



UNIVERSITÉ  
DE GENÈVE



Neurobiology of  
Concepts Expression  
[noce-lab.github.io](https://noce-lab.github.io)



# Testing Cognitive Theories with Neuroimaging: Lessons from Primary Progressive Aphasia

**Dr. Valentina Borghesani**

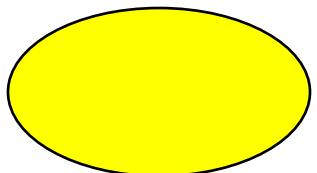
*BSc in Psychology, MSc Neuropsychology, PhD Cognitive Neuroimaging*

Assistant Professor at UNIGE/FPSE | NCCR Evolving Language

/'lɛmən/



l - e - m - o - n



Lemons are native to Asia





Channels

Labels

word forms

/'lɛmən/

spoken word

L-shaped wrist mov.

signed word

LEMON

written word

Concept

taste,  
smell

emotion  
associations

unlabeled concepts

e.g., *learning*

ungrounded labels

e.g., *patients*

n° labels ≠ n° concepts

e.g., *bilinguals*



3

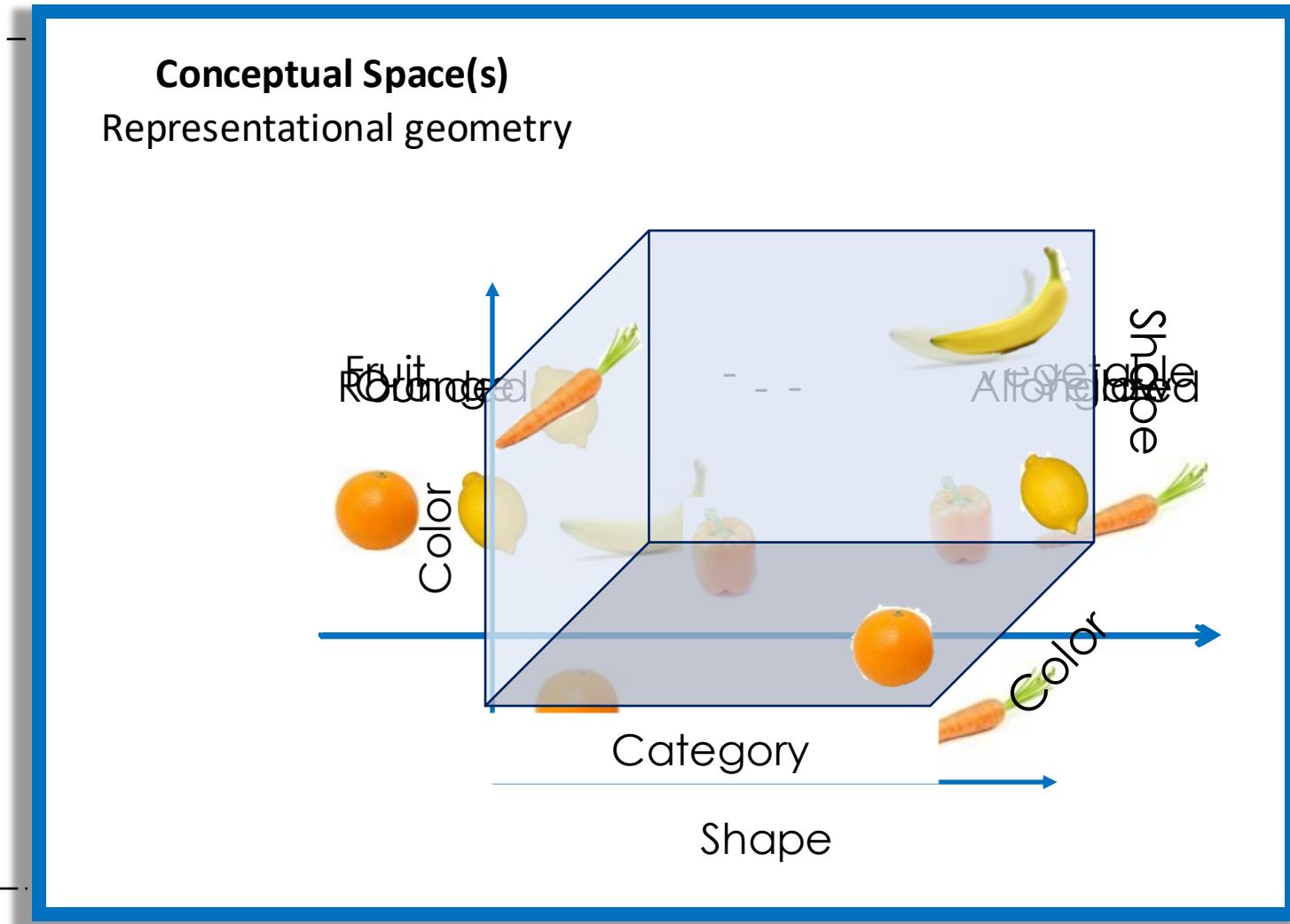
Lemons are native to Asia

Concept Features

word meaning



	/'ləmən/ spoken word
L-shaped wrist mov.	signed word
LEMON	written word



---

Channels

---

Labels

---

word forms

Concept Features

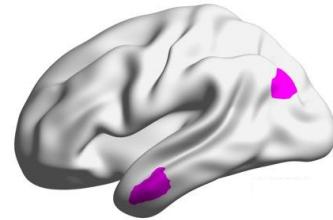
---

word meaning

Goal: to understand

**localization** & **dynamics** of neural signatures **causally** linked to semantic representations.

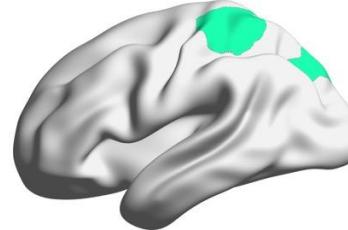
e.g., **BOLD signal** with **fMRI**



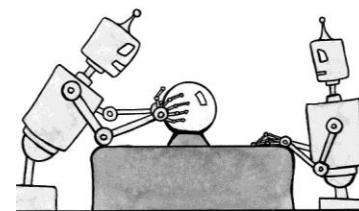
measure / describe



e.g., **spectral power** with **MEG**



predict



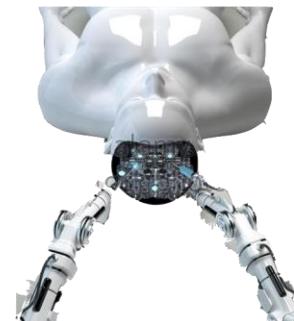
imitate / outperform



e.g., **volume** with **MRI**



