

Enabling the Nervous System to Repair Itself

CORPORATE PRESENTATION

February 14, 2023

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NervGen's Clinical Trials are Led by Experienced Drug Developers



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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



UNOVARTIS

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)

Biogen

RESPONSE BIOMEDICAL





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**

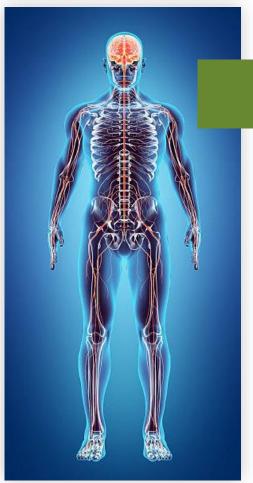
Target indications address **very attractive commercial opportunities** with **significant unmet medical needs** in spinal cord injury, Alzheimer's, multiple sclerosis and stroke

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



SEVERE SPINAL CORD INJURY MODEL

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**

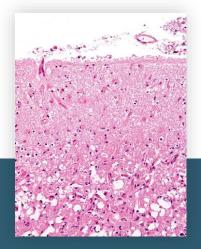
Intellectual property protection on NVG-291 until 2037



The Foundation of Our Technology

Pre 1990

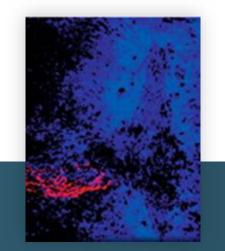
It was demonstrated that **glial scars** form at the site of injury to the nervous system and that scars in the brain cause neurons to be dysfunctional. Scars were later identified as the primary impediment of recovery



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

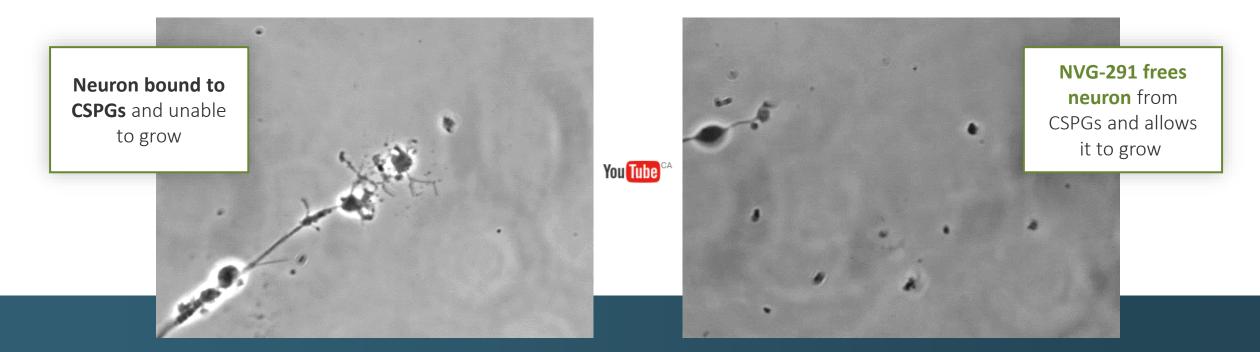


2015

Dr. Silver's team designed a peptide derived from PTPσ and shown to relieve CSPGmediated 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



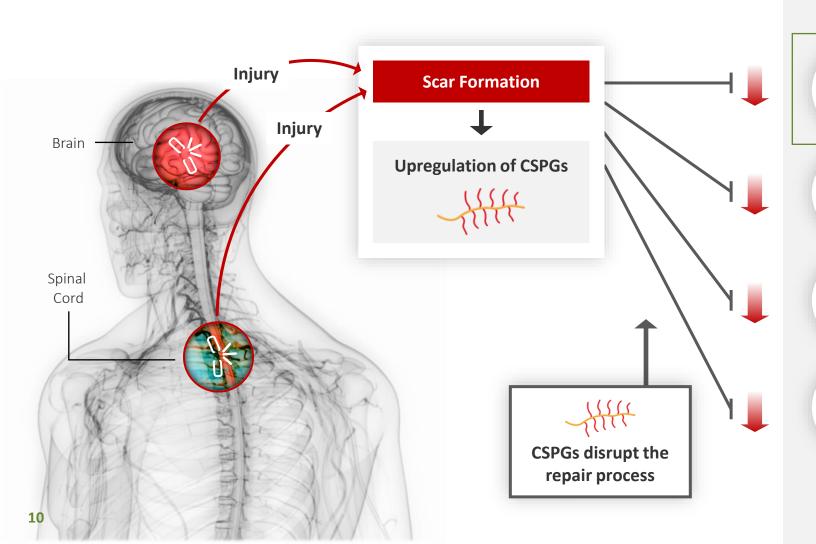
NVG-291, a 35 amino acid peptide, produced dramatic recovery in a spinal cord injury animal study: the results published in Nature¹ are now cited in over 327 publications

