



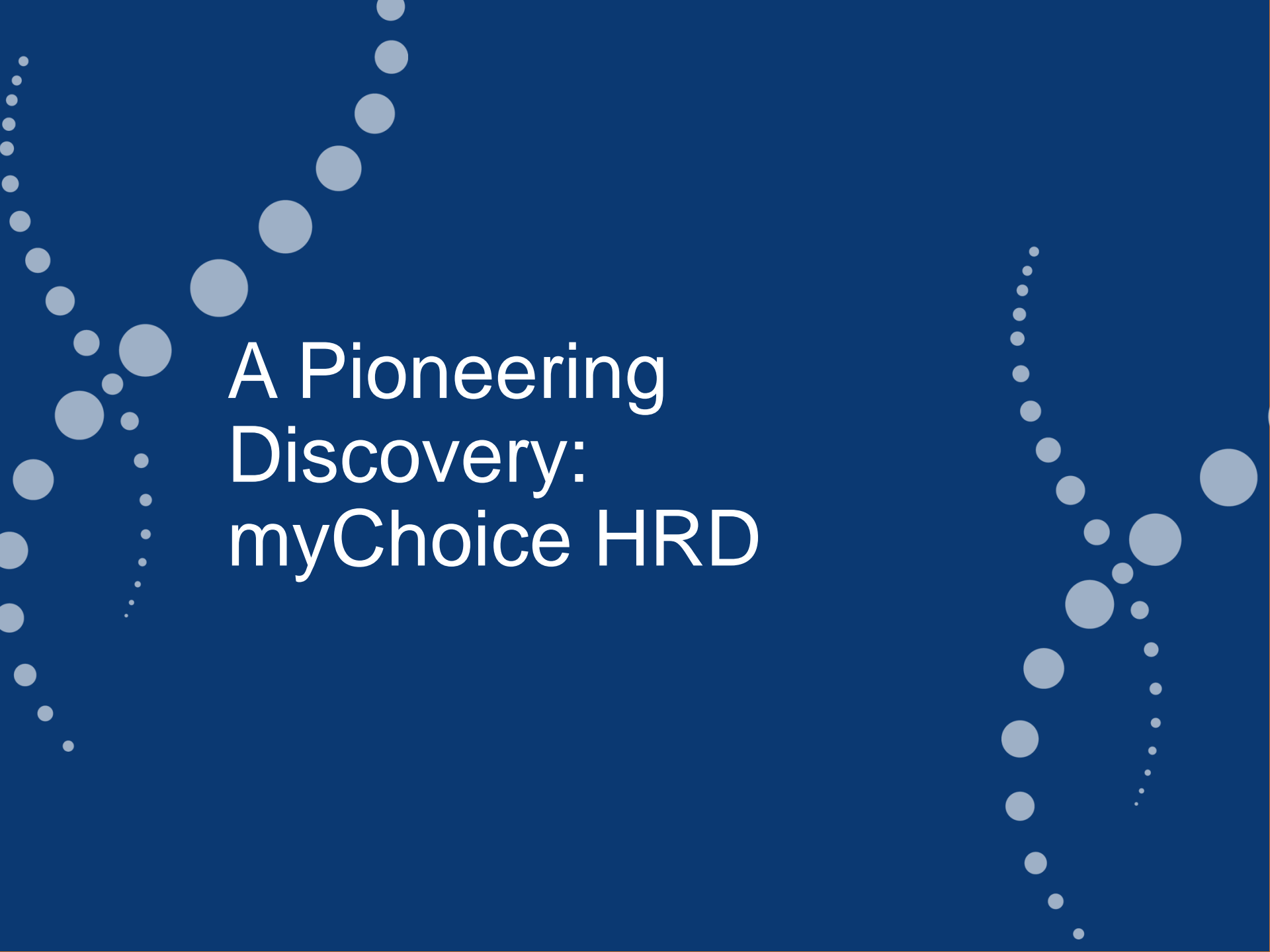
# **Myriad Genetics** **myChoice HRD<sup>®</sup> Update**

