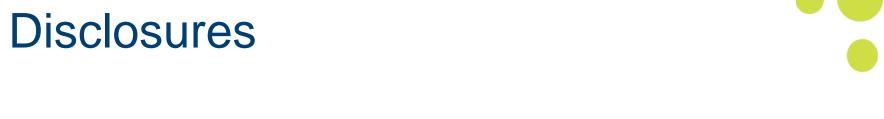
Scientific Wellness and the Future of Health and Nutrition

Nathan Price, PhD
Institute for Systems Biology
Seattle, WA

American College of Nutrition November 9, 2016





 Dr. Price is a Co-Founder of Arivale, which partially funded and may license discoveries resulting from the Hundred Person Wellness Project (to be described).

 Dr. Price is a Scientific Advisor to Habit, a new personalized nutrition company

Presentation Learning Objectives



After participating in this presentation, learners should be better able to:

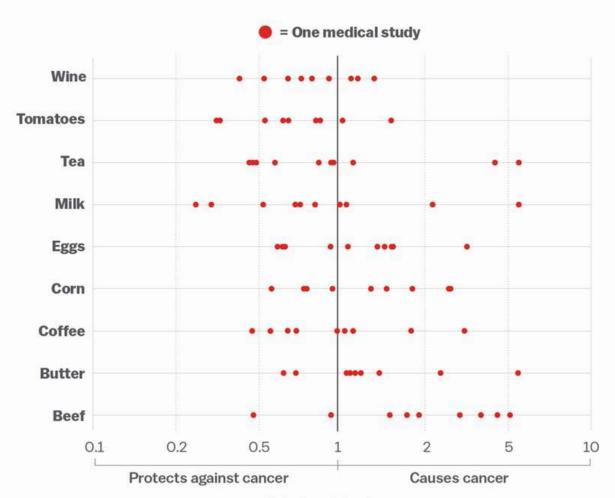
- Understand the scale of omic data collection that can inform nutrition and wellness.
- Understand the role of wellness coaches in interpreting data and motivating action.
- Understand new emerging field of 'scientific wellness'.

