



# Immediate and long term cognitive improvement after cognitive vs. emotion management psychoeducation programs

# A randomized trial in covid patients with neuropsychological difficulties

## **COVCOG Study**

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# Long COVID

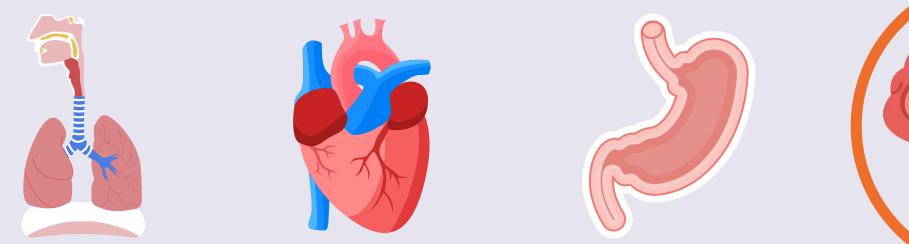
Introduction





# Manifestations of Long COVID

• Multisystemic syndrome affecting several organic systems :



- Fatigue and cognitive difficulties are among the most common reported symptoms
  - Subjective complaints (i.e. concentration, memory, multitasking)
  - Objective impairment (i.e. attentional, memory, executive)

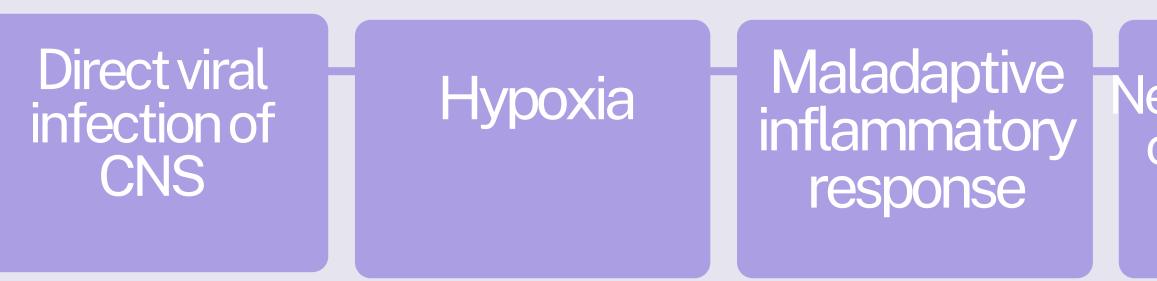
Mehandru et Merad, 2022 ; Salamanna et al., 2021; Liu et al., 2023 ; Han et al., 2022; Tavares-Júnior et al., 2022





# Pathophysiology of Long COVID

Complex interplay of factors from different aetiologies



Politi et al., 2020; Dondaine et al., 2022; Najjar et al., 2020; Wu et al., 2024; Molnar et al., 2024; Poletti et al., 2022; Dani et al., 2021

### Neuropsychiatric comorbidities

Dysfunction of the autonomic nervous system



# How symptoms evolve?

Persist even 2 years after infection

 Improvement observed but 30% still report symptoms affecting everyday life (related to cognition, sensorimotor function and mental fatigue)

Wahlgrend et al., 2023; Han et al.,



# **Different cognitive profiles**

 Distinct clinical phenotypes of Long COVID

Lack of awareness for memory dysfunction Lack of awareness for anosmia

Voruz et al., 2022; 2024

**Greater memory** impairment ; fewer psychiatric symptoms and better quality of life

 Distinct recovery trajectories 1 year post-infection

Improved performances

Persisting or increased neuropsychological deficits



# What are the treatment options?

- Adaptation of pre-existing therapies (i.e. cognitive rehabilitation programs) in ABI or PCS)
- Multiplicity of symptoms, including psychological factors = Cognitive Behavioural Therapy (CBT)

Mathern et al., 2022; Tay et al., 2021

# Objective

What is the most effective psychoeducational intervention (cognitive vs. affective) for Long COVID patients with cognitive complaints? (2 months follow-up evaluation)

Hypothesis:

Superior efficacity expected with a cognitive approach





# COVCOG Study

Effects of two interventional programs (cognitive vs. affective psychoeducation) in Long COVID patients with cognitive difficulties

### COVCOG Studv

Willems et al. BMC Neurology (2023) 23:307 https://doi.org/10.1186/s12883-023-03346-9

