



David Stamler, MD
CEO

AGM
22 November 2024



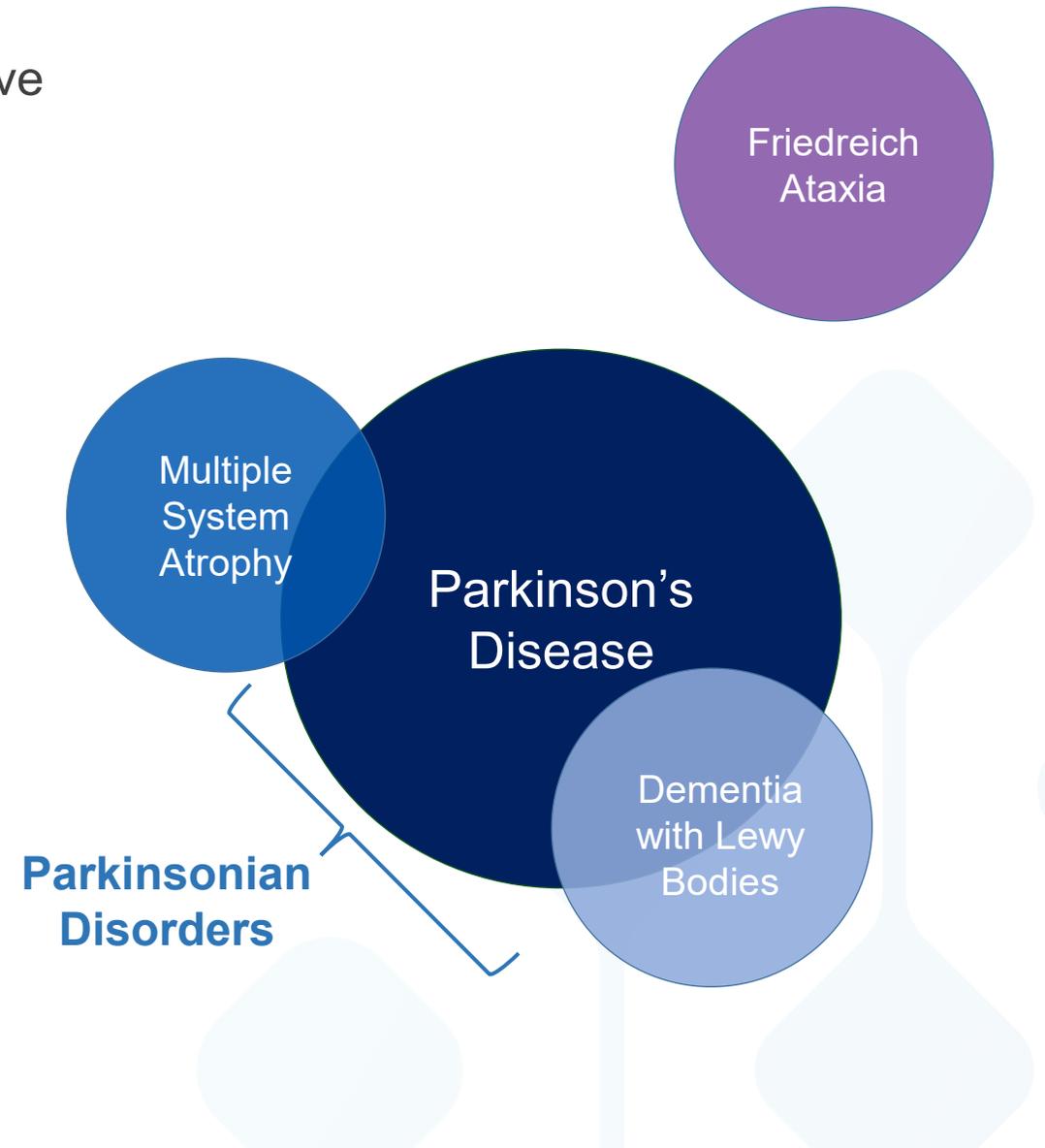
Alterity
THERAPEUTICS

◆ Forward Looking Statements

This presentation may contain some statements that may be considered “Forward-Looking Statements”, within the meaning of the US Securities Laws. Thus, any forward-looking statement relating to financial projections or other statements relating to the Company’s plans, objectives, expectations or intentions involve risks and uncertainties that may cause actual results to differ materially. For a discussion of such risks and uncertainties as they relate to us, please refer to our 2023 Form 20-F, filed with US Securities and Exchange Commission, in particular Item 3, Section D, titled “Risk Factors.”

