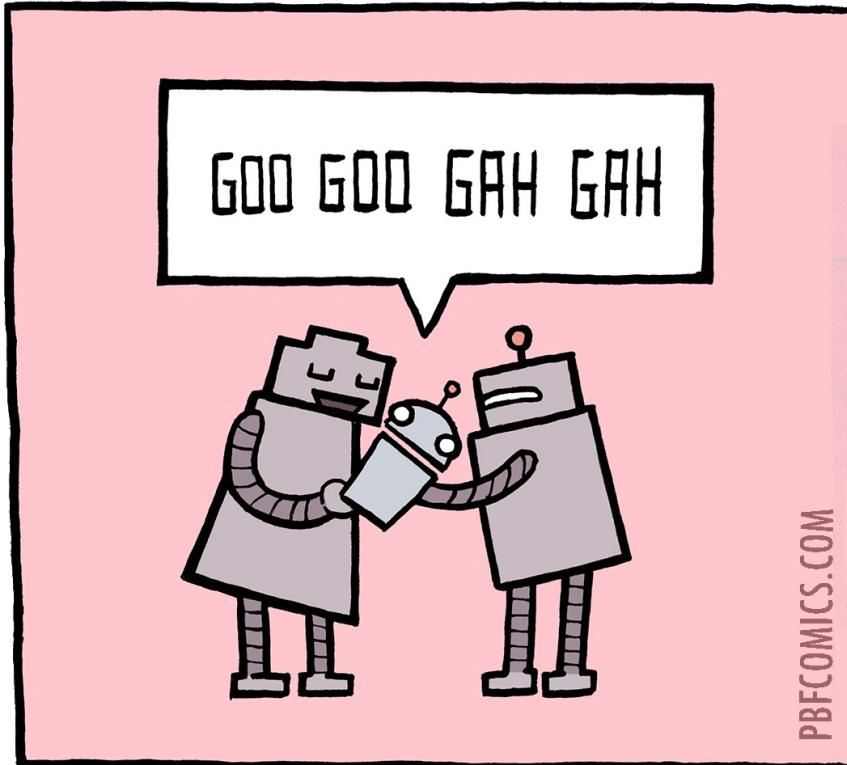


PSYC301: Factors influencing recovery following brain dysfunction

Jay Hosking, PhD



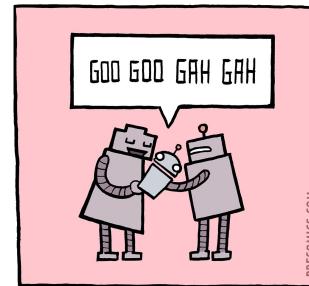
Lecture outline



- A. Common misconceptions
- B. Anosognosia
- C. Effect of age
- D. Effect of lesion size and location
- E. Chronic vs. acute dysfunction
- F. Secondary effects
- G. Environmental factors
- H. Compensatory strategies
- I. Neuroplasticity
- J. Therapy dose

Learning objectives

1. To consider some common misconceptions about brain injury and how those misconceptions might affect recovery of function in a brain-injured person.
2. To review and appreciate evidence supporting the notion that self-awareness is an important factor related to successful recovery.
3. To consider how age, the extent of damage, and the secondary effects influence recovery of function across multiple domains.
4. What is the relationship between environment and recovery?
5. What does restitution vs. substitution mean, in the context of recovery? What sort of recovery are we usually seeing?
6. Describe neuroplasticity as related to both recovery and the healthy brain.
7. How could you test whether anosognosia is real or is simply the participant refusing to admit their dysfunction? (Hint: Ramachandran has some ideas.)
8. (Required) Read the paper by Duque *et al.* 2022 on what we know about adult neurogenesis in humans. Based on this information, what role do you think neurogenesis plays in recovery? What do you think about its therapeutic value?



It's true that chronic stress is a risk factor for brain dysfunction, depression, anxiety, cancer malignancy, and autoimmune disorders, BUT that unfortunately leads to quotes like:

“Cancer and ALS and MS and rheumatoid arthritis and all these other conditions, it seems to me, happen to people who have a poor sense of themselves as independent persons... they have a poorly differentiated sense of self. ... The people I see with cancers and all these conditions have difficulty saying no and expressing anger.”

– Gabor Maté, *When the Body Says No*

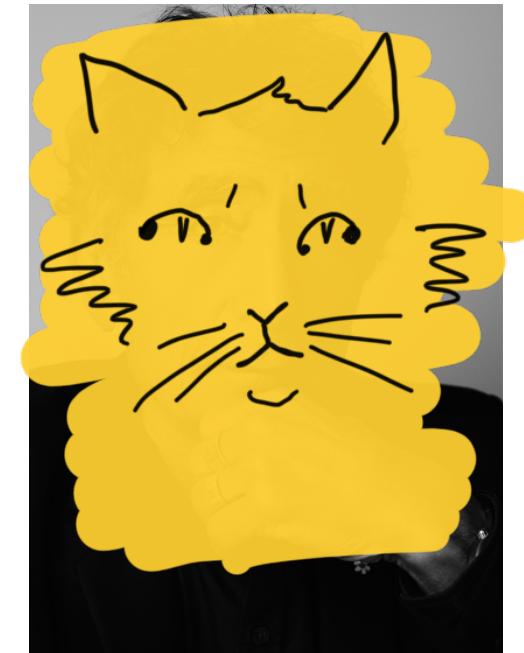
It happens in both brain dysfunction and also recovery.

Be careful of these logical fallacies!

Yes, both positivity/attitude and effort/motivation can have mild beneficial effects on recovery,

BUT

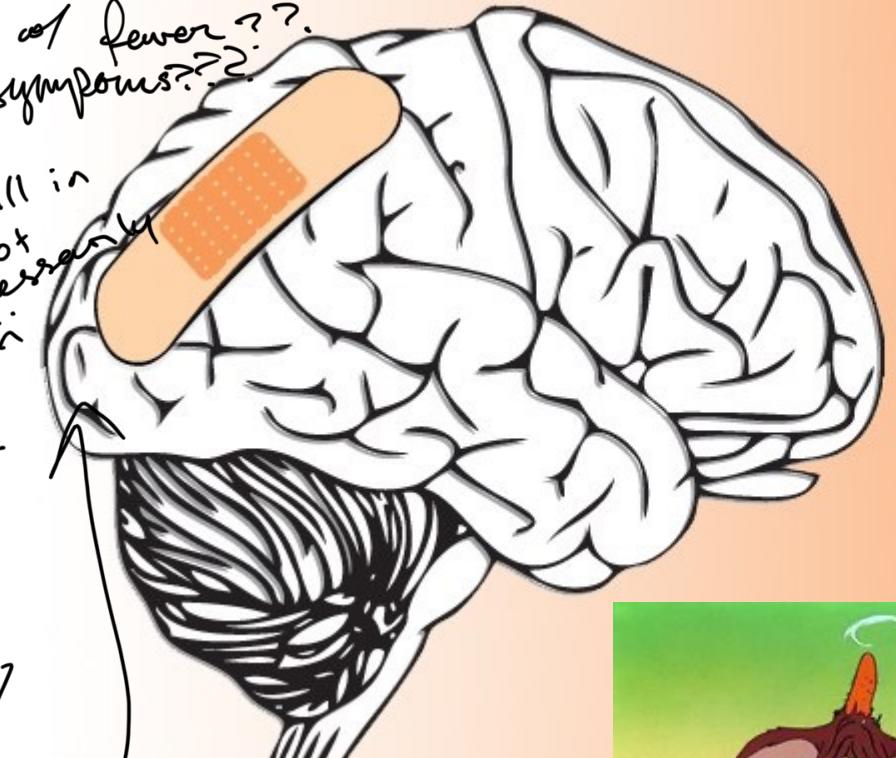
If someone has poor recovery/outcome, this does not indicate a failure of positivity/attitude or effort/motivation!



Common misconceptions about brain dysfunction

maybe it's easier to
have a positive attitude w/ fewer??
symptoms??

1. Many people falsely believe that recovery from brain dysfunction depends primarily on the injured person's efforts or attitude.
*"his" still in
there were not necessary*
2. Many people falsely believe that a person with even severe brain dysfunction can fully recover.
3. Many people misattribute a behaviour (e.g., fatigue, aggression) that is the result of brain dysfunction to the person's personality or life stage. → "grumpy old man" may be referred to / caused by dysfunction
4. Many people falsely believe that brain dysfunction must have **visible effects** on the person. *literally putting
bandaid makes pp' symp. affective*
5. Many people falsely believe that emotional problems after brain injury are usually tangential to the brain dysfunction. → *emotional affects can be direct effect of TBI*
6. Many people falsely believe that a second brain injury can restore lost memories in individuals with amnesia.