06/30/2016



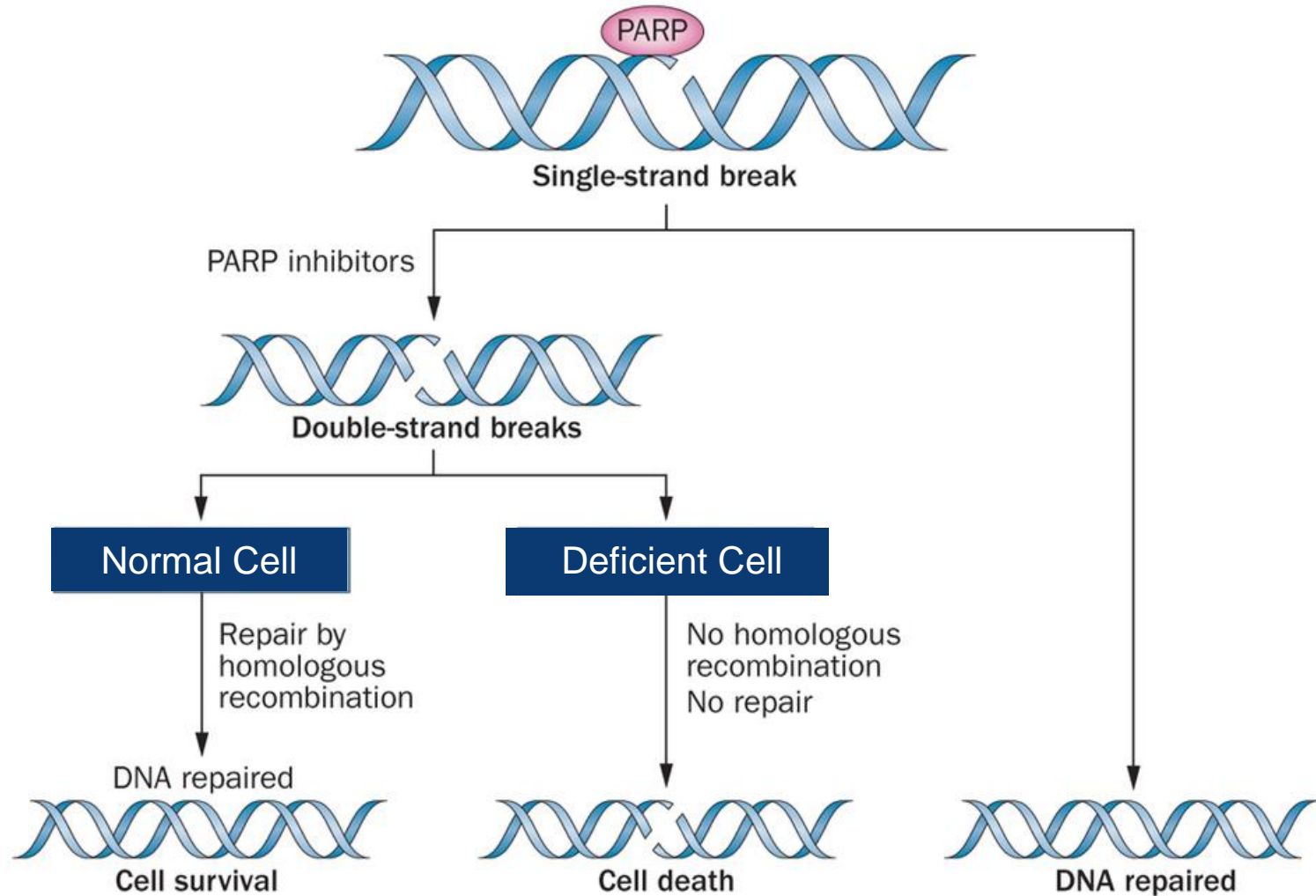
# Forward Looking Statements

Some of the information presented here today may contain projections or other forward-looking statements regarding future events or the future financial performance of the Company. These statements are based on management's current expectations and the actual events or results may differ materially and adversely from these expectations. We refer you to the documents the Company files from time to time with the Securities and Exchange Commission, specifically, the Company's annual reports on Form 10-K, its quarterly reports on Form 10-Q, and its current reports on Form 8-K. These documents identify important risk factors that could cause the actual results to differ materially from those contained in the Company's projections or forward-looking statements.