Nutrition health effects are complex: Need context and personalization





Everything we eat both causes and prevents cancer





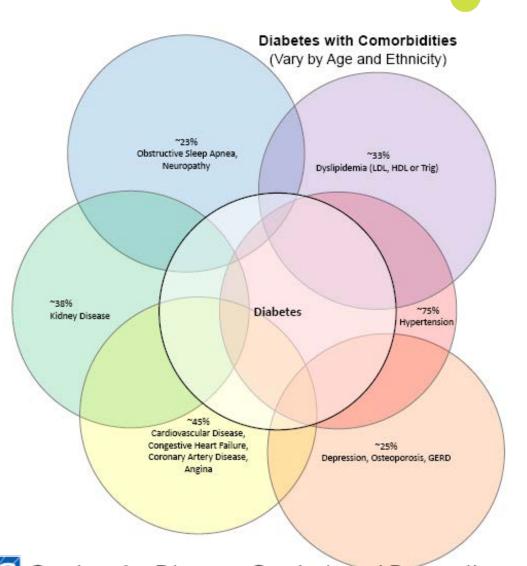




86% of Healthcare Costs Treat Chronic Disease

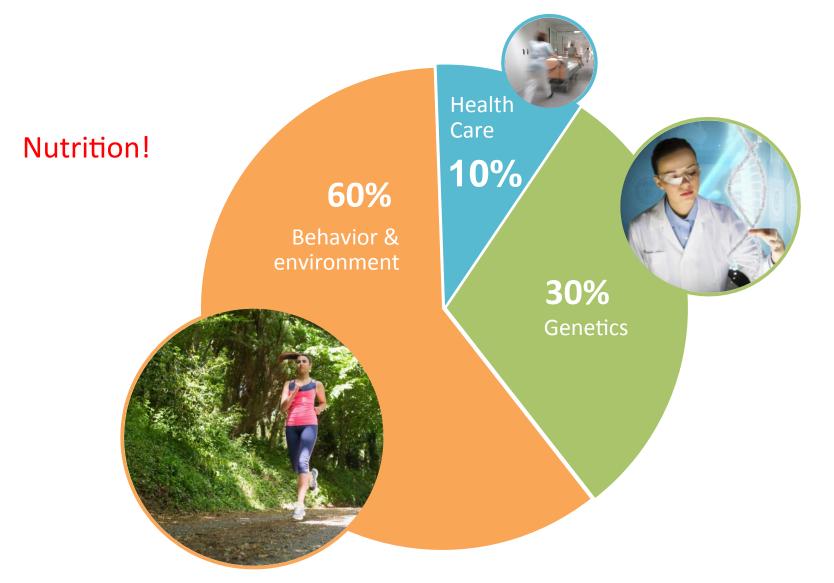


Chronic Diseases		Disease	Episodic or	Opportunities for	Cost
Prevalence (Millons)	Direct Cost (\$ Billions)	severity	Steady state	DX Monitor	Impact
Angina		High	Episodic	High	High
Anxiety disorders (social, etc.)		Medium	Episodic	High	High
Arthritis, Rheumatoid		High	Episodic	High	High
Asthma 22.2 M \$14.7 B		Medium	Episodic	High	High
Atrial Fibrillation		Medium	Episodic	High	High
Chronic Kidney Disease 26.0 M \$42.0 B		High	Steady Progression	High	High
	Heart Failure \$32.0 B	High	Steady Progression	High	High
COPD/Emphysema 12.1 M \$26.7 B		High	Steady Progression	High	High
Coronary Artery Disease 16.0 M \$87.6 B		High	Steady Progression	High	High
Depression 18.1 M \$80.0 B		High	Episodic	High	High
Dia 23.6 M	sbetes \$116.0 B	High	Steady Progression	High	High
Gastroesophageal Reflux Disease (GERD)		Medium	Episodic	Medium	High
40.0 M \$9.6 B					
Hypertension		High	Steady	Medium	High
73.0 M	\$51.0 B		Progression		
Inflammatory 1.4 M	Bowel Disease \$18.8 B	High	Episodic	High	Medium
Lupus (SLE) 1.5 M \$8.0 B		High	Episodic	High	High
Migraines		Medium	Episodic	High	High
Multiple Sclerosis		High	Episodic	High	Medium
Osteoarthritis		Medium	Episodic	High	High
Osteoporosis		High	Steady Progression	High	High
10.0 M \$14.0 B Stroke		1 Back	- J		
5.8 M	\$43.7 B	High	Episodic	High	High



Top 20 examples of chronic diseases out of 91 studied by EAC. Noted are 9 Chronic Diseases often seen as comorbidities of diabetes. Centers for Disease Control and Prevention CDC 24/7: Saving Lives, Protecting People™

Determinants of Health in U.S.



Scientific Wellness → A New Industry



Conceptual Themes of P4 Medicine



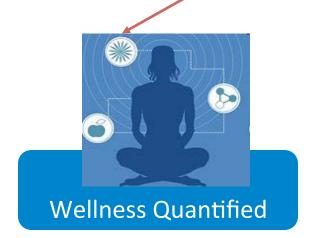
P4 Medicine

Predictive

Preventive

Personalized

Participatory



Disease Demystified

Scientific Wellness Industry

Disease Industry

Proposing the 100K Wellness Project





Clinical OMICs INNOVATOR

Promoting Wellness & Demystifying Disease:

The 100K Project

Leroy Hood, M.D., Ph.D., and Nathan D. Price, Ph.D.

Hood and Price. Clinical Omics, (2014)

EDITORIAL



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Citation:

L. Hood, N. D. Price, Demystifving disease, democratizing health care. Sci. Transl. Med. 6, 225ed5 (2014)

SYSTEMS BIOLOGY

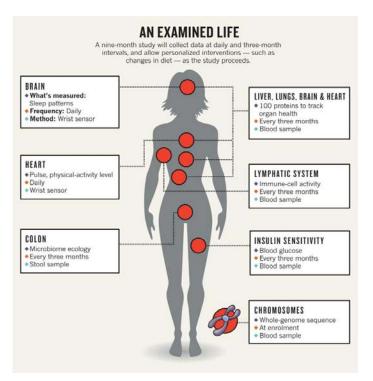
Demystifying Disease, Democratizing **Health Care**

UNSUSTAINABLE COST INCREASES THREATEN THE GLOBAL HEALTH CARE SYSTEM. and further progress is stymied more by societal than technological factors. Only by engaging health care consumers (that is, patients) as pioneers who provide both health-related data and insights into pathophysiology can we meet these societal challenges and thus accelerate the pace of biomedical innovation.