◆ Targeting Iron-Related Neurodegenerative Diseases

- Iron: central to pathology of many neurodegenerative diseases
- Parkinsonian disorders include
 - Parkinson's disease (PD)
 - Rare diseases with similar motor symptoms
 - Multiple System Atrophy (MSA)
 - Dementia with Lewy Bodies (DLB)
 - Similar underlying pathology
- Friedreich's Ataxia
 - Rare disease with uncoordinated movements
 - Genetic disorder that appears in childhood



Promising Portfolio in Neurodegenerative Diseases



ASSET		PHASE					PARTNER
PROGRAM	INDICATION	DISCOVERY	PRE-CLINICAL	NATURAL HISTORY	PHASE 1	PHASE 2	PARTNER / COLLABORATOR
ATH434-201	Multiple System Atrophy <i>Early Stage</i>	Enrollment Complete					
ATH434-202	Multiple System Atrophy <i>Advanced</i>	Enrollment Complete					
ATH434	Parkinson's Disease						
bioMUSE	Multiple System Atrophy <i>Natural History Study</i>						
ATH434	Friedreich's Ataxia						
Drug Discovery	Neurodegenerative Diseases						

◆ Significant Commercial Opportunity in Treating Multiple System Atrophy

Substantial Unmet Need

Severely debilitating illnesses with no current treatments are ripe for new entrants targeting underlying pathology of the disease.

Unique MOA

Inhibition of protein aggregation is a novel mechanism of action that may prove to impact more than motor symptoms.



Strong Intent to Prescribe

Motivated by efficacy of treating the underlying disease and not just the symptoms, clinicians intend to offer ATH434 to most of their patients with MSA.

