

The future of cell therapy.



## **Corporate Overview**

## Forward-Looking Statements

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"We aim to pioneer a new branch of medicine, based on transplanting specific cell types into the body"



**Business Overview** 

## **Company Overview**

Innovative Platform	Manufacturing and transplanting specific cell types from a single pluripotent cell line; scalable "off the shelf" cell transplants for multiple conditions		
Validating Partnerships	Genentech A Member of the Roche Group  CIRM CRUFORNIRY / TEM CELL ROENCY  CANCER RESEARCH UK		
Clinical Programs	OpRegen: Dry Age-Related Macular Degeneration with GA OPC1: Cervical Spinal Cord Injury VAC2: Non-small Cell Lung Cancer (oncology platform)		
Differentiated Data	Four cases of <u>retinal tissue restoration</u> observed in dry AMD patients One-third of spinal injury patients <u>gained at least 2 levels</u> of motor function <u>Potent induction</u> of immune responses observed in advanced cancer patients		
Market Opportunity	Billion-dollar commercial opportunities with no or few treatment options		
Financial Position	~\$65.1 million in cash and marketable securities as of Sep 30, 2021		
Market Capitalization	~\$428 million as of November 5, 2021		



## Novel Clinical Cell Therapy Pipeline

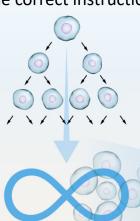
LINEAGE **PROGRAM** PHASE 1 PHASE 2 PHASE 3 **PARTNERS** Genentech 24 patients treated **OpRegen®** A Member of the Roche Group Ophthalmology Dry AMD with Geographic Atrophy (GA) 30 patients treated OPC1 Demyelination Spinal Cord Injury (SCI) 7 patients treated VAC2 Non-Small Cell Lung Cancer (NSCLC) Immuno-oncology



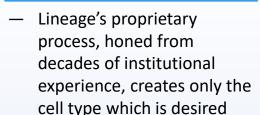
## Lineage Technology Platform – Allogeneic Cell Transplants

#### **Expansion**

- Product development starts from a frozen vial of selfrenewing stem cells
- These pluripotent cells can become any cell type in the body when provided with the correct instructions

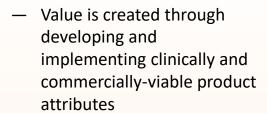


#### Differentiation



- No alterations are made to the cell's DNA
- In-house cGMP manufacturing allows for commercial-scale production from a single vial of stem cells

#### **Development**



 Pipeline expands by broadening indications or adding additional cell types



**Retinal Cells** 





→ OPC1



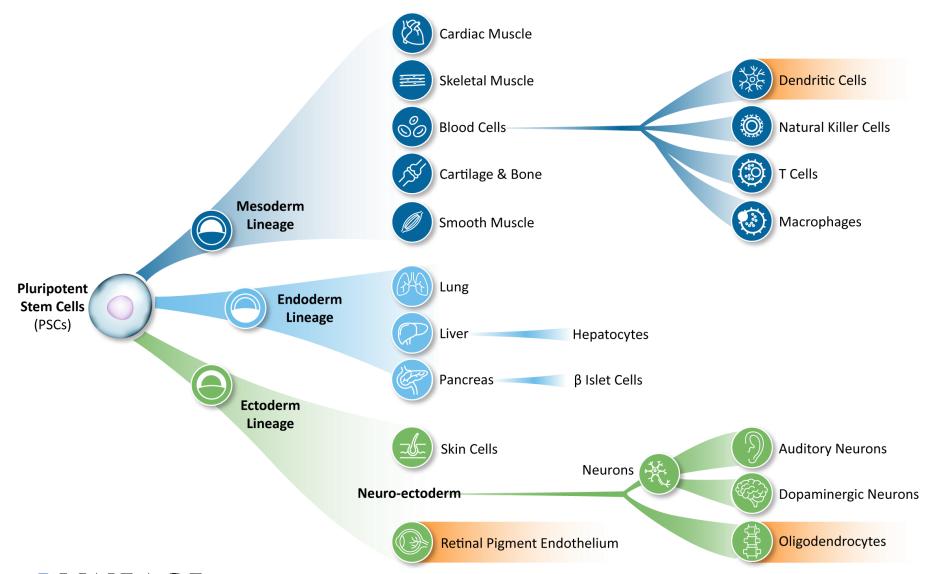
→ VAC2

 $+\ )$  Other Pipeline Programs





## **Future Product Candidate Opportunities**





## Competitive Advantage - Differentiation (Process Development)

## Lineage's competitive advantage is the *differentiation* of an *unlimited* supply of pluripotent stem cells into *specialized cell types*

#### **Capabilities**

- Source cell characterization, banking and versatile expansion systems
- Differentiation process development; culture conditions, systems, optimization of differentiation cues (growth factor selection, timing, etc.)
- Analytical method development for process control and product release
- Scale-up modalities, substrates, harvesting protocols
- Enhancements; genetic modification (optional), various expression systems
- Clinically compatible post-production processing

#### cGMP Facility



Multiple Clean Rooms for Parallel cGMP Production Runs

Extensive IP portfolio covers processes, products, and methods of use









AMD is the **leading cause** of irreversible vision loss in the US



Source: aao.org

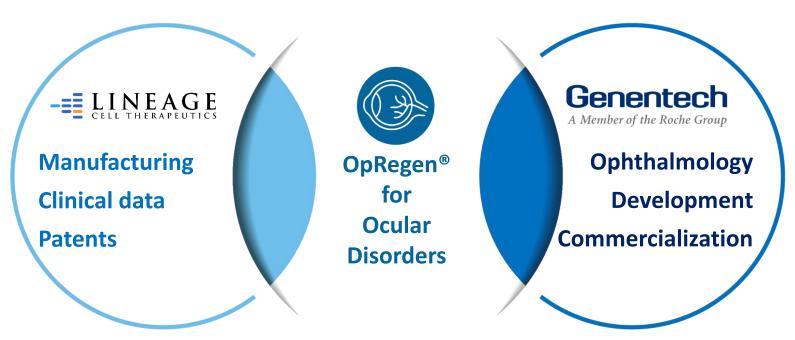
**OpRegen®: RPE Cell Transplants to Treat Dry AMD** 





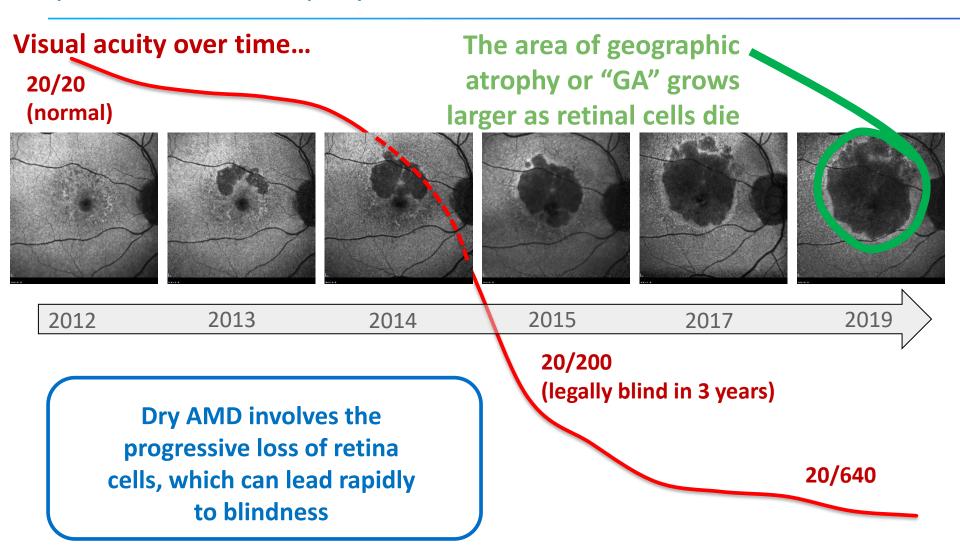
## Exclusive collaboration for the development and commercialization of OpRegen for the treatment of ocular disorders

