

Enabling the Nervous System to Repair Itself

CORPORATE PRESENTATION

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NervGen's Clinical Trials are Led by Experienced Drug Developers



Bill Radvak, BASc Executive Chairman & Interim CEO

- Co-founder of NervGen
- Previously CEO and Director of multiple start-up companies
- Founder and CEO of Response
 Biomedical, a publicly listed medical
 device company, which he led from its
 inception to a 90-employee sales and
 manufacturing company



Adam Rogers, MD
Interim President
& Board Member

- Co-founded Hemera Biosciences in 2010, a clinical stage gene therapy biotech company and assumed the role of CEO in 2017
- Oversaw all aspects of Hemera until the assets were acquired in December 2020 by Janssen Pharmaceuticals, a subsidiary of Johnson & Johnson
- Principal of Boston based PFP
 Biosciences Holdings and a boardcertified ophthalmologist



Dan Mikol, MD, PhD
Chief Medical Officer

- 25+ years pharma experience and as practicing neurologist conducting clinical research
- Joined NervGen from Amgen where he was Executive Director and Global Therapeutic Development Head, Neurology and Nephrology
- Participated in development and/or commercialization of natalizumab (Tysabri), fingolomod (Gilenya), cladribine (Mavenclad), interferon-ß-1a and erenumab (Aimovig)













NervGen Highlights

NVG-291, a first-in-class *neuroreparative* drug candidate administered by subcutaneous injection, has the potential to **redefine treatment paradigms** by **repairing nervous system damage**

Improvement demonstrated across 6 different animal models in fine and gross motor control, sensory function, autonomic functions, visual acuity, memory & learning, in many cases **unprecedented**

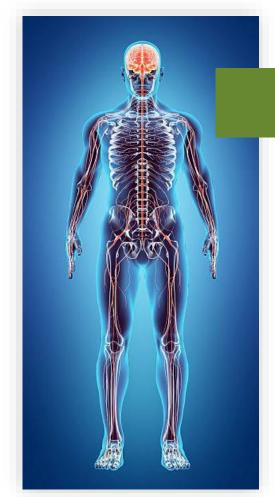
Target indications address **very attractive commercial opportunities** with **significant unmet medical needs** in spinal cord injury, Alzheimer's, multiple sclerosis and stroke

Nerve repair mechanism allows for low cost and short duration clinical trials

Phase 1b/2a clinical trial for individuals with acute and chronic spinal cord injury to be initiated in 2023 and readout in the first half of 2024



Revolutionizing the Treatment of Nervous System Damage

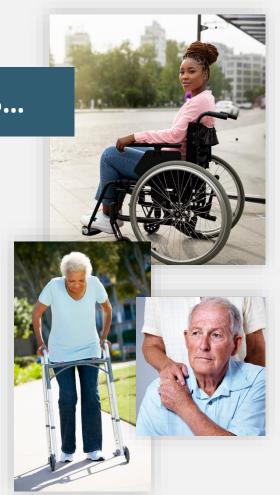


Everyone **KNOWS...**

The nervous system is a **complex system** that controls thought, movement, senses, etc.

Everyone **BELIEVES...**

The nervous system cannot repair itself





NVG-291 – First-in-Class *Neuroreparative* Drug

Representative of Placebo Group

(Back Legs and Tail Dragging)



Representative of NVG-291 Group

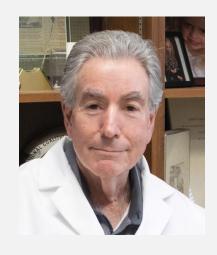
(Back Legs and Tail Active)



Remarkable and robust repair across multiple models



NervGen's Technology Was Invented by Dr. Jerry Silver Known in the Spinal Cord Injury Field as the "Oracle"



Jerry Silver, PhD

Professor and Researcher,



Adjunct Professor,



Dr. Silver's Spinal Cord Research

- Discovered why the nervous system does not repair itself
- Identified the surprising molecules responsible

