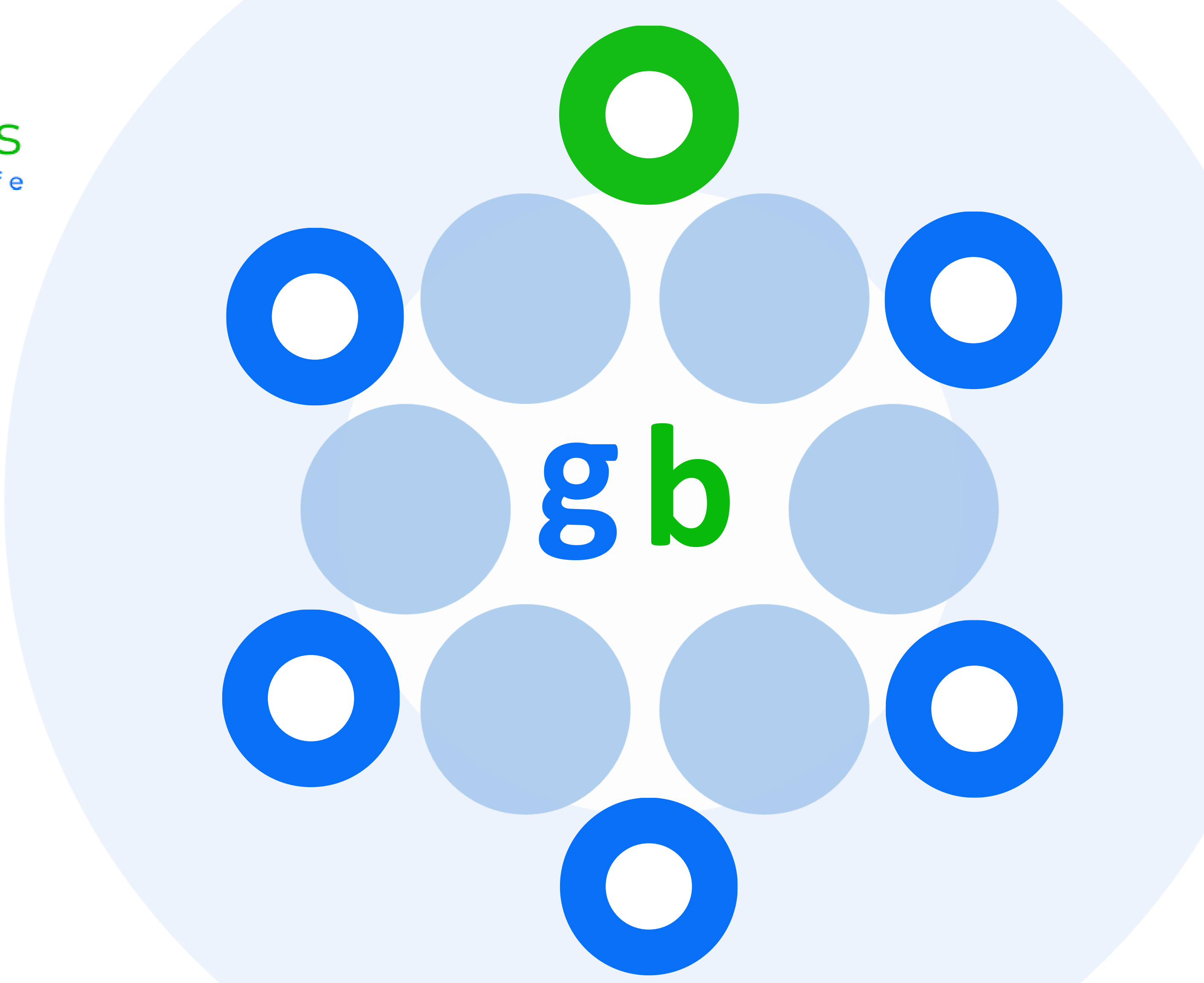


genflow
biosciences
longer better life

CORPORATE OVERVIEW

October 2024

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MARKET

Aging Is One of Our Greatest Societal & Economical Challenges

Increasing life expectancy, decreasing healthspan, rising healthcare costs – all highlight the urgent need for age-related disease treatments



88 yrs*



12-13 yrs*



2 yrs*

LIFE EXPECTANCY

***expected LE in relation to male babies born in the UK in 2018**

Source: Morgan AE, Davies TJ, Mc Auley MT. The role of DNA methylation in ageing and cancer. Proc Nutr Soc. 2018 Nov;77(4):412-422. doi: 10.1017/S0029665118000150. Epub 2018 Apr 30. PMID: 29708096

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WHO WE ARE

Pioneering Novel Gene Therapeutics for a Longer, Healthier Life

MISSION: aging is underlying risk factor for disease, and our aim is to deliver gene therapeutics that potentially halt or slow the aging process in humans and dogs