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

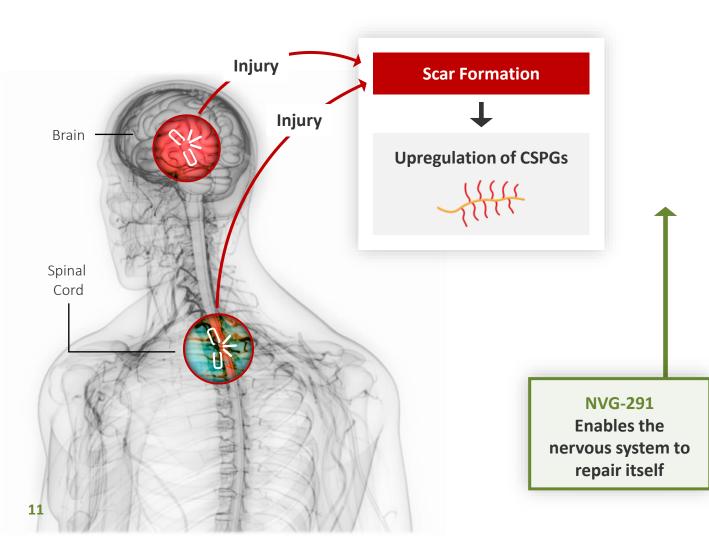


Stem cell preservation/migration Autophagy Microglial shifting



NVG-291

Takes the Brakes off Natural Repair Mechanisms



Repair Mechanisms:



Plasticity

The creation of new neuronal connections and rewiring of existing ones



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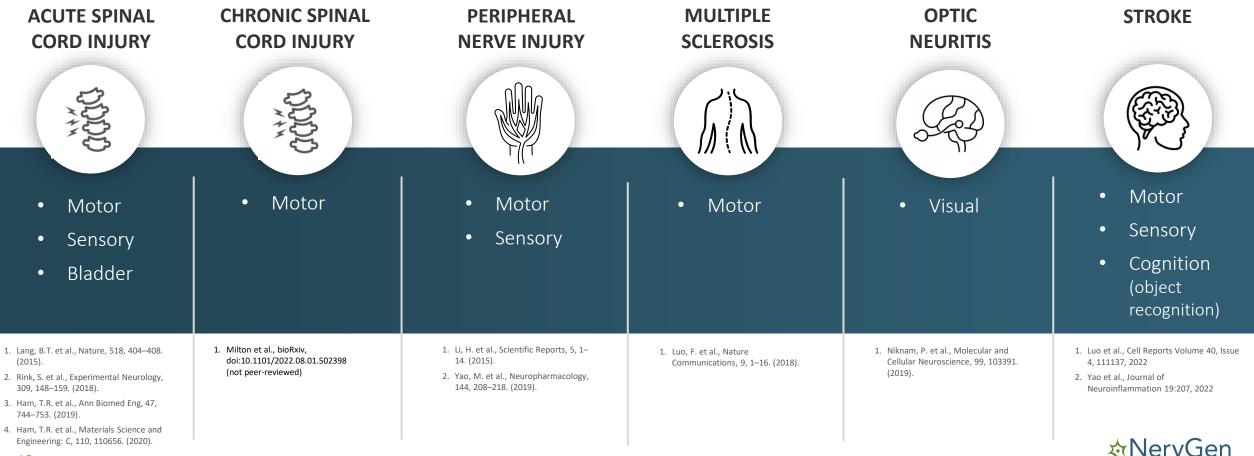
Stem cell preservation/migration Autophagy Microglial shifting



