



**RELAY**<sup>®</sup>  
THERAPEUTICS

**Corporate Presentation**  
**As of February 22, 2024**

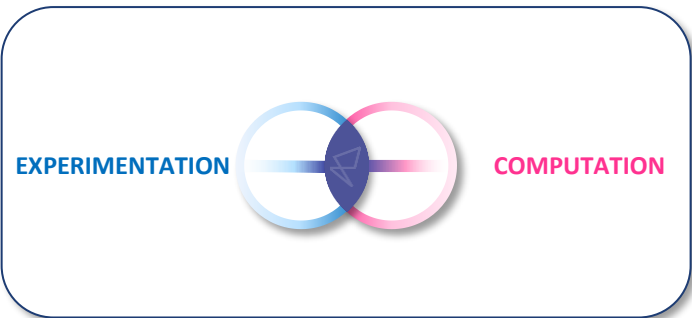
*This presentation contains forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995, as amended, including, without limitation, implied and express statements regarding the progress and timing of the clinical development of the programs across our portfolio, including the expected therapeutic benefits of our programs, and potential efficacy and tolerability, and the timing and success of interactions with and approval of regulatory authorities; the timing of clinical data updates across our pipeline, including the timing of a clinical data update for the PI3K $\alpha$  franchise, the progress of doublet and triplet combinations for RLY-2608, the timing of clinical updates for RLY-2608, and the timing of a clinical data and regulatory update for lirafugratinib; the timing of disclosure of additional pre-clinical programs; the possibility that unconfirmed results from these trials will not be confirmed by additional data as our clinical trials progress; the potential of RLY-2608 to address a major unmet medical need; expectations regarding our pipeline, operating plan, use of capital, expenses and other financial results; our cash runway projection; the competitive landscape and potential market opportunities for our product candidates; the expected strategic benefits under our collaborations; our ability to successfully establish or maintain collaborations or strategic relationships for our product candidates; expectations regarding current and future interactions with the U.S. Food and Drug Administration (FDA); our ability to manufacture our product candidates in conformity with the FDA's requirements; the capabilities and development of our Dynamo™ platform; our plans to develop, manufacture and commercialize our current product candidates and any future product candidates; and the implementation of our business model and strategic plans for our business, current product candidates and any future product candidates. The words "may," "might," "will," "could," "would," "should," "plan," "anticipate," "intend," "believe," "expect," "estimate," "seek," "predict," "future," "project," "potential," "continue," "target" and similar words or expressions, or the negative thereof, are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words.*

*Any forward-looking statements in this presentation are based on management's current expectations and beliefs and are subject to a number of risks, uncertainties and important factors that may cause actual events or results to differ materially from those expressed or implied by any forward-looking statements contained in this presentation, including, without limitation, risks associated with: the impact of global economic uncertainty, geopolitical instability and conflicts, or public health epidemics or outbreaks of an infectious disease on countries or regions in which we have operations or do business, as well as on the timing and anticipated results of our clinical trials, strategy, future operations and profitability; the delay or pause of any current or planned clinical trials or the development of our drug candidates; the risk that the preliminary results of our preclinical or clinical trials may not be predictive of future or final results in connection with future clinical trials of our product candidates; our ability to successfully demonstrate the safety and efficacy of our drug candidates; the timing and outcome of our planned interactions with regulatory authorities; and obtaining, maintaining and protecting our intellectual property. These and other risks, uncertainties and important factors are described in the section entitled "Risk Factors" in our most recent Annual Report on Form 10-K and Quarterly Report on Form 10-Q, as well as any subsequent filings with the Securities and Exchange Commission. Any forward-looking statements represent our views only as of the date of this presentation and we undertake no obligation to update or revise any forward-looking statements, whether as a result of new information, the occurrence of certain events or otherwise. We may not actually achieve the plans, intentions or expectations disclosed in our forward-looking statements, and you should not place undue reliance on our forward-looking statements. No representations or warranties (expressed or implied) are made about the accuracy of any such forward-looking statements.*

