



**brainstorm**  
cell therapeutics

# BrainStorm Cell Therapeutics

NASDAQ: BCLI

Ralph Kern MD MHSc

COO/CMO

February 11, 2019

# Forward Looking Statement

Statements in this announcement other than historical data and information constitute "forward-looking statements" and involve risks and uncertainties that could cause BrainStorm Cell Therapeutics Inc.'s actual results to differ materially from those stated or implied by such forward-looking statements. Terms and phrases such as "may", "should", "would", "could", "will", "expect", "likely", "believe", "plan", "estimate", "predict", "potential", and similar terms and phrases are intended to identify these forward-looking statements. The potential risks and uncertainties include, without limitation, risks associated with BrainStorm's limited operating history, history of losses; minimal working capital, dependence on its license to Ramot's technology; ability to adequately protect the technology; dependence on key executives and on its scientific consultants; ability to obtain required regulatory approvals; and other factors detailed in BrainStorm's annual report on Form 10-K and quarterly reports on Form 10-Q available at <http://www.sec.gov>.

These factors should be considered carefully, and readers should not place undue reliance on BrainStorm's forward-looking statements. The forward-looking statements contained in this press release are based on the beliefs, expectations and opinions of management as of the date of this press release. We do not assume any obligation to update forward-looking statements to reflect actual results or assumptions if circumstances or management's beliefs, expectations or opinions should change, unless otherwise required by law. Although we believe that the expectations reflected in the forward-looking statements are reasonable, we cannot guarantee future results, levels of activity, performance or achievements.

# Brainstorm at a Glance

Clinical stage biotechnology company developing innovative autologous cellular therapies for neurodegenerative disease

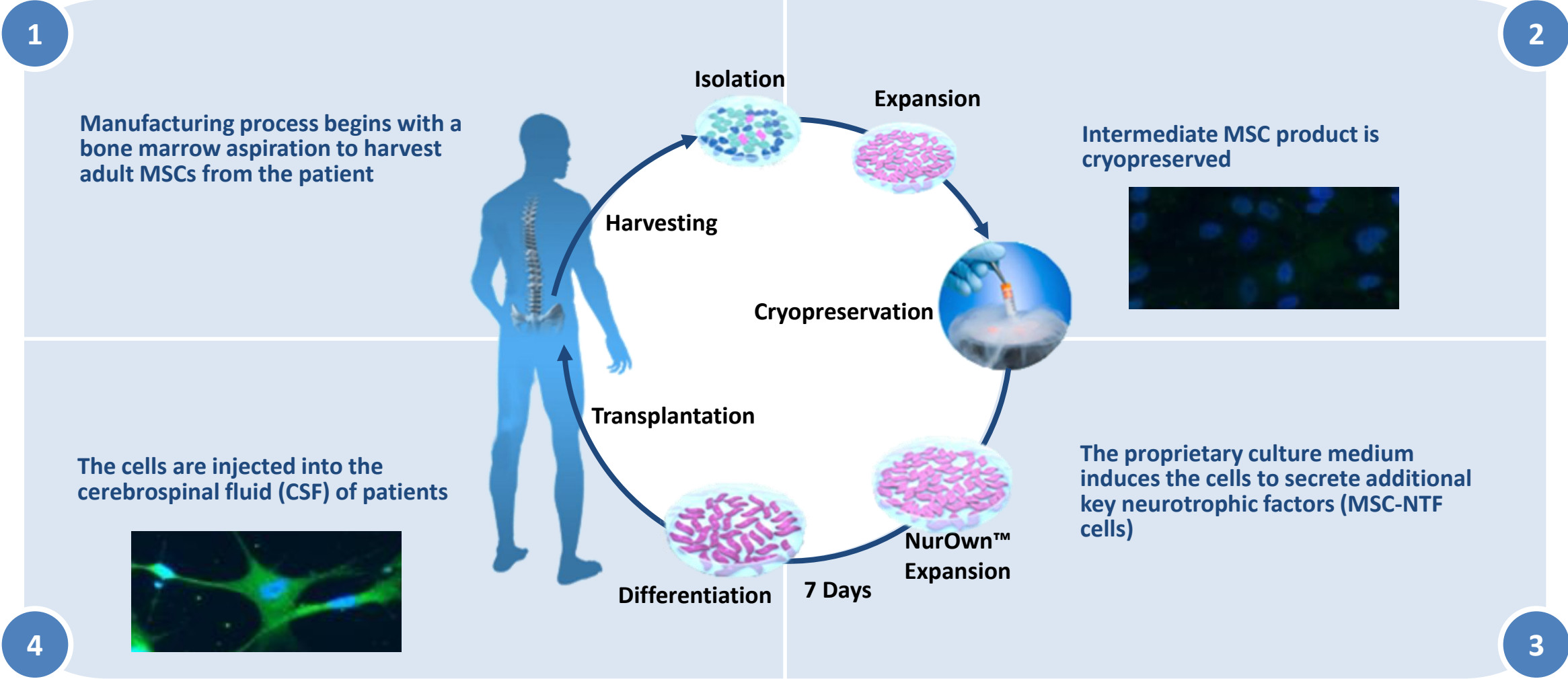
Actively recruiting phase 3 clinical trial in ALS, an orphan disease with no existing cure and limited treatment options

Large addressable markets – progressive MS (500K US) and ALS (30K US)