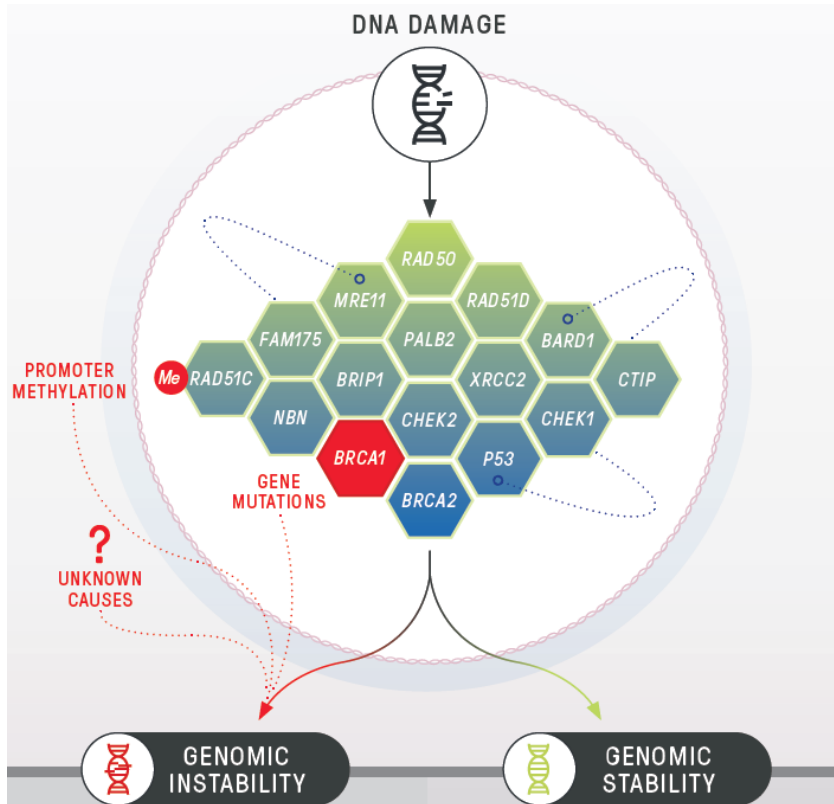


# A Pioneering Discovery: myChoice HRD

# PARP Mechanism of Action

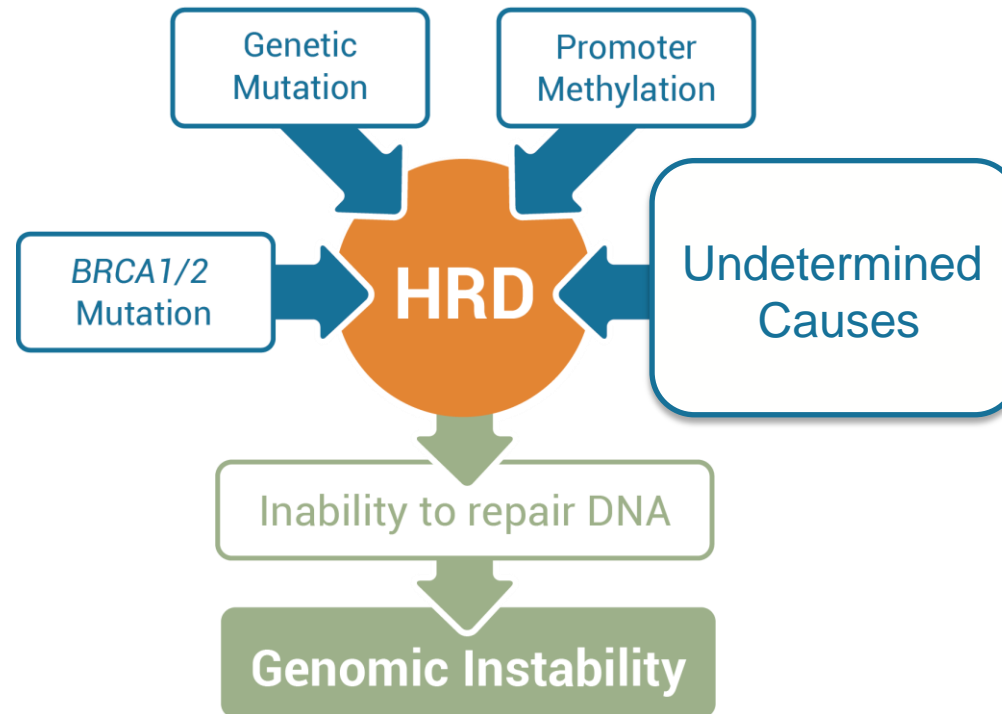


# Homologous Recombination (HR) Pathway Status Predicts Drug Response



- When the HR pathway is working properly, DNA can be repaired effectively and is error free, maintaining genomic stability
- When the HR pathway is disrupted by gene mutations, promoter methylation or unknown causes, the HR pathway stops working leading to genomic instability or homologous recombination deficiency (HRD)
- Patients who have a defective pathway are much more likely to respond to drugs that impact DNA stability such as platinum drugs and PARPs

# Most Causes of Homologous Recombination Deficiency (HRD) Are Unknown



- When the HR pathway is disrupted by **gene mutations**, **promoter methylation** or **unidentified causes**, the HR pathway stops working leading Homologous Recombination Deficiency or HRD
- Tumors with HRD are unable to repair themselves effectively after sustaining damage, leading to genomic instability

# Pioneering Discovery: Measure the Effects vs. the Causes

## The EFFECT.....



## The Cause?

Lane closure?  
Traffic light failure?  
Overturned truck?  
Fender bender?  
Presidential motorcade?

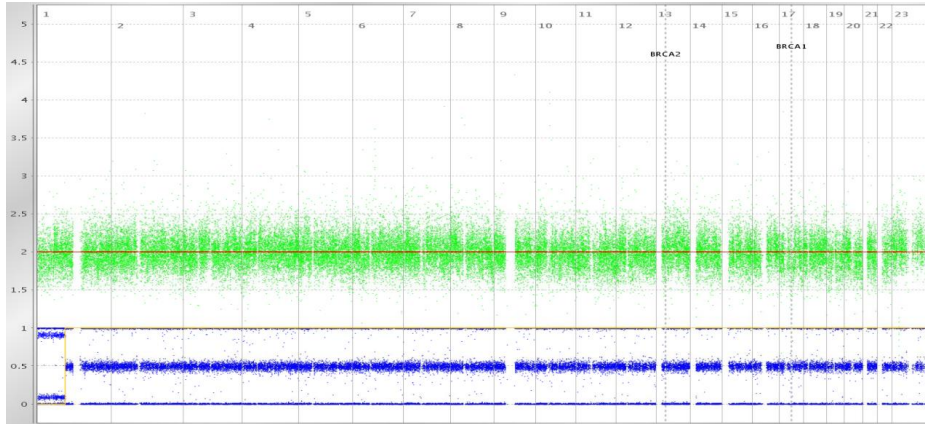
## Who cares....





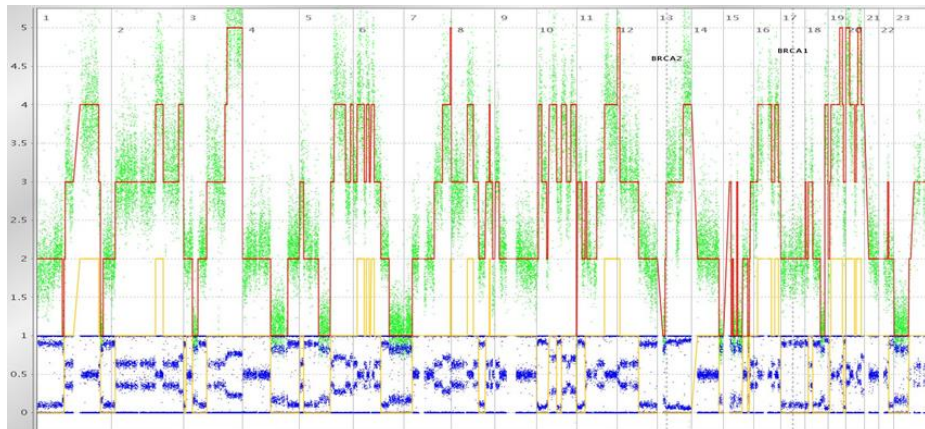
# Measuring the DNA “Scar” – With Proprietary Informatics

Genome profiles are pictures of instability in tumor DNA



myChoice HRD Negative  
(myChoice HRD Score = 3)