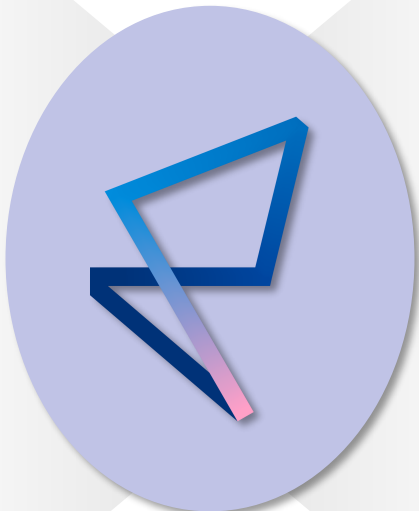
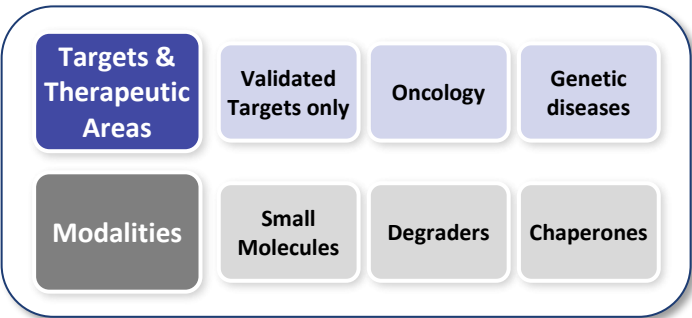
*Certain information contained in this presentation relates to or is based on studies, publications, surveys and other data obtained from third-party sources and our own internal estimates and research. While we believe these third-party studies, publications, surveys and other data to be reliable as of the date of this presentation, we have not independently verified, and make no representation as to the adequacy, fairness, accuracy or completeness of, any information obtained from third-party sources. In addition, no independent source has evaluated the reasonableness or accuracy of our internal estimates or research and no reliance should be made on any information or statements made in this presentation relating to or based on such internal estimates and research.*

*This presentation contains trademarks, trade names and service marks of other companies, which are the property of their respective owners.*

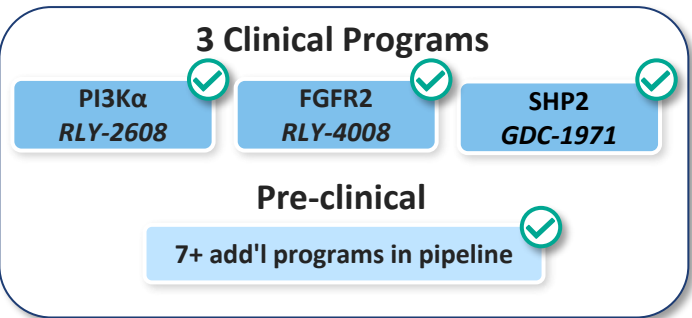
New Breed of Biotech



Clear Focus



Validated Approach



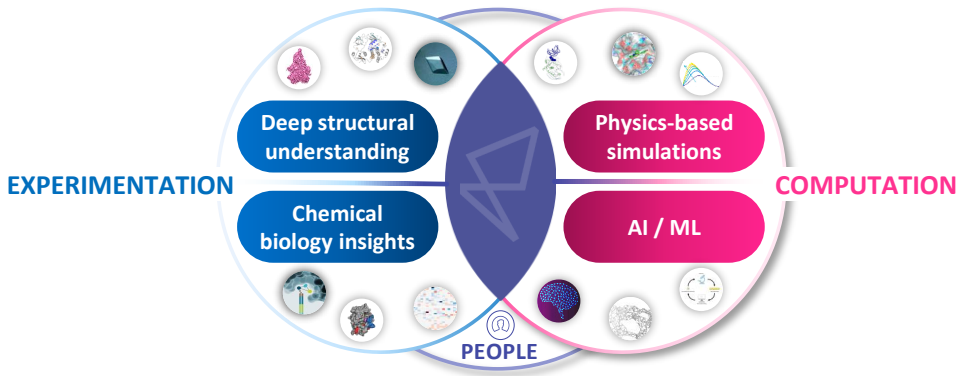
Execution-Focused

Target	Program	Preclinical	Early Clinical	Late Clinical
PI3Kα franchise	Monotherapy			
	RLY-2608 PI3Kα <sup>WT</sup>			
	Endocrine Tx (ET) doublet			
	CDK4/6 + ET triplet			
FGFR2	RLY-5836 PI3Kα <sup>WT</sup>	Deprioritized		
	PI3Kα <sup>WT</sup>			
	Lirafugratinib (RLY-4008)			
Solid Tumor	2 programs			
Genetic Disease	2 programs			
CDK2	RLY-2139	Paused, IND ready		
ERα	RLY-1013 (Degrader)	Paused at DC		
SHP2	Migaprotafin (GDC-1971) Overwatch	3 ongoing combo studies		