Proven technology platform and strong manufacturing capabilities

Robust IP portfolio

# NurOwn<sup>®</sup> Process



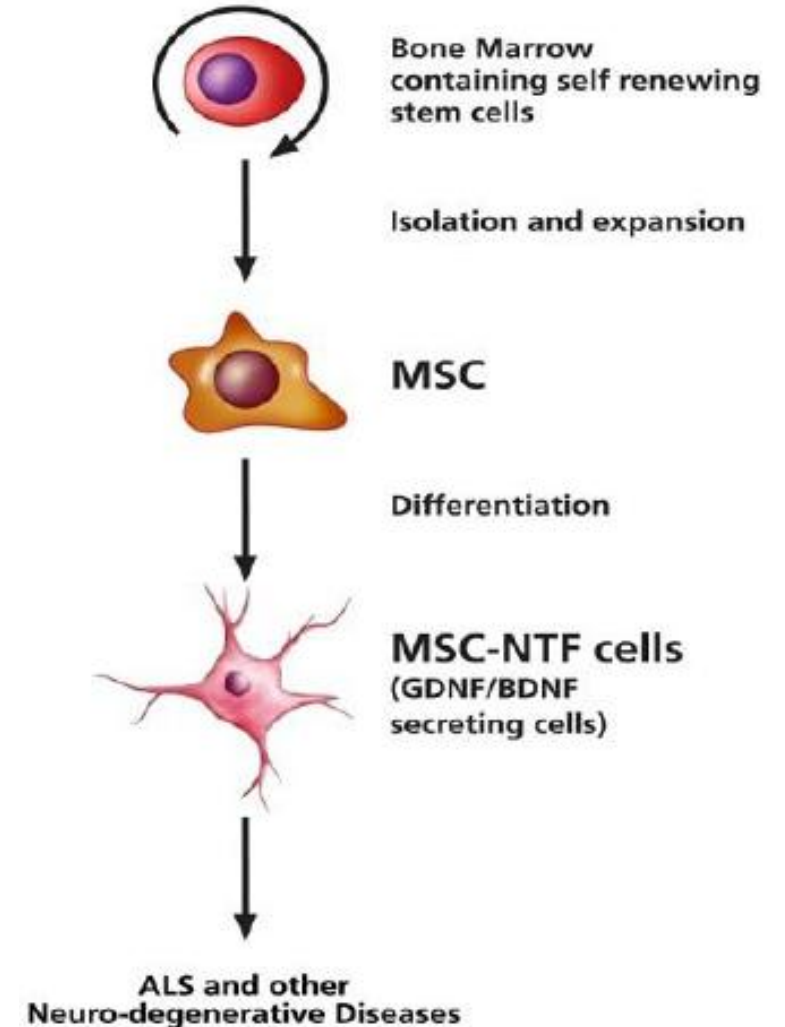
# NurOwn® Technology Platform

Autologous MSCs derived from bone marrow

Intermediate cryopreserved MSC product allows for 3 years of treatment from a single bone marrow aspiration

Final differentiated product are NTF-secreting MSCs

Dual mechanism of action- immunomodulation and neuroprotection through secreted NTFs and anti-inflammatory cytokines



# Pipeline

ALS – Amyotrophic Lateral Sclerosis (Lou Gehrig’s Disease)			
Preclinical	Phase 1	Phase 2	Phase 3

Progressive MS – Multiple Sclerosis			
Preclinical	Phase 1	Phase 3	

Autism Spectrum Disorder			
Preclinical	Phase 1	Phase 2	Phase 3

Parkinson’s Disease			
Preclinical	Phase 1	Phase 2	Phase 3

Huntington’s Disease			
Preclinical	Phase 1	Phase 2	Phase 3

Peripheral Nerve Injury			
Preclinical	Phase 1	Phase 2	Phase 3

# ALS Incidence

*It is estimated that **450,000** people worldwide are living with ALS, with over **30,000** people in the U.S. suffering from this neurodegenerative condition at any given time.*

Someone is diagnosed  
with ALS every

**90**

minutes

New US cases per year

**6,000**

**\$6B**

annual health care cost

**30**

months  
Life expectancy from diagnosis

## What is ALS?

*Amyotrophic lateral sclerosis (ALS), also known as Lou Gehrig's disease <sup>(1)</sup>, is a progressive neurodegenerative disease that affects the function of nerves and muscles. This debilitating condition usually affects people aged 40 to 70, however, individuals in their 20s and 30s have also been known to develop ALS.*