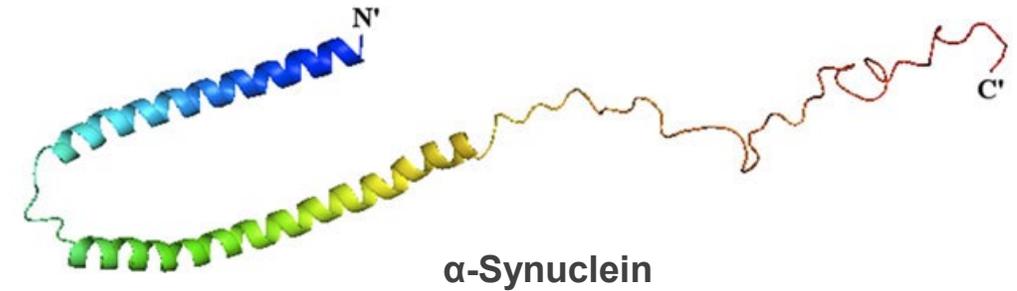
Ease of Use

Twice daily oral administration of ATH434 preferred by physicians

◆ Alpha-Synuclein and Iron in Health

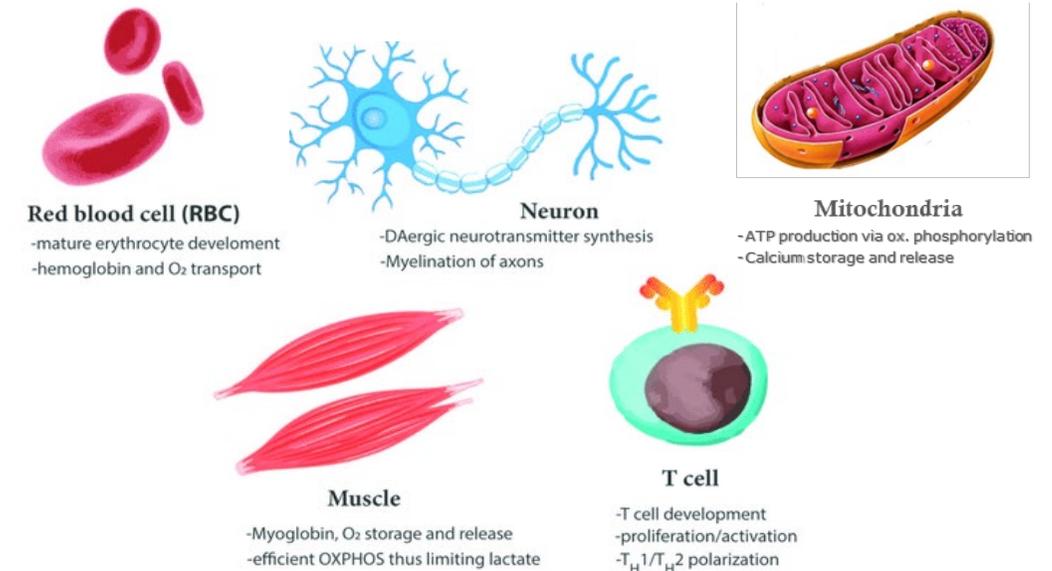
α-Synuclein protein

- Critical for normal function of neurons
- Enables nerves to communicate with each other via neurotransmitters



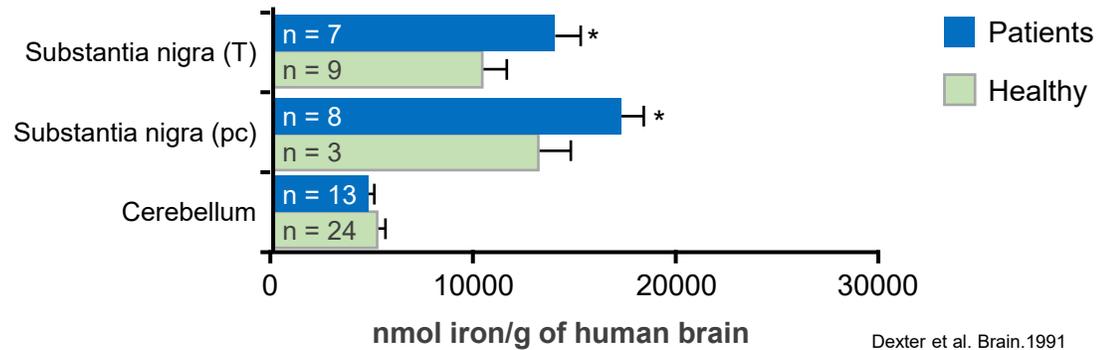
Iron is Essential for Life

- Red blood cell production, oxygen transport
- Neurotransmitter synthesis in neurons
- Energy production via ATP and activity of many enzymes