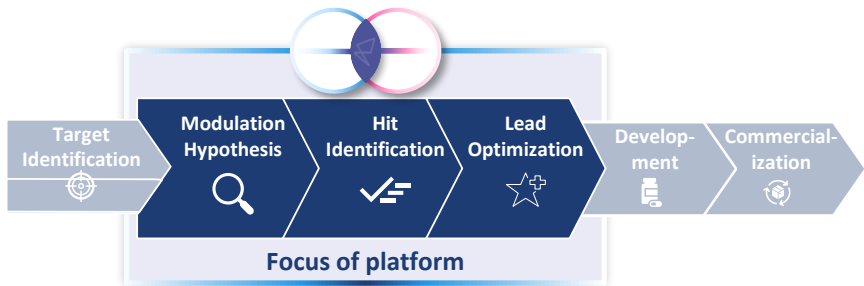
~\$750M

Cash, cash equivalents and investments as of the end of 4Q 2023

1 Dynamo™ Platform...



2 ...is focused on making medicines



3 ...aims to address selectivity on validated targets



# Relay Tx – Broad Precision Medicine Pipeline



Target	Program	Preclinical	Early Clinical	Late Clinical	Annual US Patient #
PI3Kα franchise	Monotherapy				~10-71K breast cancer ~76-243K all solid tumors
	RLY-2608 PI3Kα <sup>PAN</sup> Endocrine Tx (ET) doublet				
	CDK4/6i + ET triplet				
	RLY-5836 (PI3Kα <sup>PAN</sup> ) Dose Escalation	Deprioritized			~4-27K breast cancer ~15-50K all solid tumors
	PI3Kα <sup>H1047R</sup>				
FGFR2	Lirafugratinib (RLY-4008)				~11-35K <sup>4</sup>
Solid Tumor	2 programs				To be announced
Genetic Disease	2 programs				To be announced
CDK2	RLY-2139	Paused; IND ready			~35K <sup>2</sup>
ERα	RLY-1013 (Degradar)	Paused at DC			~30-205K <sup>3</sup>
SHP2	Migoprotafib (GDC-1971) Genentech <small>A Member of the Roche Group</small>	3 ongoing combo studies			~36-69K <sup>5</sup>

**Note:** Unless otherwise indicated, patient #'s refer to total annual number of US patients with late-line cancers compared to comprehensive annual incidence that may be amenable to treatment with our programs

1. Unless otherwise indicated, all breast cancer patient numbers refer to HR+/HER2- breast cancer tumors; 2. ~35K HR+/HER2- breast cancer patients expected to receive CDK 4/6 inhibitors in adjuvant setting, first-line setting, and second-line setting in 2024, per Decision Resources Breast Cancer Market Forecast report dated November 2023; 3. HR+/HER2- US late-line breast cancer patients compared to HR+/HER2- US incident breast cancer patients; 4. FGFR2 altered late-line solid tumors compared to comprehensive annual FGFR2 altered incident solid tumors including additional FGFR gene fusions and rearrangements resulting from truncation of the protein at exon 18 and all breast cancer patients with FGFR2 alterations; 5. SHP2 combo only includes KRAS G12C in lung and colorectal, EGFR mutations in lung, and ALK fusions in lung

# Relay Tx – Capital, Team & Execution Focus to Deliver on Milestones



## 2024 Corporate Objectives

**RLY-2608 Doublet**  
(PI3K $\alpha$ )

- Additional clinical data in 2H 2024

**RLY-2608 Triplet**  
(PI3K $\alpha$ )

- ✓ Ribociclib triplet initiation in Q4 2023
- Ribociclib triplet safety data in 2H 2024

