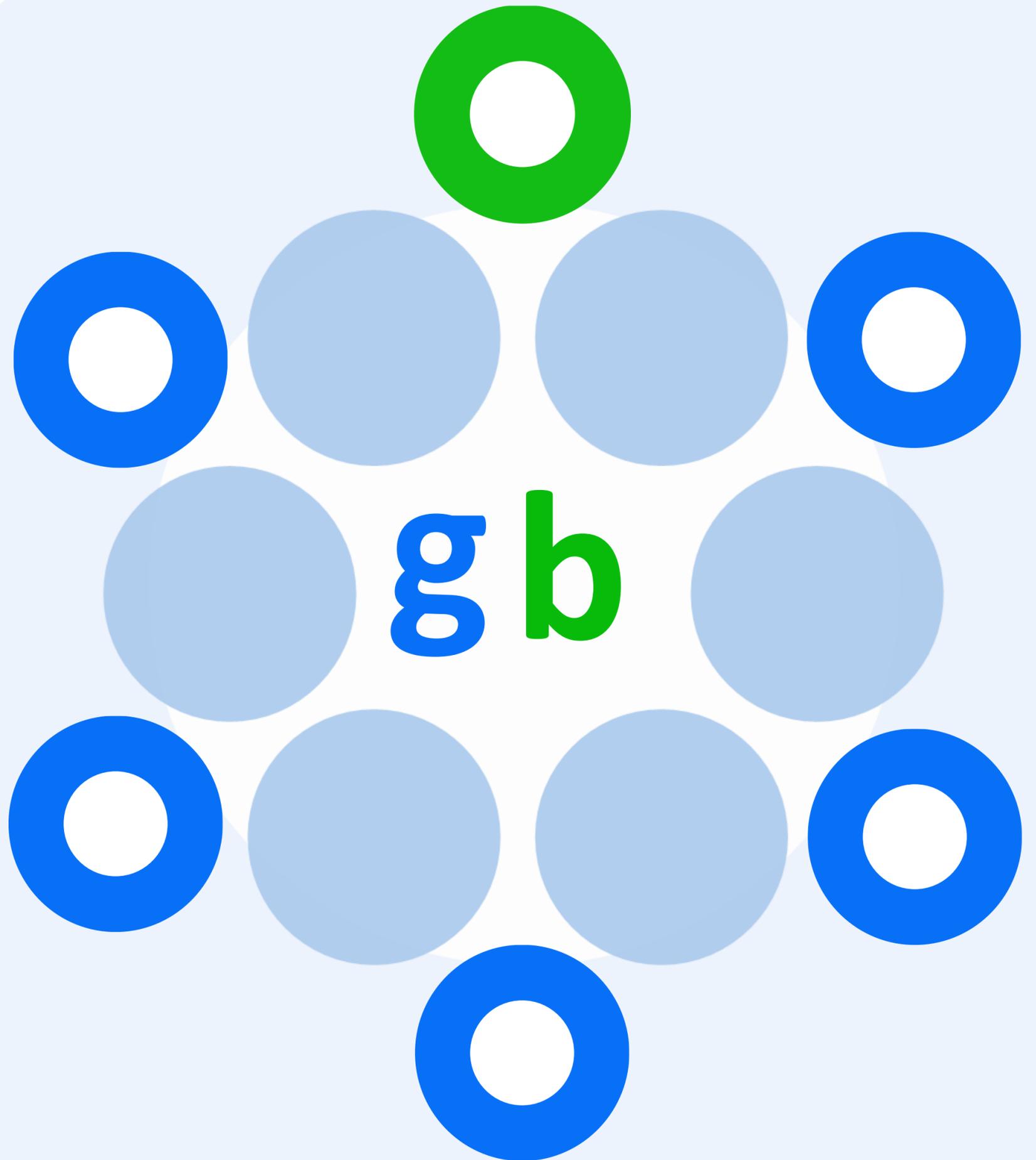


genflow
biosciences
longer better life

CORPORATE OVERVIEW

2025-2026

LSE: **GENF** - OTCQB: **GENFF**



FORWARD LOOKING STATEMENTS

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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

CENTENARIAN SIRT6 (cent SIRT6) GENE: proprietary variant of longevity gene 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 (cent SIRT6) 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, Glaucoma, Sarcopenia, Werner and life extension of dogs

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SCIENTIFIC ADVISORY BOARD



**DR. VERA
GORBUNOVA, PHD**

CO Director

Rochester Aging
Research Center
Affiliated With
Weizmann Institute
Of Science



**DR. ERIC VERDIN
MD/PHD**

CEO & President

Buck Institute
Affiliated With
UCSF School Of Medicine



**DR. MATTHEW
HIRSCHEY, PHD**

Assistant Professor

Duke University
School of Medicine
Affiliated With
American Heart Association



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VINCIGUERRA, PHD**

Principle Investigator

University of Liverpool
Affiliated With
UCL



**PROF. DR. SVEN
FRANCQUE, PHD**

NASH Expert

University of Antwerp



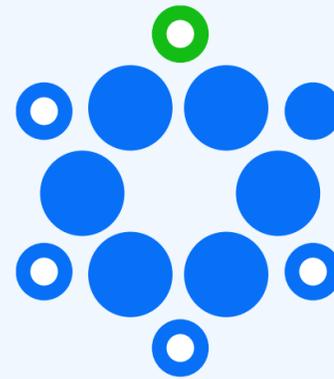
**DR. MARY
RINELLA, MD**

NASH Expert

University of
Chicago Medicine



MANAGEMENT



GAD BERDUGO Chairperson



- Managing Partner of Explorium Capital with 35+ years of leadership across global biotech business & corporate development, venture management and U.S. capital markets with a proven record at structuring and closing partnerships and capital raises
- Former C-level executive at private and public biotech companies, with financial advisory and investment management experience from Lazard and Tegriss
- Graduated from Imperial College London, University College London, and HEC Paris



DR ERIC LEIRE MD MBA Founder & CEO



- MD and MBA, Eric has been involved in biotech for over 30 years
- Research position at Harvard University. 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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SIRT6 GENE/PROTEIN REPAIRS DNA DAMAGE