NVG-291 Broadly Restores Function

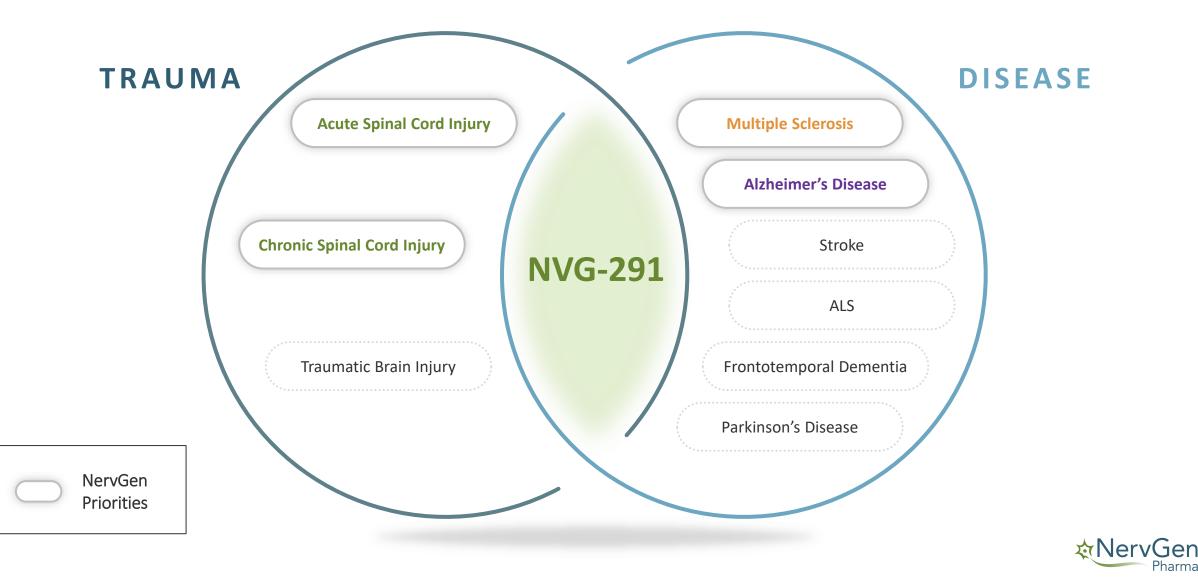
NVG-291 Has Demonstrated Dramatic Repair

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



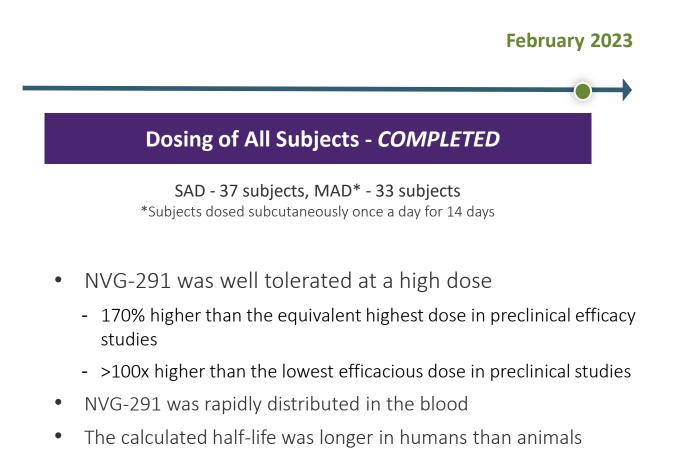
Pharma

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



Pharma

NVG-291 - Phase 1 Clinical Trial in Progress



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





Spinal Cord Injury

Acute and Chronic patients

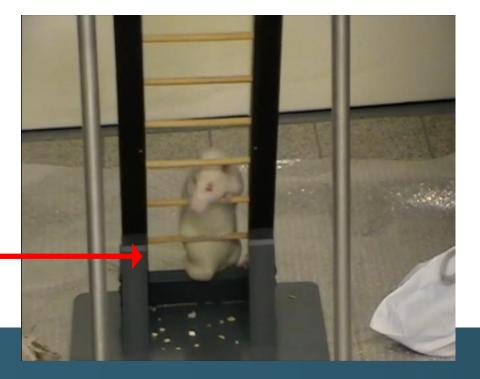
- No FDA Approved Drug that Improves Function
- NervGen's goal is to improve motor, bladder/bowel/sexual and/or sensory function *in* High unmet need coupled with potential key clinical outcomes may provide opportunity for expedited regulatory approval
- Unprecedented preclinical results