**fix / change**



## Introduction to Primary Progressive Aphasia

The synergy of **clinical & cognitive** neuroscience

### Q1: How are semantic representations encoded in the brain?

In **distributed yet specialized** cortical areas

**Rapidly** retrieved via symbols

As required by the **task** and afforded by the **stimuli**

With the **anterior temporal lobe** as a critical hub

**Borghesani et al.** 2016 *NeuroImage*

**Borghesani et al.** 2018 *Journal of Cognitive Neuroscience*

**Borghesani et al.** 2018 *Cortex*

**Borghesani et al.** 2019 *Journal of Cognitive Neuroscience*

**Borghesani et al.** 2019 *Cortex*

**Vonk, Borghesani et al.** 2019 *Aphasiology*

**Borghesani et al.** 2020 *NeuroImage:Clinical*

### Q2: What is the relation with other cognitive functions, namely language?

Critical behavioral **dissociations**

**Preserved percepts, eroded concepts**

Can be (partially) **compensated** by spared functions

Can be (partially) recovered from **language use alone**

**Borghesani & Gorno-Tempini** 2022 *HCN*

**Borghesani et al.** 2020 *Brain*

**Lukic, Borghesani, et al.** 2021 *Cortex*

**Borghesani et al.** 2021 *eLife*

**Younes, Borghesani et al.** 2022 *Brain*

**Lukic, Licata,.....& Borghesani** 2022 *Fr.Psy*

**Borghesani et al.** 2023 *Scientific Data*



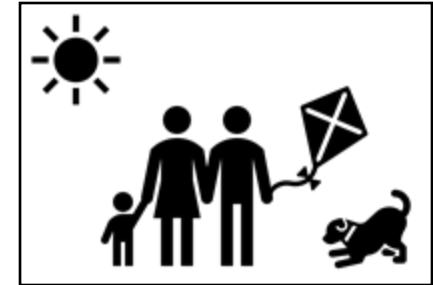
Why are you here today?

*Sometimes I have difficulty **finding** the words.*

### Primary Progressive Aphasia (or PPA)



Can you please look at this picture and tell me what you see using full sentences?