Dr. Silver Has Received Numerous Prestigious Awards Including

- Ameritec Prize
- Christopher Reeve-Joan Irvine Research Medal
- Jacob Javits Neuroscience Investigator Award

Dr. Silver's research revolutionized the understanding of the nervous system

STRONG IP PORTFOLIO

NervGen licensed the technology from Case Western and

owns global rights for all indications

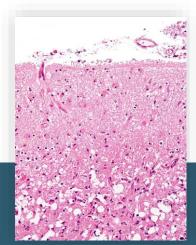
Intellectual property protection on NVG-291 until 2037



The Foundation of Our Technology

Pre 1990

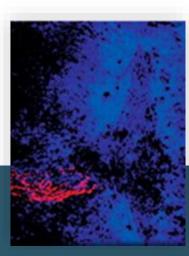
It was demonstrated that **glial scars** form at the site of injury to the nervous system and that scars in the brain cause neurons to be dysfunctional. Scars were later identified as the primary impediment of recovery



Micrograph of a glial scar

1990s

Dr. Silver identified a class of molecules called **CSPGs**, present in scars in the brain and spinal cord, that stop the body's natural repair mechanisms



Spinal cord nerve (red) trapped in the scar by CSPGs (blue)¹

2009

Dr. Silver and collaborators from Harvard co-discovered that CSPGs bind to **PTPo**, a receptor present in the brain and spinal cord and that this interaction stops cells from repairing damage

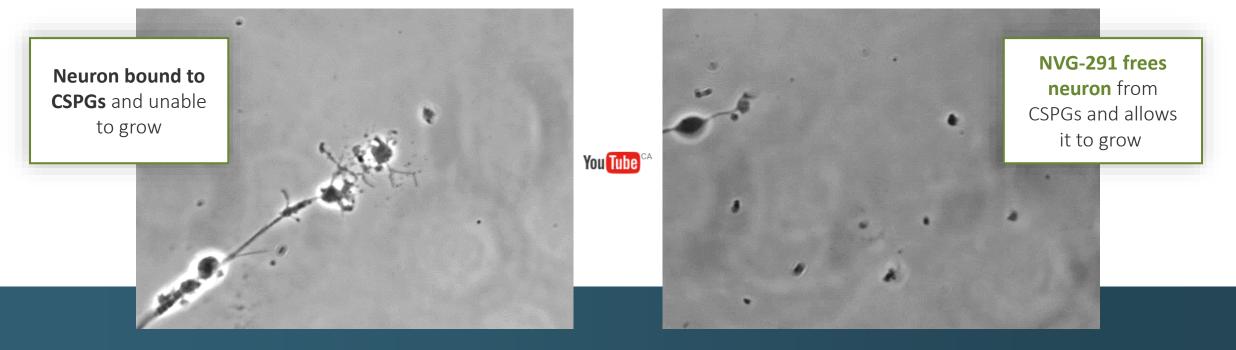


2015

Dr. Silver's team designed a peptide derived from PTPo and shown to relieve CSPG-mediated inhibition of nervous system repair. **NVG-291** enables the nervous system to repair damage by inhibiting CSPG signaling.