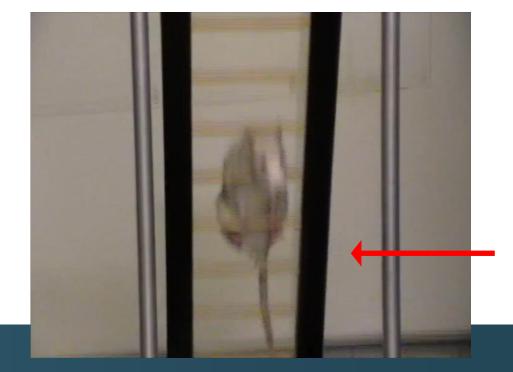
NVG-291 – Dramatically Repairs Spinal Cord Injury SEVERE SPINAL CORD INJURY MODEL

You Tube^C

Representative of Placebo Group



Representative of NVG-291 Group



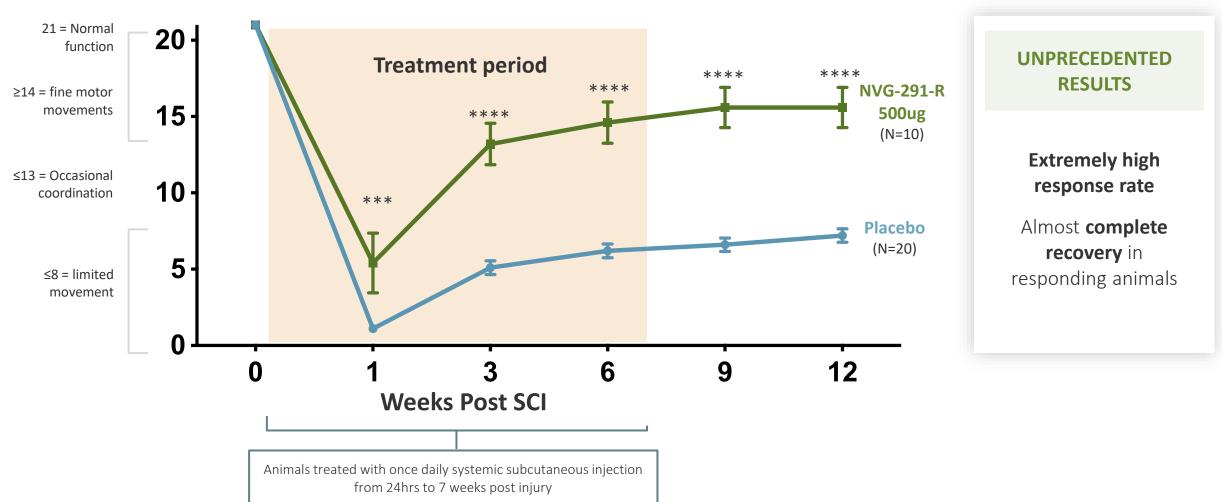
Hind legs are immobile

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



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

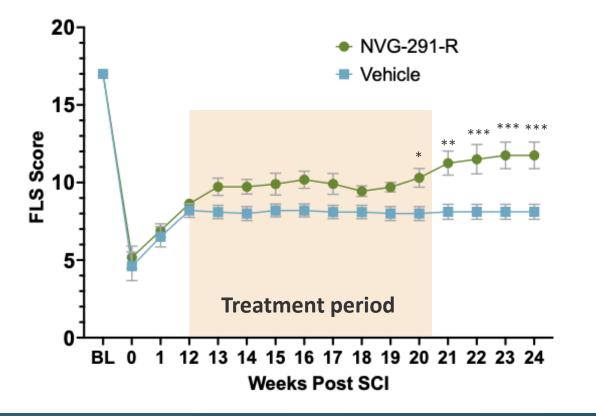
BBB Scale = Standard measure of mobility





