



Investor Day Presentation

September 7, 2023



Transforming **patient lives** across the continuum of care

Helmy Eltoukhy, PhD

co-Chief Executive Officer, co-Founder
& Chair of the Board

AmirAli Talasaz, PhD

co-Chief Executive Officer & co-Founder

Safe harbor and non-GAAP disclosures

Certain statements in this presentation and the accompanying oral commentary are forward-looking statements within the meaning of federal securities laws. These statements relate to future events or Guardant Health, Inc. (the “Company”)’s future results and involve known and unknown risks, uncertainties and other factors that may cause the actual results, levels of activity, performance or achievements of the Company or its industry to be materially different from those expressed or implied by any forward-looking statements. In some cases, forward-looking statements can be identified by terminology such as “may,” “will,” “could,” “would,” “should,” “to,” “target,” “expect,” “plan,” “anticipate,” “intend,” “believe,” “estimate,” “predict,” “potential” or other comparable terminology. All statements other than statements of historical fact could be deemed forward-looking, including any expectations regarding the Company’s commercial engine as a force multiplier for research and development initiatives; any projections of market opportunities; statements about the Company’s ability to assess potential market opportunities or any statements about the Company’s ability to successfully develop new products and services; any statements regarding expectations for future reimbursement opportunities; any statements regarding the Company’s long-term expectations, including with respect to oncology, liquid biopsy, and other aspects of the Company’s industry; any statements about launching planned new products and additional laboratories, including with respect to Guardant Reveal, CGP tissue assay, and laboratories outside the United States; any statements about the Company’s ECLIPSE study; any statements regarding expectations for future regulatory approvals; any statements about historical results that may suggest trends for the Company’s business; any statements of the plans, strategies, and objectives of management for future operations and directions; any statements of expectation or belief regarding future events, opportunities to drive future growth, potential markets or market size, or technology developments; and any statements of assumptions underlying any of the items mentioned. The Company has based these forward-looking statements on its current expectations, assumptions, estimates and projections. While the Company believes these expectations, assumptions, estimates and projections are reasonable, such forward-looking statements are only predictions and involve known and unknown risks and uncertainties, many of which are beyond the Company’s control. These and other

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This presentation also contains estimates and other statistical data made by independent parties and by the Company relating to market size, penetration and growth and other data about the Company’s industry, which involve a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. In addition, projections, assumptions, and estimates of the Company’s future performance and the future performance of the markets in which the Company operates are necessarily subject to a high degree of uncertainty and risk.

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This presentation includes references to certain financial measures that are not calculated in accordance with GAAP. Reconciliation to the most directly comparable GAAP financial measure may be found in our most recent earnings release furnished to the SEC on August 3, 2023.

Today's agenda

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DAY | 20
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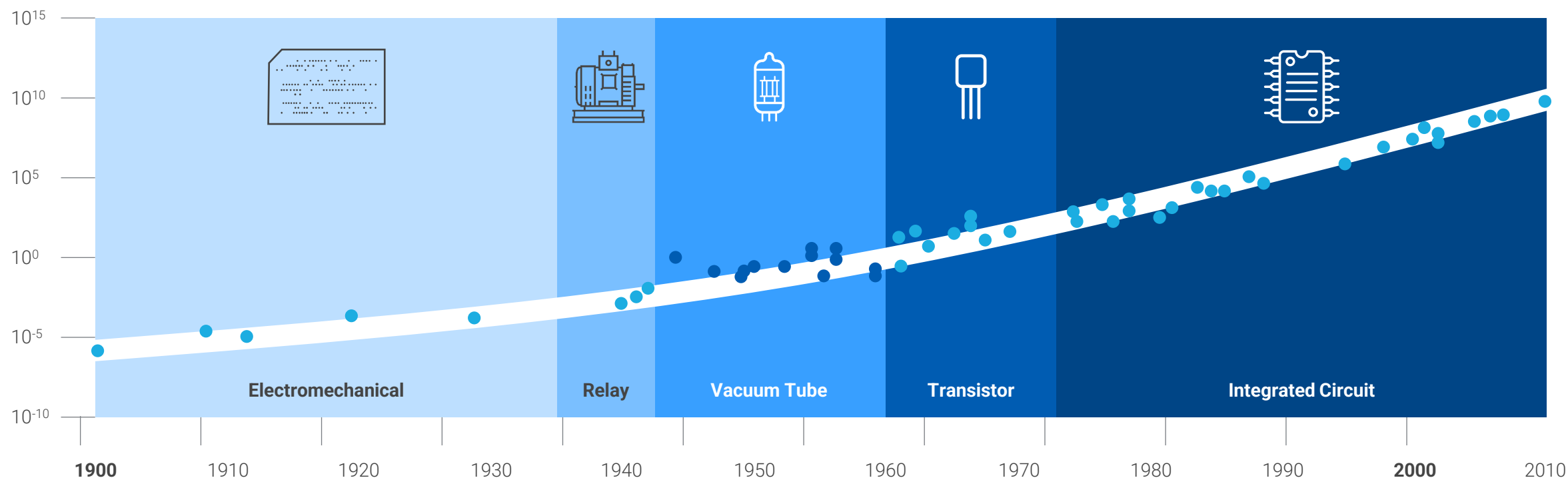
8:00 am	Welcome and opening	Helmy Eltoukhy (co-CEO) and AmirAli Talasaz (co-CEO)
8:15 am	Data generation driving future expansion	Darya Chudova (Chief Technology Officer)
8:45 am	Leading in liquid biopsy for therapy selection	Craig Eagle (Chief Medical Officer)
9:05 am	Paving the way for tissue-free MRD and surveillance	Helmy Eltoukhy (co-CEO)
9:25 am	Commercial Platform	Chris Freeman (Chief Commercial Officer)
9:50 am	Q&A	
10:10 am	Break	
10:25 am	Screening	AmirAli Talasaz (co-CEO)
11:00 am	Panel: Seeing deeper with epigenomics	Hosted by Justin Odegaard (VP, Product Development)
11:55 am	Financials	Mike Bell (Chief Financial Officer)
12:10 pm	Future / AI / What's ahead	Helmy Eltoukhy (co-CEO) and AmirAli Talasaz (co-CEO)
12:20 pm	Q&A	
12:45 pm	Lunch	

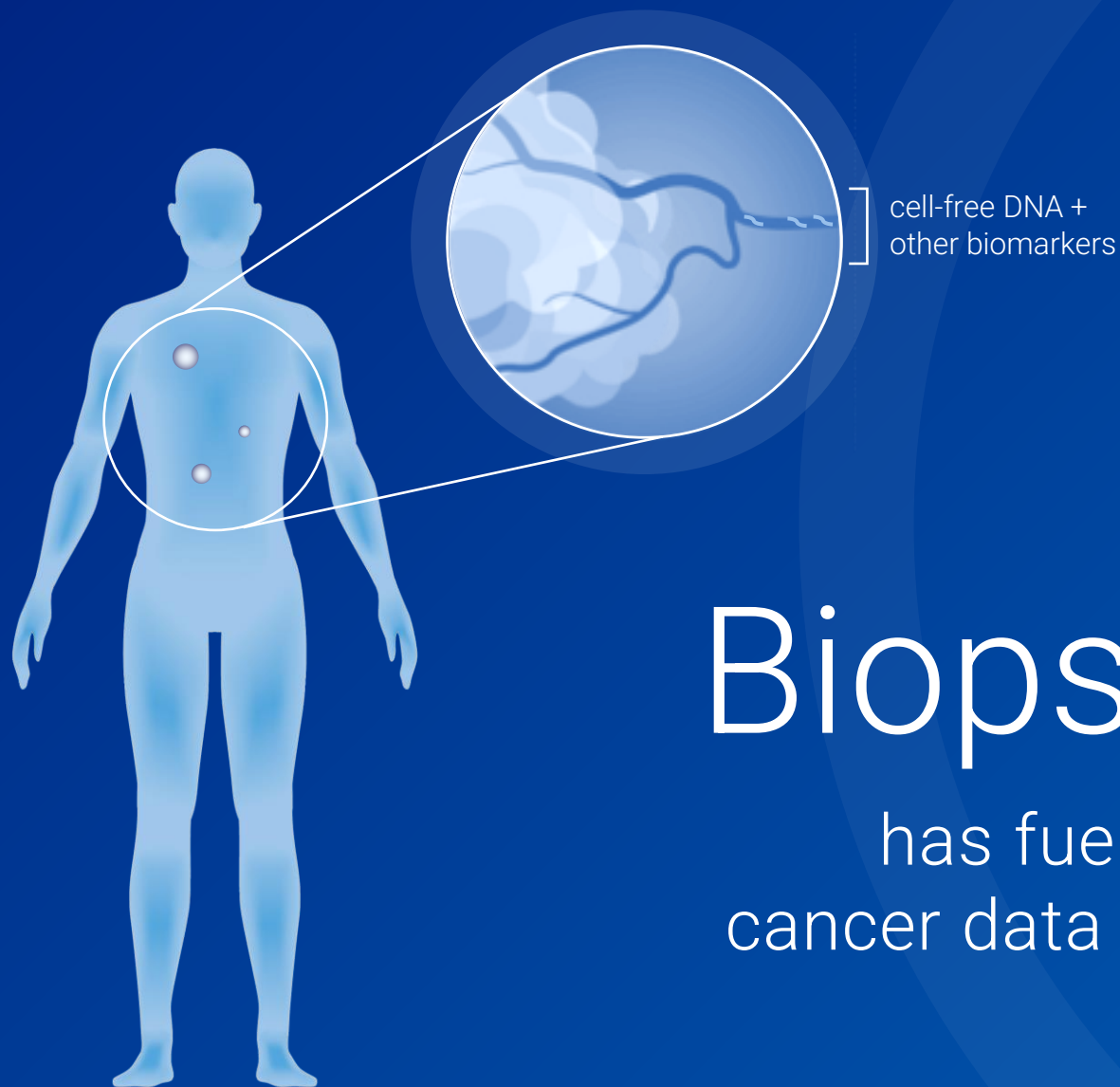


Guardant Health was
founded with the
mission to give us all
more time free from
cancer

On this mission
to conquer
cancer, our
weapon of
choice is **data**

Paradigm shifts have driven exponential increases in data generation and acquisition





The Liquid Biopsy Paradigm

has fueled **exponential increases** in cancer data and outcomes improvement

Paradigm shifts are not easy

Liquid biopsy...

“

...cannot detect late-stage cancer in blood.”

“

...is too expensive to run at scale.”

“

...is unproven and unreliable.”

“

...only detects uninformative cancer information in blood.”

“

...will only be used for niche applications.”

“

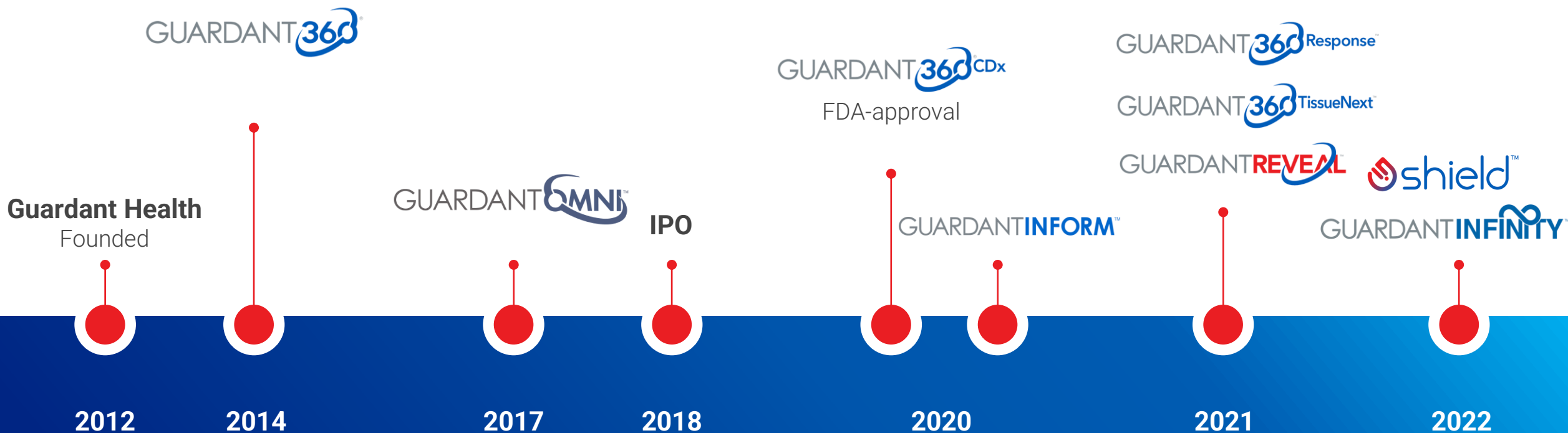
...cannot achieve high performance for MRD without tissue.”

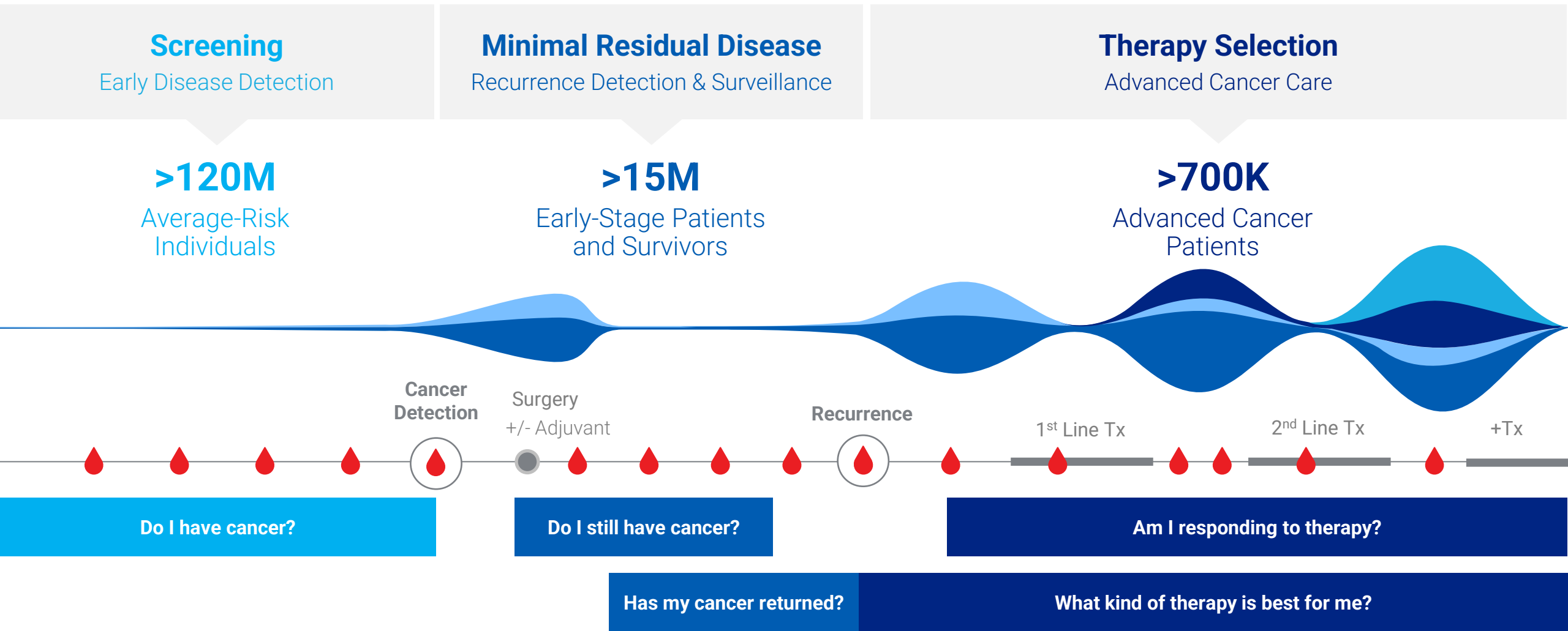
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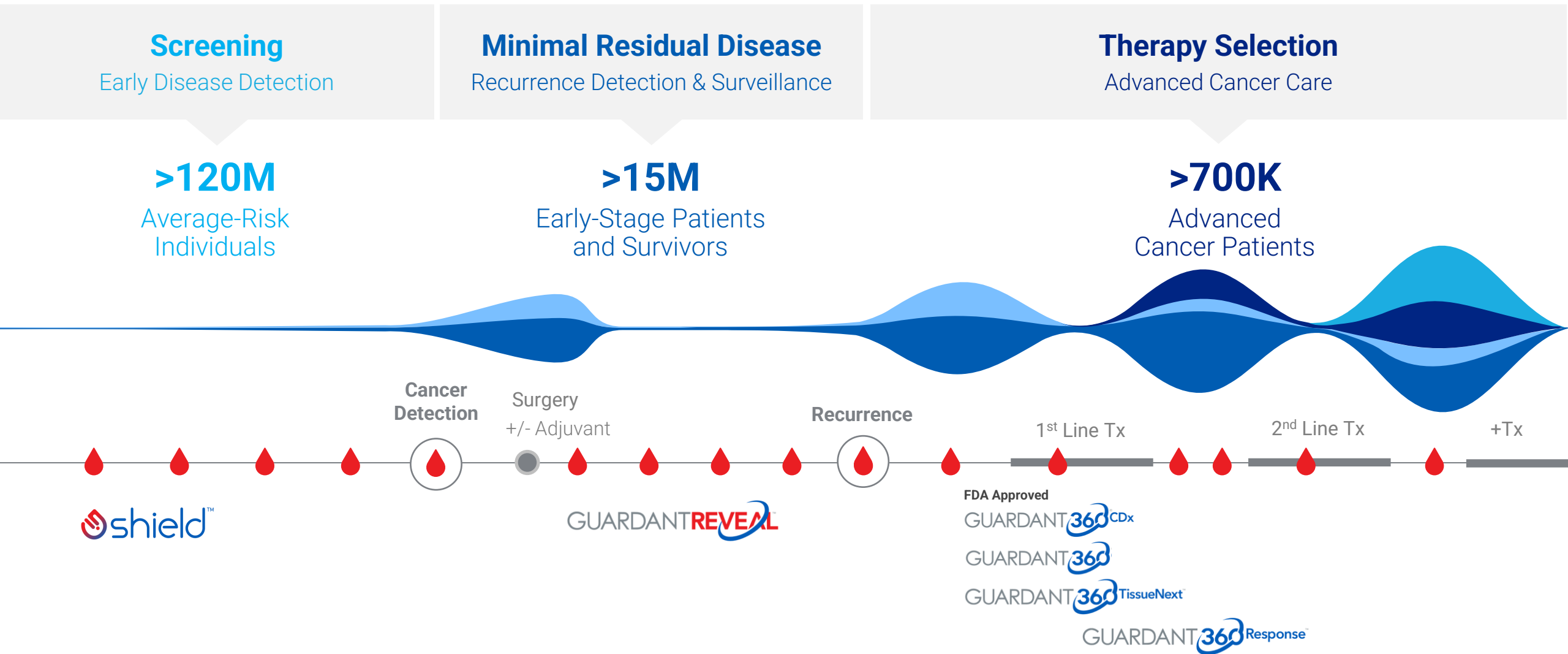
...cannot detect Stage I-III cancers with high sensitivity.”

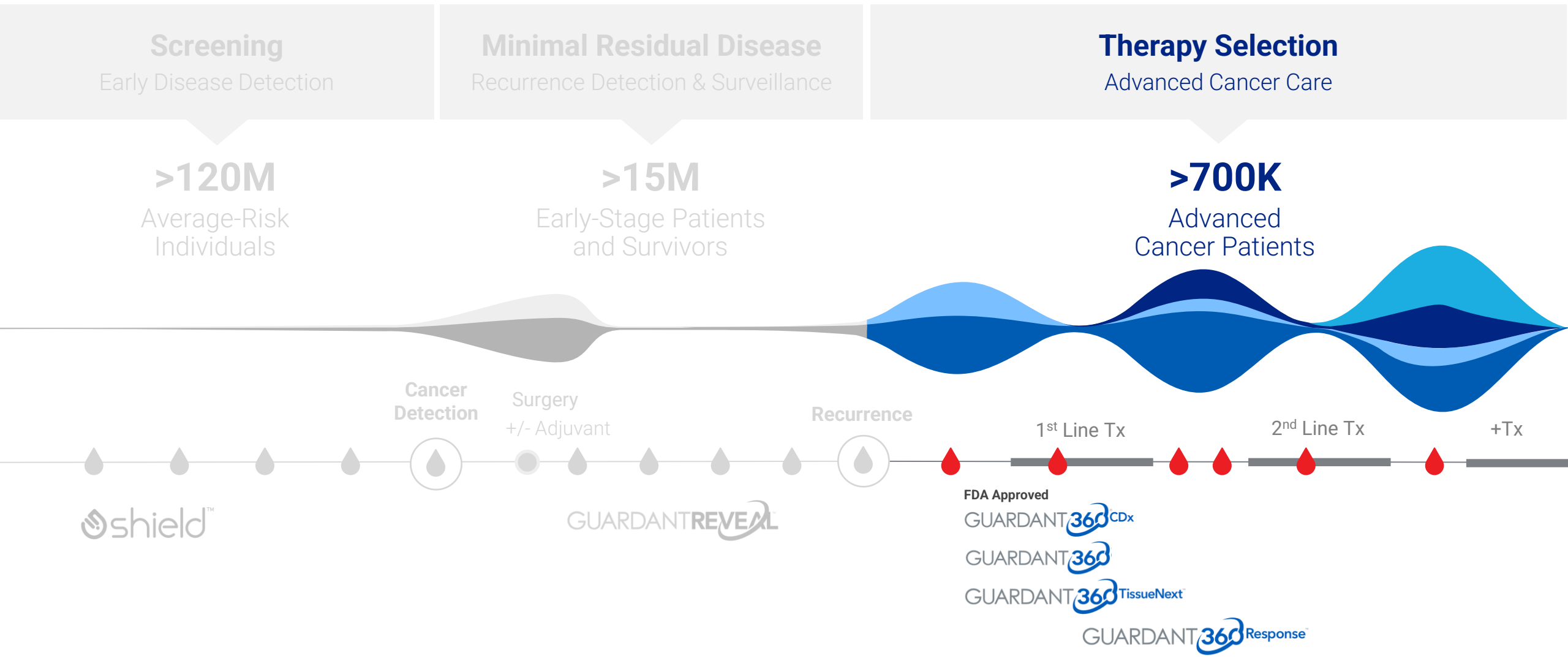
Product launches across the cancer care continuum

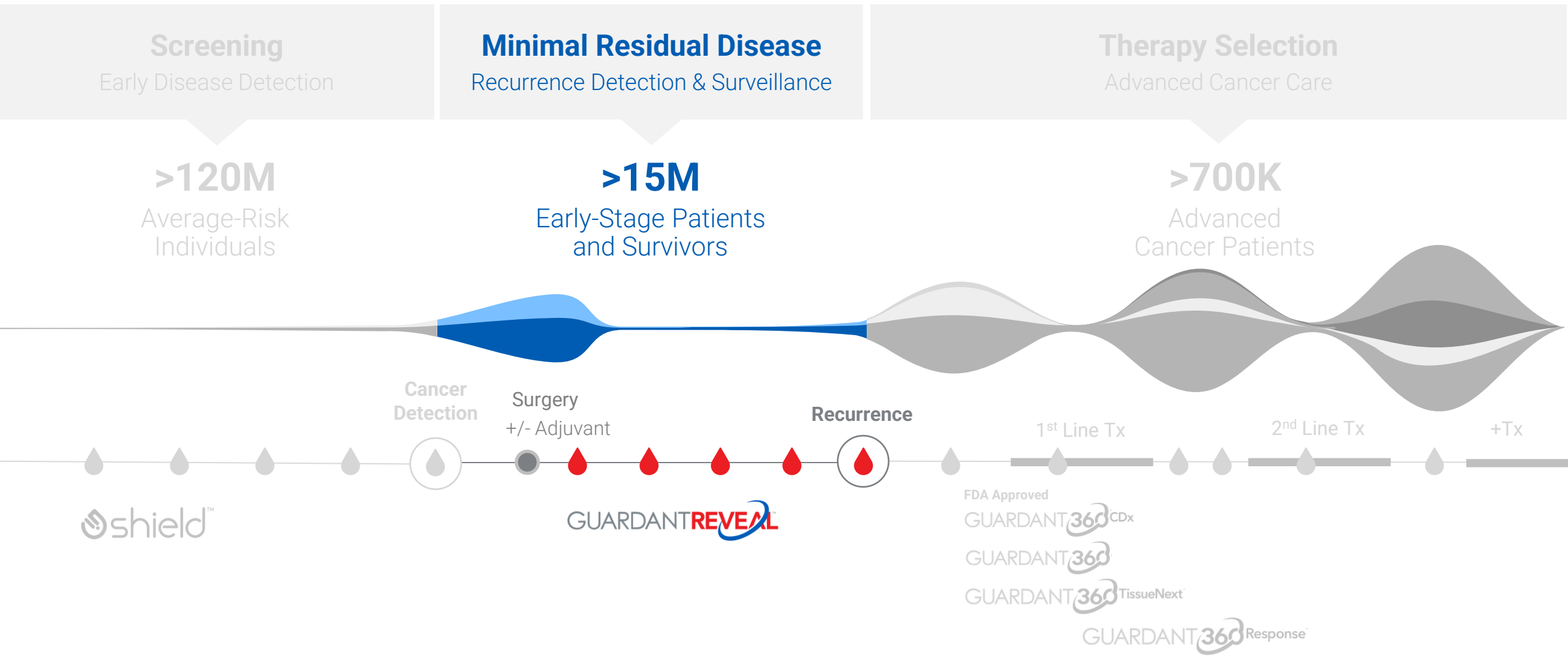
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23

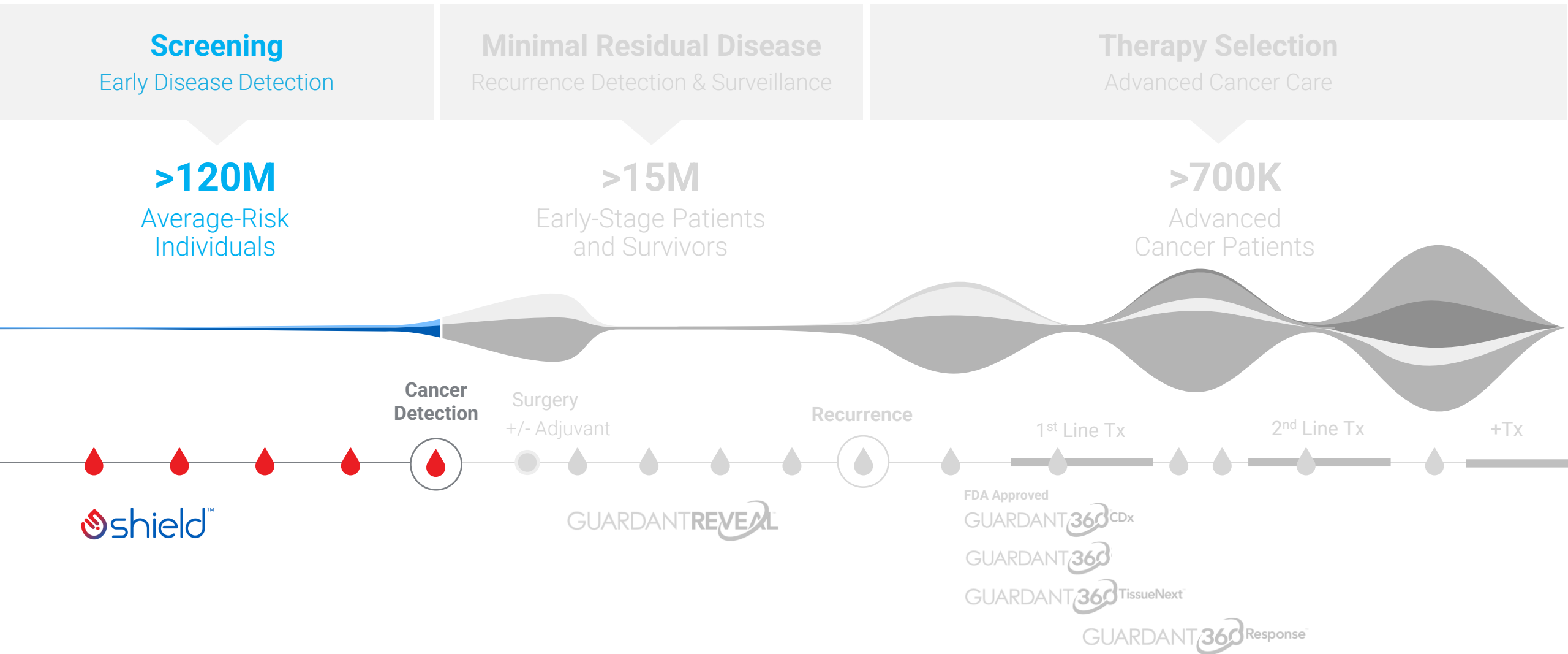












We have helped hundreds of thousands of patients and unlocked petabytes of clinical data



Cumulative
patient tests

>500K



Current ordering
oncologists

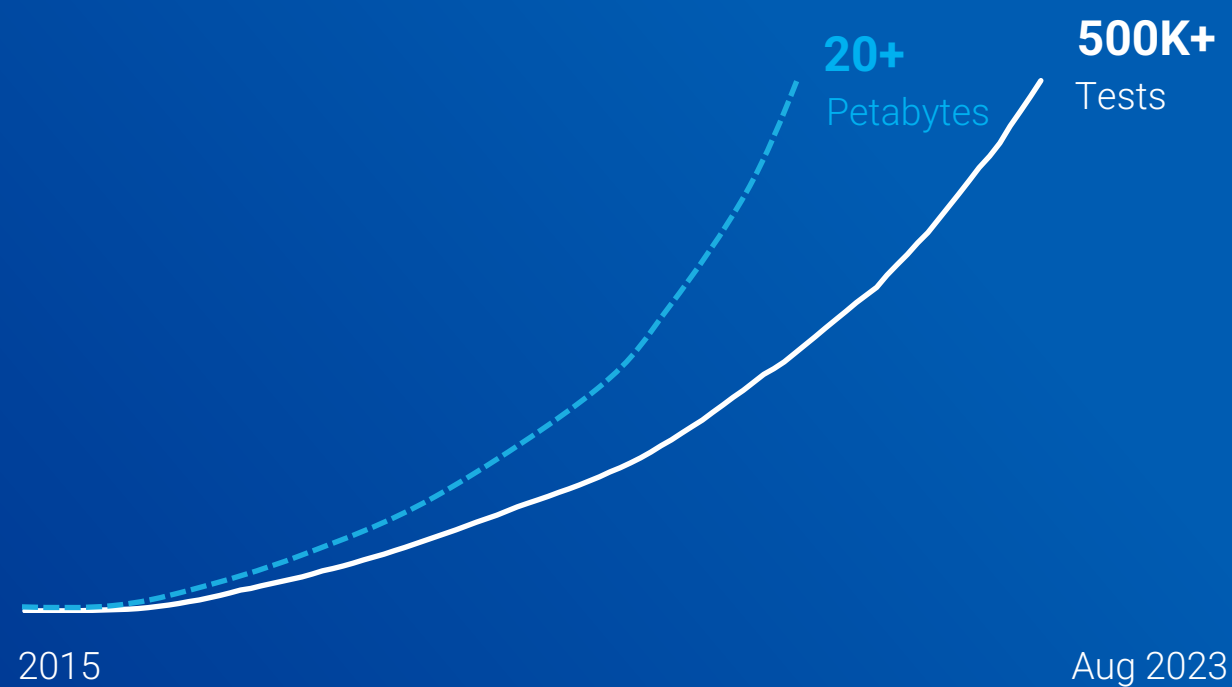
>12K



Bio-banked
samples

>500K

Exponential genomic data generation¹



Guardant today

\$548M

2023 Revenue
Guidance Midpoint¹

160+

Biopharma
Partners

60+

Countries

600+

Global Patents &
Patent Applications

Our values drive us forward



Put patients first

Our commitment to treat them as our own family



Be stronger together

Our commitment to teamwork & caring for each other



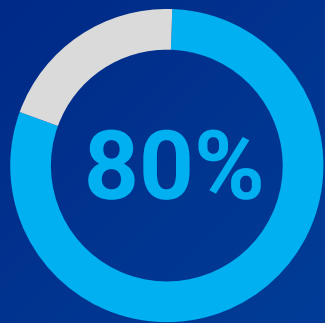
Blaze a trail

Our commitment to innovation



Make every moment matter

Our commitment to excellence, velocity and impact



Employee approval
rating

Great
Place
To
Work®
Certified
MAY 2023–MAY 2024
USA

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195+
PhDs & MDs

1,700+
Employees

56%
Female

60%
Ethnically diverse



As of December 31, 2022



This is just the
beginning...