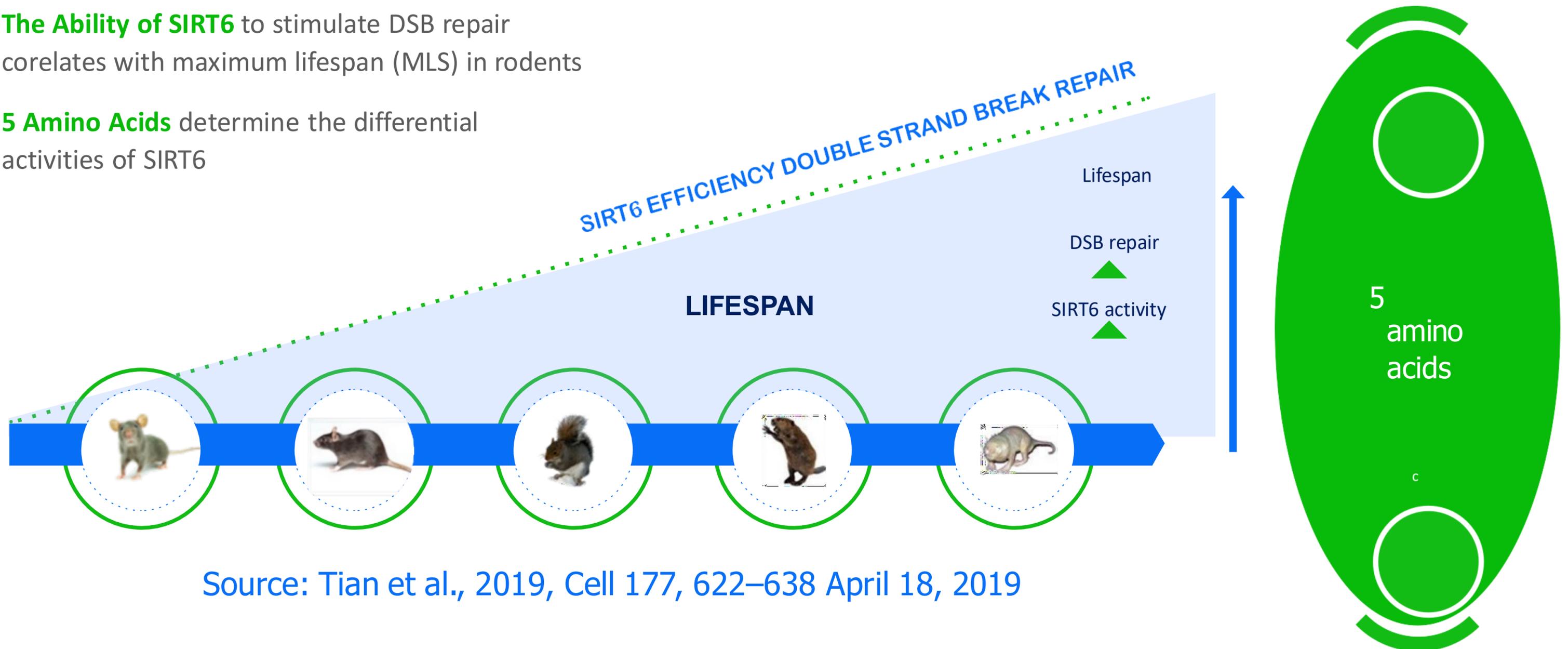
SIRT6: REPAIRING DNA

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

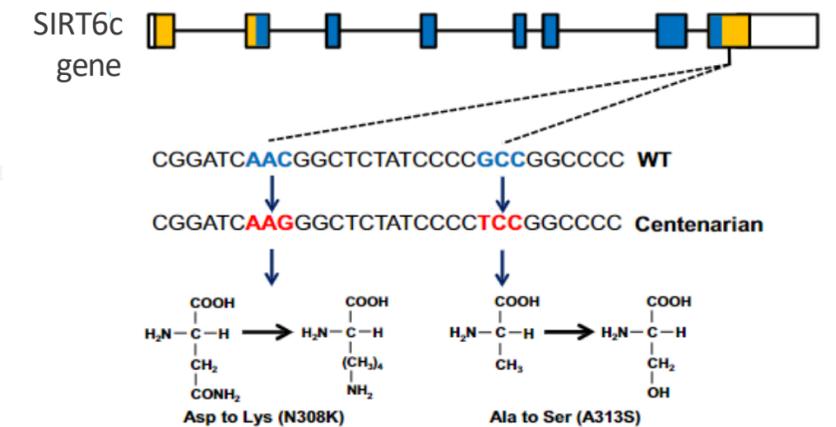
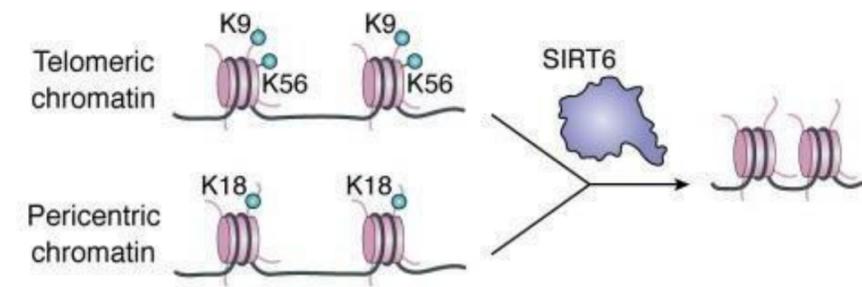
5 Amino Acids determine the differential activities of SIRT6



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

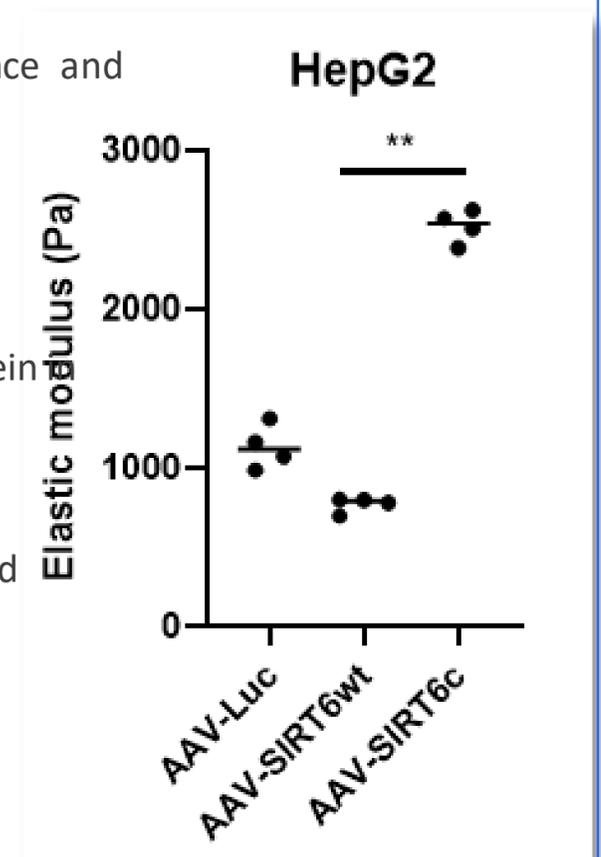
SIRT6CENT EDGE:CENTENARIAN GENETICS VALIDATED BENEFIT

- ❑ Sirtuins are a family of highly conserved signaling proteins involved in metabolic regulation and implicated in influencing cellular processes incl. aging,
- ❑ Sirtuin 6 (SIRT6) is a stress responsive NAD⁺-dependent histone deacetylase (HDAC) promoting increased longevity
- ❑ A new allelic variant of human SIRT6 with two point mutations (N308K/A313S) was recently associated with the longevity in Ashkenazi Jews (SIRT6cent)
- ❑ SIRT6cent confers various benefits, such as increased DNA repair capacity, enhanced metabolic regulation, and improved stress resistance.



SIRT6c vs SIRT6wt properties

- Enzymatic activity: SIRT6c displays weaker deacetylase activity and stronger mono-ADP-ribosyl transferase.
- Genomic stability/ DNA repair: Improved genome maintenance and DNA repair (Simon et al. EMBO, 2022).
- Anti-fibrotic effects:
 - Down-regulation of profibrotic genes expression (minor)
 - Reduced Col1A1 deposition in fibrotic conditions (minor)
 - Reduced hepatocytes cell stiffness
- Anti-inflammatory effects: Down-regulation of IL-1b and IL-6 protein stimulated culture conditions (n=3)
- HCC:
 - Modulation of ECM-related genes expression
 - Increased cell stiffness in hepatoma cells, associated to reduced invasives
- Transcriptomic analysis: Differential modulation of b-catenin/TP63 and glucocorticoids pathways



DEVELOPMENT PIPELINE

GF-1002 (Pre-Clinical) AAV8 cent SIRT6 vector for intravenous infusion, expressing cDNA of centenarian variant of SIRT6 (cent SIRT6) in liver

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

GF-1005 (Pre-Clinical/Development) Mitochondrial dysfunction: Myoblast progenitors loaded by photoporation with cent SIRT6

GF-1004 (Clinical Proof of Concept) cent SIRT6 for intravenous infusion for life extension and prevention of age-related diseases

GF-1006 Ophthalmology (pre-clinical) mRNA cent SIRT6 delivered by LNP for the treatment of cornea pathologies and glaucoma

Phase I/II MASH

IND-Enabling Phase

18 months to first-in-human PoC in 36 patients

Werner Syndrome

36 months to first-in-human

Sarcopenia

Pre-clinical

Anti-aging for dogs

w/ Vet Partner

Ophthalmology

w/ Partner

COLLABORATIONS



NEW PARTNERSHIPS EXPANDING PIPELINE



OPHTHALMOLOGY PLATFORM

New Ophthalmology Program: Initiated development of a novel gene therapy targeting ocular diseases, including corneal pathologies and glaucoma, using Genflow's proprietary Centenarian SIRT6 (cent SIRT6).