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

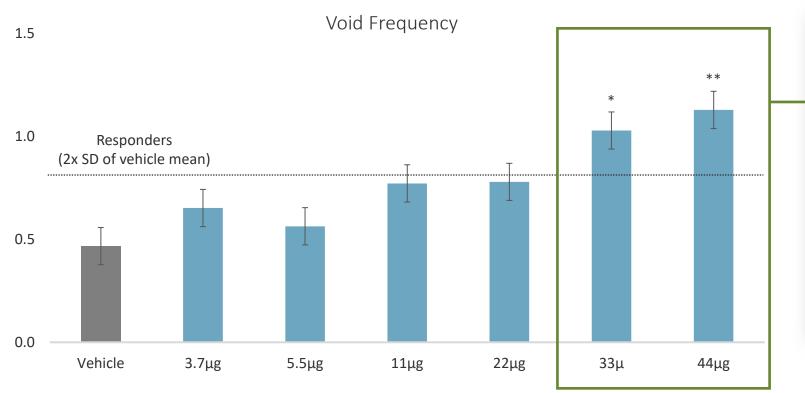
Animals treated with NVG-291-R show **significant improvements** in forelimb recovery at 24 weeks post injury

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



NVG-291 Improves Bladder Function

BLADDER DOSE RESPONSE



100% of animals in the two highest dose groups had improved bladder control function

NVG-291 treatment resulted in a dose dependent improvement in 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

- 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

Daniel Lammertse, MD University of Colorado School of Medicine World renowned surgeon/scientist; authored >240 scientific publications, >35 textbook chapters

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





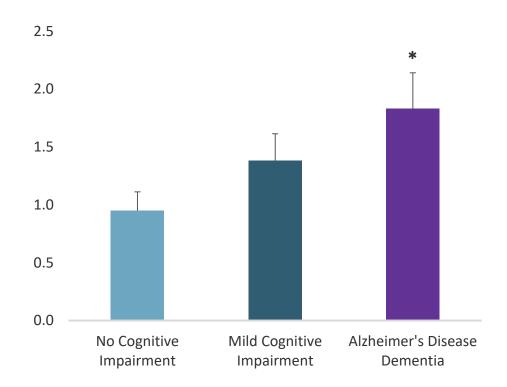
Alzheimer's Disease No FDA Approved Drug that Results in Sustained Improvement in Cognitive Function

- 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

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

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



* p<0.05 compared to NCI

23 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)	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
Martin Farlow, MD Indiana University School of Medicine	Led/contributed to >230 clinical trials; authored 493 peer reviewed research papers and 509 abstracts	Michael Weiner, MD University of California, San Francisco	Leader in development of MRI and PET for investigating and diagnosing neurodegenerative diseases
Bruce Lamb, PhD Indiana University School of Medicine	World-expert on biological underpinnings of Alzheimer's disease and related dementia	Henrik Zetterberg, MD, PhD University of Gothenburg, University College London	World expert in blood-based biomarkers in neurological disorder
George Perry, PhD University of Texas, San Antonio	Current and founding Editor-in-Chief of the Journal of Alzheimer's Disease		

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

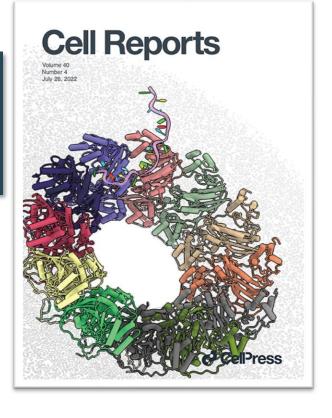


You Tube

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



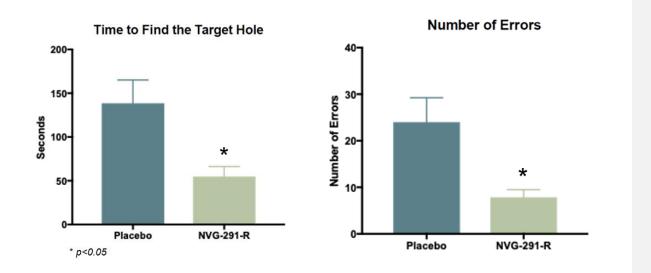
Pioneering preclinical study in the peer-reviewed scientific journal Cell Reports demonstrating NVG-291-R promotes nervous system repair and significant functional recovery in a mouse model of severe ischemic stroke

