# SLEEP 2024

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A JOINT MEETING





# **Effects of Solriamfetol** on Cognition in **Patients with Excessive Daytime Sleepiness Associated** With Narcolepsy

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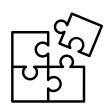
### **Background: Narcolepsy and Cognition**



 Narcolepsy is a chronic sleep disorder characterized by excessive daytime sleepiness (EDS)<sup>1</sup>



- Brain fog and difficulty concentrating are common complaints among patients and significantly impact their quality of life<sup>2</sup>
- Patients often exhibit deficits in processing speed and attention, core cognitive functions<sup>3</sup>



 Cognitive impairment persists despite treatment with wake-promoting agents, and remains an unmet need



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Rosenberg R. et al. J Clin Sleep Med. 2024; 20(4): 643-651

## **Background: Solriamfetol and Cognition**



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 Solriamfetol (Sunosi®) is a dopamine-norepinephrine reuptake inhibitor with agonistic properties at the trace amine-associated receptor 1 (TAAR1) and serotonin 1A (5HT1<sub>a</sub>) receptor<sup>1</sup>



 Solriamfetol is approved for treatment of EDS associated with narcolepsy or obstructive sleep apnea (OSA)<sup>2,3</sup>

 Solriamfetol improved cognitive performance in a clinical study of patients with OSA and EDS with cognitive impairment<sup>4</sup>



 Here we present cognitive outcomes of patients with narcolepsy and EDS treated with solriamfetol in a real-world setting

<sup>1.</sup> Gursahani H, et al W. Sleep. 2022;45(suppl 1): A329.

<sup>2.</sup> Sunosi® (solriamfetol) [Prescribing Information]. New York, NY. Axsome Therapeutics, Inc.

<sup>3.</sup>Sunosi™ (solriamfetol) tablets Summary of Product Characteristics. Waterford, Ireland: TMC Pharma (EU) Limited; 2022.

<sup>4.</sup> Van Dongen H, et al. Effects of Solriamfetol on Cognitive Function in Participants With Cognitive Impairment Associated With Excessive Daytime Sleepiness in Obstructive Sleep Apnea: Results of the SHARP Study; 2023 November 9-12; Neuroscience Education Institute Congress; Colorado Springs, CC

### **Methods: SURWEY Study**



- **SU**nosi **R**eal **W**orld **E**xperience Stud**Y** (SURWEY) was a real-world, retrospective chart review among physicians in Germany of patients prescribed solriamfetol for EDS associated with narcolepsy type 1 and 2
- SURWEY enrolled sleep clinics with physicians experienced in treating EDS and in prescribing solriamfetol

- Key patient inclusion criteria: age >18 years; stable dose of solriamfetol; ≥6 weeks treatment completed
- Physicians completed surveys based on patient records (sociodemographic and clinical characteristics) and solriamfetol dosing information

#### Methods



- Analyzed a subgroup of patients (n = 52) with narcolepsy who underwent cognitive assessments prior to initiating solriamfetol and 3 months following
- Assessments were self-reported and objective, across multiple cognitive domains
- Results are pooled across dosages; most patients received less than 150 mg/day (maximum recommended dose)

Baseline (n = 52)

Solriamfetol
Initiation

3-month Follow-up (n = 52)

Assessment	Domain	Modality
British Columbia Cognitive Complaints Inventory (BC-CCI)	Cognitive impairment	Self-reported
Test of Attentional Performance (TAP) Alertness subtests	Alertness	Objective
Wechsler Adult Intelligence Scale-IV (WAIS-IV) Coding subtest	Processing speed	Objective
Wechsler Memory Scale-IV (WMS-IV) Visual Reproduction I and II	Visual memory	Objective
Regensburger Word Fluency (RWT) S Words and Animals	Verbal fluency	Objective
Epworth Sleepiness Scale (ESS)	Sleepiness	Self-reported