Precision Delivery Platform: Employing a specially engineered non-viral vector for targeted SIRT6 delivery to the eye.

Strategic Partnership: Signed a Material Transfer Agreement in April 2025 with a leading ophthalmology company to co-develop the cent SIRT6 therapy and delivery system.

LEVERAGING AI FOR GENOMICS

New AI Partnership: Signed Master Service Agreement in April 2025 with Heureka Labs, a spin-out of Duke University, to use their proprietary AI platform to analyze complex genomic data, including RNA sequencing and gene expression profiles.

Enhancing Therapeutic Insights: Expected to drive a deeper understanding of gene regulatory networks and biological responses to optimize therapy design and personalize outcomes.

Pipeline Advancement: Initially supporting GF-1002 (cent SIRT6-based gene therapy in preclinical development) and future programs, including candidates for MASH.



GLAUCOMA RGC NEUROPROTECTION: THE RACE IS ON

The Problem

80M people worldwide with glaucoma; leading cause of irreversible blindness Current treatments only lower IOP

- they don't protect retinal ganglion cells (RGCs) RGC death is irreversible; unmet need for neuroprotection

Significant market opportunity

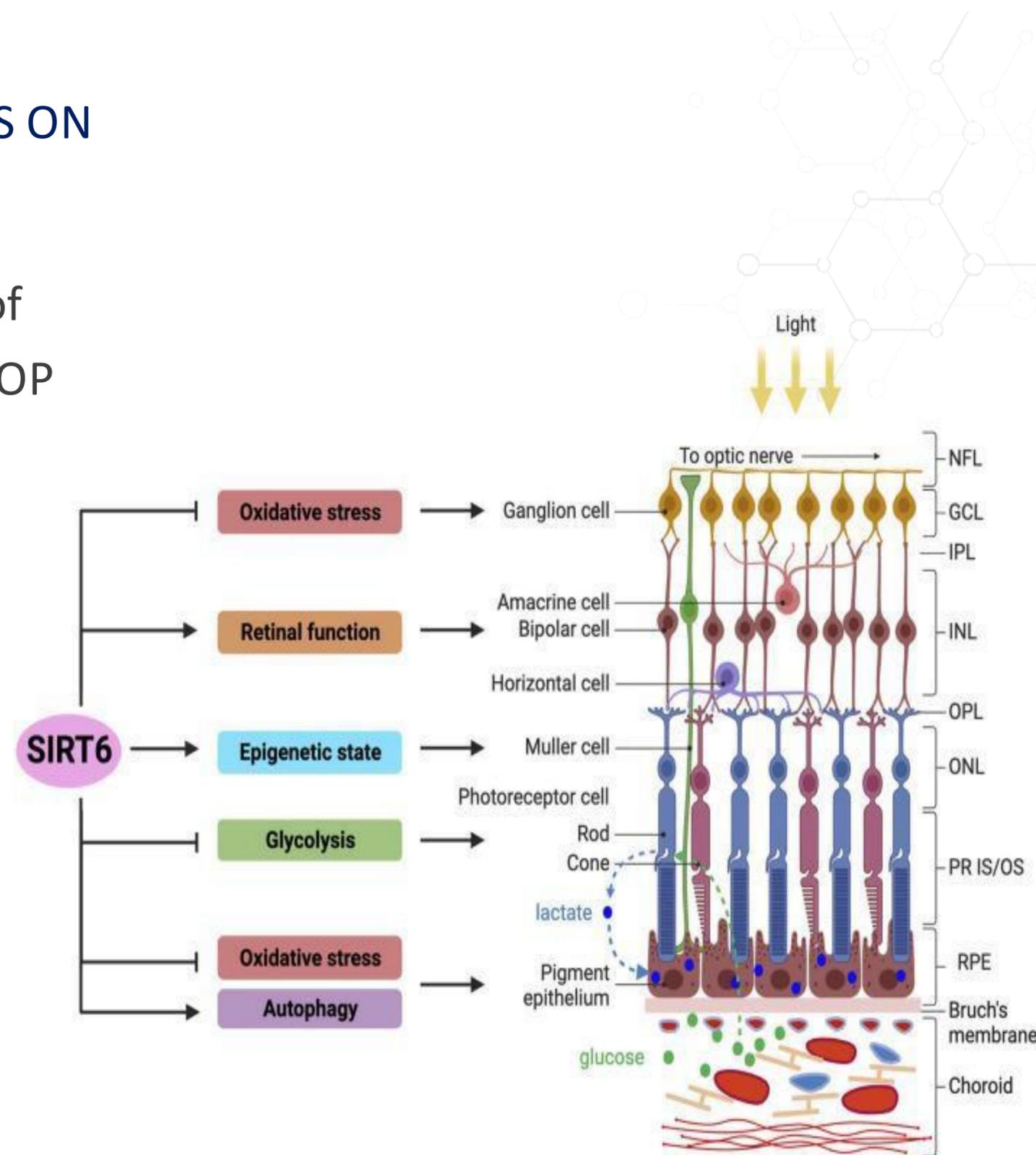
The Gap

No approved neuroprotective therapies

The mRNA Opportunity - Wide Open

LNP targeted delivery to retina – now possible

NO APPROVED OR CLINICAL-STAGE MRNA THERAPIES FOR GLAUCOMA



Cheng J, Keuthan CJ, Esumi N. The many faces of SIRT6 in the retina and retinal pigment epithelium. *Front Cell Dev Biol.* 2023 Nov 1;11:1244765. doi: 10.3389/fcell.2023.1244765. PMID: 38016059; PMCID: PMC10646311.

PARTNERING WITH LNP PROVIDER

Paradigm shift to retinal protection

- Pre-clinical evidence links SIRT6 activity with RGC protection in glaucoma/optic neuropathy models
- Demonstrating RGC rescue and preserved function would be a strong disease-modifying signal

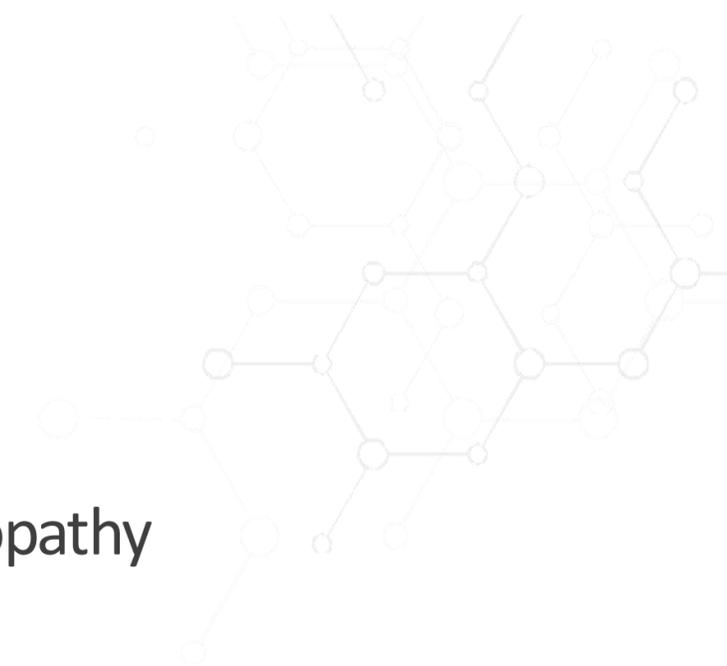
Delivery is now tractable

- Targeted LNP (tLNP) platforms are removing a major historical barrier for in vivo RNA therapeutics to reach retina/RGCs

Clear decision point

A single well-designed rodent POC that shows

- focal RGC expression of SIRT6 cent
- preserved RGC counts and
- preserved RGC function with acceptable ocular safety, gives a clean go/no-go for IND enabling work.



