



**Phase II Alzheimer's &  
Parkinson's Data**

**CTAD 2021**

# WHAT IS ANVS401?

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- ANVS401 (aka Posiphen) is a translational inhibitor of neurotoxic aggregating proteins (TINAPs).
- Lowering levels of neurotoxic proteins leads to improved axonal transport and nerve cell life.
- In turn, that results in improvement in several neurodegenerative diseases, shown here in Alzheimer's and Parkinson's disease.

# TWO PHASE 2 CLINICAL TRIALS – PART I & II

	AD Trial	PD Trial
<b>Therapeutic Area</b>	Early to Moderate AD and PD	
<b>Phase</b>	2	
<b>Patients</b>	14	14+40
<b>Design</b>	Double-Blind, Placebo-Controlled, Biomarker Study	
<b>Endpoints</b>	Evaluation after 25 days	

# ANVS401 MET THESE PRIMARY, SECONDARY & EXPLORATORY ENDOINTS

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- **Primary Endpoint** -- SAFETY  
ANVS401 is safe
- **Secondary Endpoint** -- PHARMACOKINETICS  
as expected from previous experiments
- **Exploratory Endpoints:**
  - BIOMARKERS  
reduced as expected
  - EFFICACY  
improved ADAS-Cog and WAIS in AD patients  
and UPDRS and WAIS in PD patients