TBI = traumatic brain injury

Common misconceptions about brain dysfunction

“...a brain dysfunctional patient does not appear to be aware of impaired neurological or neuropsychological functioning which is obvious to the clinician...”

Visible evidence contradicts pt's description → are they in denial or actually unaware?

ie paralyzed arm

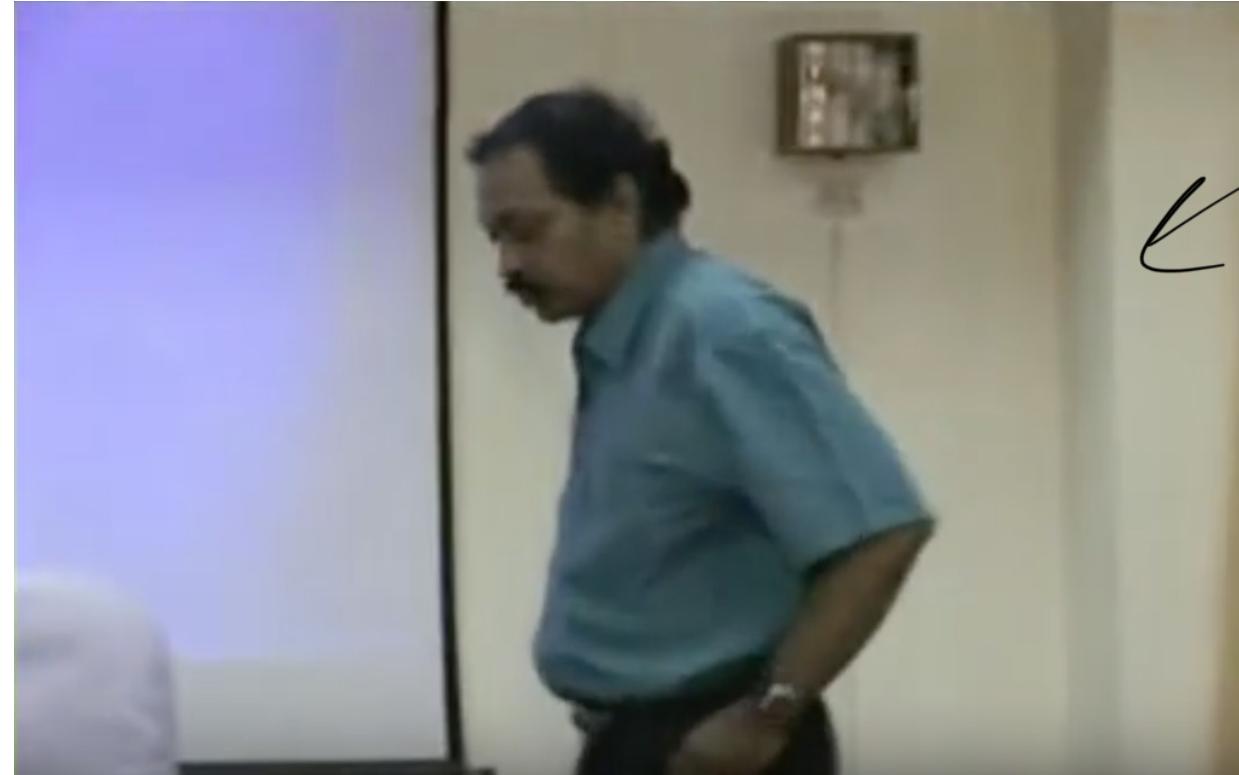
pt: arm is fine

dr: move it then

pt: I don't want to

present of 1 vs 2

handed task: choose
2 handed → truly unaware



good
neuro
researcher!
Now we experiment

<https://www.youtube.com/watch?v=MDHJDKPeB2A>

Anosognosia

Links to Anosognosia

These are all v. different disorders!
→ No one thing tied to "self-awareness"

Observed in various forms of brain dysfunction:

Psychiatric disorders (schizophrenia, bipolar)

Movement disorders

Contralateral neglect → failure to pay attn to L/R side
of world

Memory disorders

Stroke

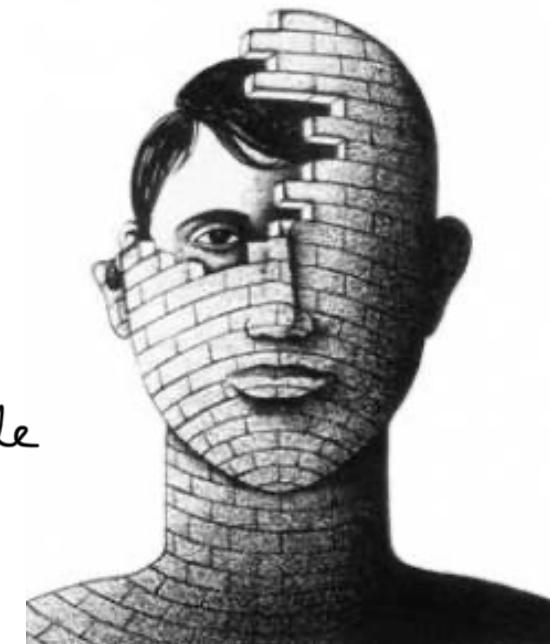
Dementia

Dysexecutive syndrome ("frontal lobe syndrome"):

Disinhibition, aggression, impulsiveness,
disruptions in planning, working memory

Executive functions &
other "higher" (v's)
of cogn. (judgment)

Anosognosia



Prigatano & Wong (1999)

- When prediction of performance improves, the likelihood of achieving rehabilitation goals improves

→ ↑ Anosognosia aka ↓ self-awareness = ↓ recovery

Cognitive and Affective Improvement in Brain Dysfunctional Patients Who Achieve Inpatient Rehabilitation Goals

George P. Prigatano, PhD, Jane L. Wong, PhD

ABSTRACT. Prigatano GP, Wong JL. Cognitive and affective improvement in brain dysfunctional patients who achieve inpatient rehabilitation goals. *Arch Phys Med Rehabil* 1999;80: 77-84.

Objective: To evaluate whether improvements in specific cognitive and affective functions are associated with achieving inpatient rehabilitation goals after the acute onset of brain dysfunction.

supervised care facility.¹⁻³ Priority goals include increased mobility, self-care, bowel and bladder management, and communication.¹ Opinions, however, have differed about the impact of cognitive impairment on neurorehabilitation outcome. Diamond and colleagues⁴ reported that improvement on the Functional Independence Measure (FIM) from admission to discharge was unrelated to cognitive status, as measured by the Mini-Mental State Examination (MMSE), in a group of geriatric patients (38% of whom had CVAs). These patients evi-

Impairs functional recovery
measure of w/ of anosognosia

Impairs functional recovery

Jehkonen et al. (2000)

- Those with initial anosognosia had poorer functional outcomes

10 day - presence of
anosognosia @
(0 day = worse
outcome 1 yr
(after
EVEN THOUGH
none showed anos-
@ 3 mo mark
↓
early indicator

Anosognosia



[Explore this journal >](#)

Unawareness of deficits after right hemisphere stroke: double-dissociation of anosognosias

M. Jehkonen, J.-P. Ahonen, P. Dastidar, P. Laippala, J. Vilkki

First published: December 2000 [Full publication history](#)

DOI: 10.1034/j.1600-0404.2000.102006378.x [View/save citation](#)

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[View issue TOC](#)
Volume 102, Issue 6
December 2000
Pages 378-384

Abstract

Objectives- The aim was to study whether anosognosia for hemiparesis, anosognosia for neglect and general unawareness of illness double-dissociate, indicating that anosognosias are specific and independent impairments of awareness. On the other hand, anosognosias may be associated with one another and with general cognitive dysfunction, which decreases awareness of deficits. The persistence and predictive value of anosognosias was examined during a 1-year follow-up. **Patients and methods-**

Impairs functional recovery

Gialanella et al. (2005) • Rehabilitation progress not as strong when anosognosia is present

even w/ all
other symptoms
the same

Functional Recovery After Hemiplegia in Patients With Neglect The Rehabilitative Role of Anosognosia

Bernardo Gialanella, MD; Vittoria Monguzzi, MD; Raffaele Santoro, MD; Silvana Rocchi, PD

Background and Purpose—The aim of this study was to verify whether the presence of anosognosia (A) affects the rehabilitative prognosis of hemiplegic subjects with neglect (N).

Methods—This study was carried out on 30 patients with left hemiplegia: 15 patients had neglect (group N) and 15 had neglect and anosognosia (group N+A). Mean age was 68.2 ± 6.3 in group N (9 men and 6 women) and 72.1 ± 6.4 in group N+A (7 men and 8 women). The average interval from onset of stroke to admission for rehabilitation was 23 and 23.6 days, respectively, in group N and in group N+A. Patients were assessed through the Mesulam test, Bisiach test, Wechsler Adult Intelligence Scale, Fugl-Meyer scale, Functional Independence Measure (FIM), and Rankin scale.

