# Optimising healthspan using gerotherapeutic drugs in middle aged individuals







NUHS Centre for Healthy Longevity

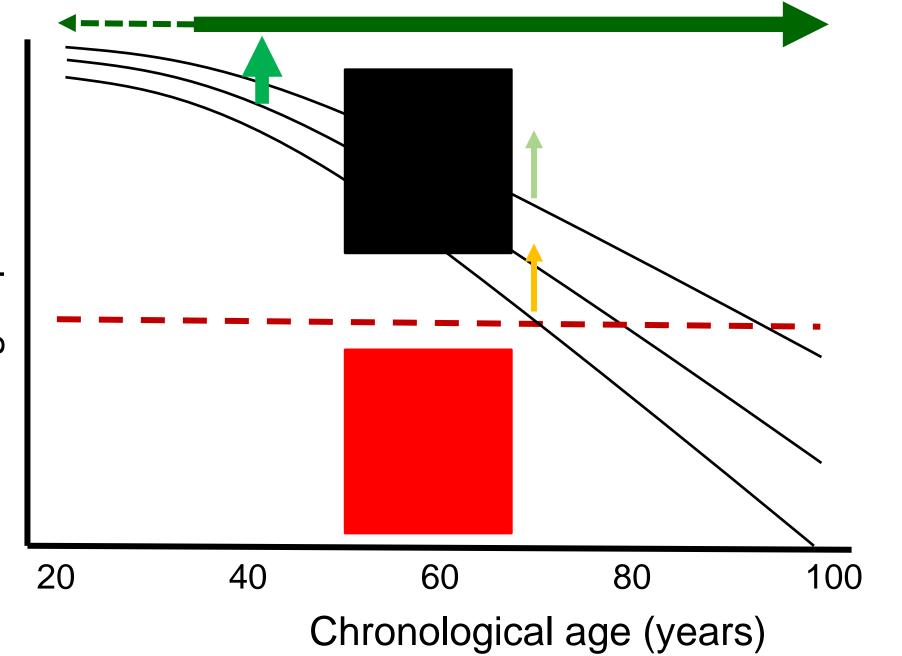
### Andrea Maier

# Longevity dividend:

health and economic gains by slowing the biological processes of aging



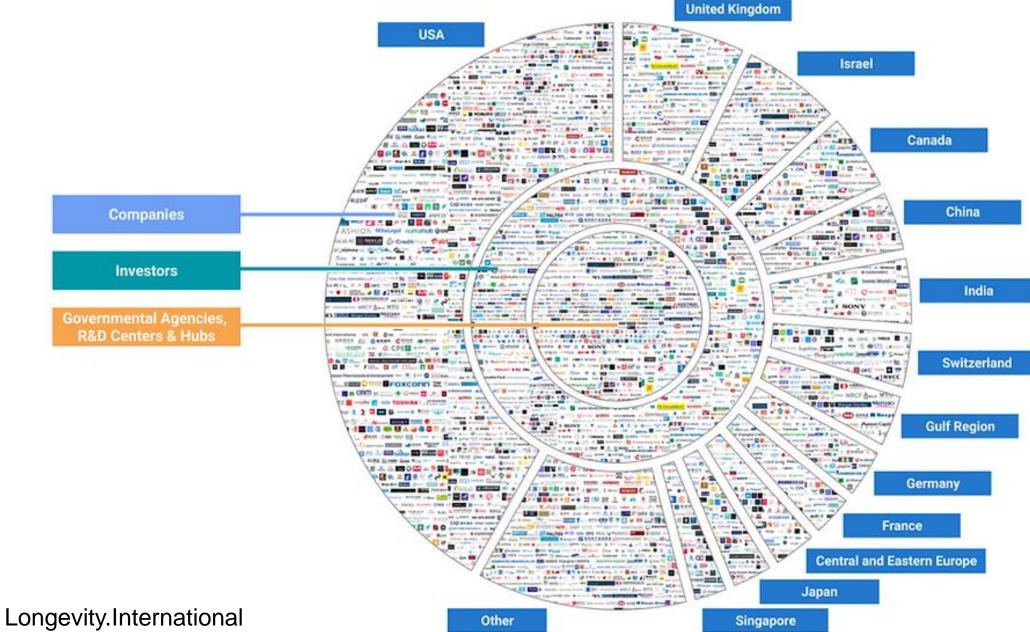
costs	value	enabler			
Picco et al. BMC Health Services Research (2016) 16:173 DMC Health Services Research   D101186/s12913-016-1421-7 Den Access   RESEARCH ARTICLE Open Access   Goonomic burden of multimorbidity among older adults: impact on healthcare and access adults: impact on healthcare and access adults. Impact on healthcare and access adults.   Moise Picco <sup>1°</sup> , Evanthia Achila <sup>2</sup> , Edimansyah Abdin <sup>1</sup> , Siow Ann Chong <sup>1</sup> , Janhavi Ajit Vainganka <sup>1</sup> , Paul McCrone <sup>2</sup> , Borg Choon Chua <sup>3</sup> , Derrick Heng <sup>4</sup> , Harish Magadi <sup>5</sup> , Li Ling Ng <sup>6</sup> , Martin Prince <sup>7</sup> and Mythily Subramaniam		CORRESPONDENCE   VOLUME 3, ISSUE 1, E12, JANUARY 01, 2022   Advanced pathological ageing should be represented in the LCD   Evelyne Bischof + Andrea B Maier + Kai-Fu Lee + Alex Zhavoronkov + David Sinclair C   Open Access + Published: January, 2022 + DOI: https://doi.org/10.1016/52666-7568(21)00303-2			
pp/a no chronic condition: SGD\$2,806 one chronic condition: SGD\$5,610 multimorbidity: SGD\$15,148	slowdown in aging that increases life expectancy by 1 year is worth US\$38 trillion	XT9T			