# REVERSAL OF TOXIC CASCADE

**LOWER LEVELS OF NEUROTOXIC PROTEINS**

**IMPROVED AXONAL TRANSPORT**

**LOWER INFLAMMATION**

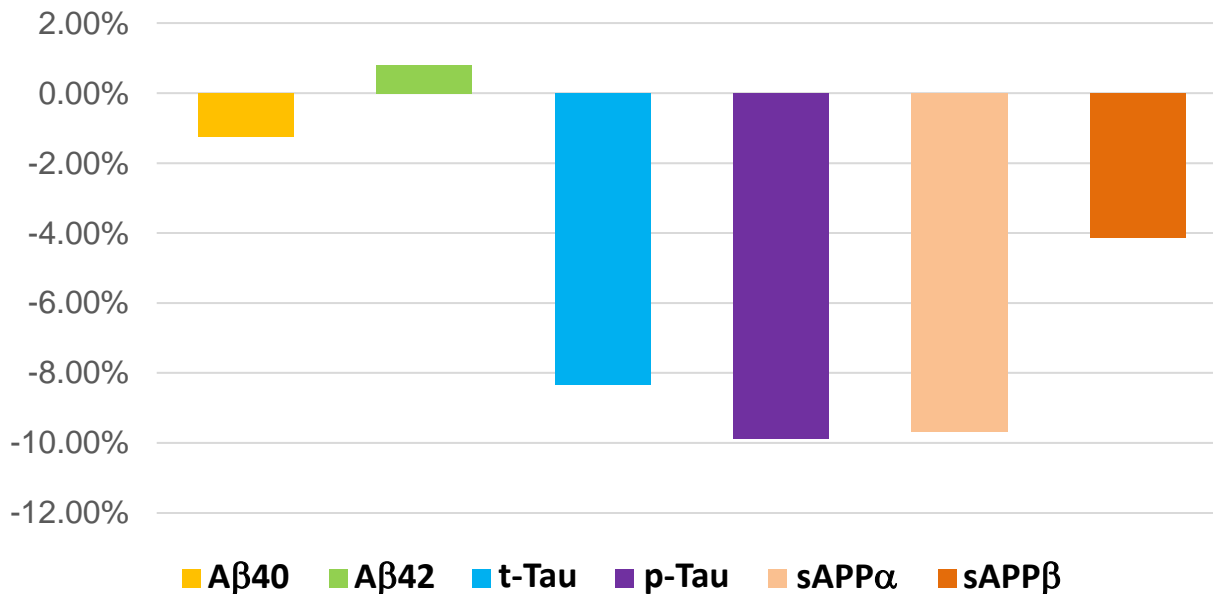
**LOWER NERVE CELL DEATH**

**IMPROVED COGNITION AND MOTOR FUNCTION**

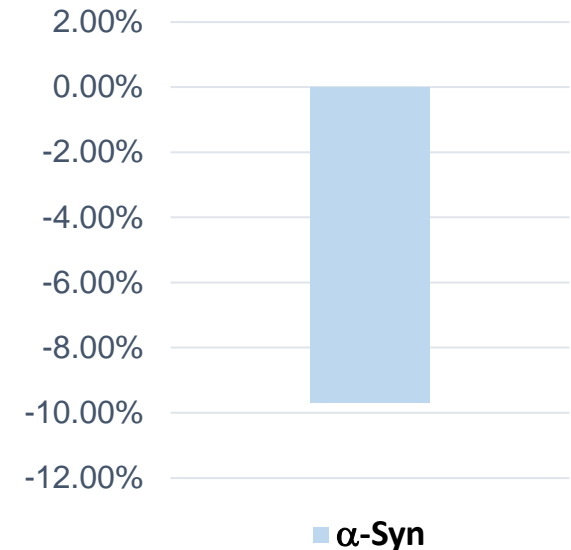
**Please visit our poster [LP13](#) for more information on ANVS401's mechanisms of actions.**

# REDUCED NEUROTOXIC PROTEINS IN BOTH AD AND PD PATIENTS

14 AD Patients-CSF % Change of Neurotoxic Proteins\*



14 PD Patients-CSF % Change of α-Synuclein\*

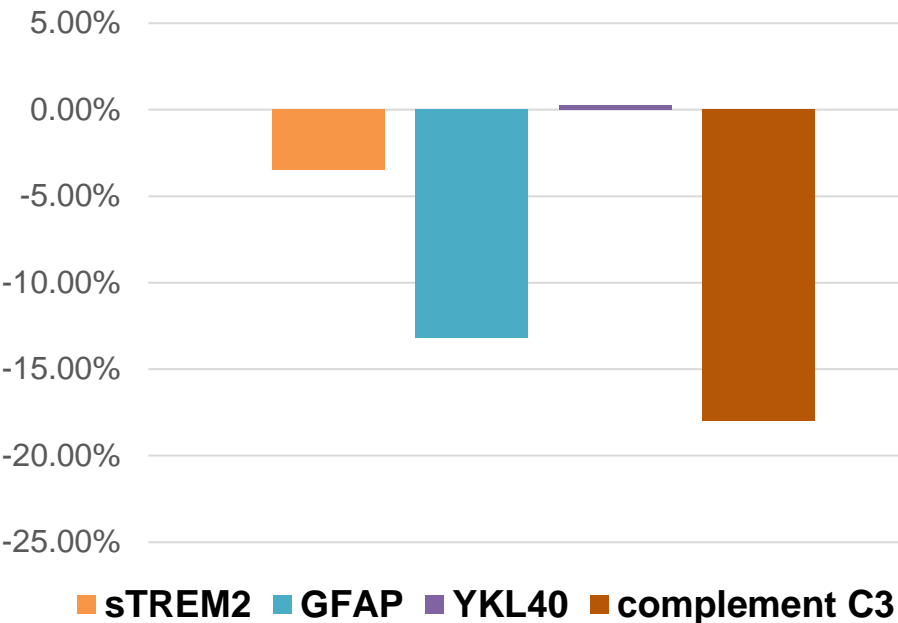


APP (and its downstream products), and p-Tau are the neurotoxic proteins involved in AD, while α-Synuclein is the neurotoxic culprit of PD. The reduction compares well to the reduction seen in animals at full efficacy.