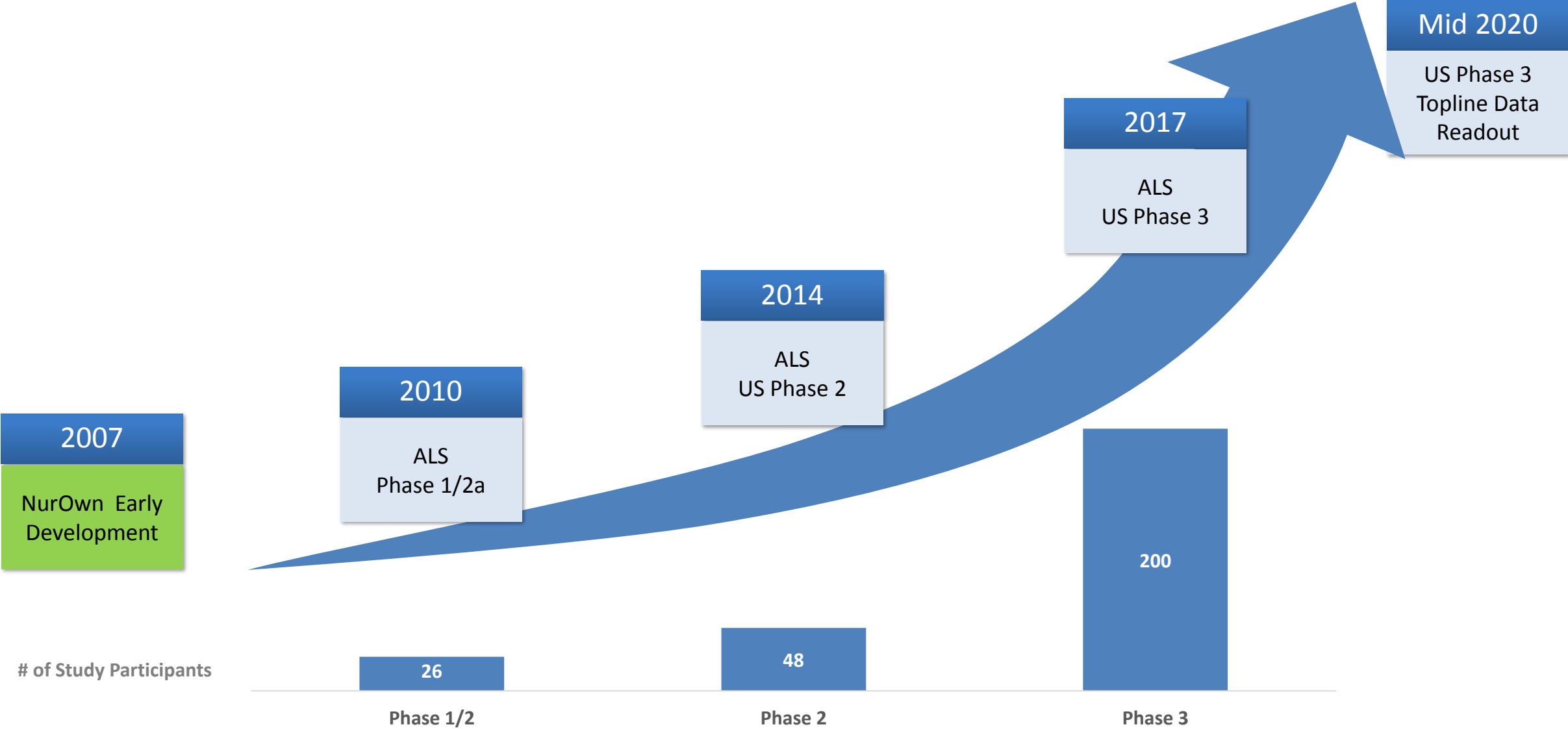
## Future Predictions

*Researchers have predicted that the number of worldwide ALS cases will increase by 69% in 2040, compared to 2015. The main cause of this projected increase is due to an ageing population, particularly in developing nations.*

(1) Lou Gehrig was a famous American baseball player diagnosed with ALS in 1939.

