

Corporate Presentation

April 2025

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NervGen Highlights

Multiple preclinical studies using NVG-291-R reported improvement in axonal repair, plasticity, and motor, sensory and cognitive function

Q1 2025 – NVG-300

preclinical data readout

in models of ischemic

stroke, ALS and SCI

Q2 2025 – Phase 1b/2a
proof-of-concept
readout in people living
with chronic SCI



History of NervGen Technology



Jerry Silver, PhD

Case Western Reserve



Dr. Silver discovered glial scars contain chondroitin sulfate proteoglycans (CSPG), a group of molecules known to inhibit cellular events central to neural tissue repair



Dr. Silver and Harvard collaborators co-discovered that CSPGs bind to protein tyrosine phosphatase sigma (PTPo), a receptor present in the brain and spinal cord, involved in CSPG-dependent inhibition of neuroplasticity



Dr. Silver's team designed a peptide (NVG-291-R) derived from PTPo shown to relieve CSPG-mediated inhibition of nervous system repair; **NVG-291** is the humanized version of NVG-291-R



NervGen licensed NVG-291 global rights for development and commercialization in all indications from Case Western Reserve University with intellectual property protection until 2037

Phase 1b/2a Trial





NervGen initiated a **Phase 1b/2a** placebo-controlled proof-of-concept trial (NCT05965700) to evaluate the efficacy of NVG-291



NVG-291-R Improves CNS/PNS Repair in Preclinical Efficacy Models

Enhanced Plasticity, Repair (Axonal, Myelination), and Recovery of Function

Conditions Modeled

Functional Endpoints ACUTE SPINAL CORD INJURY

MOTOR SENSORY BLADDER

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).

Ham, T.R. et al., Materials Science and Engineering: C, 110, 110656. (2020).

Wang, H et al., Molecular Neurobiology, s12035-024-04304-3 (2024)

CHRONIC SPINAL CORD INJURY



MOTOR

Milton et al, Journal of Neurotrauma, (2023) doi:10.1089/neu.2023.0117

Luo et al., Cell Reports Volume 40, Issue 4, 111137, (2022)

Yao et al., Journal of Neuroinflammation 19:207, (2022)

Wang, R et al., Experimental Neurology, 114564, (2023)

Zheng, W. et al., Chemical Engineering Journal 483:149225, (2024) MULTIPLE SCLEROSIS (EAE)

STATE OF THE PARTY OF THE PARTY

MOTOR



STROKE

(Ischemic,

Hemorrhagic)

MOTOR
SENSORY
OBJECT RECOGNITION

Luo, F. et al., Nature Communications, 9, 1–16. (2018). PERIPHERAL NERVE INJURY



MOTOR SENSORY

Li, H. et al., Scientific Reports, 5, 1–14. (2015).

Yao, M. et al., Neuropharmacology, 144, 208–218. (2019).

Lv, S. et al., Neural Regeneration Research 16, no. 8:1598, (2021) OPTIC NERVE DEMYELINATION



VISUAL FUNCTIONAL

Niknam, P. et al., Molecular and Cellular Neuroscience, 99, 103391. (2019).



Product Pipeline

Multiple development opportunities

CANDIDATE	PROGRAM	DISCOVERY	PRECLINICAL	PHASE 1	PHASE 1b/2a
NVG-291	Spinal Cord Injury				
	Ischemic Stroke				
NVG-300*	Ischemic Stroke				
	Amyotrophic Lateral Sclerosis				
	Alzheimer's Disease				
	Multiple Sclerosis				



Markets and Opportunity in Nervous System Injury & Disease

Significant medical costs and morbidity

	Spinal Cord Injury	Ischemic Stroke	ALS	Multiple Sclerosis	Alzheimer's Disease
Incidence*	18,000	~690,000	~7,000	10,000	500,000
Prevalence*	291,000	9.4M	~25K-30K	~1M	6.7M
Lifetime Cost*	\$1M-\$4M+	\$140,000+	\$1.4M	\$4M+	\$400,000
System Cost*	\$50B+	\$57B	\$250M-\$1.0B	\$85B	\$320B-\$345B
Current Treatment*	Decompressive surgery and rehabilitation	TPA must be given within hours of stroke; rehabilitation	Disease modifying agents (e.g. riluzole, edaravone) to slow progression – none stop progression	Immunomodulatory/ immunosuppressive therapies to reduce relapses and/or slow progression	Symptomatic therapies (e.g. cholinesterase inhibitors) to temporarily improve cognition; anti-beta mAbs to slow progression
Unmet Needs*	Effective treatments to enhance recovery	Effective treatments to enhance recovery	Treatment that improve function	Treatments to remyelinate axons and improve function	Treatments to effect enduring improvements





