



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

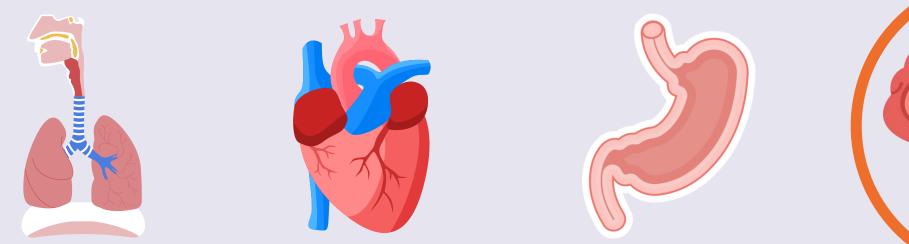
Introduction





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)

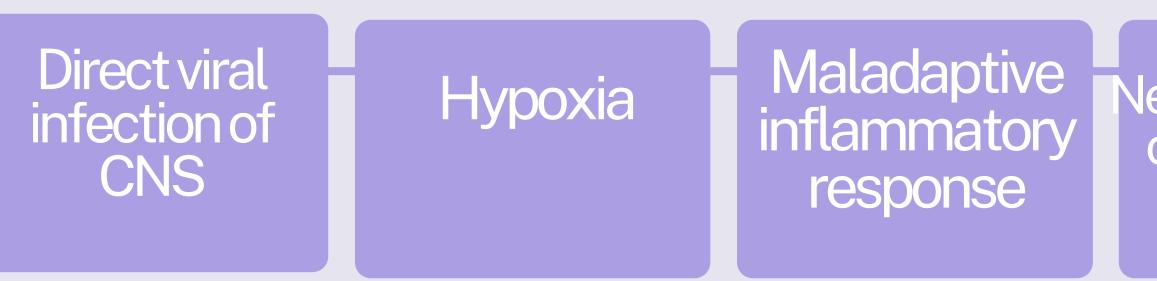
Mehandru et Merad, 2022 ; Salamanna et al., 2021; Liu et al., 2023 ; Han et al., 2022; Tavares-Júnior et al., 2022





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





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

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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

⁻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

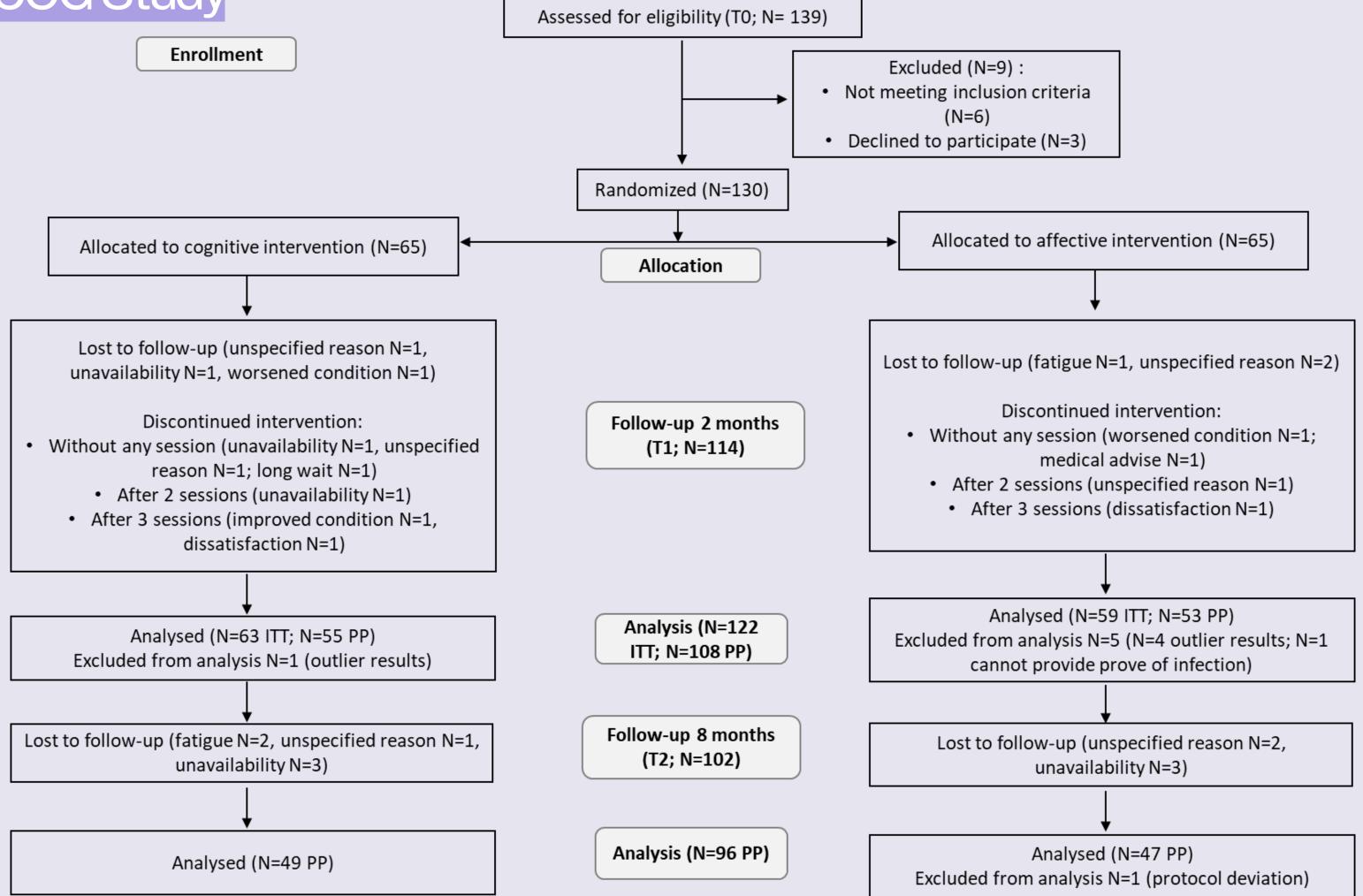
- 4 sessions of 1h30 + reactivation session of 30 min (after 1 month)
- Intensified by relaxation exercises, notes and home exercises
- Psychoeducation targeting selfefficacy for emotions management and regulation of behaviours impacting the perception of difficulties on daily living activites