Cutoff = 42



myChoice HRD Positive  
(myChoice HRD Score = 81)

genomic instability  
caused by HRD



# myChoice HRD Identifies the Most Patients with Ovarian Cancer Potentially Eligible for a Drug

15% of patients are positive for BRACAnalysis CDx<sup>1</sup>



22% of patients are positive for Tumor BRACAnalysis CDx<sup>2</sup>



≈25% of patients are positive for a broad tumor HR panel<sup>3</sup>



≈50% of patients are positive for myChoice HRD<sup>4</sup>

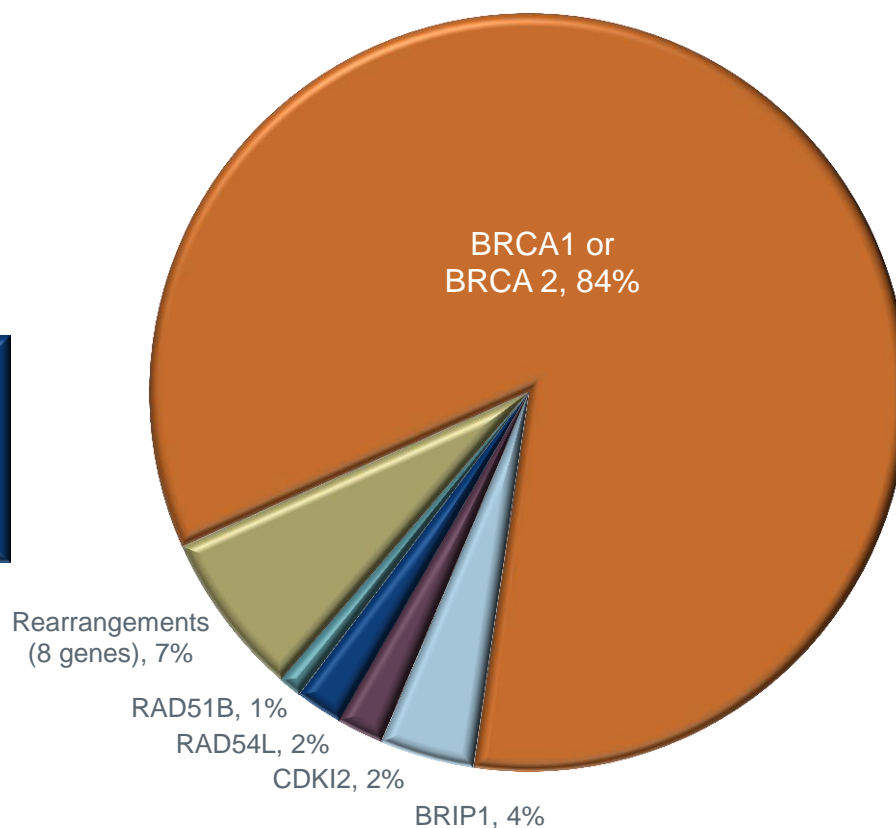


Detects twice as many patients as any other test

# Comprehensive Tumor Profiling Identifies Few Additional HRD Patients

AstraZeneca Study 19: 135 patients with tumor mutations as profiled with 315 gene Foundation Medicine Panel

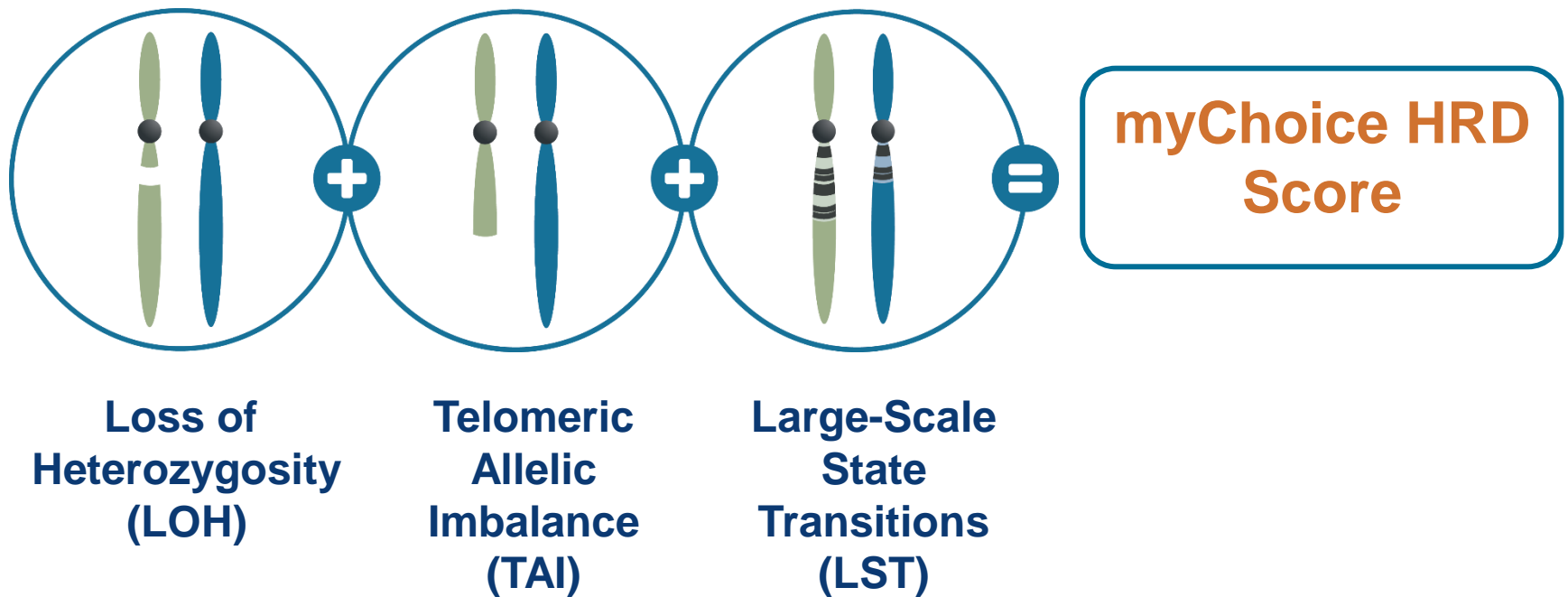
- Only 4 genes mutated more than once
- Only 12 genes had any mutations



