



# An Overview of Genetype

ERIKA SPAETH, PhD SCIENTIFIC/CLINICAL AFFAIRS



### **About Rhythm Biosciences**



We are committed to saving lives through the evaluation of cancer risk and early detection of cancers



Early detection of cancer gives individuals the best possible opportunity for favorable treatment outcomes.

Addressing large under-served global markets worth >USD\$50bn and heavily supported by public and private health care systems.

geneType<sup>TM</sup> commercially available now and ColoSTAT<sup>®</sup> anticipated commercial launch in 2H CY2025.

Patent protected, fully characterised proteomic, genomic technology that can be readily adopted in routine laboratories.

A valuable development pipeline of high quality, impactful cancer risk assessment and detection products

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### Corporate overview



#### **CORPORATE SNAPSHOT**

ASX Code	RHY
Share Price (17th June 2025)	\$0.060
Shares on Issue	283,597M
Unlisted & Listed Options	53.4M
Market Capitalisation	\$17.61M
Cash in Bank (31 March 2025)	\$2.3M
Top 20 Shareholders	43.3%

#### **SHARE PRICE CHART - ASX:RHY**



#### **BOARD AND MANAGEMENT**

#### David Atkins, PhD CEO & MD



Otto Buttula Non-Executive C



Sue MacLeman
Non-Executive Director



**Gavin Fox-Smith**Non-Executive Director



Former CEO of Congenica (UK) & Synevo Diagnostics, Sr. Executive at Johnson & Johnson and Danaher.

Founder of Veridex – cancer molecular and cellular diagnostics (USA).

Experience in product development and commercial leadership in the global medical device and diagnostic industry.

#### Non-Executive Chairman

Extensive financial, investment, IT & biotech experience.

Co-Founder and CEO of IWL (ASX: IWL); Founder / former CEO of Investors Mutual

Formerly a Director of Imugene (ASX:IMU), Chairman of Investorfirst, now HUB (ASX: HUB), HITIQ (ASX: HIQ) & Oncosil Medical (ASX: OSL). 30 years in Pharma, Biotech and Medtech including Amgen, BMS and Merck and SME's.

Experienced Board member, former CEO of NASDAQ, ASX, & AIM entities. Currently Chair of Medicines Australia, NED at Viral Vector Manufacturing Facility, Smartways Logistics, Healthcare Translation Group, ATSE & OMICO & member of the National Research Infrastructure (NRI) Advisory Group.

38 years as a leader and champion of Medical Technology in Australia/NZ and Asia.

Gavin is Chair of ANDHealth (Australia's National Digital Health Initiative), Board Director for Bowel Cancer Australia and Board Director for SAN Foundation.

Gavin was previously CEO of Omnigon, a private Australian company in the Ostomy Care market.

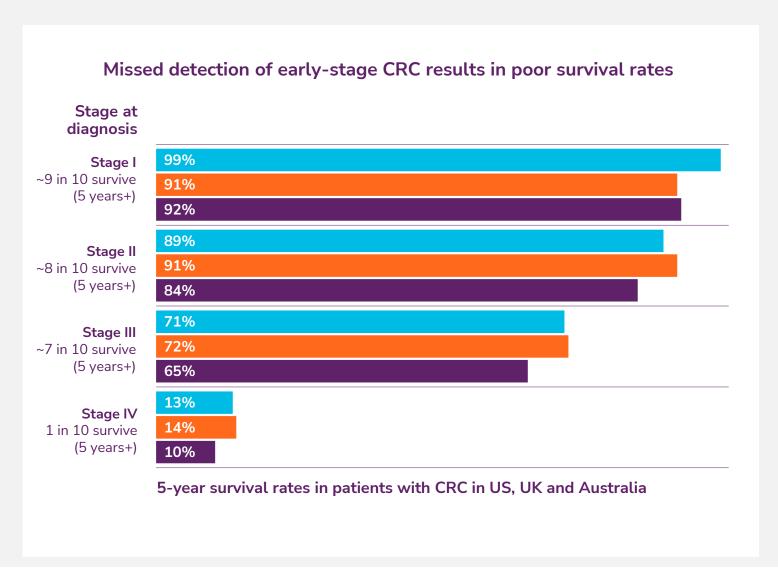
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### When diagnosed early, cancer can be successfully treated



Disease stage for colorectal cancer diagnosis is the key predictor of survival

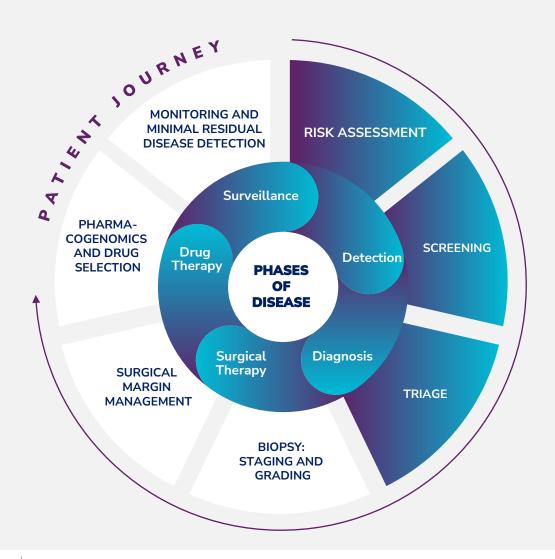
Diagnoses at Stage I or II only represent less than half of all CRCs diagnosed in:						
			Stage I		Stage II	1
<ul><li>AUSTRAL</li></ul>	-IA		22%	0	23%	0
• US			37%	0	37%	0
• UK			16%	0	21%	0
Disease treatment costs increase with later stage#						
				\$256k		
\$1	l11k	\$128k	\$145k		Total 1 in USA	.st year costs
Stage at diagnosis	1	Ш	Ш	IV		