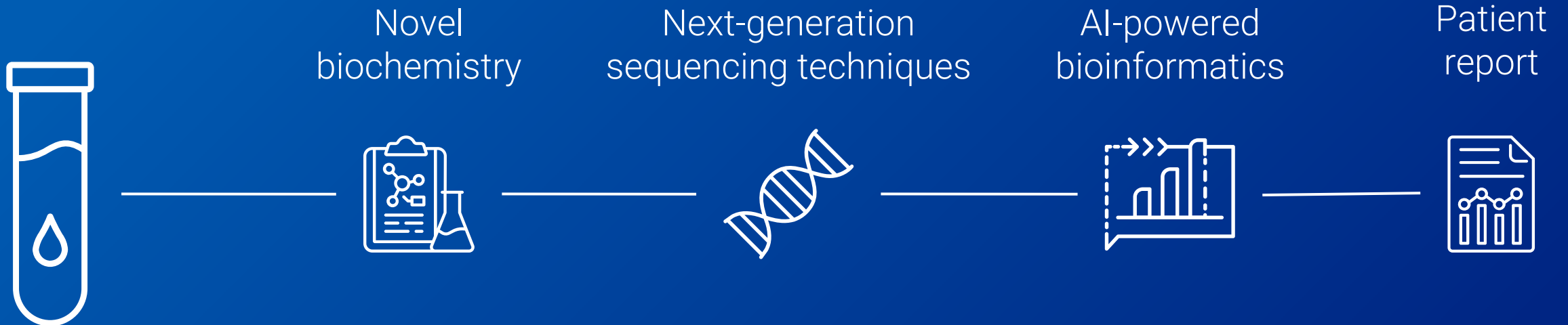


Data generation driving future expansion

Darya Chudova

Chief Technology Officer

Guardant Health pioneered liquid biopsy



Extending from advanced disease to early stage and screening required a **technology paradigm shift.**

Smart Liquid Biopsy platform



The power of liquid biopsy

- Simple blood draw
- Tissue-site agnostic
- Captures whole body heterogeneity
- Captures temporal dynamics



Enhanced by epigenomics

- Deeper tumor sensing
- Wider biological view



Delivering performance & scale

- Proprietary biochemistry with high sensitivity / specificity
- Efficient COGS
- Powered by data generation at scale

Delivers **100X** genomic breadth and **>50X** sensitivity with **no additional cost**



There are **37.2 trillion cells**
in the human body and they
all share **one genome**



Epigenomics

is the missing link in
understanding human disease

Methylation is a key determinant of the epigenomic state

Epigenomic state controls how our cells function

Single Genome



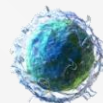
Variable methylation state



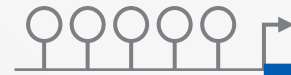
100s of cell types

Gene

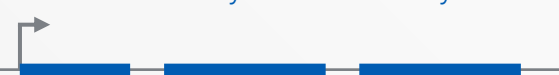
Blood cell



Unmethylated



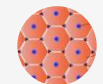
Gene A may be functionally active



ON



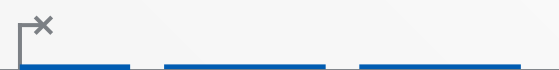
Liver cell



Methylated



Gene A is silenced

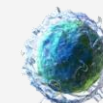


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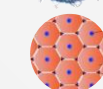


Genome

Blood cell



Liver cell



Liver cancer cell



10,000s of epigenomic switches

0	0	1	0	...	1	0	1	0	1
0	1	1	0	...	0	0	1	0	1
0	1	1	0	...	0	0	1	1	1

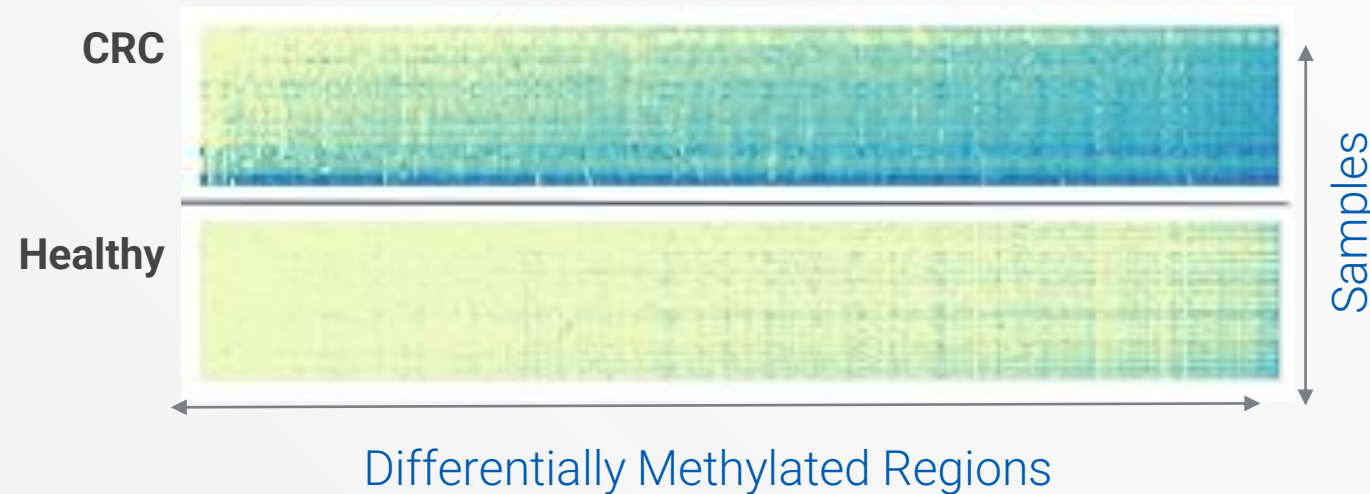
Unique fingerprint of disease

Epigenomic state change is wide throughout the genome

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1,000s

Of **differentially methylated regions** form cancer or cell type specific fingerprints



Number of detectable tumor alterations drives sensitivity

Genomic analysis



Tissue Informed Assays

Up to 50
Tumor Alterations

Epigenomic analysis

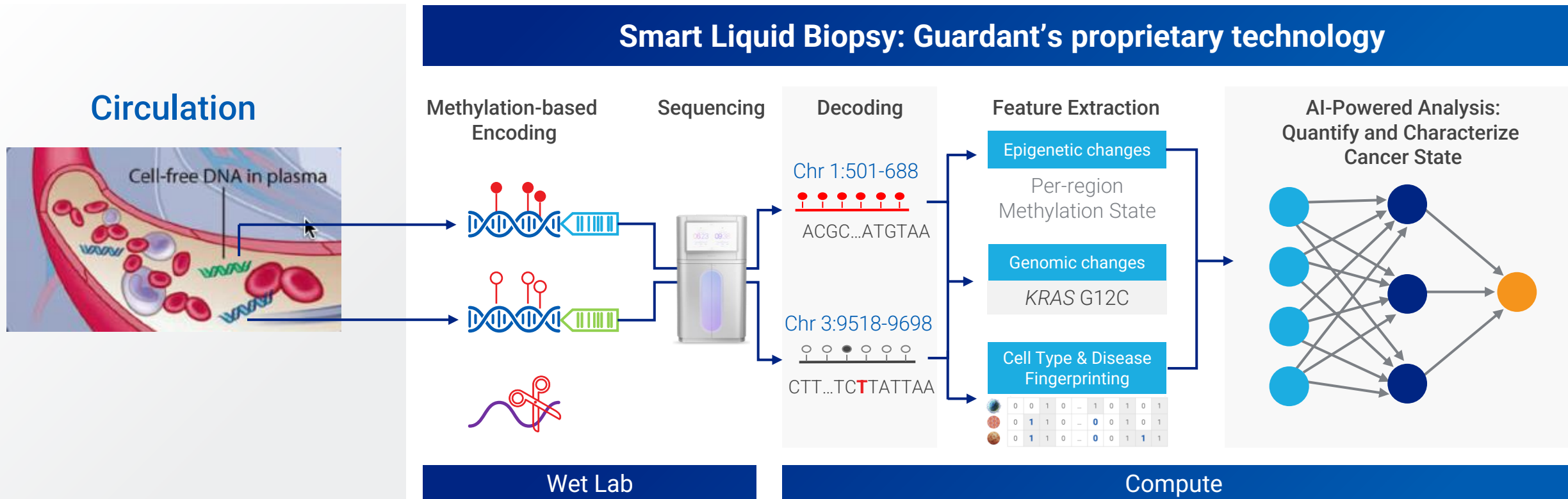


Smart Liquid Biopsy

>1,000 Tumor Alterations

Smart Liquid Biopsy unlocks both genomic and epigenomic state

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Smart Liquid Biopsy: key technology differentiators

Single sample workflow for efficiency

Genomic mutations + Epigenomic profiles

Genome wide profiling

>20X more detectable alterations¹

Highly efficient molecule capture for optimal detection

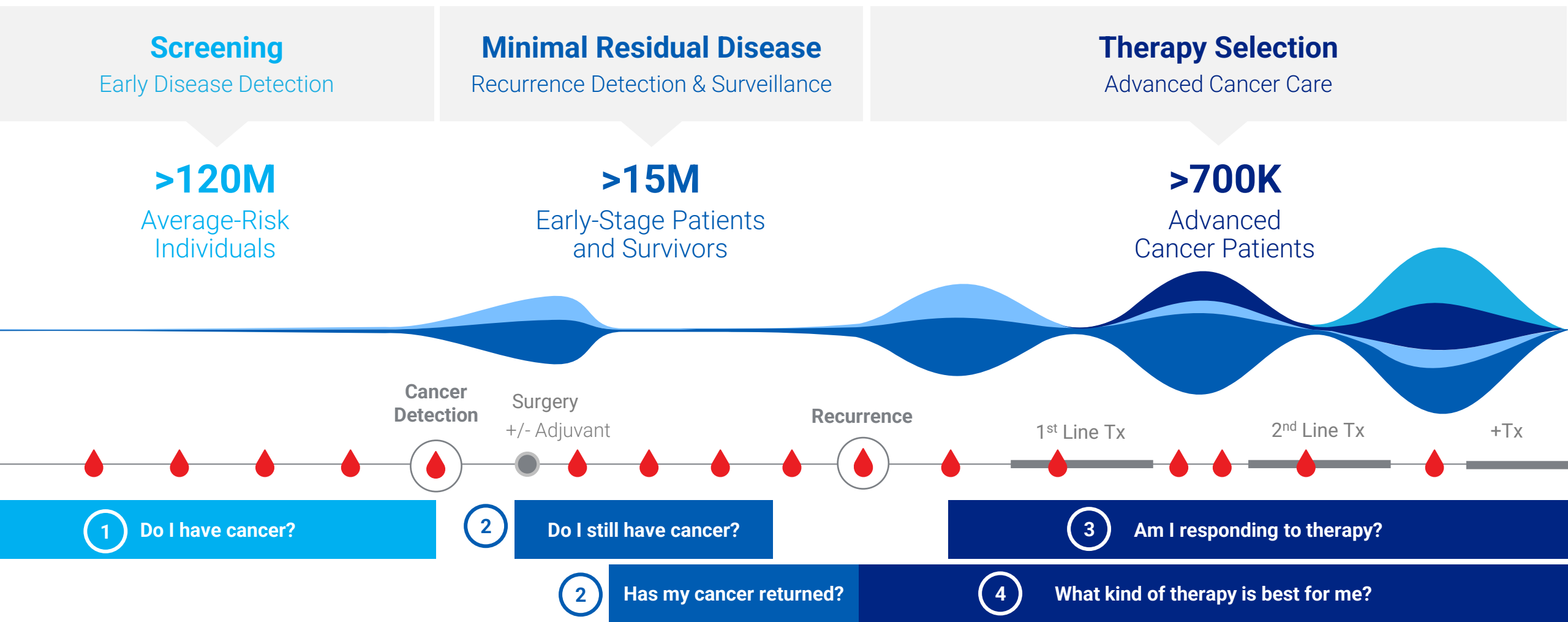
3X more signal molecules captured²

Background depletion for optimal cost

99% depletion for **7X** sequencing cost savings²

Continuous improvement powered by samples & data bank

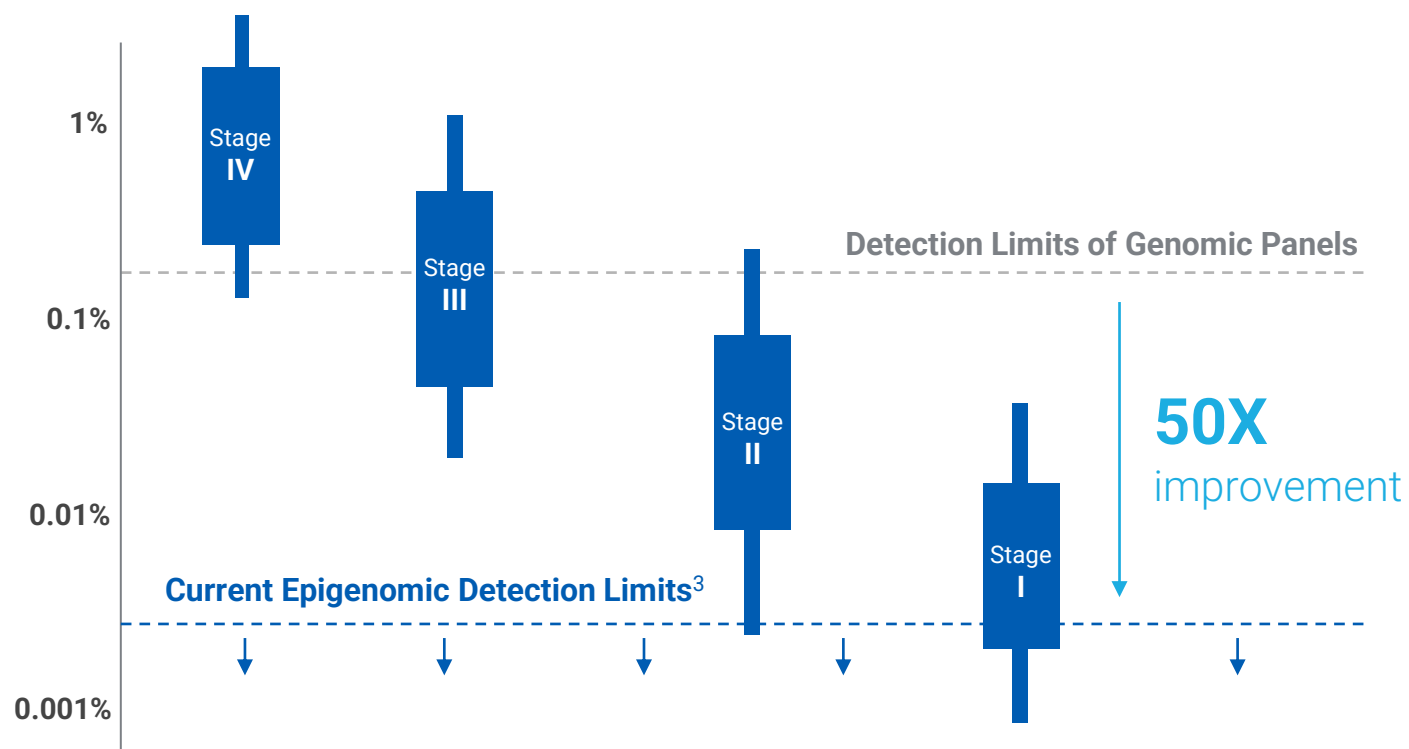
500K samples, multiple cycles of **2X** performance gains³



1

Do I have cancer?

**Clinically relevant CRC
detection capability** achieved
by epigenomics assay for
screening applications^{1,2}






2

Do I still have cancer?
Has my cancer returned?

Clinically relevant **Minimal Residual Disease (MRD)**

detection capability for tissue-free assay achieved by epigenomics **at high specificity (>98%)**



Cancer type	Tumor Fraction Level for Reliable Detection ¹ at the most challenging input level of 5ng
 CRC	0.01%
 Lung	0.01%
 Breast	0.015%

Typical cfDNA amounts in early stage clinically relevant samples²

Above 5ng
≥95% samples

Above 30ng
≤20% samples

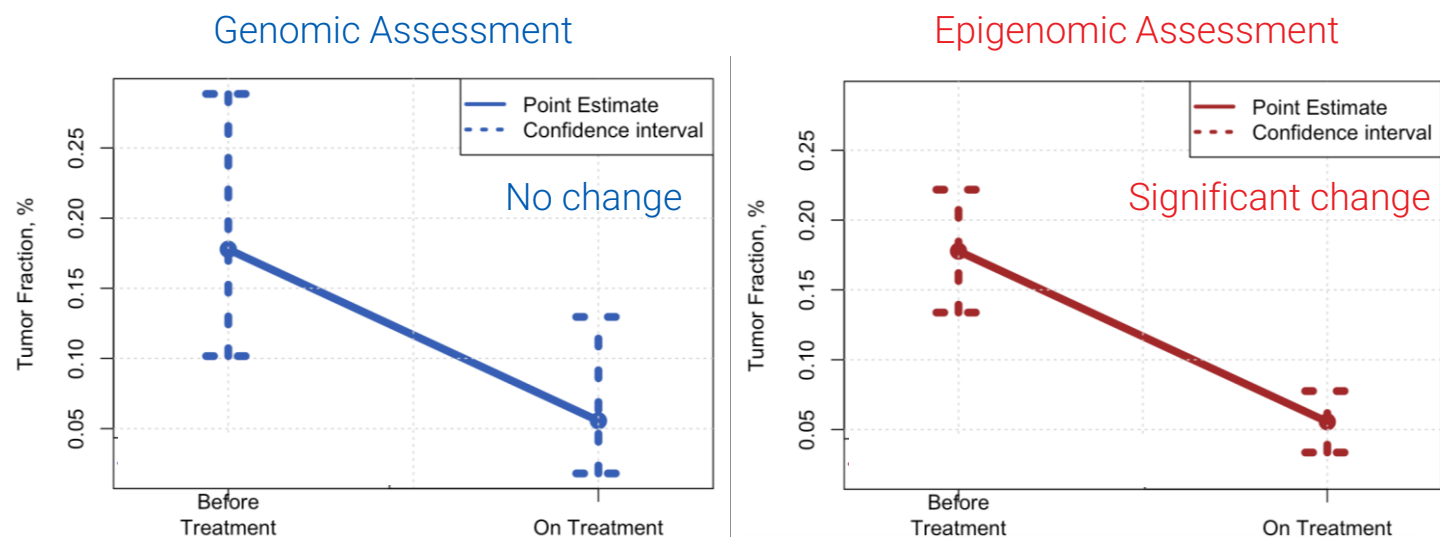
Above 60ng
≤ 5% samples

3

Am I responding to therapy? Should I switch therapy?

High precision of tumor fraction estimates is attained in epigenomic assay based on 1,000s of alterations assessed

3X higher precision¹ of epigenomics enables to see therapy response clearer and make better patient management decisions



Response in early phase trials

Surrogate Endpoint Development

Switching therapy based on lack of response

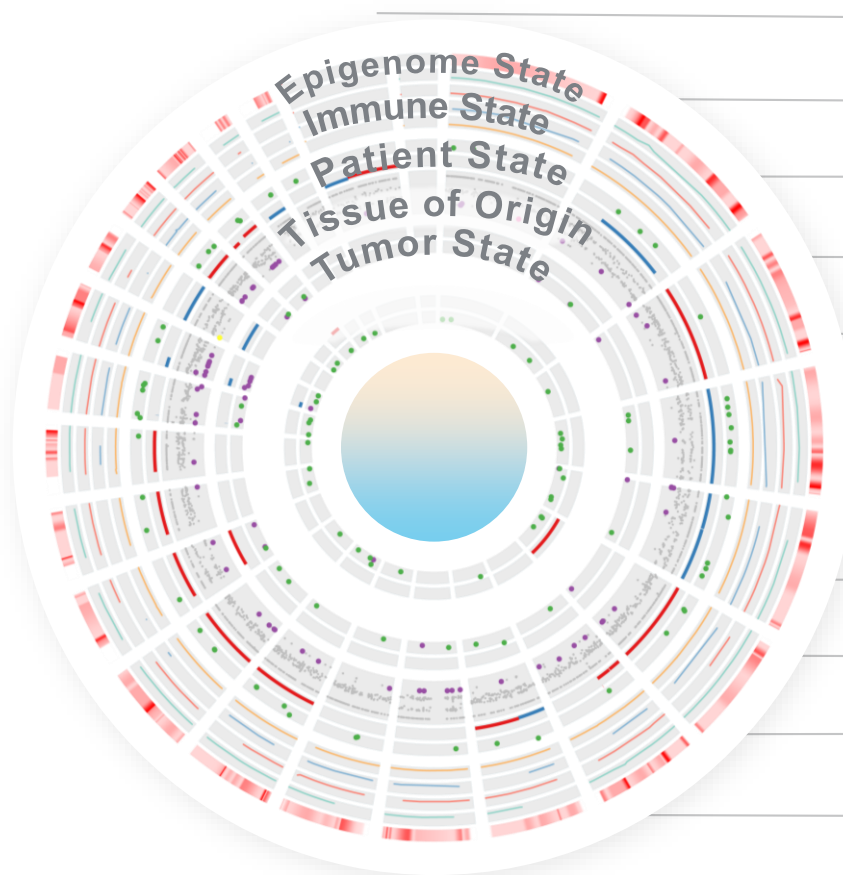
4

What kind of therapy is best for me?

Guardant360 powered by our Smart Liquid Biopsy platform will offer the most **comprehensive liquid biopsy** for characterization of cancer

Current best-in-class liquid biopsy for therapy selection and translational research

Expanded genomic and epigenomic capability offered with Smart Liquid Biopsy



SNVs/Indels

Amplifications

Copy Number Loss

Fusions

Large Rearrangements

bTMB

MSI

HRD

Virus Detection

Regulatory Epigenome

Immune State

4

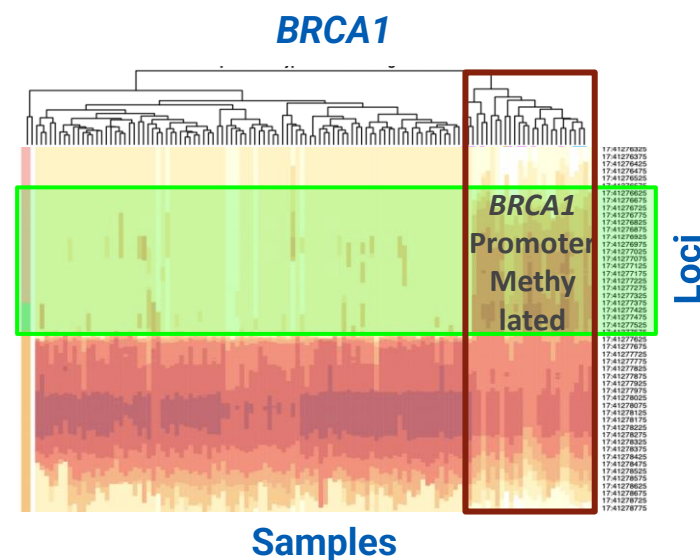
What kind of therapy
is best for me?



Epigenomics unlocks
**mechanisms of regulation that
are invisible** to genomic testing

Breast Cancer

Expanded patient eligibility for targeted therapy through promoter methylation



**~24% of TNBC patients are
associated with promoter
methylation¹**

Without analyzing both
genomic and epigenomic
alterations, patients potentially
eligible for PARPi therapies
would be missed²

4

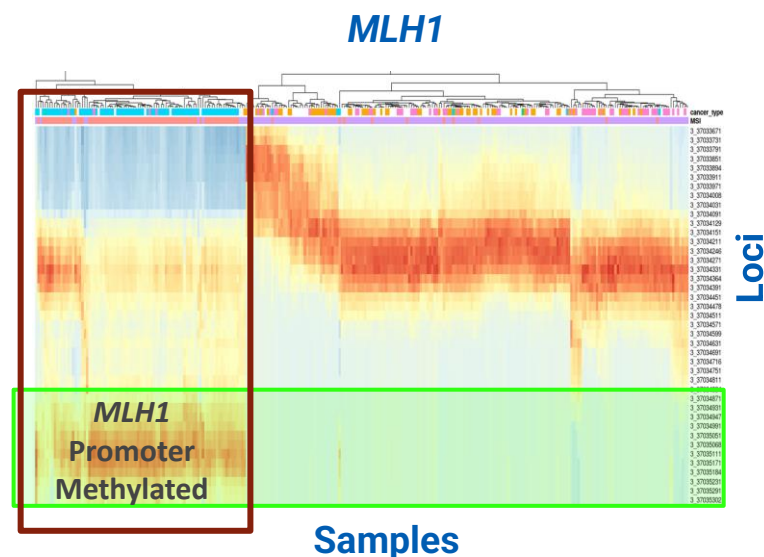
What kind of therapy
is best for me?



Epigenomics unlocks
**mechanisms of regulation that
are invisible** to genomic testing

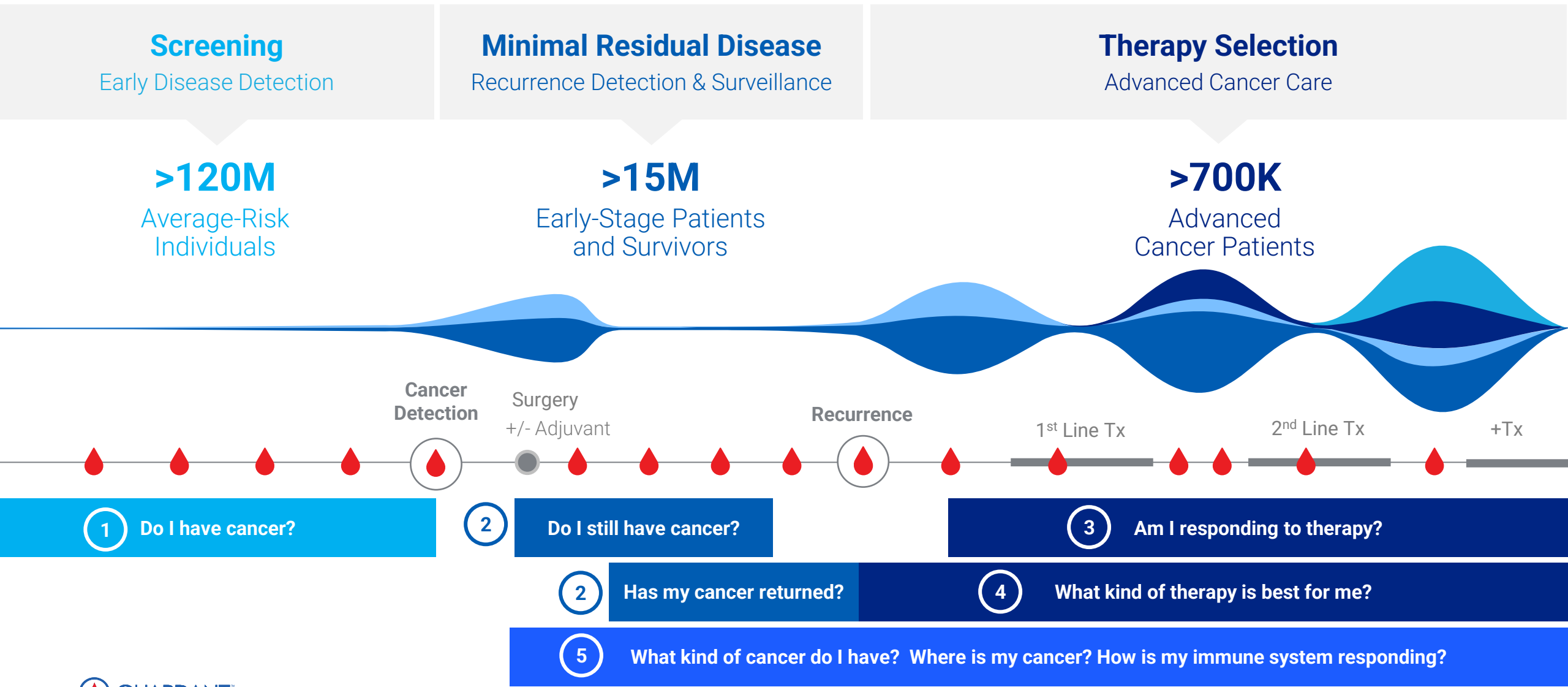
Colorectal Cancer

Expanded patient eligibility for targeted therapy through promoter methylation



MLH1 promoter methylation identified additional **7% of MSI-H patients** missed by genomic testing alone

These patients would be eligible for FDA-approved I/O therapy options.¹



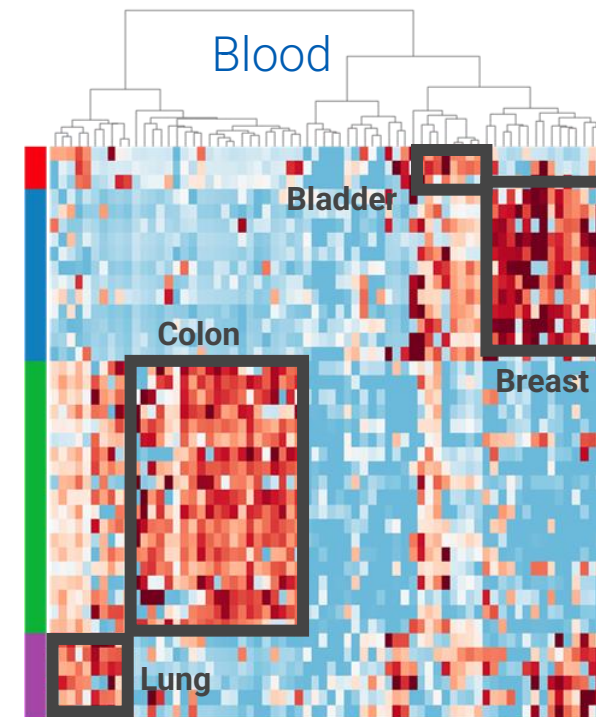
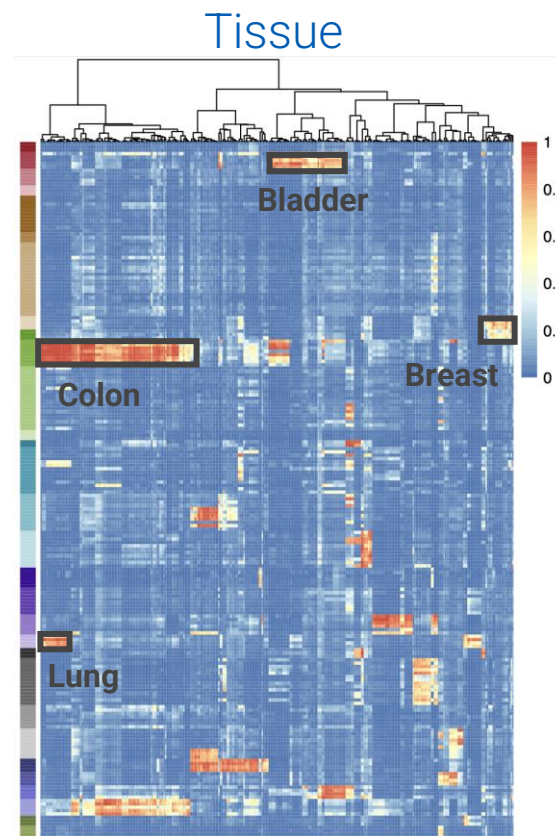
5

Where is my
cancer?