### STUDY PROTOCOL

COVCOG: Immediate and long-term cognitive improvement after cognitive versus emotion management psychoeducation programs - a randomized trial in covid patients with neuropsychological difficulties

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- Pre-registration (clinicaltrials.gov: NCT05167266)
- Randomized Control Trial (RCT)
- Data collection between March 2022 and June 2024
- N=130 randomized in either cognitive or affective intervention (ratio 1:1)
- Cognitive complaints at least 3 months after SARS-CoV-2 infection

**BMC Neurology** 





Willems et al., 2023



# Chronology of the study (10.5 months)

Randomization

Baseline evaluation

Phone screening : pre-inclusion INTERVENTION Cognitive vs. Affective (4 sessions of 1h30) Follow-up evaluation 2 months postintervention

> Follow-up evaluation 8 months postintervention



# Neuropsychological evaluation

### Cognitive assessment

- Memory
- Attention
- Cognitive control
- Langage



- (BRIEF-A)
- (MMQ)

### Self-reported questionnaires

- **Primary outcomes =**
- cognitive complaints
  - Cognitive control
  - Memory functioning

- Fatigue
- Sleep difficulties
- Quality of life
- Psychological distress
- Impact daily on activites

# **Cognitive intervention**

- 4 sessions of 1h30 + reactivation session of 30 min (after 1 month)
- Intensified by videotherapy and home exercices
- Psychoeducation targeting metacognition to teach appropriate behaviours and strategies

<sup>-</sup>atigue and sleep (i.e. recognition and management of energy)

**Cognitive control** (i.e. planification)

Short-term memory and attention (i.e. environmental adjustments and internal strategies)

Long-term memory (i.e. internal strategies and external aids)

# Affective intervention

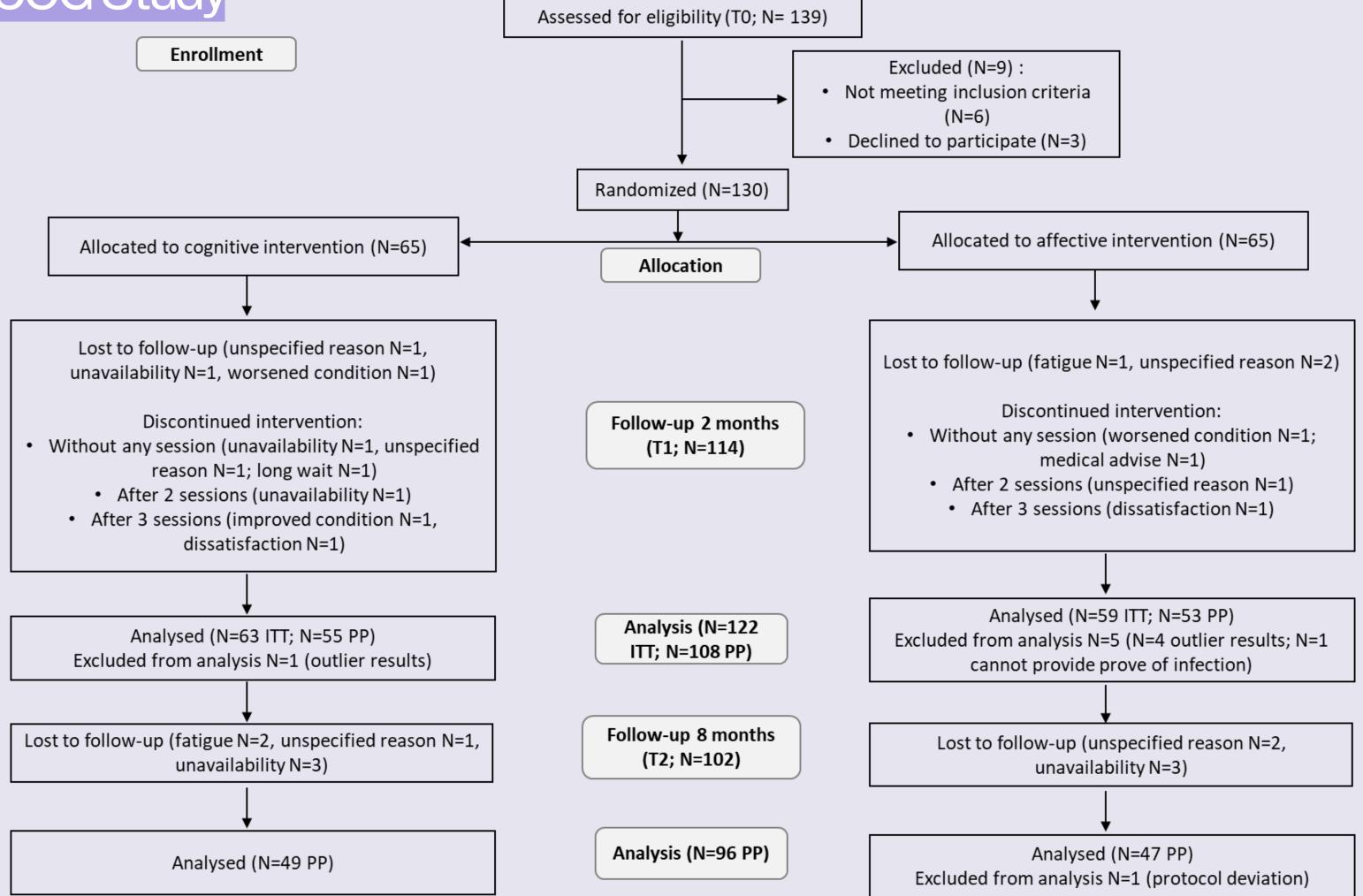
- 4 sessions of 1h30 + reactivation session of 30 min (after 1 month)
- Intensified by relaxation exercises, notes and home exercises
- Psychoeducation targeting selfefficacy for emotions management and regulation of behaviours impacting the perception of difficulties on daily living activites