# Portfolio of Tests for Assessing DNA Repair Pathway

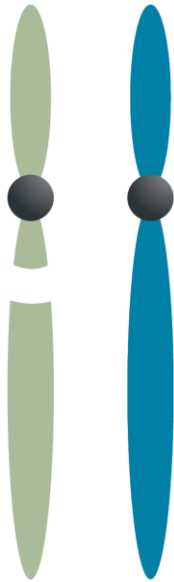
	BRACAnalysisCDx™	Tumor BRACAnalysisCDx™	Tumor Panel	MYRIAD myChoice® HRD
Sample	Blood	Tumor	Tumor	Tumor
Biomarkers	BRCA1&2	Tumor BRCA1&2	80+ clinically actionable oncology genes identified by pharma partners	Genome-wide assessment of DNA scar associated with DNA repair defects
Intellectual Property	Database, process, bioinformatics	Database, process, bioinformatics	Database, process, bioinformatics	MYGN has IP on three proprietary technologies (LOH, TAI, LST)
Currently Marketed	FDA approved	CE Marked	In research use with major pharma partners	Early access launch for platinum

# myChoice HRD Score Results From the Combined Analysis of Three Different Biomarkers



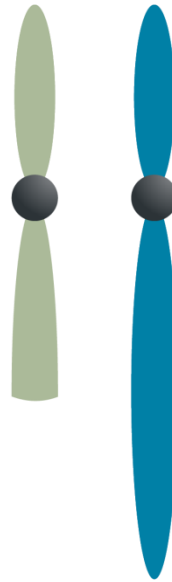


## Three Biomarkers Result in a More Discriminating Measurement of HRD



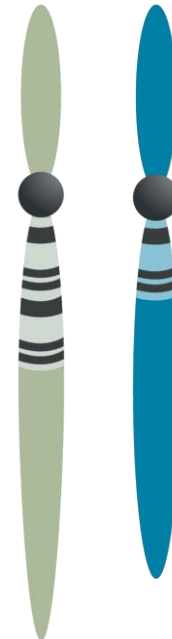
### Loss of heterozygosity (LOH)

Presence of a single allele



### Telomeric allelic imbalance (TAI)

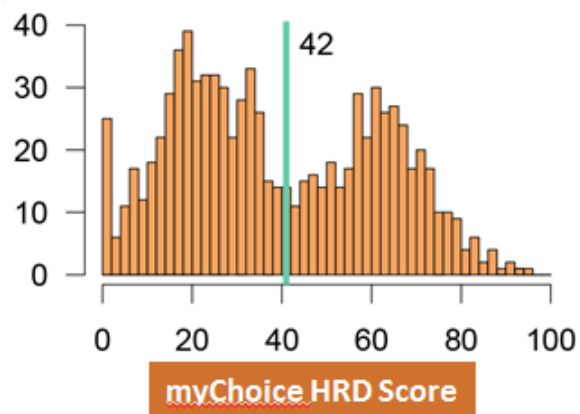
A discrepancy in the 1:1 allele ratio at the end of the chromosome (telomere)



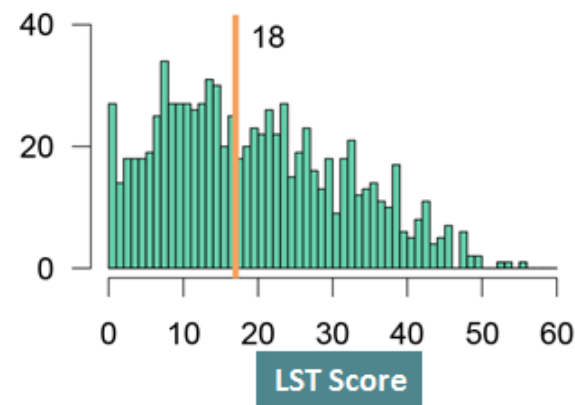
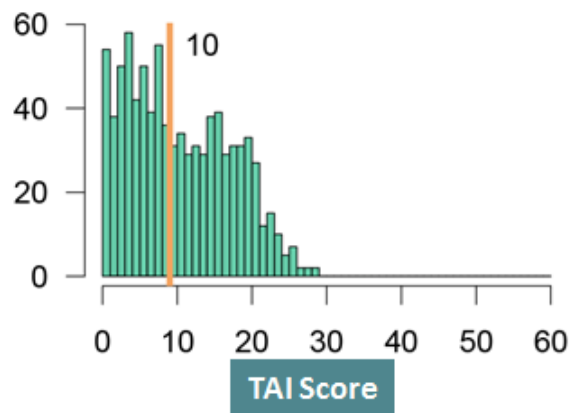
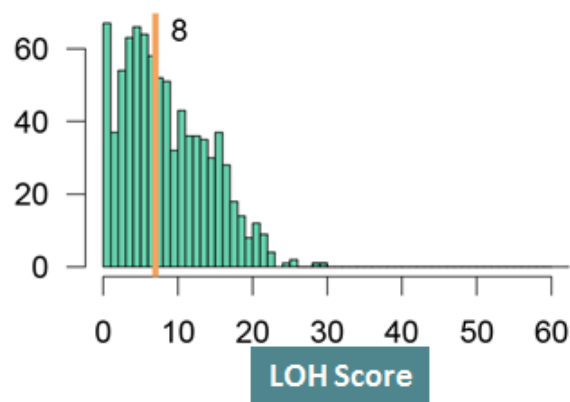
### Large-scale state transitions (LST)

Transition points between regions of abnormal and normal DNA or between two different regions of abnormality

# Single Measures of HRD are Insufficient



In a cohort of 859 ovarian tumors, only a combination of all three biomarkers (HRD score) yielded a clear cutoff.



Hennessy BT, et al. *J Clin Oncol.* 2010; 28:3570.  
TCGA Research Network. *Nature.* 2011; 474:6609.  
Bannerjee et al. *Ann Oncol.* 2013;24(3):679.