Results—Before rehabilitation, cognitive FIM scores of patients of group N were significantly higher than those of group N+A ($P=0.001$), whereas motor FIM scores and total FIM scores did not differ between the 2 groups. After rehabilitation, cognitive FIM scores ($P=0.000$) and even motor ($P=0.009$) and total FIM scores ($P=0.000$) were statistically higher in group N than in group N+A. Effectiveness ($P=0.005$) and efficiency ($P=0.012$) in the motor FIM scores of group N were significantly greater than those of group N+A. Disability was lower in group N ($P=0.040$).

Conclusions—Our study shows that the presence of anosognosia worsens the rehabilitation prognosis in hemiplegic subjects who also have neglect. (*Stroke*. 2005;36:2687-2690.)

Key Words: cerebrovascular disorders ■ neglect ■ stroke ■ rehabilitation

Interventions?

Cheng & Man (2006)

“Awareness Intervention Program” (AIP)

AIP group showed improved self-awareness but was not associated with improved functional outcome

Can we target anos. to improve recovery?
— Brain injury edu.; show PP what dysf. they were dealing w/
Decreased ↓ anos. faster
BUT
Didn't improve recovery

Anosognosia ↳ Anos. is not cause, but indicator

Original

Management of impaired self-awareness in persons with traumatic brain injury

S. K. W. Cheng & D. W. K. Man 

Pages 621-628 | Received 05 Jul 2005, Accepted 02 Mar 2006, Published online: 03 Jul 2009

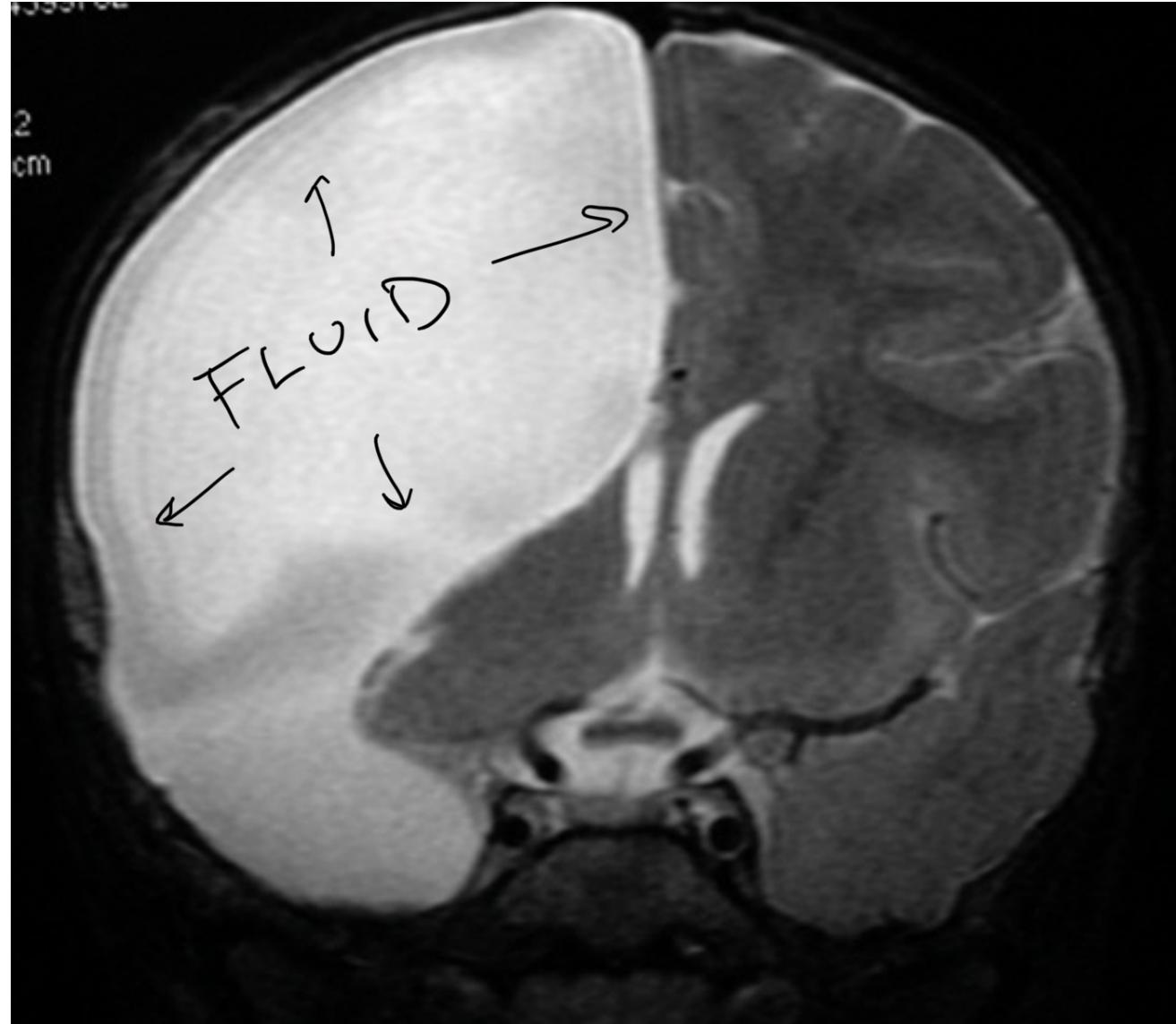
 Download citation  <https://doi.org/10.1080/02699050600677196>



 Full Article  Figures & data  References  Citations  Metrics  Reprints & Permissions  Get access

Abstract

Aim: To develop and evaluate a systematic intervention programme for the management of impaired self-awareness (ISD) in people with traumatic brain injury.



Effect of age

see
inside
csf

↓ age = ↑ outcome
neuron loss

https://www.youtube.com/watch?v=f2fCY_M7Vms

give me
an expression

"I only have
half a brain"
↓
living
relatively
normal life
If happens
very
young;
much more flexibility
torewire
12

Removal of an infant monkey's primary motor cortex results in a less dramatic impairment than in an adult monkey (Kennard, 1948)

Brain injury resulting in aphasia before age 1 is associated with the best recovery; recovery diminishes as age increases (Woods & Carey, 1979)

BUT:

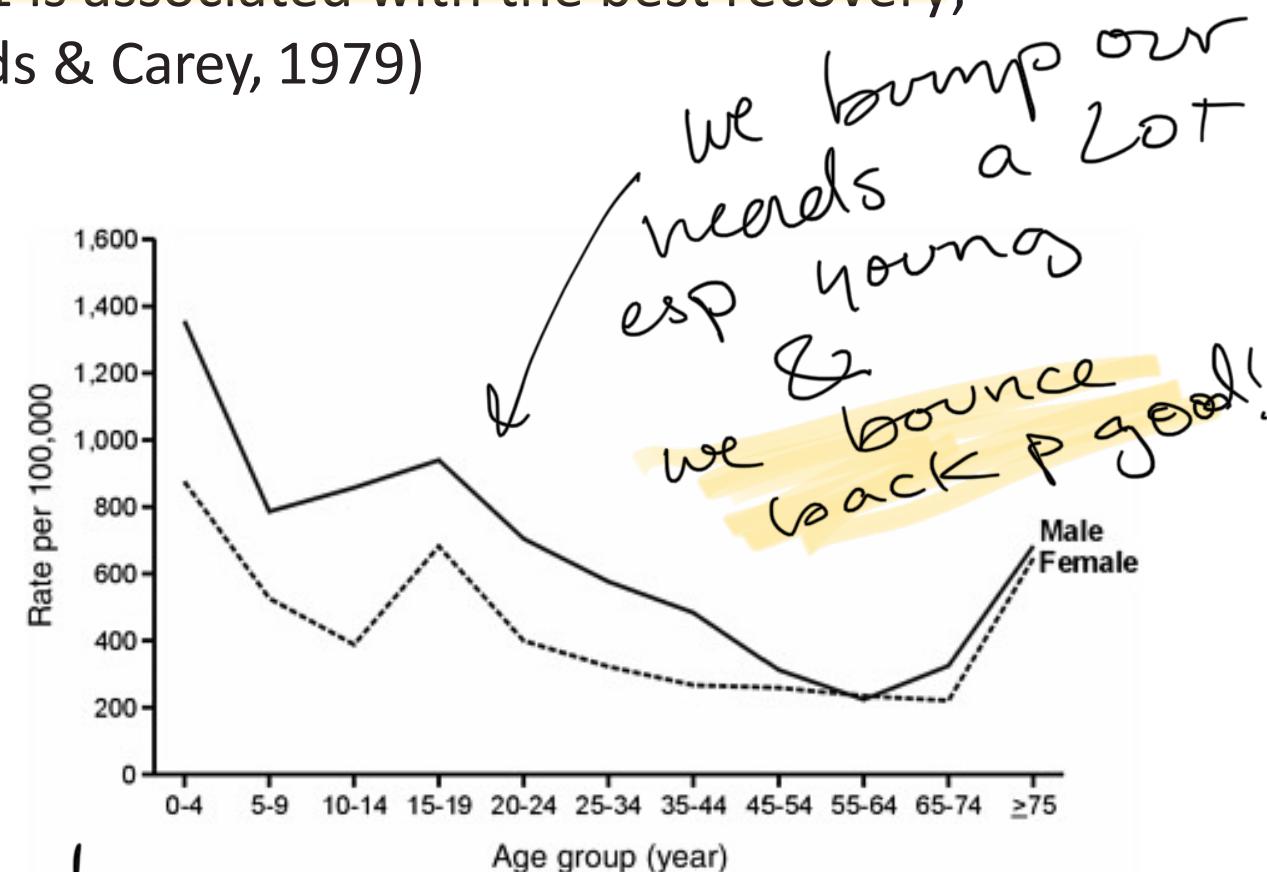
Prefrontal lesions before 16 months may be associated with an inability to learn social and moral rules later in life (Anderson *et al.*, 1999)