Recognising difficulties and emotions

Accepting and communicating emotions Tolerating uncertainty and worries

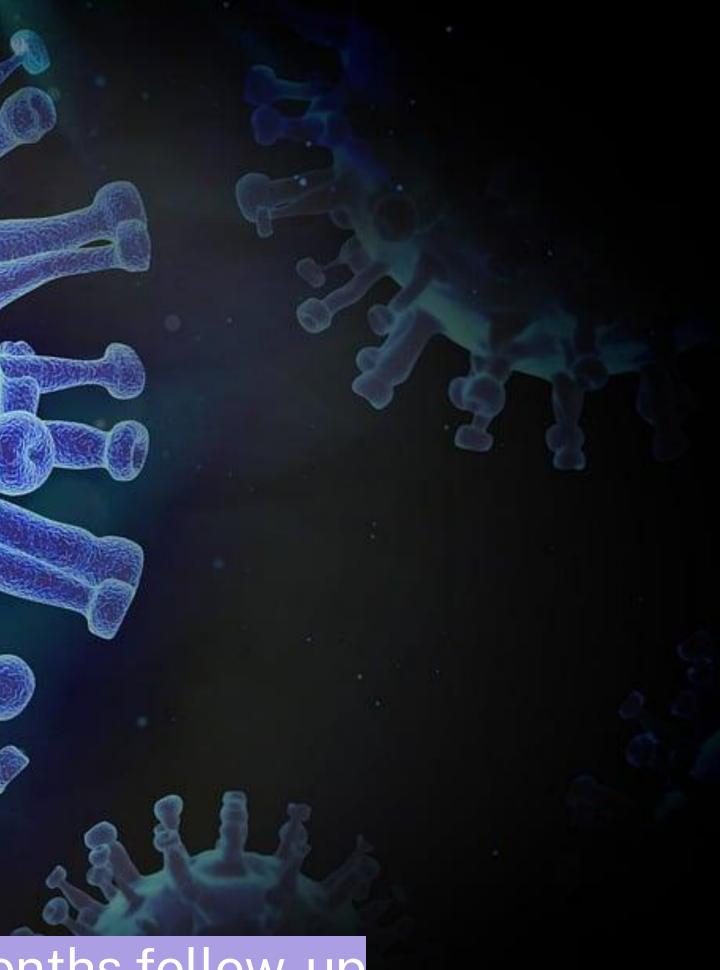
Reconnect with ourself and reactivation

### **COVCOG** Study



# Results

### Effects of psyhoeducative interventions at 2 months follow-up





		Total
Demographics	Age (mean ± SD) [range]	47 ± 10 [21-66]
	Sex (female)	85 (69.7%)
	Years of education (mean ± SD) [range]	14 ± 3 [6-17]
History of COVID-	Asymptomatic	1 (0.8%)
19	Mild infection	67 (54.9%)
	Moderate infection	41 (33.6%)
	Severe infection	13 (10.7%)
	Hospitalized	16 (13.1% ; 10 female)
	ICU treatment; mean stay	8 (6.6% ; 3 female) ; 13 days
	Number of infections (mean ± SD) [range]	1.7 ± 0.9 [1-5]
	Time since first infection (months)	20.9 ± 8.6 [4-39]