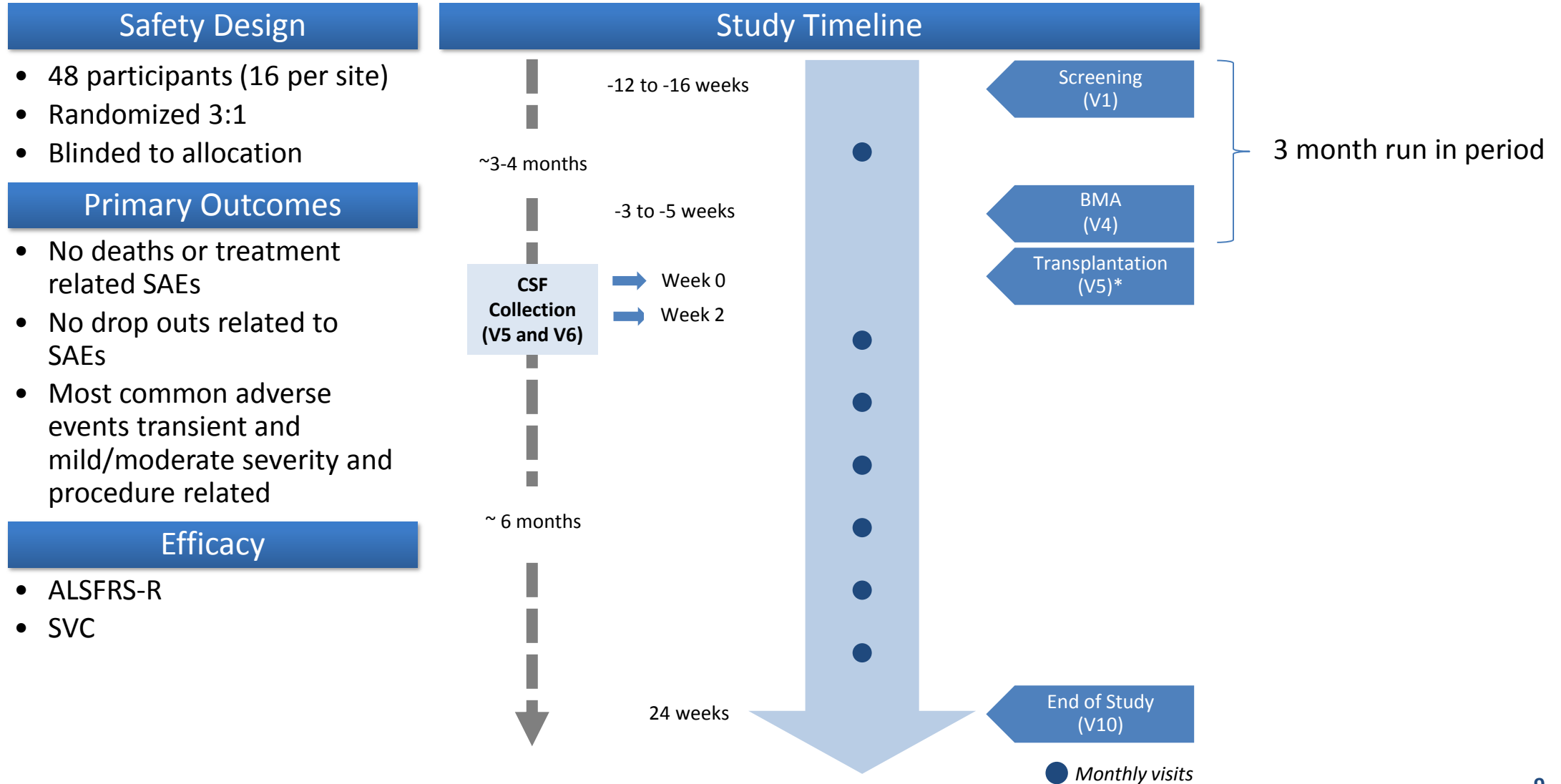


# NurOwn<sup>®</sup> ALS Development





# ALS Phase 2 Trial Design and Safety Outcomes



# Phase 2 Trial – Approach to Data Analysis

## Phase 2 Trial Efficacy Analysis

Mean change in  
ALSFRS-R slope

ALSFRS-R  
Responder  
Analysis

Rapid progressor  
subgroup  
prespecified

## ALS Rapid Progressor Subgroup

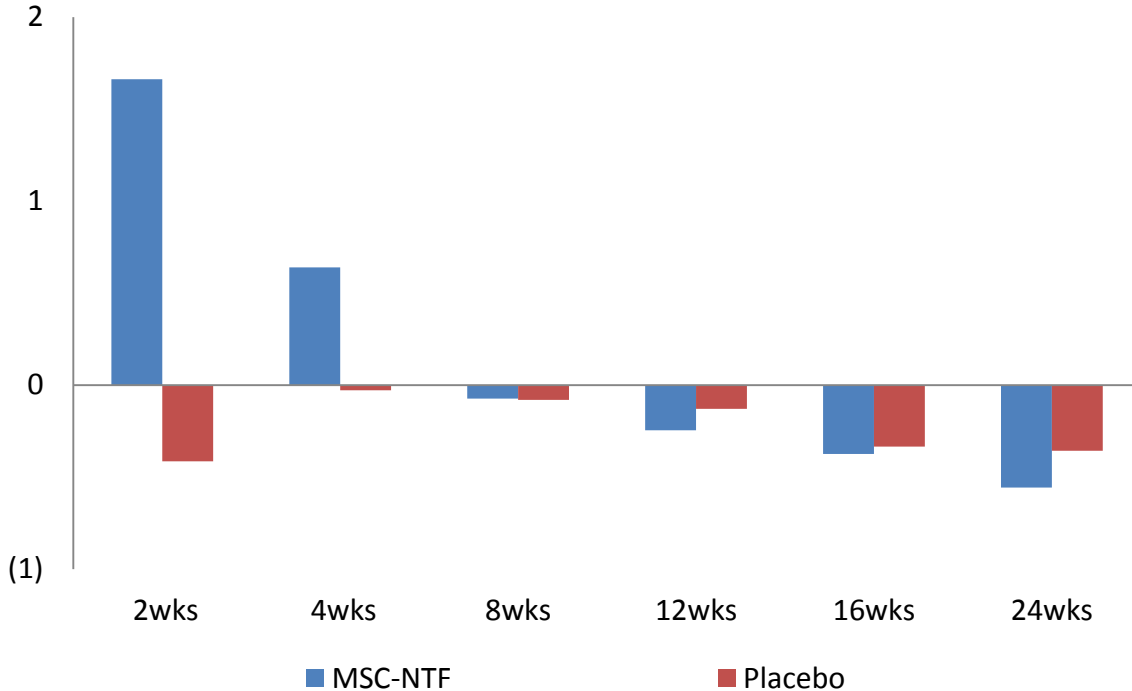
- ✓ Higher correlation of ALSFRS-R change with survival
- ✓ Higher response rate
- ✓ Higher levels of CSF inflammatory biomarkers

# ALSFRS-R slope improvement (mean pre-post LS mean slope change/month)

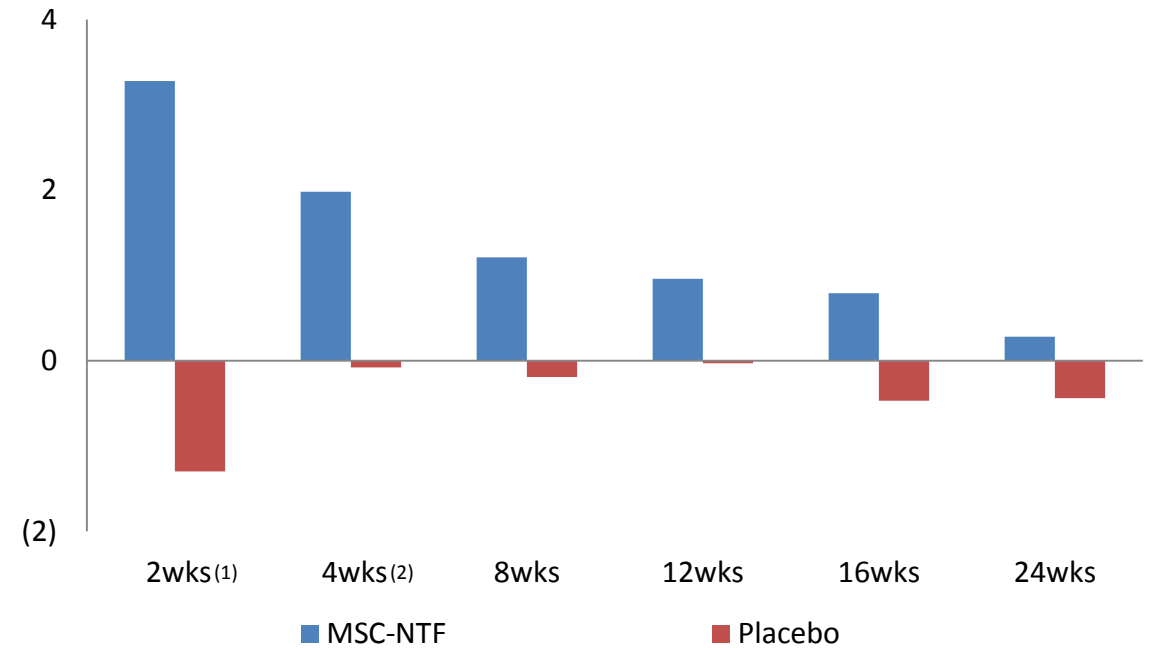
All participants (n=46)

Rapid progressors (n=21)