Tissue-specific fingerprints
are detectable with Smart
Liquid Biopsy

cfDNA methylation encodes tissue of origin



Data from Loyfer et al., *Nature* 613 (2023)

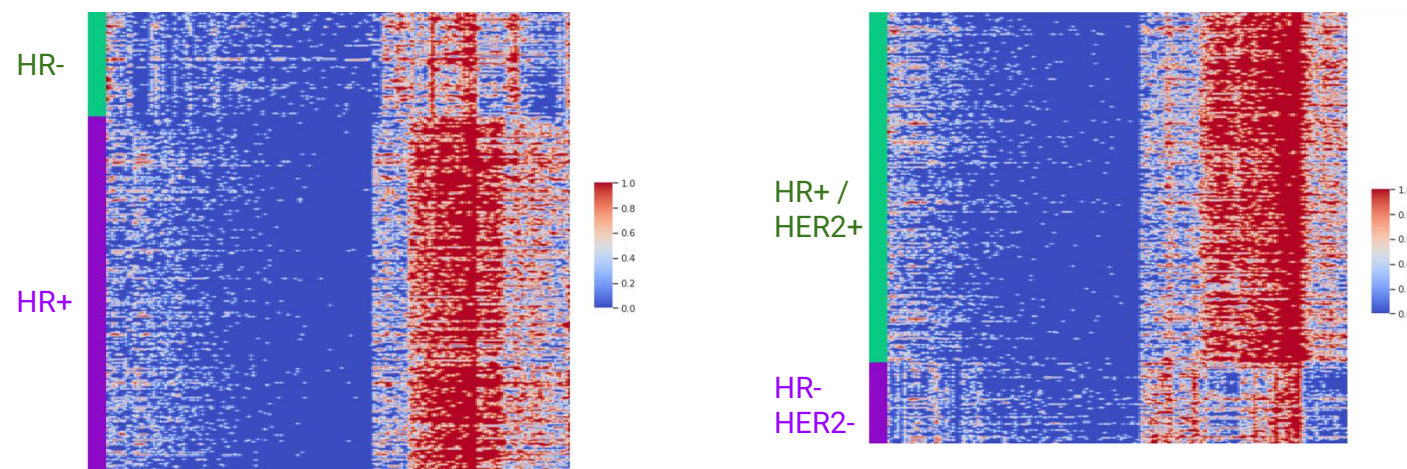
5

What kind of
cancer do I have?



Fingerprints differentiating
breast cancer subtypes are
detectable with Smart
Liquid Biopsy

cfDNA methylation encodes cancer subtype



Epigenomic profiles derived from plasma of breast cancer patients may enable precision diagnostics and potential for identifying novel response predictors

Future unified platform will drive significant R&D efficiency and operating leverage

Smart Liquid Biopsy



R&D efficiency

- Leveraged performance improvements
- Cross-development of new applications

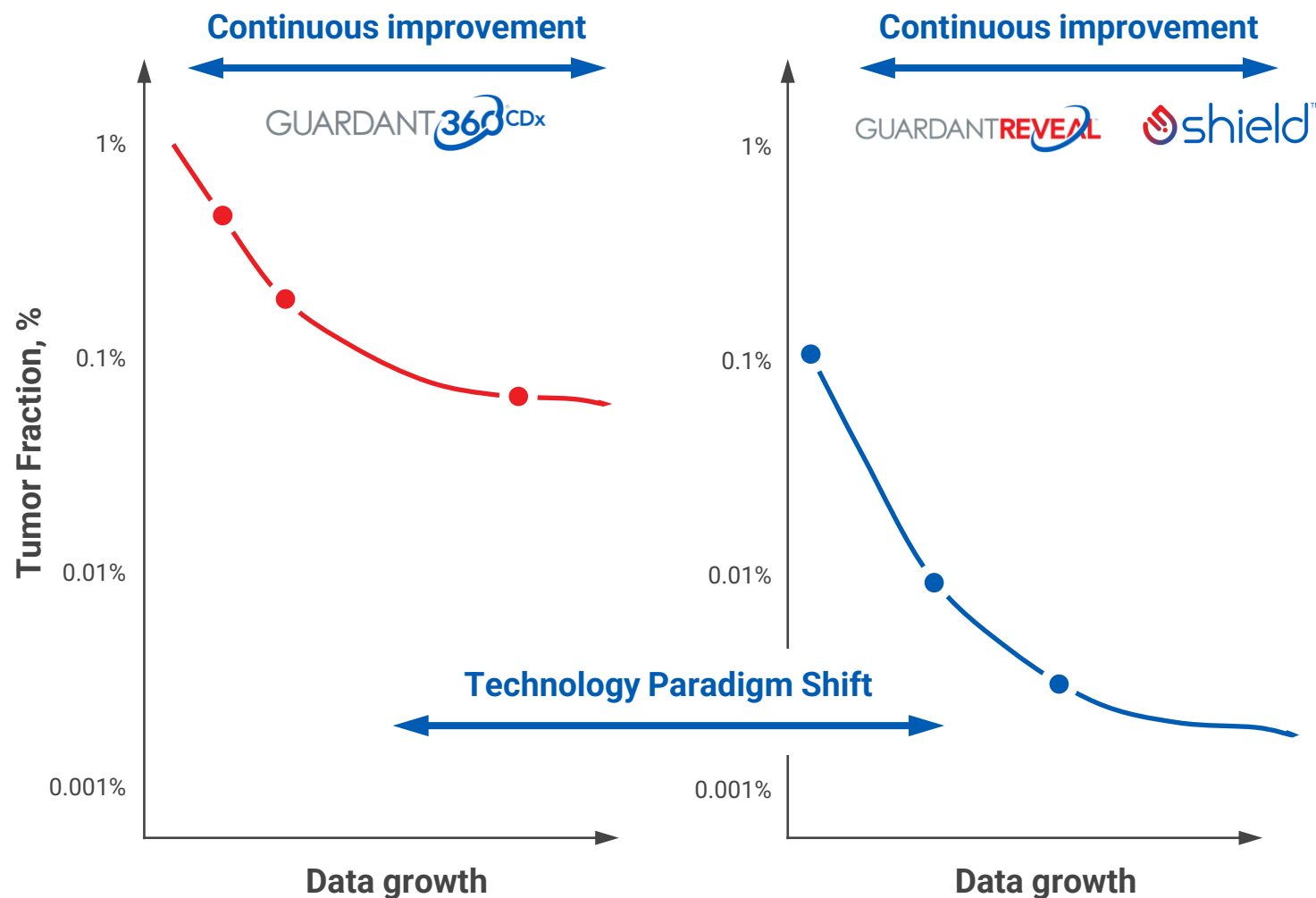
Operating leverage

- Lower COGS
- Fast turnaround time

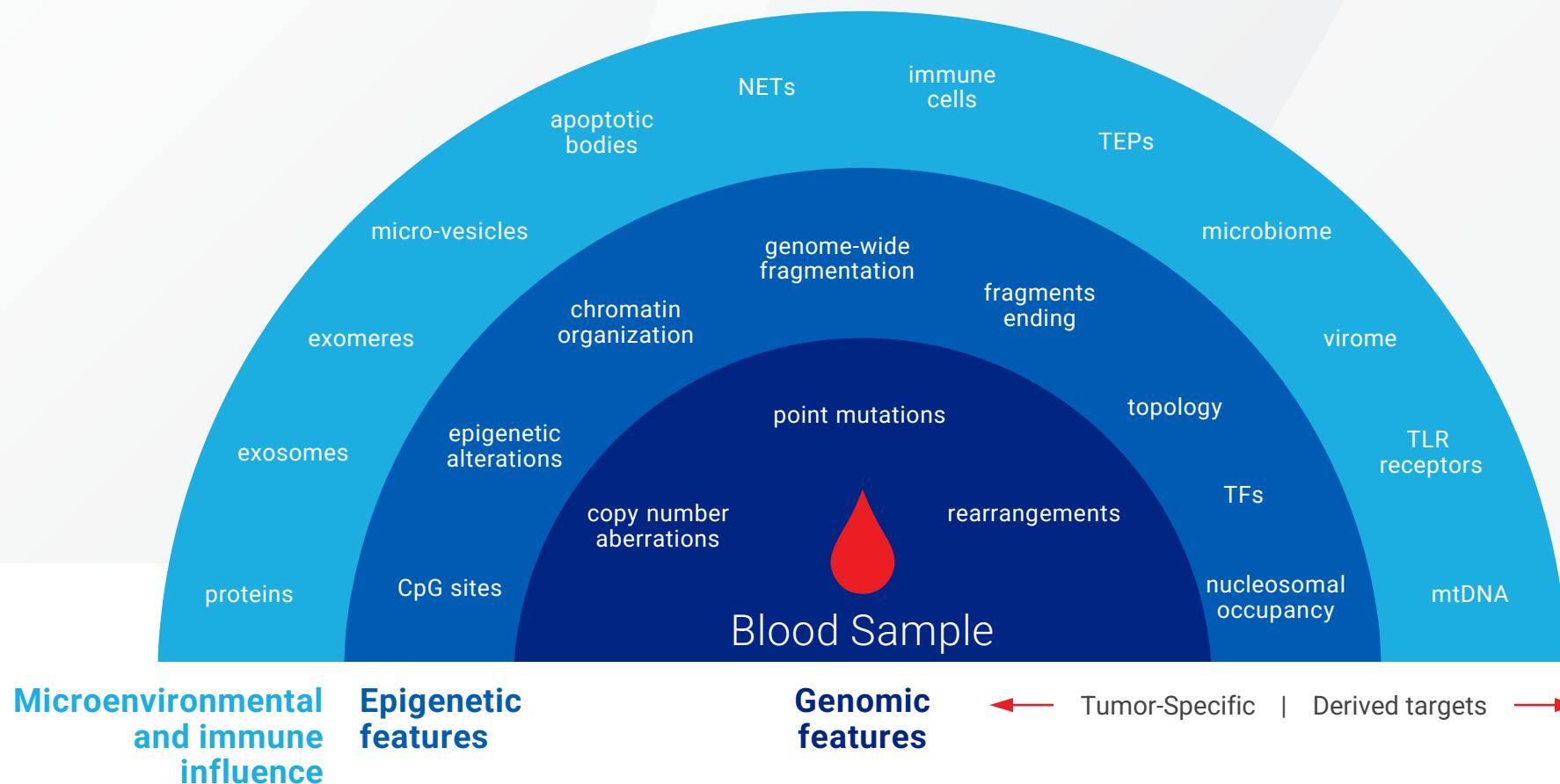
Power of More Data

Learning from data
for continuous
improvement

**Technology paradigm
shift** for radical
change in capability



Smart Liquid Biopsy will continue to unlock the rich landscape of blood-based markers

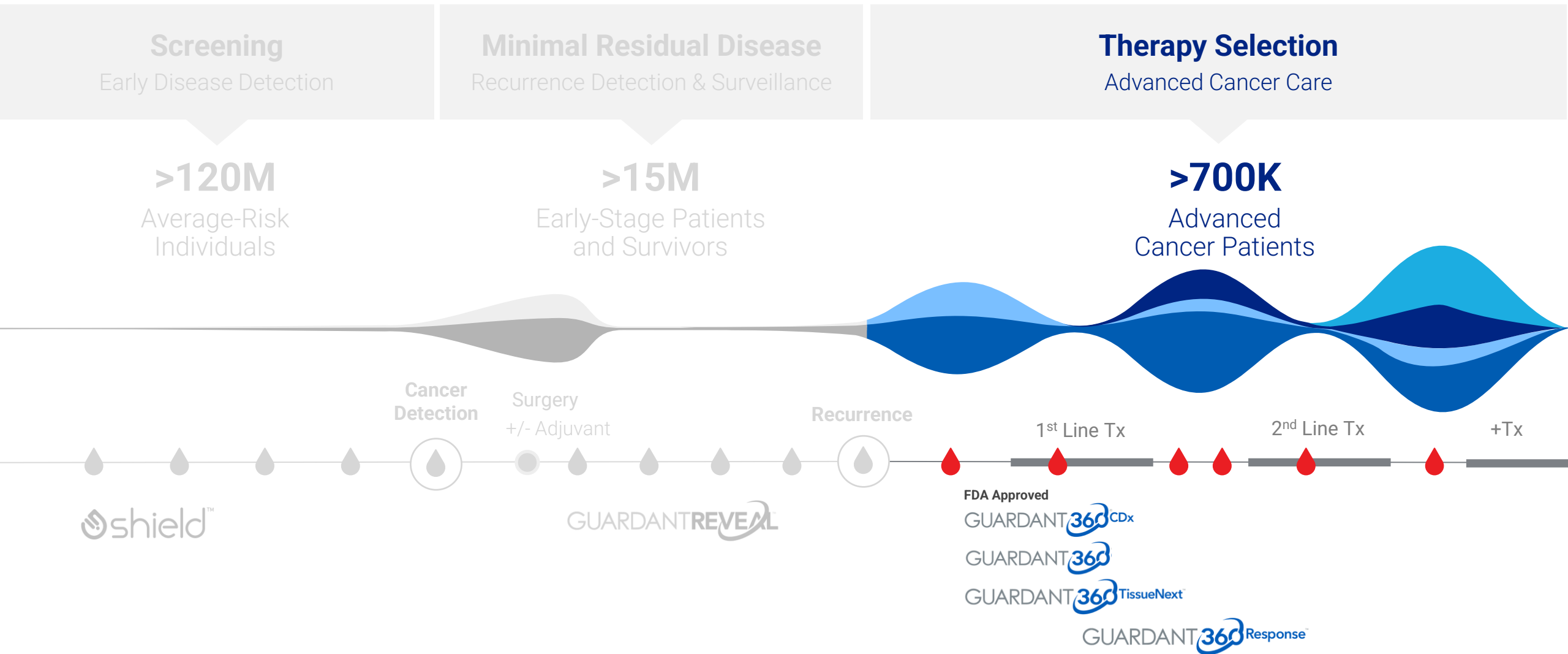




Leading in liquid biopsy for **Therapy Selection**

Craig Eagle, MD

Chief Medical Officer



Comprehensive portfolio for Therapy Selection with broad payer coverage

GUARDANT 360^{CDx}

1st FDA-Approved
Comprehensive
Liquid Biopsy



GUARDANT 360[®]

Next-Generation
Liquid Biopsy
Assay



GUARDANT 360^{Response}

1st Blood-Only
Liquid Biopsy
to Monitor
Tx Response



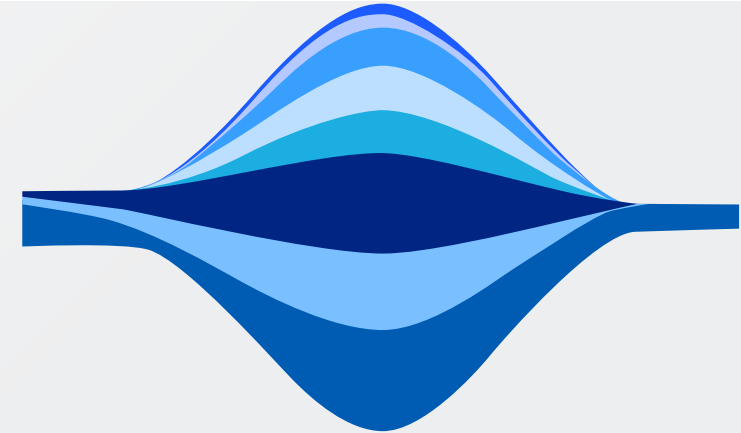
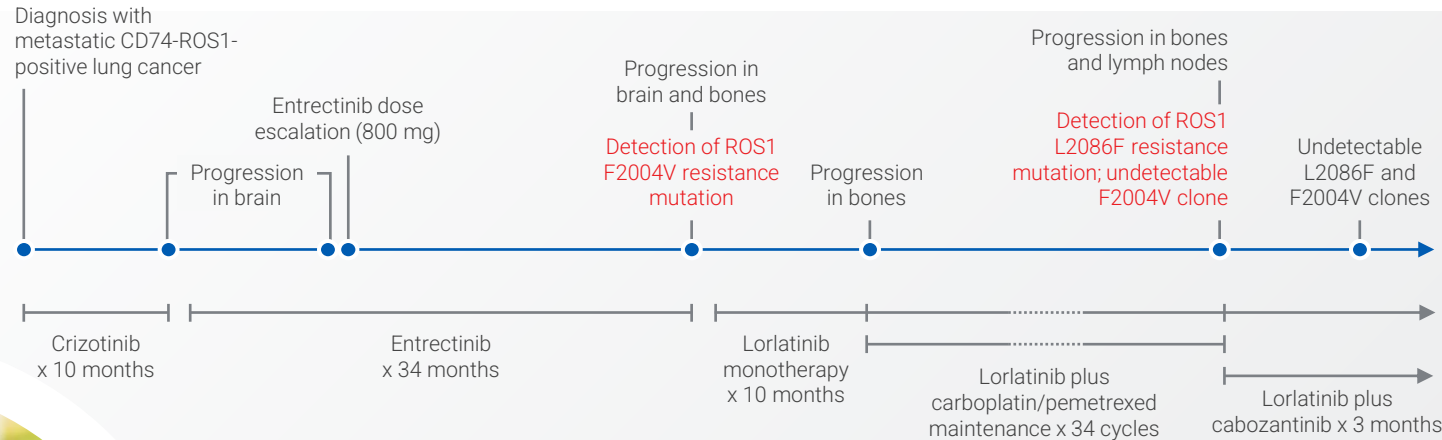
GUARDANT 360^{TissueNext}

Next-Generation
Tissue Assay



Reimbursed by Medicare

Guardant360 in lung cancer



- Diagnosed with NSCLC, treated with ROS1-inhibitors
- Guardant360 identifies acquired, targetable alterations
- Guardant360 used throughout continuum of care to assess ctDNA at time of clinical progression
- From Guardant360 results, patient treated with combination therapy
- Assessment of ctDNA 6 weeks after treatment initiation showed likely molecular response

425+ publications supporting use of Guardant360

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Partnership with Top Thought Leaders

45

NCI-Designated Comprehensive Cancer Centers involved in publication generation

Reimbursement

30

Publications used to obtain Medicare coverage

Patient Outcomes

120+

Publications with patient outcomes involving targeted therapies

Changing Guidelines

38

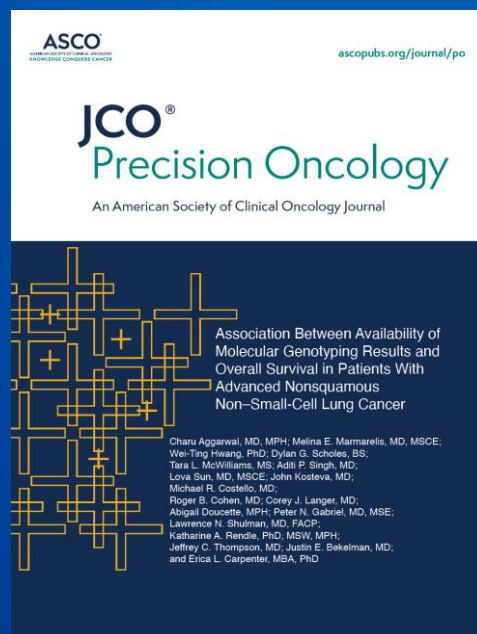
Publications included in successful NCCN submissions



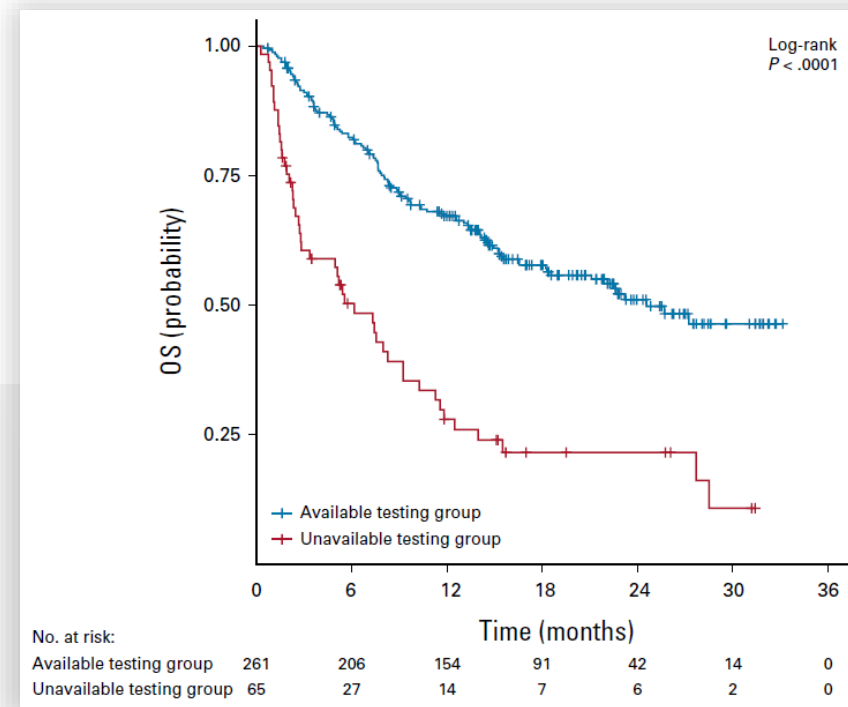
Because precision and speed impact patients

CGP results in
4x better patient
outcomes

Every month delayed
in cancer treatment
can raise risk of death
by around 10%¹



Association Between Availability of Molecular Genotyping Results and Overall Survival in Patients with Advanced Nonsquamous Non-Small-Cell Lung Cancer²



Overcomes limitations of existing tools used to measure treatment response



Accelerated insight into treatment response

Measure ctDNA change and drug efficacy earlier than standard imaging

Accelerates clinical trials and drug development



Advantages of a tissue-free approach

Overcome logistical challenges of tissue-informed approaches, such as tissue insufficiency, increased time due to procurement, and invasive procedures



Improved sensitivity and precision with methylation

Guardant360 Response on the Smart Liquid Biopsy platform will provide even deeper tumor sensing and higher precision



Real-time clinical decision making

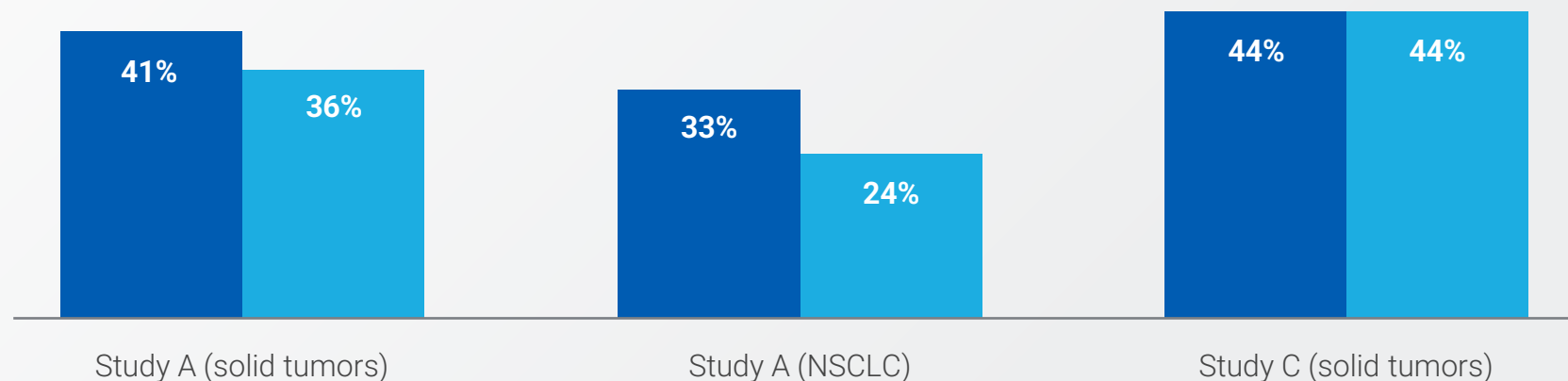
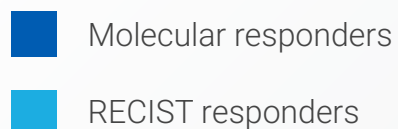
Rapidly assess drug efficacy, enabling fast-fail drug development model

Consider alternate treatments or dosing for non-responders

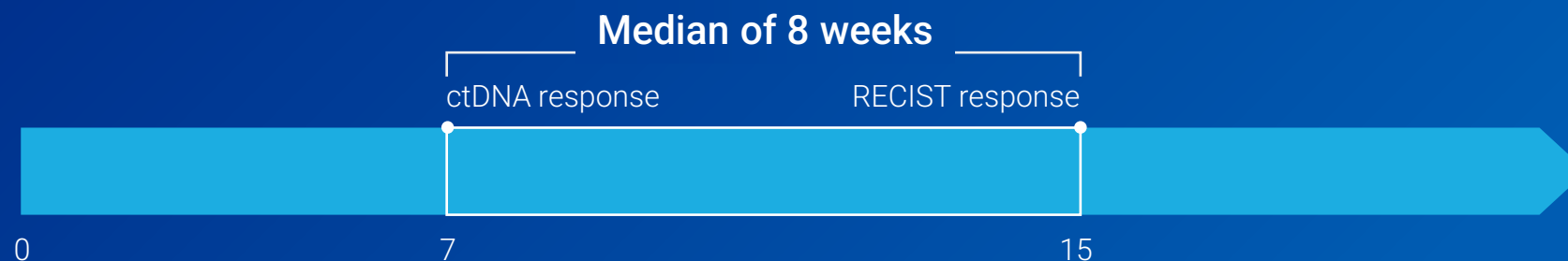
Guardant360 Response can determine which patients are responding to therapies sooner than CT-Scans

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ctDNA and RECIST
identify responders...

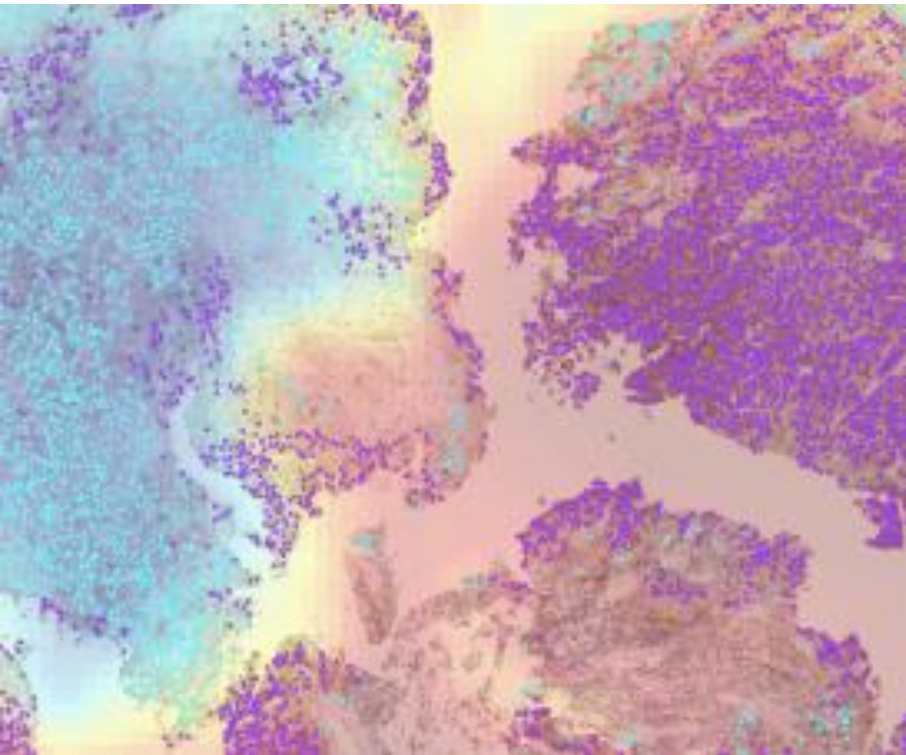


...but ctDNA identifies
them 8 weeks earlier



Advancing Guardant360 TissueNext with AI-powered digital pathology

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DAY | 20
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GUARDANT **360** TissueNext™

GUARDANT
GALAXY™ PD-L1



Only tissue CGP panel with an AI-powered scoring algorithm that **improves PD-L1 detection by >20%** in NSCLC¹



Now **validated** for multiple cancers²



Covered by Medicare for all advanced solid tumors³

FDA-approved targeted therapy options are continuing to rapidly expand across solid tumors