NVG-291 Allows Neurons to Grow in the Scar



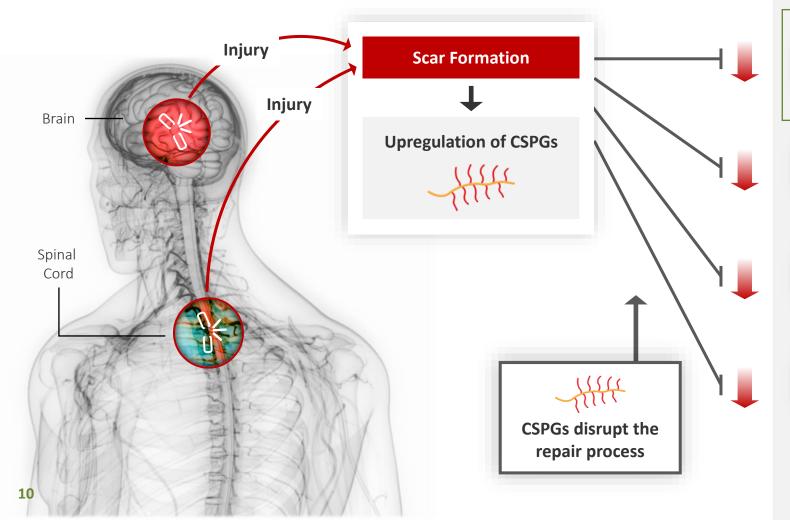
NVG-291, a 35 amino acid peptide, produced dramatic recovery in a spinal cord injury animal study: the results published in Nature¹ are now cited in over 327 publications

Administered systemically by a daily subcutaneous injection

Includes a transporter that facilitates crossing the blood brain barrier



The Body's Powerful Innate Repair Mechanisms Disrupted by CSPGs



Repair Mechanisms:



Plasticity

The creation of new neuronal connections and rewiring of existing ones



Axonal Regeneration

The ability of a severed axon to reestablish connectivity with other neurons



Remyelination

The process of repairing damaged myelin – the fatty substance that protects axons and enables fast electrochemical transmission



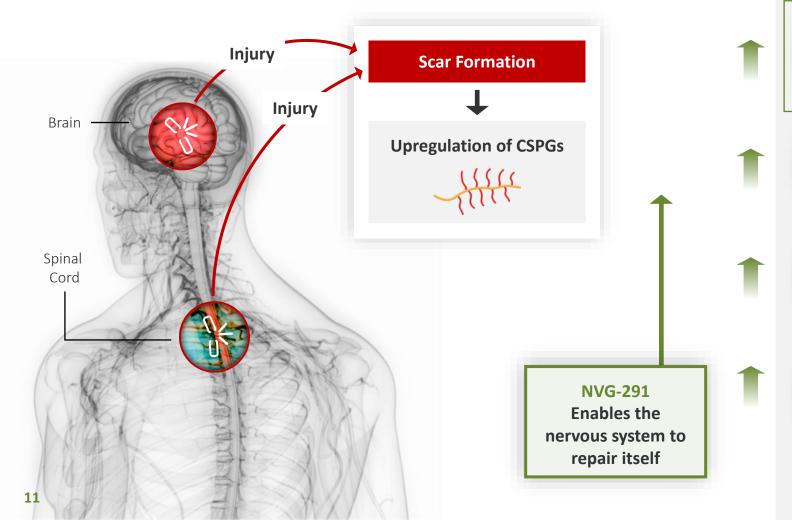
Others

Stem cell preservation/migration Autophagy Microglial shifting



NVG-291

Takes the Brakes off Natural Repair Mechanisms



Repair Mechanisms:



Plasticity

The creation of new neuronal connections and rewiring of existing ones



Axonal Regeneration

The ability of a severed axon to reestablish connectivity with other neurons



Remyelination

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Others

Stem cell preservation/migration Autophagy Microglial shifting



NVG-291 Broadly Restores Function

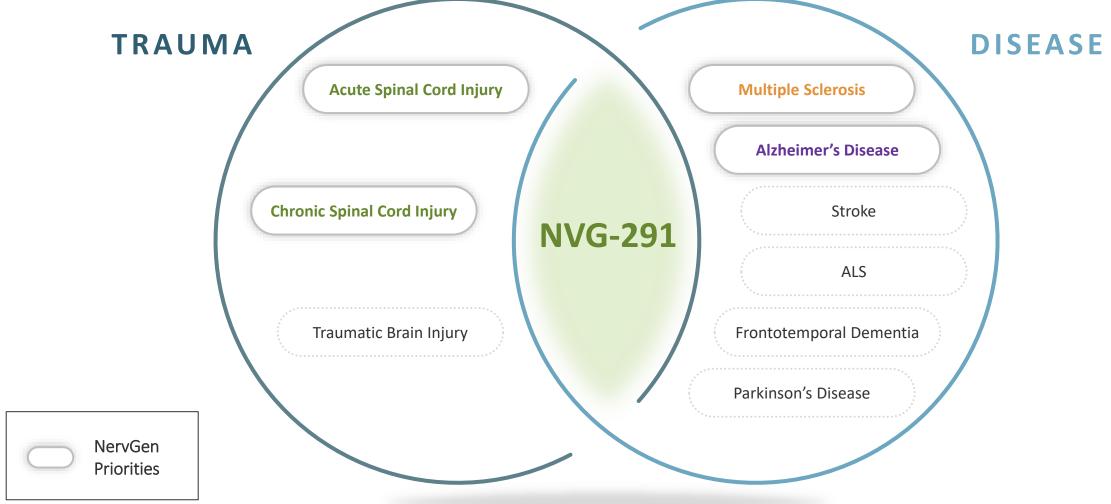
NVG-291 Has Demonstrated Dramatic Repair

in multiple animal models of neurological injury/disease, as documented in 15+ peer-reviewed papers

ACUTE SPINAL CORD INJURY	CHRONIC SPINAL CORD INJURY	PERIPHERAL NERVE INJURY	MULTIPLE SCLEROSIS	OPTIC NEURITIS	STROKE
MotorSensoryBladder	• Motor	MotorSensory	• Motor	• Visual	 Motor Sensory Cognition (object recognition)
 Lang, B.T. et al., Nature, 518, 404–408. (2015). Rink, S. et al., Experimental Neurology, 309, 148–159. (2018). Ham, T.R. et al., Ann Biomed Eng, 47, 744–753. (2019). 	1. Milton et al., bioRxiv, doi:10.1101/2022.08.01.502398 (not peer-reviewed)	 Li, H. et al., Scientific Reports, 5, 1– 14. (2015). Yao, M. et al., Neuropharmacology, 144, 208–218. (2019). 	1. Luo, F. et al., Nature Communications, 9, 1–16. (2018).	Niknam, P. et al., Molecular and Cellular Neuroscience, 99, 103391. (2019).	 Luo et al., Cell Reports Volume 40, Issue 4, 111137, 2022 Yao et al., Journal of Neuroinflammation 19:207, 2022
4. Ham, T.R. et al., Materials Science and Engineering: C, 110, 110656. (2020).					∜NervGen



NVG-291: Potential to Treat All Types of Nervous System Damage





NVG-291 - Phase 1 Clinical Trial in Progress

February 2023



Dosing of All Subjects - COMPLETED

SAD - 37 subjects, MAD* - 33 subjects
*Subjects dosed subcutaneously once a day for 14 days

- NVG-291 was well tolerated at a high dose
 - 170% higher than the equivalent highest dose in preclinical efficacy studies
 - >100x higher than the lowest efficacious dose in preclinical studies
- NVG-291 was rapidly distributed in the blood
- The calculated half-life was longer in humans than animals

Dosing of All Subjects in Phase 1 Clinical Trial completed February 2023





Spinal Cord Injury

Acute and Chronic patients

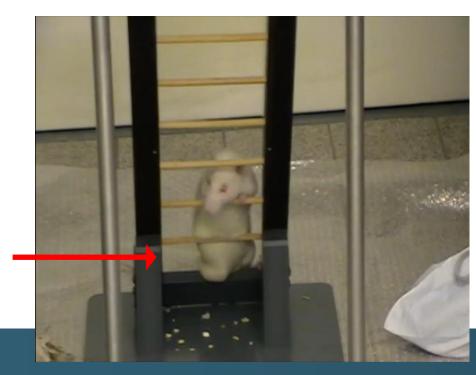
- No FDA Approved Drug that Improves Function
- NervGen's goal is to improve motor, bladder/bowel/sexual and/or sensory function in High unmet need coupled with potential key clinical outcomes may provide opportunity for expedited regulatory approval
- Unprecedented preclinical results