### Detecting cancer earlier and maintaining wellness

The power of the combination of the Rhythm capabilities and portfolio







#### Risk Assessment

Determination of risk profile

Personalised management of health

#### **COLOSTAT**®

#### **Disease Screening**

Minimally-invasive detection of disease

Early detection of disease

#### A powerful combination enabling:

- Supports preventative measures to maintain wellness
- Increase the likelihood of earlier disease detection
- Promote better treatment outcomes for detected disease
- Potential use in drug clinical studies for patient stratification
- Potential for development as companion diagnostic tests

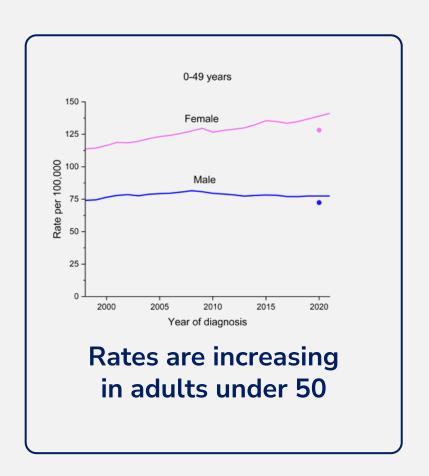
#### Suitable for 100s millions of individuals/patients globally

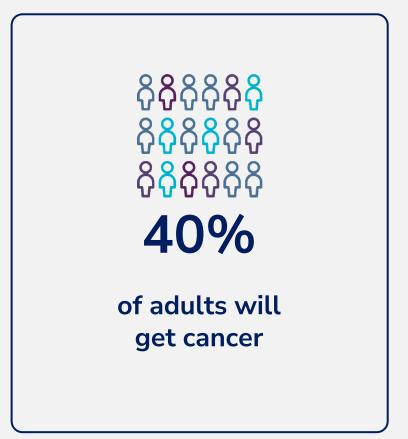
**Example**: geneType<sup>™</sup> bowel cancer risk assessment of the general population to help guide usage of ColoSTAT<sup>®</sup>. Potentially leads to improved screening compliance, earlier disease detection and improved outcomes.

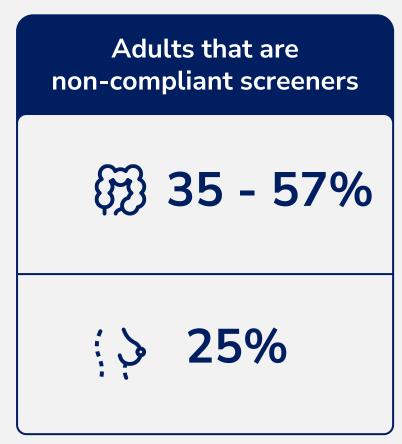
### Cancer screening is critical to managing cancer burden



Detection of cancer as early-stage disease leads to better outcomes







https://www.cancer.org/research/acs-research-news/cancer-incidence-rate-for-women-under-50-rises-above-mens.html https://www.cancer.gov/about-cancer/understanding/statistics

Kim RY, Rendle KA, Mitra N, et al. Adherence to Annual Lung Cancer Screening and Rates of Cancer Diagnosis. JAMA Netw Open. 2025;8(3):e250942. doi:10.1001/jamanetworkopen.2025.0942 https://nccrt.org/our-impact/data-and-progress/

https://www.cdc.gov/pcd/issues/2023/23\_0071.htm#:~:text=(RTI%20International).-,Results,(Tables%202%20and%203).

### **Current Screening programs**

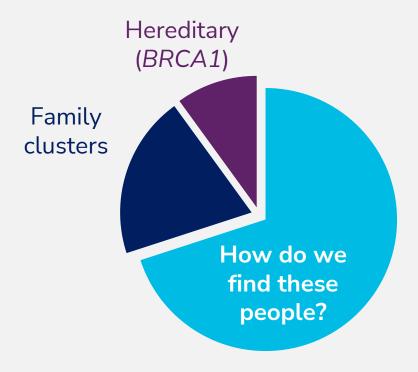
The standard of care varies in each geography





One Size Fits All







#### **Breast**

- Start at 40 or 45
- Mammogram
- Biennial



#### **Colorectal**

- Start at 45
- FIT,
   Cologuard,
   colonoscopy
- Frequency dependent on method