- \$50 million up front; double-digit tiered royalties; \$620 million of potential payments
- Lineage to complete ongoing study and continue certain manufacturing activities
- Genentech responsible for clinical development and commercialization





## Dry AMD Can Lead Rapidly to Blindness

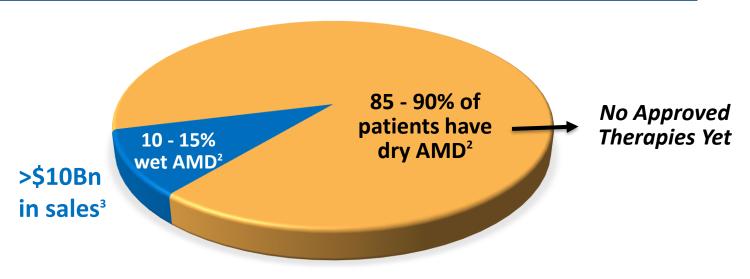




## Multi-Billion Dollar Market Opportunity in the U.S.

# Age-related Macular Degeneration (AMD) in all forms afflicts ~11 million people in the United States

Type of AMD	% of AMD Cases	FDA Approved Therapies
Wet AMD	10 – 15%	Lucentis & Eylea (\$10 Billion in annual sales)
Dry AMD	85 – 90%	None

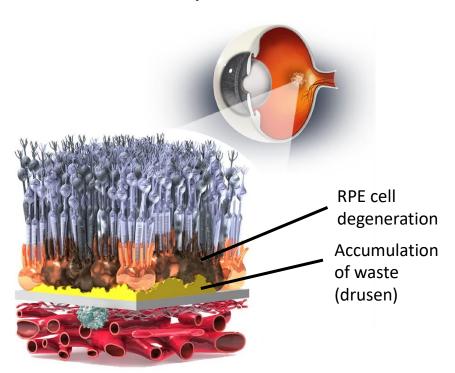




Sources: (1) Bright Focus Foundation. Macular Degeneration Facts & Statistics: Bright Focus Foundation. <a href="http://www.brightfocus.org/macular/about/understanding/facts.html">http://www.brightfocus.org/macular/about/understanding/facts.html</a>; (2) JM Seddon, Epidemiology of age-related macular degeneration. (AP Schachat, S Ryan eds.) Retina, 3rd ed. St. Louis, MO: Mosby; 2001;1039-50; (3) 2018 product sales summary based on publicly reported revenue figures for Lucentis and Eylea.

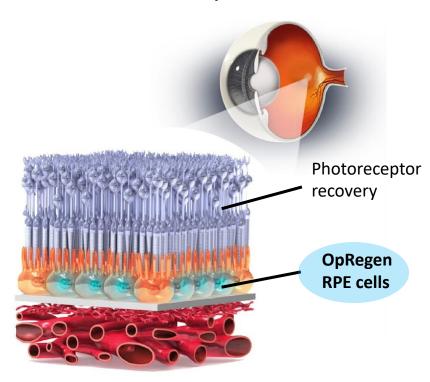
## Lineage Approach – OpRegen, an RPE Cell Transplant

## **Pre-Transplant**



Dry (atrophic) AMD involves the loss of retina cells, creating an area of geographic atrophy (GA), which causes impaired vision and blindness

## Post-Transplant



OpRegen is an injection of RPE cells beneath the retina, to replace lost retinal cells and preserve or improve vision



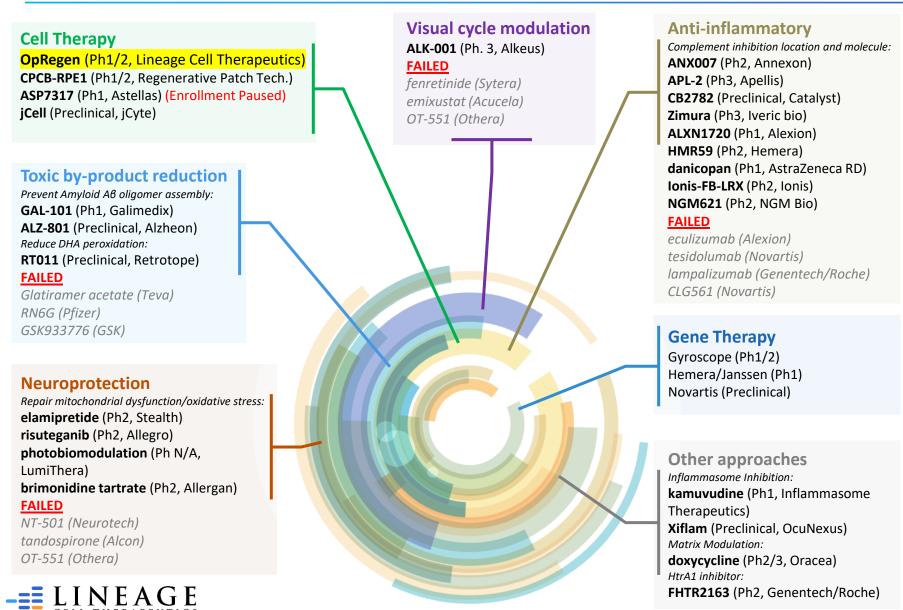
## Commercially-Suitable Manufacturing Process

- OpRegen consists of pure RPE cells >99%
  - Starts from an NIH-approved cell line established >20 years ago
  - Extensive functional and identity characterization is employed for product release
  - No genetic modifications are made to the cells
  - No residual pluripotent cells detectable in clinical material
- Clinic-ready, immediate-use "thaw and inject" formulation
  - No dose preparation required
  - From frozen cells to delivery device in 5 minutes
- Current production scale is 5 billion RPE cells per 3-liter bioreactor
  - Equal to 2,500 clinical doses/batch
  - Further scale-up can be performed in larger or parallel reactors





## **Dry AMD Competitive Landscape**









OpRegen Phase 1/2a
Clinical Trial
Interim Results



**Replace and Restore** 

## Phase 1/2a Clinical Trial - Promising Interim Results Continue (N=24)

#### **STRUCTURE:**

- 4 patients have shown evidence of <u>retinal tissue restoration</u>
  - All four cases of restoration have shown evidence of smaller or unchanged areas of atrophy at 12 months post-treatment; all four had improved BCVA at 12 months
  - 4/4 (100%) of patients with fulsome coverage across the area of atrophy showed restoration

#### **FUNCTION:**

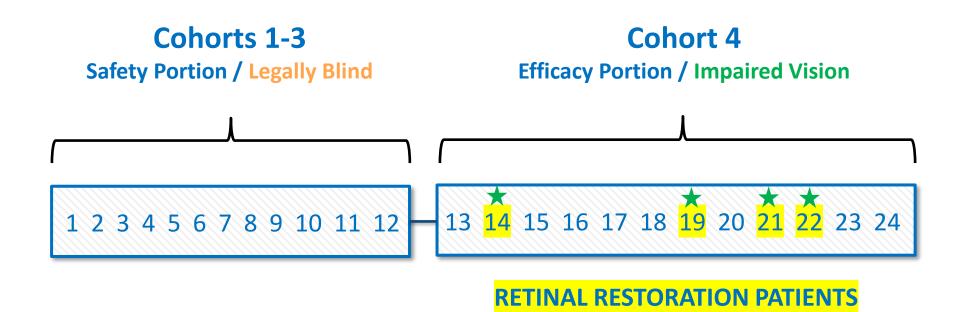
- 58% of all 12 Cohort 4 patients' treated eyes were at or above baseline visual acuity
  - 15M, or last time point available, up to >3y post-treatment
  - Visual acuity continued to decline in the majority (67%) of untreated eyes
- Statistically significant differences in visual acuity continue to be observed in Cohort 4
  patients between OpRegen treated and fellow untreated eyes (n=12)