ALSFRS-R LS mean slope change



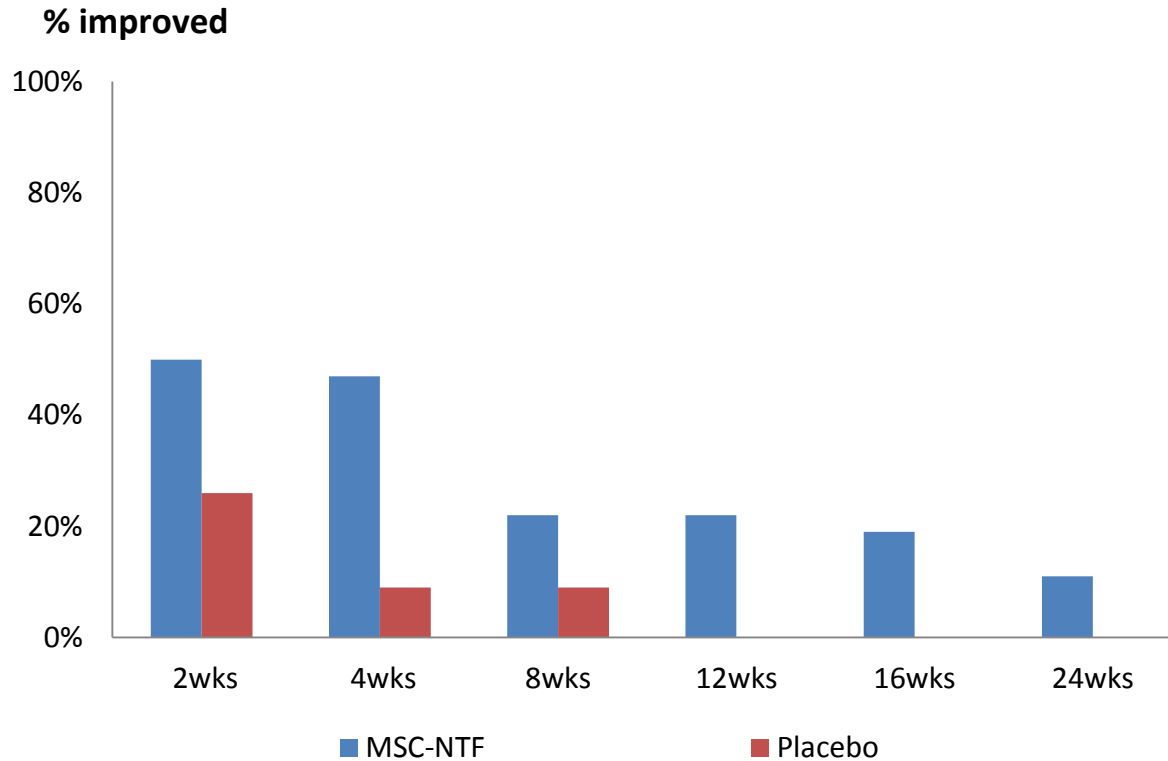
ALSFRS-R LS mean slope change



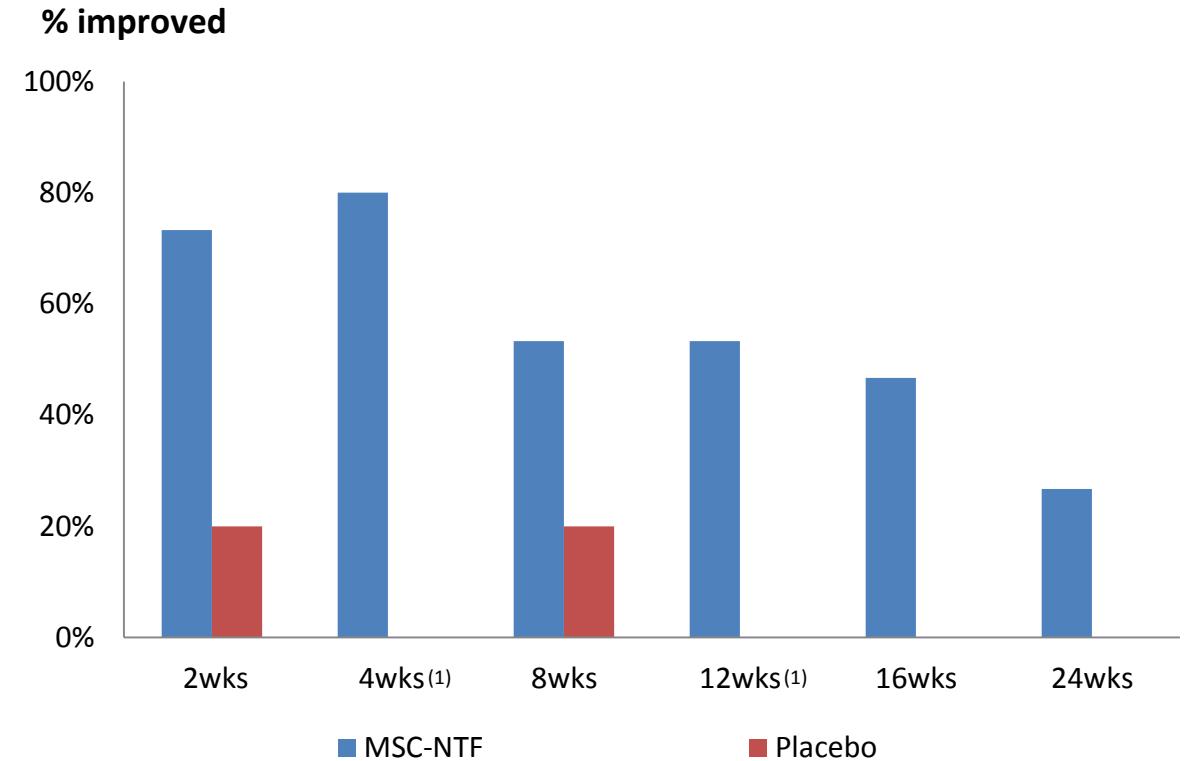
1.  $p=0.021$   
2.  $P=0.033$

# Responder Analysis: $\geq 1.5$ points/month ALSFRS-R slope improvement

All participants (n=46)