# NOVA Study and Data

# Review of the NOVA Study

## Phase 3 NOVA Trial

High-Grade Serous Ovarian Cancer, Platinum Sensitive, Relapsed

Response to Platinum Treatment  
N=490

gBRCA<sup>mut</sup>

2:1 Randomization

Niraparib  
300mg

n=120

Placebo

n=60

Endpoint Assessment

Non-gBRCA<sup>mut</sup> / HRD

2:1 Randomization

Niraparib  
300mg

n=207

Placebo

n=103

Endpoint Assessment

### Primary Endpoint:

- PFS; >90% power to detect 4.8 month improvement (HR 0.50 in both cohorts)
- Assumption: 4.8 month PFS for control arms

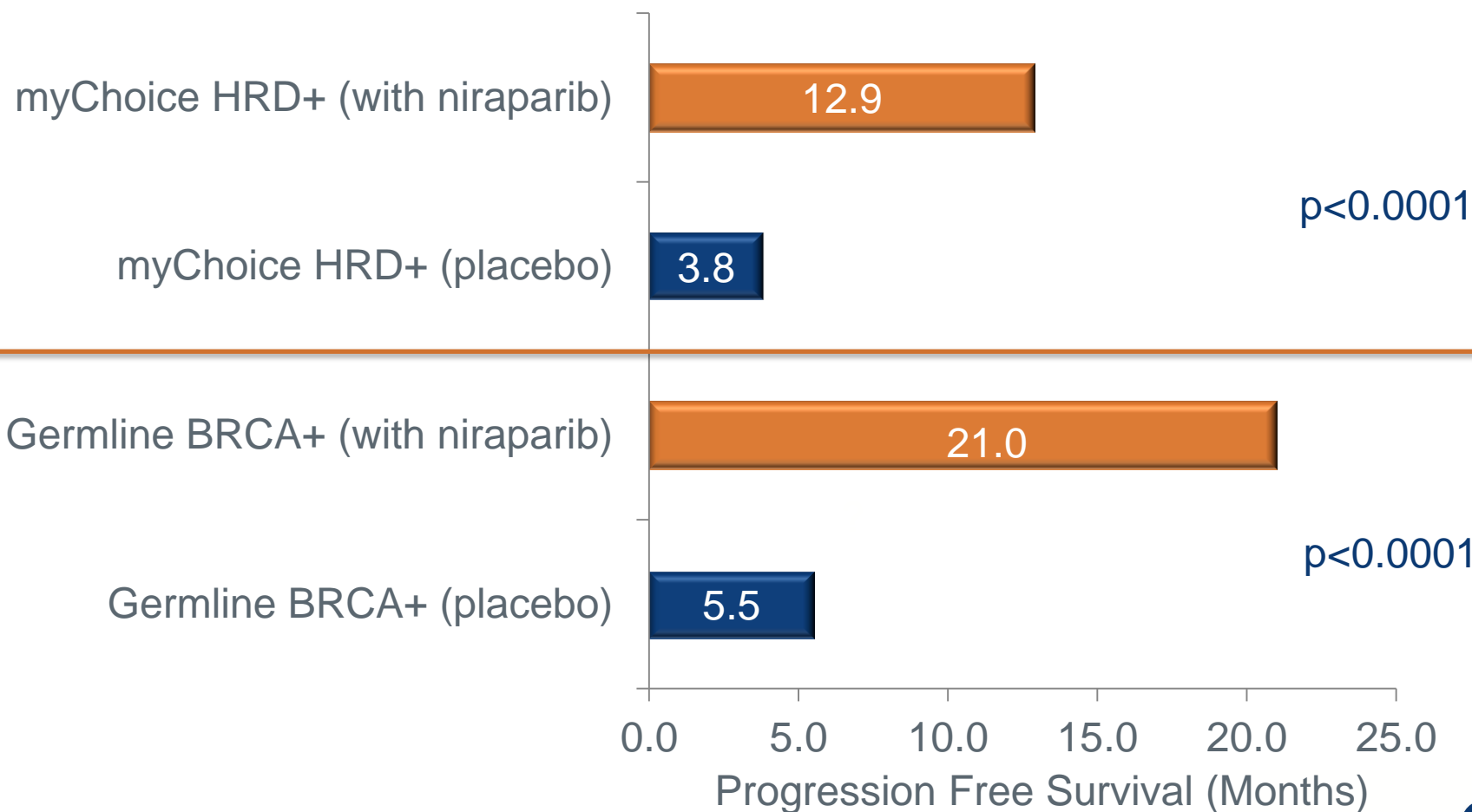
### myChoice HRD:

- First assess PFS in HRD+ subset
  - n≈165
  - >90% power to detect a HR=0.50
- If HRD+ subset has p<0.05, assess PFS for the entire cohort
  - N=310
  - >90% power to detect a HR=0.50