Spotlight on ESR1

Emerging
mutation with
high unmet need
fueling
Guardant360
CDx growth

~67%–80% of breast cancers in women are ER+, HER2-¹



Breast cancer is the
second leading cause of
cancer death in women



~298k new cases and
~44k deaths estimated
in 2023 alone



ER+ tumors have a high
likelihood of developing
ESR1 mutation

ESR1 mutations are present in up to 40% of ER+, HER2- advanced breast cancers²



ESR1 is an emergent
mutation that develops
after Breast Cancer
treatment occurs



Patients expressing
ESR1 can be put on
a new class of
targeted therapy

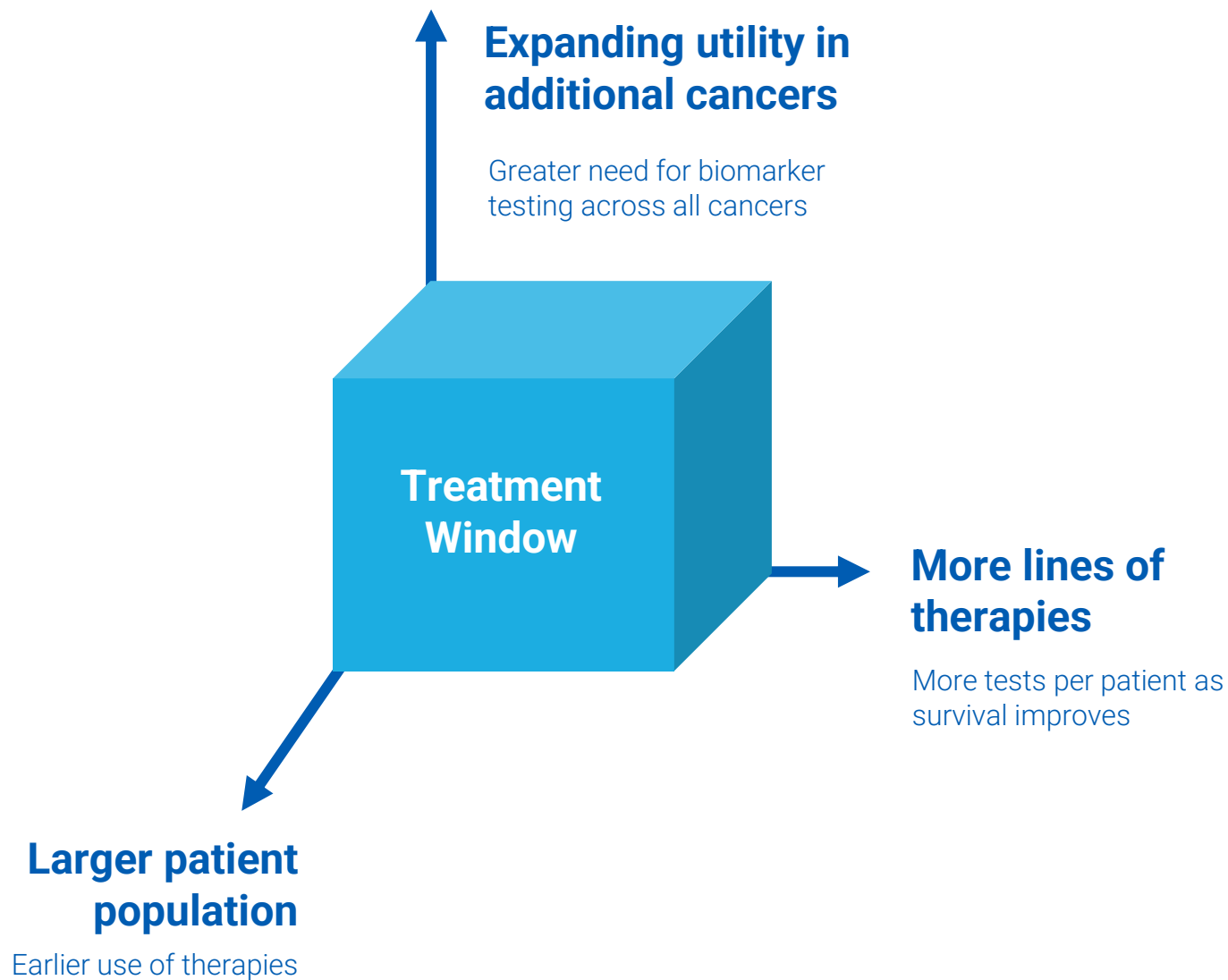


ORSERDU
(elacestrant) is the
first FDA approved
ESR1 therapy

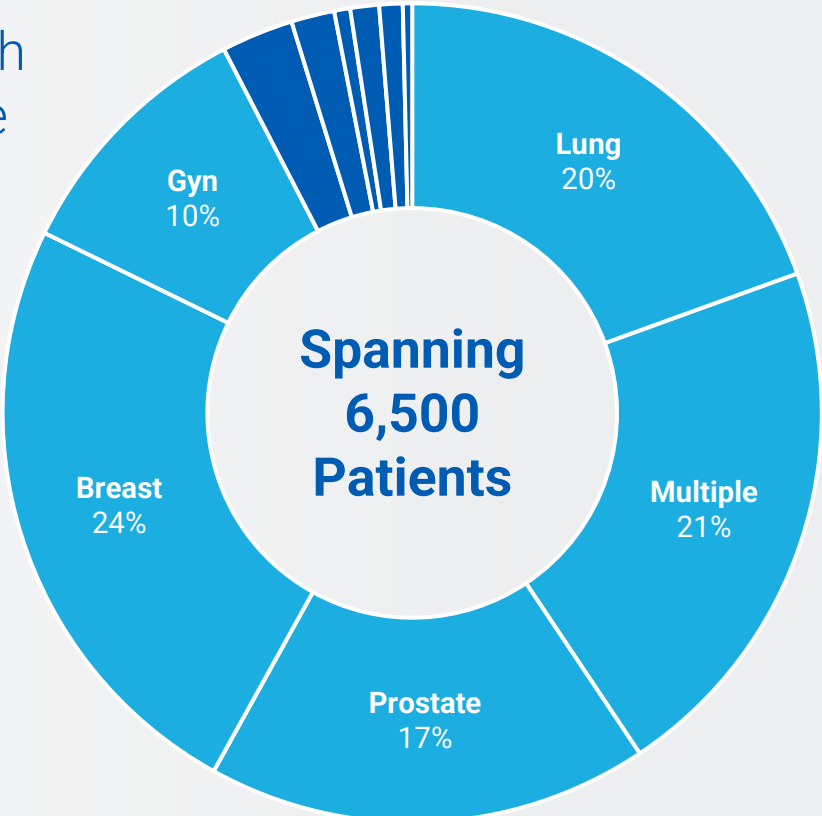
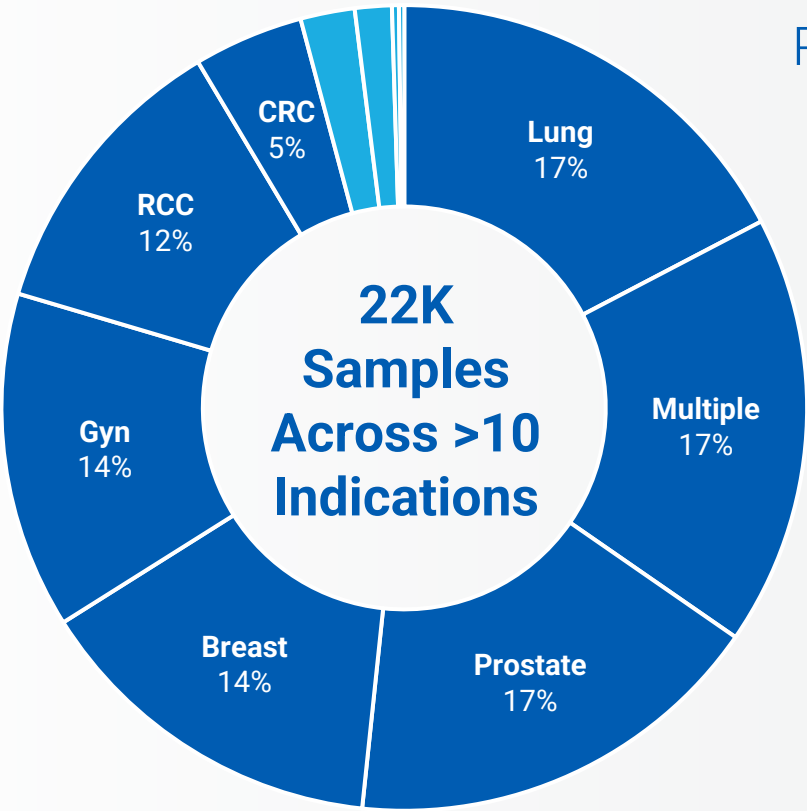


Total of 6 ESR1
programs signed
to date with
biopharma partners

The need for therapy selection in oncology is **growing rapidly**



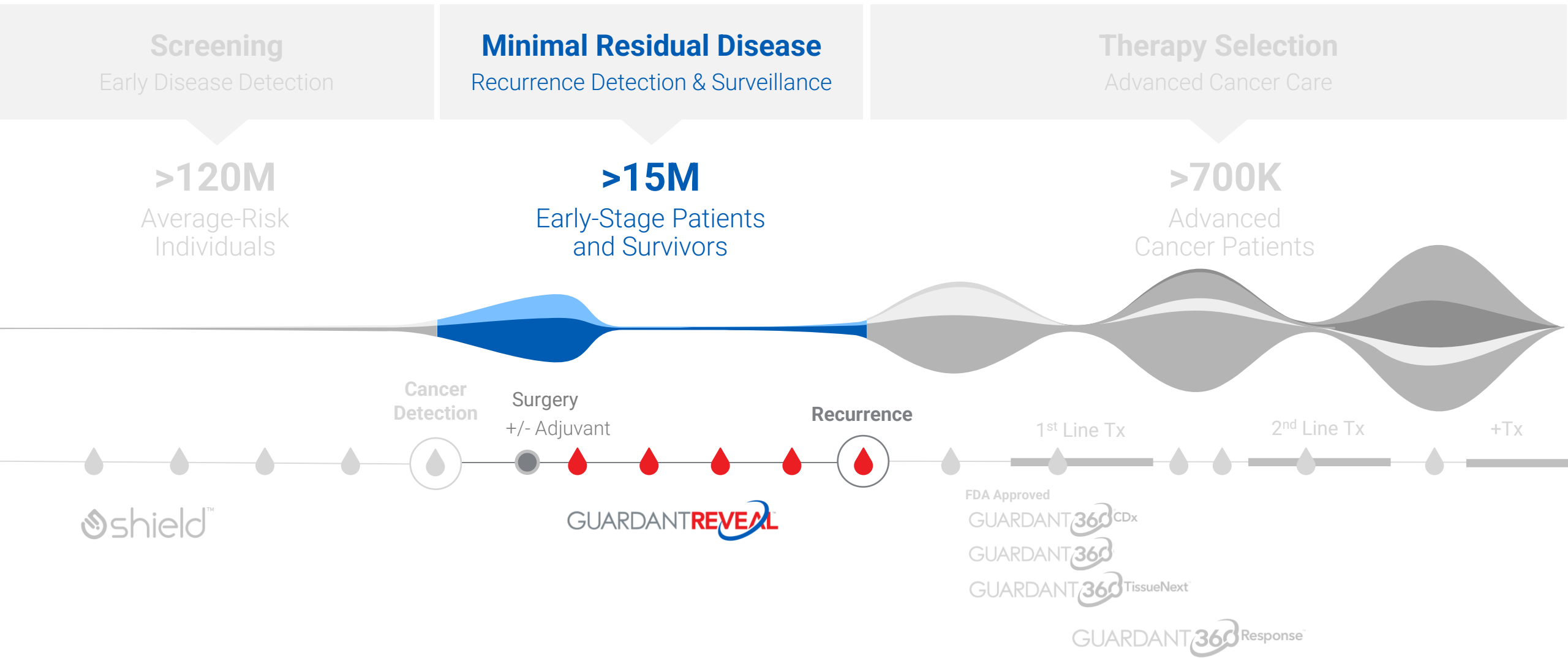
Advancing Therapy Selection with Smart Liquid Biopsy





Paving the way for **tissue-free** MRD

Helmy Eltoukhy, PhD
co-Chief Executive Officer & Chair





The only **tissue-free MRD** test available clinically

- ✓ Currently in CRC, breast and lung
- ✓ Upgrade to Smart Liquid Biopsy this year

\$20B+

U.S. market in
early development

15M+

Patients

<3%

Patients
tested today

The future of MRD is tissue-free

Faster

Decreased initial turnaround time to inform adjuvant decision making or to start initial surveillance

Easier

No need for tissue access for patients in the neoadjuvant, adjuvant settings or who are years out from surgery

Powerful

Ability to inform clinicians with precision, detection of heterogeneous cancers, tracking of tumor evolution and more

But tissue-free detection is **hard**

How can one reliably detect cancer with high sensitivity and specificity **without tissue?**

Blazing a trail for a true liquid MRD test

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23

1

Revolutionary Smart Liquid Biopsy technology achieving high sensitivity with >98% clinical specificity

2

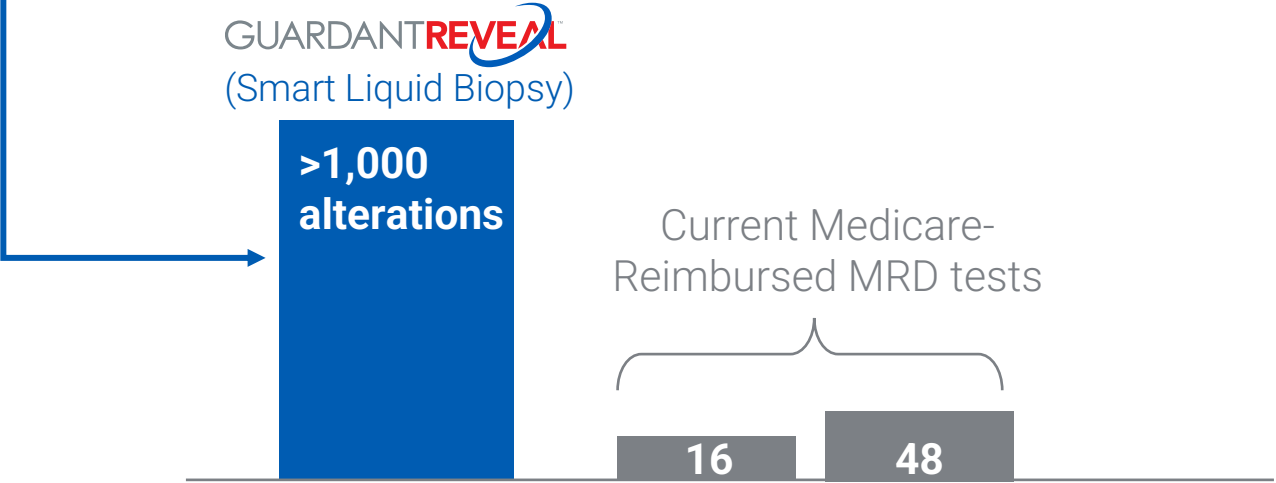
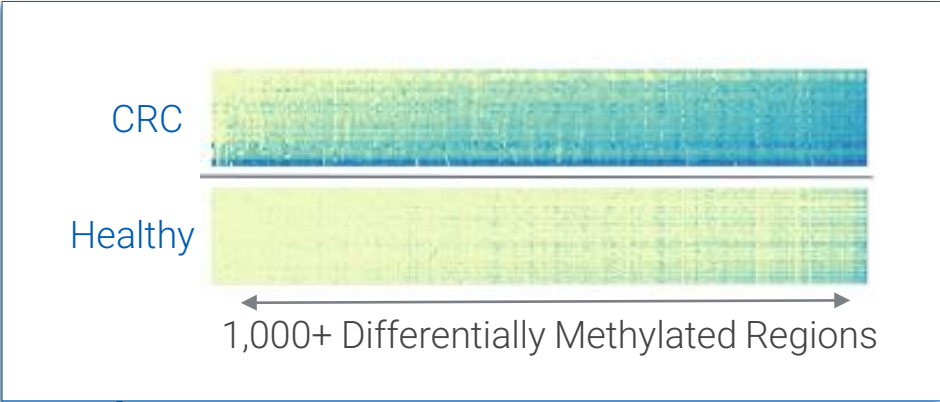
Clinical validation on thousands of patients across dozens of tumor types to secure reimbursement

3

Leverage existing market-leading oncology commercial infrastructure to scale with speed

1

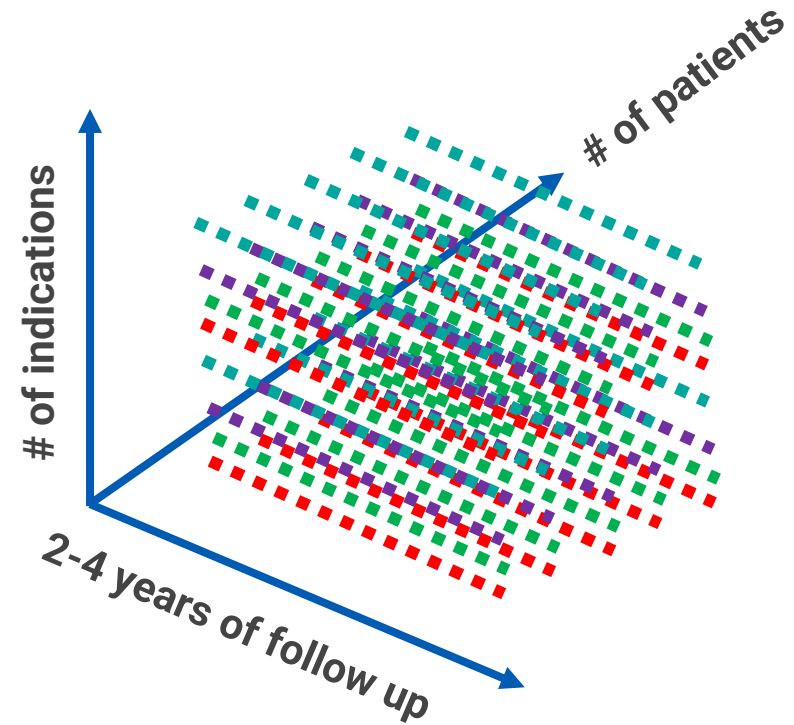
How does our Smart Liquid Biopsy technology achieve **high performance for MRD?**



Low background methylation chemistry + Genome-wide detection + Machine-learning on 1,000s of samples > **High sensitivity & high specificity**

2

Clinical validation of any new MRD test requires a **significant number of samples** with years of follow-up



10,000s of clinical samples

2

80K

MRD samples for Smart Liquid Biopsy clinical studies

Near-term Clinical Validity & Utility Cohorts

Indication	Description	Patients	Plasma Timepoints
CRC	COSMOS	>300	>2,000
IO monitoring	Multi-center	>1000	>3,000
Breast	Multi-center	>300	>300
Lung	Single center	>300	>700
Breast – TNBC	Single center	>100	>500
Breast – Mixed	Single center	>200	>1,500
Gastric	Multi-center	>400	>2,000
Pancreatic	Multi-center	>400	>2,000
Head & Neck	Multi-center	>115	>700

17
Solid tumor types

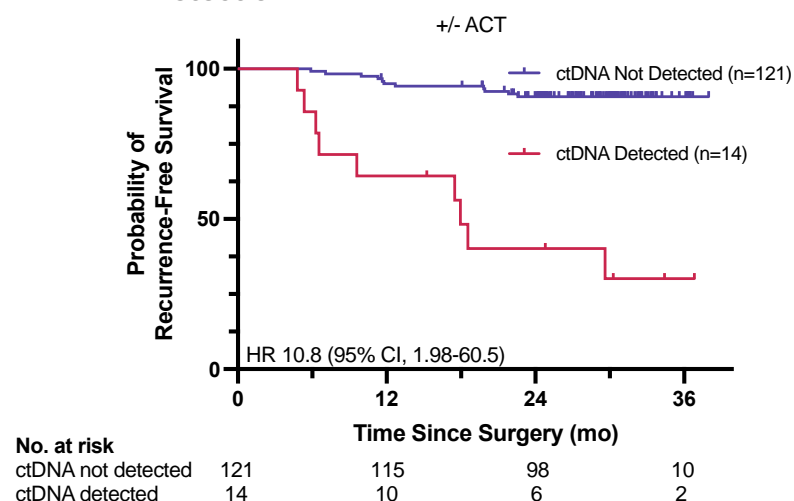
18K
Patients

New data

Smart Liquid Biopsy MRD

COSMOS-Colon Update: Stage II/III completely resected (R0) population

RFS Based on 4 Week Post-Surgery ctDNA Detection



ctDNA status	No. of Events / Total No.	Median RFS	12M RFS (95% CI)	24M RFS (95% CI)	36M RFS (95% CI)
Not Detected	11 / 121	NR	95.0% (89.3-97.7)	90.7% (83.9-94.8)	90.7% (83.9-94.8)
Detected	9 / 14	18.0mo	64.3% (34.3-83.3)	40.2% (15.1-64.4)	30.1% (8.4-56.0)

80%

Sensitivity
prior to or at recurrence

99%

Specificity
(sample-level)

Data to be included in Medicare CRC surveillance submission

Highlights of additional MRD studies underway

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Near-term readouts

Breast Cancer Clinical Validity Study

- Post-adjuvant chemoRx cohort
- Submitted to San Antonio Breast Cancer Symposium (SABCS)
- > 300 early breast cancer patients across all subtypes
- Median follow-up of 66 months

PEGASUS De-Escalation Clinical Utility Trial (Standard of Care Chemotherapy) CRC

- Ph II single-arm study
- Data readout submitted to ESMO
- 140 high-risk stage II & stage III colon cancer patients after surgical resection
- Study fully enrolled in 2022

Long-term readouts

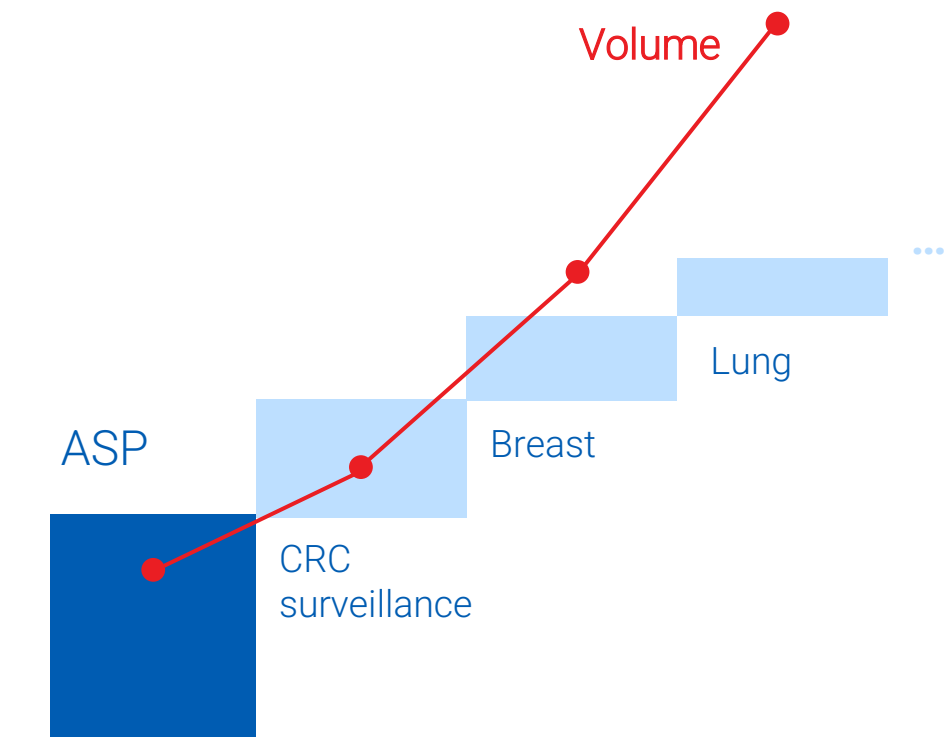
TRACC Part C De-Escalation Clinical Utility Trial (Standard of Care Chemotherapy) CRC

- Ph III randomized control trial
- Study enrollment ongoing
- 1621 high-risk stage II & stage III colon cancer patients after surgical resection
- Expect to complete enrollment in early 2027

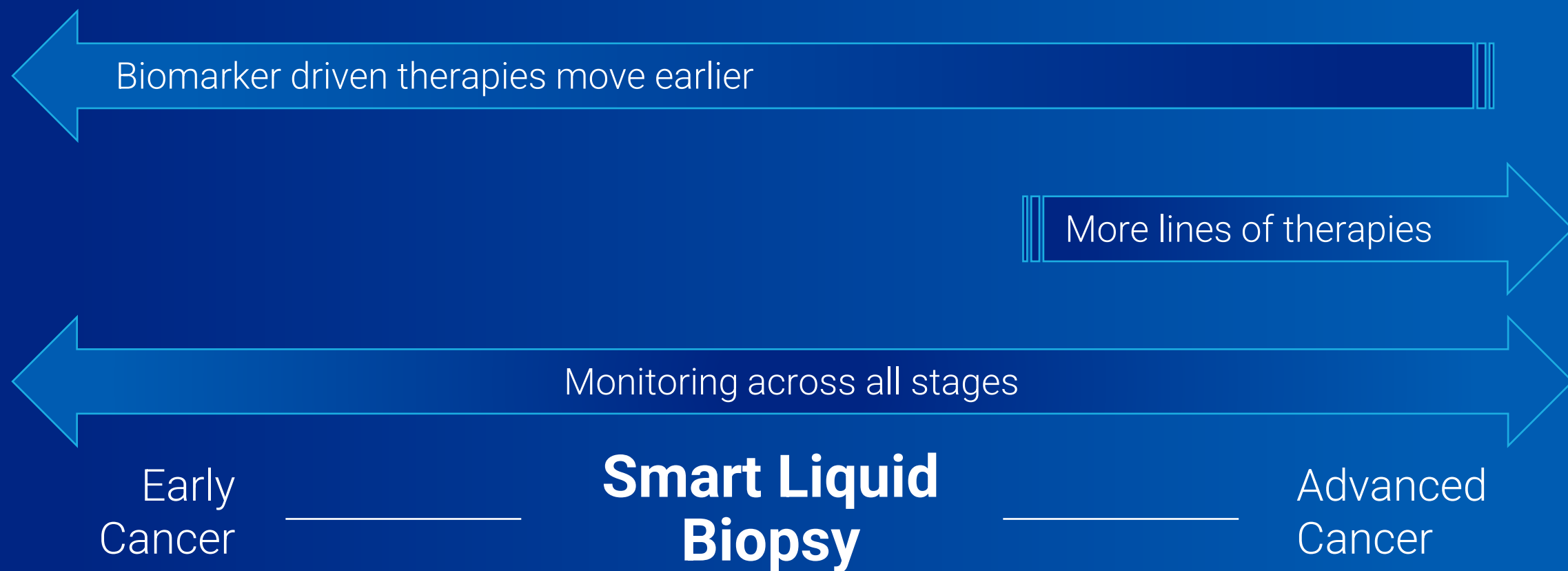
3

We can tap into our 380+ Oncology, and, over time, Screening commercial teams to **efficiently commercialize MRD**

Reimbursement-gated Volume Trajectory



The future of oncology is one precision medicine paradigm for all patients



Imagine a future in
which a simple
blood test offers
comprehensive
**molecular, spatial and
prognostic information**
streamlining patient care



Our Smart Liquid Biopsy Platform can make that
vision a reality



A **leading** commercial platform in oncology

Chris Freeman

Chief Commercial Officer

Setting a new standard for patient care and customer experience

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A **shared commercial infrastructure** across the end-to-end product portfolio

Significant untapped potential in the Therapy Selection market

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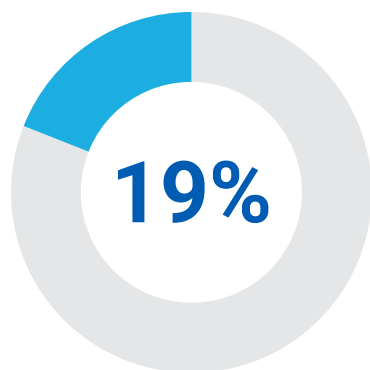
700K

Total Patients

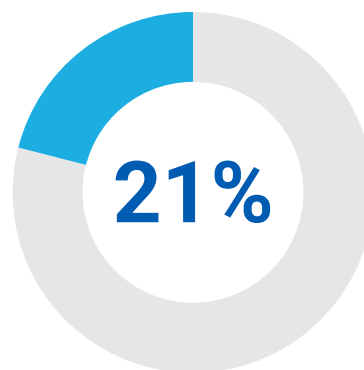
\$10B

Total Addressable
Market

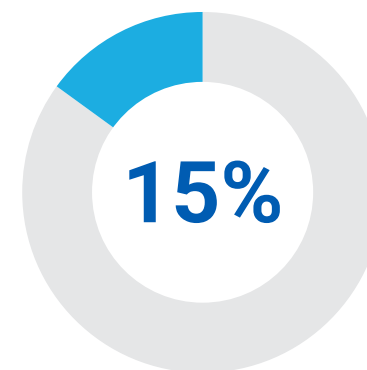
Guardant360 has established footholds in the major tumor types for Therapy Selection



NSCLC

**240K**

Breast

**119K**

CRC

**103K****Total Patients**

Therapy Selection utilization will continue to grow

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1

Increase liquid and tissue adoption

2

Unlock new actionable biomarkers and use cases

3

Physicians are using CGP testing in earlier lines

Industry-leading product portfolio with revolutionary technology

GUARDANT **360**^{CDx}

1st FDA-Approved
Comprehensive Liquid Biopsy



GUARDANT **360**[®]

Next-Generation
Liquid Biopsy Assay



GUARDANT **360**^{TissueNext}

Next-Generation
Tissue Assay



GUARDANT **REVEAL**[™]

1st Blood-Only Recurrence Monitoring
Assay



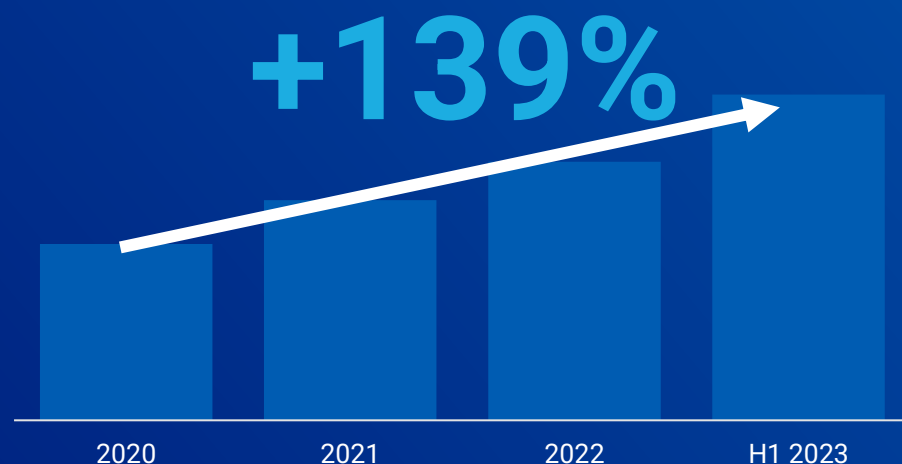
GUARDANT **360**^{Response}

1st Blood-Only Liquid Biopsy
to Monitor Tx Response



Reimbursed by Medicare

Portfolio offering provides strategic advantage

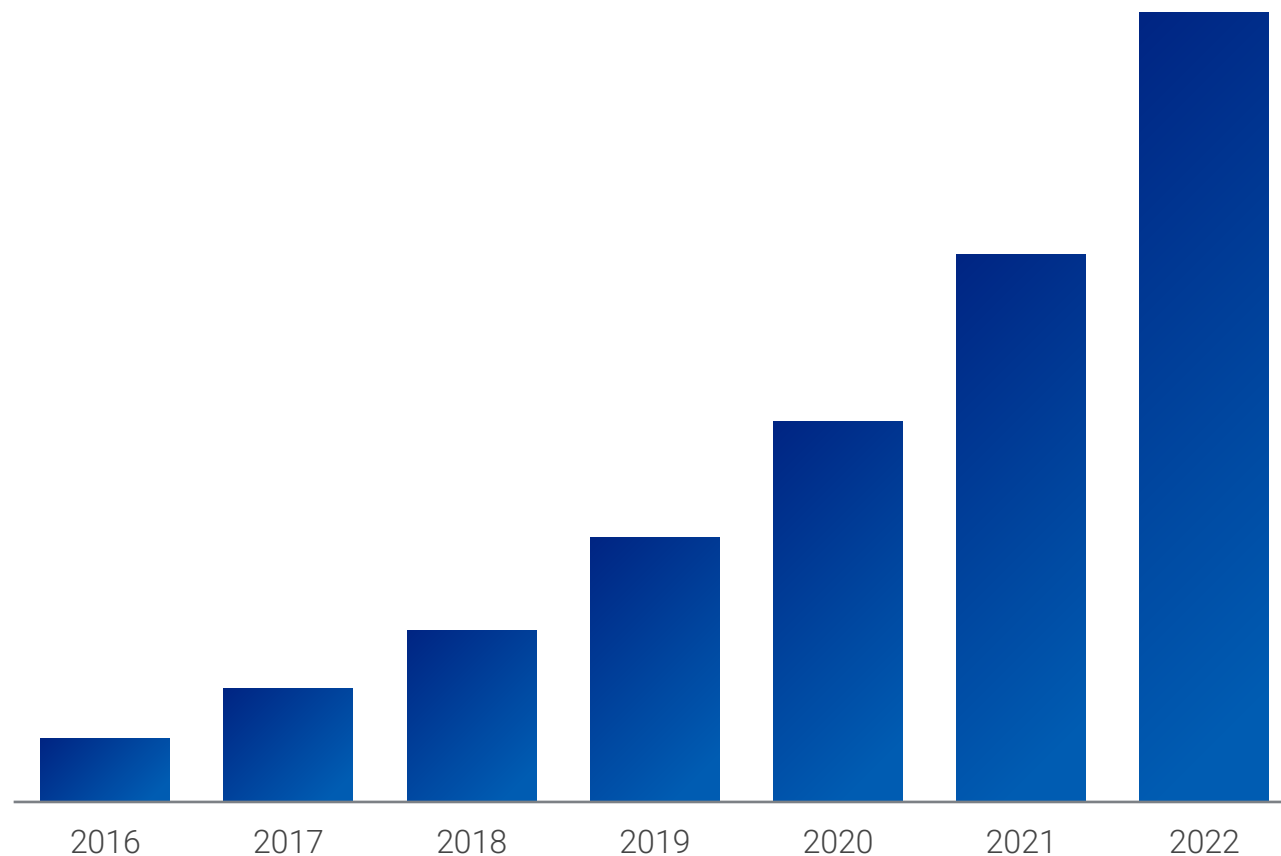


Market adoption of concurrent
blood and tissue testing

69%

Of oncologists choose their
testing partner based on
“one-stop-shop” ordering¹

Guardant
innovations
have helped
influence
~**500K** cancer
care decisions



Best-in-class products are supported with best-in-class operations

5 days

Median Guardant360 CDx turnaround time¹

>60%

of oncologists are motivated by a FAST turnaround time when choosing a testing partner²