AGED DOGS COMPARATIVE TRIAL

Proof-of-Concept comparative clinical trial of in-vivo naked DNA gene therapy in aged dogs

Conducted at Syngene's strategic partner site in Morocco, AAALAC and GLP certified
26 (24+2) aged Beagles dogs (age>10 years)

Group 1 (n=6): untreated

Group 2 (n=6): treated with gene therapy IV bolus of SIRT6 AAV

Group 3 (n=6): treated with gene therapy IV infusion of naked DNA (low dose)

Group 4 (n=6): treated with gene therapy IV infusion of naked DNA (high dose)

Blinded: first read-out in January 2026

- Treatment period: 180 days with follow-up period: 90 days
- Gene therapy has been administered to all dogs with demonstrated excellent safety profile

Endpoints

- Pan-mammalian methylation clock (Steve Horvath),
- sarcopenia (including muscle biopsies) and other markers of aging



MASH PROGRAM

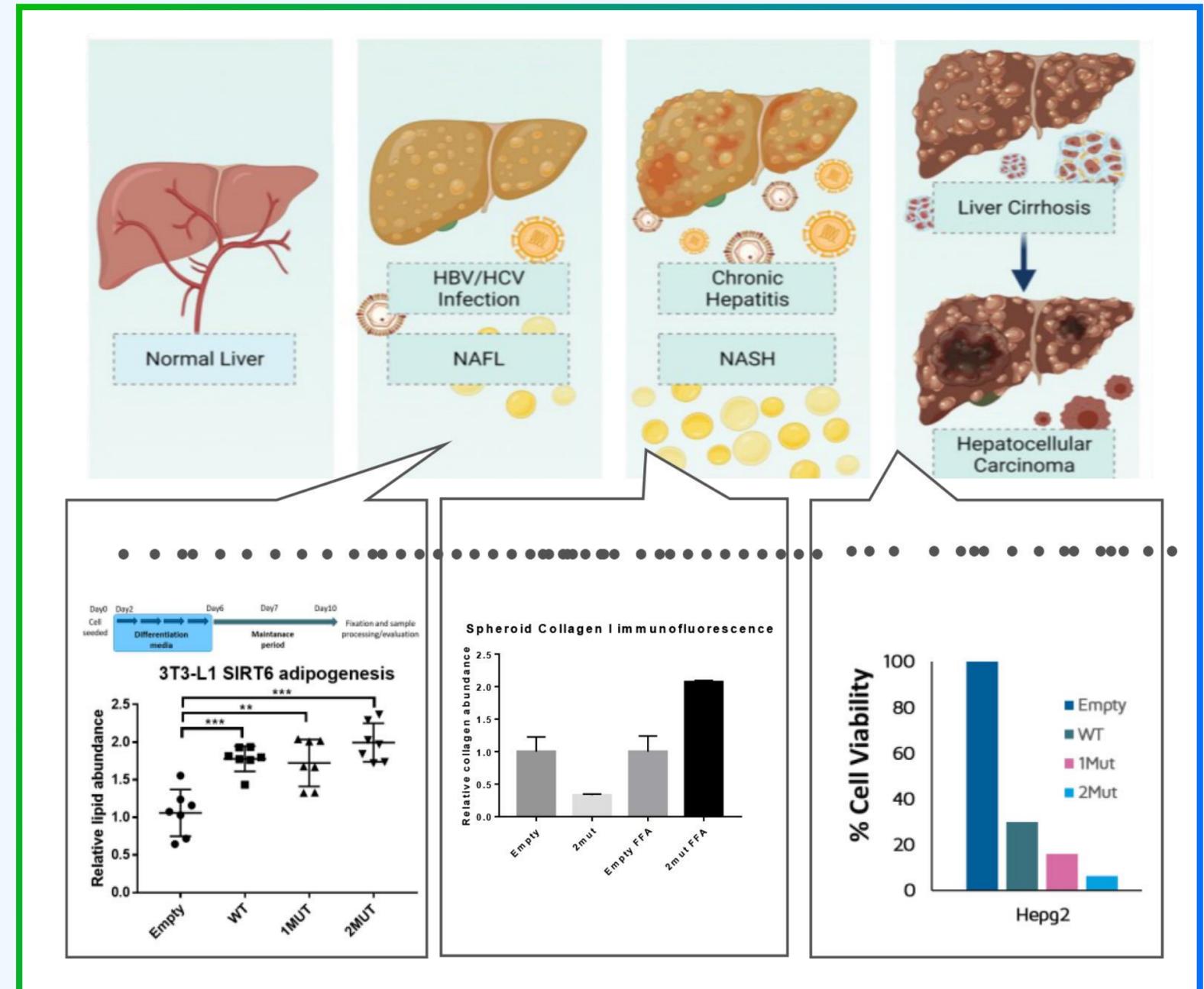
Affects est. 35 million people globally

- 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, Ratzu V, Conti F. NAFLD and liver transplantation: Current burden and expected challenges. *J Hepatol.* 2016 Dec;65(6):1245-1257.
Vlad Ratzu, Sven Francque, Arun Sanyal, Breakthroughs in therapies for NASH and remaining challenges, *Journal of Hepatology*, Volume 76, Issue 6, 2022