SIRT6 GENE: longevity protein that widely regulates aging and immunity and is a potential therapeutic target for the treatment of diseases

PROMISING PRECLINICAL RESULTS: lead drug candidate GF-1002 delivers a centenarian variant of the SIRT6 gene for treatment of MASH

SEASONED TEAM: experienced and proven management with extensive experience in public & private pharma and biotech

ROBUST PIPELINE: advancing multiple development programs over 24 months: MASH, Sarcopenia, Werner and life extension of dogs

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DEVELOPMENT PIPELINE

GF-1002 (Pre-Clinical) –Exo-AAV vector for intravenous infusion, expressing cDNA of centenarian variant of SIRT6 in liver

GF-1003 (Pre-Clinical/Development) Suspension of exosomes, expressing mRNA of centenarian variant of SIRT6 in fibroblasts

GF-1005 (Pre-Clinical/Development) Mitochondrial dysfunction: Myoblast progenitors loaded by photoporation with centSIRT6

GF-1004 (Pre-Clinical) SIRT6c/ NAD+ for intravenous infusion for life extension and prevention of age-related diseases

Phase I/II MASH

PoC in 36 patients
IND-Enabling Phase
18 months to first-in-human

Werner Syndrome

36 months to first-in-human

Sarcopenia

Pre-clinical

Anti-aging for dogs

w/ Vet Partner

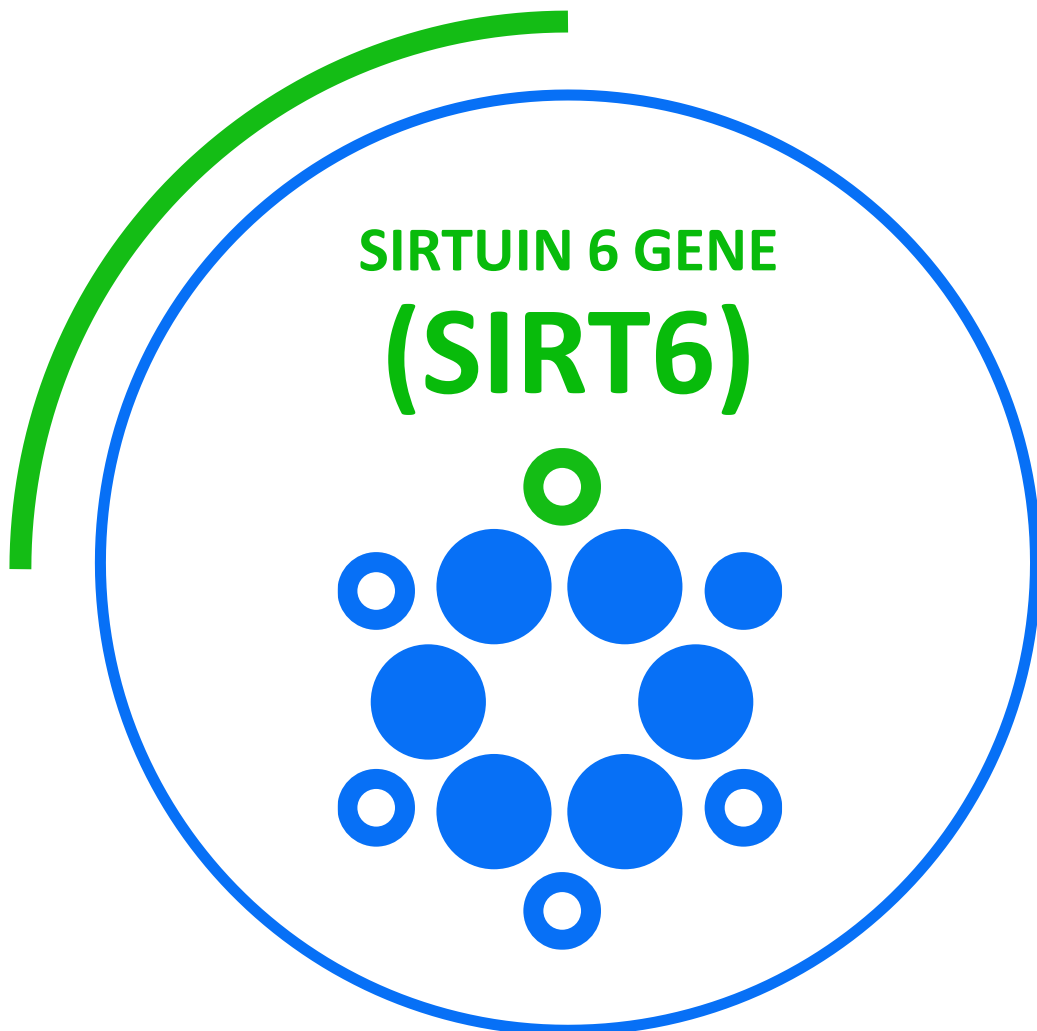
COLLABORATIONS



GENE REGULATION IN AGING

Aging is a function of overworked epigenetic regulator genes unable to respond to cellular DNA damage