**Lirafugratibnib (RLY-4008)**  
(FGFR2)

- Tumor agnostic data and regulatory update in 2H 2024

**Pre-clinical Pipeline**  
(Targets unnamed)

- New program(s) to be disclosed in 2024
- 7+ undisclosed programs in preclinical development and additional early-stage efforts across platform

**Migoprotafib (GDC-1971)**  
 (SHP2)

- Three ongoing combination trials  
\*Genentech controls data disclosures

Goal is a first- or best-in-class profile

## Significant Capital to Achieve Goals

~\$750M

Cash, cash equivalents and investments as of the end of 4Q 2023

Expected to be sufficient to fund current operating plan  
into 2H 2026

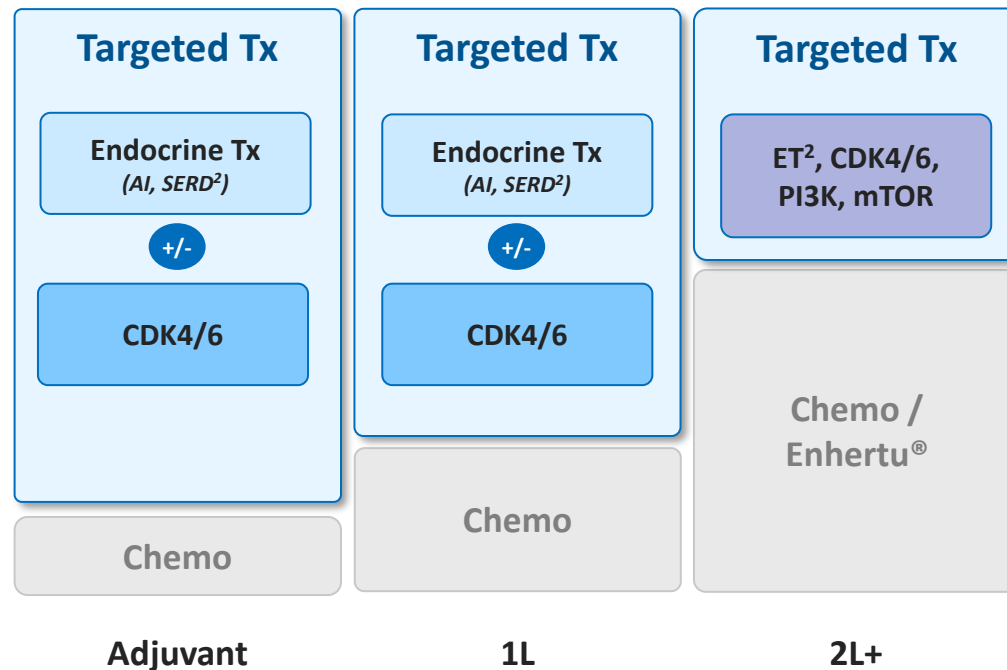
# Relay Tx – Broad Precision Medicine Pipeline



Target	Program	Preclinical	Early Clinical	Late Clinical
PI3Kα franchise	Monotherapy			
	RLY-2608 PI3Kα <sup>PAN</sup> Endocrine Tx (ET) doublet			
	CDK4/6i + ET triplet			
	RLY-5836 (PI3Kα <sup>PAN</sup> ) Dose Escalation	Deprioritized		
	PI3Kα <sup>H1047R</sup>			
CDK2	RLY-2139	Paused; IND ready		
ERα	RLY-1013 (Degradar)	Paused at DC		
FGFR2	Lirafugratinib (RLY-4008)			
Solid Tumor	2 programs			
Genetic Disease	2 programs			
SHP2	Migoprotafib (GDC-1971) Genentech <small>A Member of the Roche Group</small>	3 ongoing combo studies		