#1

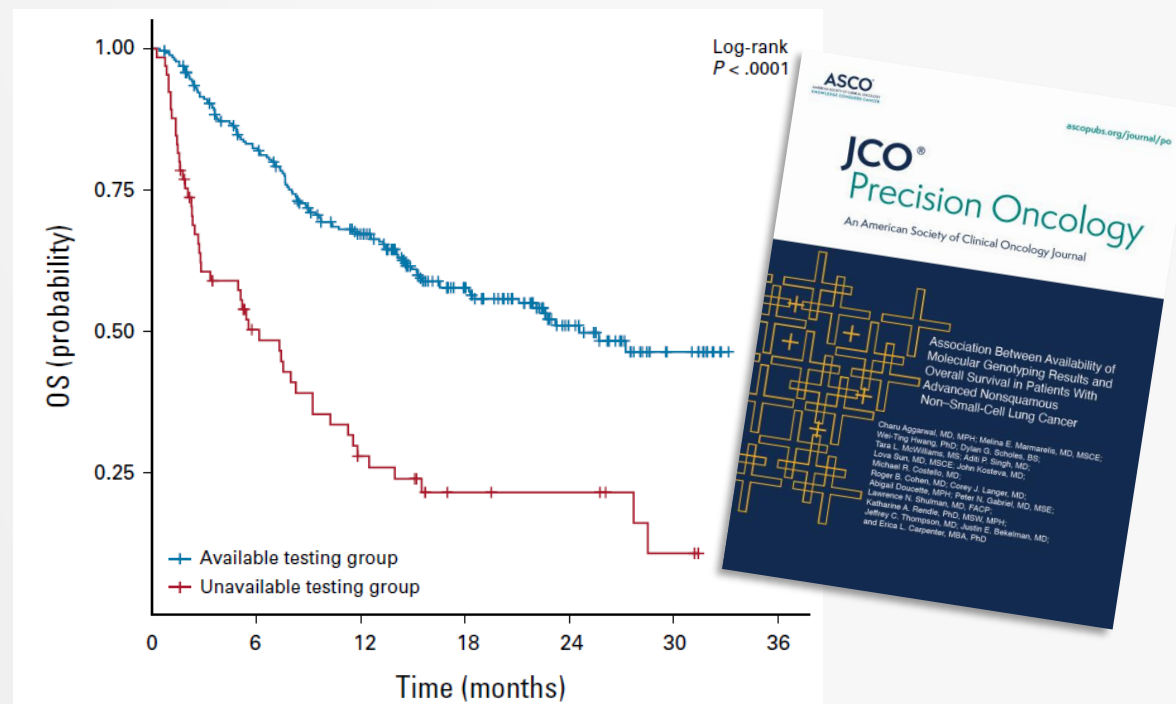
Industry leading turnaround time for an FDA approved CGP

Actionable information and speed saves patient lives

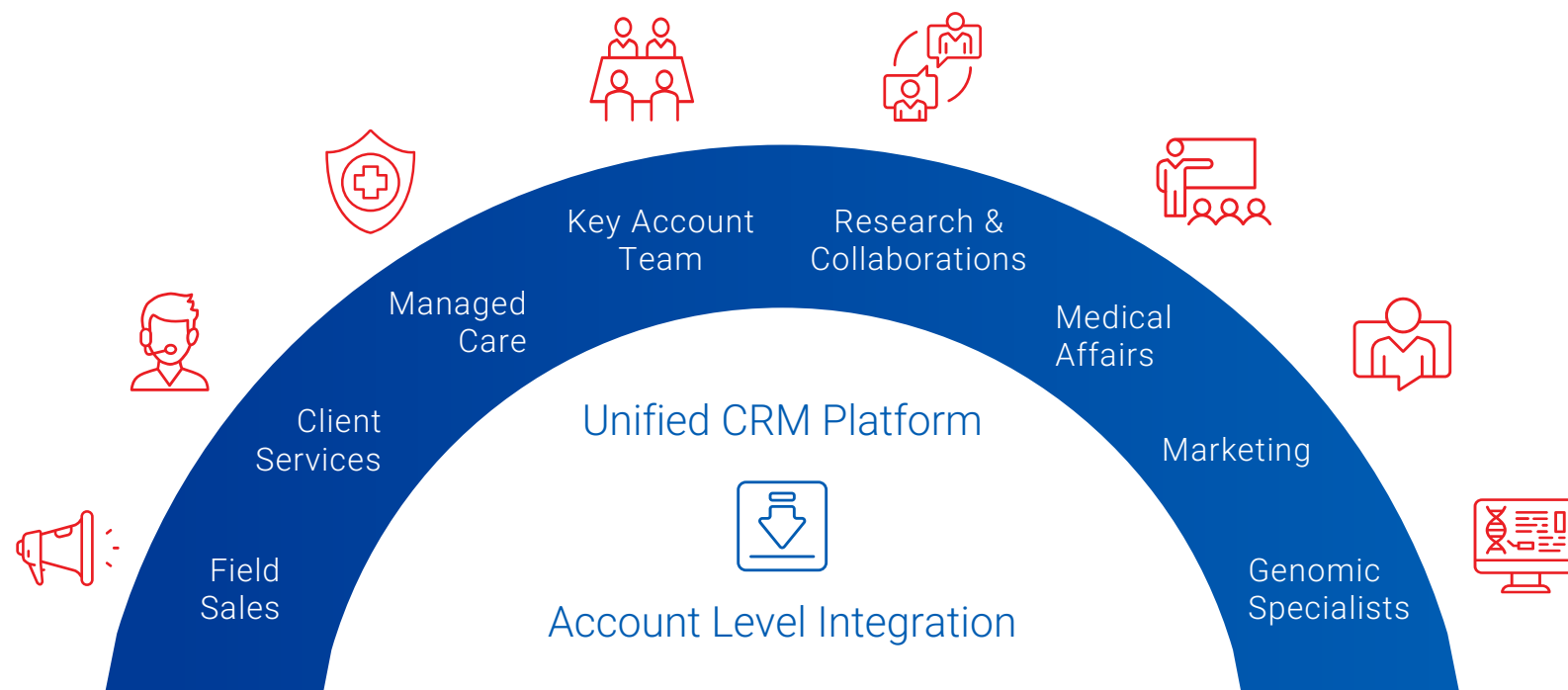
4x Better patient outcomes with CGP testing

Every month delayed in cancer treatment can raise risk of death by around 10%¹

Association Between Availability of Molecular Genotyping Results and Overall Survival in Patients with Advanced Nonsquamous Non-Small-Cell Lung Cancer



Commercial excellence begins with a **world-class** field team



#1 Rated Field Team¹

#1 Rated Support¹

#1 In-Person Share of Voice¹

#1 Remote Share of Voice¹

EMR integration accelerates access to critical information

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Our EMR partnerships will cover **65%** of all oncologists in the U.S.¹

EMR integration reduces oncologist ordering time by **75%**²

Guardant will be EMR integrated in **>400 accounts** by end of 2023

Broad coverage provides seamless access for patients

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300M+

Covered lives for
Guardant360

Approaching

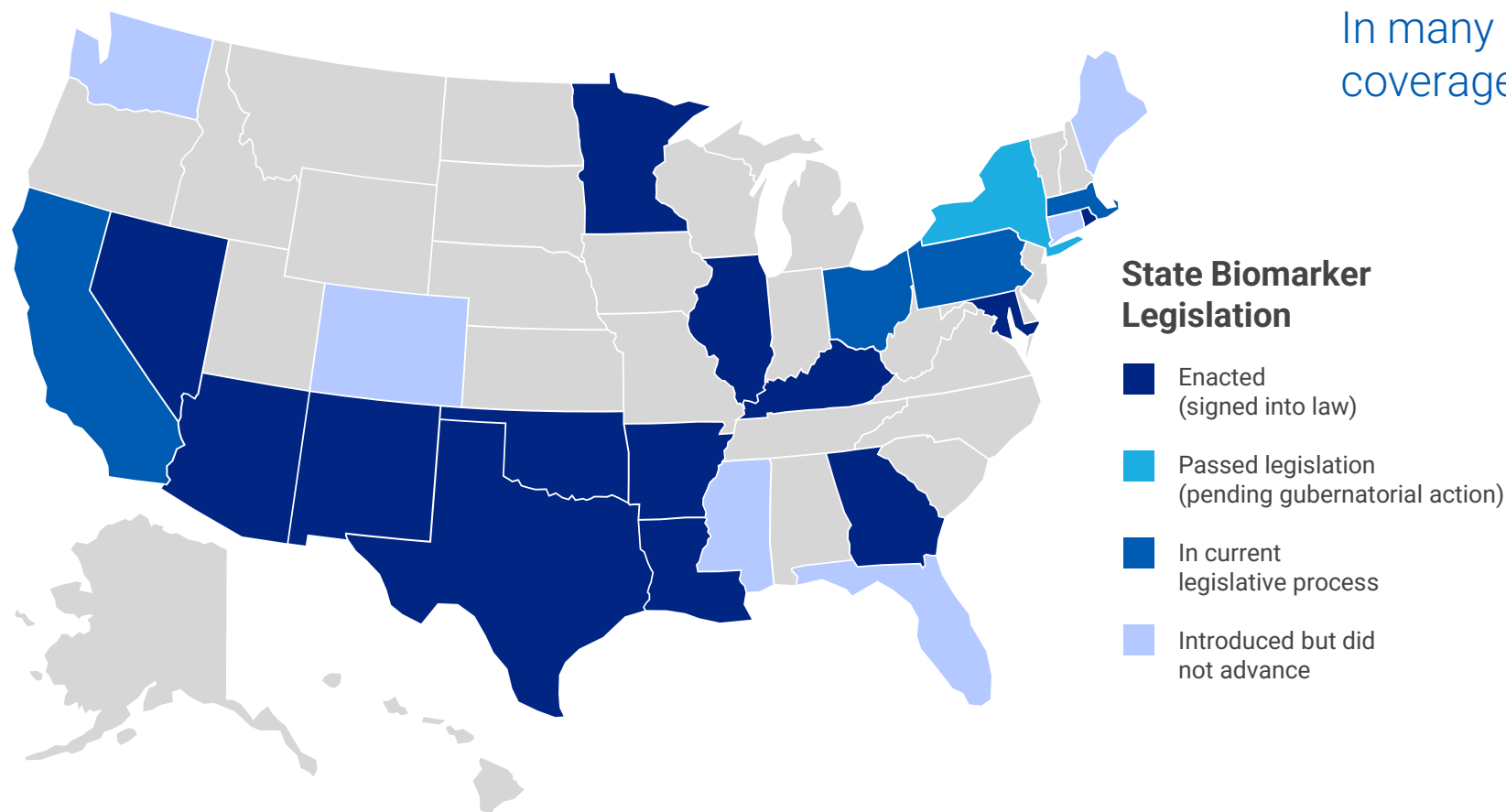
200M

Covered lives for TissueNext



Guardant has created momentum for National Biomarker Testing Coverage

In many cases will support private payer coverage for both Therapy Selection and MRD



12 states enacted **laws** for biomarker testing

5+ states have current legislation in progress

MRD market is largely untapped

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15M+

Total Patients

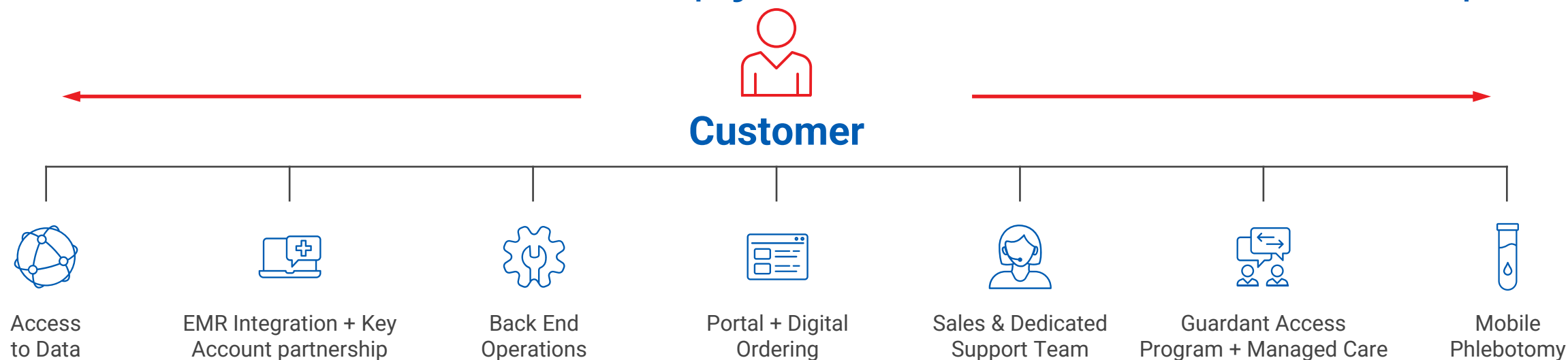
\$20B+

Total Addressable
Market

<3%

Current Testing
Penetration

MRD business can leverage the commercial infrastructure of Therapy Selection to scale at speed



Customer base of
12,000
Oncologists

#1
Field Force + Customer Satisfaction¹

Commercial excellence unlocks additional growth opportunities



Integrated
account-level
partnerships



Biopharma
collaborations



Global
expansion

Biopharma partnerships fuel clinical growth

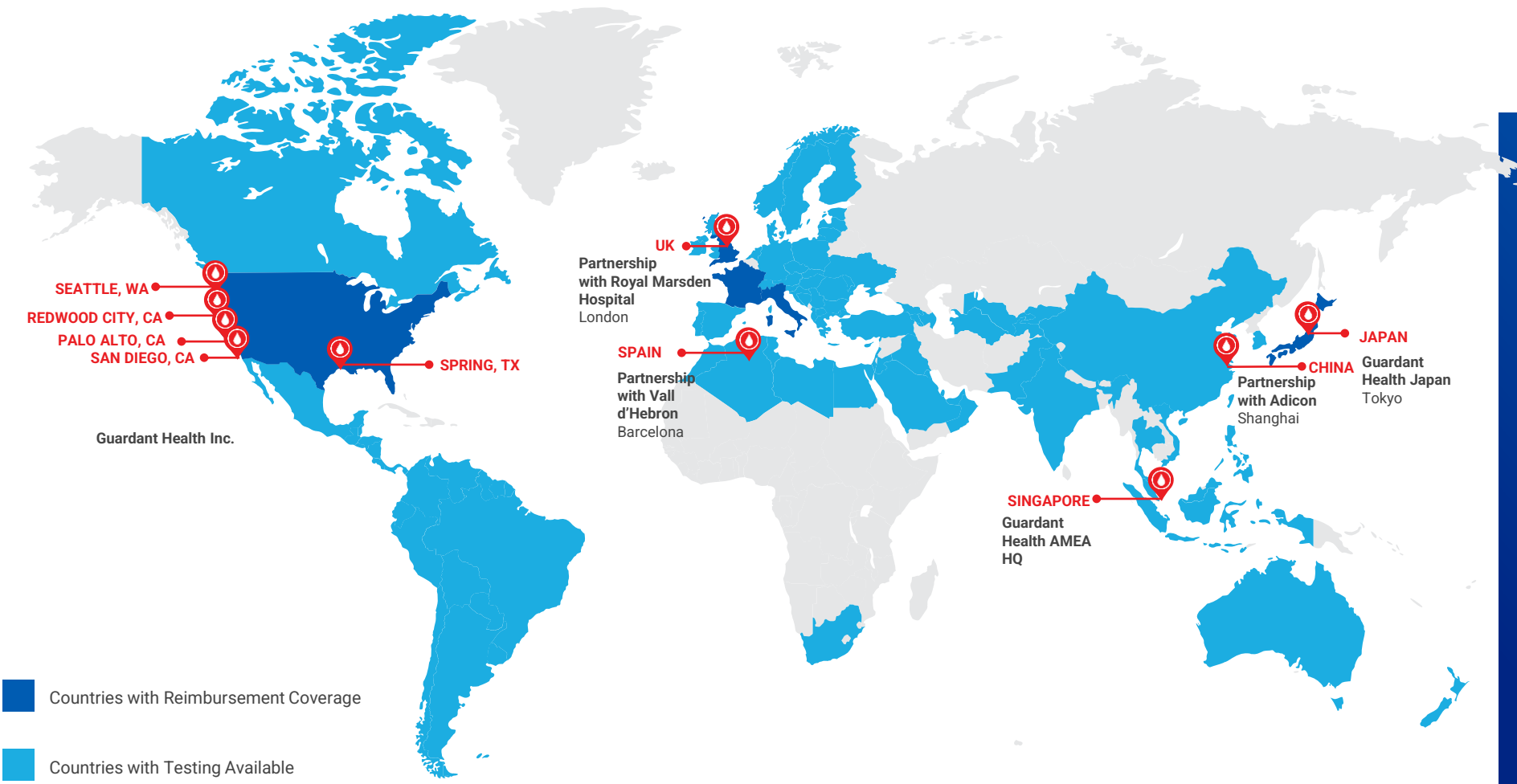
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160+

**Lifetime biopharma
partners**

- ✓ Including 19 of top 20 pharma
- ✓ Multiple active CDx collaborations
- ✓ CDx approvals help drive clinical growth (e.g., ESR1 for breast cancer)

Global commercial organization



60+
Countries

Lab footprint
Spain, UK, China & Japan

380+
Oncology
commercial team

National reimbursement in Japan & lab launch in China taps into large global market

Japan

- ✓ **Potential for rapid growth:** 1M new cancer cases per year¹, centralized care, <10% penetration of CGP market²
- ✓ **Driving business:** National reimbursement approval from the Japanese MHLW for Guardant360 CDx + efficient local sales model + strong ASP

China

- ✓ **Point of differentiation:** China is a critical, strategic market for biopharma partnerships
- ✓ **Strategic partnership for growth:** 30+ biopharma clients have engaged with Guardant since partnership with Adicon lab

Commercial excellence positions Guardant to lead the future of oncology testing

#1



Liquid biopsy CGP market share¹



Field team overall satisfaction²



Share of voice¹

1. Guardant360 Brand Tracker Q4 2022; 250+ team rated first in customer satisfaction by HCPs for past 7 quarters; 50% more oncology interactions compared to closest CGP company. *Growth is ordering oncologists/quarter and tests per oncologist/quarter, 2 Q2 2023 Brand Tracker

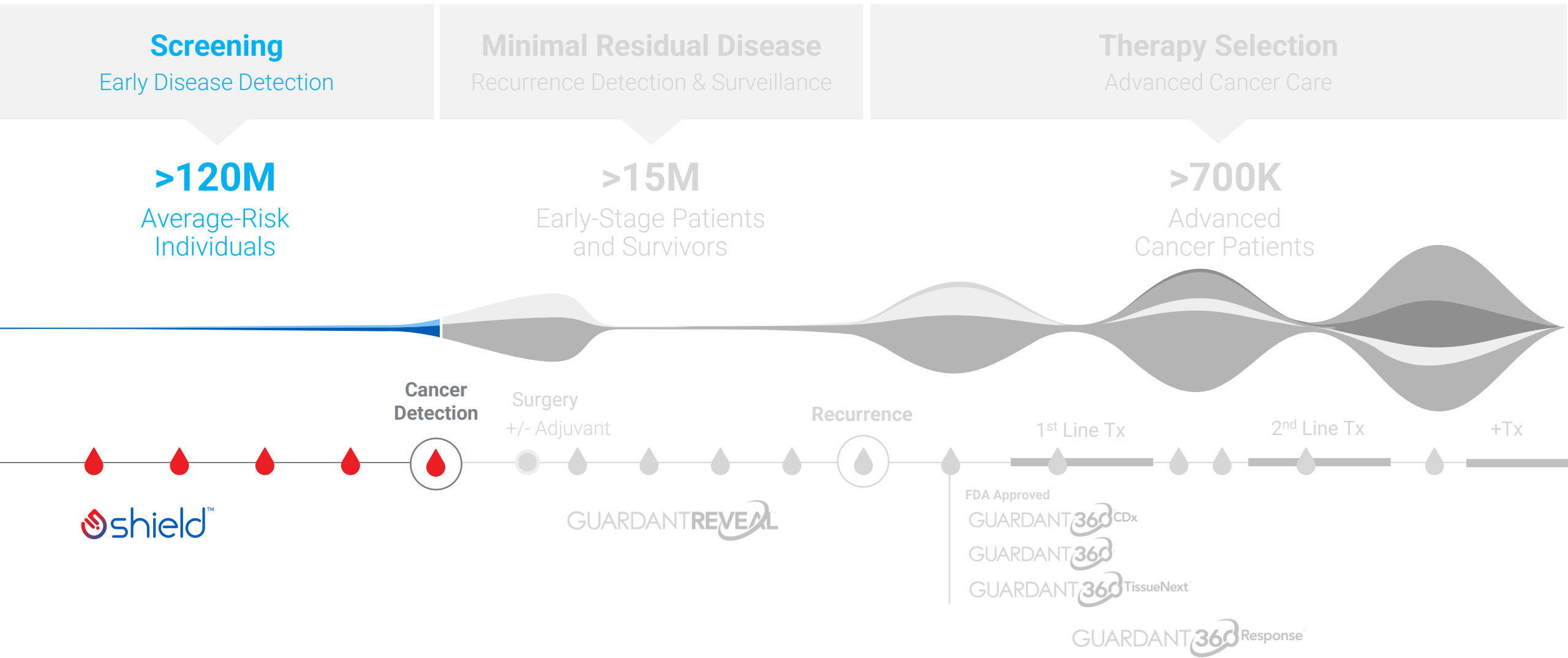
Q&A



Pioneering a **new category** of cancer screening

AmirAli Talasaz, PhD

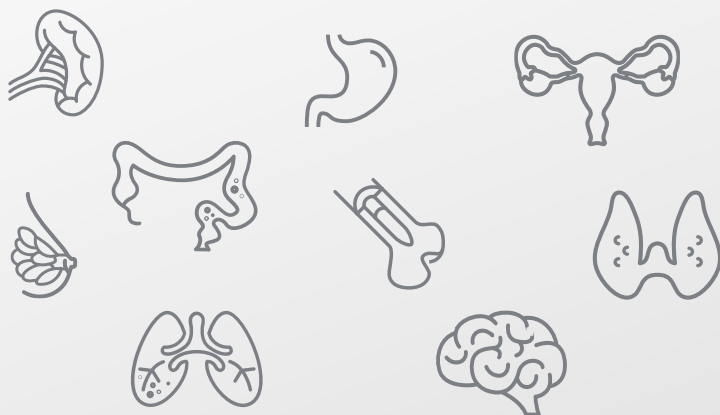
co-Chief Executive Officer & co-Founder



Blood-based screening will change the cancer screening landscape

Metastatic **14%** Survival

Statistics from 26 different cancers comprising 87% of all cancers in U.S.



Pleasant Consumer Experience

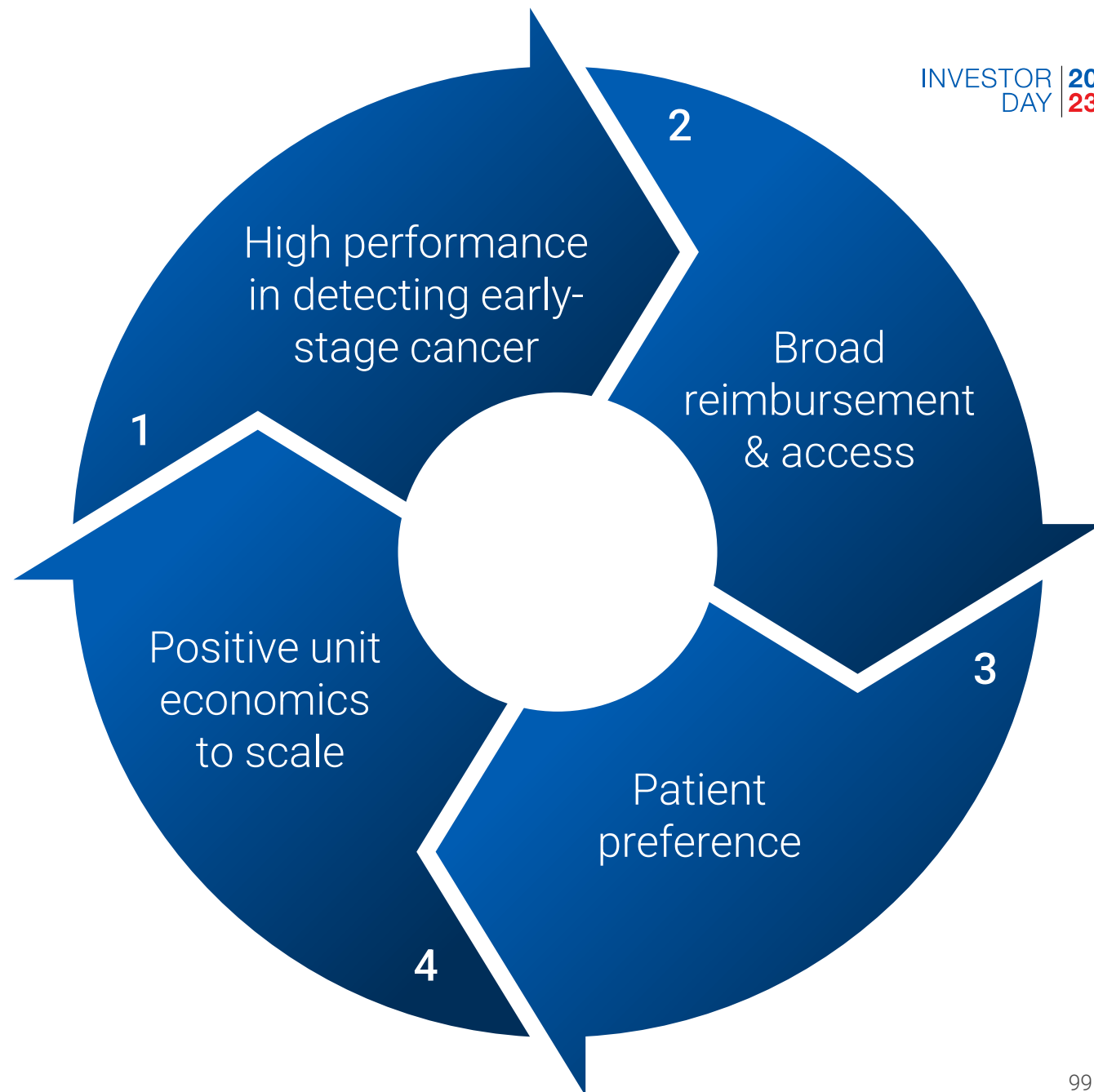


High Performance MCD Test

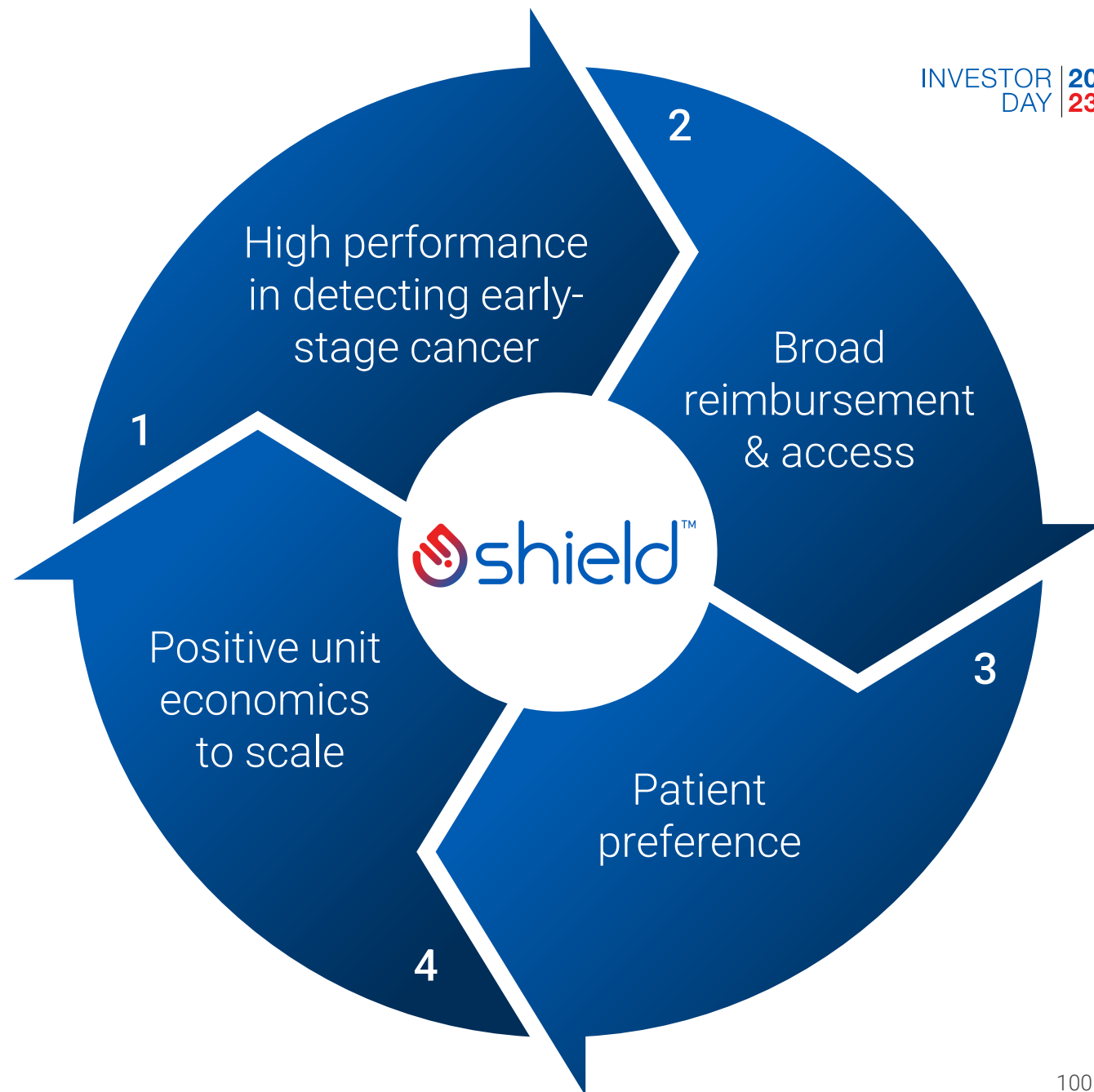
Local
90% Survival

Regional
65% Survival

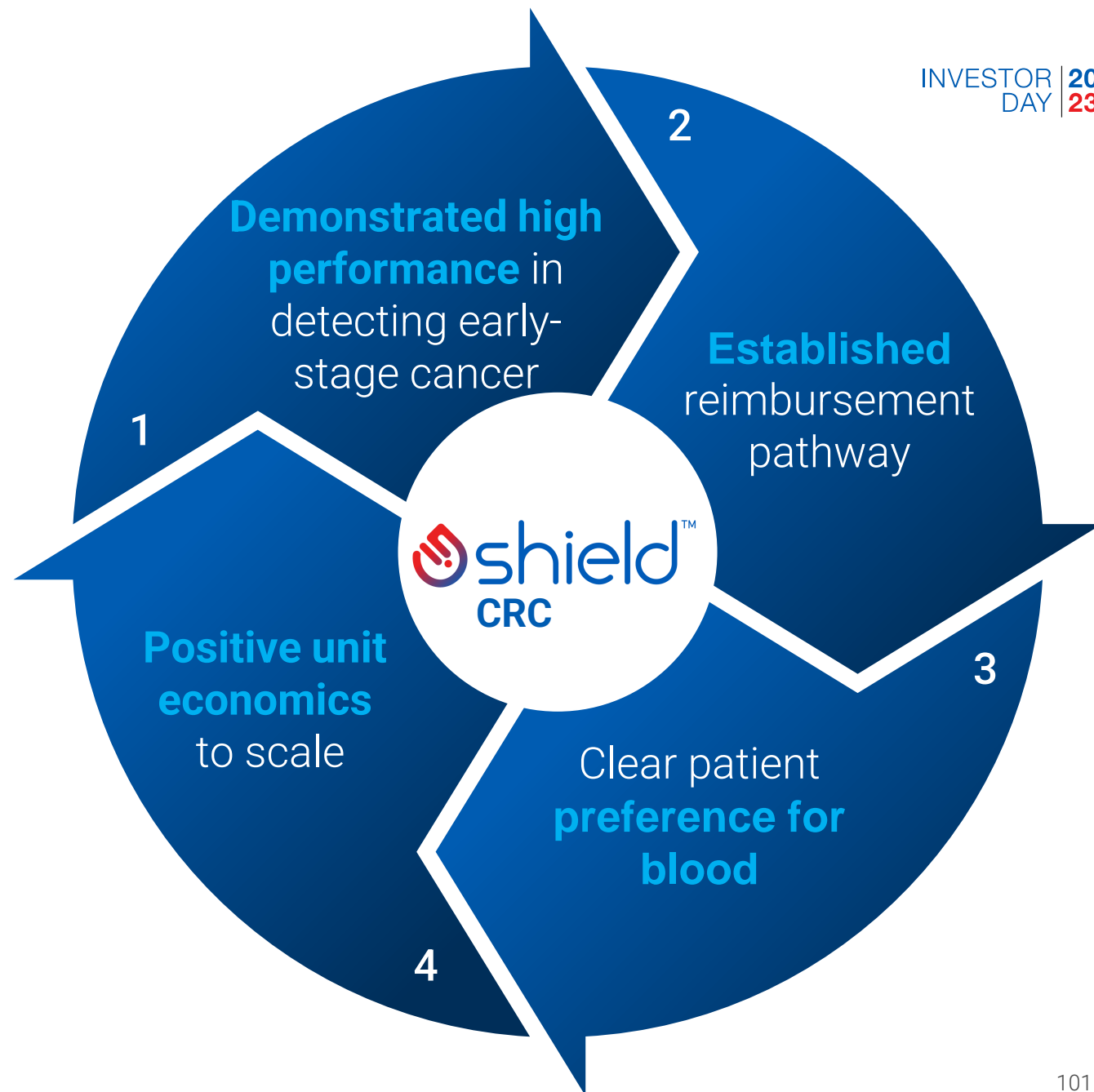
A successful
screening modality
requires **four**
components