# Organ performance

# Longevity ecosystem





50 000+ Companies 10 000+ Investors 1 000+ R&D Centers

# Increasing <u>health</u>span of SG by ...

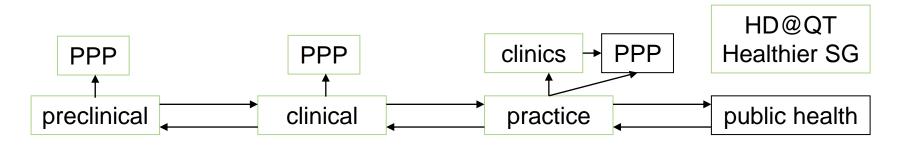


building the ecosystem

at least 3 year in 10 years

Healthy Longevity Medicine Society

Healthy Longevity Academy: executive, master, incubator

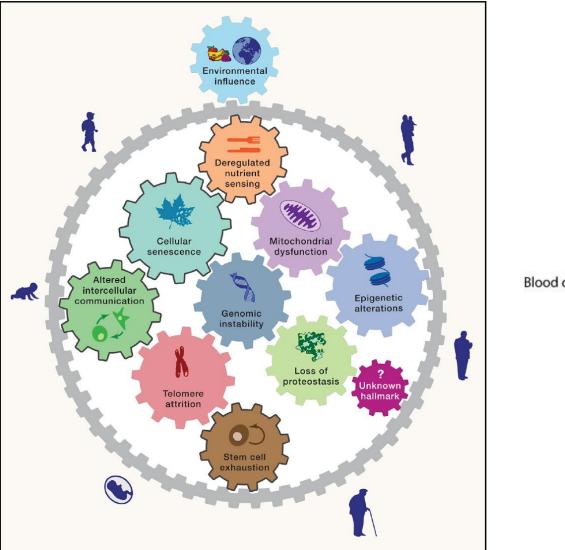


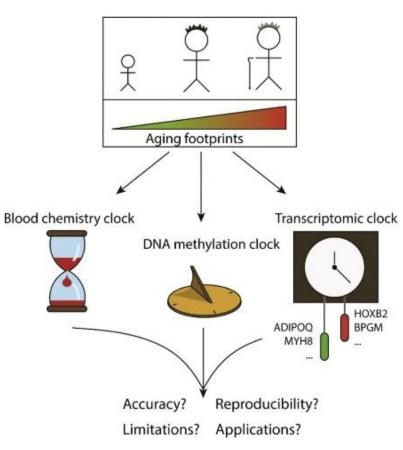
diagnostics: core measuring biological age

interventions: core 'treating' biological age



# Diagnostics





Tuttle et al., Exp Gerontol 2018; Singh et al., Cell 2019; Shahmirzadi et al., Cell Metabolism 2020; Galkin et al., Ageing Res Rev 2020; Tuttle et al., Aging Cell 2020



### **Biomarker of Ageing Consortium**

### 2023 Executive Committee



Mahdi Moqri, PhD, MBA STANFORD & HARVARD UNIVERSITIES

Dr. Moqri is a Research Fellow in Aging Research at Stanford School of Medicine and a Visiting Scholar at Harvard Medical School.