# Breast Cancer – Limitations of Current Standard of Care

## HR+/HER2- breast cancer standard of care<sup>1</sup>...



## ...is limited by efficacy of available treatments



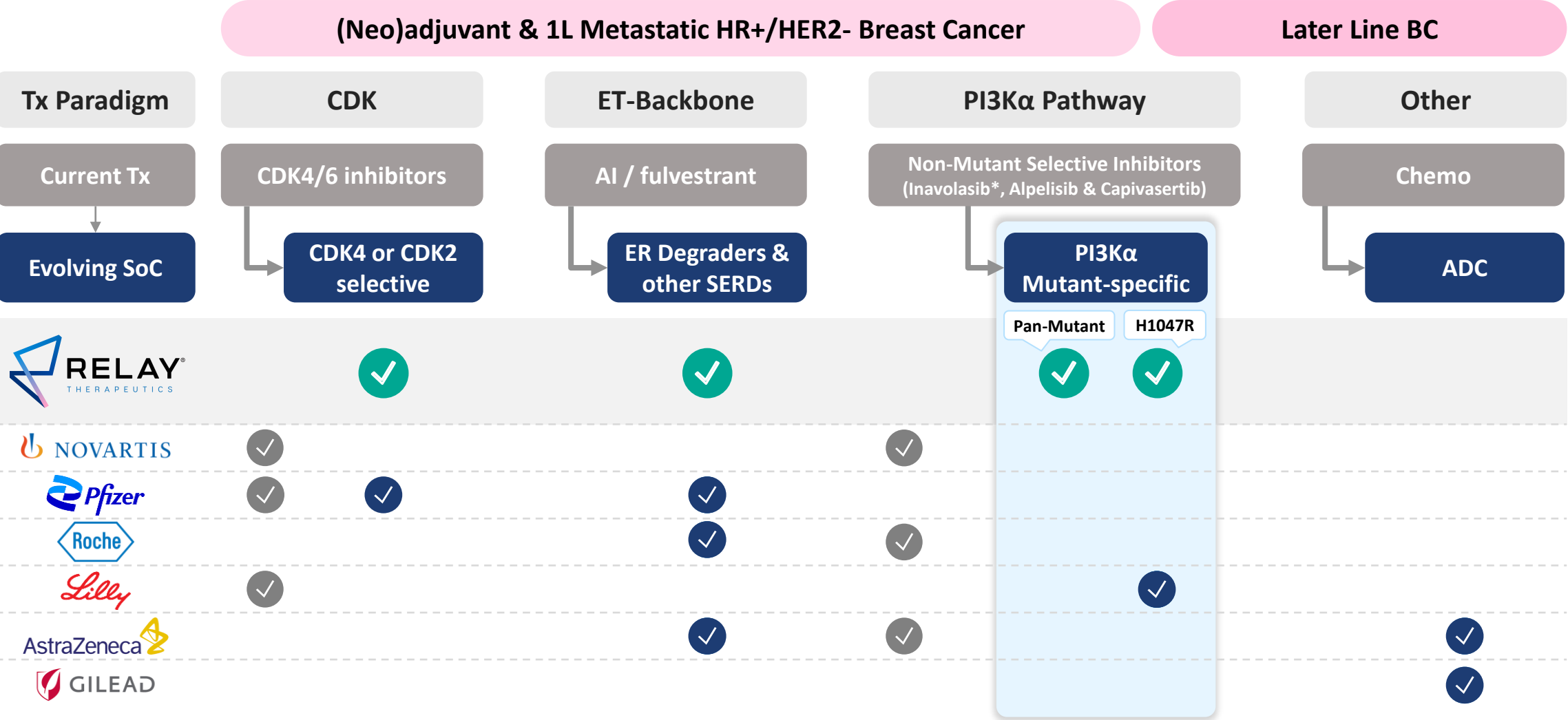
Source: Internal analysis based on third party industry data

1. Standard of care for HR+/HER2- breast cancer is illustrative; 2. AI = Aromatase Inhibitor; SERD: Selective Estrogen Receptor Degradar; ET = Endocrine Therapy