#### **SAFETY, TOLERABILITY, DURABILITY:**

- OpRegen transplants have been well tolerated with no unexpected AEs or SAEs
- Earliest grafts have persisted for more than 5 years
- Zero cases of rejection (N=24)



## Phase 1/2a Clinical Trial of OpRegen — Enrollment Complete



Purpose: To evaluate the safety and efficacy of transplanted RPE cells

in patients with dry AMD with geographic atrophy

Design: Open label, single arm, international, multi-center

Dose and Administration: One 50-100 ul dose of cells injected into the subretinal space



**★** = OpRegen placed across area of atrophy

Imaging the Retina - Fundus Autofluorescence (FAF) and Optical Coherence Tomography (OCT)

FAF

OCT

- A flash of light causes cells to fluoresce, which is recorded in a single plane and with minimal structural resolution
- OpRegen cells lack lipofuscin, the material which fluoresces, so OpRegen cells appear as atrophic areas
- Differences in light returned to a detector creates a 3D images of all retinal layers
- The AAO considers OCT the "gold standard" for imaging the retina
- Offers much greater detail of anatomical structures



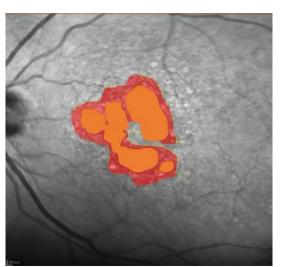
## High Resolution OCT Provides Resolution Close to Histology

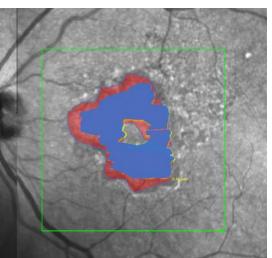
Retinal L	ayers	1100	013
Abbr.	Name	RNFL	
ILM	Internal Limiting Membrane	GCL	
RNFL	Retinal Nerve Fibre Layer	IPL IPL	
GCL	Ganglion Cell Layer	INL	
IPL	Inner Plexiform Layer		NAME OF TAXABLE PARTY.
INL	Inner Nuclear Layer	To Sales	
OPL	Outer Plexiform Layer	ONL	
ONL	Outer Nuclear Layer	ELM	
ELM	External Limiting Membrane	PR	MICHAEL MAN
PR	Photoreceptor Layers	RPE BM	
RPE	Retinal Pigment Epithelium	AND DESCRIPTION	
вм	Bruch's Membrane	cs	
СС	Choriocapillaris		
CS	Choroidal Stroma		

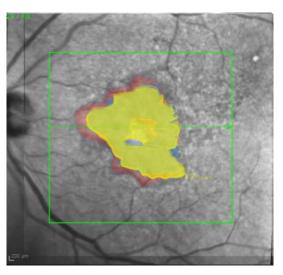


# First Case of Retinal Restoration – *Smaller* Area of GA, Maintained for ~3 Years

Date	Time in Study	Colored area on Figure below	Area mm² (SQRT)	Changes in rate of progression from previous	Changes in rate of progression from baseline
May 2017	Minus 1 year	Orange	4.21 mm² (2.05)	N/A	N/A
July 2018	Baseline	Red	7.90 mm² (2.8)	+ 0.64 mm sqrt/yr	N/A
April 2019	Month +9	Blue	5.74 mm² (2.39)	- 0.61 mm sqrt/yr	- 0.61 mm sqrt/yr
October 2019	Month +15	Green	6.48 mm² (2.54)	+ 0.30 mm sqrt/yr	- 0.20 mm sqrt/yr
June 2020	Month +23	Yellow	6.52 mm² (2.55)	+ 0.015 mm sqrt/yr	- 0.13 mm sqrt/yr

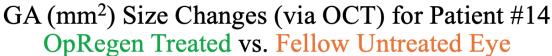


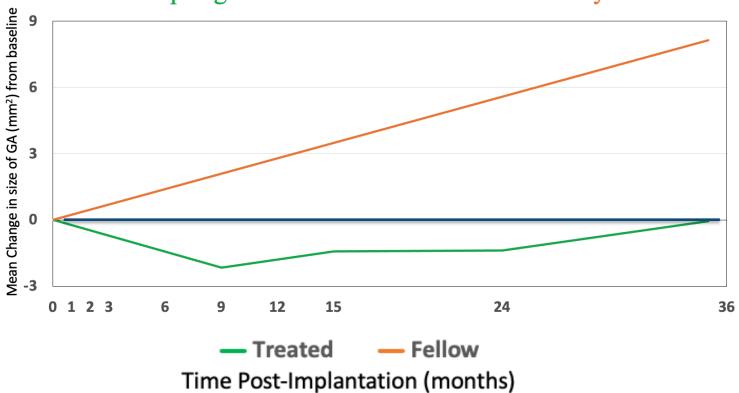






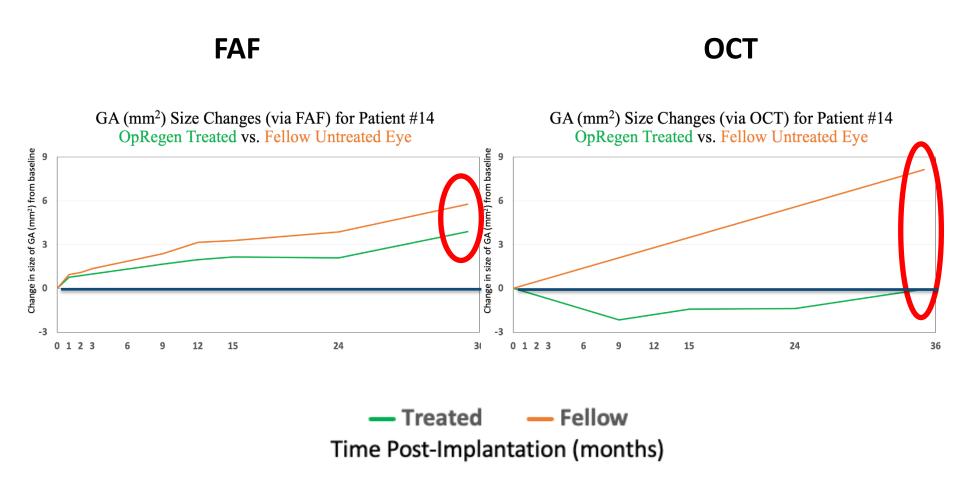
## First Reported Case of Retinal Restoration – GA Measurements