Relevant expertise:

Biomarkers of aging and rejuvenation and their assessments.



Jesse Poganik, PhD HARVARD UNIVERSITY

Dr. Poganik is Research Fellow at Brigham and Women's Hospital and Harvard Medical School.

Relevant expertises Epigenetic clocks of aging in human and model animals.



Allison Duettman FORESIGHT INSTITUTE

Allison Duettmann is the President and CEO of Foresight Institute

Relevant expertise: Advanced biotech, nanotech, computing for the long-term benefit of life.

### 2023 Scientific Committee

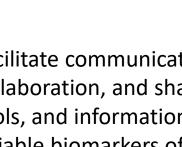


Dane Gobel METHUSELAH FOUNDATION

Program Director for the Methuselah Foundation.

Relevant expertises Non-profit management, business development, deep tech open innovation, dealflow.

Dane Gobel is the Co-Founder and



Facilitate communication, collaboration, and sharing of data, tools, and information to establish reliable biomarkers of aging, particularly for the identification and evaluation of longevity interventions.



Vadim Gladyshev, PhD HARVARD UNIVERSITY

Dr. Gladyshev is a Professor of Medicine at Harvard Medical School and Director of Redox Medicine at Brigham & Women's Hospital.

Relevant expertise:

Aging clocks and other biomarkers of biological age.



Michael Snyder, PhD STANFORD UNIVERSITY

Dr. Snyder is the Chair of the Department of Genetics at Stanford School of Medicine and the Director of the Center for Genomics and Personalized Medicine

Relevant expertise: Longitudinal biomarkers of aging and Multiomic Ageotyping.



Vittorio Sebastiano, PhD STANFORD UNIVERSITY

Dr. Sebastiano is an Associate Professor in the Department of Obstetrics & Gynecology at Stanford School of Medicine and the Co-Founder of Turn Biotechnologies.

Relevant expertise: Biomarkers of cellular rejuvenation



Andrea Maier, MD NATIONAL UNIVERSITY OF SINGAPORE

Dr. Maier is a Professor in Medicine and Healthy Ageing and the Co-Director of Center for Healthy Longevity at the National University of Singapore and a Fellow of the Royal Australasian College of Physicians.

Relevant expertise: Healthy aging and longevity biomarkers.



BUCK INSTITUTE FOR RESEARCH ON AGING

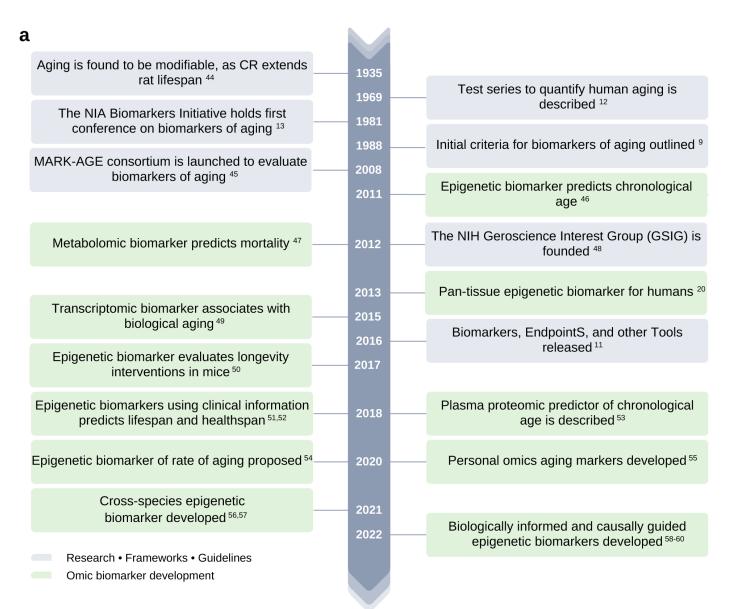
Dr. Verdin is President and Chief Executive Officer of the Buck Institute for Research on Aging, as well as Professor of Medicine at University of California, San Francisco.

Biomarkers of immune aging.