# Breast Cancer – Evolving Landscape With Very Large Market Opportunity

## \$27B Market Size of (Neo)adjuvant and 1L Metastatic HR+/HER2- Breast Cancer



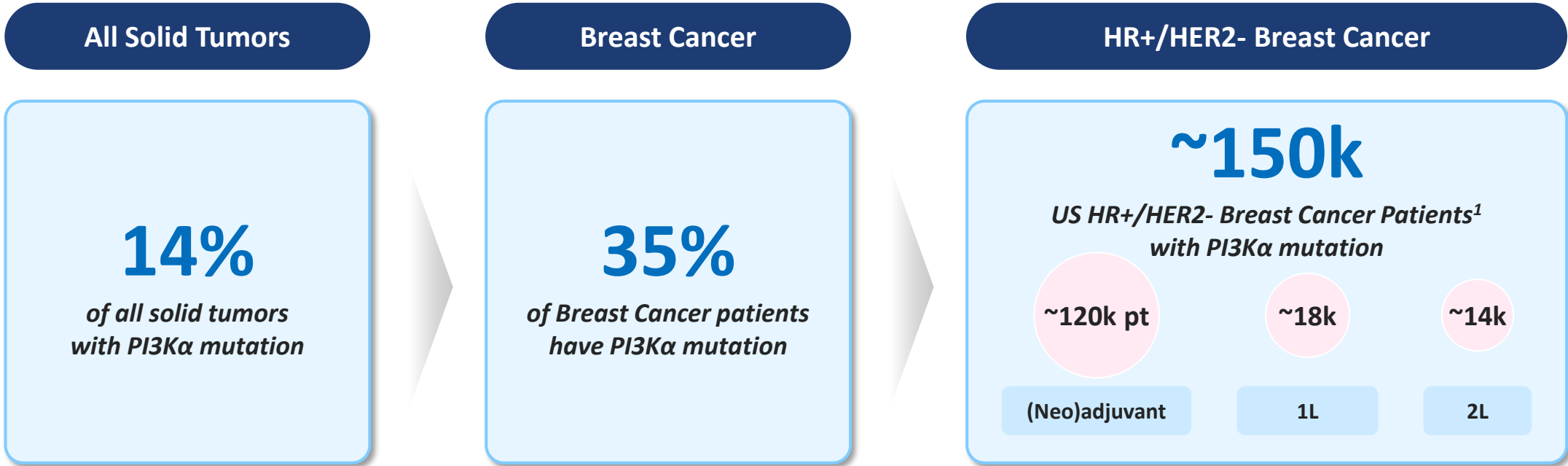
\* Inavolisib is an investigational therapy in Ph3 studies  
Source: Decision Resources Group – Breast Cancer Disease Landscape & Forecast (Nov 2023). 2031 Projection  
© 2024 Relay Therapeutics

# Relay Tx – Broad Precision Medicine Pipeline



Target	Program	Preclinical	Early Clinical	Late Clinical
PI3Kα franchise	Monotherapy			
	RLY-2608 PI3Kα <sup>PAN</sup> Endocrine Tx (ET) doublet			
	CDK4/6i + ET triplet			
	RLY-5836 (PI3Kα <sup>PAN</sup> ) Dose Escalation	Deprioritized		
	PI3Kα <sup>H1047R</sup>			
CDK2	RLY-2139	Paused; IND ready		
ERα	RLY-1013 (Degradar)	Paused at DC		
FGFR2	Lirafugratinib (RLY-4008)			
Solid Tumor	2 programs			
Genetic Disease	2 programs			
SHP2	Migoprotafib (GDC-1971) Genentech <small>A Member of the Roche Group</small>	3 ongoing combo studies		