MANY GENES REGULATE AGING.
OUR FOCUS IS THE **SIRT6 GENE**



genflow biosciences

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Aging is driven by interlinked Hallmarks, all rooted in DNA damage. Targeting one individual factor is unlikely to be effective



SIRT6: REPAIRING DNA

SIRT6 gene/protein repairs DNA damage (especially double strand breaks (DSB)) and prevents senescence of our cells

SIRT6 gene codes for SIRT6 protein

Stronger SIRT6: Longer lifespan

The Ability of SIRT6 to stimulate DSB repair correlates with maximum lifespan (MLS) in rodents

5 Amino Acids determine the differential activities of SIRT6

SIRT6 EFFICIENCY DOUBLE STRAND BREAK REPAIR

LIFE **LIFESPAN**

Lifespan
▲
DSB repair
▲
SIRT6 activity

5 amino acids



Source: Tian et al., 2019, Cell 177, 622–638 April 18, 2019

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FOCUS ON CENTENARIAN SIRT6

SIRT6 centenarian variant gene has more efficient DNA repair properties

HOMOLOGOUS

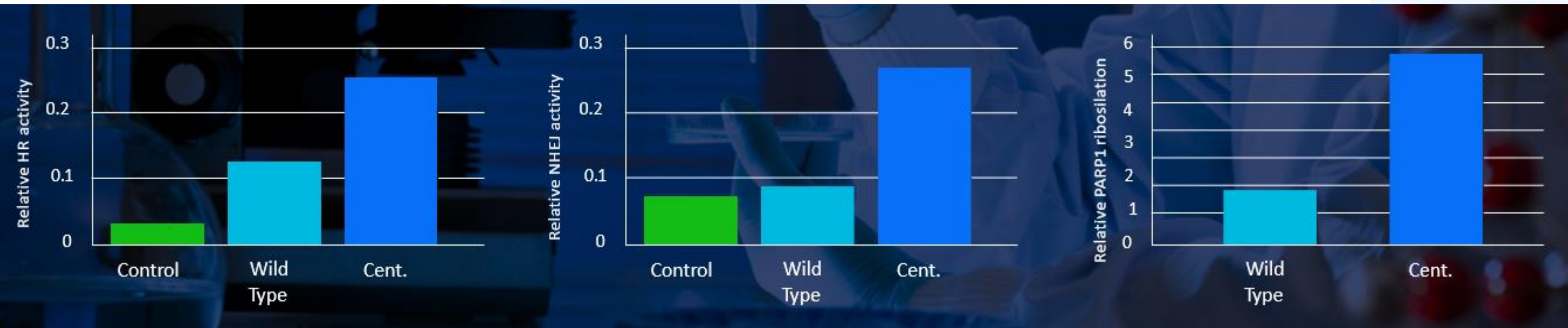
Recombination Repair