2025 KEY HIGHLIGHTS

LARGE AND EXPANDING MARKET OPPORTUNITY

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

Awarded to new grants in 2024 expanding research pipeline

Ophthalmology: retina preservation

GROWING IP PORTFOLIO WITH LONG LIFE

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

MRNA/LNP GENE DELIVERY SYSTEMS

Innovative gene delivery systems: targeted LNPs delivering cent SIRT6 mRNA

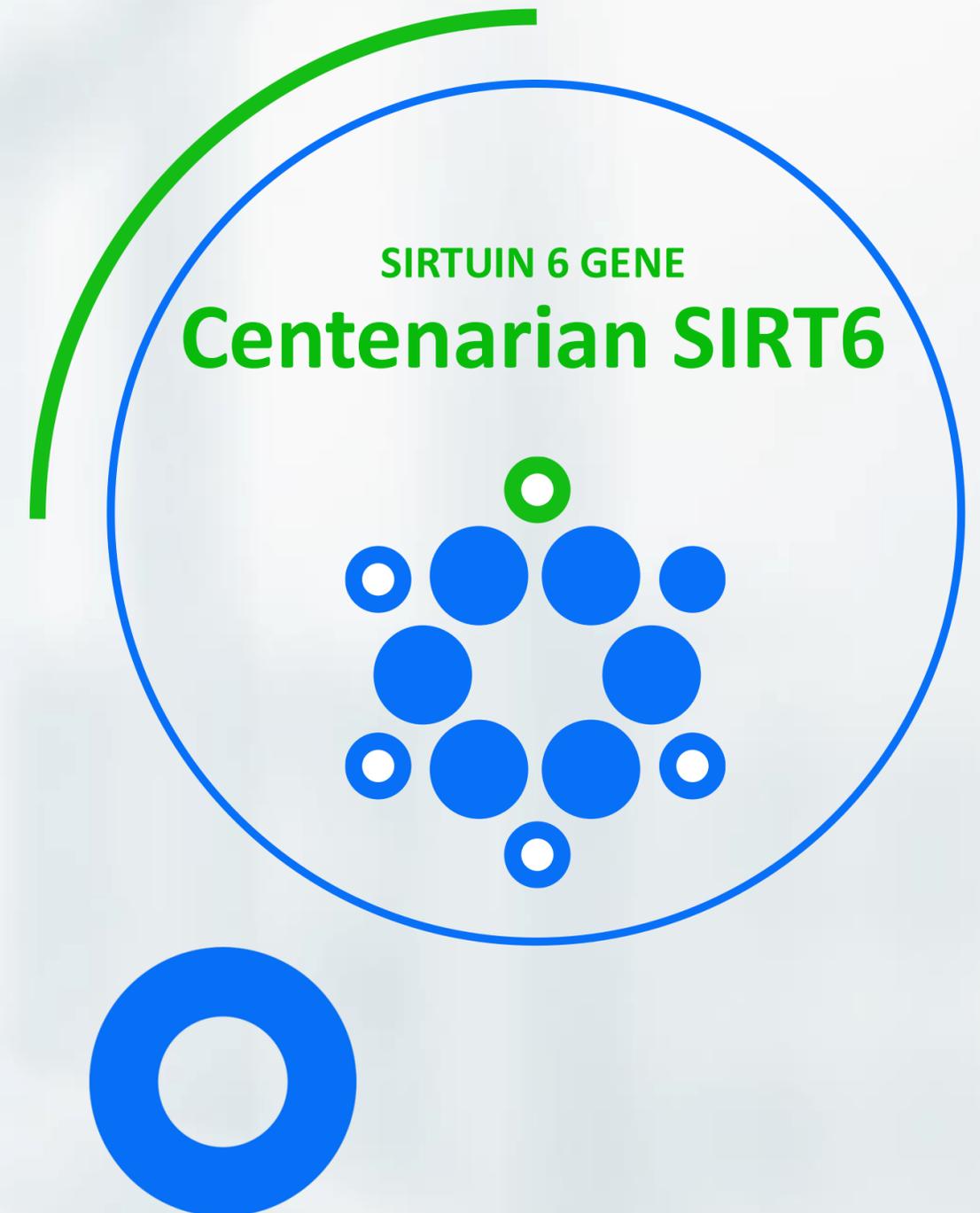
GF-1002

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

GF-1004: Pet life extension / sarcopenia

Initiated discussion under CDA with several animal health companies

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THANK YOU

Contacts

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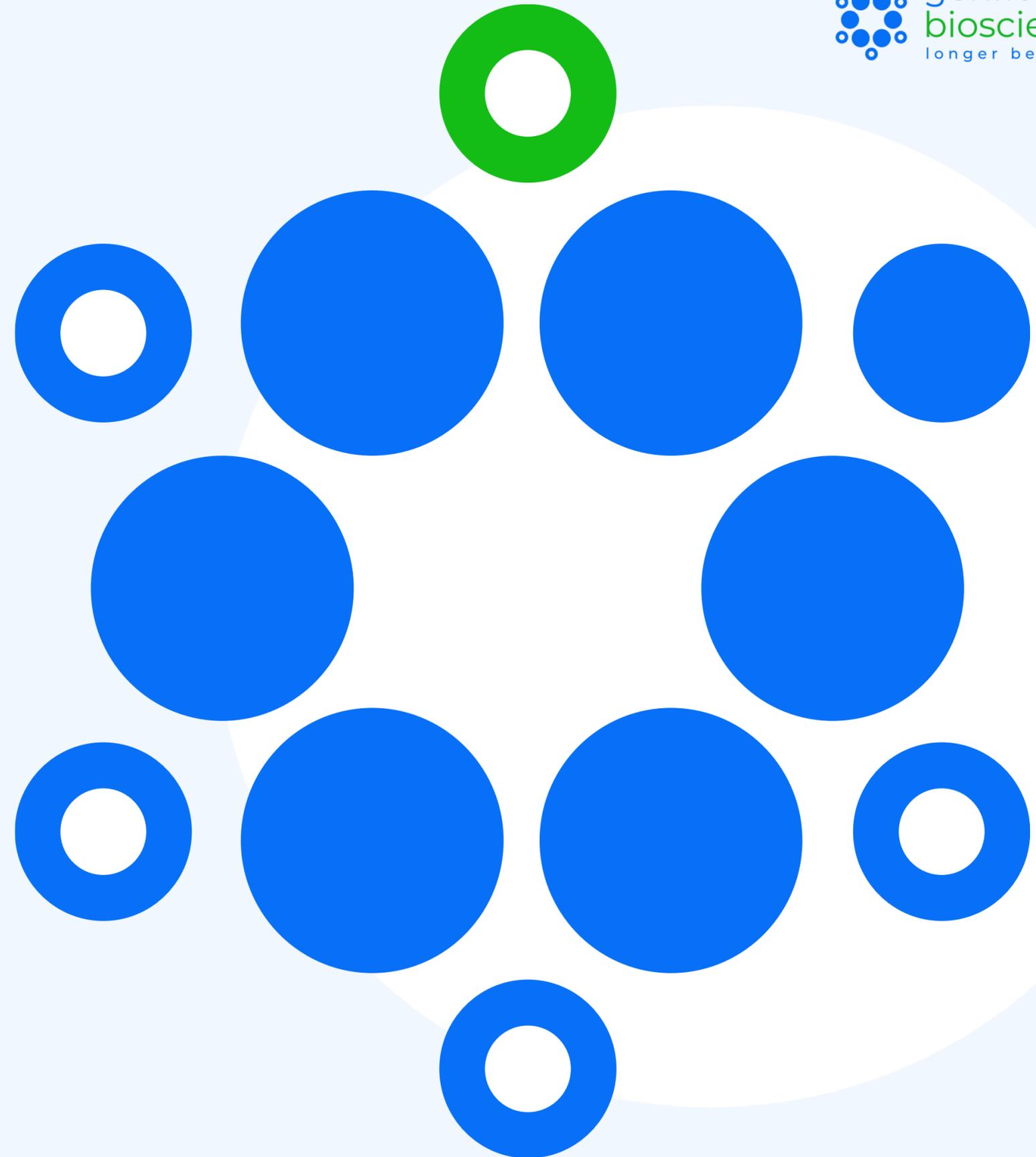
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LONGEVITY LANDSCAPE

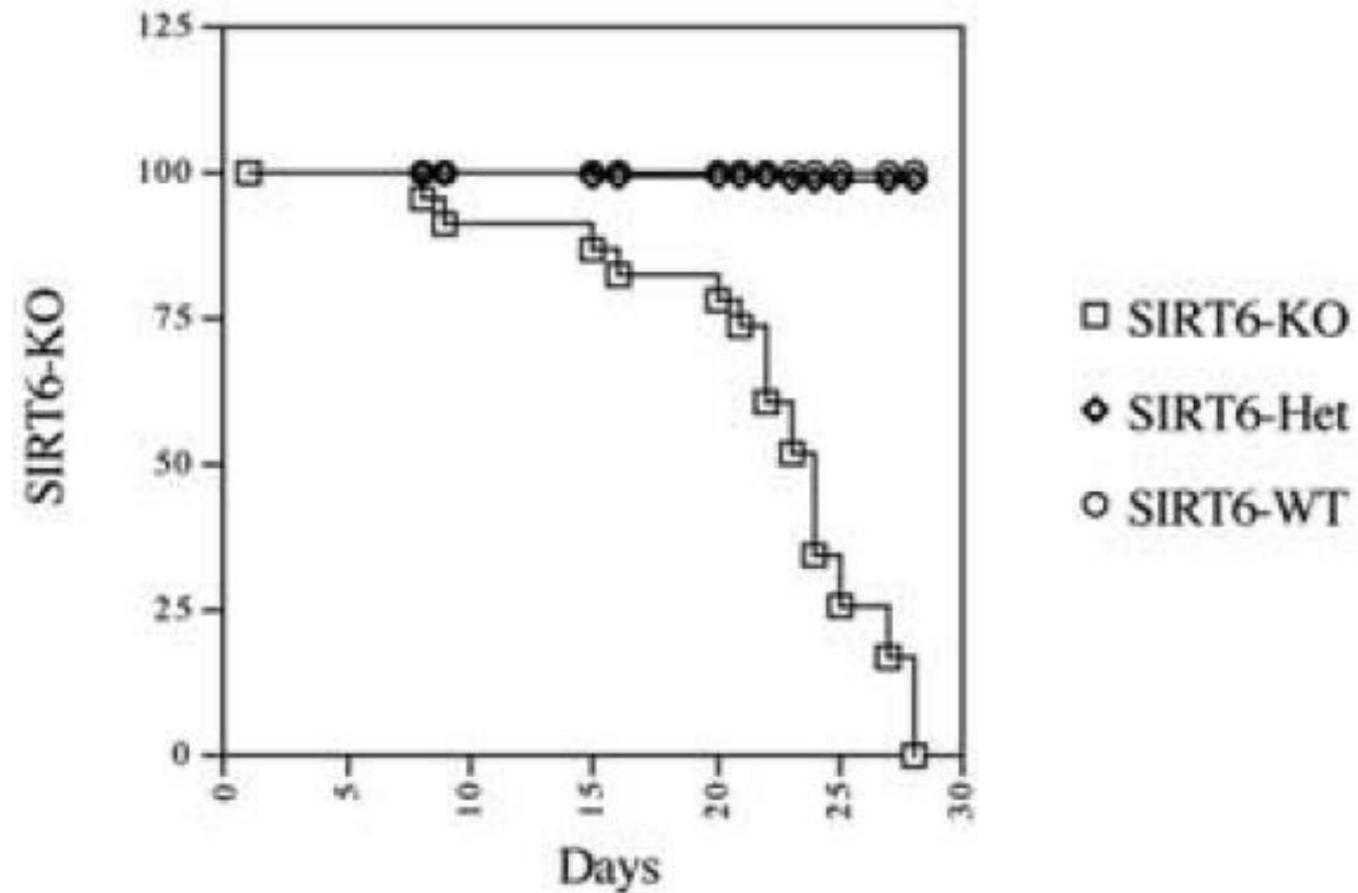
COMPANY	OVERVIEW	TECHNOLOGY	FOCUSED ON	LOCATION
	CLINICAL STAGE, PHASE II NASDAQ (UBX) MKT CAP \$785M	Small Molecules Senolytic	Senescence	USA, San Francisco, CA
	PRE-CLINICAL STAGE NYSE (AGE) MKT CAP \$25M	Therapeutics that seek to address human aging	Stem cells	USA, Alameda, CA
	CLINICAL STAGE, PHASE 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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RATIONALE FOR SIRT6 USE TO EXTEND LIFESPAN