# Niraparib Highly Efficacious in Both myChoice HRD+ Patients and Germline BRCA+ Patients

## Progression Free Survival





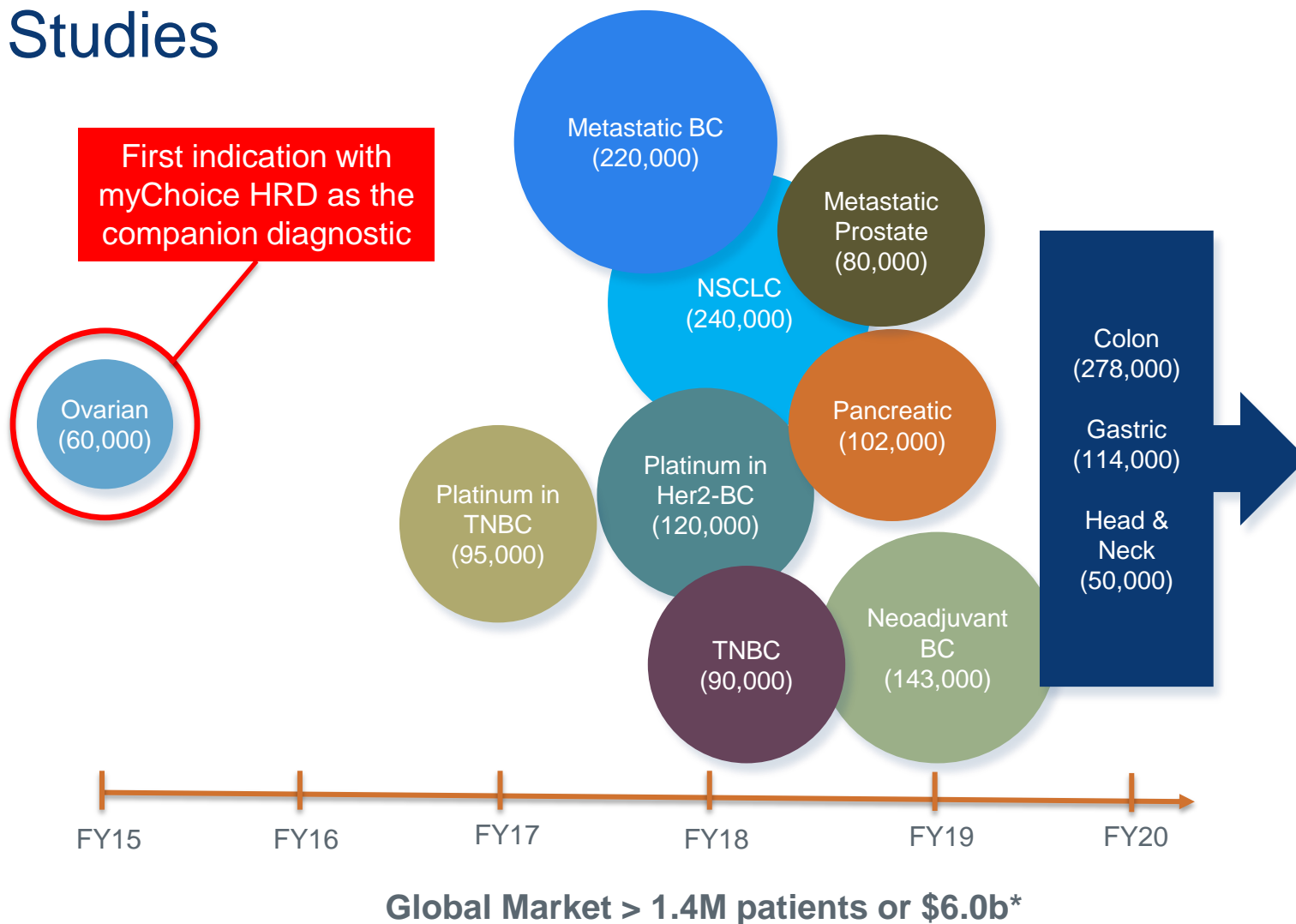
## Summary of the NOVA Data

- 1) Niraparib is highly effective in patients with germline BRCA mutations and those with myChoice HRD positive test results
- 2) myChoice HRD identifies  $\approx 2x$  as many patients who would benefit from Niraparib compared to the FDA approved BRACAnalysis CDx test.
- 3) The myChoice HRD+ patients demonstrated an incremental 9.1 months of progression free survival benefit, which is higher than seen in previous PARP studies