In March 2014, the Institute for Systems Biology will launch a longitudinal, Framinghamlike study (www.framinghamheartstudy.org) of 100,000 (100K) healthy individuals that we believe will be instrumental in bringing predictive, preventive, personalized, and participatory (P4) medicine to patients. Participatory medicine means that patients, researchers, physicians, and the entire health care community join forces to transform the practice of medicine to make it more proactive than reactive-and, in turn, less expensive and more

A systems approach is necessary for the effective management of complex diseases (1). This fundamental component of P4 medicine is built on two central features. The first is a conviction that, in 5 to 10 years, each patient will have a dynamic data cloud consisting of billions of diverse types of data points and that medicine will be informed by computational analyses that reduce high-dimensional data to actionable hypotheses designed with the intent of optimizing wellness and minimizing disease in individual patients. The second feature is that integration of patient data will reveal biological networks that specify health and are altered in disease, and that through an understanding of these differences, one can gain fundamental insights into disease mechanisms. Such insights are essential for developing more effective diagnostic and therapeutic approaches. Indeed, such an approach has already provided powerful new technologies and strategies (2) that have brought us to the brink of

At its foundation, P4 medicine is about quantifying wellness and demystifying disease. Individual data clouds will let us predict future wellness and disease. The preventive element focuses on how well we can improve individual wellness and take actions to stop or deNature. News piece, (2014)



Hood and Price, Science Translational Medicine (2014)

PIONEER 100 PROJECT

Principal Investigators: Lee Hood and Nathan Price

The 100K Wellness Project was initiated in 2014 with the generation of dynamic data clouds for 108 individuals. These data provided spectacular insights into what it is to be well and the nature of wellness to disease transitions (and vice versa).





- 108 participants
- Age range: 20s to 88+
- 9-month study
 launched March 2014
- IRB approved
- Evaluation / insights for next phase

- Whole genome sequence
- Detailed blood, urine, saliva measurements 3x
- Gut microbiome 3x
- Continual self-tracking and lifestyle monitoring
- Data integration & correlations
- Monthly coaching sessions on actionable data
- Discovery research
- Events and education

Assays / Measurements—108 Pioneers



Creating dense and dynamic personal data clouds

GENOME

Whole Genome Sequencing. **SNPs Millions**

Continual

& lifestyle monitoring

self-tracking

Database of actionable possibilities that

will grow over

time

LABS

Detailed lab tests 3x (blood, urine, saliva) Clinical chem. 150 **Metabolites 700 Proteins 400**

MICROBIOME

Gut Microbiome 3x



Wellness coaching for participants



Wellness Coach



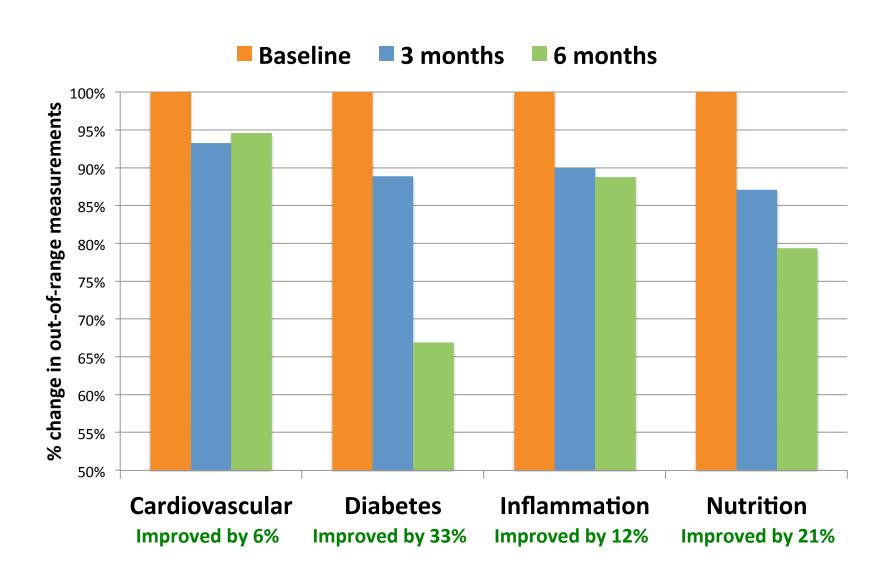
Sandi Kaplan, MS, RD

Study Physician



Craig Keebler, MD

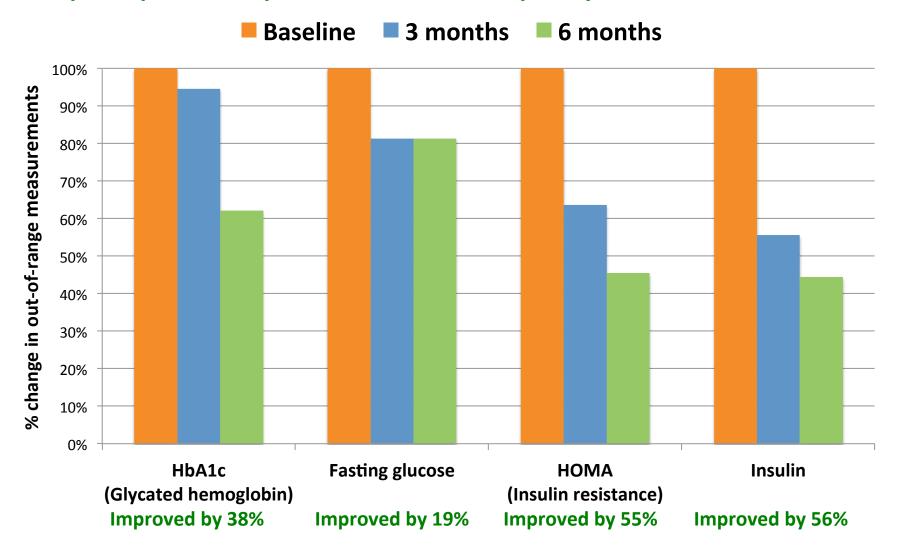
Clinical Labs Discovery: Improvements in blood health with behavioral coaching