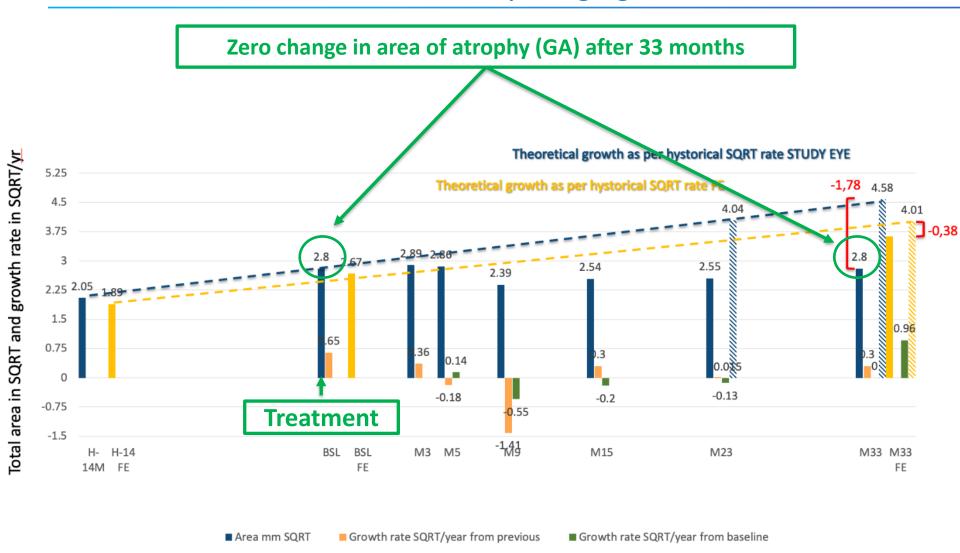


# First Case of Retinal Restoration - Utilizing OCT to Collect GA Measurements





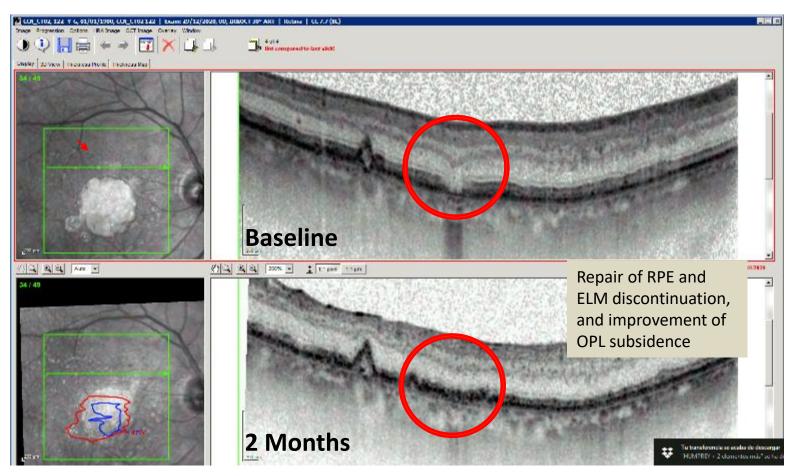
# First Case of Retinal Restoration – Assessing GA Progression Using FAF Alone or OCT and Multimodality Imaging





### Second Case of Retinal Restoration – Evident at 2 Months

Evidence of outer retinal regeneration and *complete resolution* of iRORA lesion



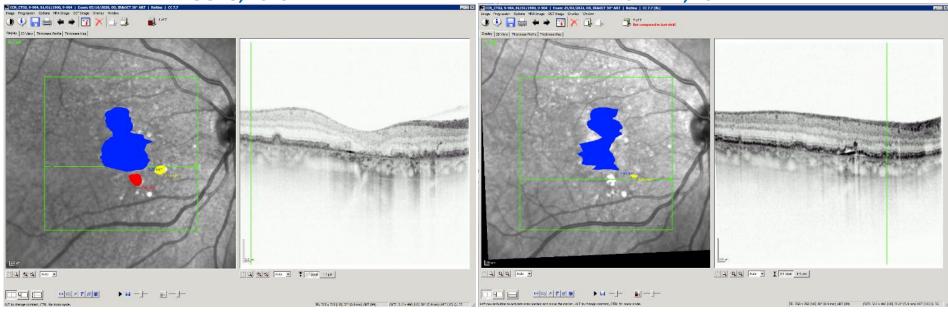
**iRORA** = Incomplete Retinal Pigment Epithelial and Outer Retinal Atrophy



#### Third Case of Retinal Restoration – Evident at 3 Months

### **ELM-based Area of Atrophy (Baseline to 3 Months)**

OCT 5, 2020 JAN 21, 2021



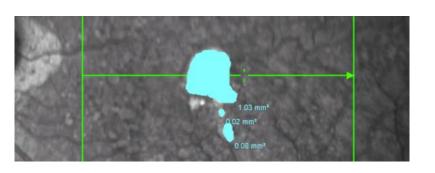
TOTAL AREA: 3.56 mm<sup>2</sup>
TOTAL AREA: 2.69 mm<sup>2</sup>

Total area 3M GROWTH RATE: -0.87 mm<sup>2</sup> (ANNUAL RATE - 3.48 mm<sup>2</sup>)

SQRT transformation 3M GROWTH RATE: -0.23 mm (ANNUAL RATE -0.92 mm)



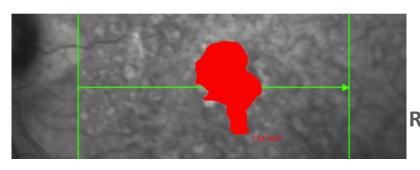
## Fourth Case of Retinal Restoration – No GA progression after >1 year



1.06 mm SQRT (1.13 mm²)

<u>Historical Image</u>

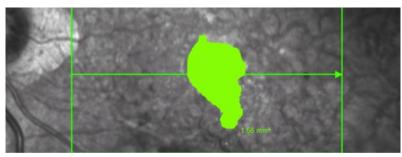
Obtained 16 Months before Baseline Visit



1.28 mm SQRT (1.64 mm<sup>2</sup>)

<u>Baseline Image</u>

Rate of Growth from Historical Image = +0.165 mm/yr



1.25 mm SQRT (1.56 mm<sup>2</sup>)

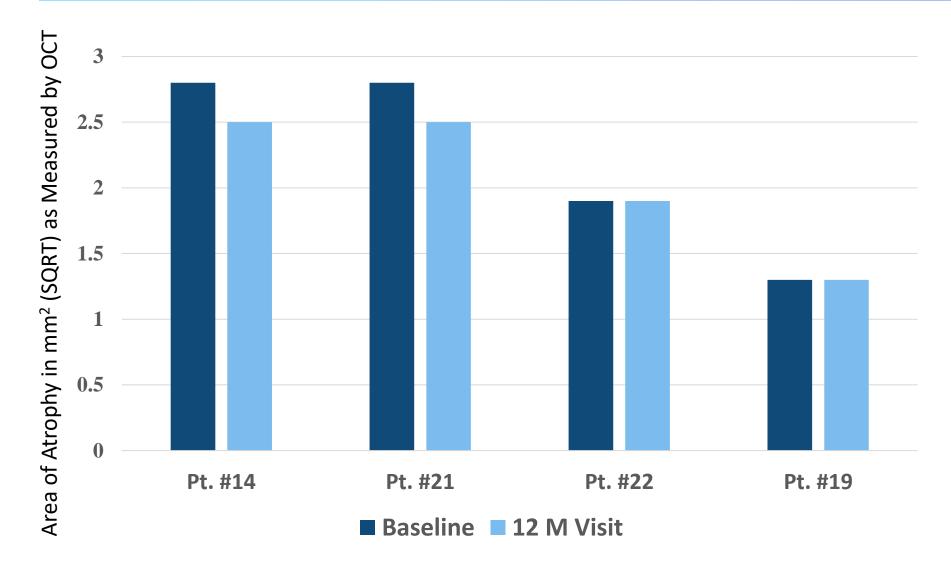
13.5 Months post-OpRegen

Rate of Growth from Baseline Image = - 0.026 mm/yr



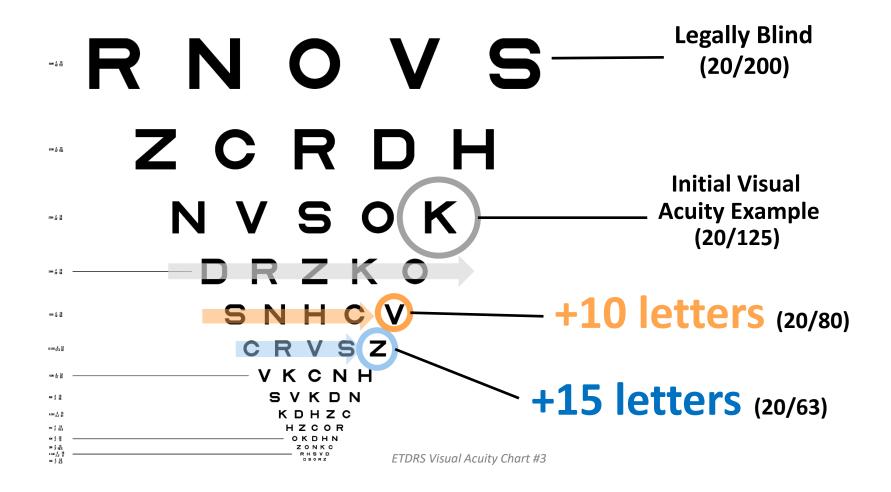
The SQRT area of atrophy decreased by 2.34% 13.5 Months post-OpRegen from the size of the GA observed at Baseline