Recognising difficulties and emotions

Accepting and communicating emotions Tolerating uncertainty and worries

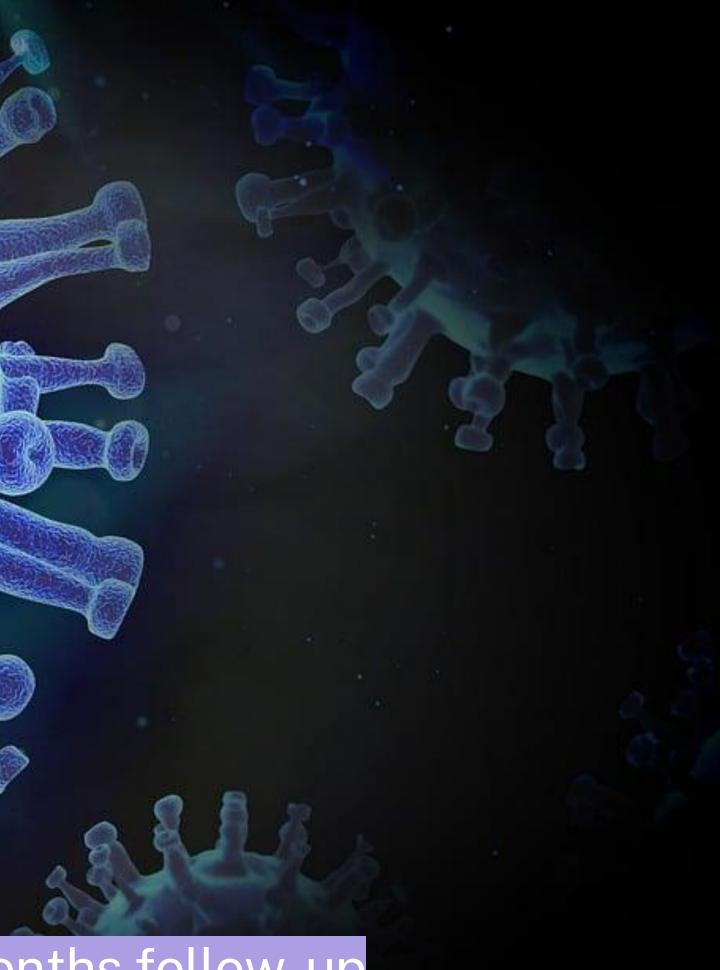
Reconnect with ourself and reactivation

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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] |