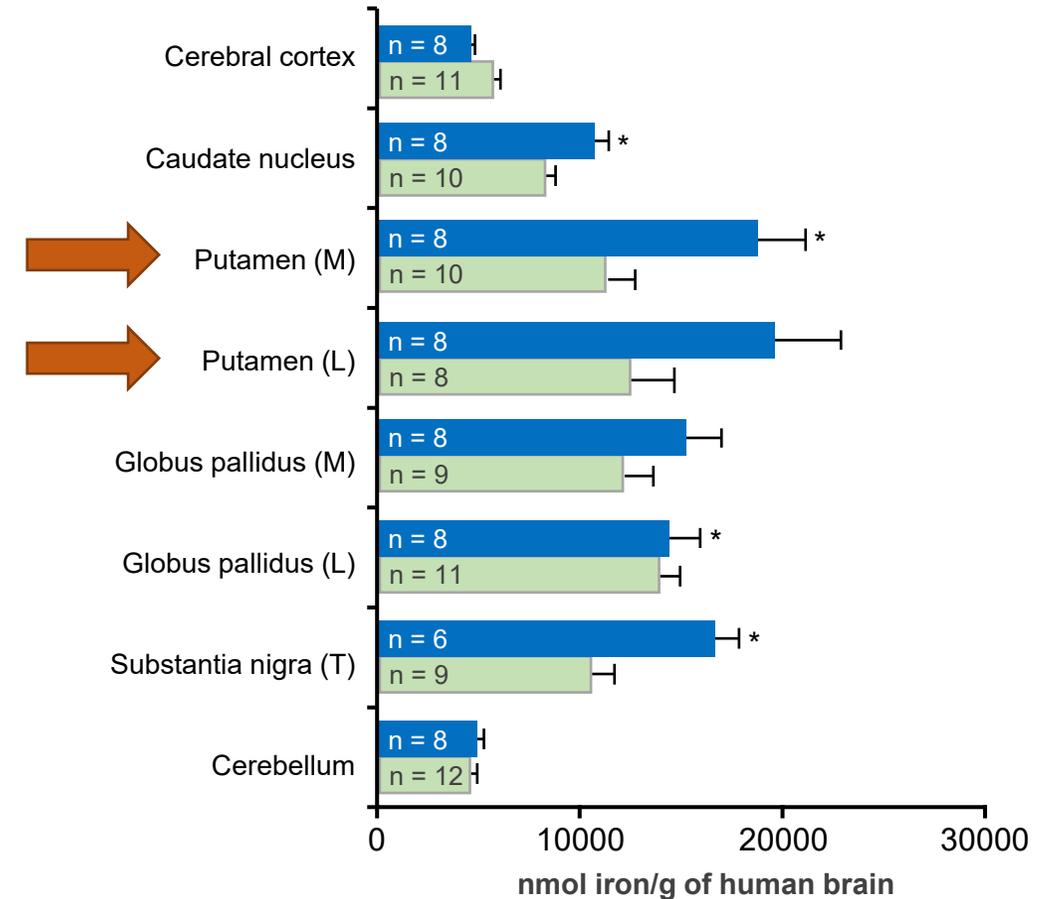


◆ Increased Brain Iron in Parkinson's Disease and Multiple System Atrophy

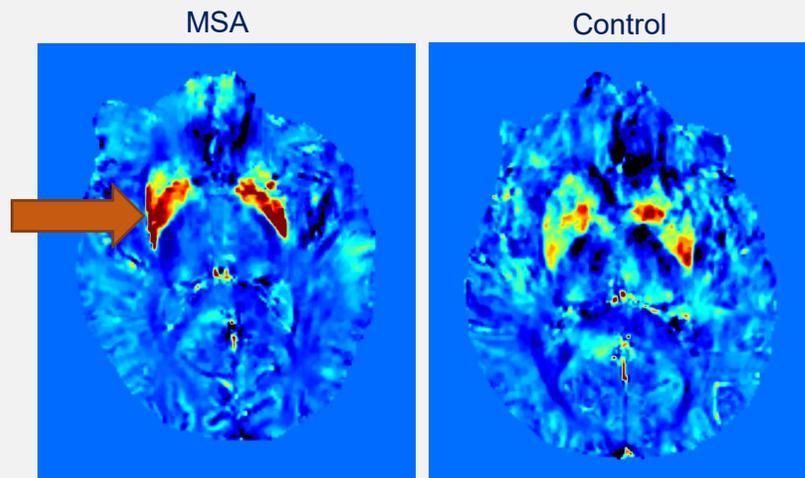
Parkinson's disease



Multiple System Atrophy

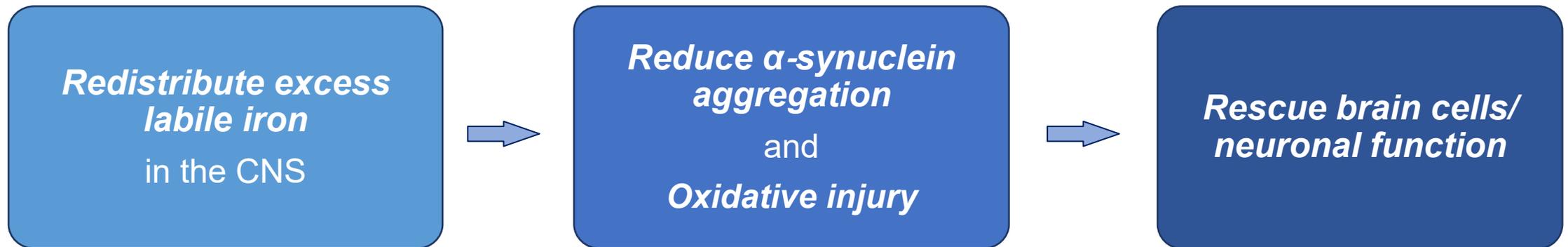


Advanced Quantitative MRI to measure brain iron



Courtesy of P. Trujillo, D. Claassen

◆ **Treatment Approach: Address Underlying Pathology**



Potential Disease Modifying Therapy for MSA

◆ ATH434: Potential Disease Modifying Therapy

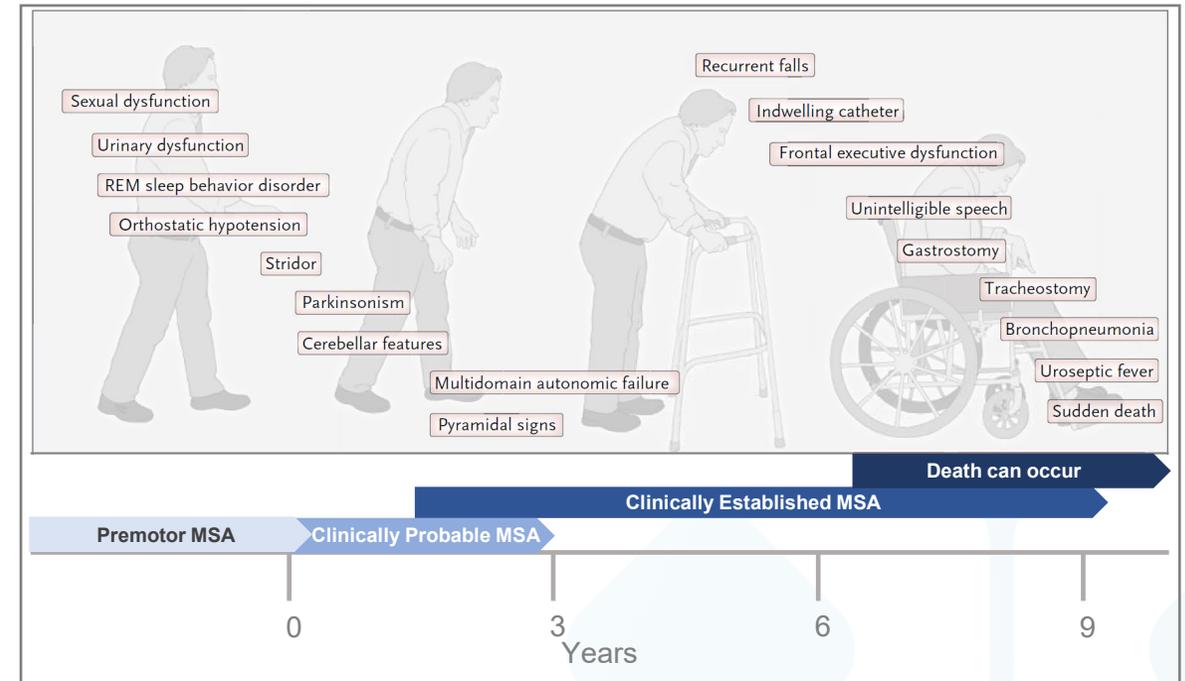
- Small molecule drug candidate
 - Decreases pathologic protein aggregation (α -synuclein) in CNS
 - Iron chaperone: redistributes excess labile iron in CNS
- Oral medication
 - Preferred over infusions and injections
- Potential to treat iron-related neurodegenerative disease
- Orphan Drug Designation in the US and EU for MSA treatment
- Development pathway endorsed by FDA and EMA

ATH434 binding to iron



◆ Multiple System Atrophy (MSA): Rare, Highly Debilitating and Rapidly Progressive Neurodegenerative Disease

- Parkinsonian disorder with no approved treatment
- Orphan disease: up to 50,000 patients in U.S.
- Disease characteristics
 - Motor: Parkinsonism, uncoordinated movements, balance problems and falls
 - Autonomic dysfunction: blood pressure maintenance, bladder control, bowel function
 - Brain atrophy and α -synuclein accumulation in multiple regions
- Over 50% require wheelchair in 5 years
- Median survival 7.5 years after symptom onset