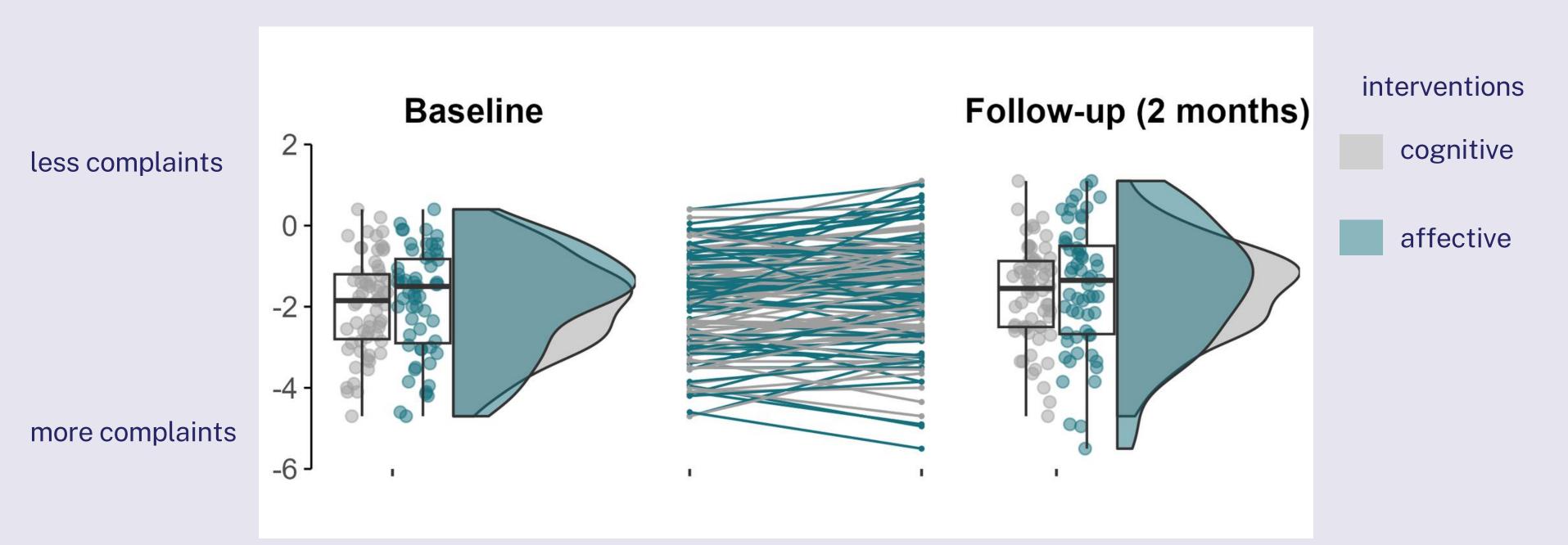


# **Baseline evaluation - Primary outcomes**

- 40% (N=49) of patients meet the difficulty threshold at the baseline for cognitive control complaints
- 35% (N=43) of patients were severily dissatisfied about their memory functioning at baseline
- No difference observed between groups prior to the interventions (all ps>0.124)