SCI Demographics & Characteristics

- Average age: ~43
- Male (78%), female (22%)
- Cause: vehicle (38%); fall (33%); violence (15%); sports (8%)
- Annual hospitalization (30%): UTI, pneumonia, decubitus ulcers
- Duration of hospitalization and rehabilitation: 2 to 3 months
- Chance of depression: 25%
- Significant urinary and sexual dysfunction

TREATMENT

Surgery (decompression)

Rehabilitation (regain function)

No FDA approved drugs to enable sustained functional recovery



SCI Facts and Figures

Incidence and Prevalence

~18,000

Spinal cord injuries every year in the US¹

~300,000

People living in the US who have suffered a spinal cord injury in 2019¹

up to 500,000

Worldwide, the estimated annual incidence of spinal cord injury²

Economic Impact

Individuals with SCI face a difficult and expensive journey through the healthcare system; that journey begins with **2-3 months in rehabilitation** and **costs \$200,000 or more per patient**³

Individuals with SCI face an expected **lifetime cost of care between \$1M and \$4M**, depending on severity and age at injury⁴

In addition to the enormous economic costs, individuals with SCI face a shorter expected lifespan, higher unemployment, higher chance of bankruptcy⁵



NVG-291-R Promotes Recovery in Acute SCI

Overview

T8 compression injury

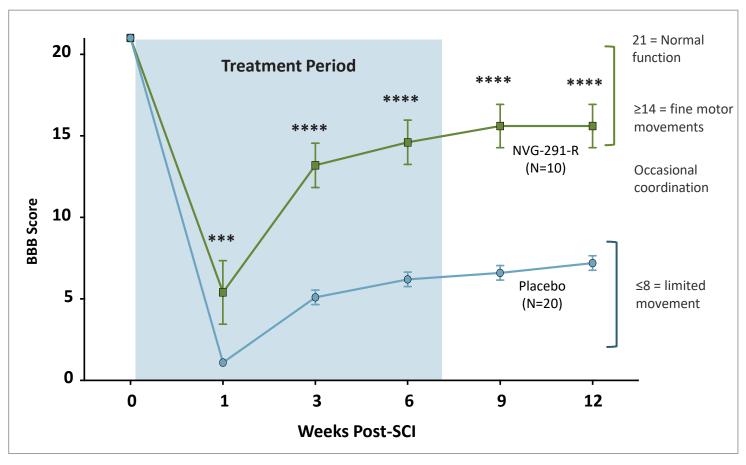
Treatment start: 1 day post-injury

Dose: 500 μg/day x 7 weeks

Results

- Significant recovery of locomotor and bladder function
- Functional improvements persist after treatment
- Enhanced neuroplasticity (i.e. axonal sprouting) near and far from injury