◆ Ongoing Treatment Studies with ATH434

Study	ATH434-201 – Phase 2	ATH434-202 – Phase 2
Design	Randomized, double-blind	Single arm, open-label
Objectives	Efficacy and safety of ATH434	Efficacy and safety of ATH434
Population	Early-stage MSA	Advanced MSA
Sample Size	N = 77	N = 10
Treatment	12 months	12 months
Brain MRI Biomarkers	Iron, volume	Iron, volume, glial pathology
Fluid Biomarkers	NfL [^] , Aggregated α -synuclein	NfL [^] , Aggregated α -synuclein
Other Biomarkers	Wearable movement sensors	—
Clinical Measures	Motor exam, autonomic function, activities of daily living, global measures	Motor exam, autonomic function, activities of daily living, global measures

[^]NfL: Neurofilament light chain

◆ ATH434-202 Interim Clinical Data Confirms Development Approach

Population: 10 with advanced MSA

Treatment: ATH434 75 mg bid

Assessment: 6 months

Safety: ATH434 well-tolerated and had no serious adverse events related to study drug

UMSARS: Unified MSA Rating Scale (Part I)

Activities of Daily Living Scale

- Group change (n=10)[^] compares favorably to patients with similar disease severity*

Global Impression of Change

- Overall rating from patient or clinician perspective

Clinical Responders

- 30% of participants showed stable or improved neurological symptoms, based on UMSARS I and Global Impression of change scores

Areas Assessed

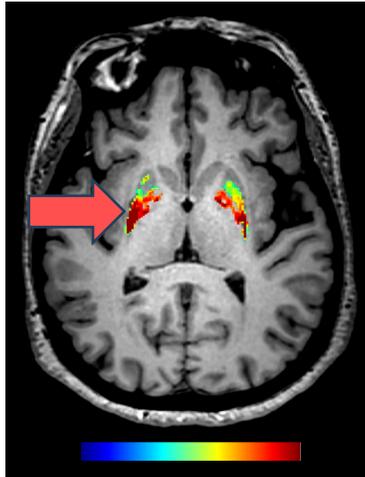
- | | |
|----------------|------------------------|
| • Speech | • Walking |
| • Swallowing | • Falling |
| • Handwriting | • Orthostatic symptoms |
| • Cutting food | • Urinary Function |
| • Dressing | • Sexual Function |
| • Hygiene | • Bowel Function |

[^] Claassen. Mov. Disord. 2024.

* Wenning. Lancet Neurol 2013;12: 264–74.

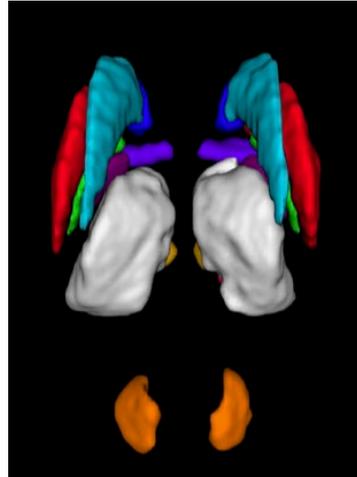
◆ ATH434-202 Interim Biomarker Data Support Clinical Observations

Brain Iron



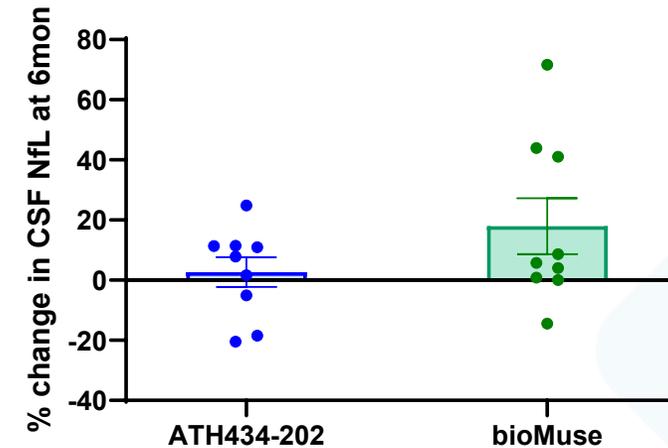
Iron content was stable in the s. nigra, putamen and globus pallidus over 12 mo in clinical responders