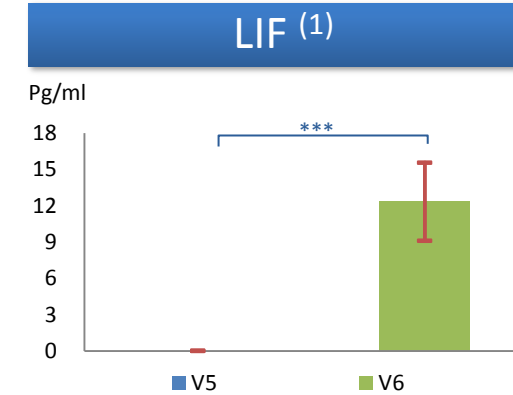
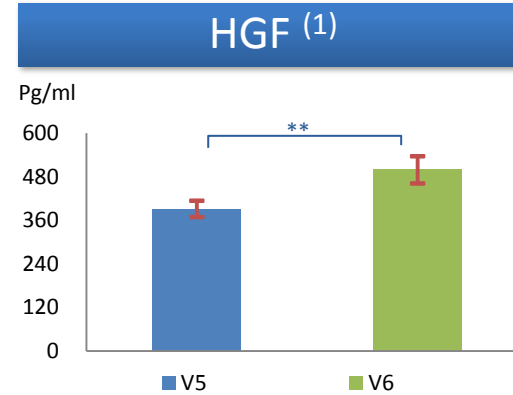
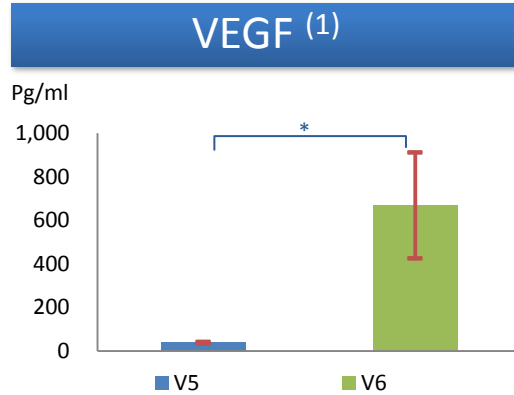
Rapid progressors (n=21)



1.  $p < 0.05$  (two-sided Fisher's exact test)

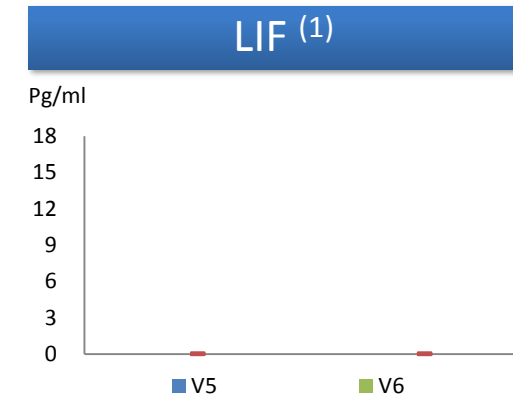
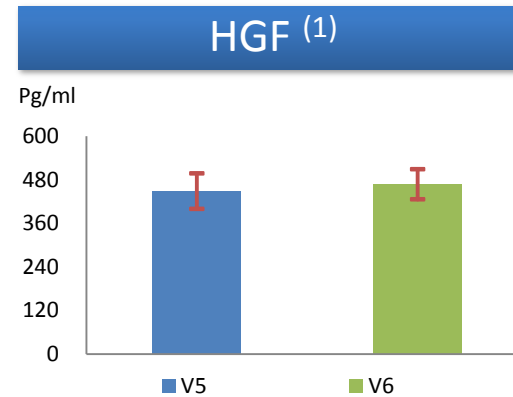
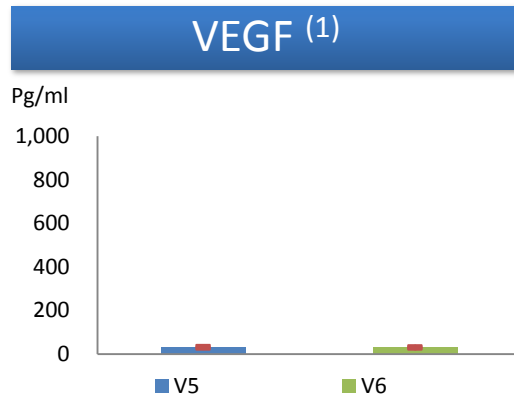
# NurOwn® Phase 2 ALS Biomarkers: CSF NTFs Significantly Increased 2 Weeks Post- Treatment Compared to Baseline

**NurOwn®**  
(n = 26)



■ Pre transplantation  
■ Post transplantation

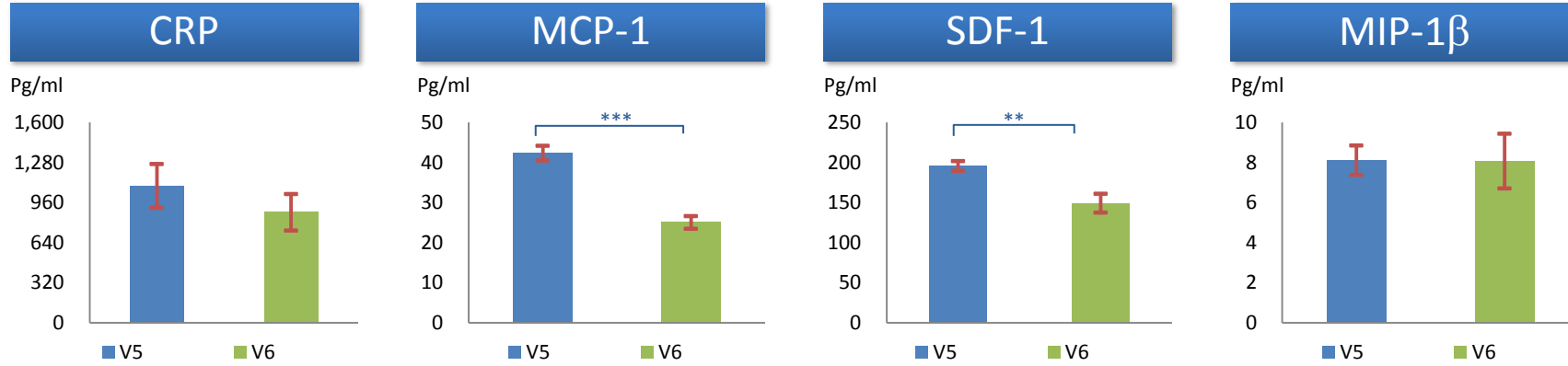
**Placebo**  
(n = 9)