Relevant expertise:

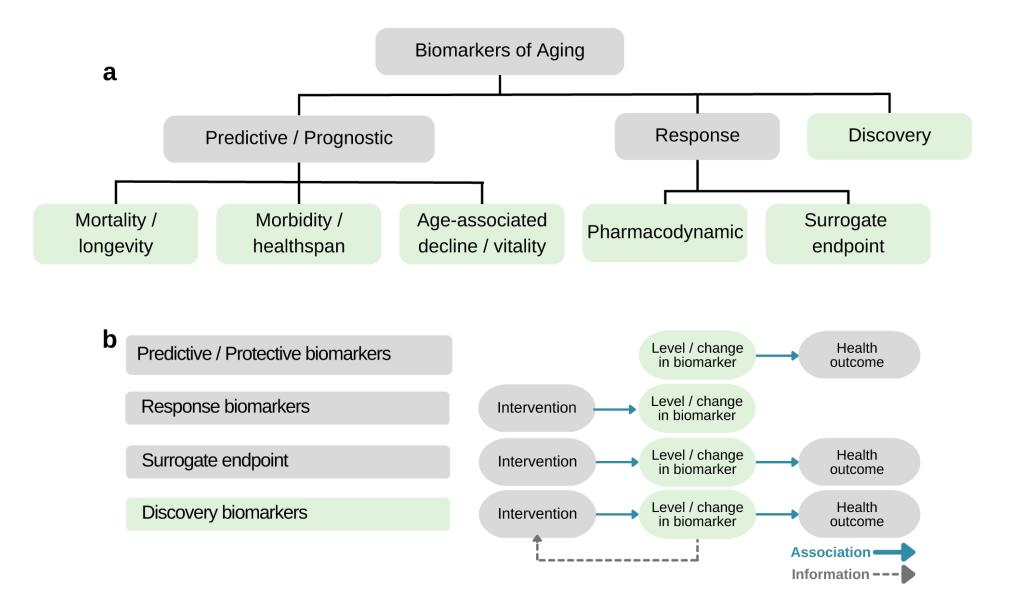
Eric Verdin, MD

# Timeline - biomarkers of aging

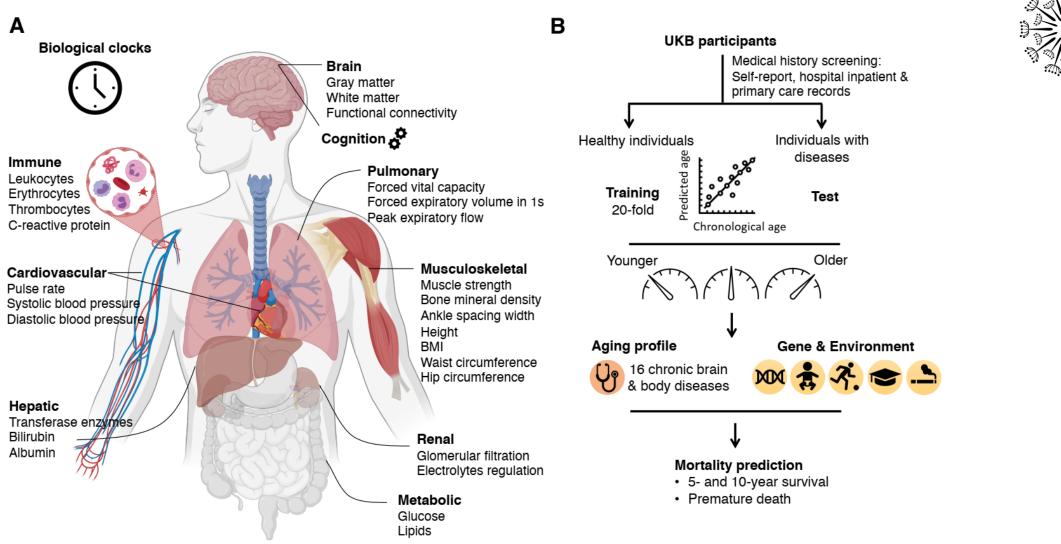




# Categories of biomarkers of ageing







143,423 individuals (age range 39-73 years, mean 56.7±8.2, 79,980 males)

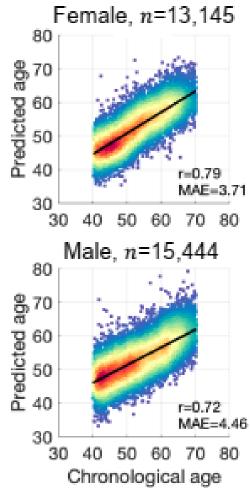
- Support vector machines (SVMs) were trained to predict chr age
- 20-fold cross-validation
- Variables standardized by weighted column mean and standard deviation