The future of screening is a high-performance **blood-based test for multi-cancer**



We strategically started with **CRC** as **the first indication** for Shield



Lack of screening compliance significantly impacts lives lost to colorectal cancer

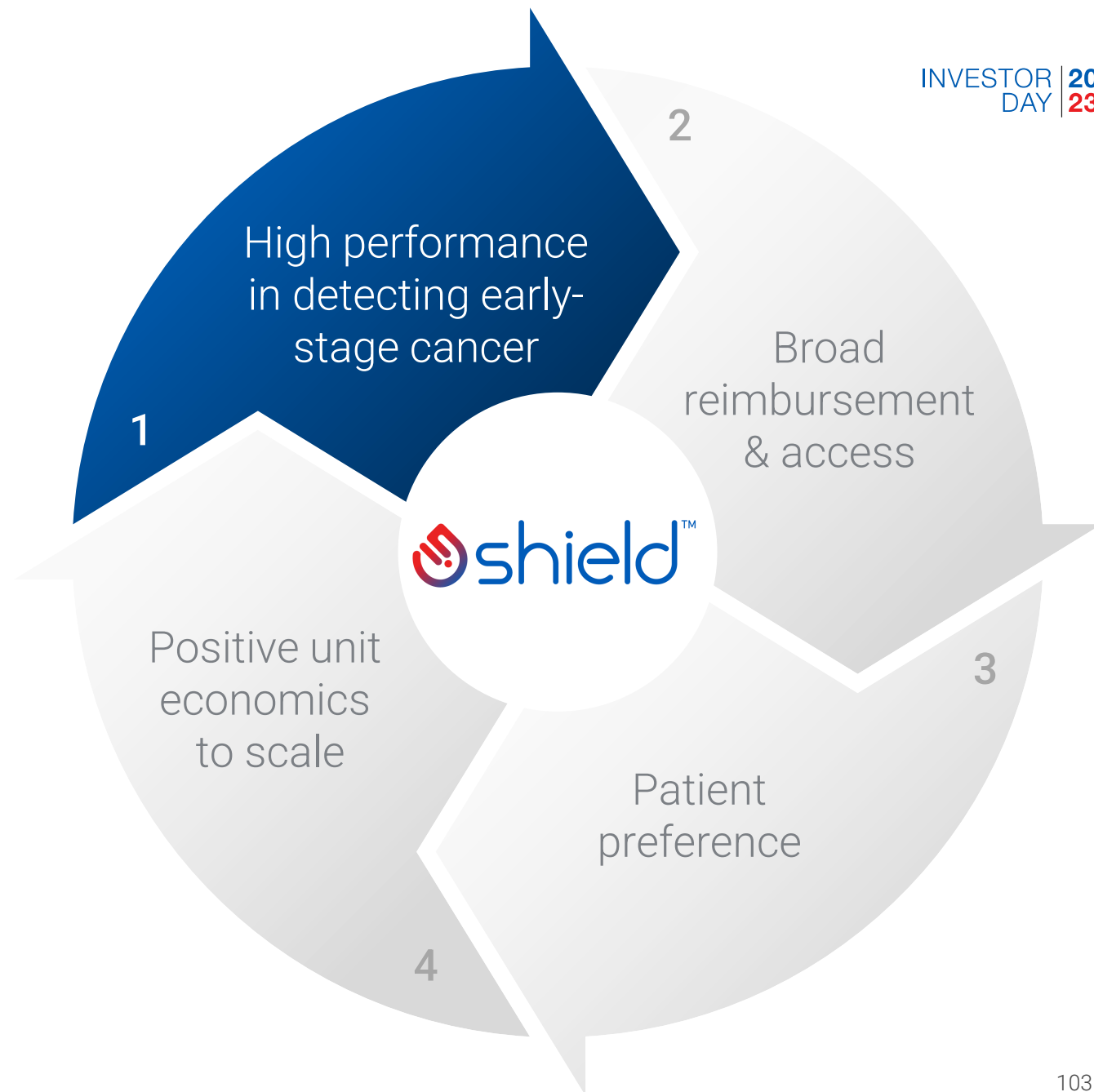
50K

Colorectal cancer deaths per year^{1,2}

76%

of CRC deaths are in individuals not up to date with screening³

Smart Liquid Biopsy analyzes 1,000s of regions in cfDNA with leading sensitivity at low-cost



Shield is the first easy to use, highly accurate blood-based cancer screening test with **proven efficacy in a pivotal trial**



A new screening **category**

ECLIPSE trial demonstrated efficacy

83%

CRC Sensitivity

90%

Specificity

- 100% sensitivity stage II and stage III/IV
- 55% sensitivity stage I
- 81% CRC sensitivity at stage I – III
- 13% advanced adenoma sensitivity

Shield CRC performance is in range with other non-invasive guideline-recommended modalities

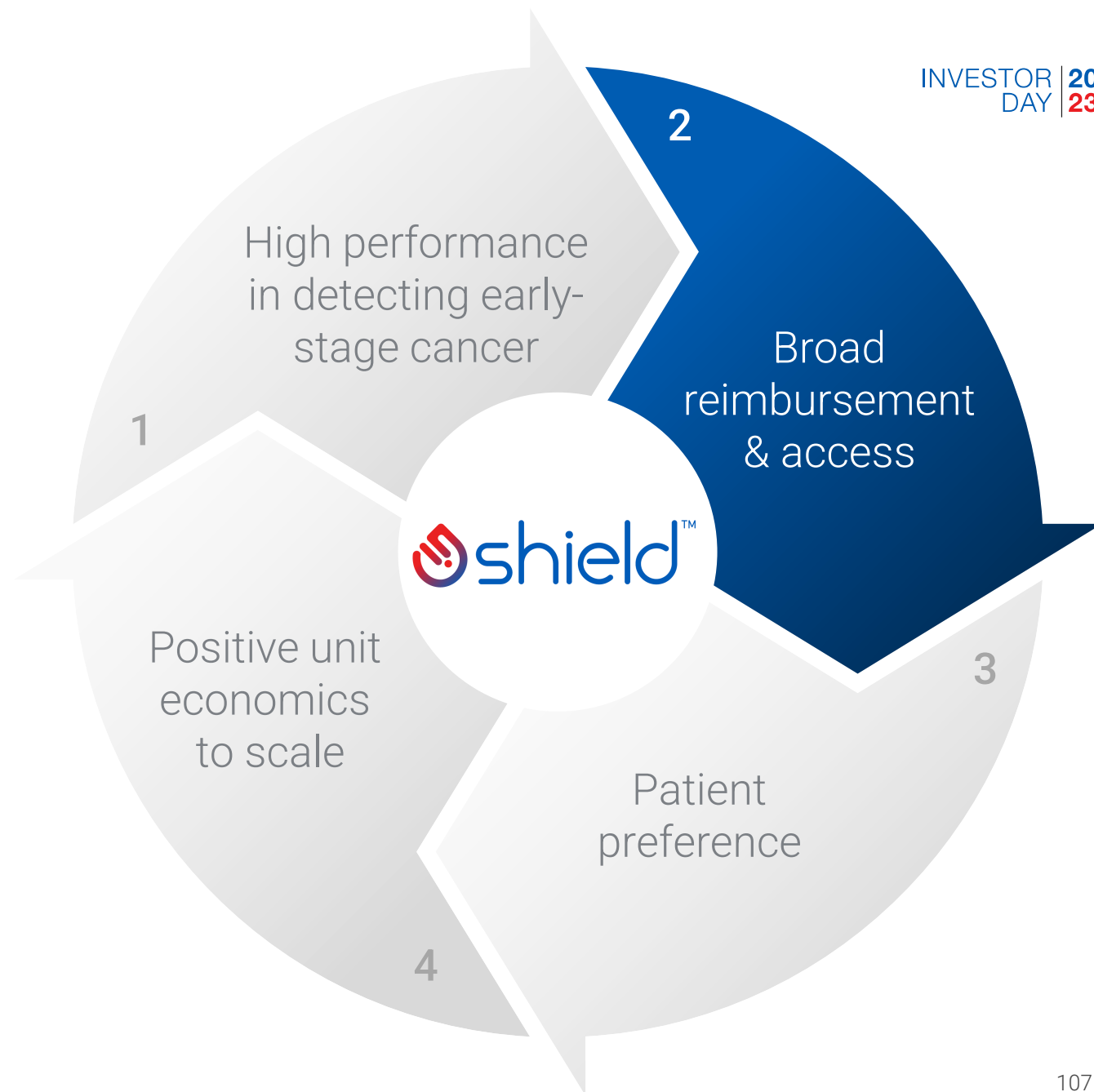
74–92%

CRC sensitivity for current guideline-recommended tests^{1,2}

87–96%

Specificity for current guideline-recommended tests^{1,2}

Broad and equitable access
is key to durable
adoption and
outcome
improvements



Reimbursement pathway for Multi-Cancer Shield starts with colorectal cancer indication



 shield™ Multi-Cancer



...



 shield™ CRC

Reimbursement pathway for Shield starts with colorectal cancer indication



shield™ CRC

Clear pathway starting with CRC



Medicare Coverage

Blood-based CRC screening NCD
with 3-year interval



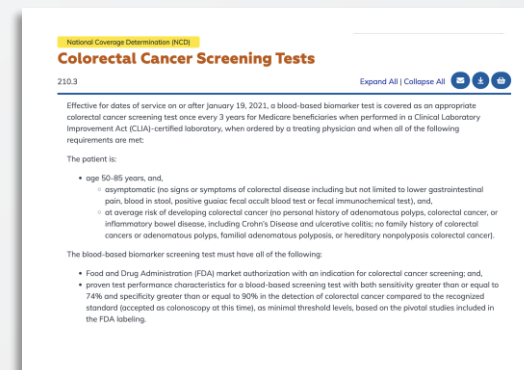
Private Coverage

ACS and USPSTF guidelines



Access

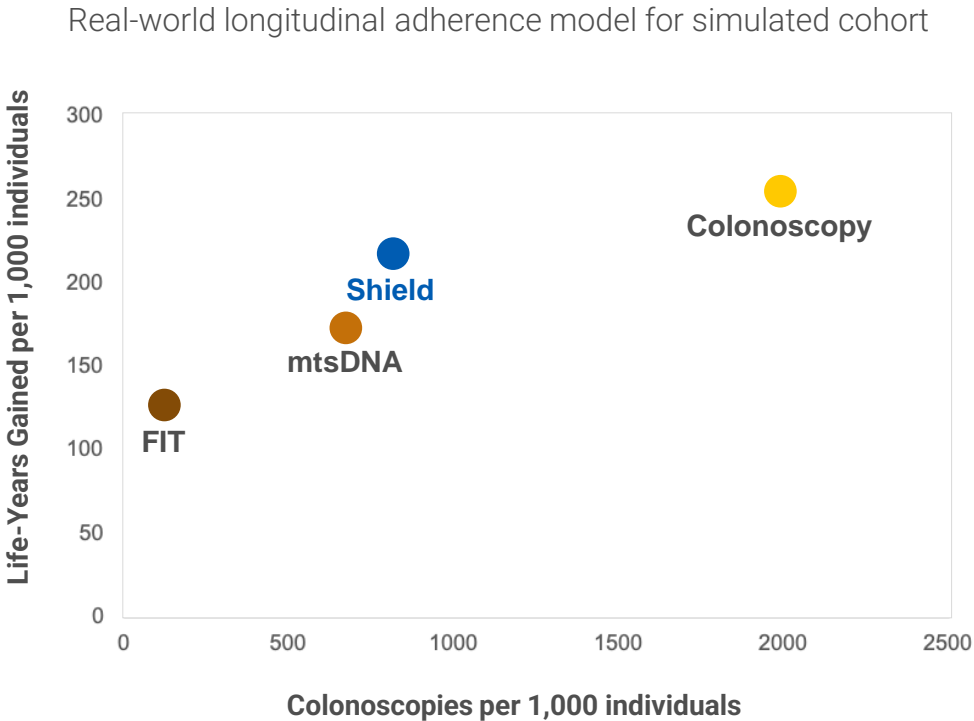
~120M individuals




National Coverage Determination by CMS

USPSTF recommendations are based on the harms and benefits of each testing modality

Lifetime Colonoscopies and Life-Years Gained



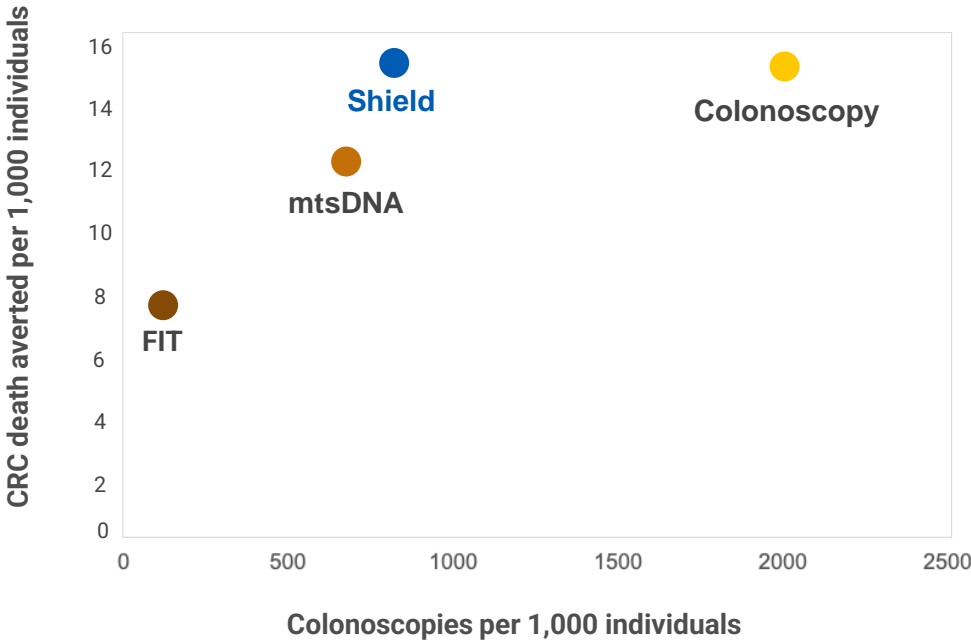
Model Parameters

	Screening Interval ¹⁻³	CRC Sensitivity ^{2,3}	Specificity ^{2,3}	One-Time Adherence ³⁻⁶
Colonoscopy	10 years	95%	86%	38%
FIT	1 year	74%	96%	43%
mtsDNA	3 years	92%	90%	65%
 shield™	3 years	83%	90%	90%


Outcome model demonstrates that Shield has the highest averted rate of CRC deaths

Lifetime Colonoscopies & CRC Deaths Averted

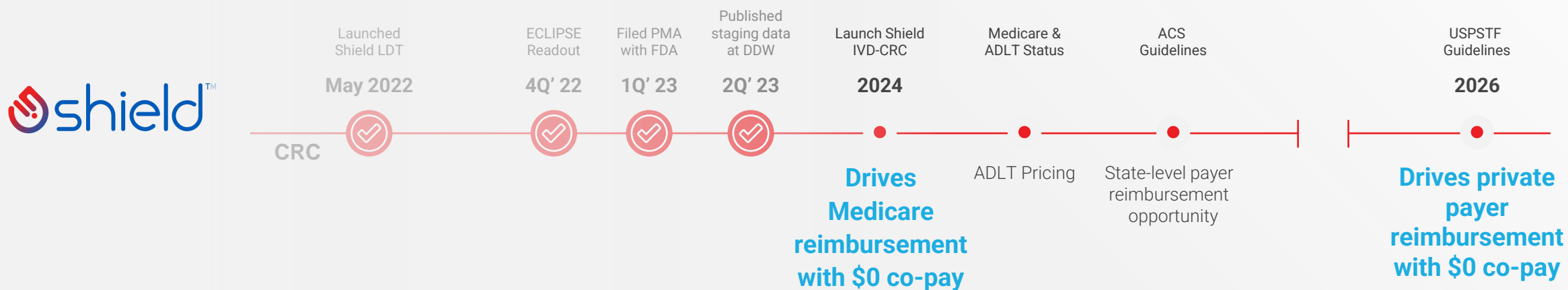
Real-world longitudinal adherence model for simulated cohort



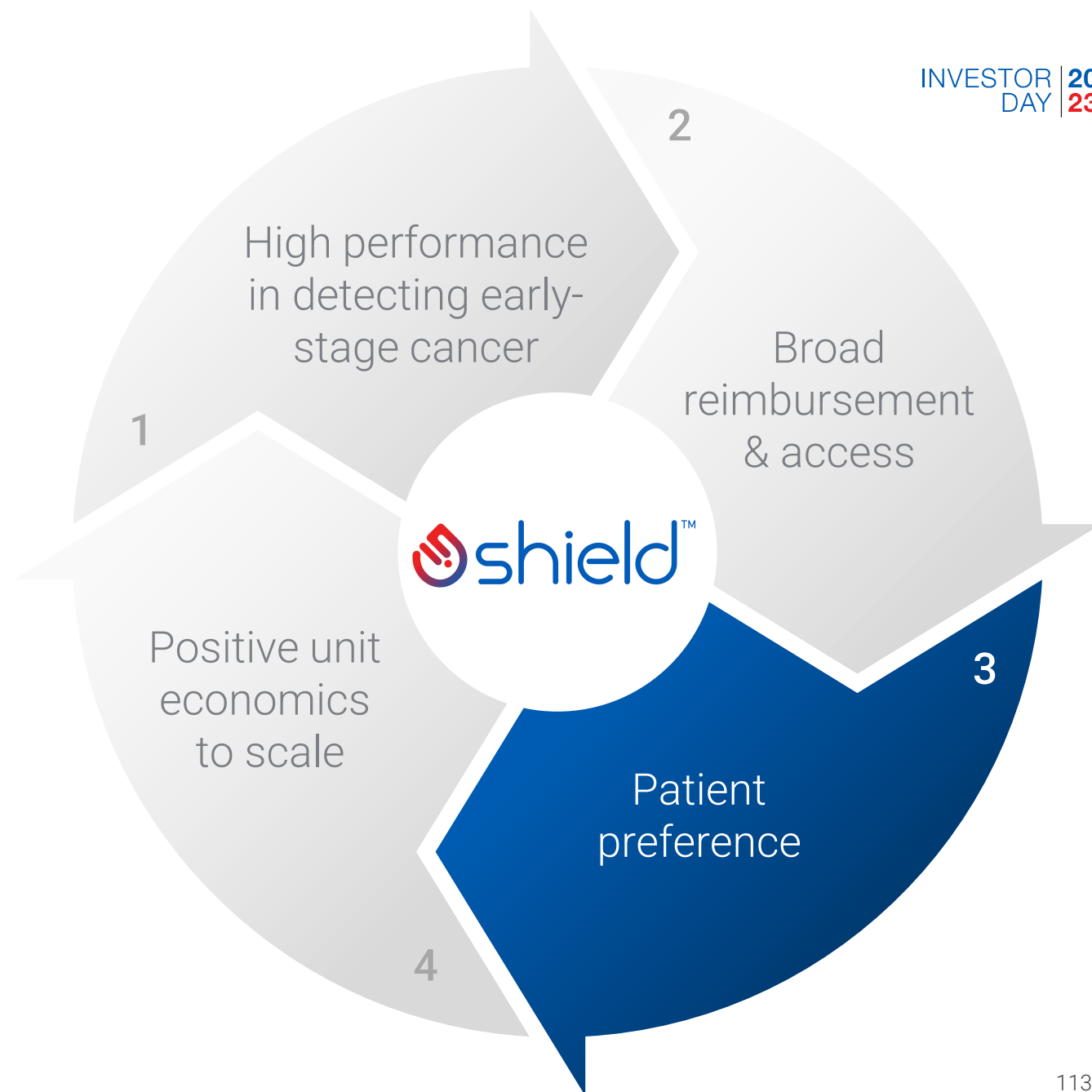
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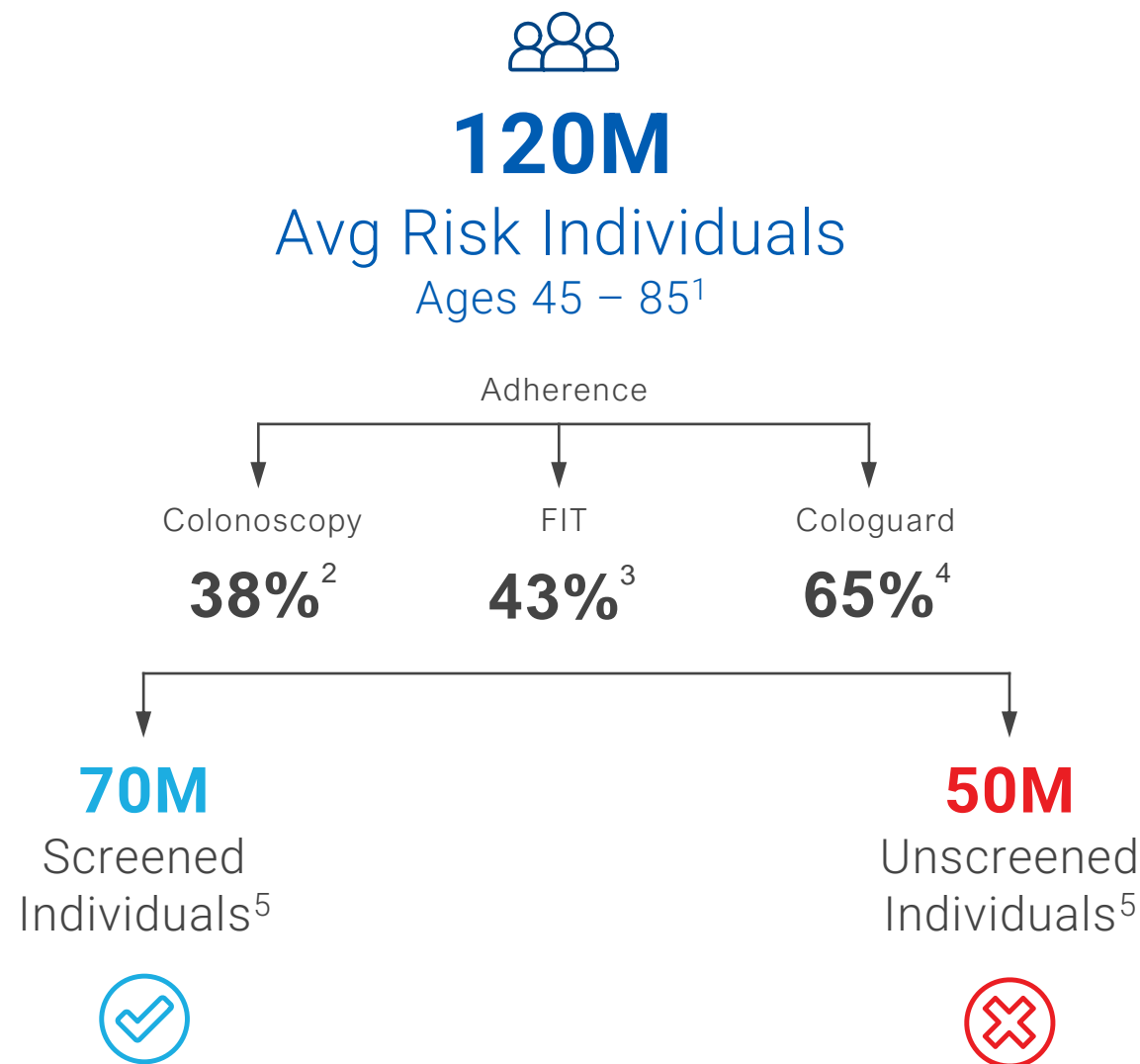
Blazing a trail to the first FDA approved and reimbursed CRC screening test



People want a
**more pleasant
and convenient**
cancer screening
option



Successful
screening requires
both **adherence**
and **performance**



1. AACR Study, Cancer Prev Res (2020) 13 2. Singal AG, Gupta S, Skinner CS, et al. Effect of Colonoscopy Outreach vs Fecal Immunochemical Test Outreach on Colorectal Cancer Screening Completion: A Randomized Clinical Trial. JAMA. 2017;318(9):806–815. doi:10.1001/jama.2017.11389 3. Cancer Prevention Research. Impact of Patient Adherence to Stool-Based Colorectal Cancer Screening and Colonoscopy Following a Positive Test on Clinical Outcomes, <https://aacrjournals.org/cancerpreventionresearch/article/14/9/845/666815/Impact-of-Patient-Adherence-to-Stool-Based> [aacrjournals.org]. ACS Colorectal Cancer Facts 2020-2022, 4. EXACT Sciences presentation at Goldman Sachs 44th Annual Global Healthcare Conference, June 2023 5. ACS Colorectal Cancer Facts 2020-2022

Shield addresses existing challenges to CRC screening

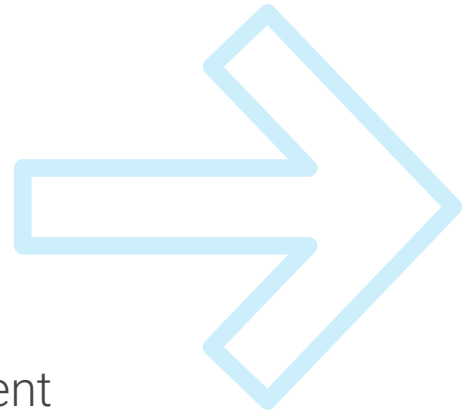
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Challenges in CRC screening¹⁻³

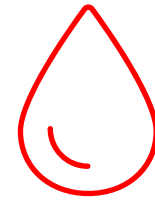
Long Colonoscopy wait times

Unpleasant experience

Time consuming, multi-step patient participation



Addressed by Shield



Simple blood draw

A **new category** of test will address the unmet need for patient preference



7 out of 10

Individuals who had a stool test would **NOT** choose stool again¹



5 to 1

Unscreened individuals **prefer blood** over stool¹



90%

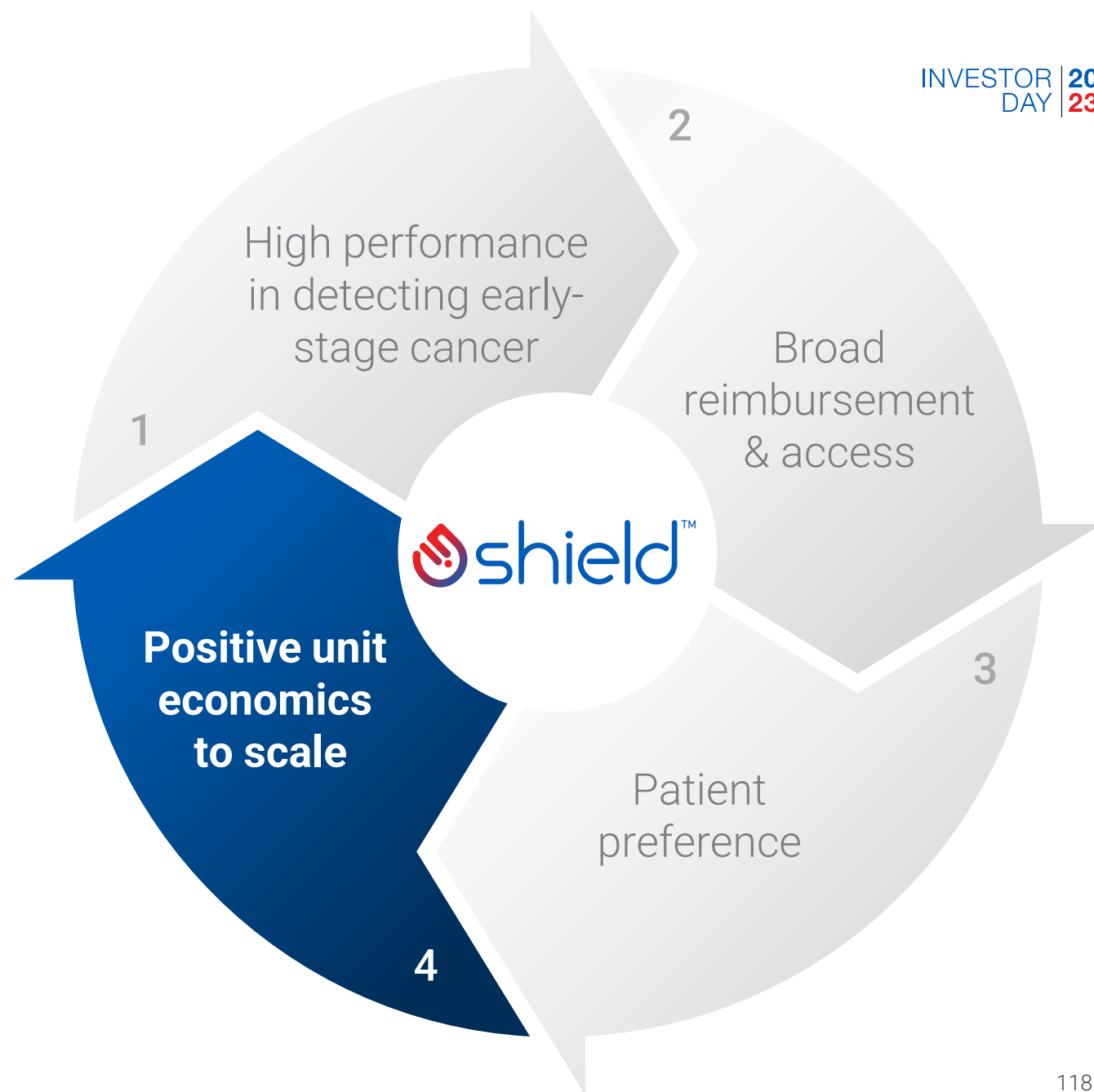
Patient adherence to Shield LDT in >10K patients across >1,000 clinics²