1. Mean ± SEM. p < 0.05, p < 0.01, p < 0.001 for VEGF, HGF and LIF, respectively.

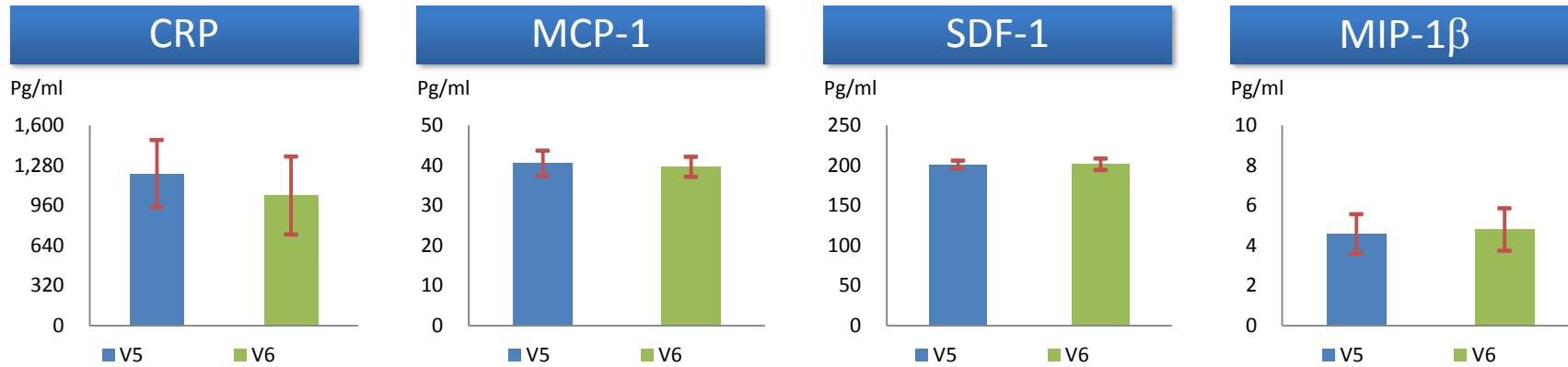
# NurOwn<sup>®</sup> Phase 2 ALS Biomarkers: CSF Inflammation Significantly Decreased 2 weeks Post-Treatment Compared to Baseline

**NurOwn<sup>®</sup>**  
(n = 26)



■ Pre transplantation  
■ Post transplantation

**Placebo**  
(n = 9)



1. Mean ± SEM.  $p < 0.01$ ,  $p < 0.001$  for [], respectively.

# NurOwn® Phase 2 ALS Trial Summary

Intrathecal treatment was safe and well tolerated

Repeat dosing is needed

Rapid progressor subgroup showed higher efficacy

Biomarkers confirmed mechanism of action



# NurOwn<sup>®</sup> ALS Phase 3 Clinical Trial

## Pre-Treatment

- Inclusion criteria
  - Less than 60 years of age
  - SVC > 65%
  - ALS ≤ 2 years
  - Rapid progressors
- Exclusion criteria
  - Edaravone
  - Ventilation
  - Feeding tube
- Randomization
- Bone Marrow Aspiration

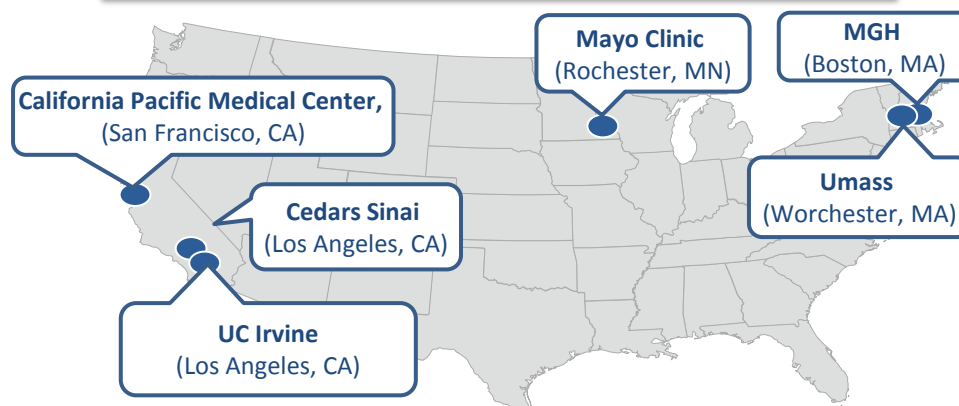
## Treatment

- N=200 patients
  - Enrollment completed by mid-2019
- 1:1 randomization
- Study duration: 11.5 months
  - Seven months post-first transplantation
- Top-line data expected mid 2020

## Outcomes

- ALSFRS-R responder analysis (post-treatment slope change)
- Safety
- ALSFRS-R change from baseline
- SVC
- Tracheostomy-free survival
- CSF/biomarkers (seven samples over six months)