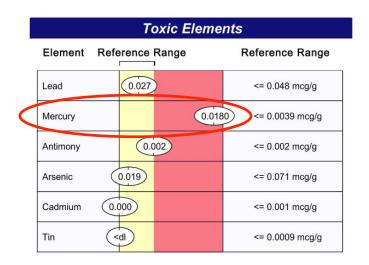
Clinical Labs Discovery: Significant pre-diabetes improvements

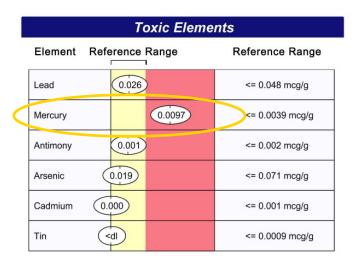


Seven participants with pre-diabetes were completely normalized in six months



Diet modification to reduce heavy metal toxicity





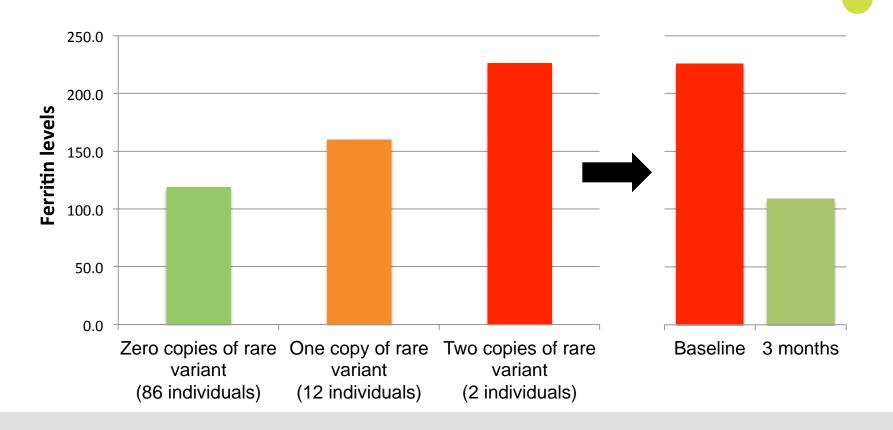


- 1. Baseline: High mercury levels in blood
- Coached to modify diet

 eight weeks of eating salmon sushi vs. tuna sushi (3x a week)
- 3. Reduced mercury levels in three months

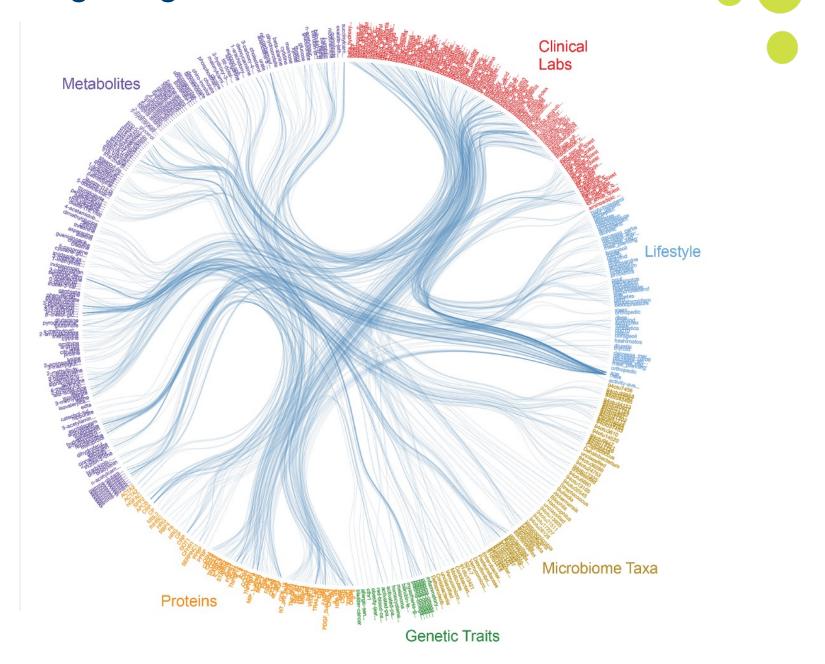
A wellness to disease transition—genetics plus environment—an actionable possibility