NON-HOMOLOGOUS End

Joining Repair

RELATIVE PARP1

Ribosilation



MASH PROGRAM

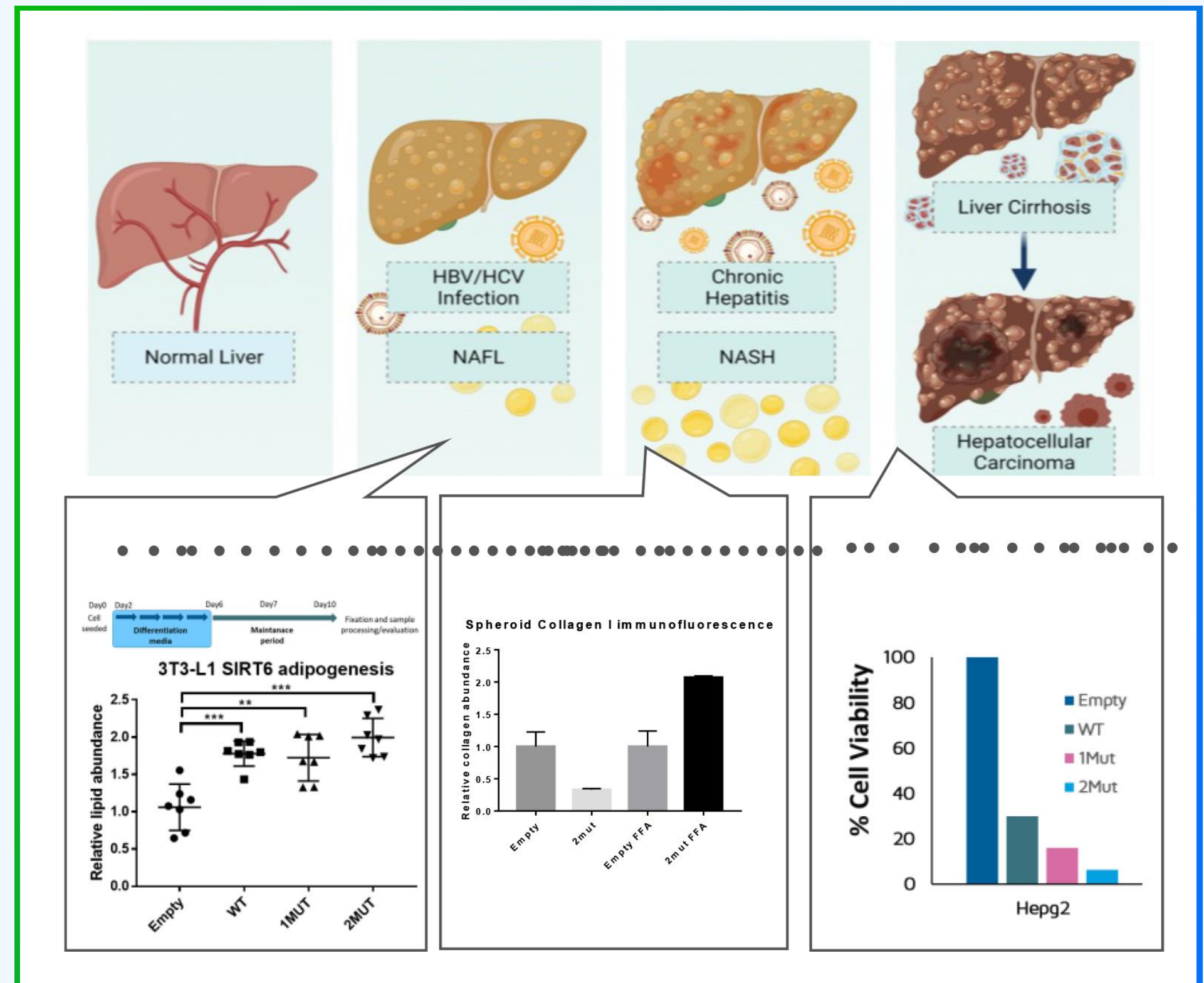
Affects est. 35 million people globally

- Increasing prevalence
- Leading cause of chronic liver disease and liver transplant

Significant unmet medical need

Clear regulatory accelerated development pathway. EMA and FDA guidelines accept:

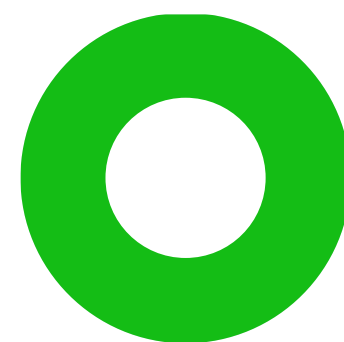
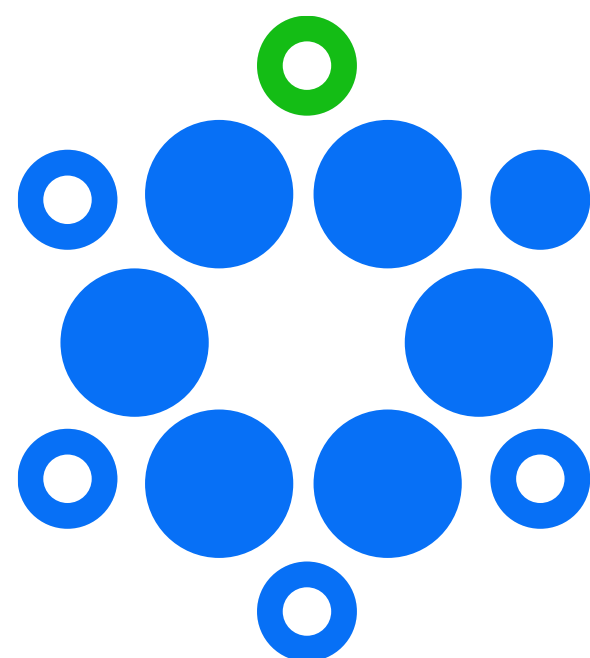
- ✓ Key surrogate outcomes for therapeutic trials: regression of fibrosis or resolution of NASH
- ✓ Histological changes are achievable within a 12-18-month time-frame
- ✓ Placebo control
- ✓ Conditional fast-track approval



Pais R, Barritt AS 4th, Calmus Y, Scatton O, Runge T, Lebray P, Poynard T, Ratziu V, Conti F. NAFLD and liver transplantation: Current burden and expected challenges. *J Hepatol.* 2016 Dec;65(6):1245-1257.
Vlad Ratziu, Sven Francque, Arun Sanyal, Breakthroughs in therapies for NASH and remaining challenges, *Journal of Hepatology*, Volume 76, Issue 6, 2022