Prefrontal damage
exception: younger brain recovers worse

Effect of age



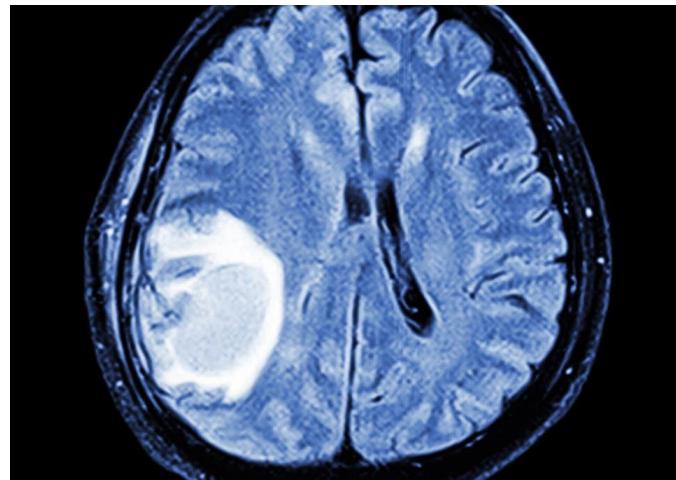
Dusty requested
Ted Cruz dog



Average annual TBI-related rates for emergency department visits, hospitalizations, and deaths, by age group and sex, U.S., 1995–2001.
Langlois *et al.* 2004

There is a direct relationship between the size of a brain lesion and the extent of recovery:

Larger lesions generally result in more functional impairments.



unilateral
lesion >
bilateral

In addition, patients with bilateral lesions show less recovery than patients with unilateral lesions (e.g. hippocampal lesions; Scoville & Milner, 1957; Zola-Morgan, Squire & Amaral, 1986)

Effect of lesion size and location

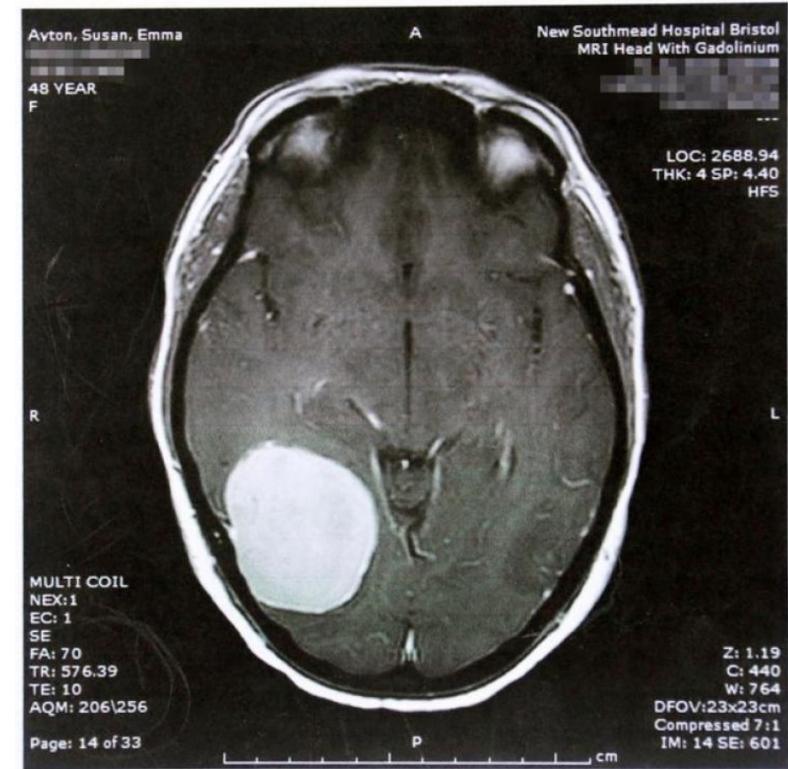
Our redundant
symmetrical
structures

Massive tumour, little impairment

vs.

Sudden stroke with associated damage (of a comparable size), functionally devastated

Sudden / is far worse
rapid
than slow



Chronic vs. acute dysfunction

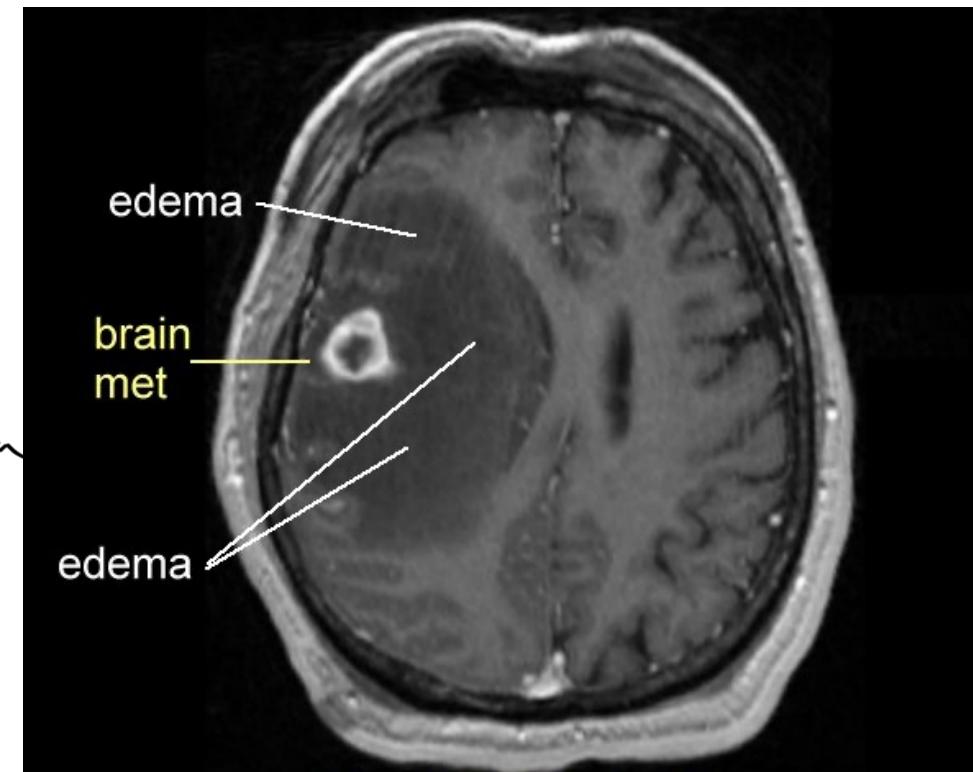
triggered by potassium signalling loop
causes TNF&interstitial fluid Edema

Swelling (edema) after stroke may mask or distort functions in essentially intact regions of the brain

Dissipation of these effects may account for substantial early recovery

doesn't always go away

~2 wk after stroke:
when swelling ↓, healthy neurons able to function again
"bounce back" of function



Secondary effects

Loss of inputs = Diaschisis

A brain lesion can cause dysfunction in an area remote to it because of its strong connections with that area.

Thus, recovery can occur because of a spontaneous reduction of diaschisis (i.e. reduction in metabolic depression in cerebral areas remote to the lesion).