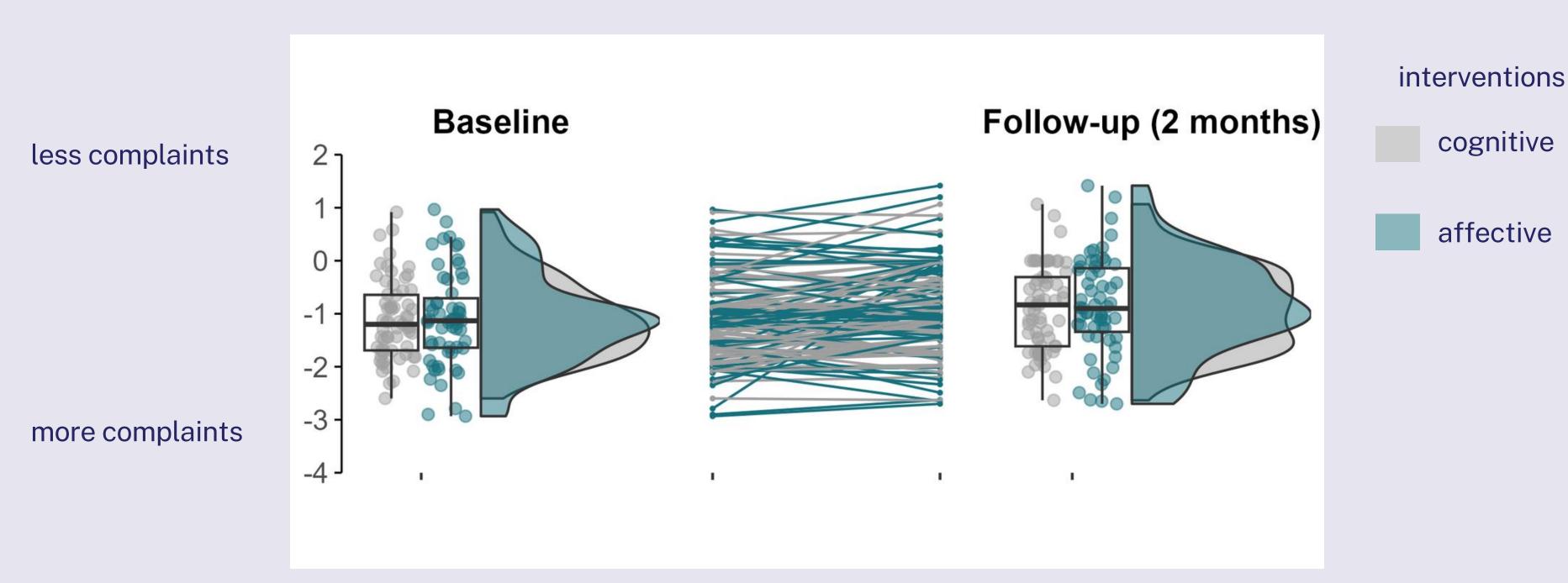
## 2 months follow-up : cognitive control



- Cognitive control complaints decreased at 2 months FU for both intervention groups (F = 17.417, p = .008, SES = -0.14 [95% Cl: -0.21, -0.07])
- No moment\*group interaction (F = 0.173; p=0.67)



# 2 months follow-up: satisfaction with memory



- Memory complaints decreased at 2 months FU for both intervention groups, (F = 16.325 ; p < .001, SES = -0.11 [95% CI: -0.16, -0.06])
- No moment\*group interaction effect (F=0.034; p=0.8)

### Results

## 2 months follow-up: secondary outcomes

	Time effect on secondary outcomes			
	F value	P value	SES	95% IC
Cognitive Tests				
Attention	9.861	.002	-0.15	[-0.24, -
Memory	10.218	.002	-0.13	[-0.21, -0
Executive	3.742	.055	-0.09	[-0.18, 0
Quality of Life	12.873	<.001	0.16	[ 0.07, 0
Fatigue				
Physical Fatigue	13.304	<.001	0.15	[ 0.07, 0
Cognitive Fatigue	20.630	<.001	0.22	[ 0.12, 0
Psychosocial Fatigue	2.8315	.09	0.08	[-0.01, (
Sleep	5.4345	.02	0.10	[ 0.01, C
Psychological distress	3.5096	.06	0.07	[ 0.00, 0
Activities				
activity_impairment	3.289	.07	0.07	[-0.01, (
work_impairment	7.578	.007	0.13	[ 0.04, 0

-0.06]
0.05]
0.00]
0.24]
0.23]
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0.18]
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### Results

# Reliable change TO-T1 (primary outcomes)

Intervention	Memory functioning	Cognitive control
Cognitive	29 (46%)	23 (37%)
Affective	23 (39%)	23 (39%)

N total = 122 (63 cognitive + 59 affective)



# Influence of sponteneous recovery?

Linear regressions :

- No time effect between first infection and baseline evaluation (cognitive control, p=0.77; memory, p=0.64)
- No time effect between first infection and follow-up evaluation (cognitive control, p=0.69; memory, p=0.15)



Sponteneous recovery is highly unlikely

### **Conclusions and perspectives**

- For both intervention groups : decrease in cognitive complaints; attentional and memory domains; quality of life; fatigue and sleep difficulties; and impairment on work
- Significant improvement due to specific aspects of the interventions? general effect? placebo response?