Blood-based screening is patient preferred and fits **seamlessly into routine care**

Screening eligible individuals regularly engage with healthcare providers



Our technology, combined with a patient preferred modality, allows for a **scalable business**



Favorable gross margin pathway enabled by Medicare ADLT pricing



\$500 **ASP**

One-year post-FDA approval, enabled by favorable ADLT pricing for Medicare beneficiaries

\$200 **COGS**

When processing 1 million samples per year, enabled by Shield technology and high-throughput automation

High adherence and patient preference drives ~40% increased S&M productivity

		
Orders Placed	100	100
Adherence	65% ¹	90% ²
Orders Resulted	65	90

Targeted commercial strategy for Shield launch in 2024



High concentration of Medicare lives



States with coverage mandates post ACS guidelines



Strong utilization of non-invasive screening



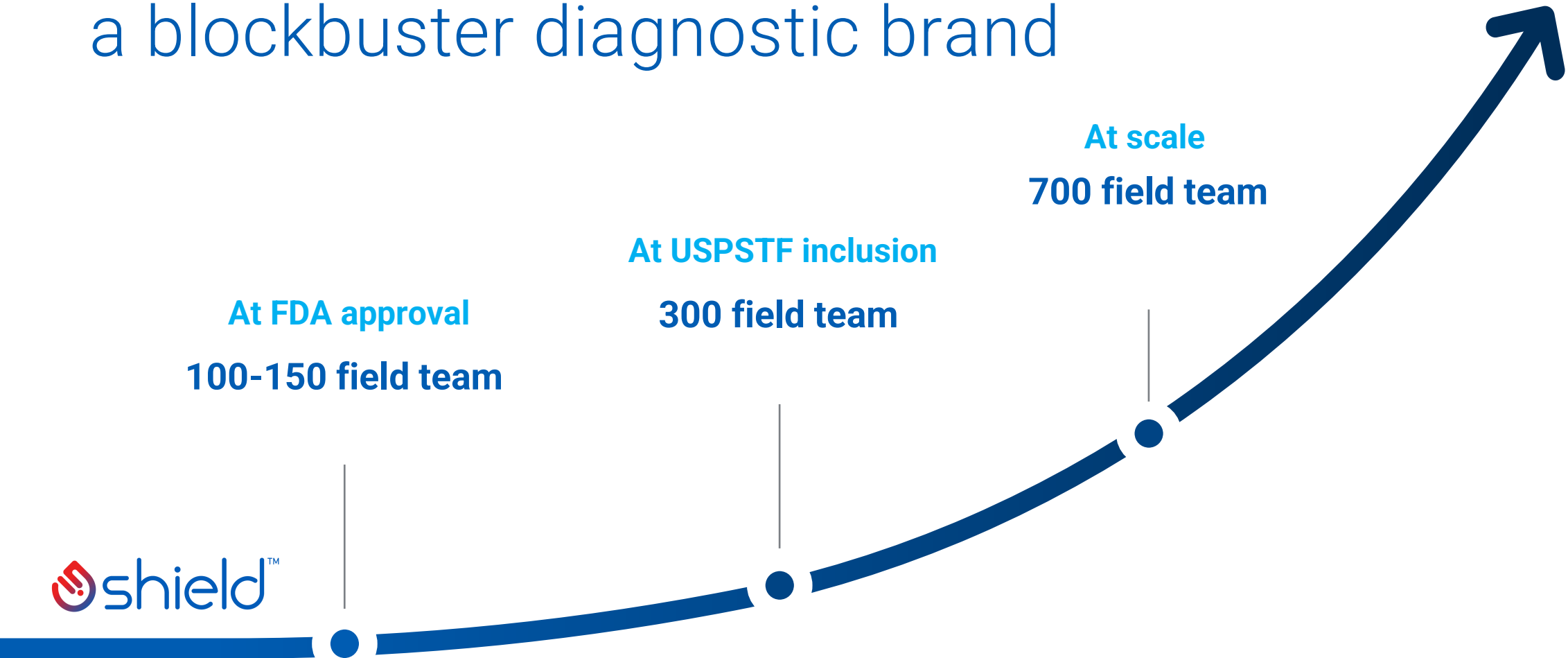
Phlebotomy access

Maximize ASP

Maximize efficiency & productivity

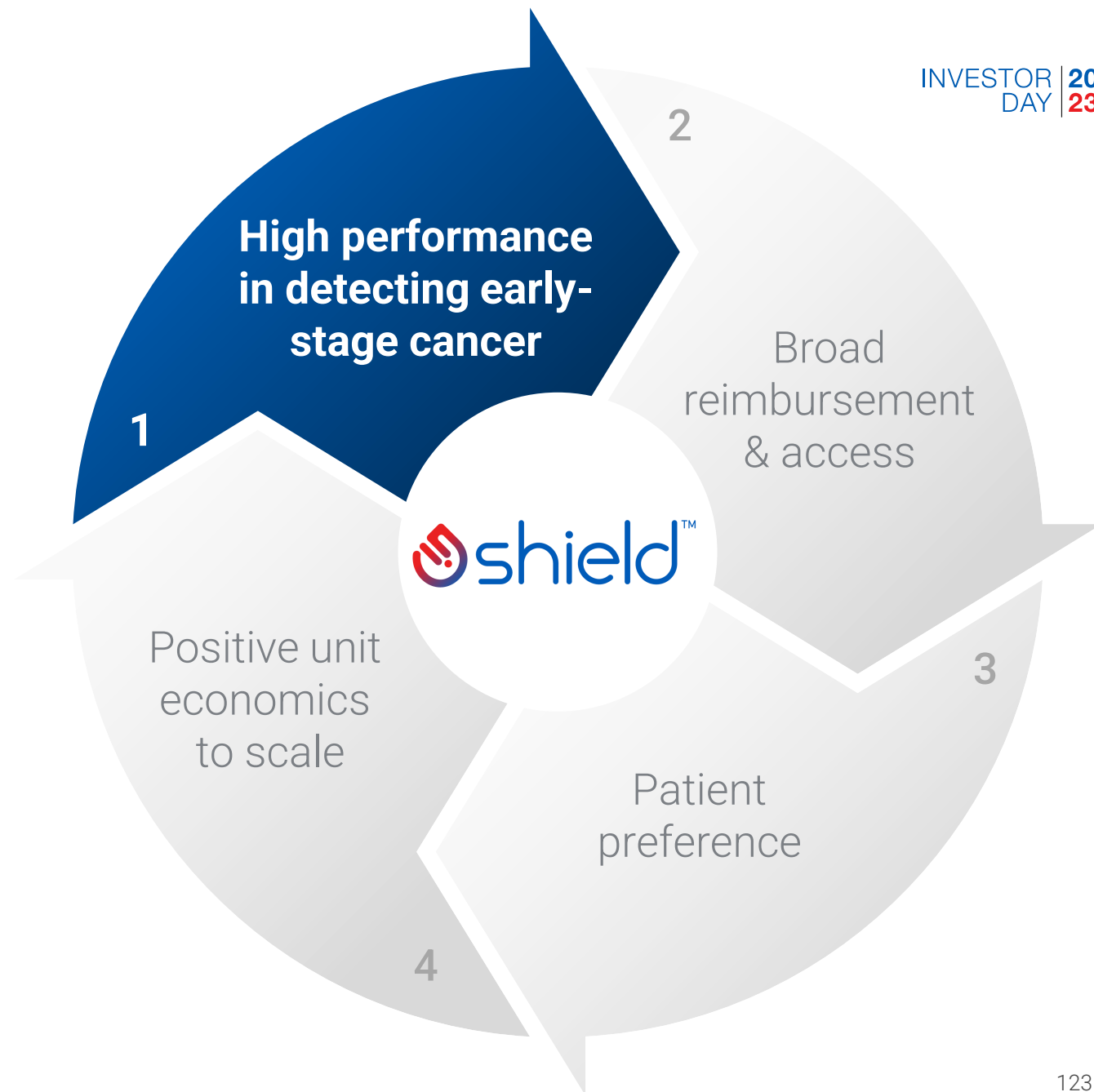
Milestone-gated commercial roadmap to a blockbuster diagnostic brand

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shield™

Scalable
commercial
growth enables
**continuous
improvement**
through data,
biobank, clinical
insights and real-
world evidence



Upgraded algorithm with demonstrated **improved analytical sensitivity** in detecting tumor signals at ~2x lower levels



Version 2

New data

Shield V2 clinical data

Clinical sensitivity of Shield V2 in a U.S.-based colonoscopy-screened cohort

Prospective Study Results

	CRC Sensitivity N = 45	Specificity N = 1,458
Shield CRC V1 – PMA device	84% (38/45)	91%
Shield CRC V2	91% (41/45)	91%

	CRC Sensitivity N = 65	Specificity N = 6,680
ECLIPSE	83%	90%

Sensitivity in Stage I/II:

V1: 76% (19/25)

V2: 88% (22/25)

Similar AA performance

Continuous improvement of Shield IVD – best-in-class blood-based CRC screening



**Submit V2 supplemental PMA
following approval of Shield***

Potential for V2 approval in **2025**

Next step to Multi-Cancer is adding lung indication





Same test. Simple software upgrades.
Enables **new cancer indications.**

Reinventing screening for lung cancer with a simpler and accessible option

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150K

Lung cancer deaths per year, leading cause of cancer-related death¹

86%

Of eligible individuals are unscreened¹

12M

Unscreened individuals²

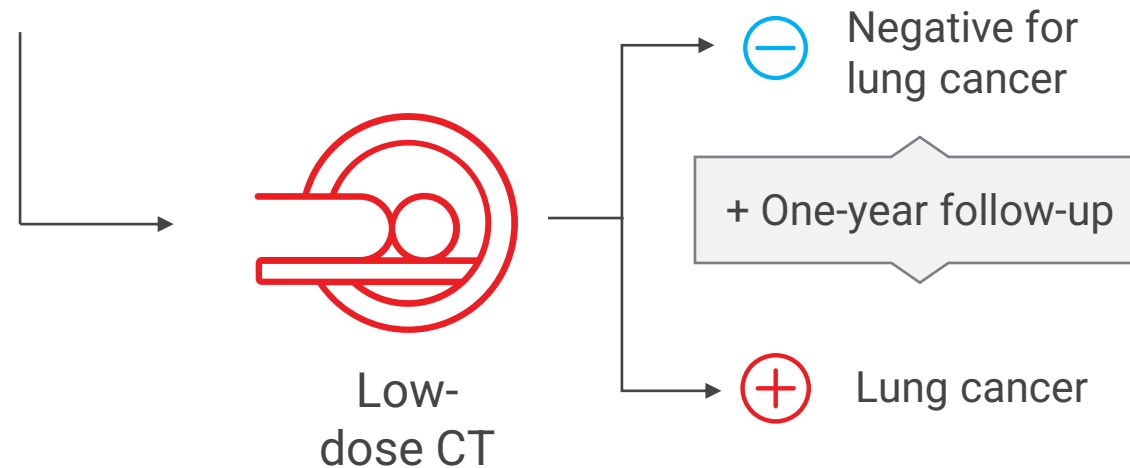
On track to
complete
Shield-Lung
enrollment in
2025



>7K

Currently enrolled

Enrolling individuals who meet
USPSTF Criteria
for High-Risk Lung Cancer
Screening¹



First endpoint readout in **2026**

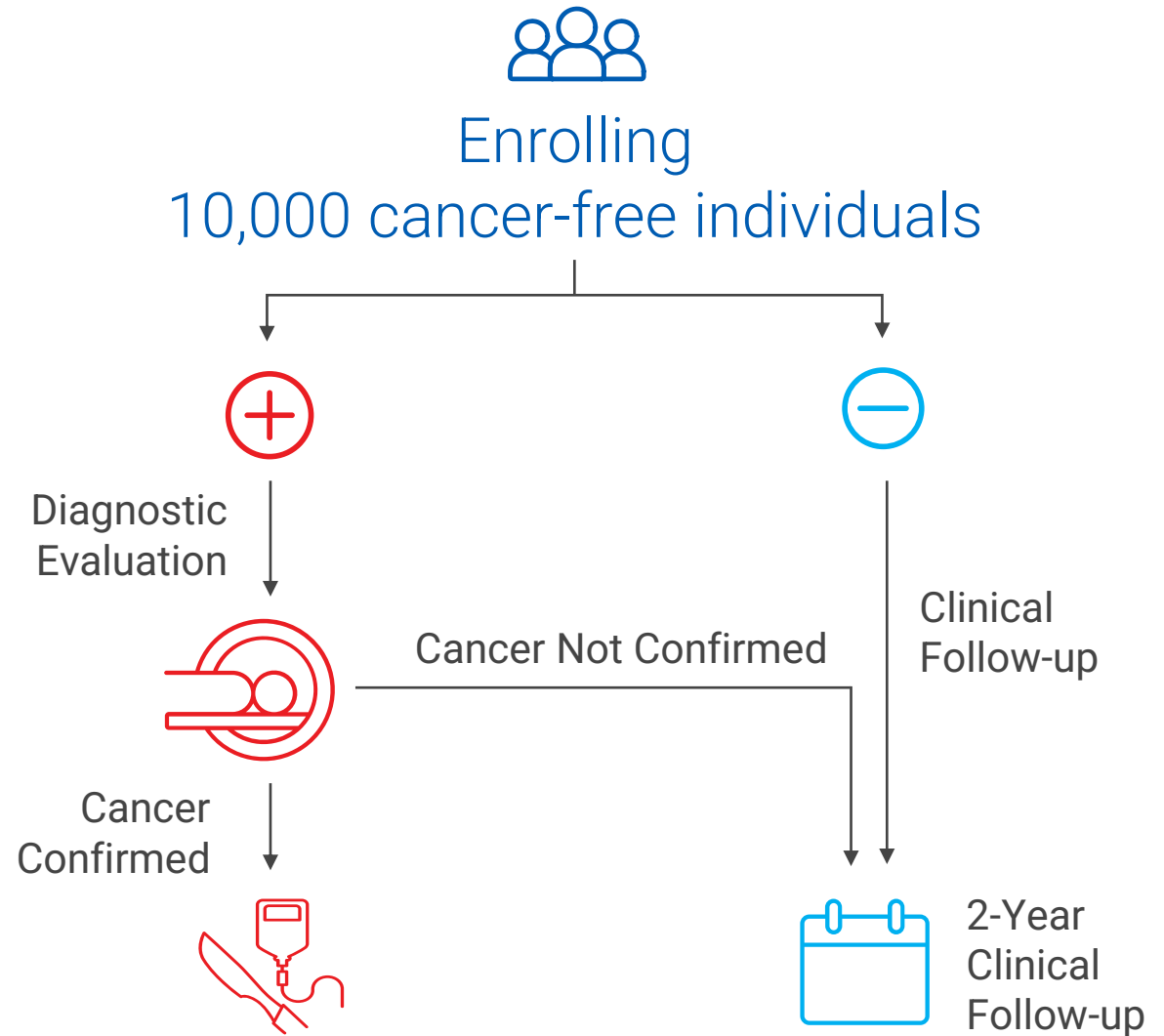
1. Nguyen, et al. 2023. Screening for high frequency malignant disease (SHIELD) J Clin Oncol 41, 2023 (suppl 16; abstr TPS1610). 2. <https://clinicaltrials.gov/study/NCT05117840>

Prospective interventional multi-cancer detection study



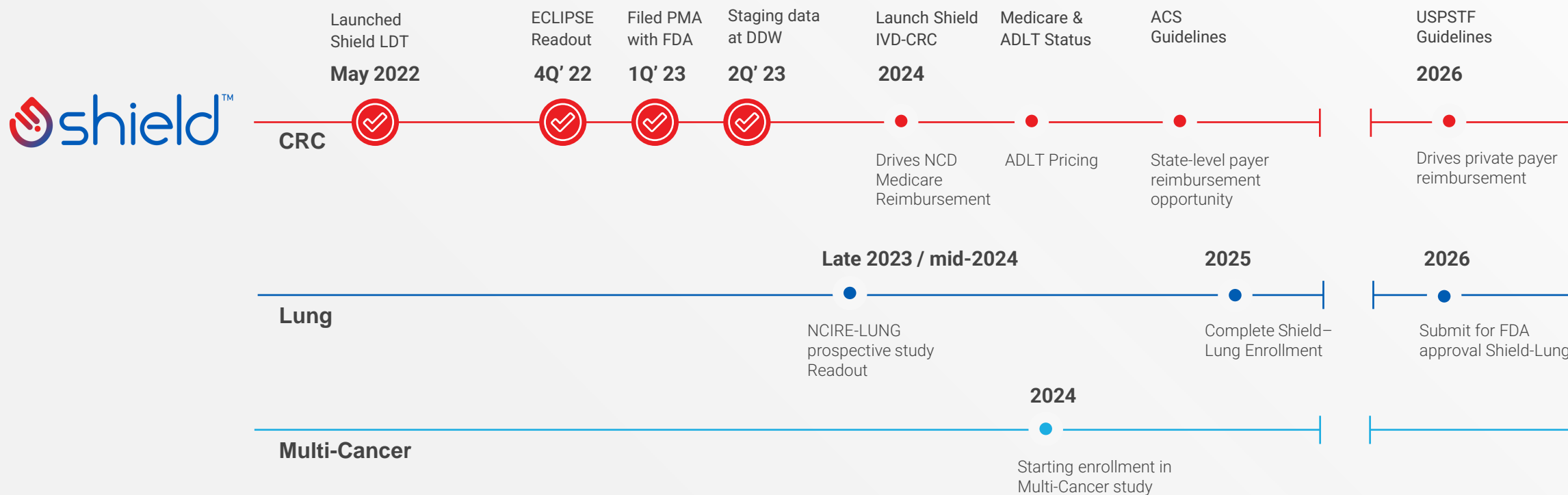
2H 2024

Expected first patient
enrollment



Blazing a trail to the first FDA approved and reimbursed Multi-Cancer test

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A once in a lifetime
opportunity to **reinvent
cancer screening**



Seeing deeper with **epigenomics**

Dr. Justin Odeggaard

Vice President, Product Development

Research Focus

Epigenomics

Single-cell method development

High-throughput biochemistry

Bioinformatics for understanding gene regulation



Dr. William Greenleaf

Professor of Genetics, Stanford University

Research Focus

Epigenetic cfDNA-based biomarkers for advanced cancer patients

Epigenetic mechanisms of therapeutic response and resistance in Prostate Cancer



Dr. Jacob Berchuck

Assistant Professor of Medicine, Harvard Medical School

Research Focus

Innovative biomarkers in lung cancer

Liquid-biopsy assay development

Data science



Dr. Simon Heeke

Assistant Professor, MD Anderson Cancer Center

Research Focus

Diabetes and pancreatic cancer

Epigenetic liquid biopsies

DNA methylation clocks



Dr. Yuval Dor

Professor, The Hebrew University of Jerusalem

Today's panelists

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Dr. Will Greenleaf
Professor



Dr. Jacob Burchuck
Assistant Professor



Dr. Simon Heeke
Assistant Professor



Dr. Yuval Dor
Professor



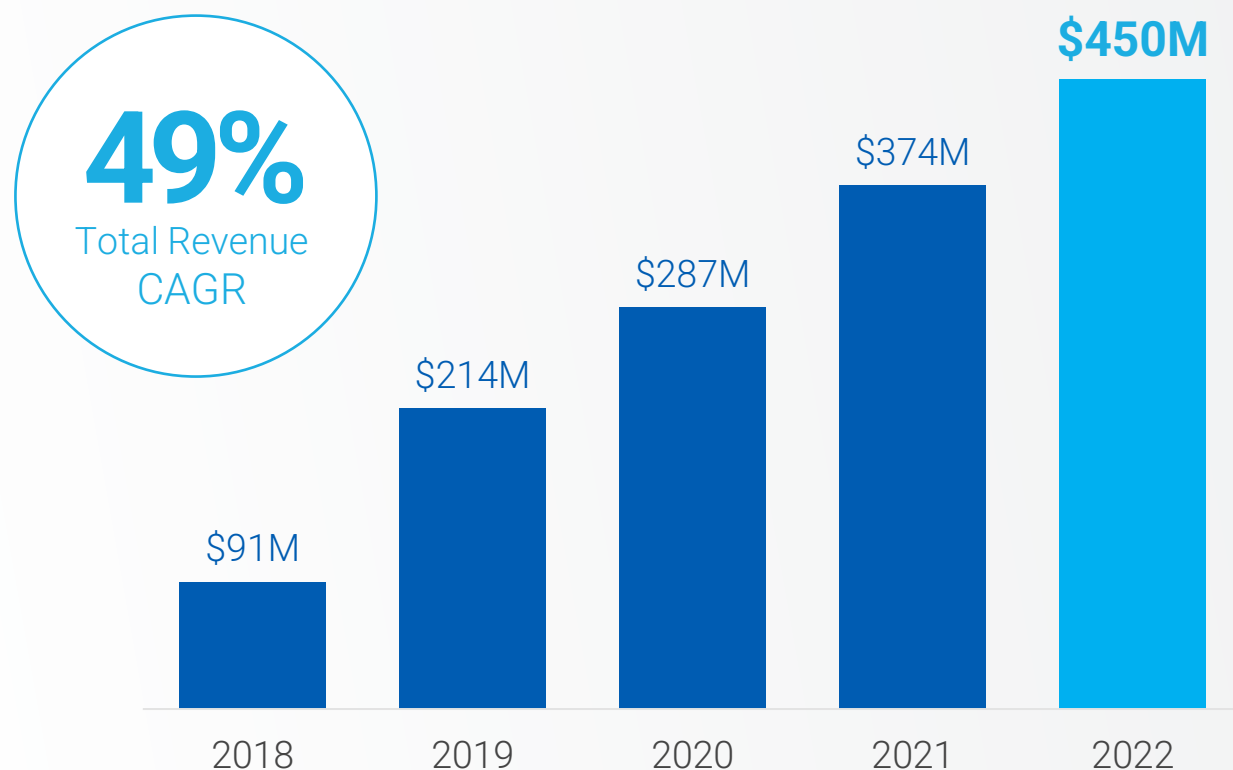


Balancing rapid revenue growth with **financial discipline**

Mike Bell

Chief Financial Officer

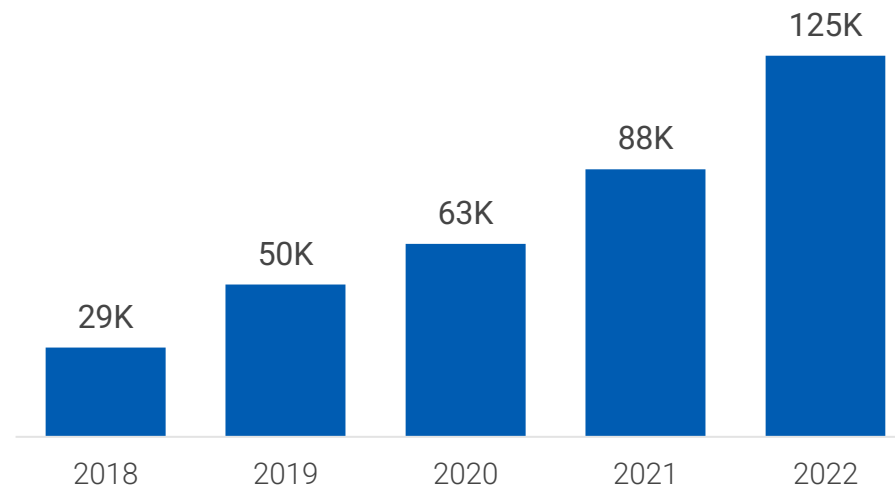
Historical revenue growth fueled by core Therapy Selection business



Therapy Selection growth drivers

- ✓ Reimbursement coverage
- ✓ Regulatory approvals
- ✓ Clinical data
- ✓ Commercial strength
- ✓ Biopharma partnerships
- ✓ Portfolio expansion

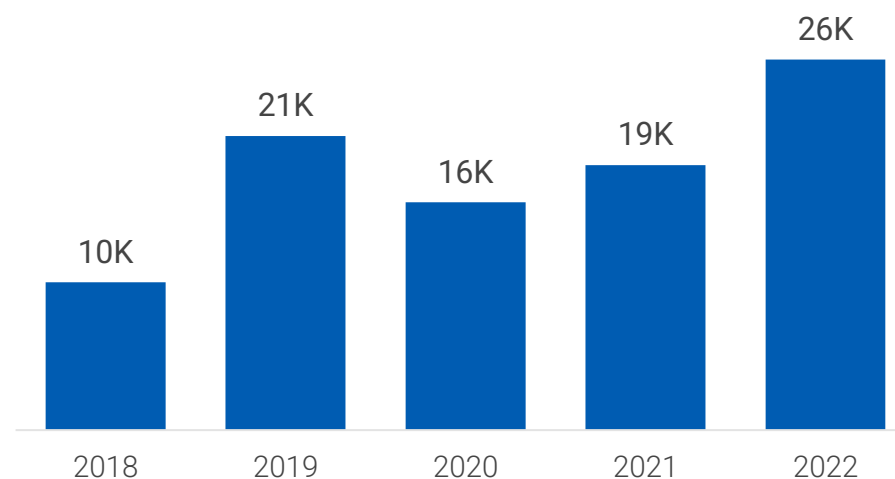
Guardant360 has
been the primary
**volume growth
driver**



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44%

Clinical Testing
Volume CAGR

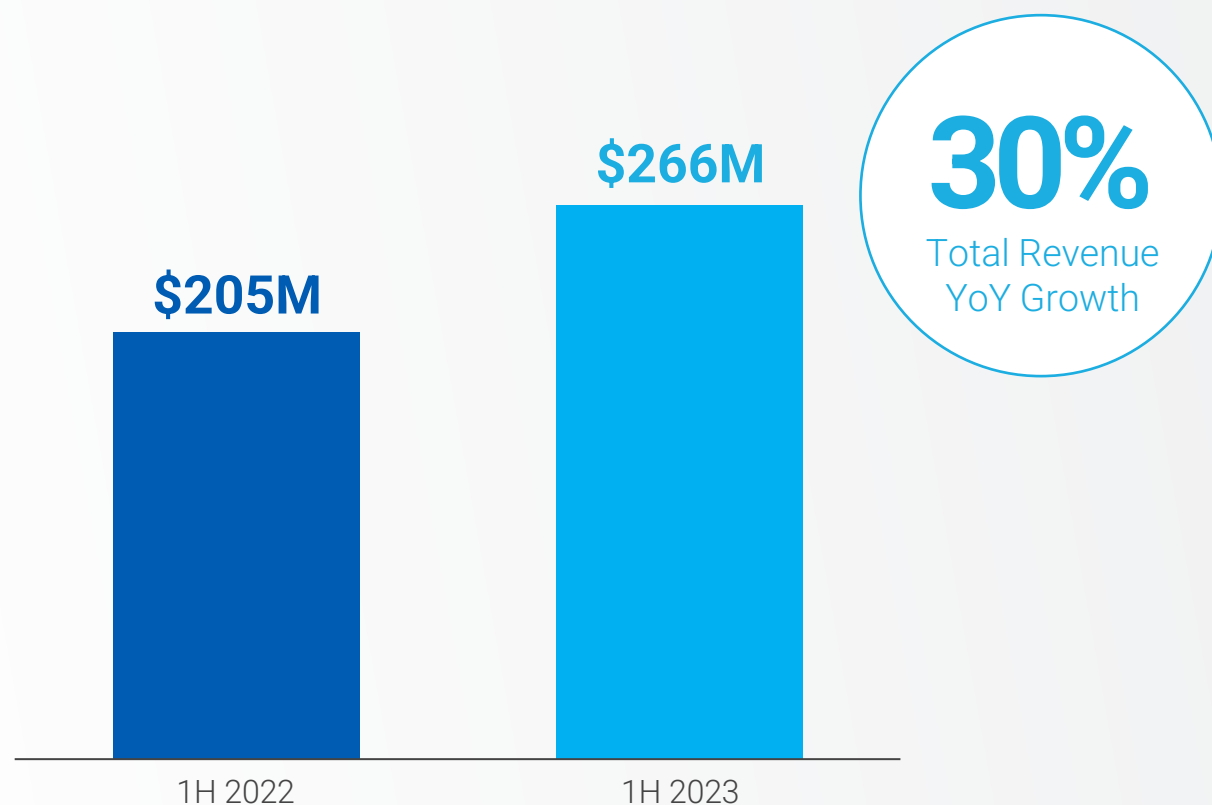


26%

Biopharma Testing
Volume CAGR

Strong revenue growth and major Therapy Selection tailwinds in 1H 2023

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23



Recent Highlights

- ✓ Guardant360 CDx breast FDA approval
- ✓ Guardant360 expanded reimbursement
- ✓ Guardant360 Japan reimbursement
- ✓ Response Medicare coverage
- ✓ EMR integrations

Therapy Selection has multiple growth drivers well beyond 2023



Portfolio ASP
increases

GUARDANT **360** CDx

Future Guardant360
CDx approvals

GUARDANT **360** TissueNext[™]
GUARDANT **360** Response[™]

TissueNext &
Response attachment



Market expansion

GUARDANT **INFINITY**

Infinity offering to
biopharma partners



International expansion,
e.g., Japan, UK, China

Leveraging existing infrastructure will enable Therapy Selection profitability



Commercial

- 380+ oncology commercial team
- 160+ biopharma partnerships



Regulatory

- Proven IVD and reimbursement expertise
- Robust, well-established quality systems



Operations

- Scaled logistical and operational backbone
- Existing lab capacity for future growth



Research & Development

- Clinical data development engine
- Unified platform strategy



International

- In-country operations
- Top-tier ex-US partners: UK, Spain, China

Therapy Selection on track to be cash flow **breakeven by YE 2023**

FY 2023 financial targets focused on revenue growth and financial discipline

INVESTOR
DAY | 20
23

Revenue

\$545 – \$550M

21% - 22% y/y
growth

OpEx

< FY 2022

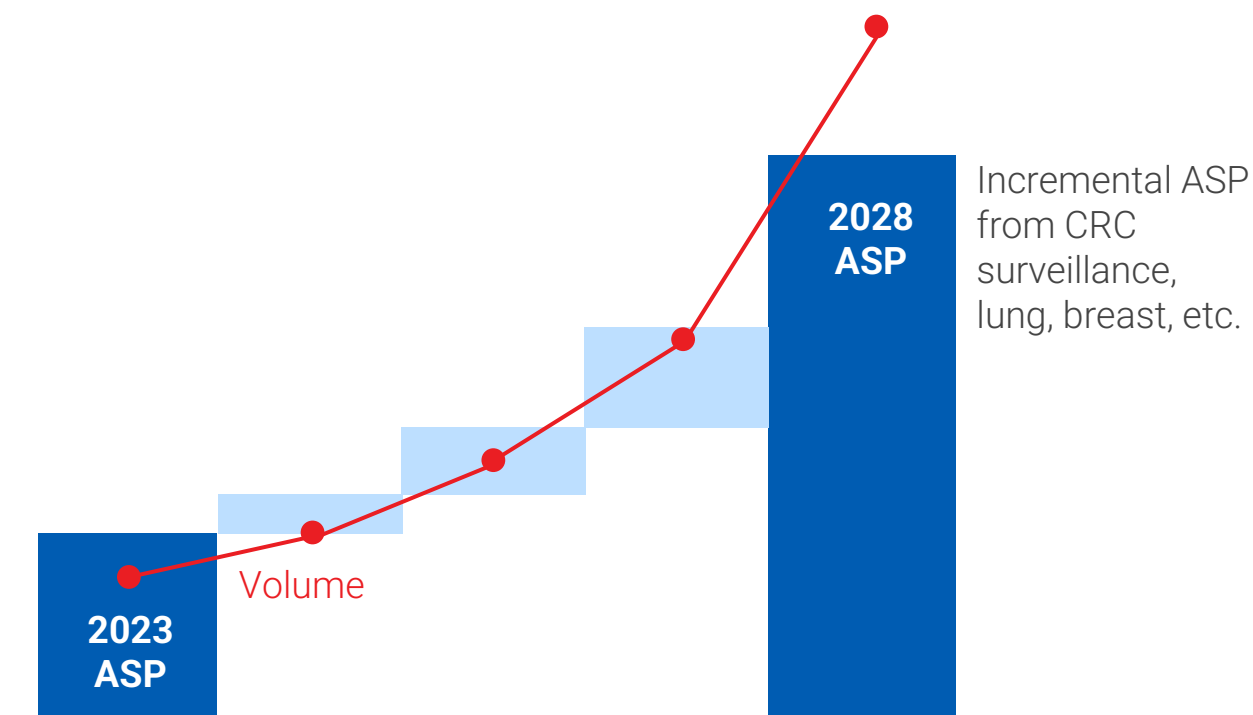
Leveraging existing
infrastructure

Free Cash Flow

~(\$350M)