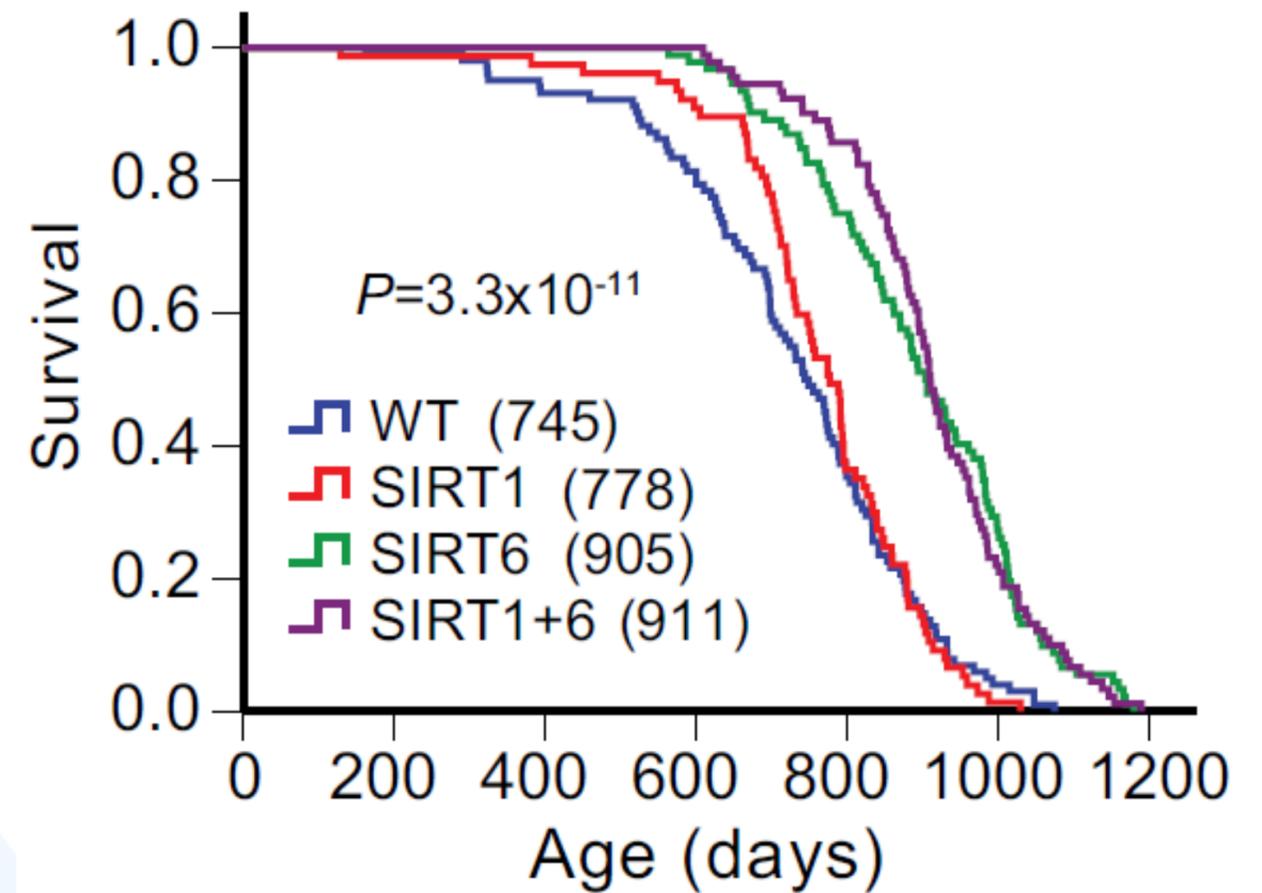
SIRT6 transgenic mice

SIRT6 knockout(KO)