Cerebellum becomes more metabolically active: find "next best input" when input lost → can make full recovery

Toxic restitution

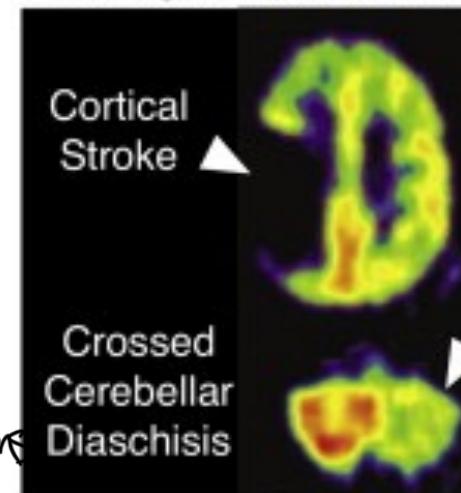
Secondary effects

(altered new)

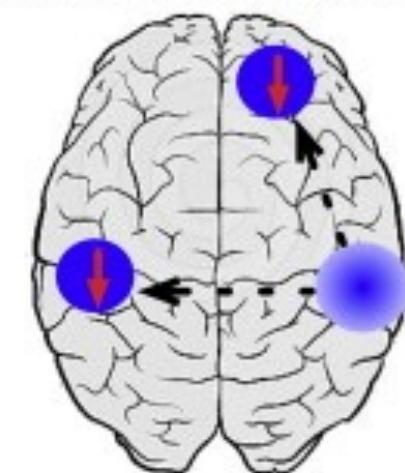
connections

made, neurons not lost forever?
google ↗

A Diaschisis in distal brain regions (cerebellum)



B Diaschisis in functionally connected cortical areas



Environmental factors may affect the amount of functional recovery following brain injury.

e.g. *occlude good hand*

Animals models, enriched environments, **constraint-induced movement therapy**

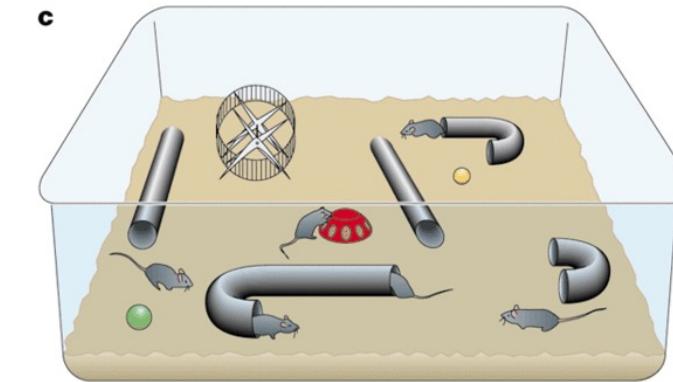
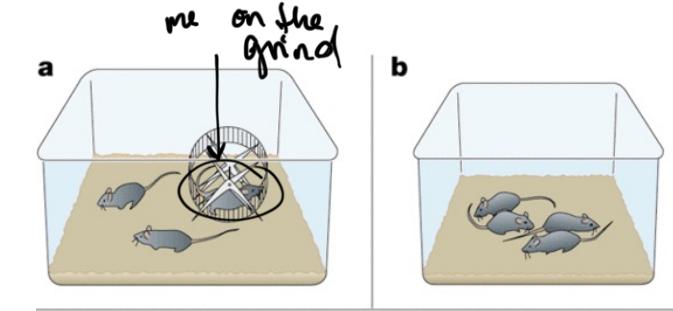
↪ "rest" can ↓ recovery

Monkeys with focal strokes to the motor cortex that were retrained in skilled hand use

Environmental factors

Social
Community
Physical therapy

use it or lose it



Compensatory strategies are a mainstay of rehabilitation of patients with brain dysfunction

So recovery may not represent true *restitution* of function, but *substitution* of a new behavior Often, neurons die → other neurons substitute

So, a patient learns strategies to adapt to his or her new motor, sensory or cognitive impairments

✓ SUPER good
at recovery

e.g. Beam-walking of rats with lesions to sensorimotor cortex → completely diff run to before:
substitution

It is not a failing if you find a new way to succeed at doing an old thing! e.g. Carrying a notebook, walking with an aid, using notation software, etc.

↑
Off

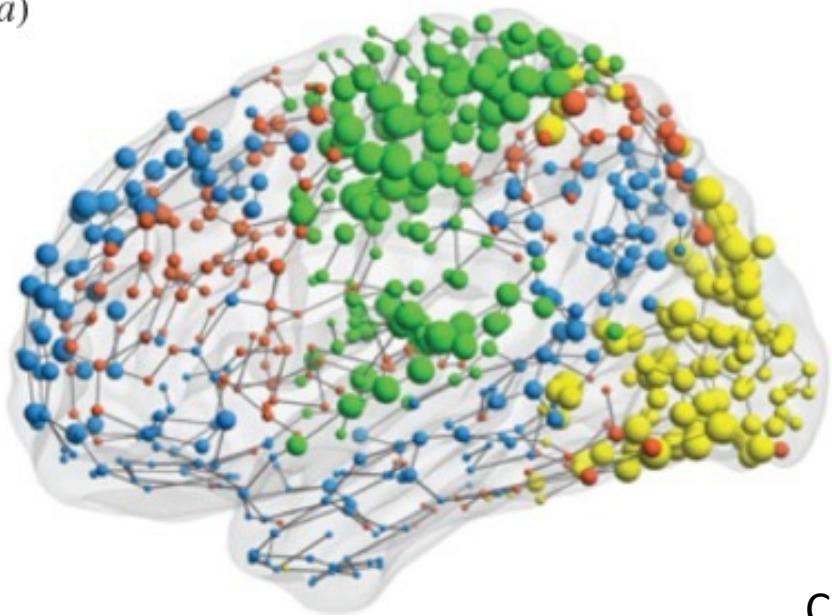
Compensatory strategies



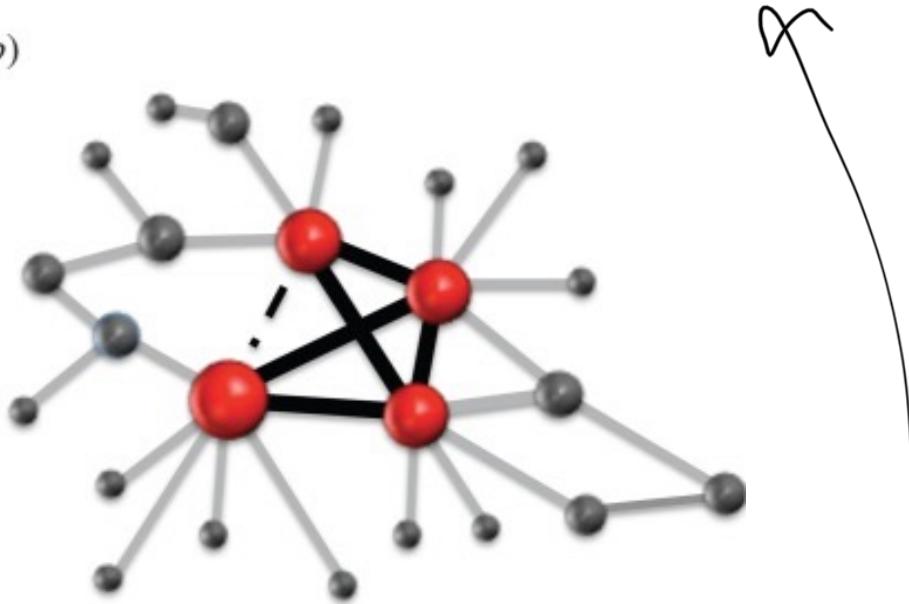
"rewired"

New wiring for old jobs

(a)



(b)



Crossley et al. 2013, Vértes et al. 2014



Reminder: your brain has ~80 billion neurons,
and more connections between them than is
strictly necessary

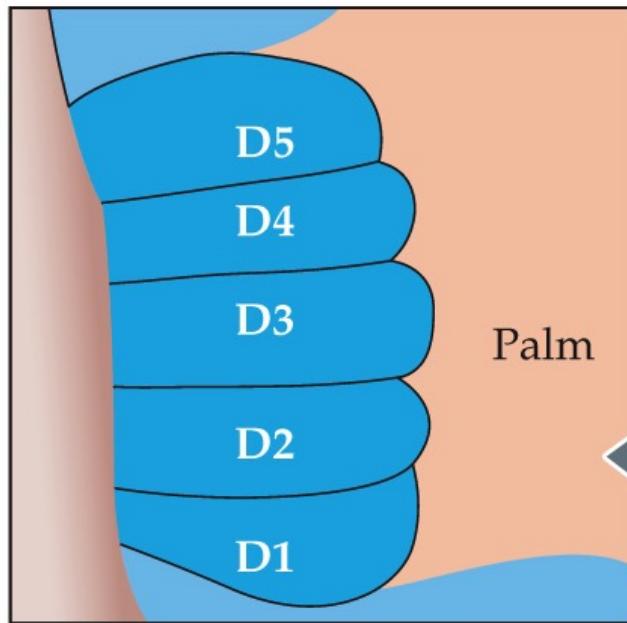
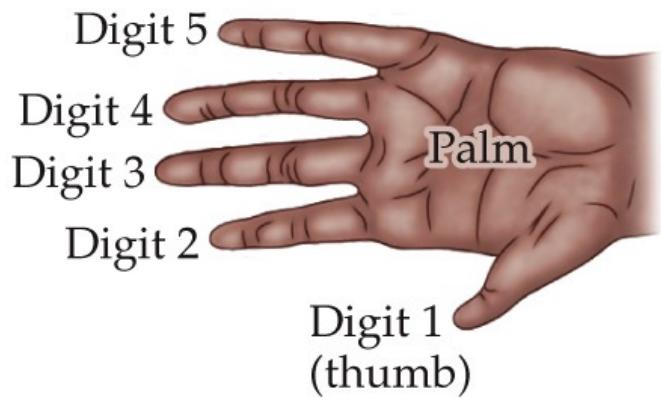
"Rich club" vs. "Poor club" of neurons (Buzsaki 2022)