Yi Tian et al., Nature Medicine 2023

# Body clock

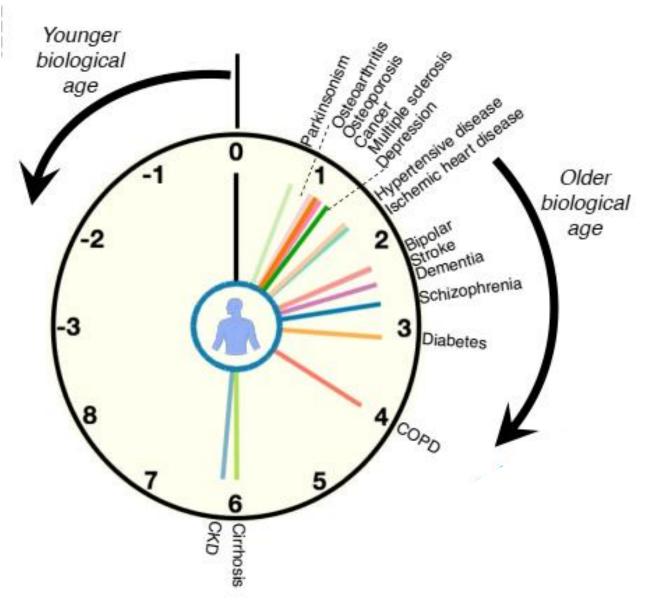


78 physiological measures across 7 organ systems



Yi Tian et al., Nature Medicine 2023

# Body clock – healthy $\rightarrow$ diseased

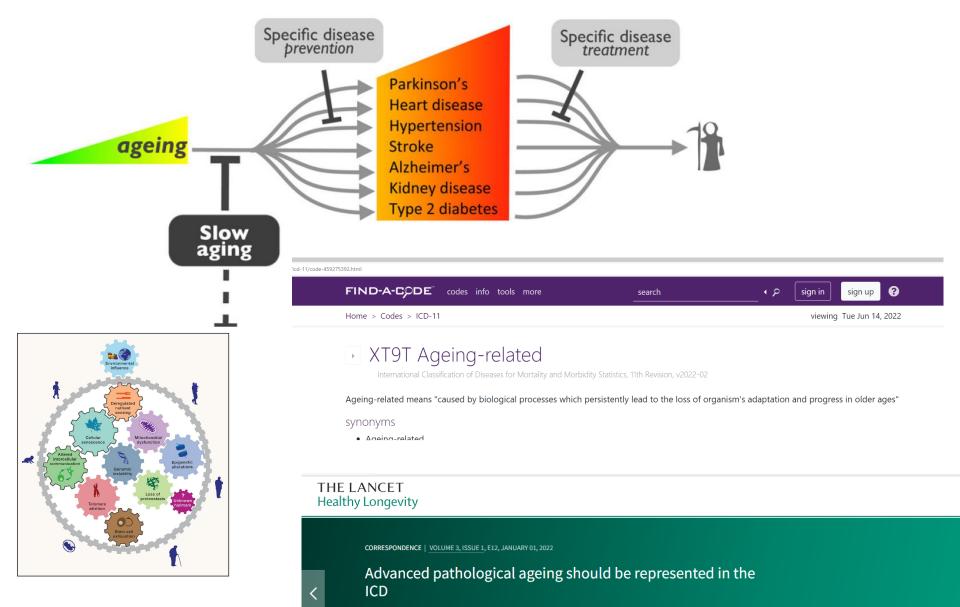




Yi Tian et al., Nature Medicine 2023