**P1.** *I s- see a family of three. [pause] Th- the father has a [pause] kite. They have a dog.*

**False starts and pauses, but correct grammar and content words.**

**P2.** *Sunny day. Baby. Mother. Fa-d-er. He...kite. Also, a dog.*

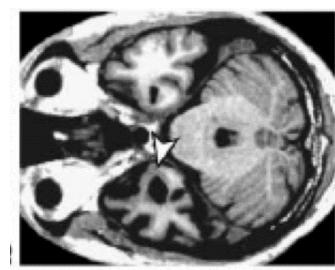
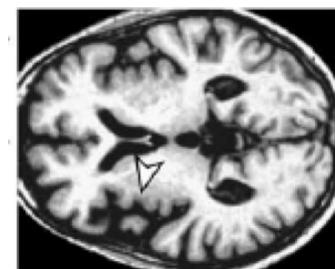
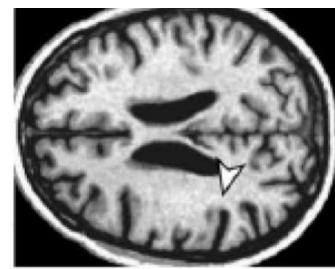
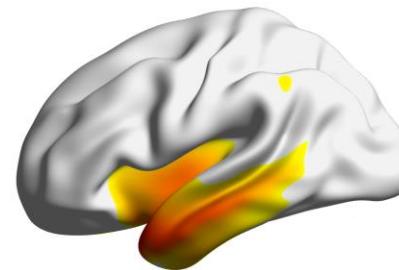
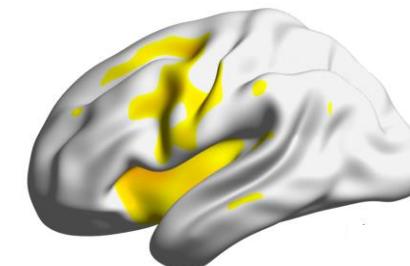
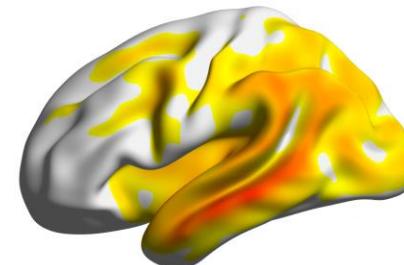
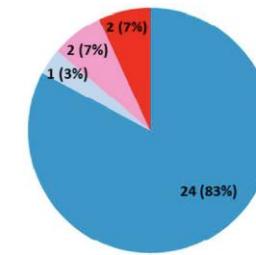
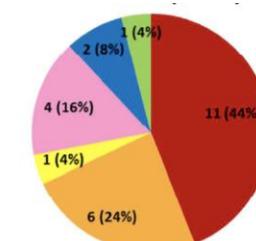
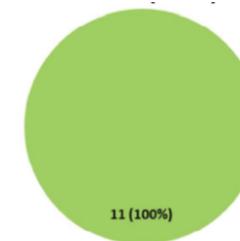
**Grammatically incorrect and sounds distortions, but right content words.**

**P3.** *There are three people and one of them is holding an object. There is also an animal.*

**Good fluency and grammar, but inadequate content words.**

**Language Profile**

**P1.** phonology and/or short term verbal memory deficits  
= logopenic variant (lvPPA)

**Anatomical Substrate****Pathological Correlates**

**P3.** semantic memory deficits  
= semantic variant (svPPA)

Legend:

- CBD
- PSP
- tau 4R uncl
- PiD
- TDP-A
- AD + CBD
- TDP-C
- TDP-B + tau
- PiD
- GGT

### Computational Models:

Dual-route cascaded model - *Coltheart et al., 1993*

Triangle model - *Plaut et al., 1996*

### Neuropsychology :

Phonological vs. surface dyslexia - *Marshall & Newcombe, 1973*

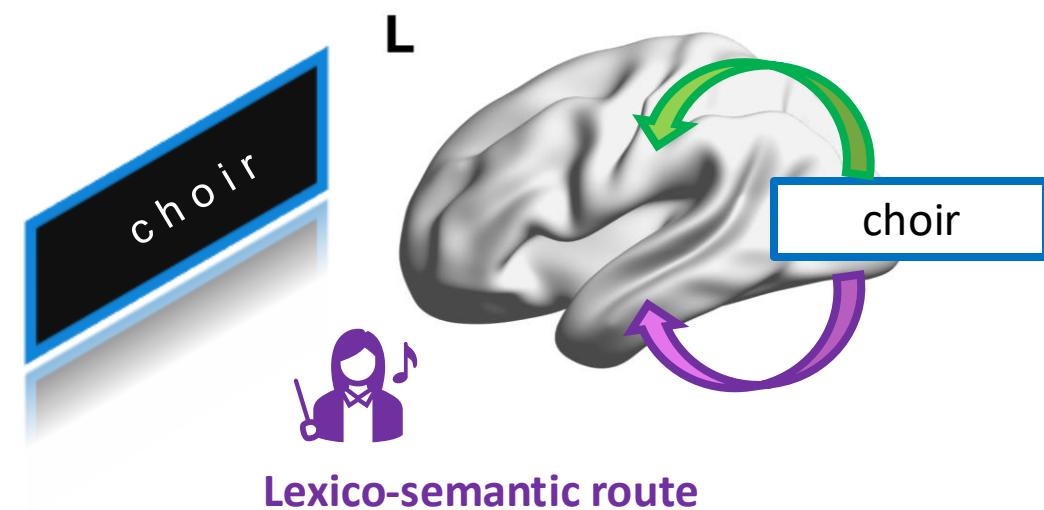
### Neuroimaging studies:

Parietal structures activates for pseudowords reading vs.