## Four Cases of Retinal Restoration – No GA progression after 1 year





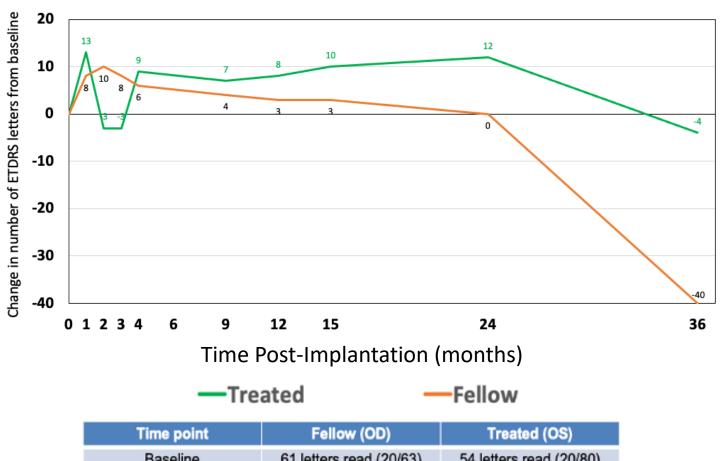
## Real-World "Letters of Improvement"

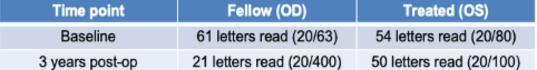




## First Case of Retinal Restoration - Durable Improvements

## BCVA Changes for Patient #14 – Treated vs. Fellow Eye

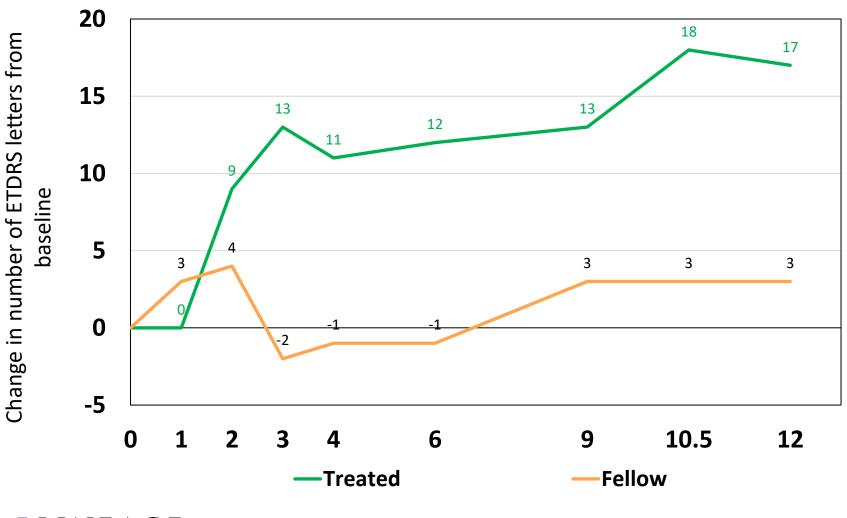






# Third Case of Retinal Restoration - Vision in Treated Eye Dramatically Improved

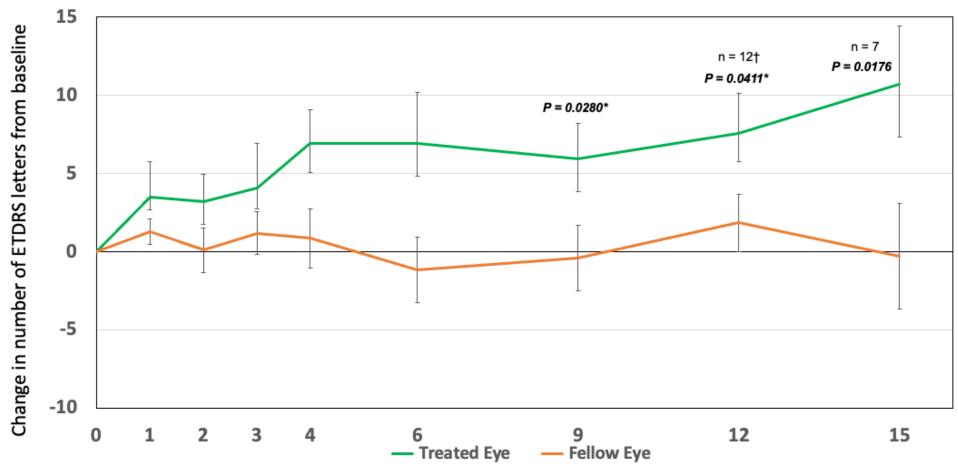
## BCVA Changes Treated vs. Fellow Eye





Time Post-Implantation (months)

## Mean Change (SEM) in Cohort 4 BCVA – Treated and Fellow Eye



<sup>\*2-</sup>sided Wilcoxon Signed Rank (NCSS)
† Reflects monitored data changes since AAO 2021 presentation

Time Post-Implantation (months)



## Positive Patient-Reported Outcomes (Quality of Life Questionnaire)

- National Eye Institute (NEI) Visual Function Questionnaire (VFQ-25)
  - 25 vision-related questions reported across 11 constructs
- Improvement occurred in 10 of 11 categories and remained unchanged in one category in Cohort 4 patients

Category	n (%) Change from Screening to 1 Year post-Treatment (n = 11 available to date)
General Vision	8/11 (73%) of patients reported improvement
Ocular Pain	1/11 (9%) of patients reported improvement
Near Activities	8/11 (73%) of patients reported improvement
Distance Activities	4/11 (36%) of patients reported improvement
Vision Specific: Social Functioning	5/11 (45%) of patients reported improvement
Vision Specific: Mental Health	7/11 (64%) of patients reported improvement
Vision Specific: Role Difficulties	4/11 (36%) of patients reported improvement
Vision Specific: Dependency	4/11 (36%) of patients reported improvement
Driving	0/11 (0%) of patients reported improvement (most patients were not driving at screening)
Color Vision	1/11 (9%) of patients reported improvement (most patients reported highest possible score, so no improvement was possible)
Peripheral Vision	2/11 (18%) of patients reported improvement



## OpRegen Represents a Multi Billion-Dollar Commercial Opportunity

- Transplanting RPE cells may provide transformational benefits beyond the reach of other approaches
- Market opportunity is not limited by monogenic deficiencies (e.g. gene therapy)
- Four clinical cases of retinal restoration reported (first known clinical cases)
- Treatment to date has been well-tolerated
  - Some patients have exhibited functional meaningful improvements in clinically-relevant metrics such as visual acuity, GA growth, and reading speed
- Potential for recurring revenues, but with multiple treatments years apart
- Potential application in other retinal diseases (example: Stargardt's Disease)
- Issued patents cover aspects of production, product characterization, and formulation
- Fast Track designation from FDA
- Validating development partnership with global ophthalmology leader, Genentech









Source: christopherreeve.org



## Why Spinal Cord Injury (SCI) Matters



Lucas Linder, an OPC1 clinical trial participant, was paralyzed from the neck down.

The next year, he threw out the first pitch at a Major League Baseball game.