# Interventions



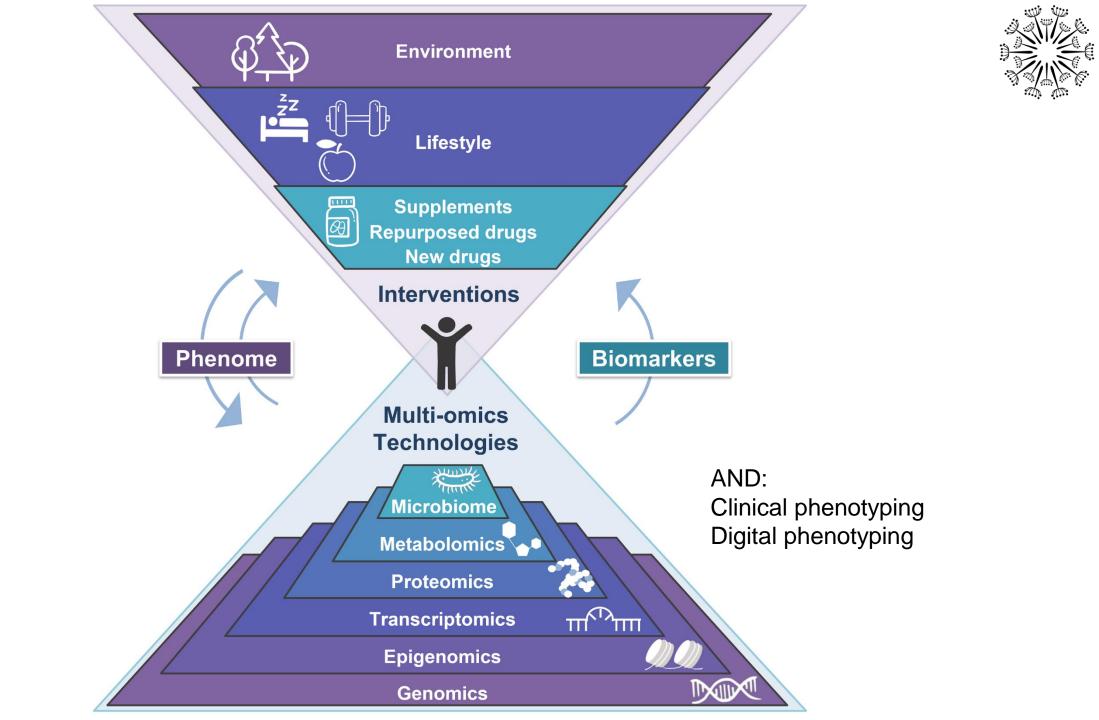


Seals et al., J Physiol 2015

Evelyne Bischof + Andrea B Maier + Kai-Fu Lee + Alex Zhavoronkov + David Sinclair 🖂

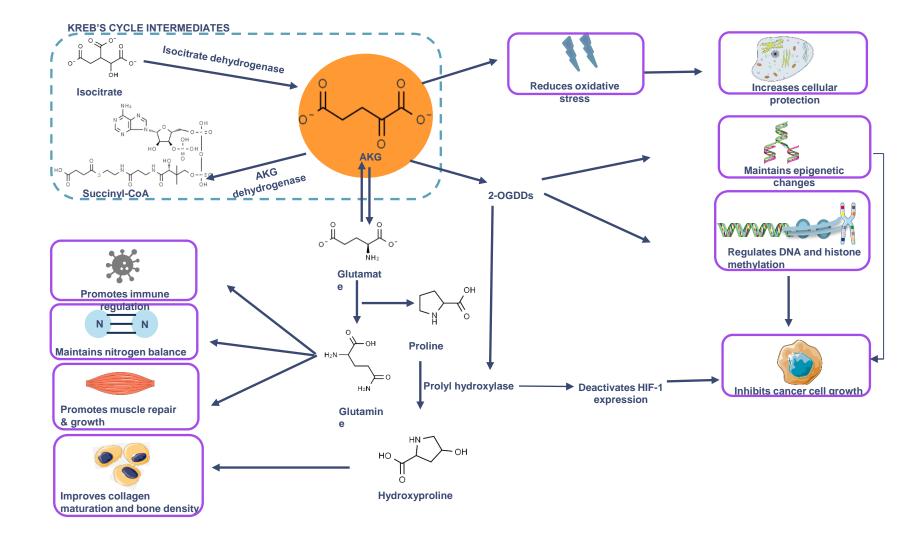
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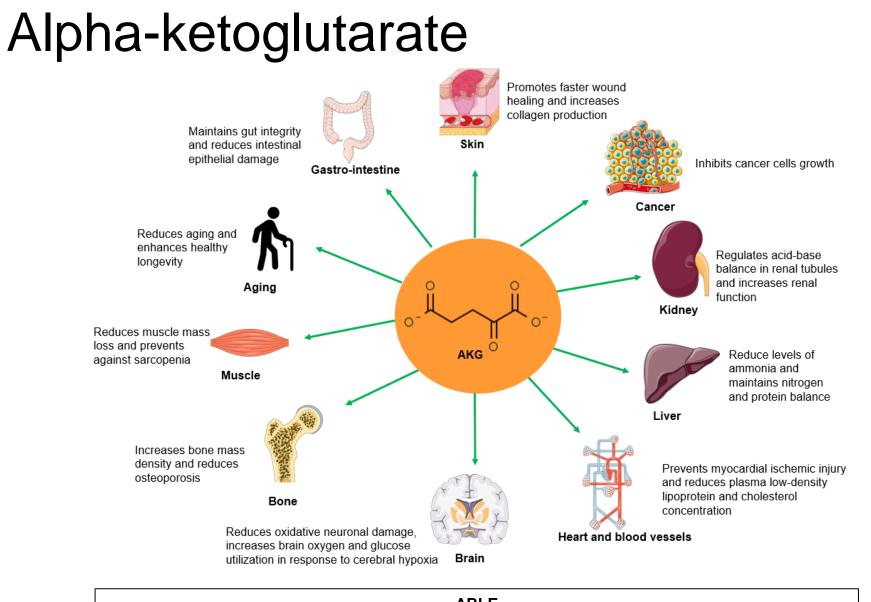