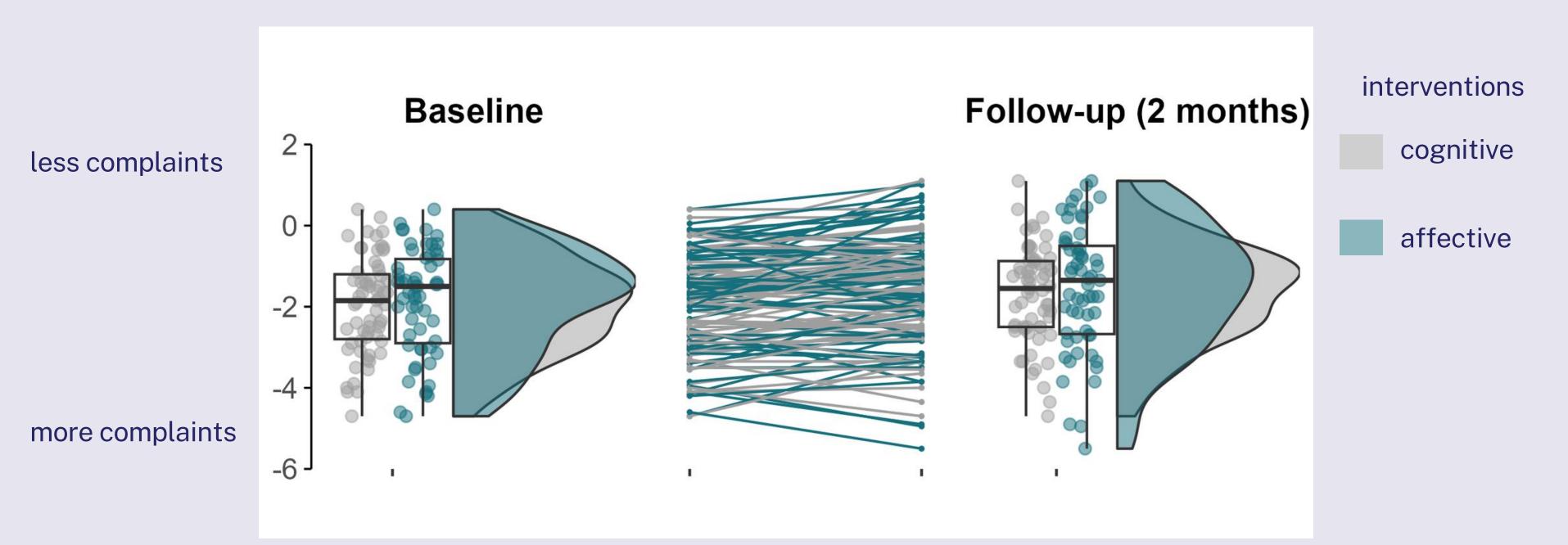


Baseline evaluation - Primary outcomes

- 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)



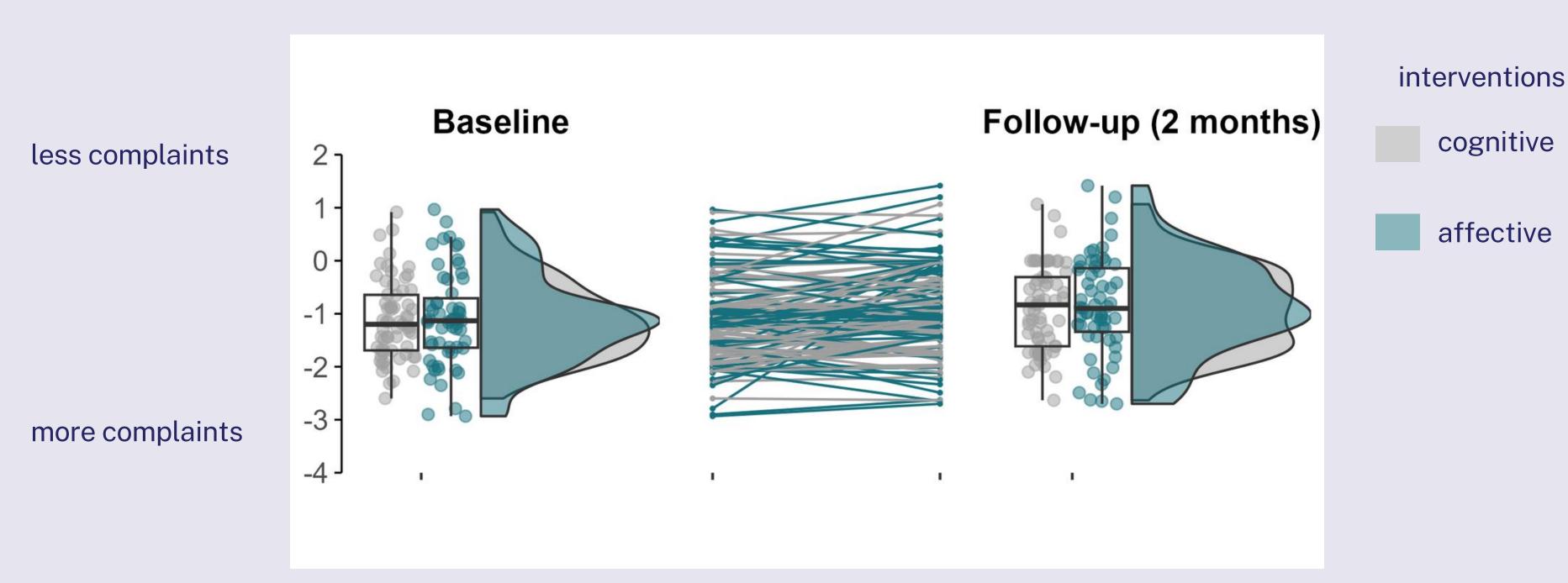
2 months follow-up : cognitive control



- 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 |
| | | | | |

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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