#### **Prostate**

- Start at 55
- PSA
- Frequency/ start dependent on risk

### Cancer is unique to each individual

RHYTHM"
BIOSCIENCES

Detection of disease needs to reflect this uniqueness



Improve the current standard of care from one-size-fits-all screening

### Personalised screening approaches

- Precisely identify who is at risk
- Provide tailored risk-reduction options

### Risk Assessment is the Gateway to Precision Diagnostics



The first step to employing personalised screening approaches

Risk assessment

Screening

Diagnosis

Personalised treatment

Prognostic response

Recurrence

## WHAT?

Predicting a person's chance of developing a condition/disease

### WHY?

To provide more appropriate screening/risk-reduction to those who need it

### HOW?

Validated Assessment tool that integrates genomics

### geneType<sup>TM</sup> sets the stage for personalised screening options



Risk assessment allows us to move away from a one-size-fits all screening approach



### Our understanding of the range of risk factors has advanced



Each individual's risk is a factor of multiple variables

#### Scientists can assign VALUEs to risk factors:

- Family history
- Low BMI
- High BMI
- Cholesterol levels
- Blood pressure
- Number of live births
- Smoking habits
- Use of aspirin
- Last colonoscopy
- Last mammogram
- Genetic variants





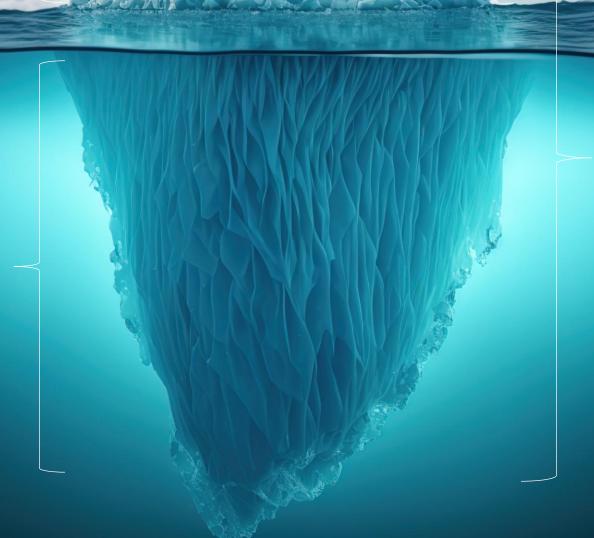
### Traditional Risk Assessment

Easy-to-identify risk factors



### Genetic Risk

Polygenic risk score



### geneType<sup>TM</sup>

Looks at the bigger picture of risk

### geneType<sup>TM</sup> is different from other genetic testing



Genetype detects many common low penetrance genetic variants vs. individual mutations.



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### BRCA1/2

- 1 mutation
- Very high risk
- Rare in the population (0.2-1%)
- Sonic, Healius, LabCorp, Myriad

### geneType<sup>TM</sup>

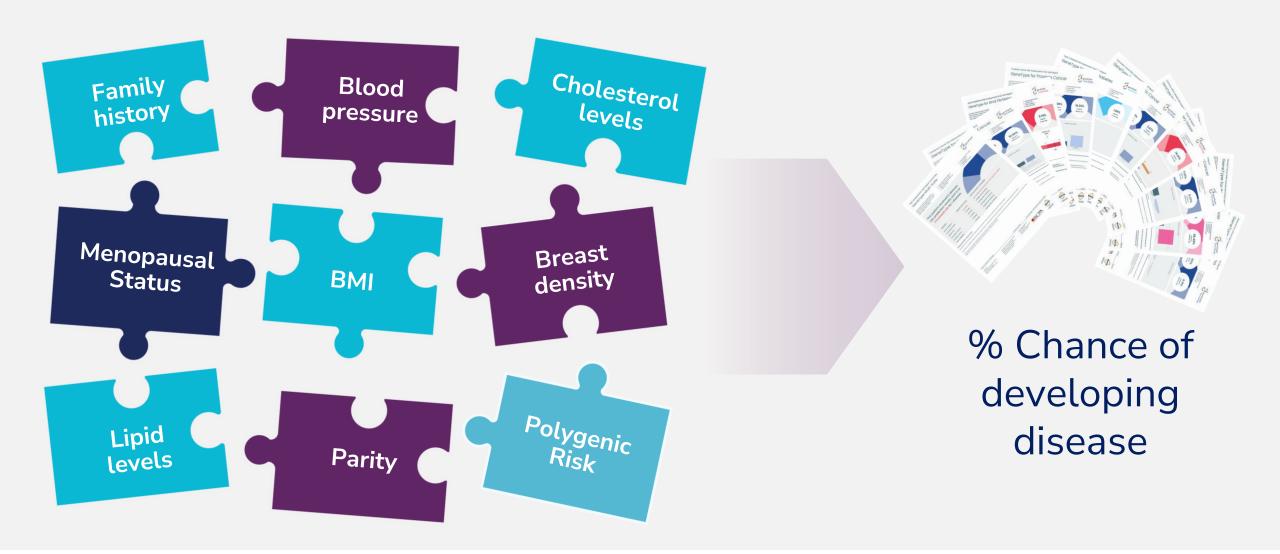
- Many risk factors: genetic and clinical
- increased risk
- Up to 20% of the population

Hickmann, E., et al. BMC Health Serv Res 22, 1116 (2022). Venner, E., et al Commun Biol 7, 174 (2024). HU et al. N Engl J Med 2021;384:440-451