# Business Impact

# Extensive Collaborations With >22 Clinical Studies



\*Includes U.S., Canada and EU6



# myChoice HRD Addressable Market in Ovarian Cancer



U.S.  
22,000  
patients



EU5 &  
Canada  
38,000  
patients

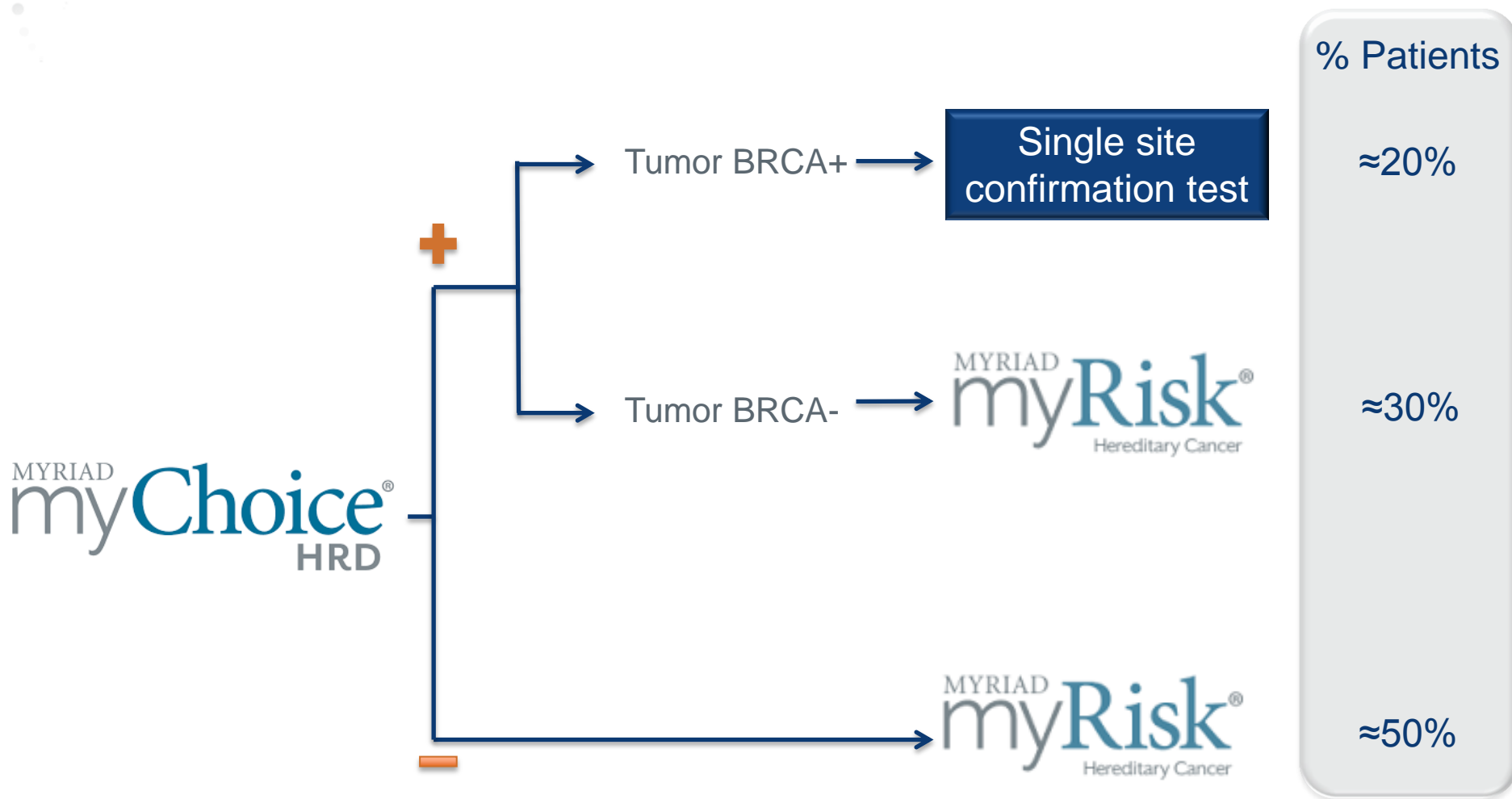
**>\$200M  
Global  
Market**

Differentiated diagnostic  
that is strategically  
important to pharma

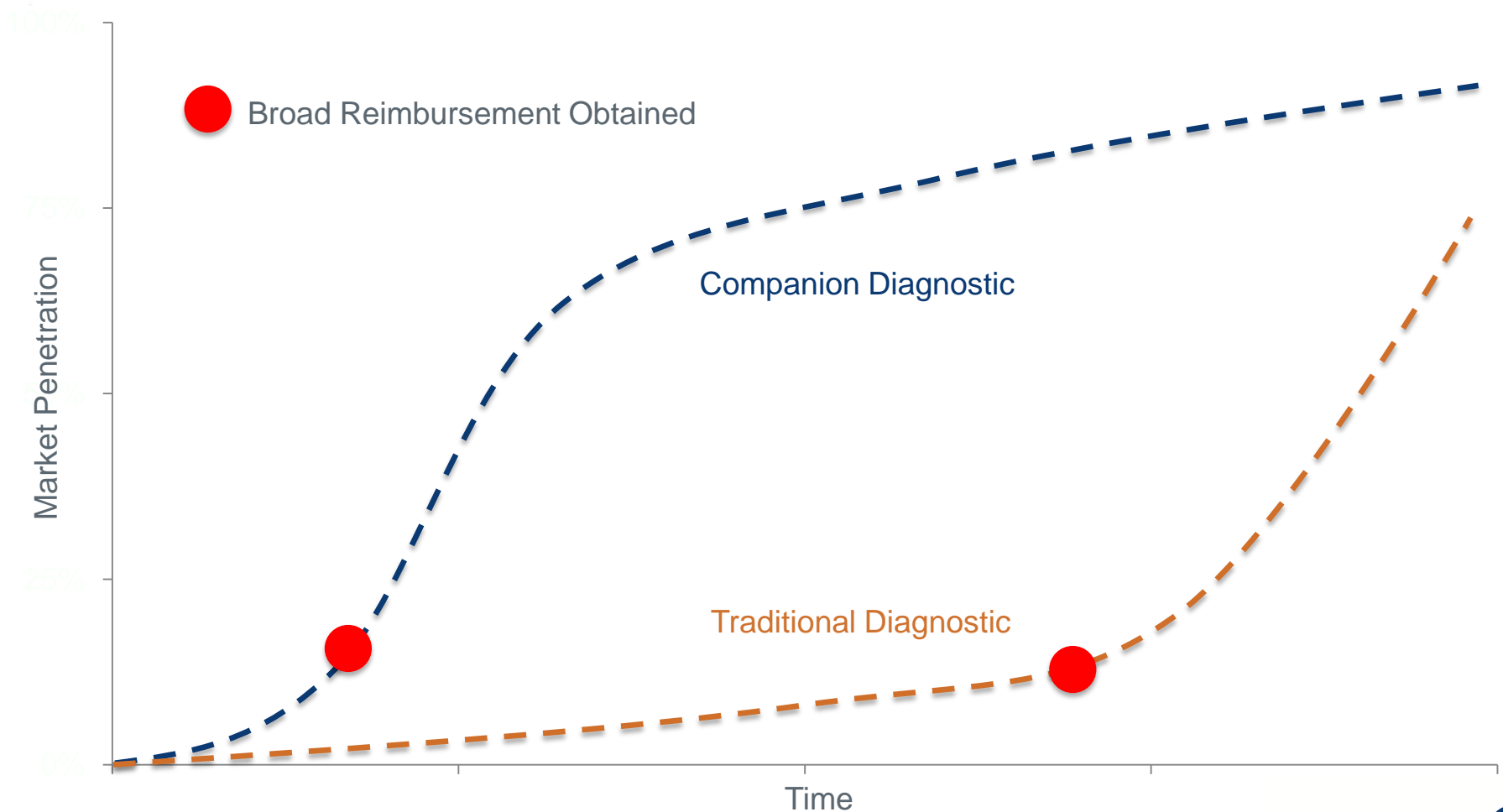
Protected by strong  
intellectual property with  
both patents and trade  
secrets

Unlike BRACAnalysis  
CDx, this is an additive  
opportunity

# myChoice HRD is Additive To Hereditary Cancer Testing



# Revenue Curve for Companion Diagnostics is Much Steeper Than for Traditional Diagnostics





## Other Strategic Considerations

- First prospective data and major validation of myChoice HRD – reduces risk for pharma partners and likely drives additional collaborations
- Growing core competency in FDA approved, high-value companion diagnostics
- Intellectual property around myChoice HRD is significant and provides global differentiation