EXOSOME DELIVERY SYSTEM: SAFE AND COST-EFFECTIVE

Genflow's patent-pending technology has already been tested in several preclinical studies



**Non-Integrating,
No Risk
of Insertional
Oncogenesis**

**Non-Replicating,
Safe Transient
Expression**

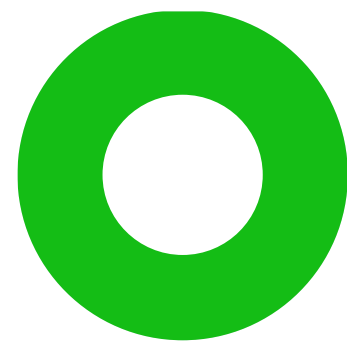
**Reduced
Potential for
Immunogenicity**

ADVANTAGES: EXOSOME DELIVERY

Exo-AAV can mediate efficient, specific, and more durable SIRT6 expression in liver compared to conventional AAV

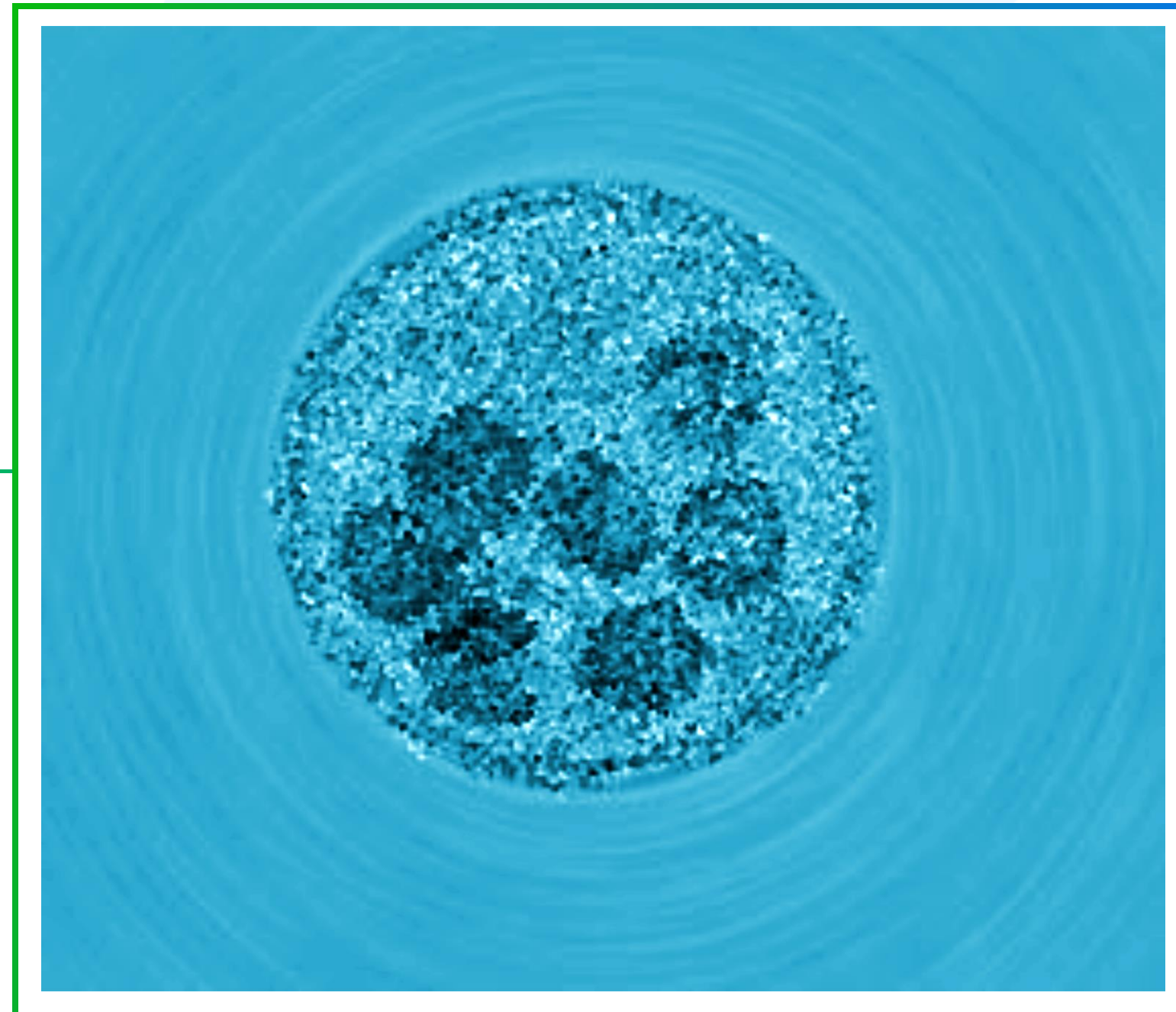
NO IMMUNOGENICITY

Lack of Local Systemic Immunogenicity



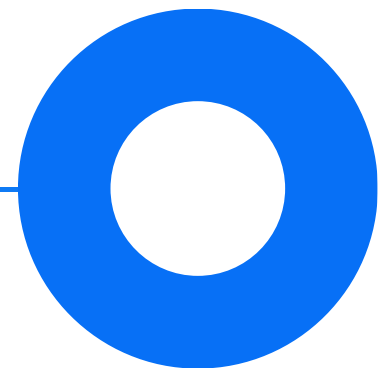
TARGETED DELIVERY

Engineered Exosome
To Direct to Specific
Cell Types



POTENCY ADVANTAGE

Improved Transduction
Verses Free AAVs Rapid Uptake
Sustained



THERAPEUTIC WINDOW

Potency Improvement,
Local Retention, Lack
of Systemic Leakage

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Hudry E, Martin C, Gandhi S, György B, Scheffer DI, Mu D, Merkel SF, Mingozi F, Fitzpatrick Z, Dimant H, Masek M, Ragan T, Tan S, Brisson AR, Ramirez SH, Hyman BT, Maguire CA. Exosome-associated AAV vector as a robust and convenient neuroscience tool. *Gene Ther.* 2016 Apr;23(4):380-92. doi: 10.1038/gt.2016.11. Epub 2016 Feb 2. Erratum in: *Gene Ther.* 2016 Nov;23 (11):819. PMID: 26836117; PMCID: PMC4824662.

2023 KEY HIGHLIGHTS & 2024 PRIORITIES

GROWING IP PORTFOLIO

Provisional patent application in 2023 for editing **SIRT6** gene, linked to longevity and age-related diseases

EXPANDED MARKET

Awarded to new grants in 2023 & 2024 expanding research pipeline and size of therapeutic markets