Brain Volume



Brain volume stable between 6 and 12 mo in clinical responders

Neuronal Injury Marker



Mean levels of NfL, a marker of neuronal injury only ↑2.7% in ATH434-patients compared to ↑17.9% in untreated patients (bioMUSE)

Participants who stabilized or improved had less advanced disease

◆ Target Measures for Success

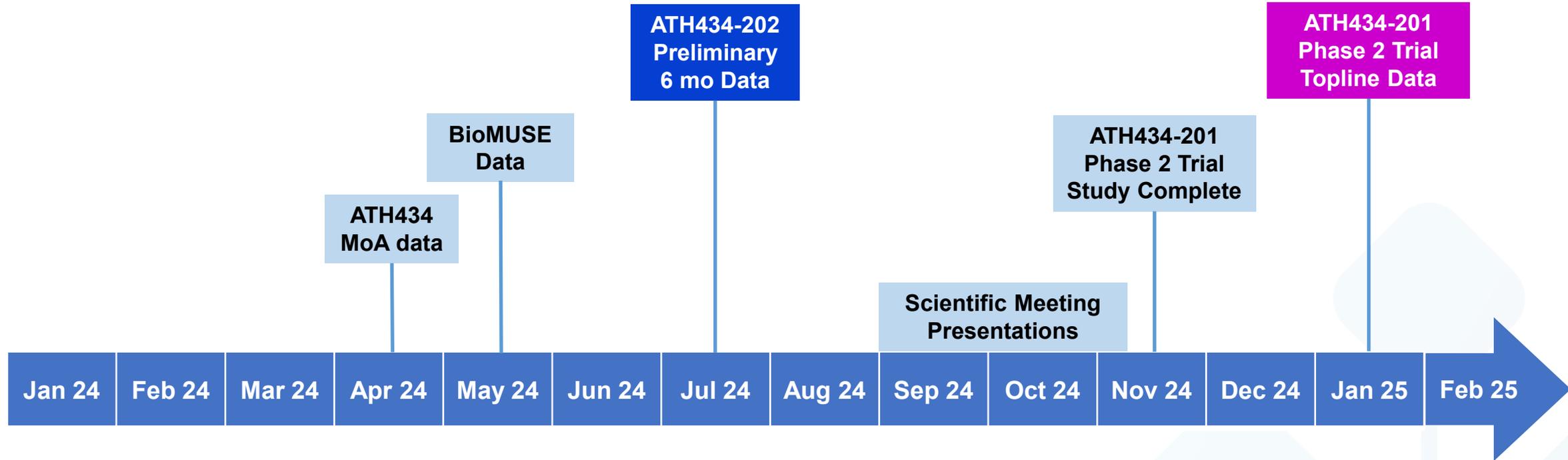


**Favorable Safety Profile /
No Drug Related Serious Adverse Events**

**Evidence of Drug Effect on Biomarker Endpoint or
Evidence of Benefit on Clinical Endpoint Relevant to MSA**

Safety and Efficacy Data Support Advancing to Phase 3

◆ Key Milestones



◆ Alterity: Poised for Progress

- Targeting Orphan disease with no approved treatments
- Natural History Study de-risked Phase 2
- Positive interim data in open label study confirms clinical development approach
- Phase 2 double-blind data expected in January 2025
- Development team with multiple FDA approvals
- Cash balance of AU\$9.28M as of 30 Sept 2024

Catalysts

MSA Natural History Study

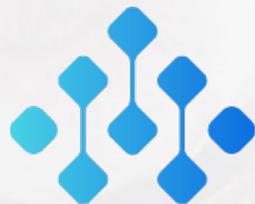
- ✓ H1 2024: Present new biomarker data

ATH434-201 Phase 2 Double-Blind Trial

- ✓ Nov 2023: Enrollment Complete
- Nov 2024: Study Complete
- Jan 2025: Topline Data

ATH434-202 Phase 2 Open label Trial

- ✓ Jul 2024: Preliminary 6-mo Data
- H1 2025: 12-month Data



Alterity

THERAPEUTICS

ASX:ATH | NASDAQ:ATHE