Neuroplasticity

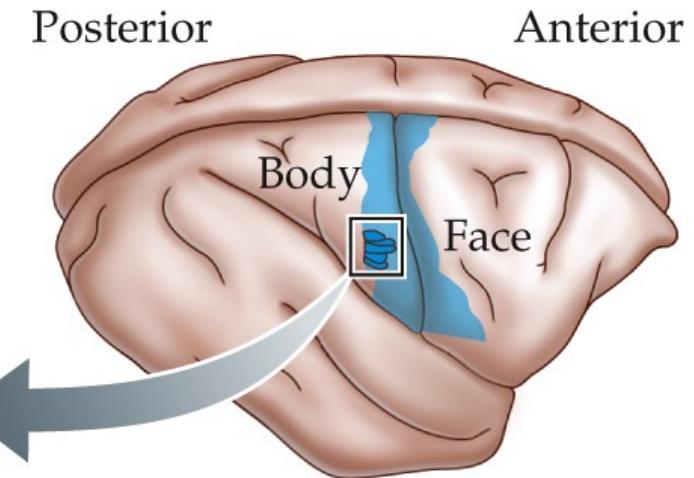
some neurons are used way more / connected
way more than others

Cortical representation

(A) Representation of the left hand in primary somatosensory cortex in right hemisphere of monkey brain



Details of cortical map
(D5 = digit 5, etc.)



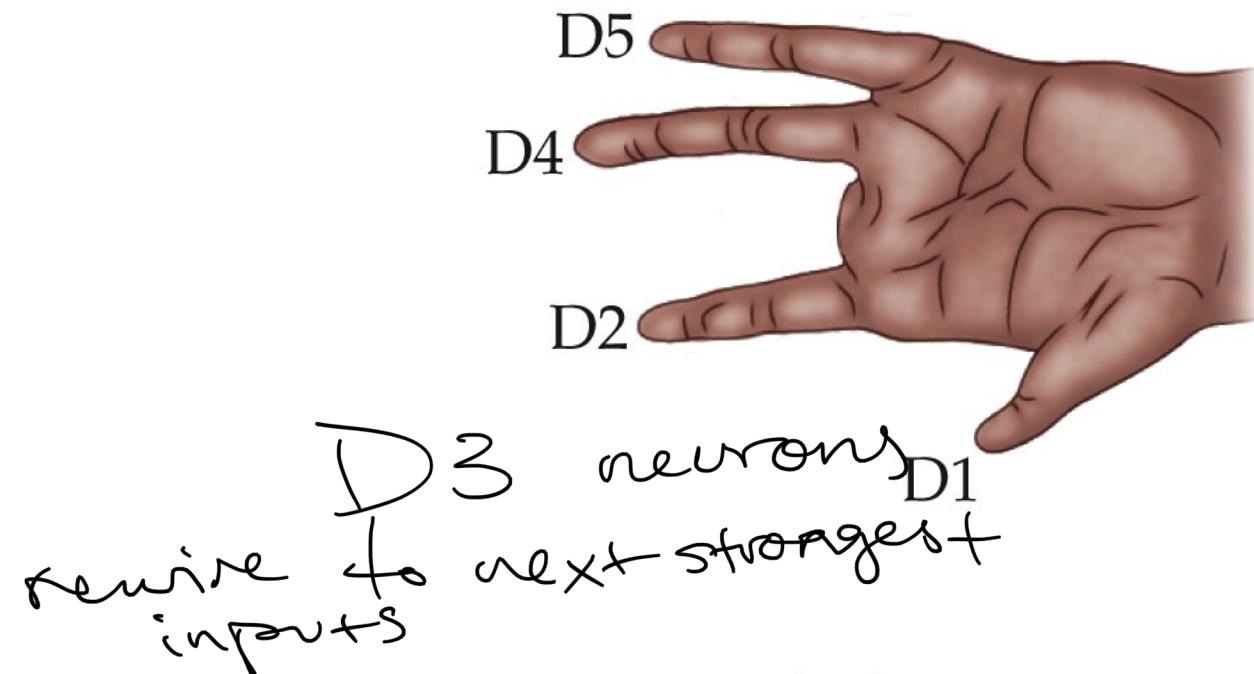
BEHAVIORAL NEUROSCIENCE 8e, Figure 8.17 (Part 1)
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Functional areas are flexible
Representation is based on use

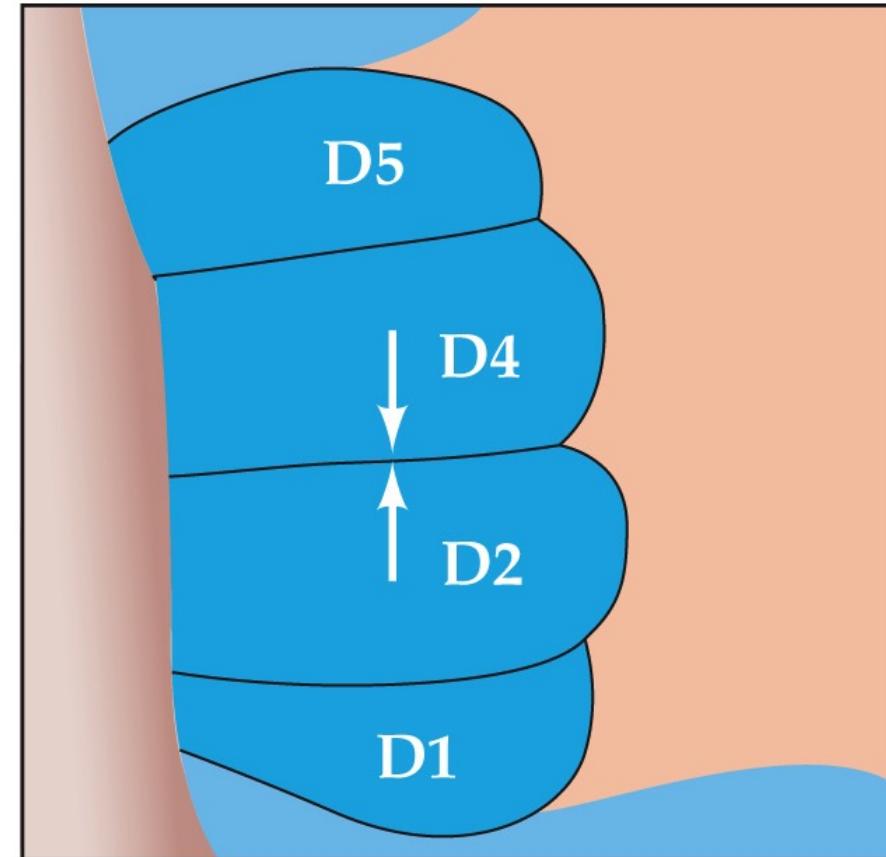
Neuroplasticity due to damage

“The brain abhors an unused neuron.”