# Alpha-ketoglutarate





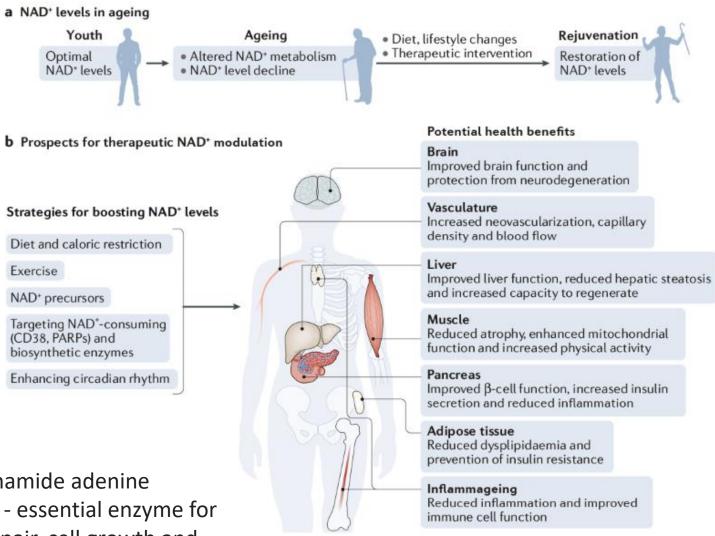
Gyanwali et al., Trends Endocrinology Metabolism 2022



ABLE double blinded RCT, 1g Ca-AKG / placebo, 6m + 3 m follow up 40-60 year-old healthy individuals with higher DNA methylation age compared to their chronological age primary outcome DNA methylation age

Gyanwali et al., Trends Endocrinology Metabolism 2022

### Nicotinamide mononucleotide (NMN)



precursors of nicotinamide adenine dinucleotide (NAD<sup>+</sup>) - essential enzyme for metabolism, DNA repair, cell growth and survival.

Verdin, Science 2015

### Nicotinamide mononucleotide (NMN)



	Placebo, n = 20					300mg NMN, n = 20				
	baseline	day 30	day 60	p1	p2	baseline	day 30	day 60	p1	p2
NAD, mean ± SD (pmol/mL)	8.1±5.2	9.8±8.4	11.8±9.4	0.44	0.14	11.8±11.7	29.8±20.1	32.6±17.9	0.0014	<0.001
Six-minutes walking test, mean ± SD (m)	324±144	310±125	330±117	0.73	0.90	307±108	350±114	380±143	0.23	0.079
Blood biological age, mean ± SD (years)	39.80±7.2	3	45.35±8.22	2	0.029	42.15±6.03		43.65±6.73		0.46
SF-36, mean ± SD (score)	121.6±13.	8 126.9±12.1	128.2±12.9	9 0.20	0.12	123.6±12.8	131.5±11.7	136.7±12.1	0.058	0.0025