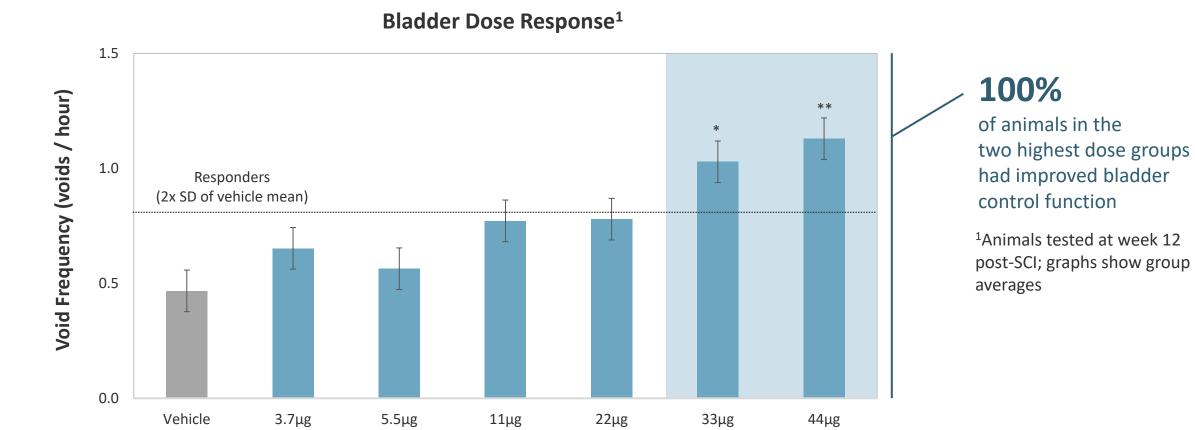
Hindlimb Function



*** P<0.001, **** P<0.0001



Improved Bladder Function After NVG-291-R in Preclinical Models



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

*p <0.05, ** p <0.01, n=5 per dosing group



NVG-291-R Improves Function in Severe Spinal Cord Injury Model

Representative of Placebo Group

(Back Legs and Tail Dragging)

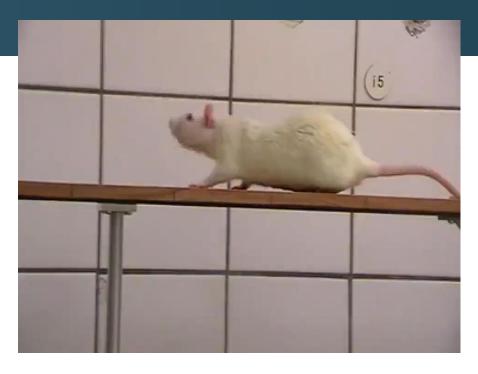
Representative of NVG-291-R Group

(Back Legs and Tail Active)





Click here to play video

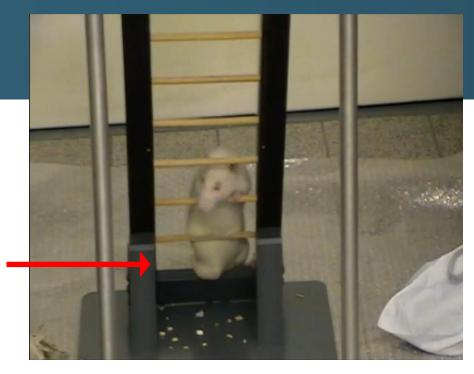


Remarkable and robust repair across multiple models



NVG-291-R Enables Significant Motor Recovery in Severe SCI Model

Representative of Placebo Group



Hind legs are immobile in placebo-treated animals

Representative of NVG-291 Group



Significant motor recovery with NVG-291-R: consistent coordination, toe clearance, tail held high consistently



Chronic SCI Preclinical Study

Overview

C2 lateral hemisection model of SCI

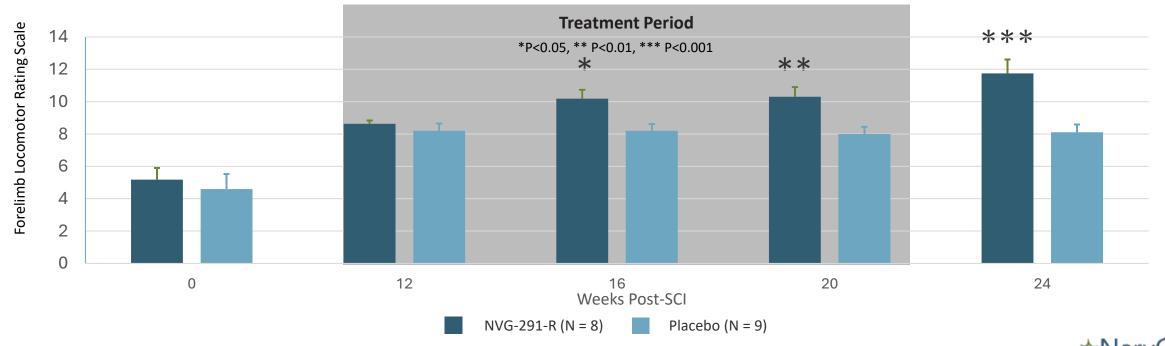
Treatment start: 12 weeks post-injury

Dose: 500 µg/day x 8.5 weeks

Results

NVG-291-R promoted recovery in chronic SCI

- Significant recovery of forelimb locomotor function
- Functional improvements persist after treatment