## Spinal Cord Injury (SCI) Overview

#### Lifetime care for an SCI patient can cost nearly \$5 million

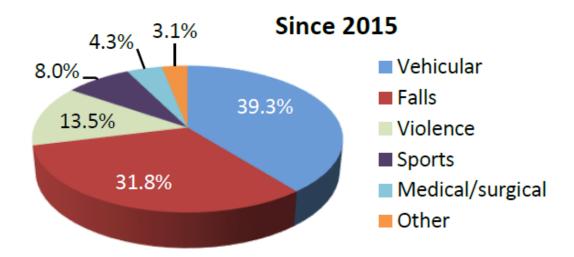
#### Incidence

Approximately 18,000 new cases in the U.S. each year

#### Prevalence

Between 249,000 and 363,000 people in the US

#### Causes





#### **SCI Burden and Unmet Needs**

#### A significant burden for patients and caregivers\*

- 67% of patients are unemployed 10 years post-injury
- Lifetime healthcare costs can reach
   \$5M for one patient

#### Potential lifelong impairments

- Mobility (wheelchair)
- Pain
- Re-hospitalizations
- Infections
- Ventilator dependency
- Depression
- Shortened life expectancy

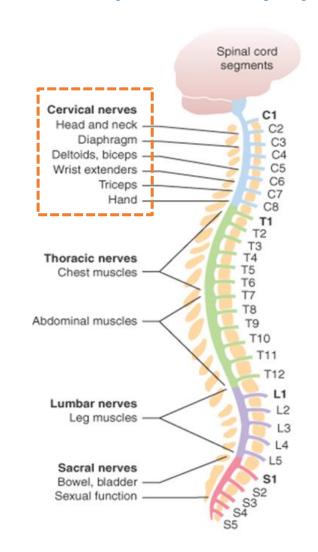




#### **SCI Treatment Objectives**

#### Loss of movement is the primary feature of a spinal cord injury

- Higher-level injuries result in more extensive impairments
- Gains in motor function, particularly in the upper extremities, can provide significant benefits in self-care and lower costs of care
- The goal of Lineage's cell therapy is to provide additional arm, hand, and finger function, increasing independence and quality of life

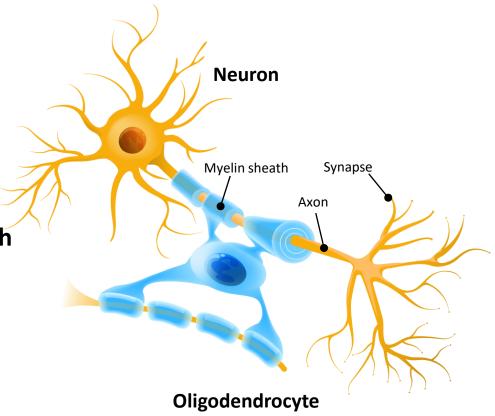




### OPC1 cells for Spinal Cord Injury

## Transplanting oligodendrocytes may provide additional upper extremities function (arms and fingers) and improve quality of life

- OPC1 is comprised of OPCs (oligodendrocyte progenitor cells)
- OPCs are precursors to
   Oligodendrocytes, the myelinating
   cells of the central nervous system
   which provide insulation to nerve
   axons in the form of a myelin sheath
- Myelin is essential for proper function of neurons
- OPC1 cells are implanted into the spinal cord at the injury site





#### **OPC1** Asset Overview

- OPC1 utilizes targeted cell replacement (similar approach as OpRegen)
- OPC1 is covered by multiple issued patents
- OPC1 has RMAT Designation
- OPC1 has Orphan Drug Designation
- OPC1 has received >\$14M in support from CIRM (California Institute for Regenerative Medicine)
- OPC1 has application to other demyelinating conditions

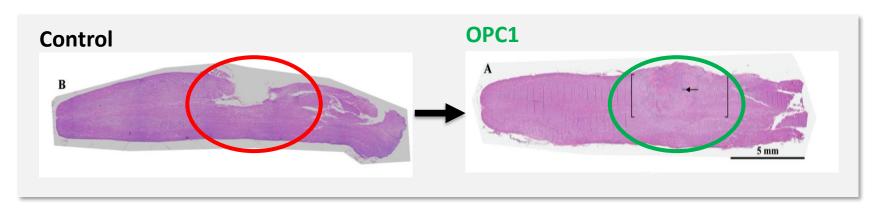


**OPC1 Transplant Procedure** 



#### **OPC1** Mechanisms of Action

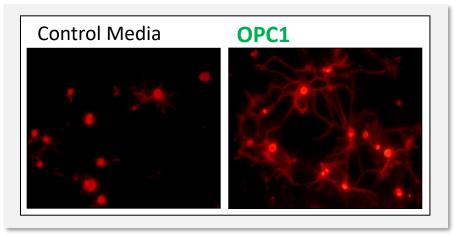
## **Suppression of Cavitation**



## **Myelination of axons**



## **Secretion of neurotrophic factors**





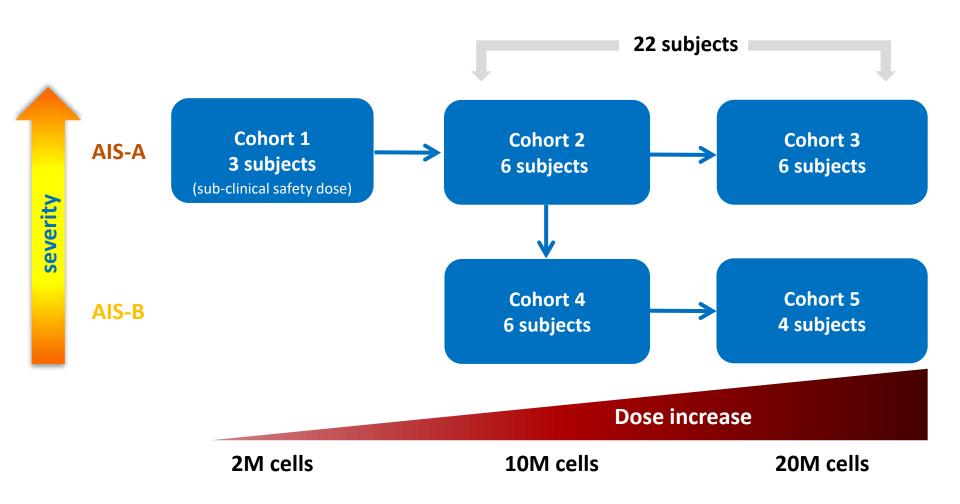
## **OPC1** for Spinal Cord Injury

- Lineage's OPCs are derived from an NIHregistered cell line
- The OPCs are allogeneic ("off the shelf"), and not taken from the patient
- Treatment of SCI occurs <u>3-6 weeks</u> postinjury and includes short-course (60-day) immunosuppression
- The OPCs are "ready to use" in a cryopreserved thaw-and-inject formulation





## SCiStar Clinical Trial Study Design





#### SCiStar Clinical Trial - Summary of Adverse Events

#### Majority of adverse events were mild to moderate in severity

All Treated Subjects (n=25)	AEs	SAEs	
Total	534	29	
Related to OPC1	1*	0	
Related to Injection Procedure	20	1	
Related to Tacrolimus	11	1	

To date, there have been no serious adverse events related to the OPC1 cells Safety data is available for 2 to 5 years on all 25 patients



#### SCiStar Clinical Trial - Cell Engraftment

#### 12- and 24-Month MRI Scans Indicate Durable Engraftment

- Cystic cavitation (syringomyelia) occurs in ~80% of SCI cases
- MRI results suggest formation of a tissue matrix at the injury site, indicating that OPC1 cells have durably engrafted and helped prevent syringomyelia
- 96% (24/25) of OPC1 patients had serial MRI scans that indicated <u>no</u> <u>sign</u> of a lesion cavity at 12 months (or 24 months for 22 scans available)