—Your teacher, last night when he was
prepping these slides



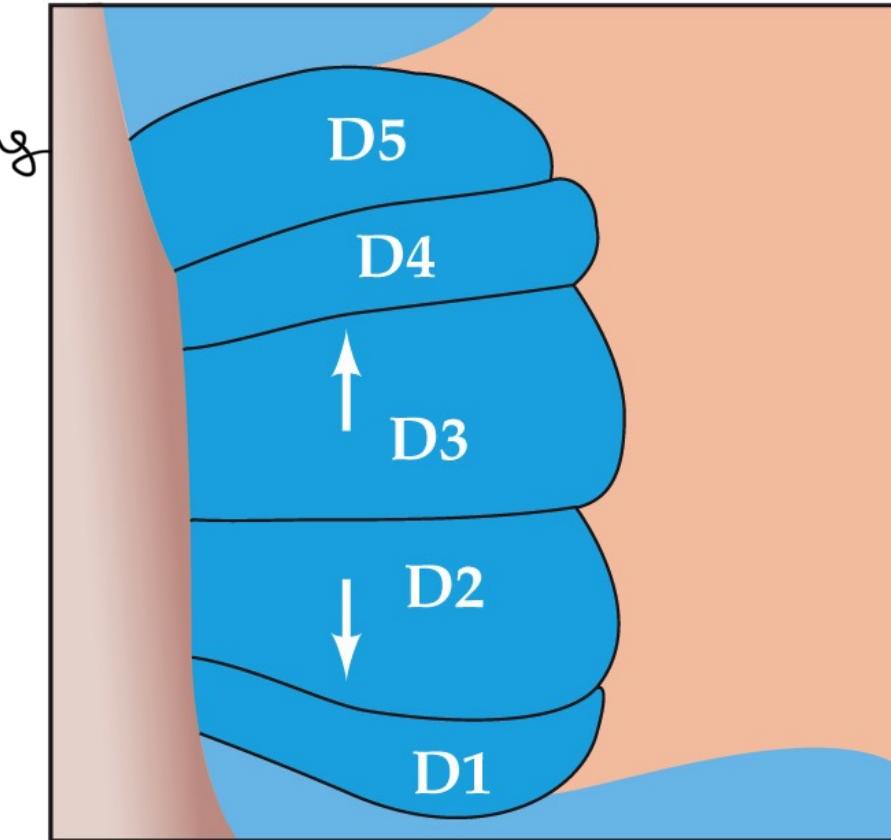
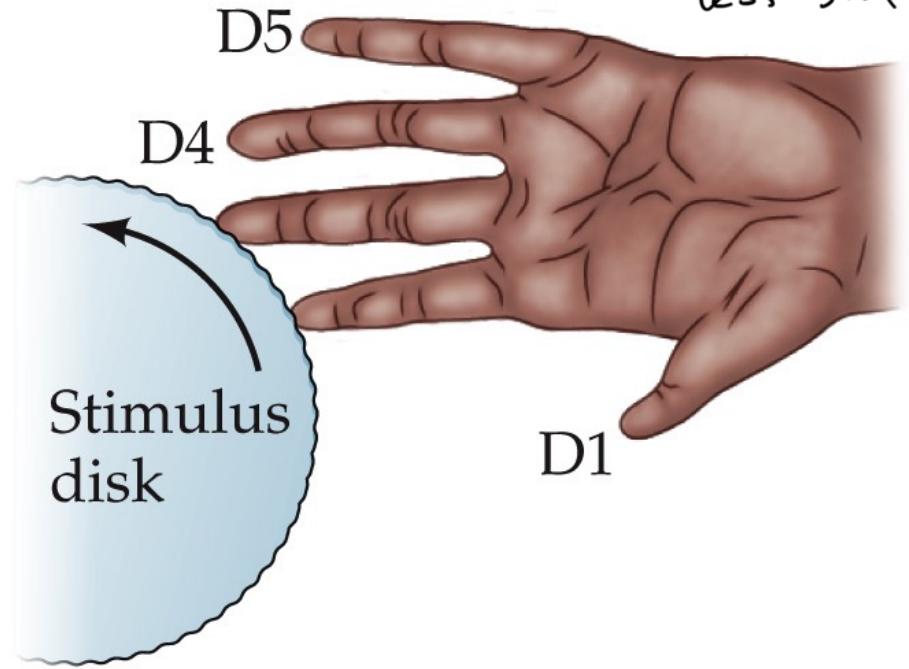
BEHAVIORAL NEUROSCIENCE 8e, Figure 8.17 (Part 2)
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Neuroplasticity 22

Neuroplasticity due to use

Implications for your life?



BEHAVIORAL NEUROSCIENCE 8e, Figure 8.17 (Part 3)
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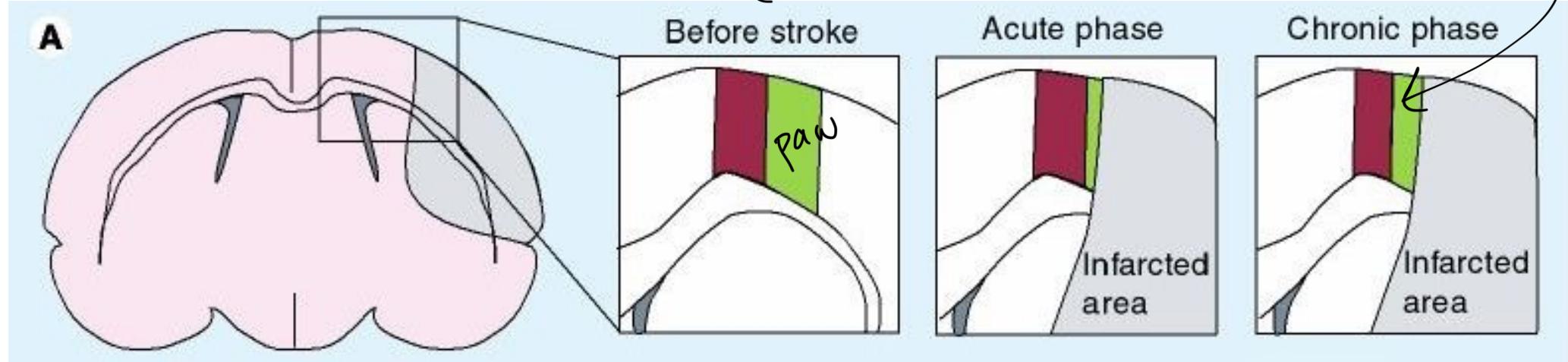
The more you use neurons, the more the region expands

↑
neurons shift roles

↑
neurons that are poor are ones that change

Neuroplasticity due to cortical damage (rat)