NVG-291 Phase 1 Clinical Trial in Healthy Volunteers

Study Design

Single Dose

- 37 subjects
- 6 dose levels
- Assessed through Day 8

Multiple Dose

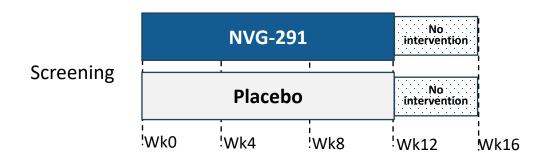
- 33 subjects
- 4 dose levels
- Subjects dosed subcutaneously once/day for 14 days
- Assessed through Day 21

Safety Results

- Well tolerated across all doses
 - Maximum tolerated dose (MTD) not reached
- No treatment discontinuations
- No serious/severe adverse events (AE) in NVG-291 group
- Most common AE was Grade 1/2 injection site reactions (ISR)
- No clinically significant effects related to NVG-291 treatment across all study parameters



Phase 1b/2a Trial of NVG-291-201 in SCI



Over 16 weeks

- Daily SQ injections (12 weeks)
- Electrophysiological assessments
- Clinical assessments
- Exercise/training: ~5 days per week

Single-center study – Shirley Ryan AbilityLab (Chicago, IL, USA)

- Uniform assessments and training regimen reduces variability of results
- Electrophysiological measurements easily standardized same assessors, equipment, technique

Two cohorts

- Approximately 20 patients per cohort
- Randomized 1:1 to NVG-291 (fixed dose) or placebo
- Weeks 1-12: blinded treatment







Phase 1b/2a Study Inclusion & Exclusion Criteria

Key Inclusion Criteria

Age 18-75

Traumatic SCI

Neurological level of injury C7 or higher

Motor incomplete with minimal/maximal level of motor function in upper and lower extremities

*Intact motor evoked potential (MEP)*¹ in two *qualifying* muscle groups:

- (1) At least 1 tibialis anterior (TA)
- (2) At least 1 first dorsal interosseus (FDI)

Key Exclusion Criteria

Non-traumatic SCI

SCI from gunshot or penetrating/stab injury

Two or more (non-contiguous) spinal cord lesions

Ventilator dependence

Cohorts of motor incomplete cervical SCI

1. Chronic: 1-10 years post-injury

2. Subacute: 20-90 days post-injury



Phase 1b/2a Trial Primary Objective and Co-Primary Endpoints

Primary Objective

To evaluate the effect of NVG-291 compared to placebo on relative percentage change in corticospinal connectivity to *qualifying* muscle groups

Co-Primary Endpoints

Relative percentage change from baseline to Week 12 in the normalized MEP amplitudes (corticospinal contribution) in the *qualifying* **FDI** and **TA** muscle groups

Ten Muscle Groups Assessed			
Upper extremity	Lower extremity		
Biceps brachii	Quadriceps		
Triceps brachii	Hamstrings		
First dorsal interosseous (FDI) Q	Tibialis anterior (TA) Q		
Flexor carpi radialis	Soleus		
Extensor carpi radialis	Abductor hallucis		

^Q Qualifying muscle group

Assuming a <u>treatment effect</u> on and <u>variability</u> of MEPs similar to that observed with electrical stimulation studies¹, with **8 subjects per arm** this study will have **280% power** to detect a difference ($\alpha = 0.025$, Student t-test 2-sided)



Phase 1b/2a Trial Secondary Endpoints (Clinical)

Change from baseline to Week 12 in **10mWT** time

O4 Change from baseline to Week 12 in **GRASSP** version 2 scores

Change from baseline to Week 12 in **9-HPT** time

Change from baseline to Week 12 in lower extremity motor scores

Change from baseline to Week 12 in **pinch** dynamometry force

Change from baseline to Week 12 in upper extremity **motor scores**

Other secondary objectives:

Changes in other electrophysiological parameters

- Change in MEP amplitudes (corticospinal) of non-qualifying muscle groups
- Change in reticulospinal MEP amplitudes
- Change in MEP latencies
- Change in maximal voluntary contractions

Safety/tolerability of NVG-291; pharmacokinetics of NVG-291

Exploratory objectives:

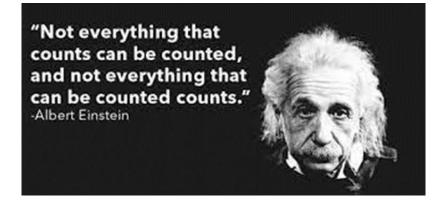
Changes in spasticity (modified Ashworth, pendulum test), SCAR, ISNCSCI sensory scores, autonomic function (ISAFSCI), mobility/ADLs (SCIM III), quality of life (SCI-QOL), advanced MRI imaging, blood biomarkers



Additional Patient-Reported Data: Qualitative Review of Subject Experience







Incorporating qualitative semi-structured subject interviews

Exploring subjects' experiences of potential beneficial treatment

Aligns with FDA Patient-Focused Drug Development Guidance



SCI Development Programs

Company	Product	Stage	Comments
AbbVie	Elezanumab: Intravenous RGMa protein inhibitors	Phase 2	 Phase 2 initiated Sep 2020, results expected H2 2025 Electrophysiology-based secondary endpoint recently added to trial design (NCT04295538)
Mitsubishi Tanabe	MT-3921: Intravenous RGMa protein inhibitors	Phase 2	 Phase 2 initiated – last update posted Apr 2024 Results expected H1 2025 (NCT04683848)
Lineage Cell Therapeutic	OPC-1: Intraparenchymal hESC-derived oligodendrocyte progenitor cells	Phase 1/2	 Amended IND to include new delivery device and initiate DOSED trial in 2024 Open label, subacute and chronic, primary endpoint is safety with new device (NCT05975424)
ReNetX Bio	AXER-204: Intrathecal Anti Nogo-A, MAG, and Omgp	Phase 1/2	 Missed primary endpoint in Ph 1/2 ITT population but showed signal in motor incomplete patients in post hoc analysis* (NCT03989440)





NVG-300, A Proprietary Molecule Diversifying Pipeline and Opportunity for Partnering

01

A new proprietary molecule discovered at NervGen in 2022



Eligible for the BLA development path



Demonstrated favorable pharmaceutical properties (solubility, metabolic stability)

 Severe injury model characterized by heightened spinal cord damage and impaired spontaneous recovery



Composition of matter IP protection expected to extend beyond 2040

Next steps

Formulation development

Initiating evaluation in preclinical models of ischemic stroke and ALS, confirmatory SCI study



Experienced Leadership Team



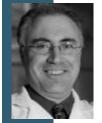
MIKE KELLY, MBA, CHIEF EXECUTIVE OFFICER

Over 30 years of pharmaceutical experience. Formerly President of US Ops for Adapt Pharma, Inc., which developed and commercialized NARCAN Nasal Spray in the US and Canada and was sold to Emergent BioSolutions for US\$735 million.



BILL ADAMS, CPA, CA, CHIEF FINANCIAL OFFICER

Over 25 years of strategic financial management experience including mergers and acquisitions, operations and capital markets in Canada and the US.



DAN MIKOL, MD, PHD, CHIEF MEDICAL OFFICER

Over 25 years of experience in neurology clinical research. Former Head of clinical development at Amgen in neuroscience and nephrology and was instrumental in the approval of Aimovig, and development lead for Tysabri at Biogen.



CHUCK OLSON, DSC, SR. VP, TECHNICAL OPERATIONS

Over 40 years of experience in process development, manufacturing and CMC associated quality and regulatory activities for many clinical and commercial products.



LIZ EBERHARDT, BSC, SR. VP, PROJECT MANAGEMENT

Over 25 years of biotech experience in product leadership and program management and has taken multiple compounds through all stages of development including preclinical and commercialization.