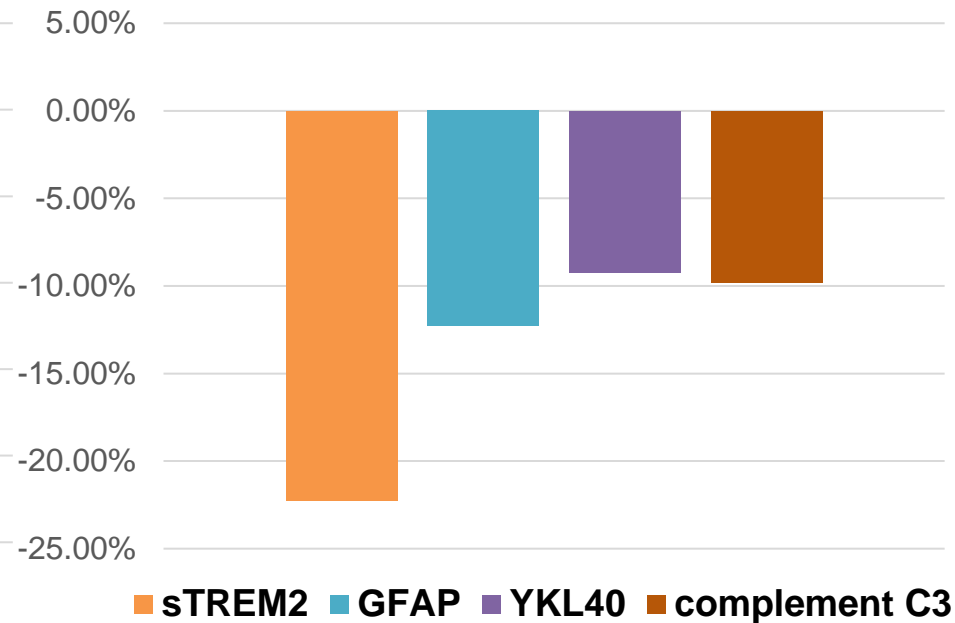
\*All values are in comparison to placebo based on all data points

# REDUCED INFLAMMATION IN BOTH AD AND PD PATIENTS

14 AD Patients-CSF% Change Inflammatory Markers\*



14 PD Patients-CSF% Change Inflammatory Markers\*

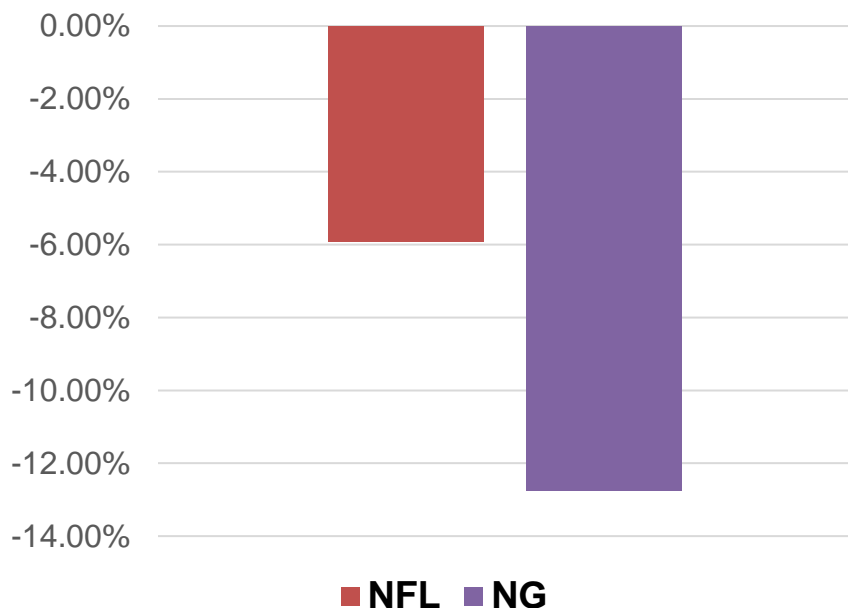


Inflammatory markers are lowered in AD and in PD patients, showing a normalization of inflammation in both neurodegenerative disorders.