### geneType<sup>TM</sup> risk assessment



Disease risk prediction is strongest when combining multiple risk factors



#### Risk Assessment Test Report Summary

#### GeneType Multi-Suite

Laboratory Accession Number: HG01342 Date of Specimen Collection: 02-Feb-2022 Date of Laboratory Receipt: 09-Feb-2022 Date of Report: 10-Jun-2025

Patient Name: Jane Doe Date of Birth: 28-Dec-1974 Patient Address: 1 Example Road Suburbia, VIC 3000

Ordering Medi Dr Sam Bloggs Sampletown, 1

Laboratory Accession Number: HG01342 Date of Specimen Collection: 02-Feb-2022 Date of Laboratory Receipt: 09-Feb-2022 Date of Report: 07-Feb-2025

Patient Name: Cynthia Richardson Date of Birth: 28-Dec-1974 Patient MRN: Potient Address: Il Bourke Street

Melbourne, VIC 3000

Ordering Medic Dr Laura Jones 12357 Sandpipe Melbourne, VIC

Laboratory Accession Number: HG01342 Date of Specimen Collection: 02-Feb-2022 Date of Laboratory Receipt: 09-Feb-2022 Date of Report: 10-Jun-2025

Patient Name: Jane Doe Date of Birth: 28-Dec-1974 Patient MRN: Patient Address: 1 Example Road Suburbia, VIC 3000

Ordering Medical Practitioner: Dr Sam Bloggs 42 Demo Street Sampletown, VIC 3000

#### This patient received 8 disease risk prediction results and is at increased risk for 2 diseases.



#### Interpretation Summary

This patient has an increased risk of developing Atrial Fibrillation, Breast Cancer. Please read the appended reports for speci

This patient is considered within the average range of risk for an adult of the same age, sex and ethnicity for the remaining

The risk scores are patient-specific and connot be used to estimate risk in relatives. Furthermore, the following results should patient's full clinical history, particularly for patients close to a threshold risk value.

#### This patient is at an **INCREASED**

risk of breast cancer

Breast Cancer Risk Assessment Final Test Report

GeneType for Breast Cancer





\*The average risk is based on the same age, biological gender and race/ethnicity as the patient from the general Australian

#### Interpretation

This patient has a 31.53% chance of developing breast cancer within her remaining lifetime up to age 90 years which is consi not mean that the patient will develop breast cancer over this timeframe.

In accordance with the RACGP Guidelines for Preventive Activities in General Practice, any patient with a remaining lifetime ris their remaining lifetime risk is between >13% and <25% they are at a moderately increased risk and if their remaining lifetime r

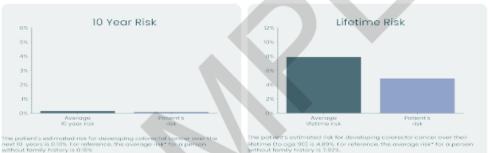
The patient should continue following general population breast screening protocols at a minimum, regardless of their estim risk scores are patient-specific and cannot be used to estimate risk in relatives. Furthermore, these results should be interprecontext of the patient's full clinical history, particularly for patients close to a threshold risk value.

#### This patient is at an AVERAGE risk of developing colorectal cancer

Colorectal Cancer Risk Assessment Final Test Report

GeneType for Colorectal Cancer





\*The average risk is based on the same age, biological gender and race/ethnicity as the patient from the general Australian population

#### Interpretation

This patient has a 0.10% chance of developing colorectal cancer within 10 years, which is considered average risk.

In accordance with Cancer Council Australia, any patient with a 10-year risk score <2x average risk is defined as average risk, between 2-4x moderately increased risk, and >4x at high risk.

The patient should continue following general population colorectal screening protocols at a minimum, regardless of their estimated risk score. Also note that the risk scores are patient-specific and cannot be used to estimate risk in relatives. Furthermore, these results should be interpreted by a medical practitioner in the context of the patient's full clinical history, particularly for patients close to a threshold risk value.

Report continued on next page

v2 May 2025

Summary pa

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Sequenced by Gene By Gene

CUA# 45DT02202 | CAP# 7212851 1445 N Loop W STE 820 Houston, TX 77008 Analyzed and Reported by ichorDX inc. CUA# 99D2023356 1300 Baxter St STE 255, Charlotte NC 28204

Reported by designated pathologist: Dr. Peter Kaub Compliant with ISO15189 and NPAAC requirements

Sequenced by Gene By Gene CUA# 45D102202 | CAP# 7212851 1445 N Loop W STE 820 Houston, TX 77008 Analyzed and Reported by IchorDX Inc. CHA# 9902023356 1300 Baxter St STE 255, Charlotte NC 28204

Laboratory Director: Angle Purvis, PhD Reported by designated pathologist: Dr. Peter Kaub Compliant with ISOI5189 and NPAAC requirements

customersupport@genetypeus US Tel: +1.704.926.5700

AU Tet +61 3 8256 2880

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Sequenced by Gene By Gene CUA# 450102202 | CAP# 7212851 1445 N Loop W STE 820 Houston, TX 77008 Analyzed and Reported by ichorDX inc. 1300 Bristor St STE 255, Chordotto NC 28204