paw important!
rewire for it



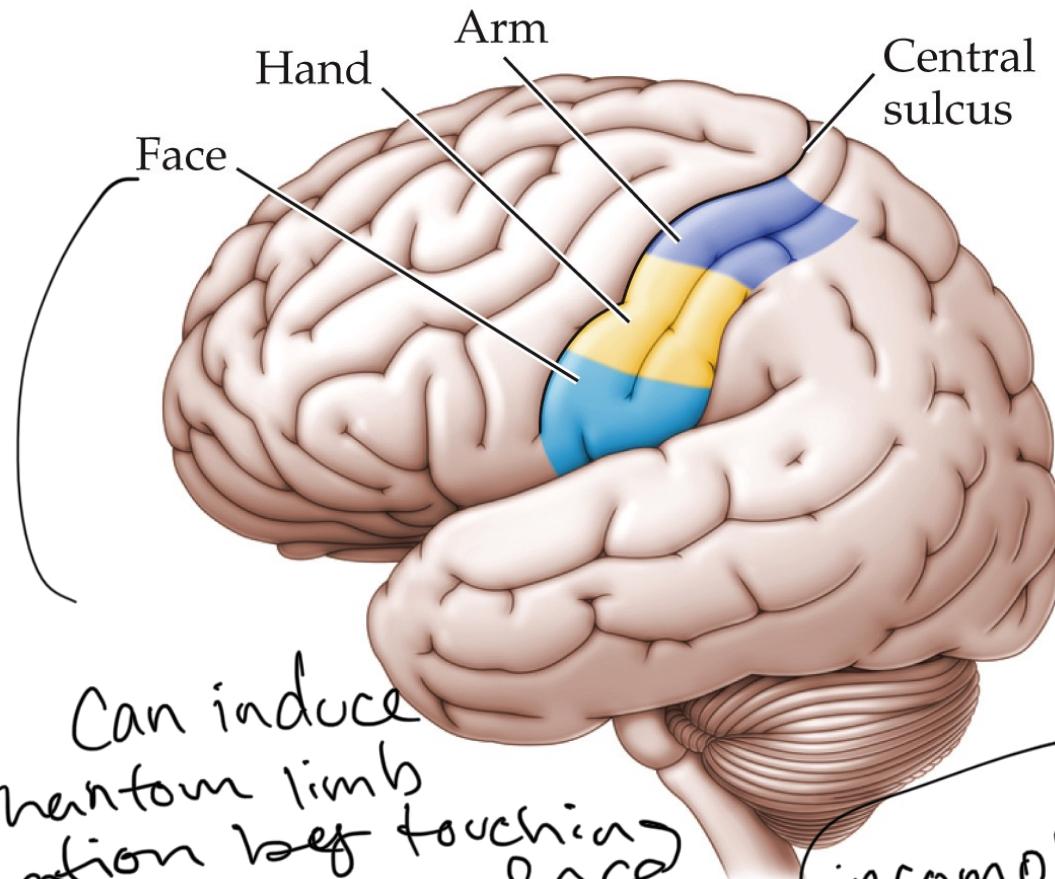
Constraint-induced therapy

Neuroplasticity



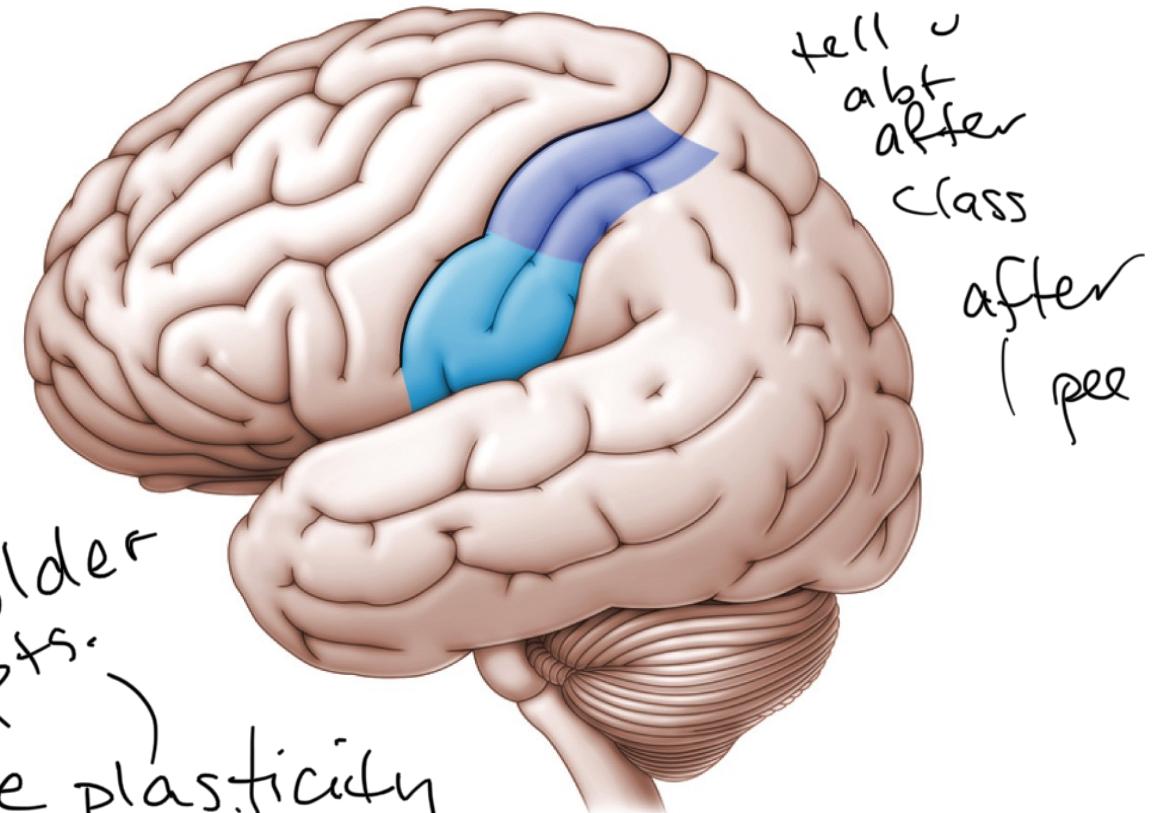
Phantom limbs: a case of large-scale plasticity

(A) Typical somatosensory cortex



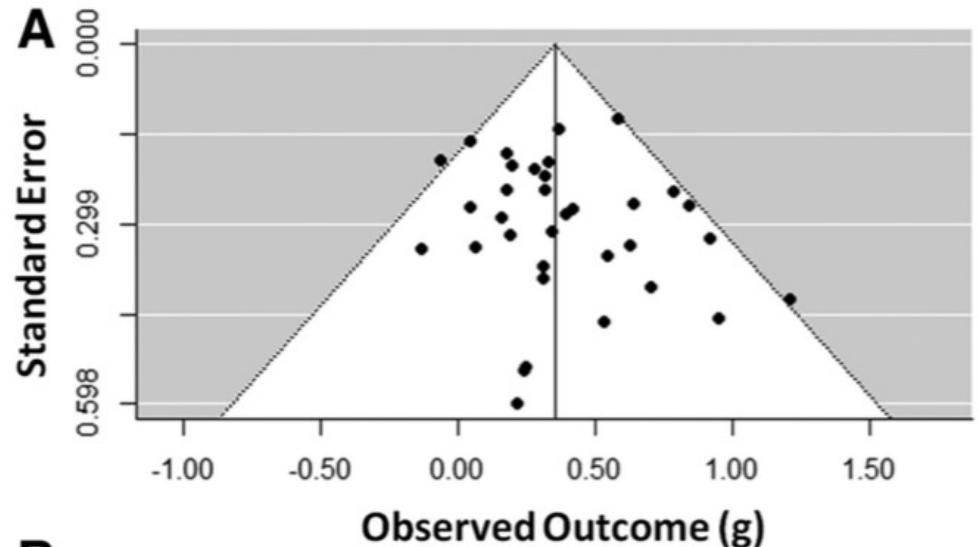
BEHAVIORAL NEUROSCIENCE 8e, Figure 8.18 (Part 1)
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(B) Somatosensory cortex reorganized after loss of hand



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Neuroplasticity 25



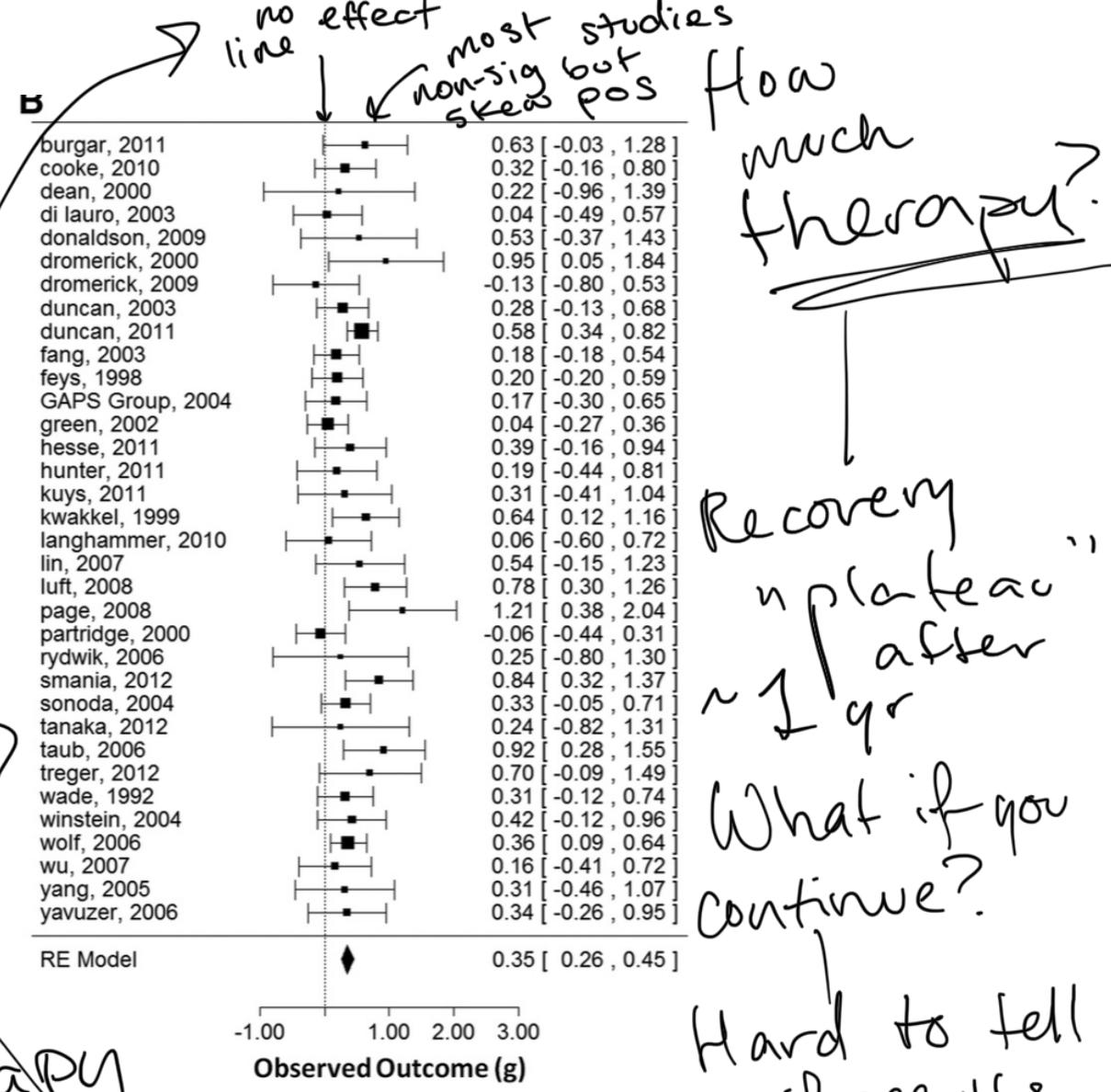
B

Intro to meta-analysis methods

Lohse meta-analysis: to understand effect of therapy, while controlling for time

Therapy dose

more therapy
is actually better!



from Lohse et al. (2014)

26