NVG-291 – Dramatically Repairs Spinal Cord Injury

SEVERE SPINAL CORD INJURY MODEL

Representative of Placebo Group





Representative of NVG-291 Group



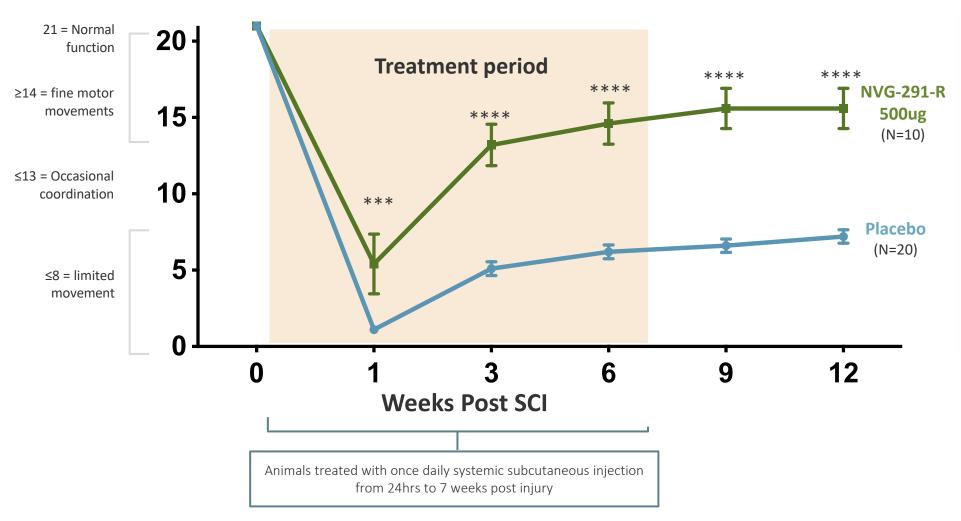
Hind legs are immobile

Significant motor recovery: consistent coordination, toe clearance, tail held high consistently



Spinal Cord Injury – NVG-291-R Promotes Functional Recovery

BBB Scale = Standard measure of mobility



UNPRECEDENTED RESULTS

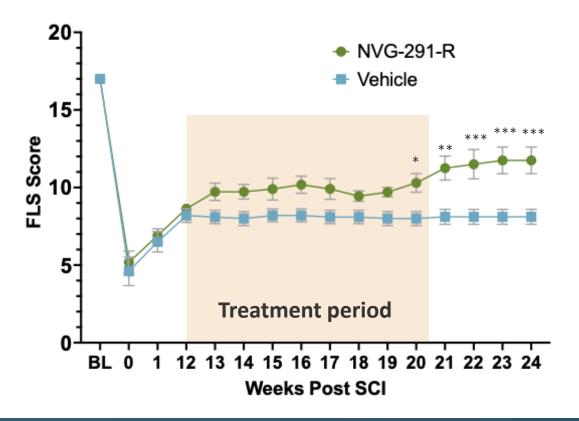
Extremely high response rate

Almost **complete recovery** in
responding animals



Chronic Spinal Cord Injury – NVG-291-R Promotes Functional Recovery

Forelimb Locomotor Scale
Treatment beginning 3 months post spinal cord injury



Animals treated with once daily systemic injection starting 3 months post injury for 60 days

Animals treated with NVG-291-R show **significant improvements** in forelimb recovery at 24 weeks post injury

Significant functional improvements observed when NVG-291-R was administered 3 months after a spinal cord injury