Genetics and Clinical Labs: Hemochromatosis Detected risk of a deadly disease in two participants

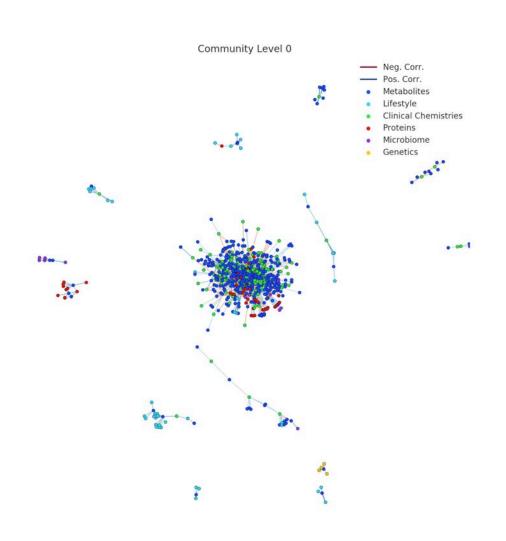


- Blood + Genetics illuminated the effects of increasing copies of the Hemochromatosis variant
- Left untreated, this disorder could lead to cartilage damage, liver cancer, diabetes, and heart disease: Easily treated by regular blood donations to reduce the iron stores
- One participant ALREADY had cartilage damage from his undiagnosed disease
- Subsequent family genetic testing detected other family members at risk