Temporal structures activates for irregular words reading -

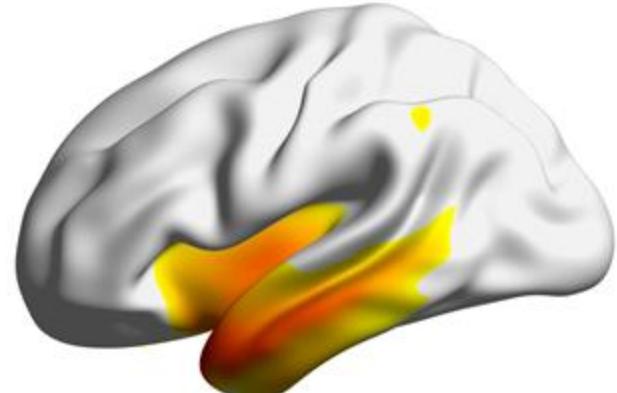
*Mechelli et al., 2003; Taylor et al., 2013*

### Sub-lexical/phonological route



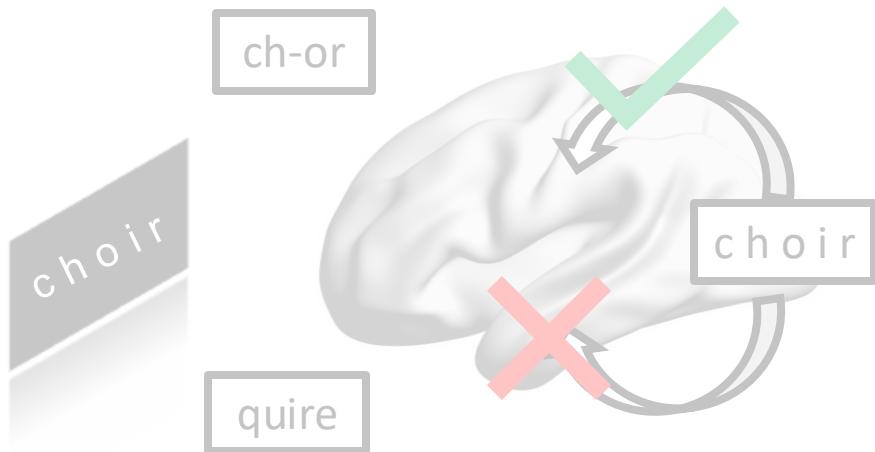
### Lexico-semantic route

**Hypothesis:** given the damage to ventral path, svPPA patients can only rely on the dorsal path & phonological information

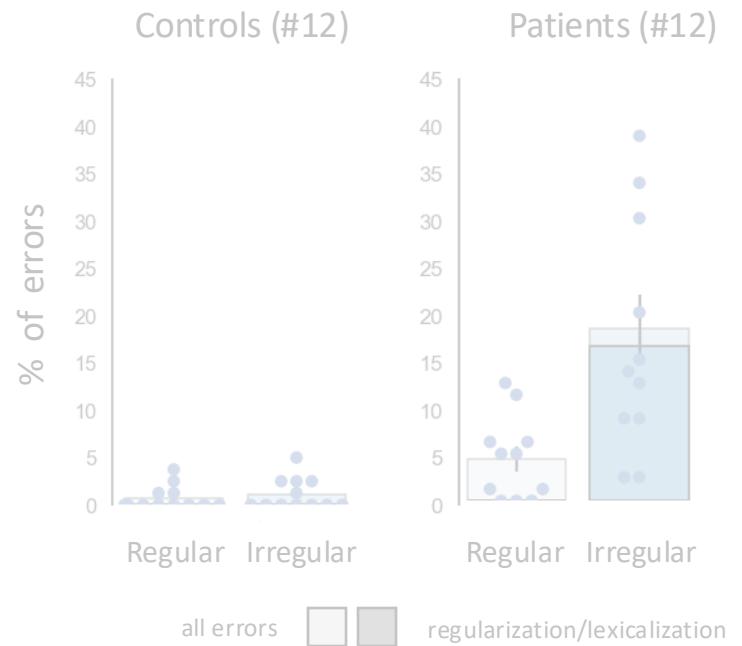




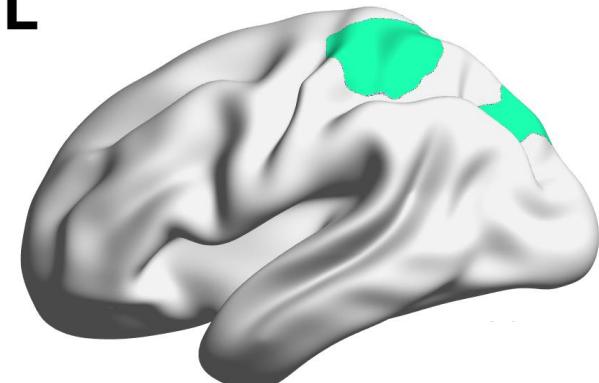
Sub-lexical/phonological route



Lexico-semantic route



L



**Results:** starting  $\sim 400$  ms post word onset, patients recruit dorsal areas to read irregular words.

**Conclusion:** dorsal structures can partially compensate ventral damage.

### Theoretical Models:

Hub-and-spoke model - *Patterson et al. 2007*

Convergence zones - *Damasio et al., 2004*

Dynamic multilevel reactivation framework - *Reilly et al., 2016*

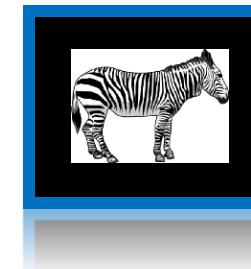
### Neuroimaging studies:

Distributed semantic representations during

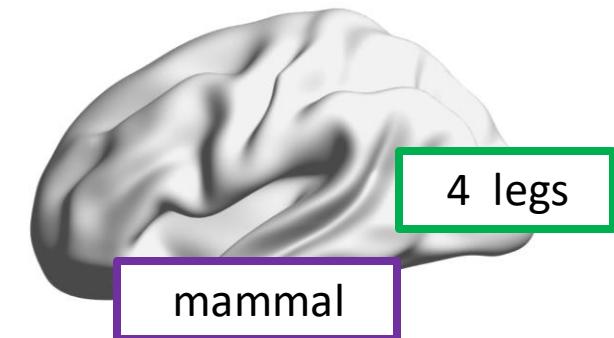
*Movie watching* - *Huth et al., 2012*

*Story listening* - *De Heer et al., 2017*

*Single words processing* - *Fernandino et al., 2016*

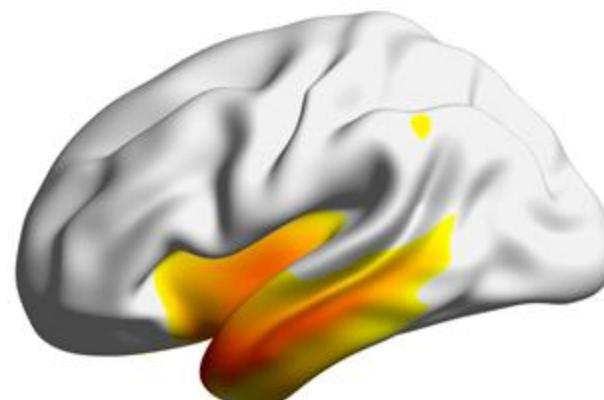


Perceptual information

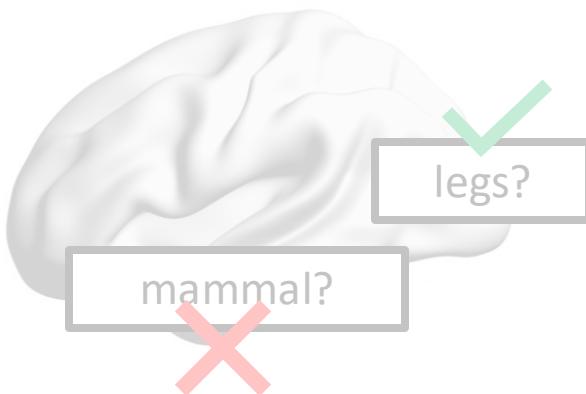


mammal

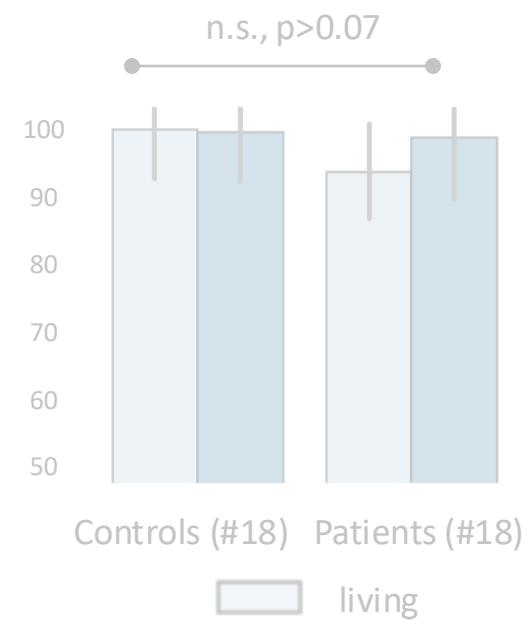
Conceptual information



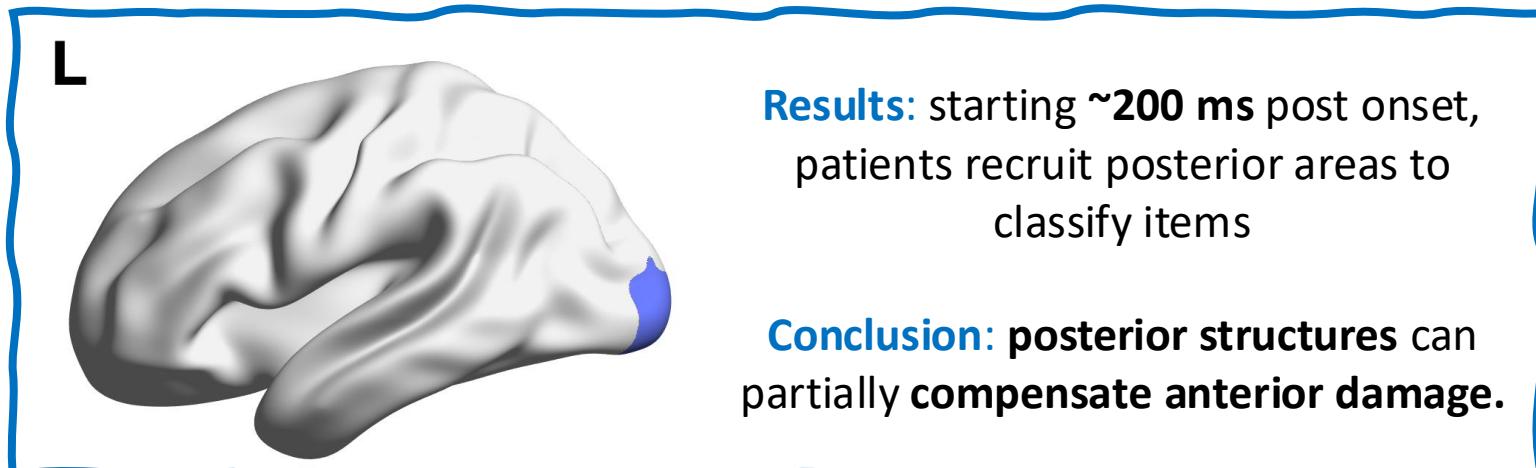
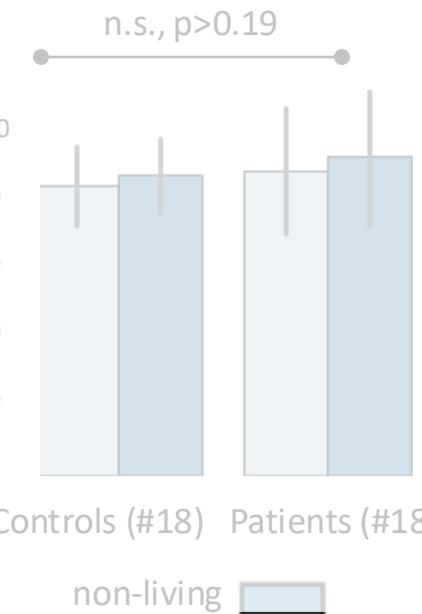
**Hypothesis:** given the damage to the anterior regions, svPPA patients can only rely on the posterior regions & perceptual information



Accuracy [% correct]:



Reaction Time [ms]:

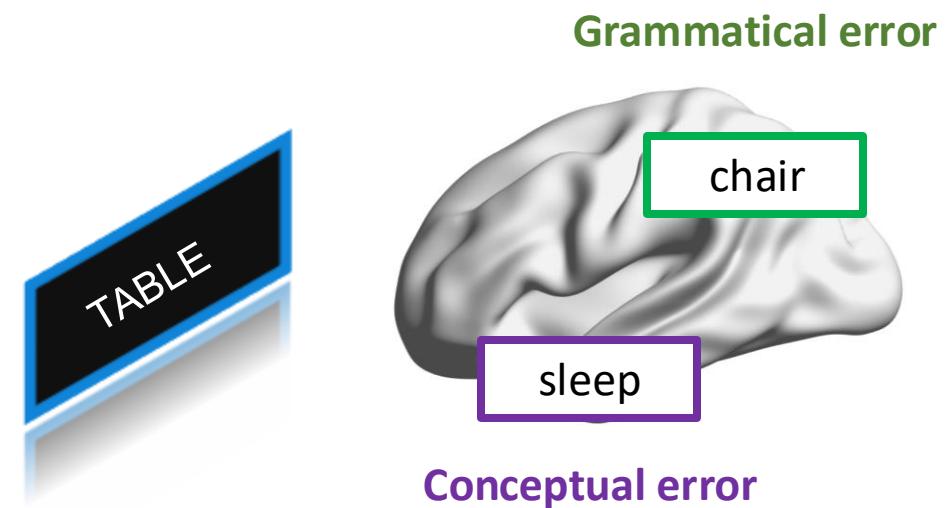


**Reviews:**

Neuropsychology - Mätzig et al. 2009

Neuroimaging - Vigliocco et al. 2011

Neurodegenerative patients - Lukic et al., 2021



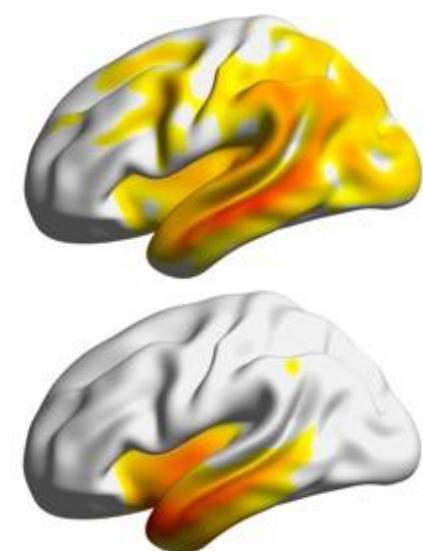
**Hypotheses:**

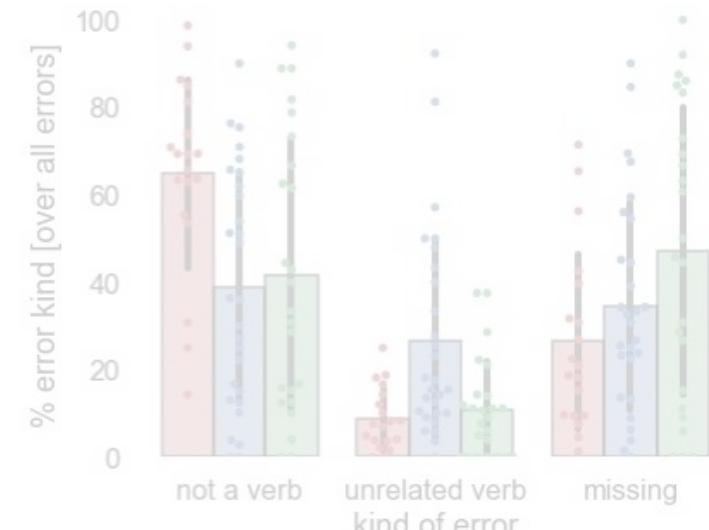
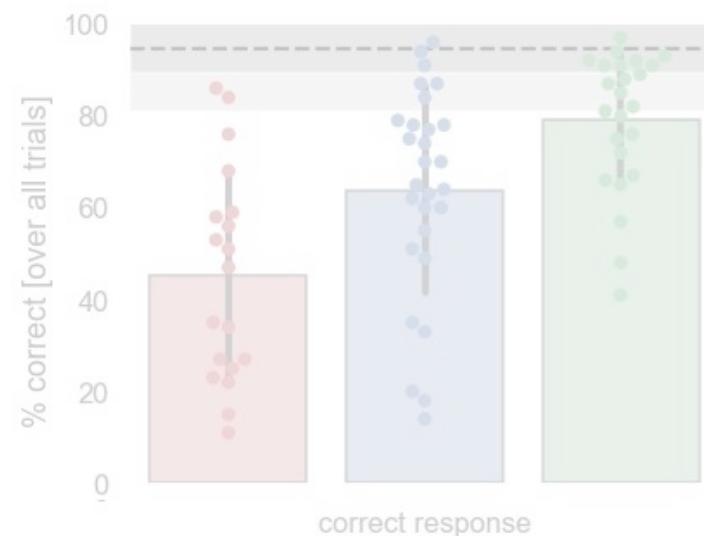
given the damage to the posterior temporal regions,

**IvPPA patients** will show **verb-related deficits**

given the damage to the anterior temporal regions,