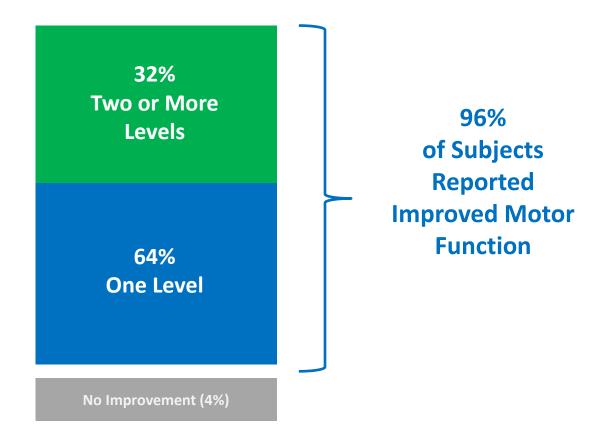


Weighted sagittal MRI



#### SCiStar Clinical Trial - Motor Function Gains

#### 22 Patients at 12 months





## INTERNATIONAL STANDARDS FOR NEUROLOGICAL CLASSIFICATION OF SPINAL CORD INJURY (ISNCSCI)



Patient Name\_\_\_\_\_\_ Date/Time of Exam \_\_\_\_\_\_

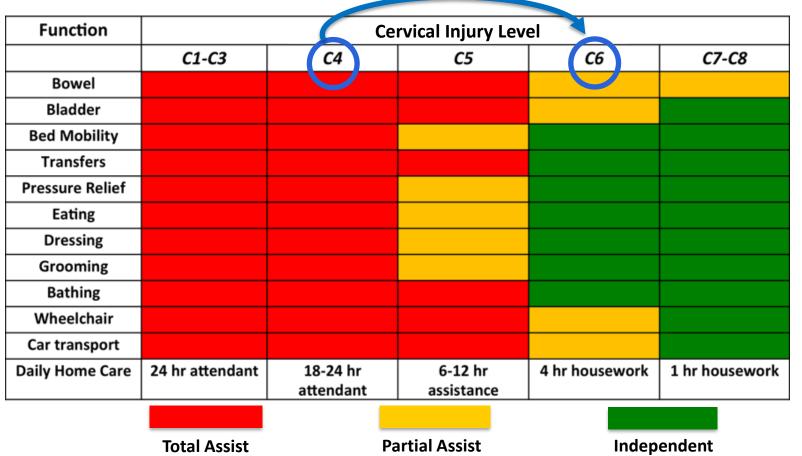
Examiner Name \_\_\_\_\_ Signature \_\_\_\_\_

RIGHT MOTOR KEY MUSCLES SENSORY KEY SENSORY POINTS Light Touch (LT) Pin Prick (PP)	SENSORY KEY SENSORY POINTS Light Touch (LT) Pin Prick (PP)  LEFT
C2 C3 C4	C2 C3 C4
UER Wrist extensors C6 (Upper Extremity Right) Elbow extensors C7 Finger flexors C8 Finger abductors (little finger) T1	C5 Elbow flexors C6 Wrist extensors UEL C7 Elbow extensors (Upper Extremity Left) C8 Finger flexors T1 Finger abductors (little finger)
Comments (Non-key Muscle? Reason for NT? Pain?):  T3  T4  T5  T6  T7  T8  T9  T10  T11  T12  L1  Hip flexors L2  L1  Long toe extensors L3  Long toe extensors L5  Ankle plantar flexors S1  S2  S3  S4-5  (VAC) Voluntary anal contraction (Yes/No)  RIGHT TOTALS  (MAXIMUM) (50) (56) (56)	MOTOR (SCORING ON REVERSE SIDE)  0 = total paralysis 1 = palpable or visible contraction 2 = active movement, against gravity 4 = active movement, against some resistance 5 = active movement, against some resistance 5 = normal corrected for pain/disuse NT = not testable  110 SENSORY (SCORING ON REVERSE SIDE)  111 SENSORY (SCORING ON REVERSE SIDE)  112 0 = absent
(50)	LT $\longrightarrow$ + LLT $\longrightarrow$ = LT TOTAL $\longrightarrow$ RPP $\longrightarrow$ + LPP $\longrightarrow$ = PP TOTAL $\longrightarrow$ MAX (56) (56) (112)
LEVELS 1. SENSORY LEVEL OF INJURY Incomplete = An	PLETE OR INCOMPLETE?  (In complete injuries only)  ZONE OF PARTIAL  PRESERVATION  MOSTOR  MOTOR  MOTOR

#### Real-World Benefit from a 2 Motor Level Improvement

Motor level gains translate into clinically meaningful improvements in self-care and reductions in cost of care

32% had +2 Level Improvement





## SCiStar Clinical Trial - Analysis of Patients with Least UEMS Recovery

## C4 or cord compressions occurred in 5 of the 7 worst patient outcomes and both issues can be addressed in the next trial

Subject	UEMS Change at 12 mo.	Cord Compression After OPC1 Injection?	NLI Baseline	Baseline AIS	Cohort	Dose	Age	Injection Days Post Injury
2207	7	N	C4	В	5	20 M	62	37
2203	6	N	C6	А	3	20 M	45	31
2105	6	N	C4	А	3	10 M	19	20
2004	5	N	C6	В	4	10 M	21	25
2007	4	N	C4	В	4	10 M	55	38
2307	4	Υ	<b>C</b> 5	В	5	10 M	19	38
2303	3	Υ	C6	В	4	10 M	22	35

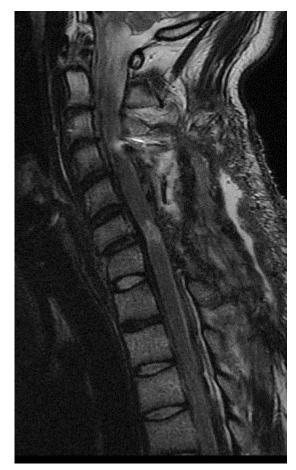
- Two patients had cord compression after OPC1 injection (2303 and 2307 at Day 30 and Day 7)
- Patients 2105, 2207, 2007 had a C4 (highest/most severe) injury level at baseline
- Patient 2105 also had a hematoma in the spinal cord at baseline & a failed graft



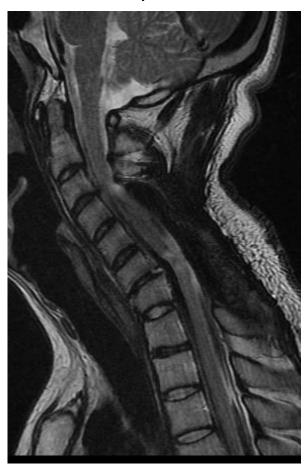
## SCiStar Clinical Trial – Cord Compression

## Subject 2303 (Cohort 4): Cord Compression at Day 30

Baseline Day 30 Day 365









#### SCiStar Clinical Trial – Takeaways

- Excellent overall safety profile
- 96% durable engraftment confirmed via MRI
- MRI scans through 24 months show no evidence of adverse changes
- No subjects had a decline in motor function from Year 1 to Year 2
- 95% of patients exhibited motor recovery in the upper extremities at 12 months (requires at least 1 motor level gain on at least 1 side)
- Significant motor improvements achieved in five of six Cohort 2 subjects
- The two worst performing subjects had spinal cord compression (can be addressed in next trial)