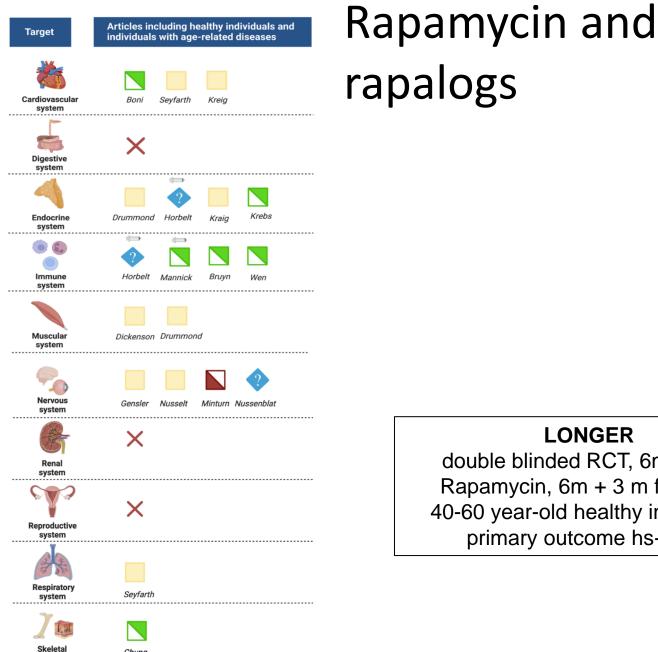
80 men
$50 \pm 6$ years
no chronic disease
NMN for 60 days

600mg NMN, n = 20			_		_				
baseline	day 30	day 60	p1	p2	baseline	day 30	day 60	p1	p2
8.0±3.3	39.0±12.6	45.3±11.8	< 0.001	<0.001	10.5±6.8	43.1±14.3	48.5±19.8	< 0.001	<0.001
289±92	400±85	435±104	< 0.001	<0.001	323±113	425±141	480±128	0.016	<0.001
45.20±6.49	)	44.05±6.37		0.57	44.30±7.28		45.30±5.91		0.64
117.6±16.2	2 129.3±12.5	5136.3±12.2	0.015	<0.001	121.9±16.7	135.7±12.0	140.3±11.1	0.0045	<0.001



Lin et al, GeroScience 2022





### LONGER double blinded RCT, 6mg/week Rapamycin, 6m + 3 m follow up 40-60 year-old healthy individuals primary outcome hs-CRP

Hodzic et al., Gerongology; Hodzic et al., in preparation

Chung

system

New longevity clinic to provide patients 'customised' health plan to slow ageing









🗇 1 of 2 Alexandra Hospital staff demonstrate the Facial Ageing test done with a special camera. ST PHOTO: GIN TAY



PUBLISHED SEP 7, 2022, 5:10 PM SGT



Healthy Longevity Medicine is optimizing healthspan by targeting ageing processes across the lifespan





### Overview

The Healthy Longevity Medicine Society (HLMS) was established in August 2022 to build a clinically credible framework and platform for longevity medicine that promotes the highest standards of interdisciplinary collaboration in the field. The HLMS is governed by a Council of elected members representing different geographical locations and sectors. The HLMS aims to educate, foster research and professional development, set recommendations and guidelines, and coordinate activities across the various domains of longevity medicine.

### **Our Mission**

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The HLMS has four main objectives:

- To identify and promote educational opportunities in longevity medicine, including accreditations and credentials
- To set and promote professional standards (including physician guidelines) in longevity medicine, thereby advancing and

### 9 Type here to search

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### Info / membership: hlms.co

Have questions? Get in touch at LongevityAcademy@nus.edu.sg



HEALTHY LONGEVITY

TALENT INCUBATOR

An Intensive Course for **Our Next-Generation Leaders** of Healthcare

3<sup>rd</sup> - 13<sup>th</sup> July 2023 Kent Ridge Campus National University of Singapore

We welcome Master/ PhD students or healthcare professionals (in training or  $\leq$  1 year of work experience) with keen interest in creating the better future of healthcare for everybody.





NUHS Centre for Healthy Longevity

Health technology is advancing. Human lifespan is extending. But healthspan is not catching up. However, healthspan extension is proven in animal models!

What do these trends mean to the future of medicine? How to optimise health across the lifespan? Is Geroscience ready to be implemented in healthcare?

### Why participate?

- Equip yourself with the knowledge & skills to become the future leader of healthcare!
- Learn the foundations of healthy longevity science from world-class experts!
- Hear inspiring stories & get career advice from thought leaders in both public & private sectors!
- Engage in thought-provoking discussions & creative activities uniquely crafted for you!
- Develop transferrable skills, e.g., leadership, communications, personal branding, & more!
- Meet like-minded talents from around the world & start building your international network today!