Reducing cash
burn

Focus on Reveal reimbursement to increase ASP and accelerate volumes



Highly efficient MRD Business Model leveraging existing Therapy Selection infrastructure



Commercial



Operations



Medical Affairs



Development



International

5-year financial targets for Therapy Selection & MRD



Revenue Growth

- >20% Combined Therapy Selection & MRD revenue CAGR
(Total revenue excluding Screening)



Clinical ASPs by 2028

- G360 > \$3,000
- Tissue > \$2,000
- Response > \$1,000
- Reveal > \$1,000



Gross Margins

- 60-70% Combined Therapy Selection & MRD gross margin
(Gross margin excluding screening)



OpEx & CapEx

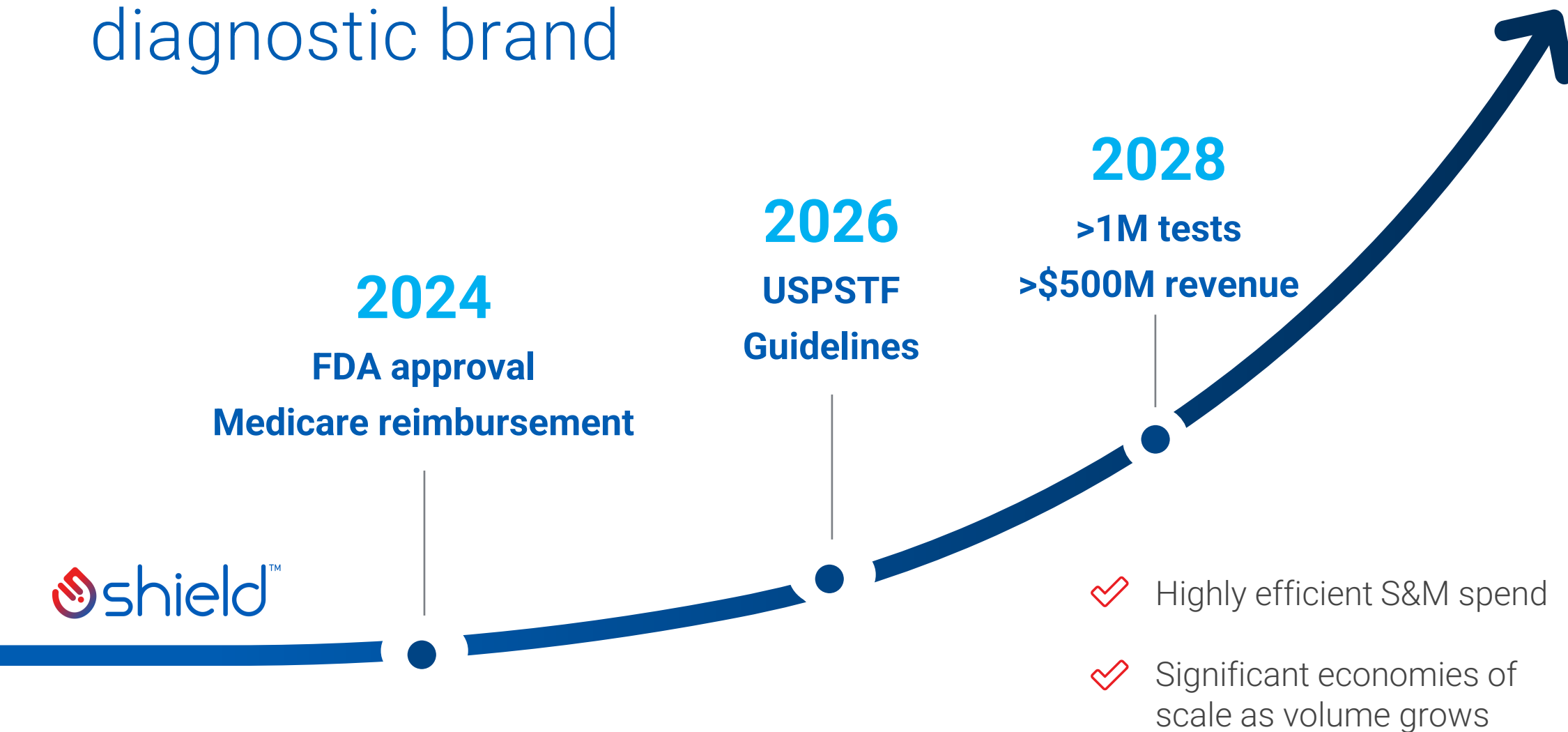
- Leverage existing infrastructure



Cash Flow

- Therapy Selection cash flow breakeven end of 2023
- Combined Therapy Selection & MRD generates positive cash flow over cumulative 5-year period
(Cash flow excluding Screening)

Shield is poised to be a blockbuster diagnostic brand



Balancing rapid Shield growth with measured spend over next 5 years

Annual **cash burn ~\$200M** over 5 years



ASP > \$500

- Medicare ADLT rate
- Incremental private payer coverage based on ACS recommendations
- Full private payer coverage following USPSTF guidelines



Gross Margins

- Positive 1 year post launch
- ~60% at 1M annual volume



CapEx

- Low cost, high volume, automated lab operations
- Phased CapEx investments



R&D

- Lung and MCED
- Leverage Smart Liquid Biopsy platform



SG&A

- Efficient sales & marketing model / milestone gated investments
- Leverage Guardant back-office infrastructure

5-year total revenue targets

>30%

5 Year CAGR

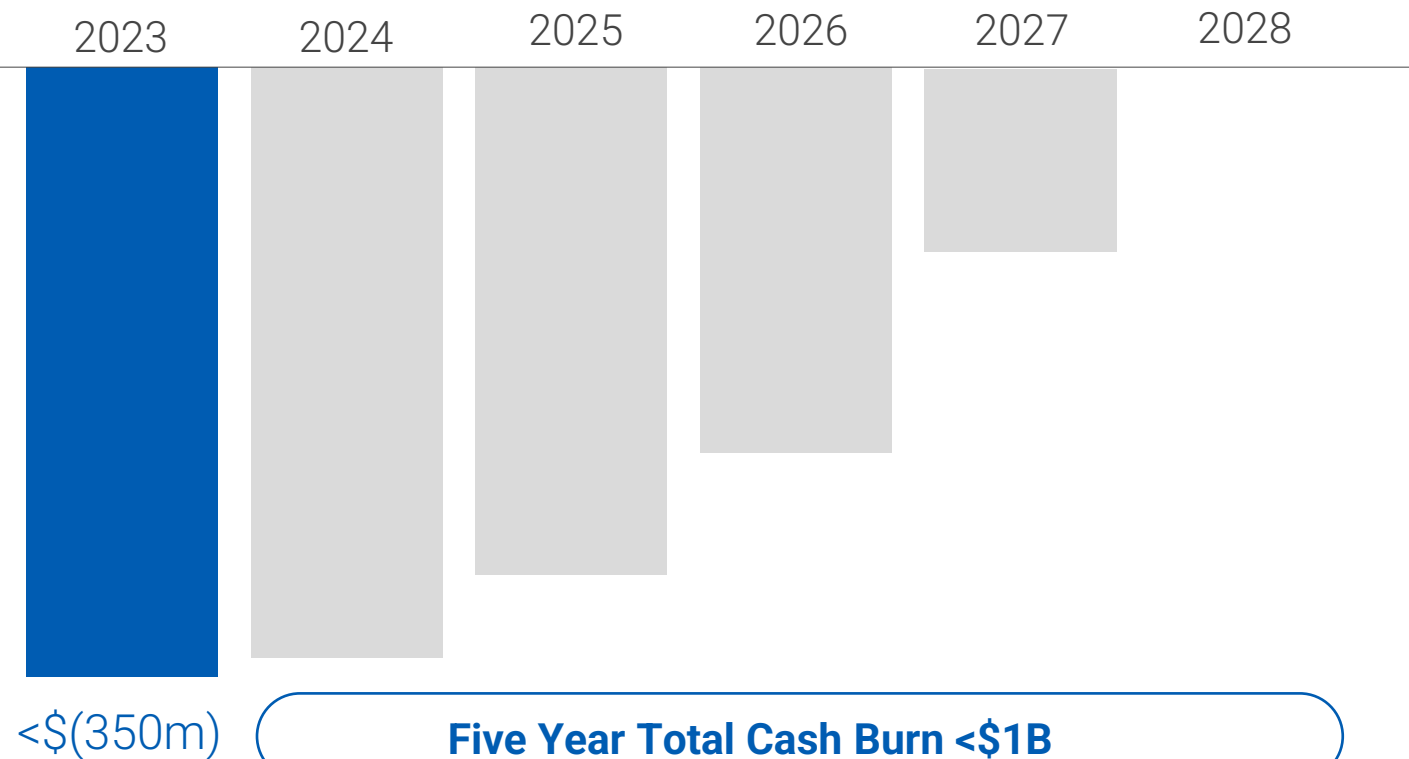
>\$2B

In 2028

Current cash
provides runway
to reach **cash flow
breakeven**

YE Cash
>\$1B

INVESTOR DAY | **20**
23
Cash Flow
Breakeven



- Therapy Selection breakeven end of 2023
- Combined Therapy Selection + MRD generates positive cash flow over cumulative 5-year period
- Screening ~\$200m annual cash burn

Key financial targets



Grow profitable Therapy Selection business



Expand **MRD** reimbursement to **increase ASP and accelerate volumes**



Balance rapid Screening revenue growth with **measured** spend



Achieve **>30% total revenue 5-year CAGR** and **>\$2B** in 2028



Reach cash flow **breakeven** by end of 2028



The future of transforming cancer care **and beyond...**

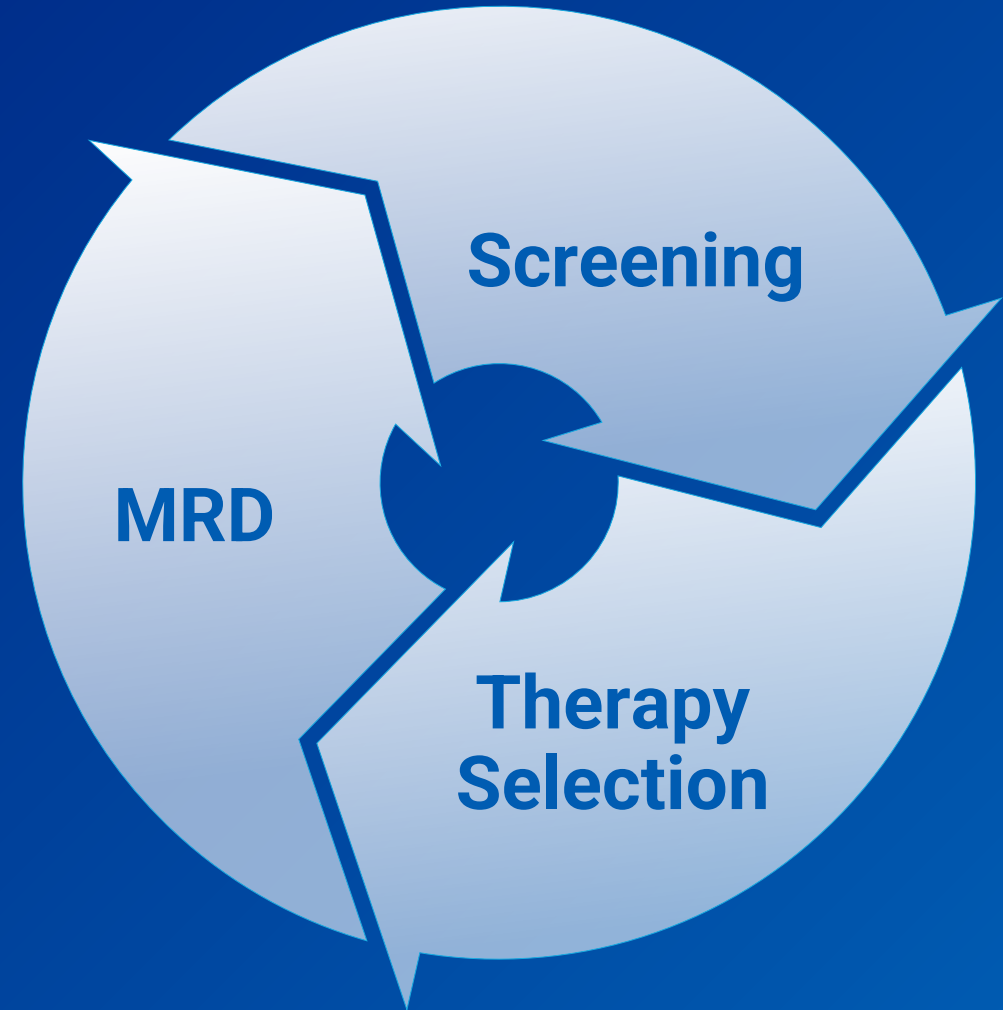


Helmy Eltoukhy, PhD
co-Chief Executive Officer & Chair

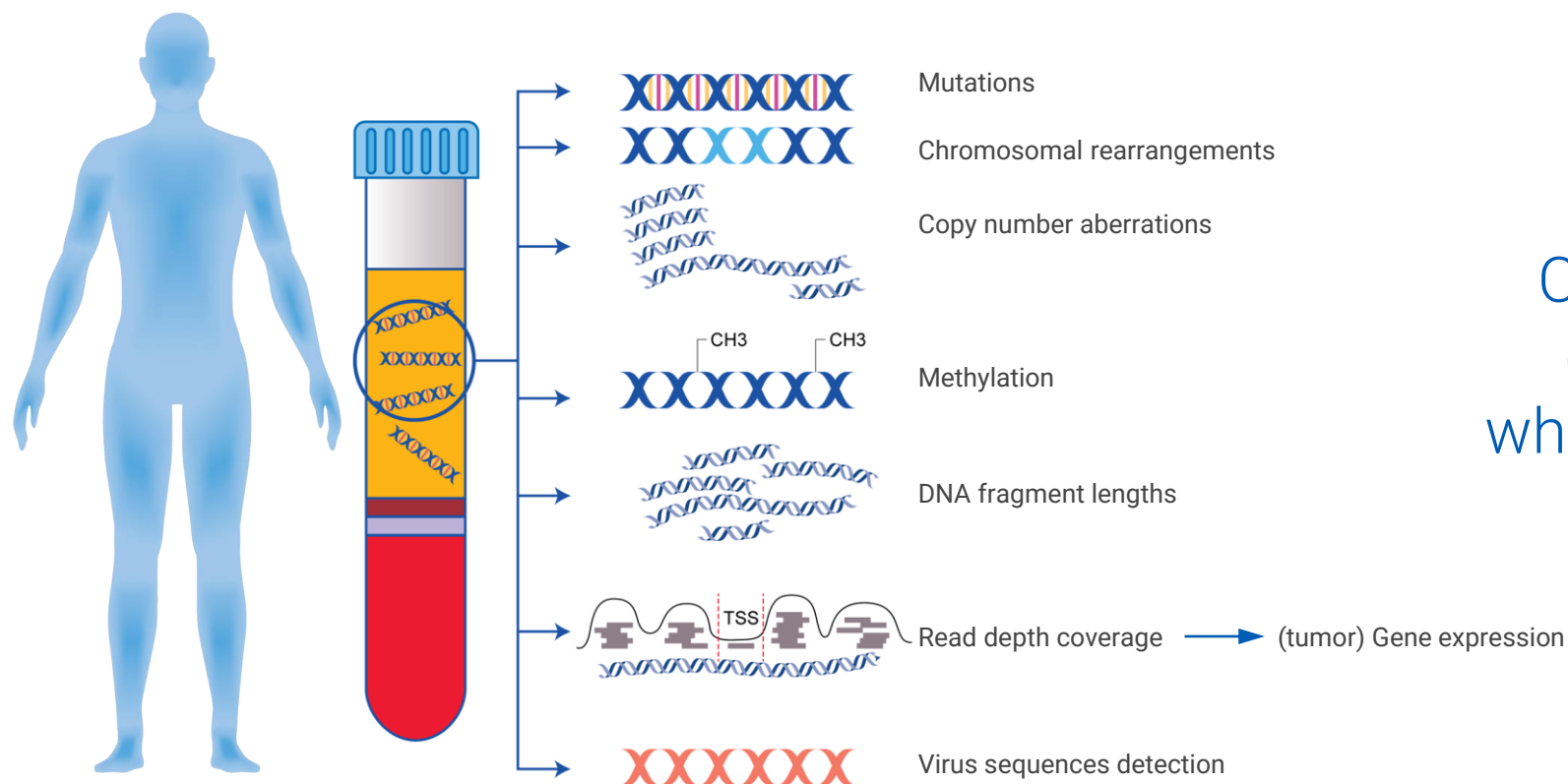
AmirAli Talasaz, PhD
co-Chief Executive Officer & co-Founder

Cancer is not a
linear journey

Our ecosystem of
tests will
**synergistically
work together**



Biologically rich data



Comprehensive longitudinal
characterization of tumor,
whole body epigenomic profiles
and immune response

GUARDANTINFORM™

>90%

of commercial samples
are linked to associated
medical claims data¹

Comorbidities



Disease progression



Lines of therapy



Time to next treatment



Treatment history



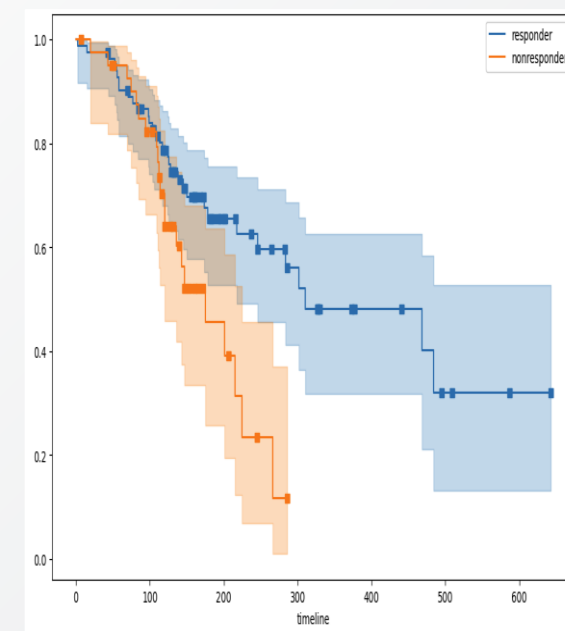
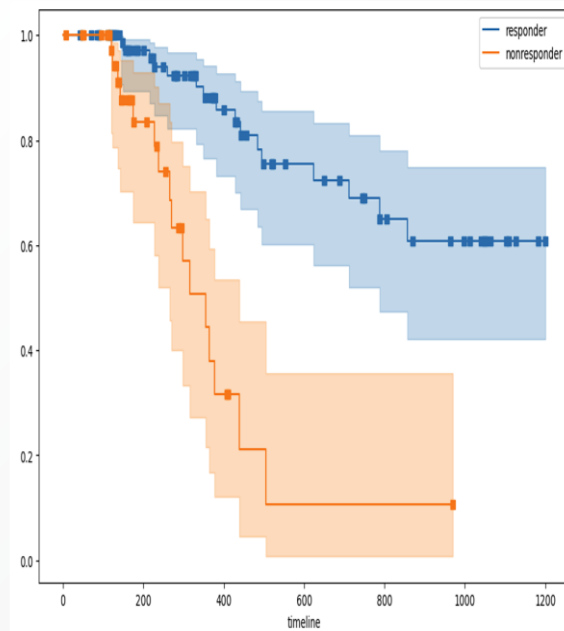
Treatment discontinuation



Survival

The power of
genomic and
epigenomic data
is amplified by
connections to
**real-world
evidence**

Real-World Analysis Shows Utility of ctDNA Kinetics in Patients With Advanced Colorectal Cancer



Kaplan-Meier in molecular responders vs non-responders for patients receiving chemotherapy +/- VEGF.

Real-world evidence from GuardantINFORM has been successfully used to validate novel signatures

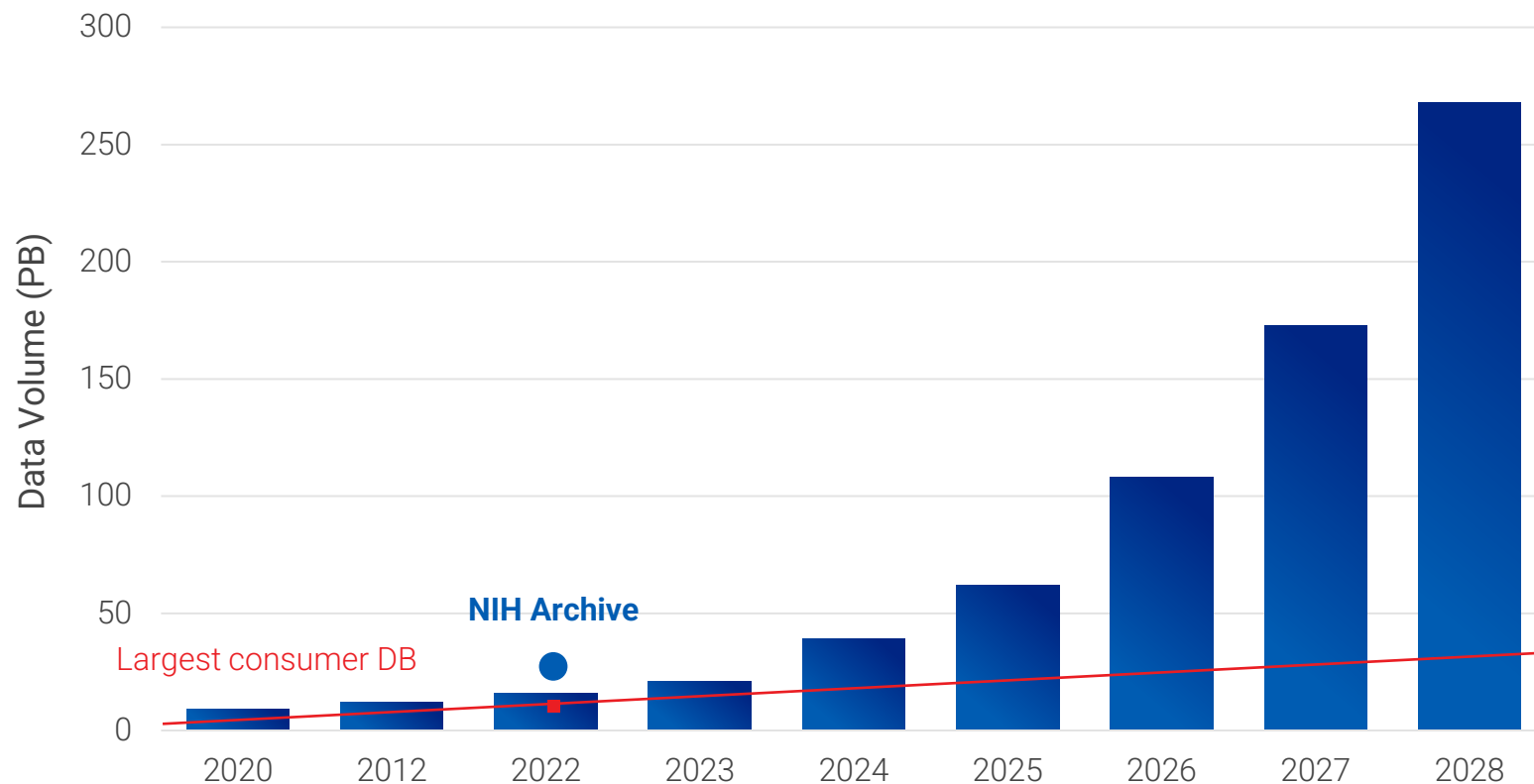
Data generation at scale

- ✓ Single Platform
- ✓ Scalable
- ✓ Cost-Efficient
- ✓ Configurable



Expecting to sequence millions of clinical samples per year across **average risk population** and **patients impacted by cancer**

Data growth powering research

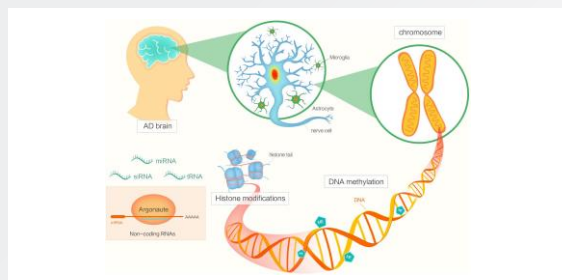


**18-month
doubling rate**

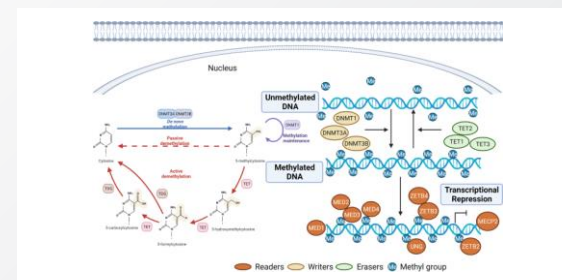
Guardant data growth is
comparable to that of the largest
NIH archive of genomic data

Almost every disease has a visible fingerprint when viewed through the lens of epigenomics

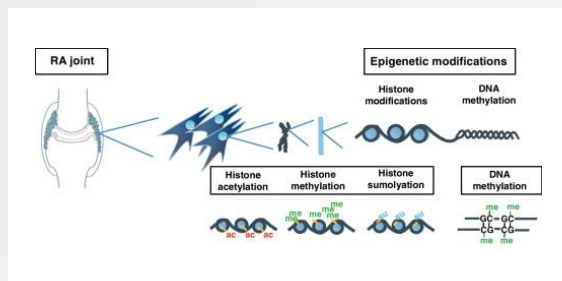
Neurodegenerative Disease



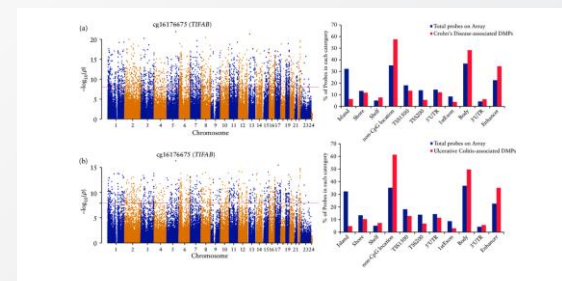
Cardiovascular Disease



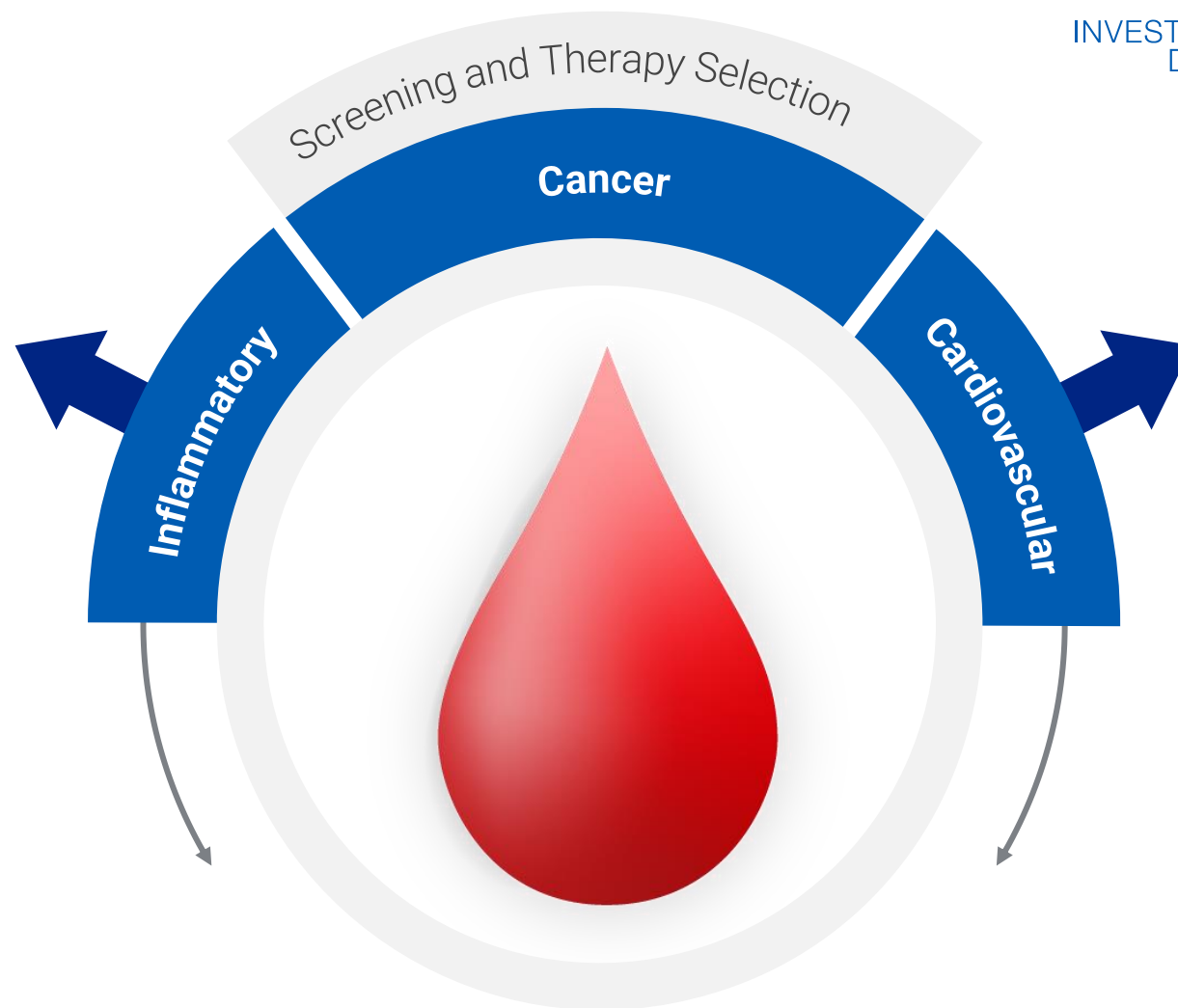
Rheumatoid Arthritis



Inflammatory Bowel Disease



Our technology platform and commercial infrastructure can be **leveraged across multiple diseases beyond cancer** and will reshape precision medicine



The promise of precision medicine will be accessed through a single comprehensive blood test

Our continued success
will expand our
mission over time
from one of
conquering cancer to
that of **guarding
wellness**



Q&A