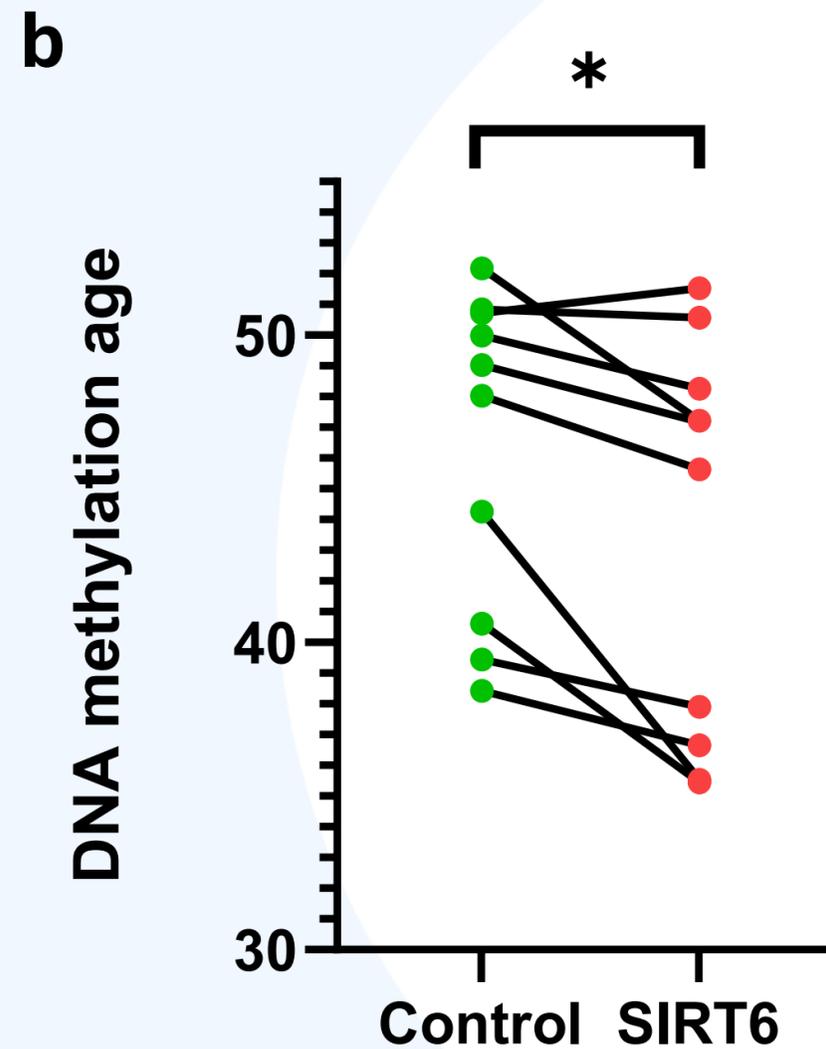
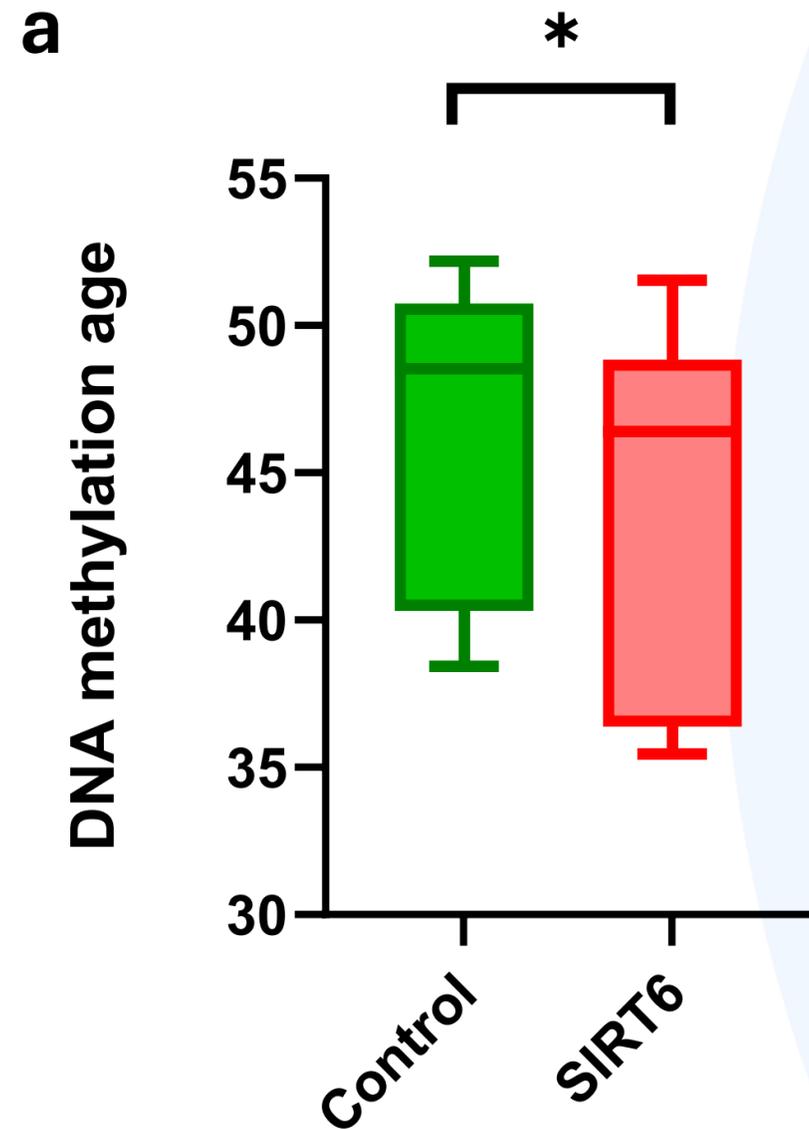
SIRT6 Overexpression(OE)

Sexes pooled



RATIONALE FOR SIRT6 USE IN AGED DOGS

SIRT6 OE in aged cells reverses epigenetic age (DNA mAge)

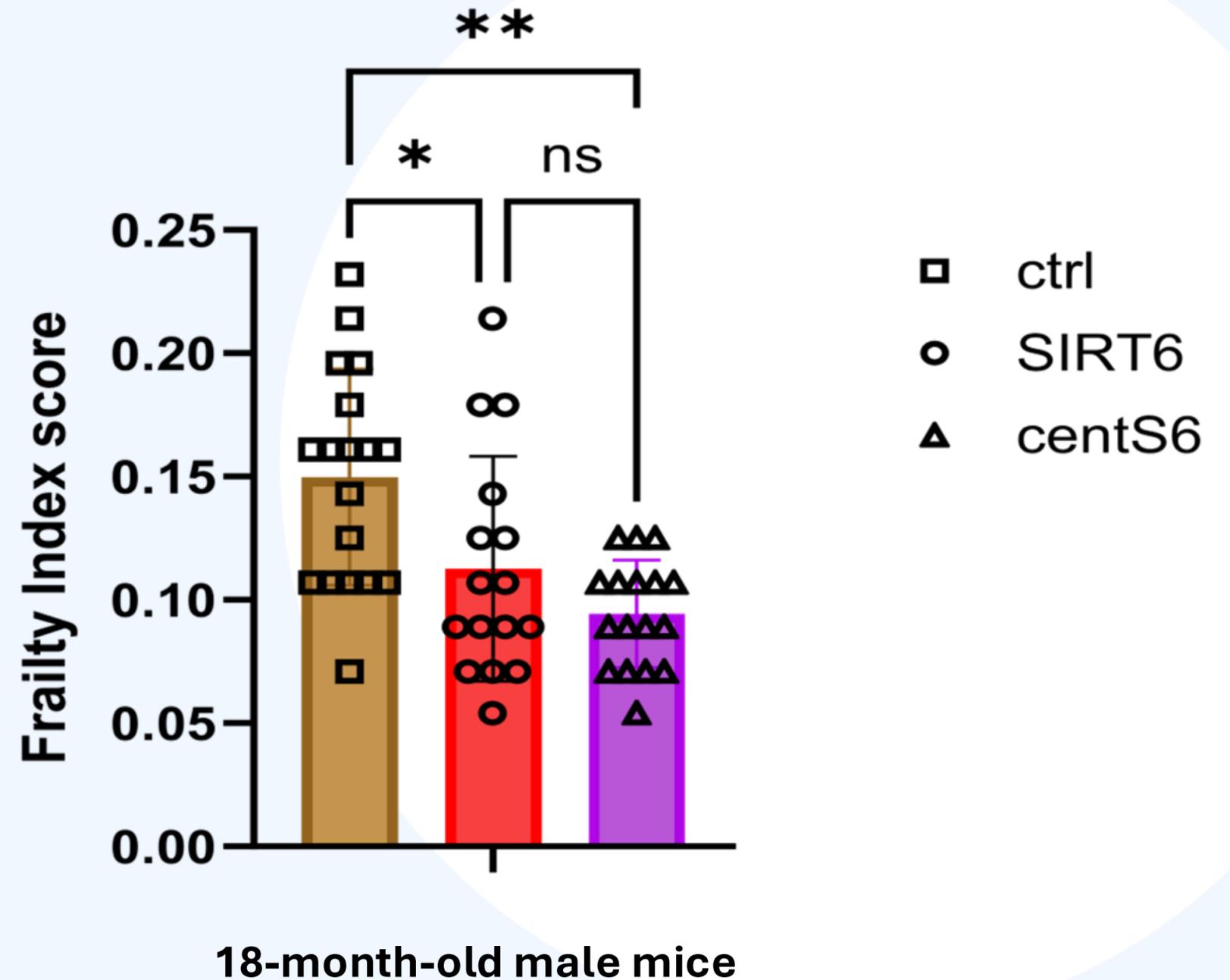


RATIONALE FOR SIRT6 USE IN SARCOPENIA

Genflow SIRT6 study conducted with Rochester University, NY

Frailty Test in Mice

Centenarian SIRT6 demonstrates statistically significant improvement



NAKED DNA DELIVERY

Naked DNA in vivo delivery in small mammals like dogs holds several advantages:

- **Safety and Simplicity:** Naked DNA lacks viral components, reducing the risk of immune responses and integration-related mutagenesis, making it safer than viral vectors.
- **Cost-Effectiveness:** Unlike complex viral vector production, plasmid DNA is relatively easy and inexpensive to produce.
- **Transient Expression:** Naked DNA delivers transgenes without permanent integration, which is ideal for temporary therapeutic needs or preclinical research where long-term expression is unnecessary.