## Site Location



# Progressive MS Unmet Need

US MS prevalence

**1M**

Progressive MS

**50%**

***Women***

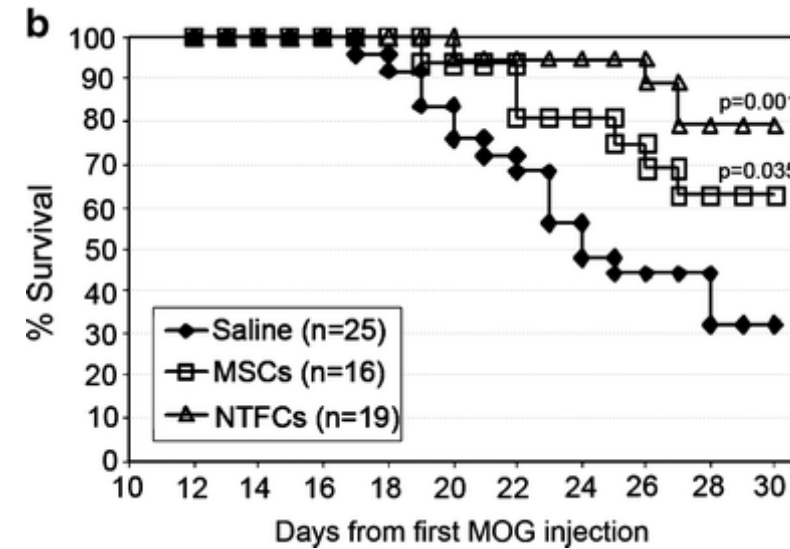
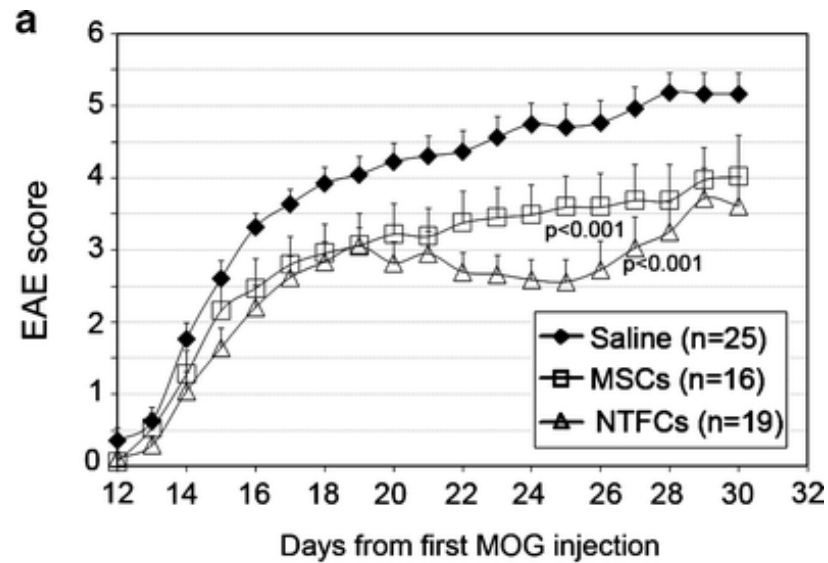
are more likely to be diagnosed  
with MS than men

**\$4.3B**

*US annual economic burden*

# MS Preclinical

## EAE Model



Intracerebroventricular injection of  $2.5 \times 10^5$  cells

# MS Phase 2 Clinical Trial

IND granted December 2018

Repeat dosing intrathecal MSC-NTF cells (NurOwn®)

5 leading US MS centers

Validated MS outcomes and biomarkers

First patient to be treated Q1 2019, top line data mid 2020

# Financial Highlights

\$US in Thousands <small>(except for Shares, Warrants and Options data)</small>	September 30, 2018
Cash and cash equivalents	697
Short-term deposit	10,194
Non-Dilutive Grants <sup>(1)</sup>	8,000
<b>Total Liquidity</b>	<b>18,891</b>
Cap Structure	
Debt	-
Shares Outstanding <small>(including outstanding Warrants and Options)</small>	20,700,713
Options / Warrants outstanding	1,457,620 / 4,578,867
Net Operating Loss	88,640

(1) Supported by \$16 million CIRM and \$2 million Israel Innovation Authorities grants

# BrainStorm Management Team

**Chaim Lebovits**  
President and CEO

**Ralph Kern, MD, MHSc.**  
COO and CMO

**Eyal Rubin**  
EVP, CFO

**Mary Kay Turner**  
VP, Patient Advocacy and Government Affairs

**Joe Petroziello BS MS**  
VP, Scientific and Corporate Communications

**Yael Gothelf PhD.**  
VP, Scientific and Regulatory Affairs

**Uri Yablonka**  
EVP, Chief Business Officer

**Yossef Levy PhD.**  
VP, Cell Production

**Revital Aricha PhD.**  
VP, R&D

**Susan Ward PhD**  
Head Clinical Research Operations



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# BrainStorm Cell Therapeutics

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**Thank You**



# Appendix

# Additional Professional Board Members

## June S. Almenoff MD PhD

- Chief Operating Officer & Chief Medical Officer of Innovate Biopharmaceutical
- Formerly President and CMO of Furiex Pharmaceuticals and Board member of Tigenix NV (TIG)

## Tony Polverino PhD

- CSO - Kite Pharmaceuticals

# Scientific Advisory Board

## Jerold Chun, M.D., PhD -Chair

- Neuroscientist, Professor and Senior Vice President of Neuroscience Drug Discovery, Sanford Burnham Prebys Medical Discovery Institute, San Diego CA.

## Stanley H. Appel, M.D

- Peggy and Gary Edwards Distinguished Endowed Chair for the Treatment and Research of ALS, Department of Neurology, Neurological Institute, Houston Methodist Hospital, Houston TX.

## Amit Bar-Or, M.D.

- Presidential Endowed Chair at the University of Pennsylvania (UPenn/CHOP), Director of the Centre for Neuroinflammation and Experimental Neurotherapeutics and Chief, MS Division, Philadelphia PA.

# IP Portfolio

Patent Name/ Int. App. No.	Pending Jurisdictions	Allowed Jurisdictions	Granted Jurisdictions	Expiry Date
ISOLATED CELLS AND POPULATIONS COMPRISING SAME FOR THE TREATMENT OF CNS DISEASES/PCT/IL2006/000699	US		Europe, US	2030
MESENCHYMAL STEM CELLS FOR THE TREATMENT OF CNS DISEASES PCT/ IL2009/000525		Hong Kong	US, Europe, Israel	2032
METHODS OF GENERATING MESENCHYMAL STEM CELLS WHICH SECRETE NEUROTROPHIC FACTORS / PCT/IL2013/050660	Europe, Hong Kong, Israel, Canada, Brazil, Japan	Israel	US, Japan	2038
METHOD OF QUALIFYING CELLS /PCT IL2015/050159	US, Europe, Hong Kong, Israel, Canada, Brazil, Japan			2040
Methods of treating ALS PCT/IL2017/050801	PCT			2042