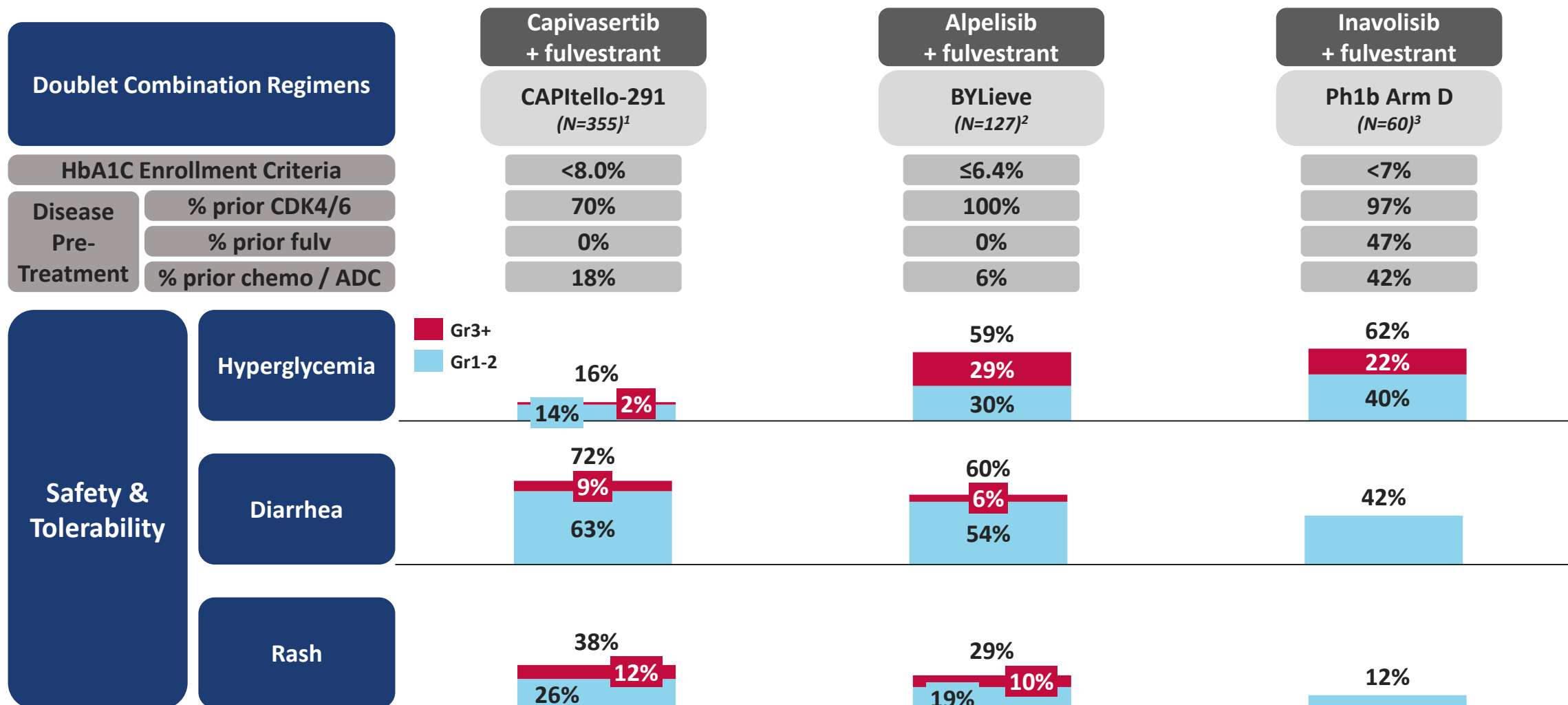
# PI3Kα Represents a Major Market Opportunity



**RLY-2608 has the potential to address very large patient population**

Sources: 3<sup>rd</sup> party data; Global Data HER2-/HR+ Breast Cancer Global Patient Forecast, October 2023;  
1. Includes prevalent PI3Kα mutated HR+/HER2- patients receiving therapy in Neo/Adjuvant setting (includes incident patients in 2023 receiving endocrine or non-endocrine therapy in Neo/Adjuvant settings [~50k], and patients diagnosed in previous years with local/regional disease receiving sequential endocrine therapy in 2023 [~69k]), and prevalent PI3Kα mutated HR+/HER2- metastatic patients receiving therapy in 1L or 2L setting; 2. Approved in combination with fulvestrant in patients with at least one prior endocrine-based regimen in metastatic setting or early progression on endocrine therapy (during or within 12 months of completing adjuvant treatment)  
© 2024 Relay Therapeutics