KEVIN ROONEY, MBA, SR. VP, CORPORATE DEVELOPMENT & STRATEGY

Over 30 years of experience in building businesses in oncology, central nervous system, diabetes, anesthesia, rare disease, cardiovascular, gastrointestinal, and infectious disease therapeutic areas.



MATVEY LUKASHEV, PHD, VP, RESEARCH & PRECLINICAL DEVELOPMENT

Over 30 years of experience in academia, industry and biotech settings focused on translational research and drug discovery.











































Board of Directors



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Former Partner, Deloitte LLP



Harold Punnett, DMD
Co-Founder



Mike Kelly
President & CEO, NervGen



Adam Rogers, MD
Former CEO & Co-Founder, Hemera



Brian BayleyDirector, Earlston Investments



Neil Klompas
President & CEO, Augurex



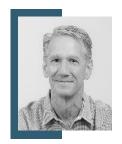
John RuffoloFounder & Managing Partner, Maverix



Randall Kaye, MD
CMO, Longboard Pharmaceuticals



Krista McKerracherFormer Global Franchise Head, Novartis



Craig Thompson
CEO, Cerevance



Upcoming Milestones

Q1 2025

Initiate dosing in subacute
cohort in Phase 1b/2a proof-ofconcept in people living with SCI

NVG-300 preclinical data readout in models of ischemic stroke and ALS

Q2 2025

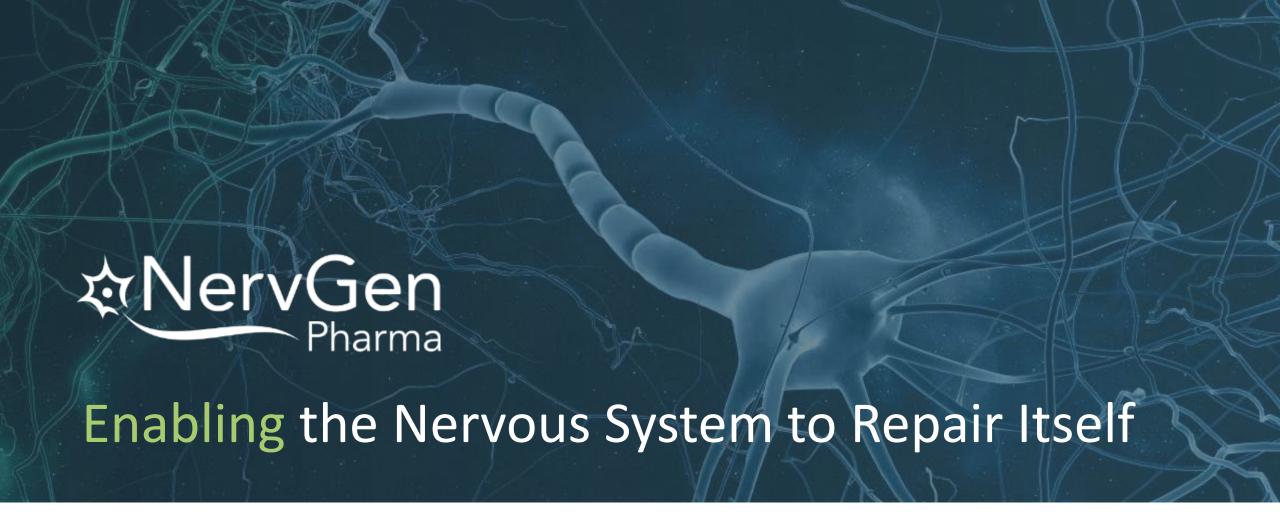
Phase 1b/2a proof-of-concept readout in people living with chronic SCI



Share and Capital Structure

Exchange/Market: Ticker TSX: NGEN.V OTCQB: NGENF CA\$3.09 / US\$2.17 Recent Share Price (Apr 3, 2025) **Shares Outstanding** 71.0 million Fully Diluted 94.8 million (~13.7 million options & retention securities, ~10.1 million warrants*) **Insider Ownership** 20.9% Cash & Cash Equivalents (Dec 31, 2024) CA\$17.3 million / US\$12.0 million





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