Deriving Insights from Data: New Frontiers

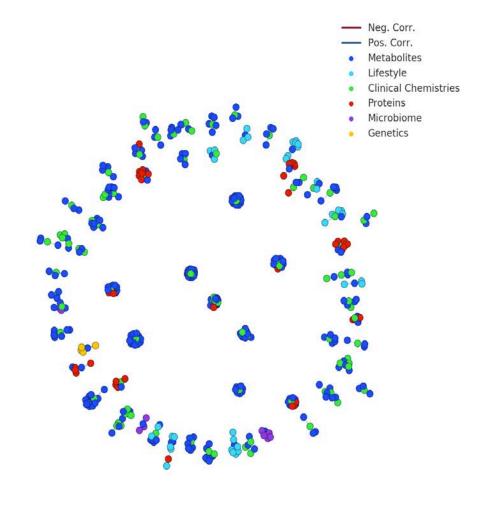


Identifying inter-related molecular modules

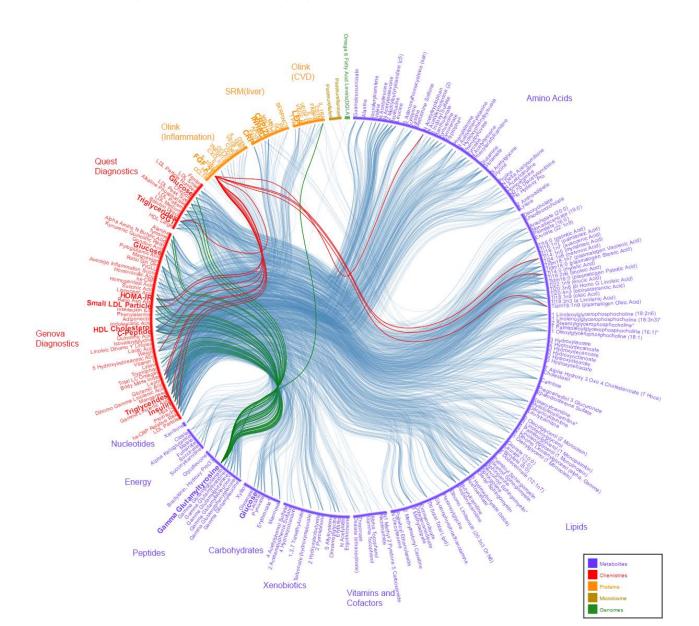


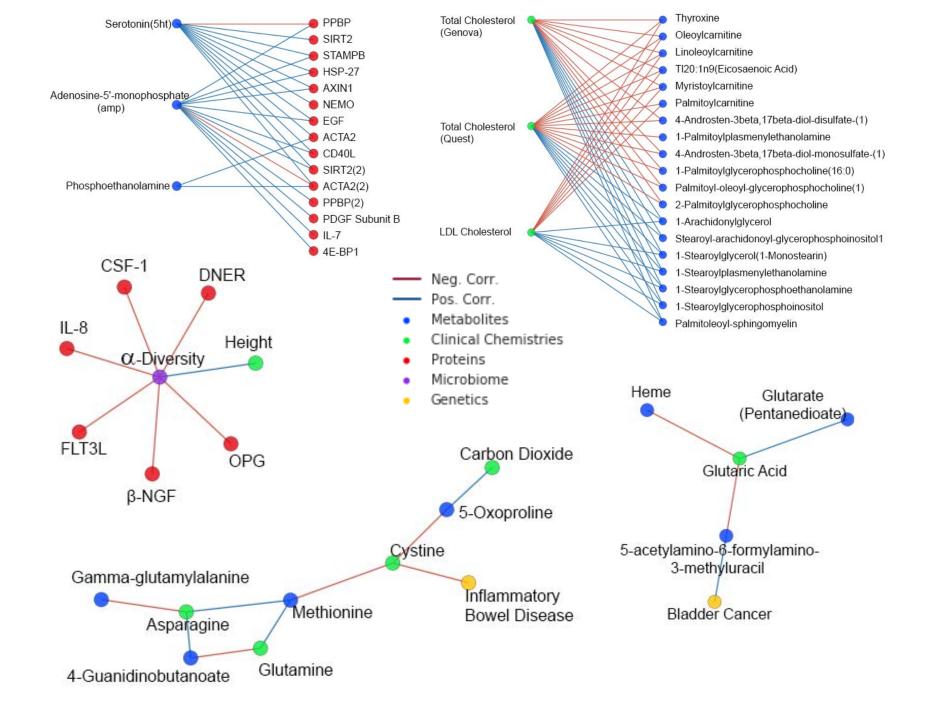
Total cholesterol community

- Cholesterol is positively associated with alphatocopherol (Vitamin E)
- Cholesterol is negatively associated with endogenous thyroxine
- A beneficial side effect of the drug thryroxine (Synthroid) is lowering LDL cholesterol



The largest molecular community: related to cardiometabolic health





We can determine your genetic risk for at least 60 diseases.



GWAS variants have been determined for about 60 diseases and traits

ADHD Alzheimer's disease Anorexia **Asthma** Atrial fibrillation **Breast cancer** Bipolar disorder Blood pressure Bone mineral density Inflammation Calcium Cardiovascular disease Celiac disease Cholesterol levels Chronic kidney disease Colorectal cancer

Coronary heart disease

COPD Crohn's disease Esophageal cancer Gout Grave's disease Hematocrit Hypertension Hypothyroidism Inflammatory bowel disease Iron levels Lung Cancer Lupus Macular degeneration Magnesium levels Metabolic syndrome Migraine Multiple sclerosis

Myopia Obesity Osteoarthritis Osteoporosis Ovarian cancer Pancreatic cancer Parkinson's disease Primary biliary cirrhosis Prostate cancer **Psoriasis** Rheumatoid arthritis Schizophrenia Stroke Type 1 Diabetes Type 2 Diabetes Ulcerative colitis **Urate levels**