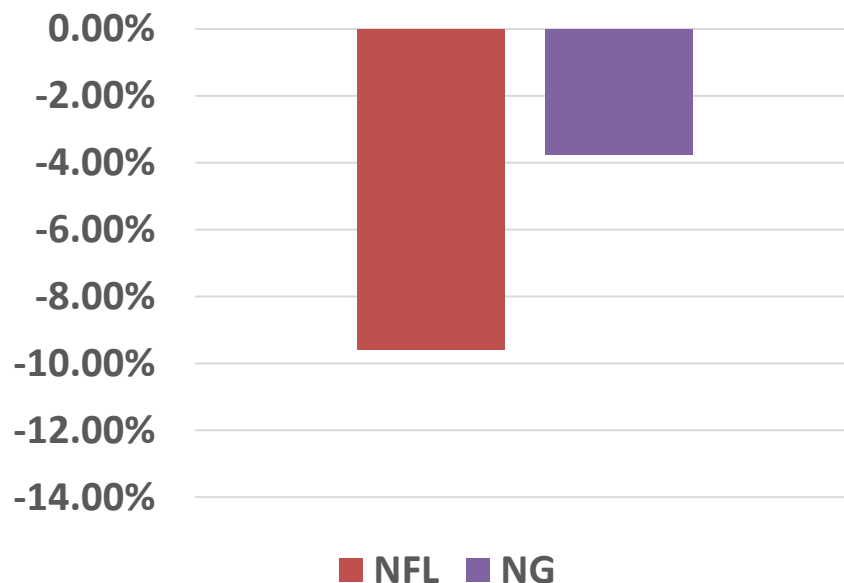
\*All values are in comparison to placebo based on all data points

# REDUCED AXONAL AND SYNAPTIC DYSFUNCTIONS IN BOTH AD AND PD PATIENTS

**14 AD Patients-CSF % Change Axonal and Synaptic Markers\***



**14 PD Patients-CSF % Change Axonal and Synaptic Markers\***

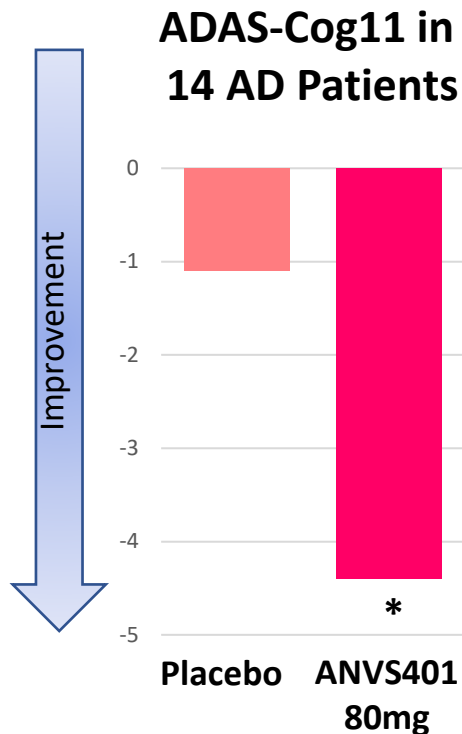


Neuronal and synaptic markers are lowered in AD and in PD patients, showing that the nerve cells are healthier.

\*All values are in comparison to placebo based on all data points.