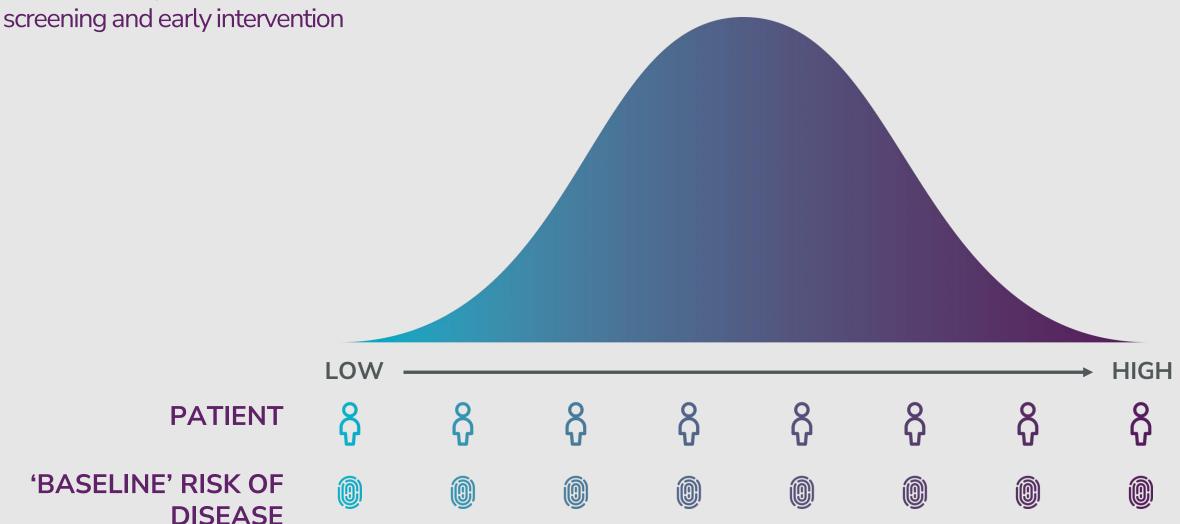
Laboratory Director: Angle Purvis, PhD Reported by designated pathologist: Dr. Peter Kaub Compliant with ISCIEI89 and NPAAC requirements

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### Inherent risk: We are all born with a polygenic risk profile



Understanding where a patient sits on the spectrum of risk unlocks new opportunities for precision



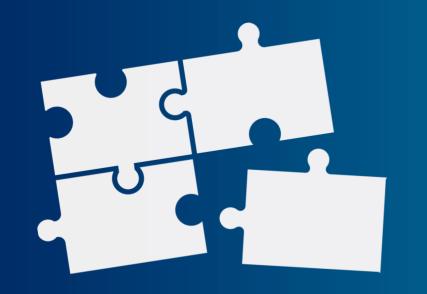
Spaeth E. Chapter 2 - Polygenic risk scores: A conceptual overview to add PRS to your clinical toolbox. In: Kelly L, Stanford WP (eds). Implementation of Personalized Precision Medicine. Academic Press, 2025, pp 23-33.

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geneType Population Risk Assessment

### The future of personalised medicine







Personalised healthcare



Risk assessment

geneType Population Risk Assessment

### geneType<sup>TM</sup> development



Creating a risk assessment that contains the fewest, but most impactful risk factors

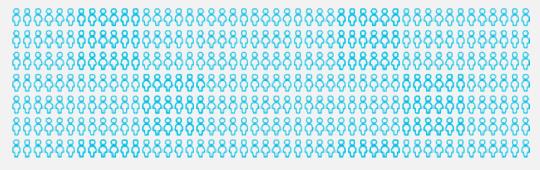
#### MAJOR CONSIDERATIONS

- Improve current binary risk (standard of care)
- Limit stress to healthcare provider (ease of implementation)

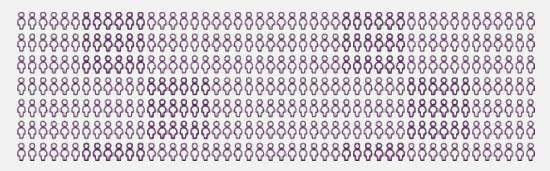
#### **PROCESS**

- Develop algorithm
  - literature review
  - access to large datasets for discovery/training
- Validate algorithm
  - access to large datasets for testing/validation
  - file IP/publish data
  - Tech transfer into accredited lab environment

#### THOUSANDS OF ADULTS WITHOUT THE DISEASE



#### THOUSANDS OF ADULTS WITH THE DISEASE



### geneType<sup>TM</sup> vs Gold Standard



Test portfolio consists of diseases that performed better than a current clinical standard tool



Across the following statistical metrics:

- ✓ Discrimination
- ✓ Calibration
- ✓ Net reclassification
- ✓ Net benefit

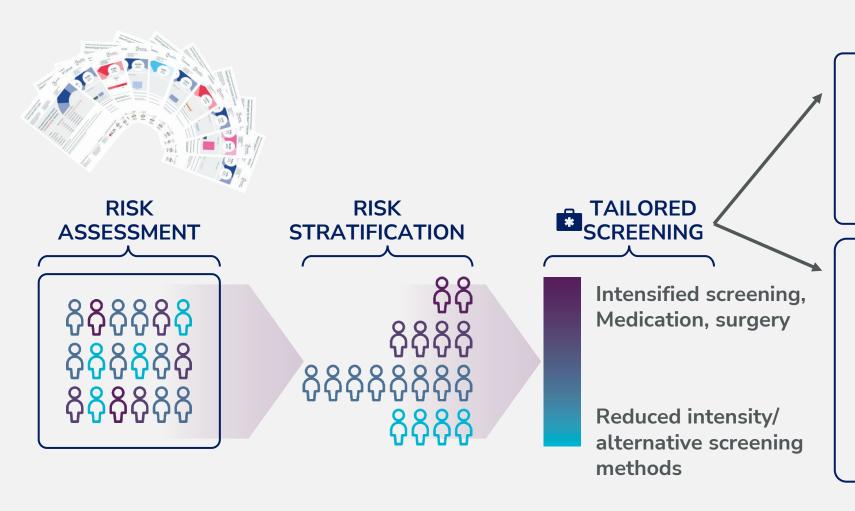
geneType <sup>TM</sup> Disease risk assessment				
Breast <sup>1,2</sup>				
Ovarian <sup>3</sup>				
Prostate <sup>4</sup>				
Colorectal <sup>5,6</sup>				
Pancreatic <sup>7</sup>				
Melanoma <sup>8</sup>				
Coronary artery disease <sup>9</sup>				
Atrial fibrillation <sup>9</sup>				
Type 2 diabetes <sup>9</sup>				

#### Peer-reviewed, published Clinical validation of the integrated cancer risk models

- 1. Allman R, Mu Y, Dite GS, Spaeth E, Hopper JL, Rosner BA. Validation of a breast cancer risk prediction model based on the key risk factors: family history, mammographic density and polygenic risk. Breast Cancer Res Treat 2023; 198: 335-347.
- 2. Spaeth EL, Dite GS, Hopper JL, Allman R. Validation of an Abridged Breast Cancer Risk Prediction Model for the General Population. Cancer Prev Res (Phila) 2023; 16: 281-291.
- 3. Dite GS, Spaeth E, Murphy NM, Allman R. A combined clinical and genetic model for predicting risk of ovarian cancer. Eur J Cancer Prev 2023; 32: 57-64.
- 4. Dite GS. Spaeth E. Murphy NM. Allman R. Development and validation of a simple prostate cancer risk prediction model based on age, family history, and polygenic risk, Prostate 2023.
- 5. Gafni A, Dite GS, Spaeth Tuff E, Allman R, Hopper JL. Ability of known colorectal cancer susceptibility SNPs to predict colorectal cancer risk: A cohort study within the UK Biobank. PLOS ONE 2021: 16: e0251469.
- 6. Spaeth Tuff EL, Gafni A, Dite GS, Allman R. Improvement of a clinical colorectal cancer risk prediction model integrating polygenic risk. Journal of Clinical Oncology 2023; 41: 81-81.
- 7. Dite GS, Spaeth E, Murphy NM, Allman R. Predicting 10-year risk of pancreatic cancer using a combined genetic and clinical model. Gastro Hep Advances 2023 in press
- 8. Wong CK, Dite GS, Spaeth E, Murphy NM, Allman R. Melanoma risk prediction based on a polygenic risk score and clinical risk factors. Melanoma Res 2023.
- 9. Wong CK, Makalic E, Dite GS, Whiting L, Murphy NM, Hopper JL et al. Polygenic risk scores for cardiovascular diseases and type 2 diabetes. PLoS One 2022; 17: e0278764.

# geneType<sup>TM</sup> risk stratification supports screening and prevention strategies





# Complementing existing screening pathways

- Age of entry
- Frequency
- Modality

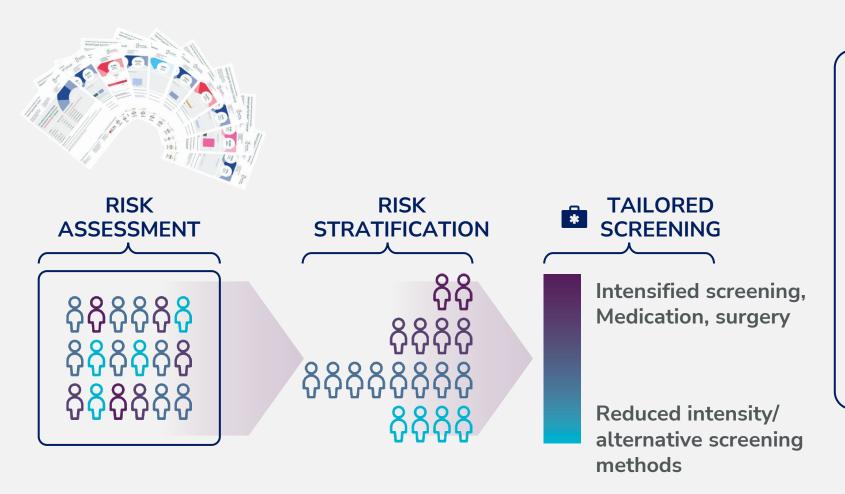
#### Novel screening technologies

- Single/Multi cancer early detection
  - ctDNA, cfDNA, protein, methylation
- Imaging
  - o Al-software, ultrasound, MRI
- Specimen types
  - Blood, saliva, tears, urine, stool