NVG-291 Improves Bladder Function



Bladder function is a key quality of life measure in the paralyzed population



NVG-291 Spinal Cord Injury Clinical Trial

Spinal Cord Injury Phase 1b/2a

~\$10 million estimated cost

Chronic arm: 20 individuals that are 1 – 10 years post injury

> 10 administered NVG-291 and 10 administered placebo

Sub-acute arm: 20 individuals that are 10 - 49 days post injury

> 10 administered NVG-291 and 10 administered placebo

- Placebo-controlled, single center trial
- Once daily subcutaneous injection for 3 months
- Primary endpoint: electrophysiology
- Secondary endpoints: numerous clinical assessments of upper and lower extremity function

World-class Advisory Board with experts in research, clinical design, functional assessments and biomarkers

James Guest MD, PhD, FACS University of Miami	World renowned surgeon/scientist; global expertise in clinical trial methodology	
Linda Jones, PT, PhD Thomas Jefferson University	Expert consultant to pharma, universities, and non-profit organizations	
Steven Kirshblum MD Rutgers New Jersey Medical	Nationally recognized expert; Spinal Cord Medicine textbook editor	

Brian Kwon MD, PhD, FRCSC University of British Columbia	World renowned surgeon/scientist; authored >240 scientific publications, >35 textbook chapters	
Daniel Lammertse, MD University of Colorado School of Medicine	Former Director and President of the American Spinal Injury Association	

Spinal Cord Injury Financial Case

RARE DISEASE PRICING IN A SUBSTANTIAL ADDRESSABLE MARKET

US MARKET			
Acute patients annually	~18,000		
Chronic patients total	~300,000		

REVENUE PROJECTIONS		
Target pricing per course	\$200,000	
Acute: US peak annual revenue	\$500M++	
Chronic: US total revenue	\$10-12 Billion	

- Pricing Drivers:
 - Rare disease, market range \$150-350K/yr
 - Orphan Status in EU
 - Lifetime cost of care ranges from \$1-5M

- Large addressable markets chronic and acute
- Short time to market, potential expedited approval
- Multiple advocacy groups support
- Grant funding available

Attractive annual revenue for Acute bolstered by rapid, large expected revenue from Chronic





Alzheimer's Disease

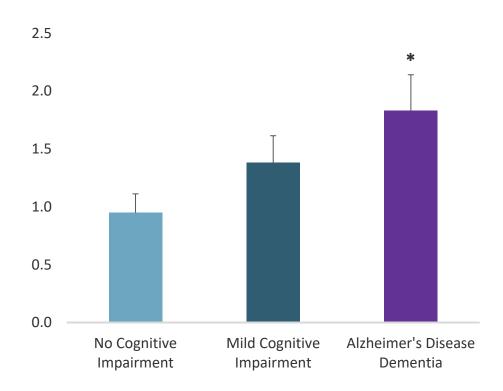
No FDA Approved Drug that Results in Sustained Improvement in Cognitive Function

- Symptomatic treatments to improve cognition in Alzheimer's disease are unsatisfactory
- A disease—modifying therapy approved in 2021 has questionable benefit in slowing cognitive decline
- NervGen's goal is to repair damage and improve cognitive function

NVG-291 Pathway to Treat Alzheimer's Disease

CSPG ACCUMULATION IN AD PATIENT BRAINS¹

(CSPGs) Brevican/GAPDH



Removing PTPo improves cognitive function in Alzheimer's models⁴

NVG-291's multiple modes of action, **plasticity**, **axonal regeneration and remyelination** have the potential to benefit patients suffering from Alzheimer's



Preclinical studies have demonstrated that breaking down CSPGs improves Alzheimer's symptoms^{2,3}

^{*} p<0.05 compared to NCI

^{1.} Howell, M.D. et al., Acta Neuropathol Commun, 3, 54. (2015). ² Yang et al., Experimental Neurology (2015).