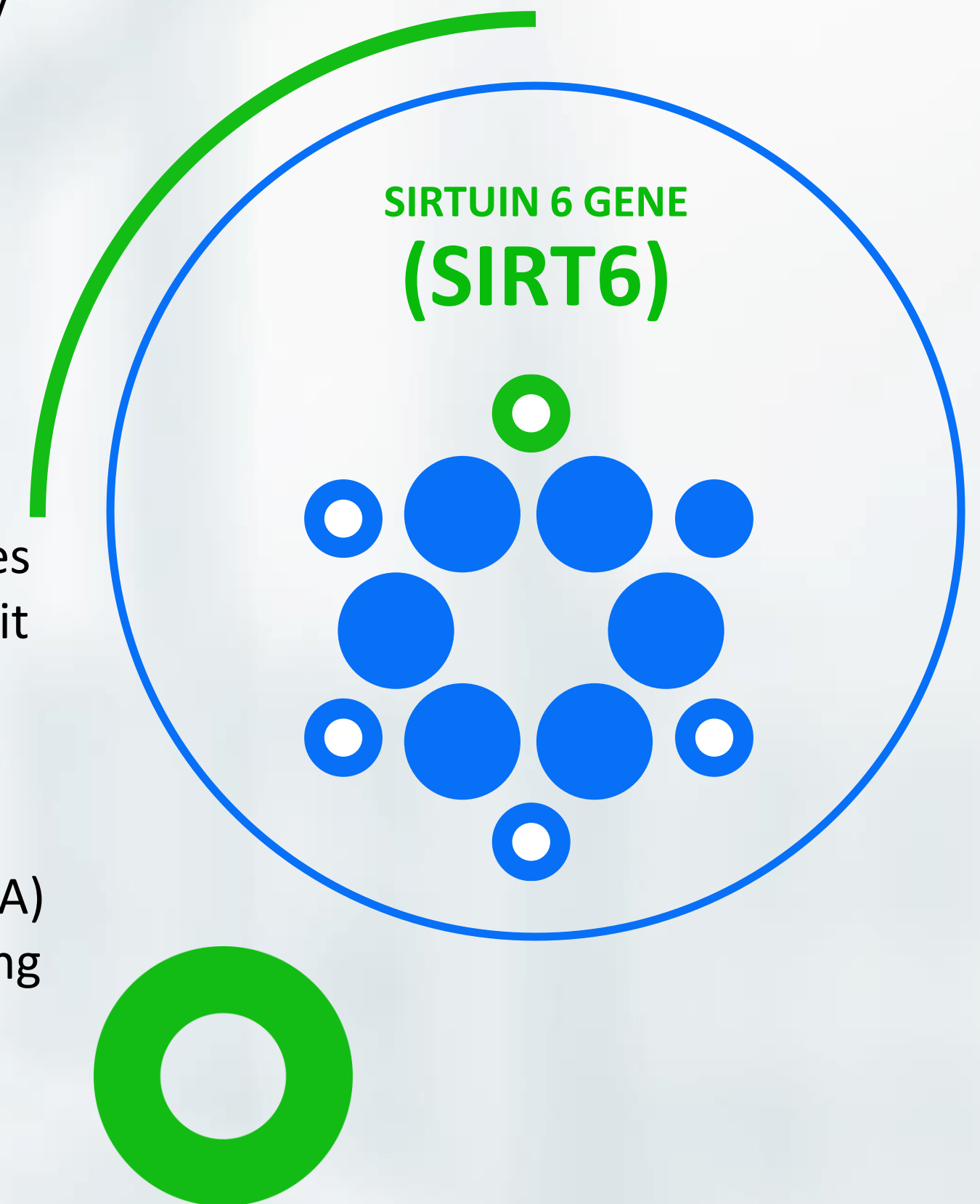
GF-1002

Undertaking key Investigational New Drug (IND) -enabling development activities to help define pharmacological and toxicological properties and potential benefit to NASH patients

GF-1003

Commencing preliminary discussions with the European Medicines Agency (EMA) on Mechanism of Action (MoA) data for Orphan Drug Application (ODA) targeting Werner Syndrome