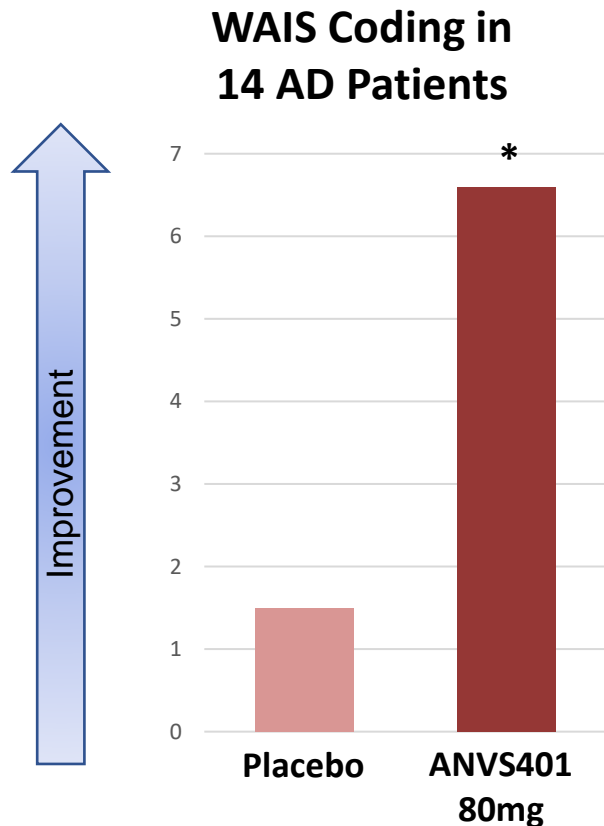
# IMPROVED COGNITION IN AD PATIENTS – ADAS-Cog11



From baseline to 25 days in the ANVS401-treated group, ADAS-Cog11 improves by 4.4 points, a statistically significant improvement of 30% ( $p < 0.05$ ). Compared to placebo at 25 days the treated group is 3.3 points better, an improvement of 22%.

\*  $P < 0.05$

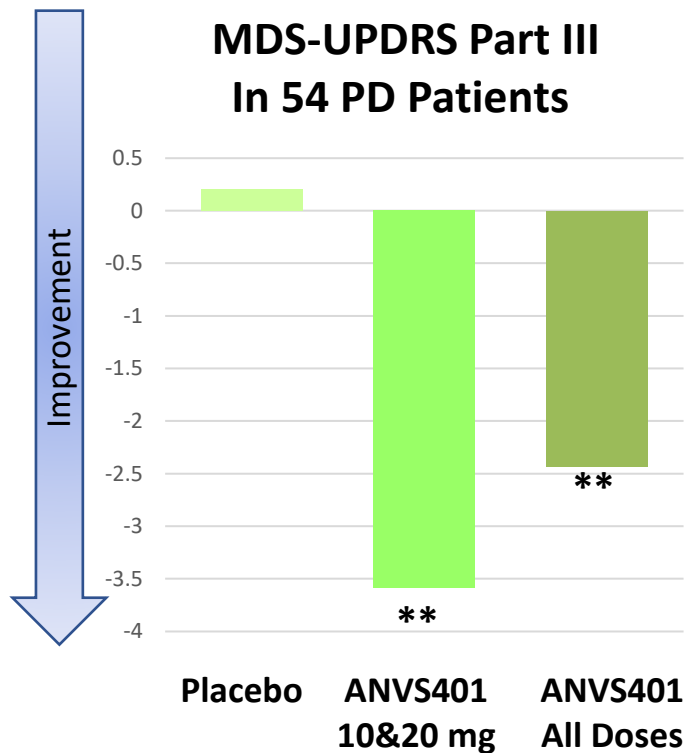
# IMPROVED CODING SPEED IN AD PATIENTS - WAIS



The WAIS coding test measures speed in movement and thinking. Treated AD patients show a statistically significant 23% improvement from baseline.