#### New Parenchymal Spinal Delivery (PSD) System

- Better performance: Greater stability and control
  - Eliminates motion between platform/XYZ manipulator/injection needle
  - Pump and syringe not in sterile field: programmed (accurate) dose rate
- Enhanced clinical usability and safety: no cessation of ventilation
  - Attaches directly to the patient, compatible with breathing motion
  - Magnetic needle provides stabilization of micromotion from heartbeats
- Enhanced end user experience:
  - Smaller and fewer components
  - Easily assembled
  - Single hand operation
  - Accurate needle depth insertion
  - Compatible with OPC1 TAI formulation
- Animal testing with OPC1 ongoing
- Clinical trial in sub-acute and chronic patients planned for 2022





### OPC1 Manufacturing Improvements Following FIM Study

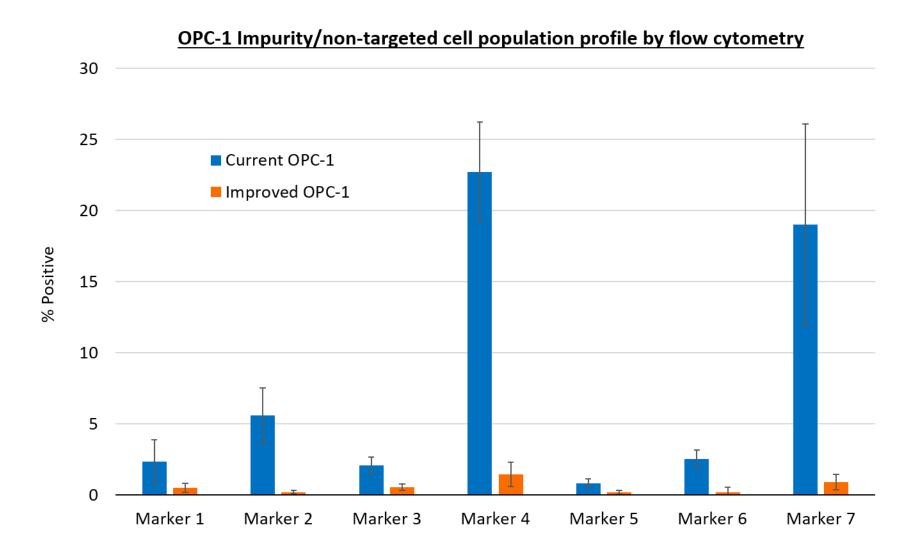
#### Lineage has made major improvements in production and quality of OPC1

- A new ready-to-inject formulation was developed
- Elimination of dose preparation achieved
- 10- to 20-fold increase in production scale
- Significant reduction in impurities
- No reduction in functional activity
- 12 new analytical and functional methods developed
- Elimination of all animal-based production reagents
- Patent applications on the process and product, if allowed, will have expiration dates of 2039 and 2040





## **OPC1** Manufacturing Improvements: Lower Impurities





#### OPC1 Program – Key Takeaways

- Excellent overall safety profile
- 96% durable engraftment confirmed via MRI
- 95% of patients exhibited motor recovery in the upper extremities at 12 months (requires at least 1 motor level gain on at least 1 side)
- 5/6 patients in cohort 2 achieved significant (2 level) motor improvements
- The two worst-performing patients had spinal cord compression, which can be addressed in the next trial
- Greatly improved product purity and scale
- Superior delivery device is entering clinical testing (safety trial to include chronic patients)
- Planning underway for a randomized, controlled and potentially registrational clinical trial







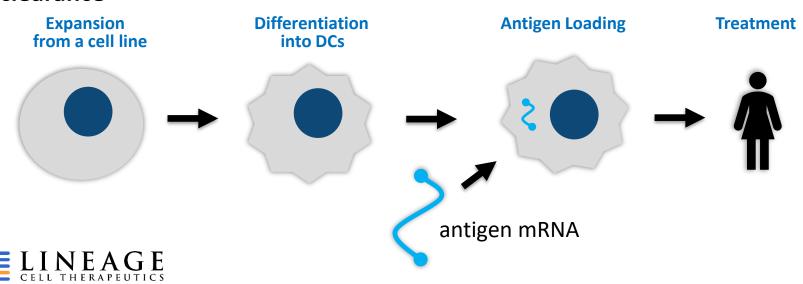
Immunotherapy is "poised to revolutionize treatment for all types of cancer"

Source: cancerresearch.org

VAC: A Cell Therapy Platform for Cancer and Infectious
Diseases

#### The VAC Platform: On demand cell therapy for cancer

- The VAC platform consists of large-scale, allogeneic ("off the shelf") production
  of mature dendritic cells (DCs). No production delay between diagnosis and
  treatment, as with autologous or patient-specific therapies.
- DCs are manufactured and loaded with either a tumor antigen (to treat cancer) or a viral antigen (as a vaccine for infectious diseases)
- Antigen presentation to the patient's T cells creates a targeted and robust immune response (up to 3%), aiding tumor cell destruction or pathogen clearance



#### VAC Development – A Platform for Multiple Product Candidates

# **Mature Dendritic** Lineage **Manufacturing Cells Platform**

#### **VAC1** and **VAC2** Highlights

- Positive phase 1 data in AML
- Positive ongoing phase 1 trial in lung cancer (NSCLC)
- Cancer Research UK alliance
- High T cell responses in clinical trials

#### VAC3, VAC4, VAC5...Opportunities

- Partnerships based on new products
- Retain highest value candidates
- Currently evaluating new antigens

#### **VAC-Infectious Diseases**

- Designed to provide long-term protection via memory T cells
- Leverages VAC clinical data



Selected

Antigen

#### VAC2 - Phase 1 Clinical Trial (Ongoing, Conducted by CRUK)

- Enrollment ongoing (7/8 patients treated to date)
- VAC2 has been well tolerated in all patients dosed; no treatment delays due to adverse events attributable to VAC2
- Directionally positive Phase 1 data
  - Induction of durable, antigen-specific linked T cell help
  - Magnitude of T cell induction 40-400 times higher than that witnessed with other approaches (DNA / RNA vaccines)
  - Adverse events suggestive of induction of an adaptive immune response
  - Injection site reactions, flu-like symptoms (all grade 1 or 2)
- Major radiological response reported in 1 patient > chemotherapy > VAC2
- Safety & preliminary immunogenicity data obtained in Q1 2020 formed basis for Lineage to exercise option to re-acquire asset and advance internally



#### **VAC Platform Next Steps**

#### **Upcoming Events and Key Considerations:**

- Complete dosing in ongoing clinical trial (1 patient remaining)
- Introduce improvements to the manufacturing process
- Design new products (i.e. VAC3, 4, 5, 6...) with newly discovered antigens
- Identify potential partnership and grant opportunities for more rapid expansion of the VAC platform
  - First strategic alliance (with Immunomic Therapeutics) announced April 2021



## Our Goal is to Provide Life-Changing Cell Therapies to Patients

# Lineage Cell Therapeutics: Bringing the Promises of Cell Therapy into Clinical Reality







World class
in-house
process
development
and GMP
manufacturing



One of the largest patent portfolios in cell therapy



Multiple validating corporate partnerships



Leader in the field of regenerative medicine



## The Patients Are Our Inspiration. View their stories at lineagecell.com/media/#patients

#### **OPC1 SCiStar Study Participants**





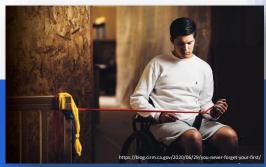
#### **Lucas Lindner**

"There's no reason to not look forward in the same way now that I had before all of this happened. I'm looking forward to driving again... it's a bright future."



#### Kris Boesen

"I couldn't drink, couldn't feed myself, couldn't text or pretty much do anything, I was basically just existing. I wasn't living my life, I was existing."



#### **Jake Javier**

"Even though it's a completely different perspective, I can still lead that way. I can just try to be the best I can and to persevere the best I can."

Diablo Magazine, Feb. 16, 2017

# The Millions Worldwide Suffering from Dry AMD Vision Loss

"Macular degeneration is a very frustrating condition which can greatly affect your day-to-day life."

- Macular Society