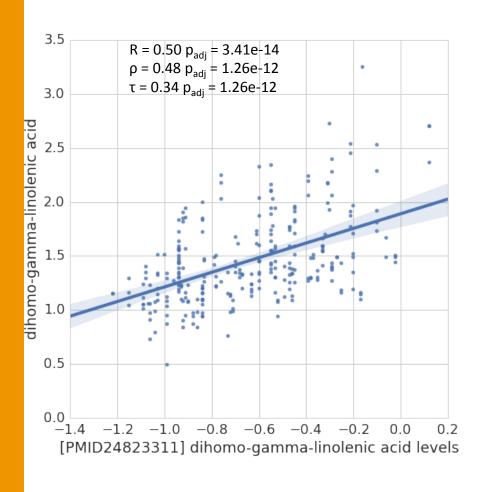
Vari rs85

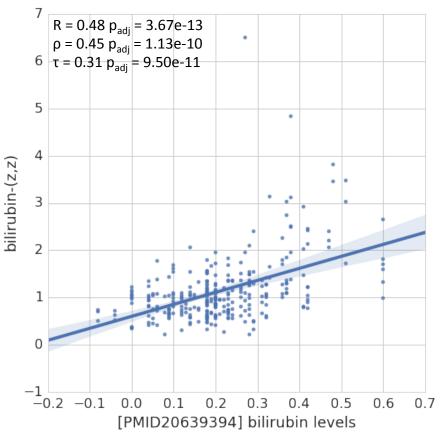


Variant rs9769

Your genome affects metabolite concentrations in your blood

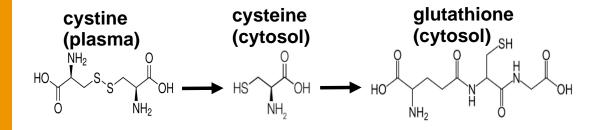




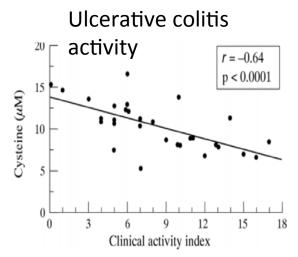


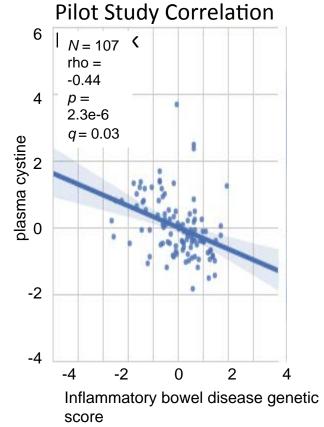
Nutrient measurements correlate with genetic predisposition for IBD





	Cystine	
Control (n=65)	61.3 (1.7)	
Crohn's disease (n=33)		
Before surgery	42.8 (2.4)***	
10 days after surgery	56.0 (3.0)	
3 months after surgery	52.7 (2.8)1-160	
Ulcerative colitis (n=33)		
Before surgery	47.3 (1.8)***	
10 days after surgery	64.3 (2.4)	
3 months after surgery	64.5 (3.6)	





Sido, B., Hack, V., Hochlehnert, A., Lipps, H., Herfarth, C., and Dröge, W. (1998). Impairment of intestinal glutathione synthesis in patients with inflammatory bowel disease. Gut *42*, 485–492.



Where are we going?

Scientific Wellness: Two Integrated Directions



Arivale

- A consumer facing scientific wellness company
- 5,000 individuals in the first 18 months
- Transform how biotech industry operates

ISB-Providence

- Dense, dynamic, personal data clouds
- Research to validate wellness metrics
- Research for better assays
- Optimize wellness
- Study wellness to disease transitions
- Study disease [progression, response to therapy and transition to wellness



Arivale: Scientific Wellness at Scale

Founders



Clayton Lewis, CEO and Co-founder
Maveron, MarketLeader, Harborview Medical Center, Capitol Hill

Lee Hood, MD, PhD, Co-founder, SAB Chair ISB, Amgen, National Academies, Presidential Medal

Nathan Price, PhD, Co-founder, BOD ISB, University of Illinois, Urbana-Champaign, UCSD, UW

Customer Acquisition & Engagement



Grant Ries, Chief Revenue Officer Founder: Bluekai; Oracle, AdReady, YuMe

Alicia Nakamoto, Director, Community RealSelf, Amazon, Bing, HP

Stewart Meyer, Director, Client Experience Amazon, Razorfish, Kinetix, Best Buy

Computational Bioscience & Software Development



Bryan Wheeler, MS, Director, Software Development Amazon, MSNBC.com, Microsoft

Andrew Magis, PhD, Senior Bioinformatics Scientist ISB, University of Illinois, Urbana-Champaign

Behavioral Coaching



Jennifer Lovejoy, PhD, Chief Translational Science Officer Alere/ Free & Clear, Bastyr Dean, Pennington Biomedical Research Center (LSU) Endowed Professorship

Sandi Kaplan, MS, RD, Director, Coaching Services Alere / Free & Clear, founder Zing Bars

Operations & Finance



Sean Bell, Chief Business Officer ISB, Alere / Free & Clear, LexisNexis

Kern Maresca, Director of Finance Alere / Free & Clear, MindMyBody

External Affairs



Gretchen SorensenISB, Clinton Administration, Turner Broadcasting, Capitol Hill

Capital Raised: \$39M







Scaling Up: Building the Research Portal

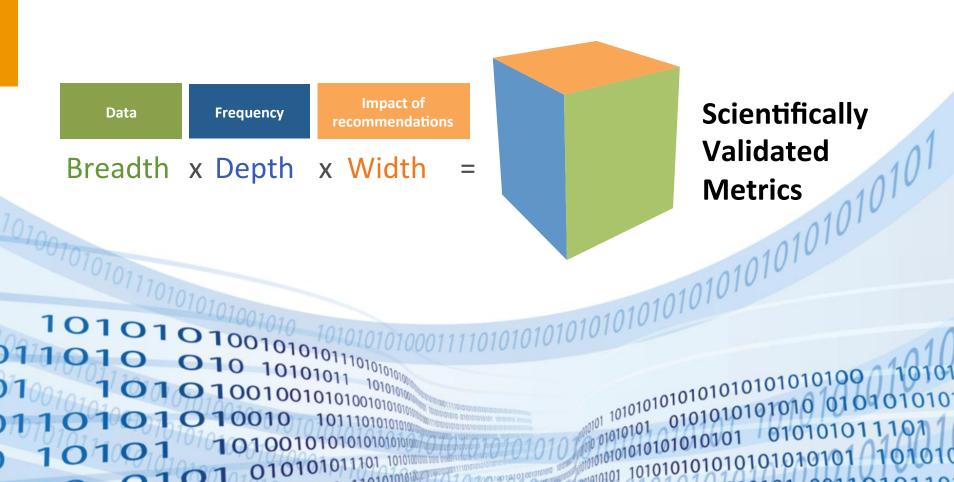


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From 108 to Thousands to Millions