**svPPA patients** will show **conceptual deficits**



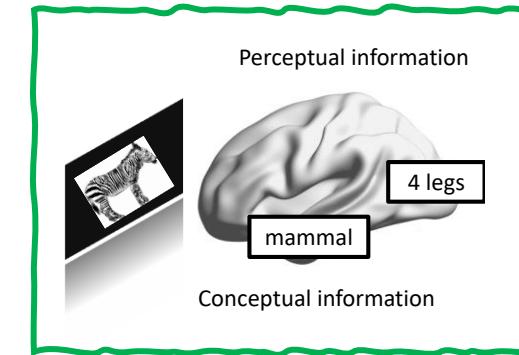
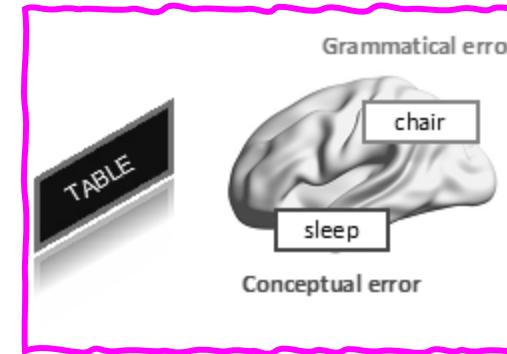
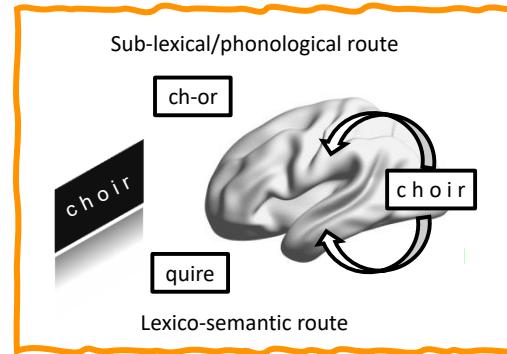
IvPPA  
(n=19)nfPPA  
(n=24)svPPA  
(n=28)

**Results:** IvPPA produce many semantically **related nouns**, while svPPA semantically **unrelated verbs**

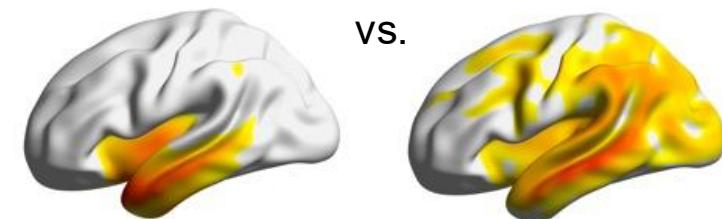
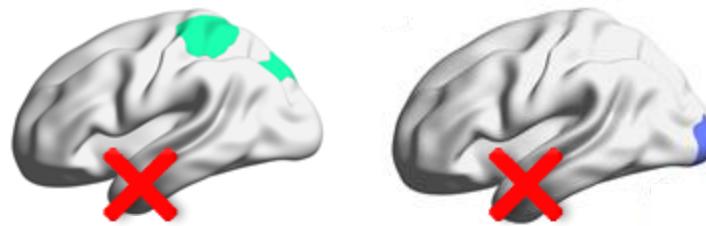
**Conclusion:** temporo-parietal regions & lexical processing, anterior temporal lobe & semantic processes.