³ Vegh et al., Acta Neuropathologica Communications (2014). ⁴ Gu et al., BioRxiv (2016)

NVG-291 Safety/Efficacy Studies in Alzheimer's Disease Patients

Alzheimer's Phase 1b/2a

~80 patients, ~\$20 million estimated cost

- Multicenter, placebo controlled
- 3 months treatment duration
- Efficacy analysis includes functional and structural imaging, cognitive assessments and fluid biomarkers

World-class Advisory Board with experts in research, clinical design, cognitive assessments and biomarkers

Jeffrey Cummings, MD, ScD University of Nevada	Originator, Neuropsychiatric Inventory (NPI)
Martin Farlow, MD Indiana University School of Medicine	Led/contributed to >230 clinical trials; authored 493 peer reviewed research papers and 509 abstracts
Bruce Lamb, PhD Indiana University School of Medicine	World-expert on biological underpinnings of Alzheimer's disease and related dementia
George Perry, PhD University of Texas, San Antonio	Current and founding Editor-in-Chief of the Journal of Alzheimer's Disease

Reisa Sperling, MD Harvard Medical School; Massachusetts General Hospital	Led NIA-Alzheimer's Assoc. guideline development group; Serves on National Institute on Aging Advisory Council
Michael Weiner, MD University of California, San Francisco	Leader in development of MRI and PET for investigating and diagnosing neurodegenerative diseases
Henrik Zetterberg, MD, PhD University of Gothenburg, University College London	World expert in blood-based biomarkers in neurological disorder

NVG-291-R – Improved Spatial Learning and Memory

STROKE MODEL



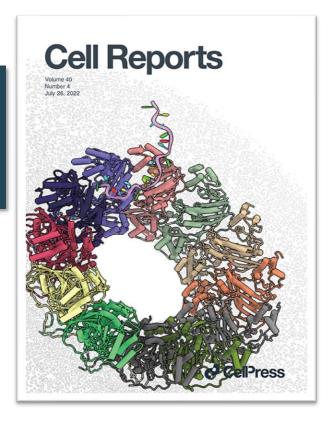
Significant improvement in cognitive function when treated 24 hours - and even 7 days after stroke



NVG-291-R in Stroke – Breakthrough Results in a New Indication

CELL REPORTS - PEER-REVIEWED PUBLICATION

"... Small Canadian
Biotech Could Challenge
Roche's Hold in Stroke
Treatment" – BioSpace
July 2022



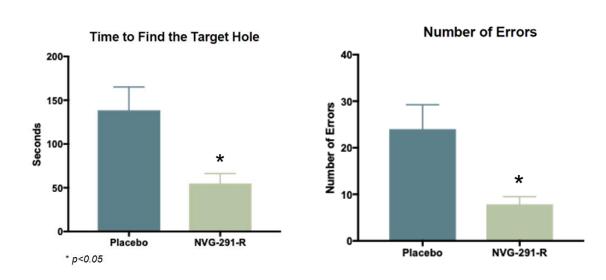
Pioneering preclinical study in the peer-reviewed scientific journal Cell Reports demonstrating NVG-291-R promotes nervous system repair and significant functional recovery in a mouse model of severe ischemic stroke

Significant functional repair from a stroke 7 days after onset in landmark preclinical study

Dramatic and Unprecedented Recovery From a Stroke

Improved Spatial Learning and Memory

Barnes Maze Test
Treatment beginning 7 days post stroke



Animals treated with NVG-291-R made **fewer errors** and identified the target hole in a **faster time** compared to placebo treated animals

Significant improvements observed when NVG-291-R was administered **up to 7 days after** an ischemic stroke

Improvements in spatial learning and memory may be relevant for other indications affecting cognition, such as Alzheimer's disease





Multiple Sclerosis *Repair*

No FDA Approved Drug that Improves Function

- MS is an autoimmune disease where the immune system attacks myelin in the central nervous system, and over time this results in increasing disability
- Approved disease-modifying drugs modulate the immune system, which can reduce relapses and slow disability progression, but none repair damage
- NervGen's goal is to repair/remyelinate the damage from MS, thereby improving function