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PROVIDENCE-ISB AFFILIATION

Tens of millions of EMRs
Have access to patient samples
Can work with Providence to design clinical trials
Provides avenue to implement P4 Medicine

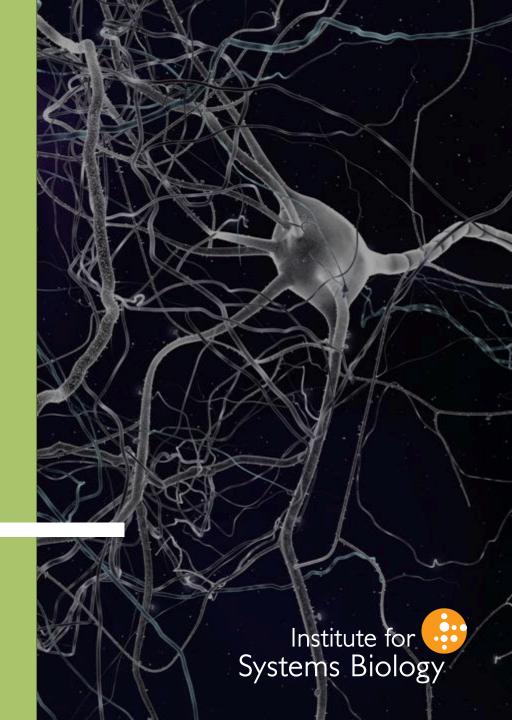
SCIENTIFIC WELLNESS



BREAST CANCER SURVIVOR WELLNESS



ALZHEIMER'S DISEASE



(PRE)DIABETES



Dense, Dynamic Personal Data Clouds

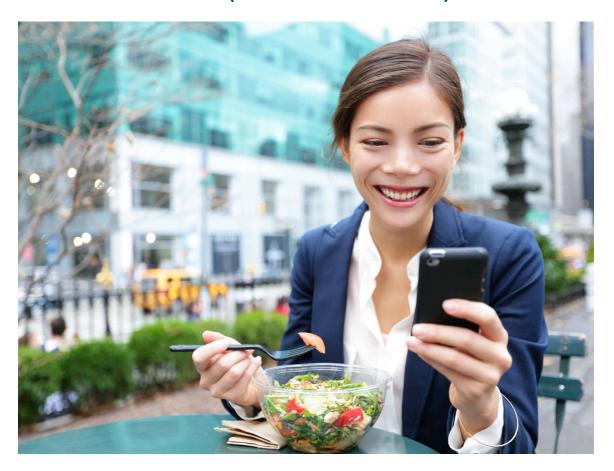


These personalized data clouds are the foundation of Precision Medicine and Precision Nutrition.



Enabling Individuals to take Responsibility for their Own Wellness (and Disease)





Individuals taking responsibility for their own health – with informed personalized nutrition – will dramatically reduce the cost of healthcare

ISB Hundred Person Wellness Project: Team

Special thanks to our funders: Robert Wood Johnson Foundation and M.J. Murdock Charitable Trust

Project Leadership

- Leroy Hood, MD, PhD
- Nathan Price, PhD
- Sean Bell, Business Director

Participant Engagement

- Jennifer Lovejoy, PhD, VP Clinical Affairs
- Sandi Kaplan, Wellness Coach
- Craig Keebler, MD, Study Physician

Data Analytics

- Nathan Price, PhD Analytics Lead
 Gustavo Glusman, PhD, Genomics
- Andrew Magis, PhD, Multi-omics
- John Earls, Data integration

Project Management

- Kristin Brogaard, PhD Project Manager
- Sara Mecca, Project Assistant
- Mary Brunkow, PhD, Project Coordinator

Communications

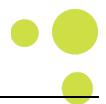
- Gretchen Sorenson, Consultant
- Hsiao-Ching Chou, Commun. Director

Medical Advisory Board

- Robert Green, MD
- Jane Guiltinan, ND
- Michael Raff, MD
- Sarah Speck, MD



Presentation Clinical Actions



After participating in this presentation, clinicians should be better able to:

 Discuss the power of data in driving clinical focus towards disease prevention and wellness optimization.