In perspective :

- Specific benefits of one or the other intervention on certain outcomes?
- Complementary qualitative study to explore the implementation process of the two interventions

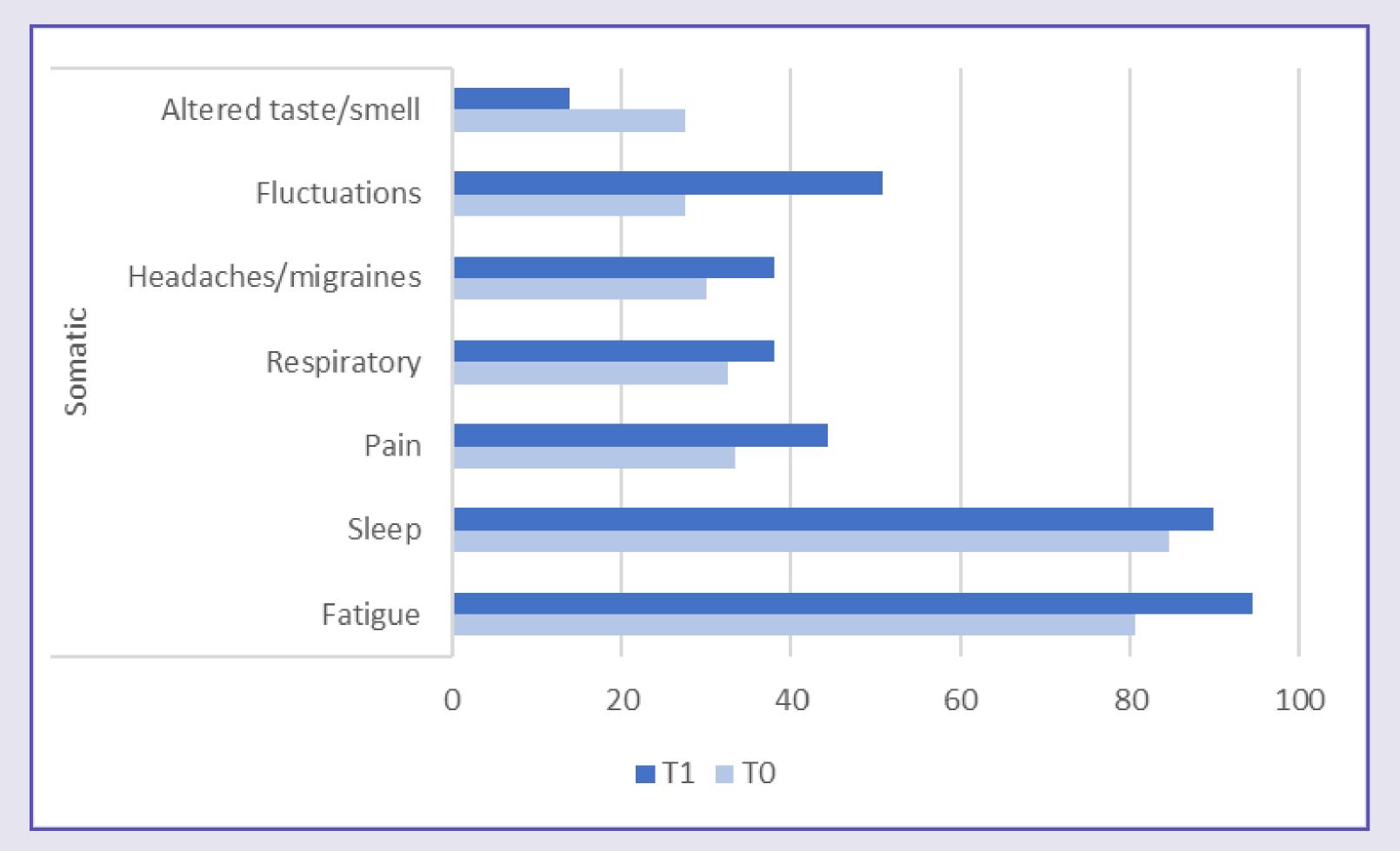


# Thank you very much!





### Somatic complaints sponteneously reported





## Cognitive complaints sponteneously reported