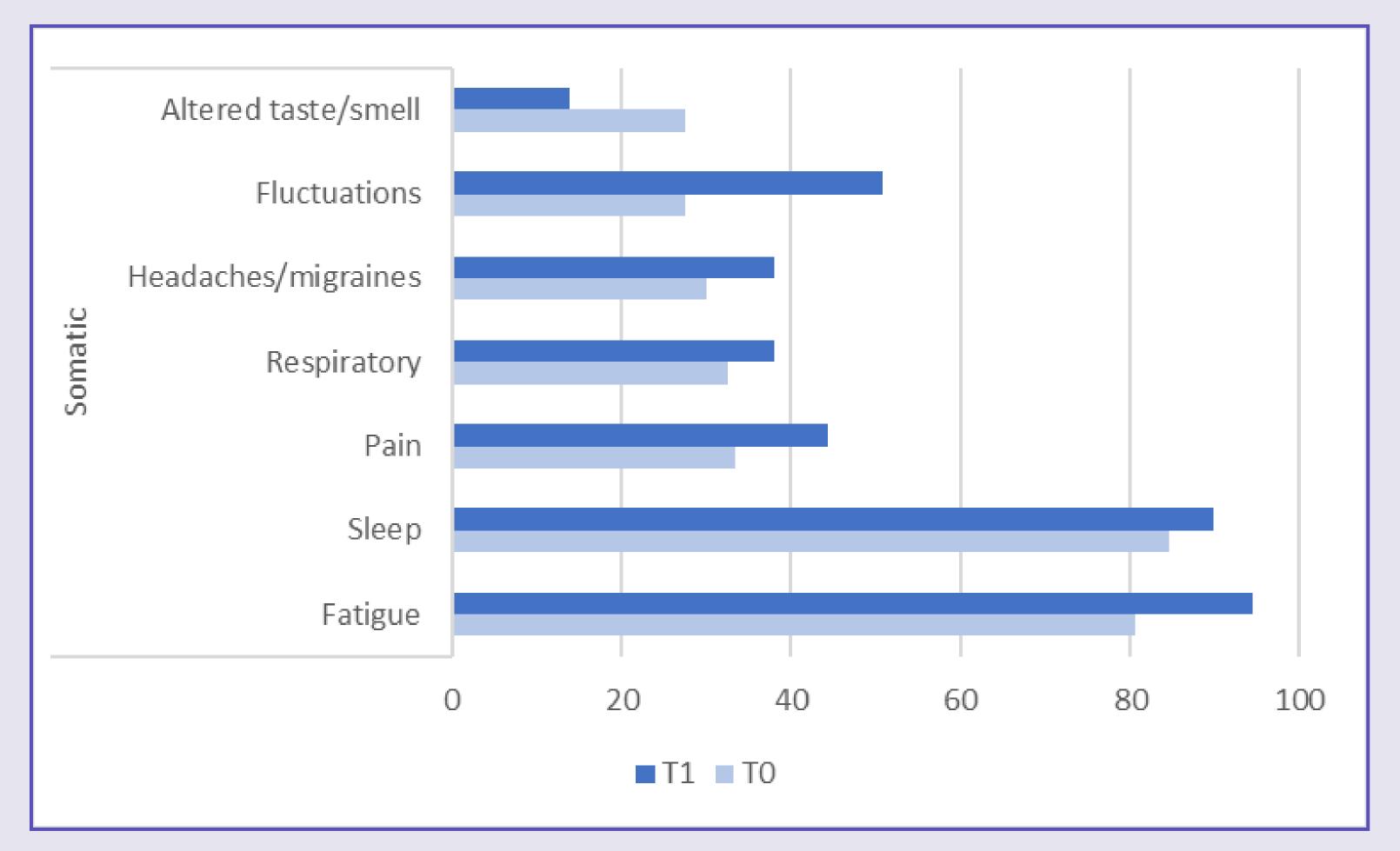


Thank you very much!





Somatic complaints sponteneously reported





Cognitive complaints sponteneously reported