Intravenous (IV) Administration:

- **Systemic Distribution:** IV injection enables the DNA to circulate widely, reaching multiple tissues, though uptake by cells is generally low.
- **Lower Expression Levels:** The naked DNA is more rapidly cleared from the bloodstream than when using intra-muscular injections, resulting in need for the veterinary to repeat the iv injections.

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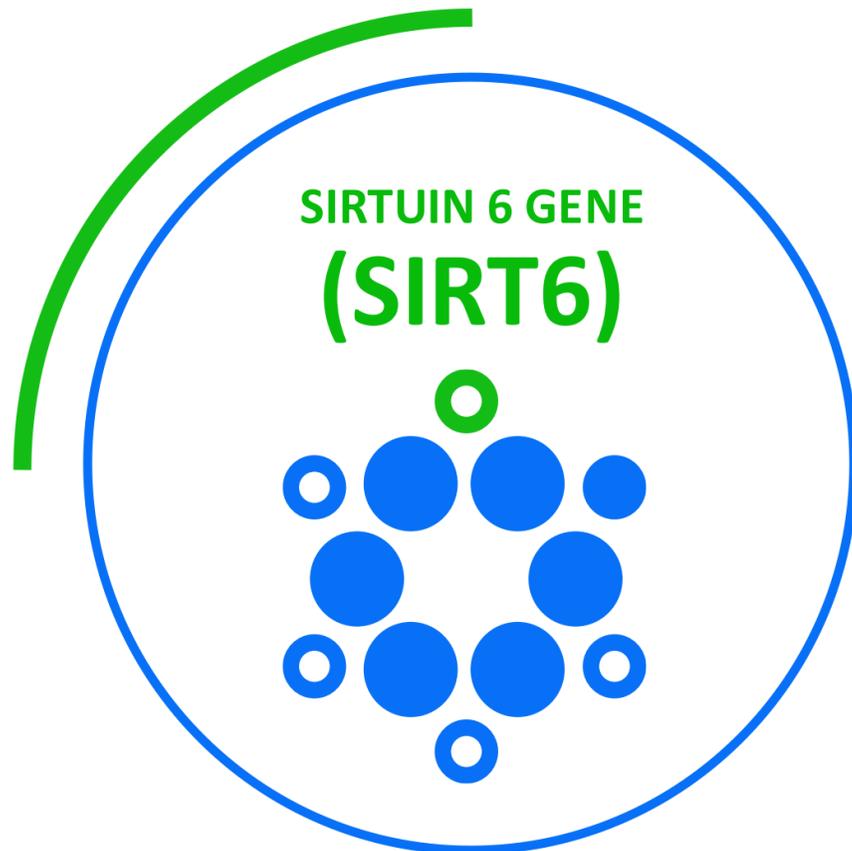


GENE REGULATION IN AGING

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

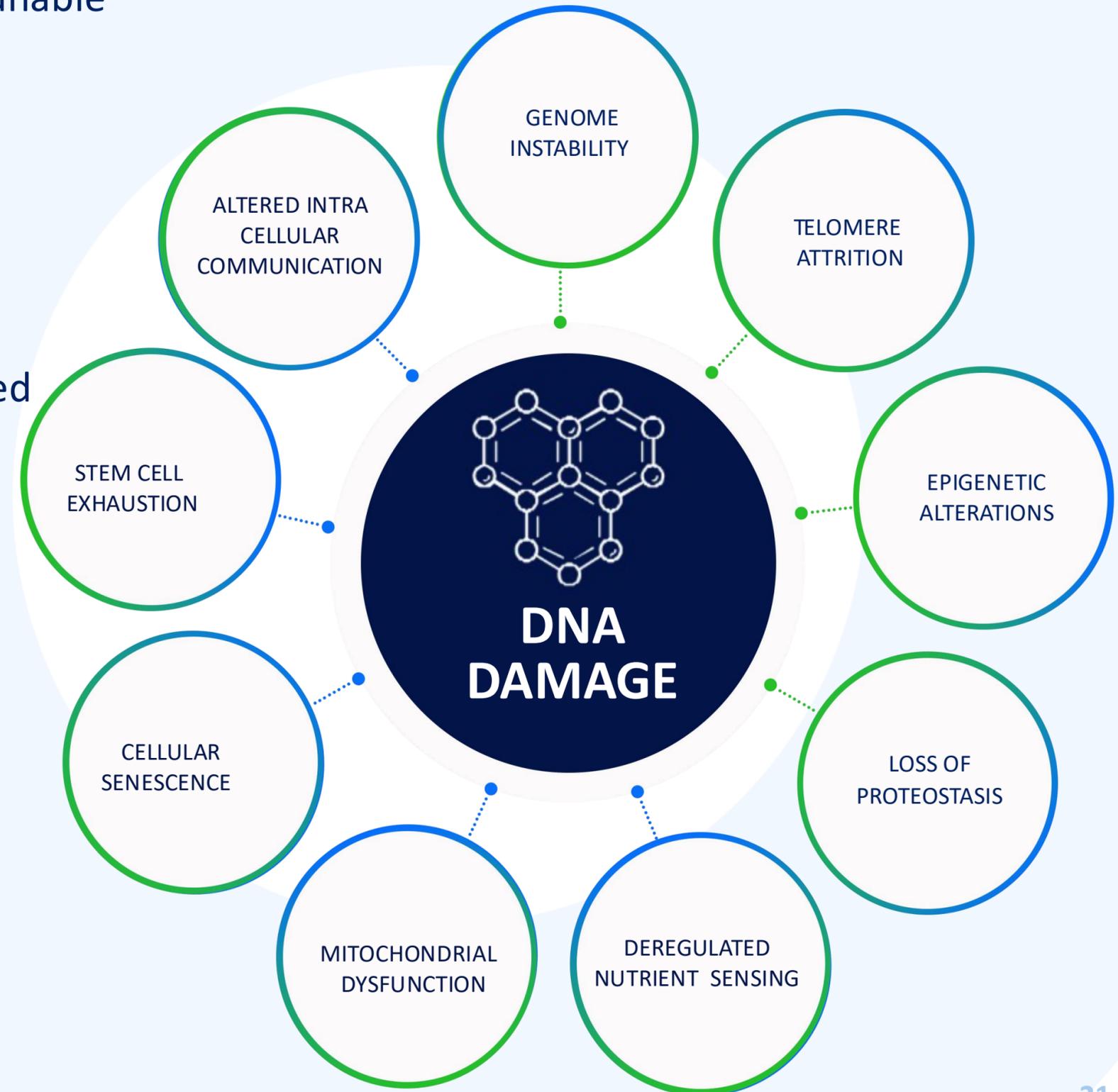
MANY GENES REGULATE AGING.
OUR FOCUS IS THE CENTENARIAN **SIRT6 (cent SIRT6) GENE**

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



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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

LIFESPAN

Lifespan
 ▲
 DSB repair
 ▲
 SIRT6 activity



5 amino acids



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

FOCUS ON CENTENARIAN SIRT6 (cent SIRT6)

SIRT6 centenarian variant (cent SIRT6) gene has more efficient DNA repair properties

HOMOLOGOUS

Recombination Repair

NON-HOMOLOGOUS End

Joining Repair

RELATIVE PARP1

Ribosilation