### Market entry: risk stratification across the healthcare ecosystem



From general practice to specialty care, and from private payors to public health programs—risk stratification enables targeted, value-driven decision-making at every level



#### **Applications**

- Applications
- Individualized patient care
- Primary care
- Specialty care services
- Imaging clinics, urologists, endoscopy clinics, family history/genetics clinics,
- Institutional/payor level care
- Self-insured employers
- US private payors
- Government sponsored care

### geneType<sup>TM</sup> performance has a material and affordable impact



Breast cancer risk assessment on a population basis and health economics

#### **Breast Cancer Risk**

- Cases diagnosed annually (incidence) = 20,640
- Individuals living with disease (prevalence) = 79,720
- Average lifetime risk = 1 in 7 or 14%

#### Risk Assessment Methods#:

Cross-validated in multiple cohorts of healthy women, age 40 – 69, followed for 5 years

Compared models: geneType<sup>TM</sup>



traditional clinical models (Gail#, IBIS^)



- **Data summary**<sup>#</sup>: **Out** o ~23.9K women of 200K women:
- classified as increased risk
- ~6.4K women classified as increased risk

Over 5 years, ~3.1K developed breast cancer:

- o 27% (861) were flagged by geneType
- o 6% of those (182) were flagged by traditional models

#### **Economics of Risk Assessment**





- Assumed 55% adoption rate
- At-risk cases receive supplemental screening (MRI) and annual mammograms
- Increase in early-stage cancers detected
- Decrease in interval cancers
- Reduction in total cost to care

Net saving of \$74 per participant per year

<sup>#</sup> Spaeth et al Cancer Prev Res 2023:16:1–12 ^Allman R. et al. Breast Cancer Res Treat 2023: 198: 335-347.

### geneType<sup>TM</sup> customer segment summary



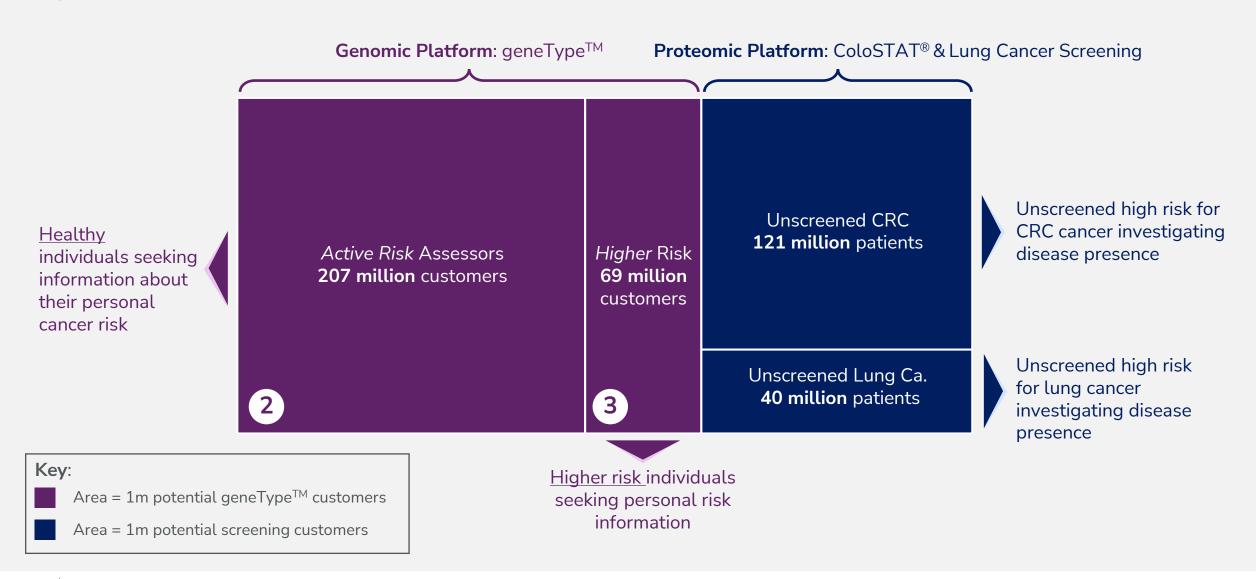
Three distinct customer segments where geneType<sup>TM</sup> would add significant value

	Worried well – Proactive Preventers	Active Risk Assessors – 2 Actively engaged in assessment	At Higher Risk – Seeking more information
Summary	A desire to understand their risks.	Encounter a reason to believe their risk is higher than average – typically age.	Urgency in need to assess risk. Available information unsatisfactory.
Use case	Otherwise, <u>healthy</u> but intervene to ensure longer healthspan.	Are engaged in population screening initiatives due to age, family or clinical history.	Possibly symptomatic or considered high risk. Reflex test for negative mutation testing or negative biopsy.
Sales channel	Corporate programs, primary carers, functional clinics, General Practice.	General Practice, Clinical geneticists, Specialists, Corporate programs.	Clinical geneticists, Specialists

### Rhythm is targeting a significant global market



A global market of >0.5bn individuals potentially served by the Rhythm product platform.