Theories on the neurocognitive correlates of language (e.g., reading, syntax) and its interplay with semantic can be empirically tested combining neuroimaging & neuropsychology



Damage to the anterior temporal lobe (and following semantic loss) can be compensated relying on dorsal (phonological) or posterior (perceptual) processes, and the analysis of error patterns can improve syndromic diagnosis





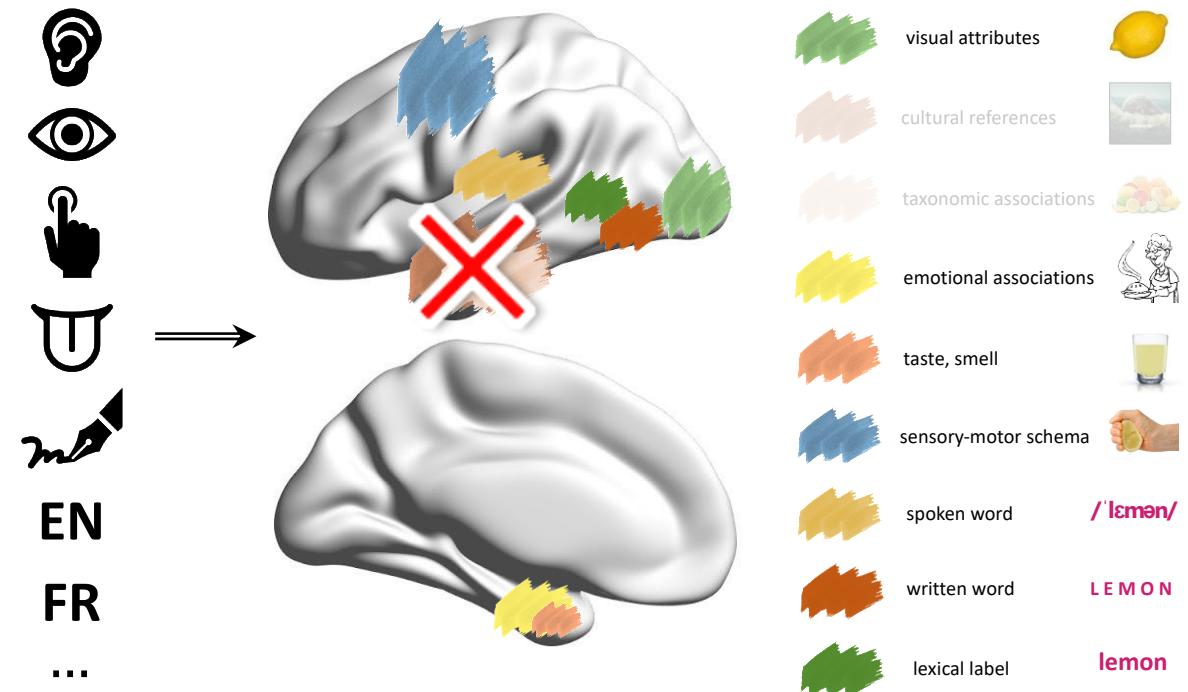
**Neurodiversity:** can we model how **language** (i.e., input/output system) interfaces with **semantic** (i.e., multidimensional concepts) **across modalities, tasks, languages, level of expertise, lifespan, etc.. ?**



Can we improve the description of the **neural topography & dynamics** of the interplay between language & semantic?



**Translational neuroscience:** can alterations in language & semantic can be used as **neuro-cognitive biomarkers** for **neuro-psychiatric conditions**? Can we leverage what we learn for **interventions**?



Thanks to colleagues, funding agencies, and to you for your attention!



UNIVERSITÉ  
FRANCO  
ITALIENNE

UNIVERSITÀ  
ITALO  
FRANCESE



Fondazione Zegna



Centre intégré  
universitaire de santé  
et de services sociaux  
du Centre-Sud-  
de-l'Île-de-Montréal

Québec

Centre de recherche  
**iugm**  
Institut universitaire  
de gériatrie de Montréal

Fondation Courtois

UNIQUE



**UPMC**  
SORBONNE UNIVERSITÉS



COLLÈGE  
DE FRANCE  
1530



**UCSF** Weill Institute for Neurosciences

Memory and Aging Center



UNIVERSITÉ  
DE GENÈVE



nccr  
Revolving  
language  
NATIONAL CENTRE OF COMPETENCE IN RESEARCH



Campus Biotech, H8.03



[noce-lab.github.io](https://noce-lab.github.io)



**Neurobiology of  
Concepts Expression**  
[noce-lab.github.io](https://noce-lab.github.io)



@NoCeLab



Mon-Thr, 9-to-5

## Can Büyükberber – Stimuli Space