\*  $P < 0.05$

# IMPROVED MOTOR FUNCTION IN PD PATIENTS – MDS- UPDRS



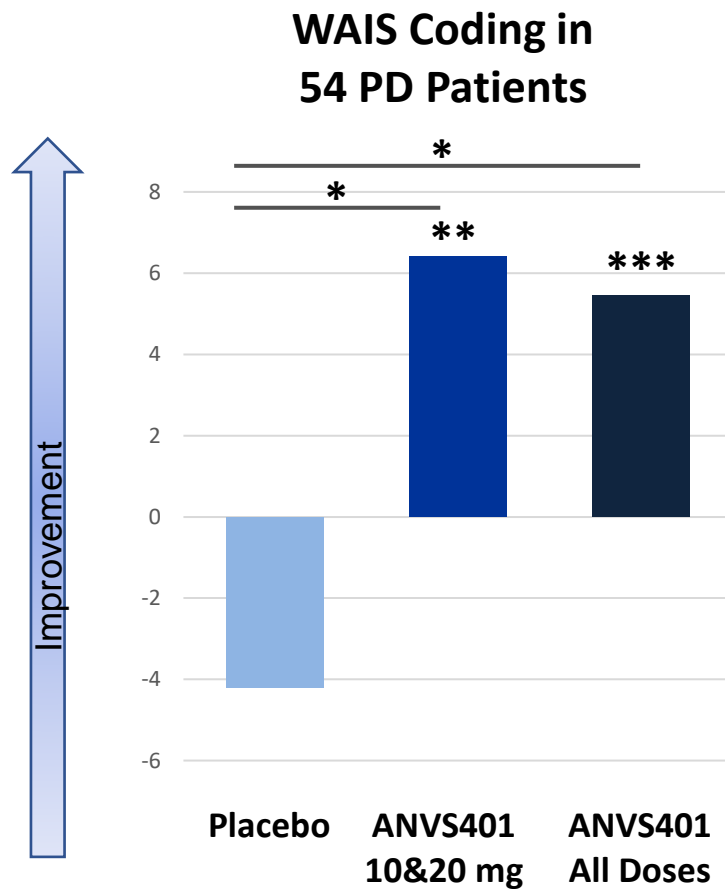
All doses of ANVS401 improved the MDS-UPDRS Part III individually. Combined as shown they improved the Part III score by 2.43 points (13.4%) compared to baseline (\*\* $p < 0.01$ ). 10mg and 20mg are the most efficacious doses and they statistically improved the Part III score by 3.59 points (16.5%) in 25 days comparing to baseline.

\*\* $p < 0.01$

Part II, IV and Total Scores are also improved.

Please visit our poster [LP13](#) for more information on data for all doses

# IMPROVED CODING SPEED IN PD PATIENTS - WAIS



The WAIS coding test measures speed in movement and thinking. PD patients show a statistically significant improvement both from baseline (17.1% for 10+20mg; 13.5% for all doses combined) and from placebo (25.1% for 10+20mg; 22.8% for all doses combined).

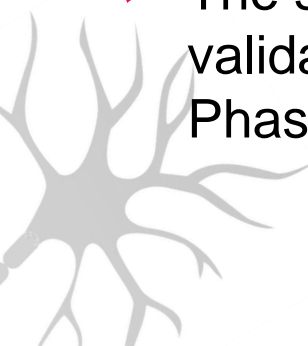
\*  $p < 0.05$  \*\*  $P < 0.01$

\*\*\*  $P < 0.001$

Please visit our poster [LP13](#) for more information on data for all doses

# CONCLUSIONS AND NEXT STEPS



- ✓ ANVS401 shows improvements in Phase 2a clinical trials:
    - Cognition in AD patients
    - Motor function in PD patients
    - WAIS coding in AD and PD patients
  - ✓ This is the first double-blind, placebo-controlled study that improves cognition in AD patients as measured by ADAS-Cog and function in PD patients as measured MDS-UPDRS.
  - ✓ This reduction of neurotoxic biomarkers observed in earlier studies correlates with the ANVS401-induced biomarker reduction with optimal positive outcomes in AD and PD patients.
  - ✓ The successful completion of our Phase 2 clinical trials is providing validation of our approach in two diseases and allows us to plan for Phase 3 trials.
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The logo for ANNNOVIS features the word "ANNNOVIS" in a white, bold, sans-serif font. A red graphic element, consisting of two curved lines that form a partial circle, is positioned behind the letter 'O'.

**ANNNOVIS**

**Thank You!**

**Questions?**