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2023 KEY HIGHLIGHTS & 2024 PRIORITIES

LARGE MARKET OPPORTUNITY

NASH: 35 Million globally. Increasing prevalence. Door opener to even broader anti-aging indication

LONG LIFE IP

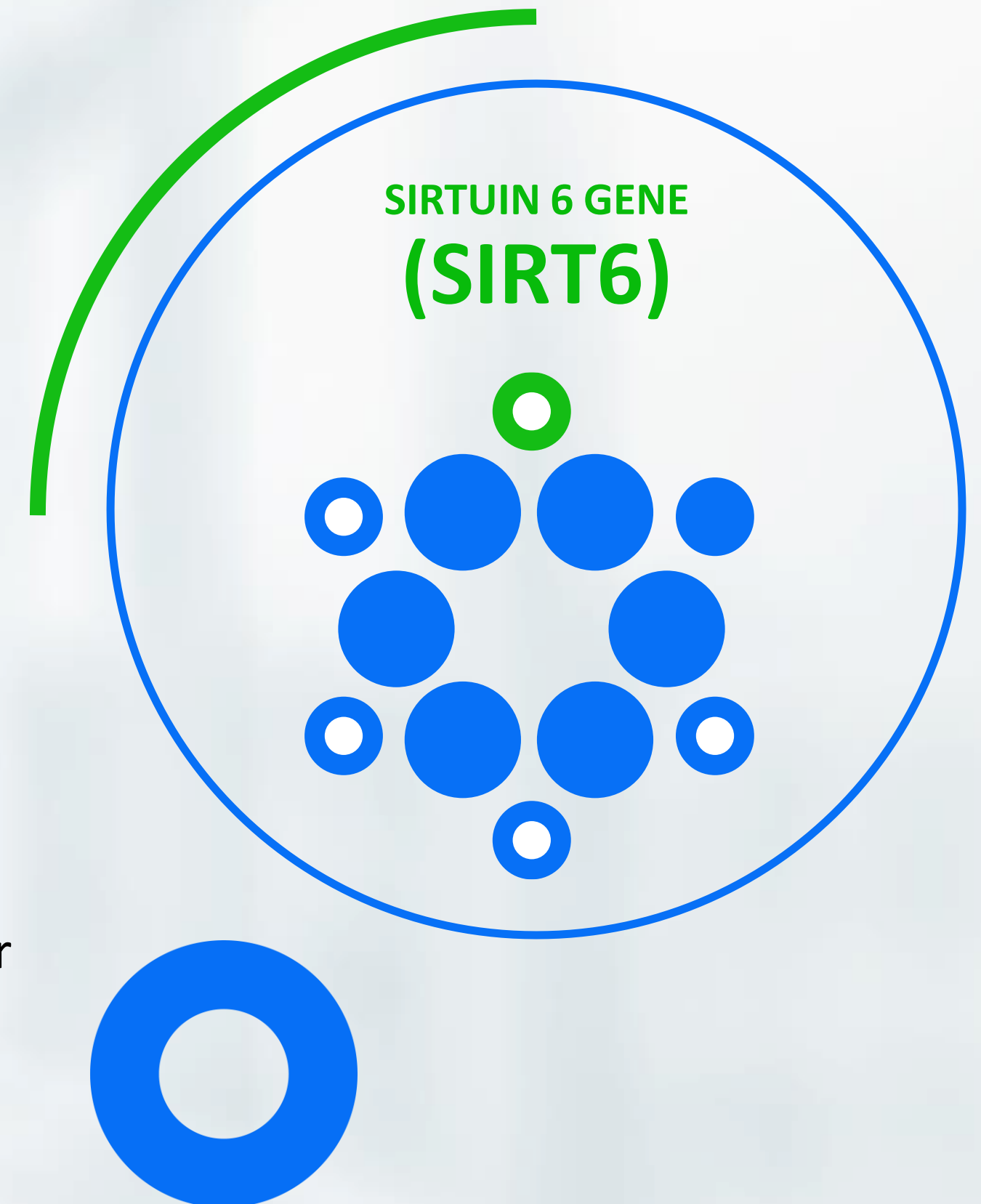
2 patent families **SIRT6** centenarian and gene delivery (entering National Phase); Additional upcoming patent applications (entering PCT)