- Mean age was 36.4 years
- 44.2% were female

# Study Findings: Solriamfetol Treatment Improved Subjective Cognitive Function



• Patient-perceived cognitive complaints were evaluated with the British Columbia-Cognitive Complaints Inventory (BC-CCI)

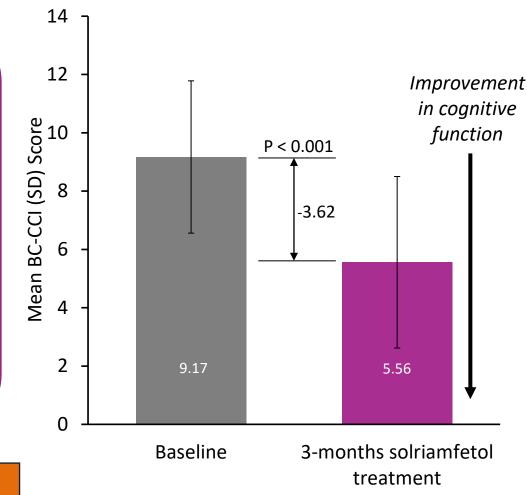
#### **6 Cognitive Complaint Items**

Participants were asked to rate their **problems** with concentration, memory, and thinking skills during the past 7 days. Questions included:

- 1. Forgetfulness/memory problems
- 2. Poor concentration
- Trouble expressing thoughts
- 4. Trouble finding the right word
- 5. Slow thinking speed
- 6. Trouble figuring things out or solving problems

A 4-point scale (0–3) was used with higher scores indicating greater cognitive impairment:

- 0 = Not at all
- 1 = Some
- 2 = Quite a bit
- 3 = Very much



Solriamfetol
significantly
improved subjective
cognitive function
by 3.62 points
(39.4%)

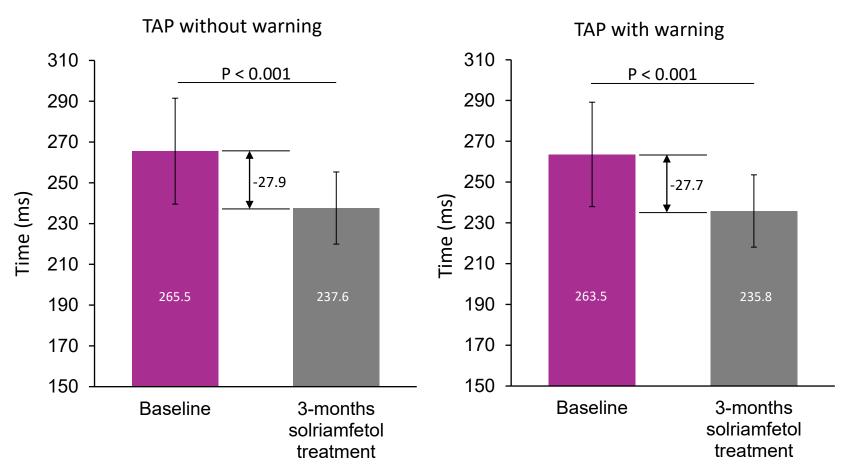
\*Baseline scores indicated moderate cognitive complaints prior to solriamfetol treatment



### **Study Findings: Solriamfetol Improved Alertness**



- Test of Attentional Performance (TAP) assesses multiple attentional functions, including alertness<sup>1</sup>
  - Previously used to assess cognitive deficits in patients with narcolepsy<sup>2</sup>



Solriamfetol significantly improved alertness in the TAP regardless of warning signal (10.5% for both)

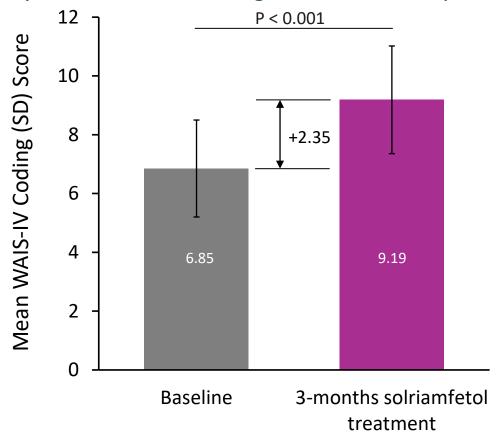
Naumann A, et al. J Sleep Res. 2006;15(3):329-38.