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

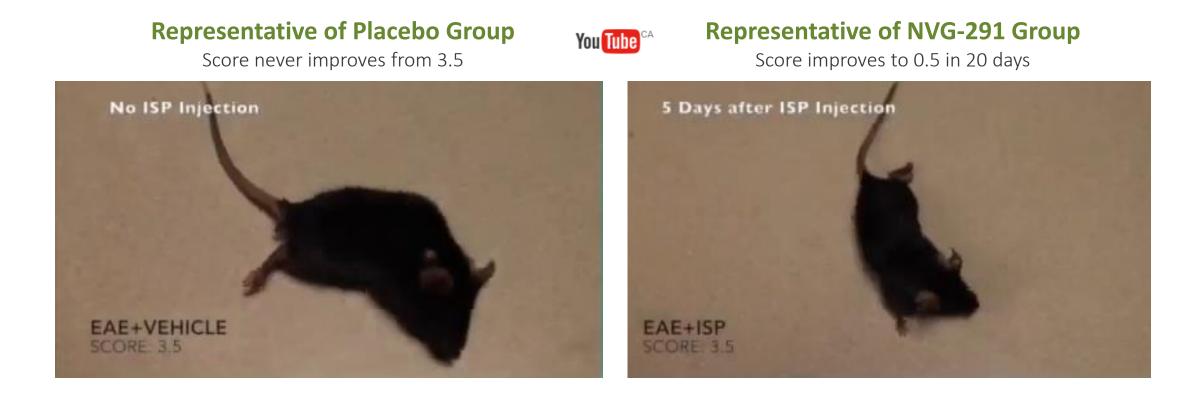


Multiple Sclerosis <u>Repair</u> No FDA Approved Drug that Improves Function

- 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



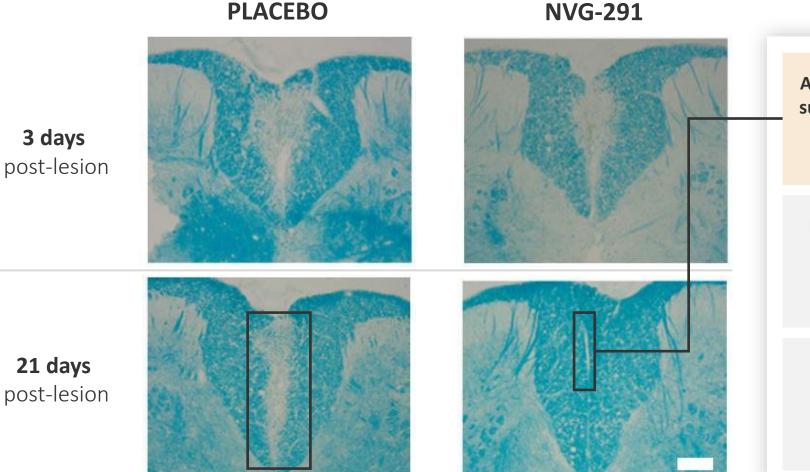
NVG-291 Restores Motor Function in Multiple Sclerosis



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



NVG-291 Remyelinates in Multiple Sclerosis



Lesion size in LPC demyelination model

POSITIVE PRECLINICAL RESULTS¹

Animals treated with NVG-291 had substantially greater remyelination compared to placebo treated animals after 21 days

Histological evidence showed that the desired types of cells were migrating to the lesion, and that these cells were forming myelin

> Increased remyelination was accompanied by improvements in nerve conduction



NVG-291 Safety/Efficacy Studies in Multiple Sclerosis Patients



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	Robert Naismith, MD Washington University	Expert in clinical trial design and clinical outcomes measures
Jeremy Chataway, MD University College London	Advanced Clinical trial design expert in MS	Anneke van der Walt, MD, PhD Monash University	Led several international studies on dig biomarkers in MS
Jeffrey Cohen, MD Cleveland Clinic Lerner College of Medicine	Ex-ACTRIMS President		

Share and Capital Structure

Exchange/Market: Ticker	TSX: NGEN.V	OTCQX: NGENF
Recent Share Price (February 10, 2023)	CA \$1.96	US \$1.50
Shares Outstanding	59.0 million	
Fully Diluted	76.2 million (~7.3 million options, ~9.9 million warrants)	
Insider Ownership	24.4%	
~Cash & Cash Equivalents (September 30, 2022)	CA \$27.7 million	US \$20.2 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





Enabling the Nervous System to Repair Itself



@NervgenP

in NervGen Pharma Corp.