### There are multiple emerging drivers of geneType<sup>TM</sup> adoption



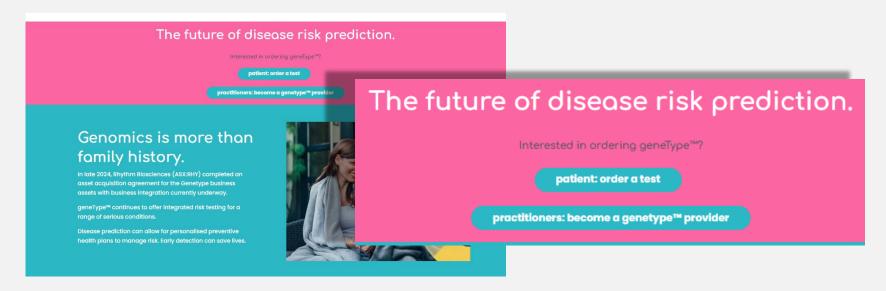
Rhythm is well positioned to participate in a major change in clinical practice

GeneType Collection Kit	Major Driver	Impact on geneType <sup>TM</sup> adoption
Clinical Trials: large cohorts	BARCODE1 <sup>1</sup> , WISDOM <sup>2</sup>	Efficacy and validation
Implementation Studies	CASSOWARY <sup>3</sup> , ROBIN <sup>4</sup> , GenoVA <sup>5</sup> , eMERGE <sup>6</sup>	Utility and usability
Clinical guidelines	Medical bodies/Gov't: US, EU, ASU	Guidance for routine use
Value Based Care	Public and private payors	Focus on outcomes
Health Economics & Reimbursement	AMA Coding, CMS coverage	Mainstream adoption
Advances in genomics	Massive growth in databases and cost o delivery	f Validation data, novel algorithms and reduced COGS
EMR Integration	Availability of test ordering in EMR platforms	Patient and physician engagement

### Ordering geneType<sup>TM</sup> in Australia using an HCP

The ordering options are continuously expanding

















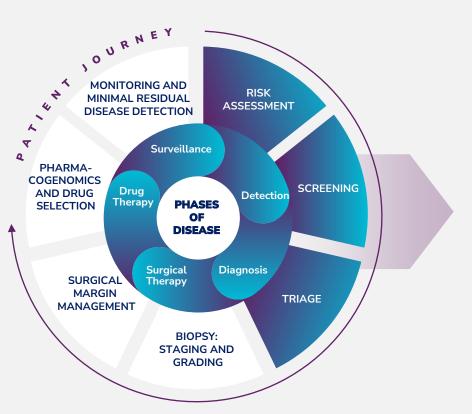




### Our combined platforms supports a highly comprehensive approach to cancer diagnostics



The potential RHY product portfolio addresses a huge total available market



Risk Assessment

Screening

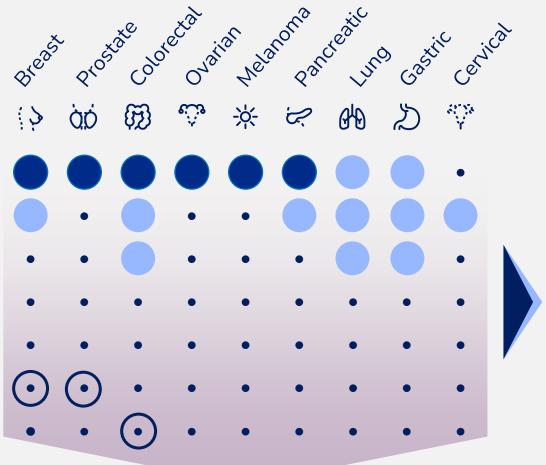
Triage

Biopsy

Surgical Margins

Drug Management

Monitoring



# Upcoming value inflection points



PRODUCT	ITEM	DESCRIPTION	COMPLETION DATE
ColoSTAT®	Kit Validation Ready Kit Verification completion, production kits re		Q2 CY 2025
ColoSTAT®	Final clinical validation	Evaluation of final system	Q3 CY 2025
geneType™	geneType™ strategic partnerships	Establishment of key commercial partners	Q3 CY 2025
geneType™	geneType™ menu expansion	Expansion of existing commercial portfolio	Q4 CY 2025
ColoSTAT®	Commercialisation	ColoSTAT® Partner's In House IVD launch	Q4 CY 2025
Lung cancer screening assay	Complete pre-clinical evaluation	Design of assay prototype	Q1 CY 2026

### Rhythm Bioscience: An attractive investment opportunity



#### **Overview**

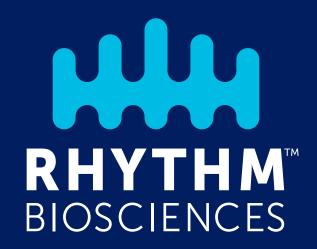
Rhythm's mission is to detect cancer earlier and reduce the burden of cancer. We are developing and selling proprietary, high value products and services designed to detect cancer earlier.

#### Growth

Rhythm is solving an important clinical and economic problem. We are preparing for rapid global commercial growth.

#### Reasons to Invest

- 1. Multi-product portfolio
- 2. Addressing a significant TAM
- 3. Planning for growth acceleration



### Thank you!

david.atkins@rhythmbio.com www.rhythmbio.com