Nin V. et al. J Neurol Disord. 2022:10:12.

# Study Findings: Solriamfetol Improved Processing Speed



- Processing speed was evaluated using the Coding subtest (a variation of the Digit Symbol Substitution Test) of the Wechsler Adult Intelligence Scale-IV (WAIS-IV), a multi-component IQ test<sup>1</sup>
  - Previously used to measure cognitive deficits in patients with narcolepsy<sup>2</sup>



Solriamfetol statistically significantly improved processing speed by 2.35 points (34.3%)

<sup>\*</sup>Baseline score indicated impaired processing speed prior to solriamfetol treatment

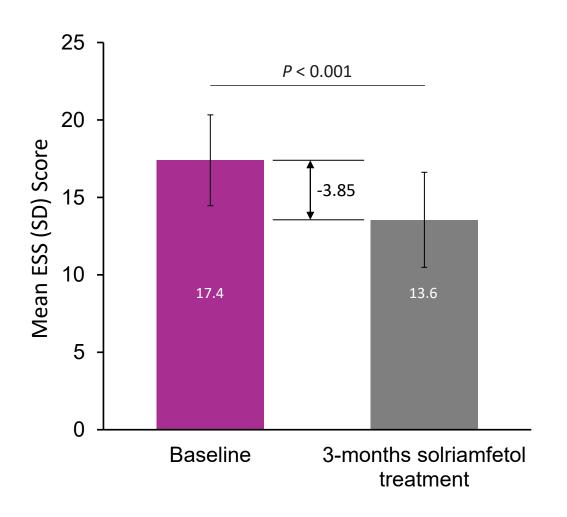
<sup>1.</sup> Wechsler D. Wechsler Adult Intelligence Scale: WAIS-IV; Technical and Interpretive Manual. San Antonio, Tex: Pearson; 2008.

Nin V, et al. J Neurol Disord. 2022;10:12.

# Study Findings: Solriamfetol Reduced Excessive Daytime Sleepiness



• EDS was assessed with the Epworth Sleepiness Scale (ESS), a self-report questionnaire



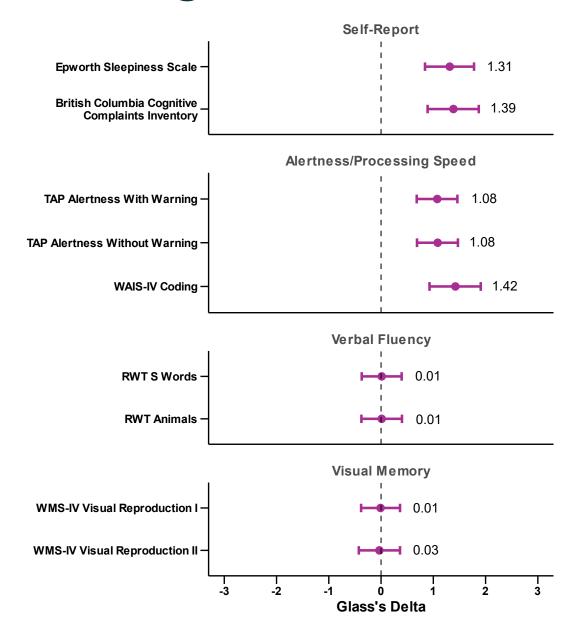
Solriamfetol statistically significantly improved excessive daytime sleepiness

### **Study Findings: Sleepiness and Cognition**



Solriamfetol improved sleepiness, subjective cognitive function, alertness, and processing speed