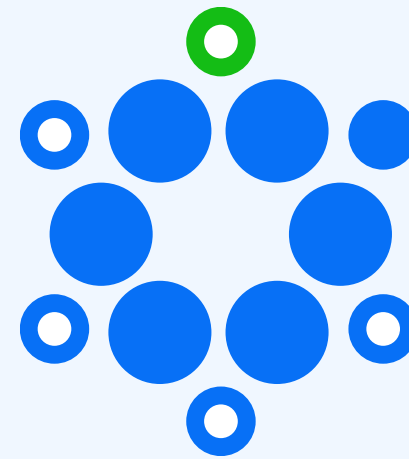
GENE DELIVERY SYSTEMS

Proprietary innovative gene delivery systems: exo-AAV, mRNA exosomes, LNPs Centenarian variant of SIRT6 gene

GF-1002 and GF-1003

Multiple key clinical and regulatory milestones expected in next 18 months
Undervalued stock opportunity; Potential acquisition by pharmaceutical partner





TAMARA JOSEPH
Chairperson

- Seasoned healthcare leader with extensive experience in both early-stage and commercial biotech companies
- Supported Nasdaq financings of over \$800m
- Currently serving as Chief Legal Officer at Spero Therapeutics Inc. (NASDAQ:SPRO)
- Served as an adviser to the boards of five US publicly traded biotechs, including Cubist Pharmaceuticals Inc.
- BA in Economics from Duke, a JD from the University of Michigan, and LLM degrees from Belgium and the University of Paris



DR ERIC LEIRE MD MBA
Founder & CEO

- MD and MBA, Eric has been involved in biotech for over 30 years
- Held senior positions including CEO of publicly traded biotech companies (Nasdaq, OTC.QB, OMX.Nasdaq)
- Inventor of several patents and author of medical peer-reviewed publications



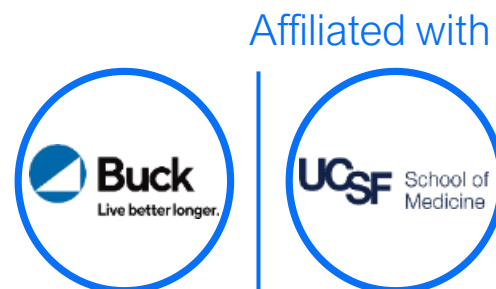
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NASH Expert

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Chicago Medicine










INTELLECTUAL PROPERTY



EFS ID	1-21069	43268050
Application Number	US 63/188,573	US 63/222,557
Title of Invention	Variants of SIRT6 for use in preventing and/or treating age-related diseases	Method of in vivo administration of the coding sequence of the SIRT6 gene via Adeno-Associated-Virus
First Named Inventor	Vera Gorbunova, Seluanov and Suh	Eric Leire
Receipt Date	May 14, 2021	July 16, 2021
Ownership	Worldwide Exclusive license from University Rochester New York / Columbia University / Albert Einstein College of medicine	Genflow Biosciences SRL

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LONGEVITY LANDSCAPE

COMPANY	OVERVIEW	TECHNOLOGY	FOCUSED ON	LOCATION
	CLINICAL STAGE, PHASE II NASDAQ (UBX) MKT CAP \$785M	Small Molecules Senolytic	Senescence	USA, San Francisco, CA
	PRE-CLINICAL STAGE NYSE (AGE) MKT CAP \$25M	Therapeutics that seek to address human aging	Stem cells	USA, Alameda, CA
	CLINICAL STAGE, PHASE I NASDAQ (VERV) MKT CAP \$885M	In Vivo LNP CRIPR Gene editing	Hypercholesterolemia	USA, Cambridge, MA
	PRE-CLINICAL STAGE NASDAQ (FREQ) MKT CAP \$16M	Small Molecules to Activate progenitor calls for MS	Stem cell exhaustion	USA, Woburn, MA
	PRE-CLINICAL STAGE PRIVATE RAISED \$124M	Epigenetic reprogramming	Mitochondrial dysfunction	USA, Boston, MA
	CLINICAL STAGE, PHASE III PRIVATE RAISED \$778M	Alternative splicing modulation to develop medicines to treat ageing-related diseases	Osteoarthritis	USA, San Diego, CA
	CLINICAL STAGE, PHASE I PRIVATE RAISED \$26M	Gene Therapy	Proteostatis	USA, San Carlos, CA

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THANK YOU

Contact

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