NVG-291 Restores Motor Function in Multiple Sclerosis

Representative of Placebo Group

You Tube CA

Representative of NVG-291 Group

Score improves to 0.5 in 20 days



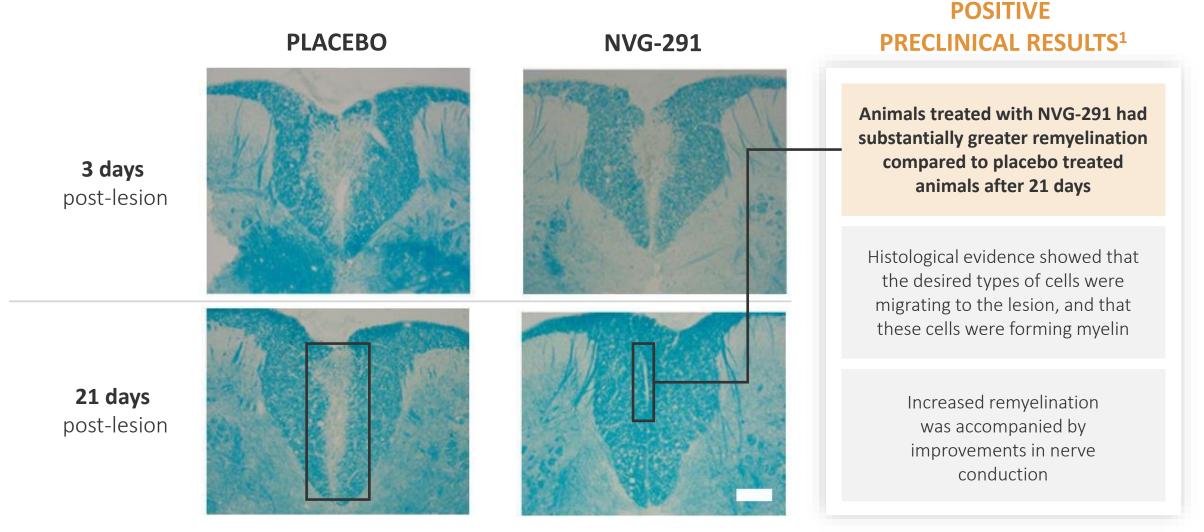




NVG-291 restored motor function in MS model¹, even when administered after symptoms were fully developed



NVG-291 Remyelinates in Multiple Sclerosis







NVG-291 Safety/Efficacy Studies in Multiple Sclerosis Patients

Multiple Sclerosis Phase 2

~80 patients, ~\$20 million estimated costs

- Multicenter, placebo controlled
- 3 months treatment duration
- Efficacy analysis includes clinical assessments, advanced structural imaging and fluid biomarkers

World-class Advisory Board with experts in MS research, clinical design, and functional assessments

Jack Antel, MD McGill University	Ex-Pres., Americas Committee for Treatment and Research in MS; Ex- Pres., International Soc. of Neuroimmunology
Jeremy Chataway, MD University College London	Advanced Clinical trial design expert in MS
Jeffrey Cohen, MD Cleveland Clinic Lerner College of Medicine	Ex-ACTRIMS President

Robert Naismith, MD Washington University	Expert in clinical trial design and clinical outcomes measures
Anneke van der Walt, MD, PhD Monash University	Led several international studies on digital biomarkers in MS

Share and Capital Structure

Exchange/Market: Ticker	TSX: NGEN.V	OTCQX: NGENF
Recent Share Price (March 31, 2023)	CA \$1.64	US \$1.23
Shares Outstanding	59.1 million	
Fully Diluted	76.8 million (~7.9 million options, ~9.8 million warrants)	
Insider Ownership	21.8%	
~Cash & Cash Equivalents (December 31, 2022)	CA \$22.5 million	US \$16.6 million



Key Upcoming Value Drivers

Phase 1b/2a spinal cord injury clinical trial to be initiated

Preclinical study results in an Alzheimer's model

Awarding of US Department of Defense and privately funded grants

New CEO