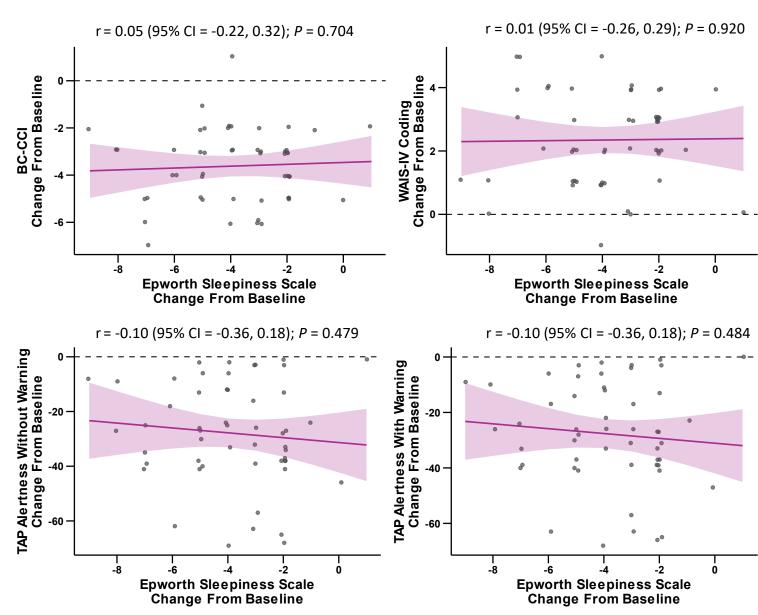
No meaningful effects on verbal fluency on visual memory



# Study Findings: Reduction in Sleepiness not Predictive of Cognitive Improvement with Solriamfetol



ess score improvements were not predictive of improvements in multiple measures of cognition



#### **Conclusions**



- In this retrospective, real-world study, cognitive performance was assessed in patients with EDS associated with narcolepsy
- Solriamfetol treatment for 3 months improved:
  - Patient-reported cognitive function
  - Alertness and processing speed
  - Excessive daytime sleepiness
  - Cognitive performance
- These results indicate solriamfetol has the potential to improve cognitive function in patients with EDS associated with narcolepsy



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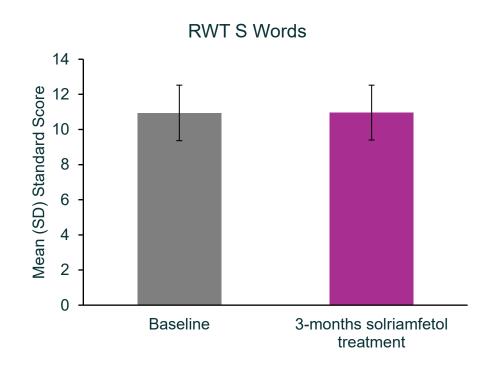
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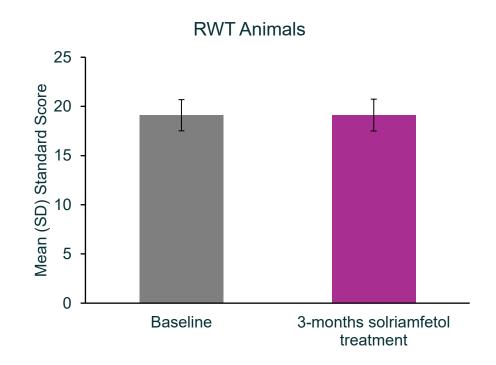
Backup slides

#### **Results: Verbal Fluency**



- Verbal fluency was assessed with the Regensburger Word Fluency Test, subtests S Words and Animals
- No impairment in verbal fluency was observed at baseline, and no change was observed with solfriamfetol treatment





#### **Results: Visual Memory**



- Visual memory and retention were assessed with the Wechsler Memory Scale Visual Reproduction subtests I and II
- No impairment in visual memory was observed at baseline, as measured by either test