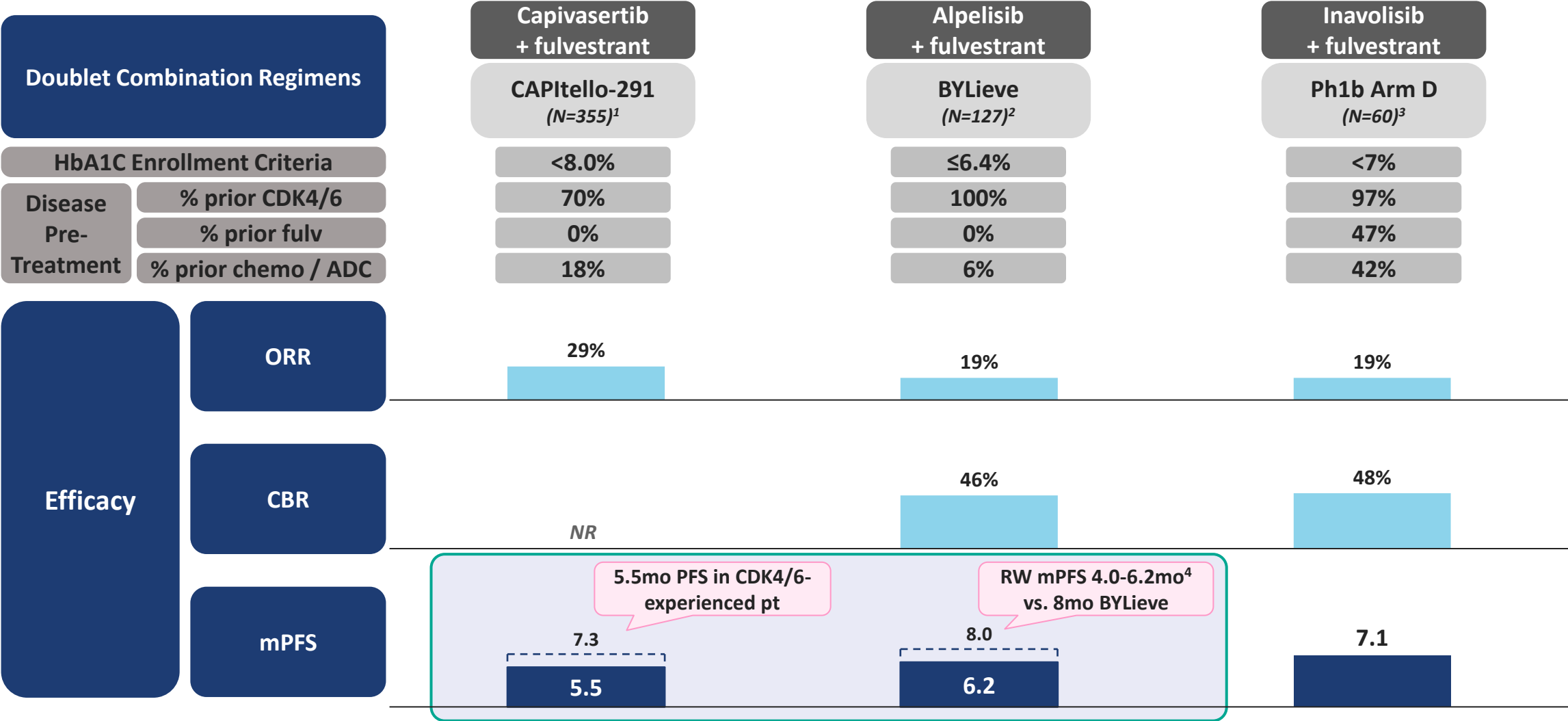
# RLY-2608 – Safety Profiles of Existing PI3Kα Pathway Compounds



Sources: 1. Turner N Engl J Med 2023; 388:2058-2070; 2. Rugo 2021 Lancet Oncol 22:489; 3. SABCS 2021 #P5-17-05

Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.  
© 2024 Relay Therapeutics

# RLY-2608 – Efficacy Profiles of Existing PI3Kα Pathway Compounds



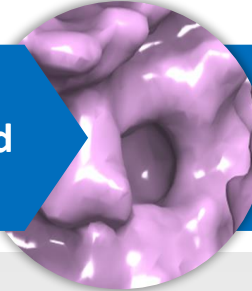
Sources: 1. Turner N Engl J Med 2023; 388:2058-2070; 2. Rugo 2021 Lancet Oncol 22:489; 3. SABCS 2021 #P5-17-05; 4. ASCO 2022 #1055 (Novartis-sponsored real-world evidence study for alpelisib + fulvestrant)  
Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.  
© 2024 Relay Therapeutics

# PI3K $\alpha$ – Proprietary Insights Unlock Novel Approaches

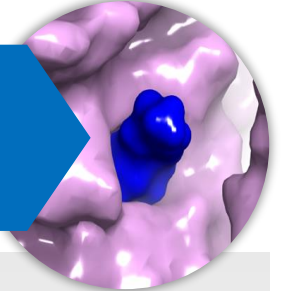
Solved first full-length  
structures of PI3K $\alpha$   
(mutant and wild-type)



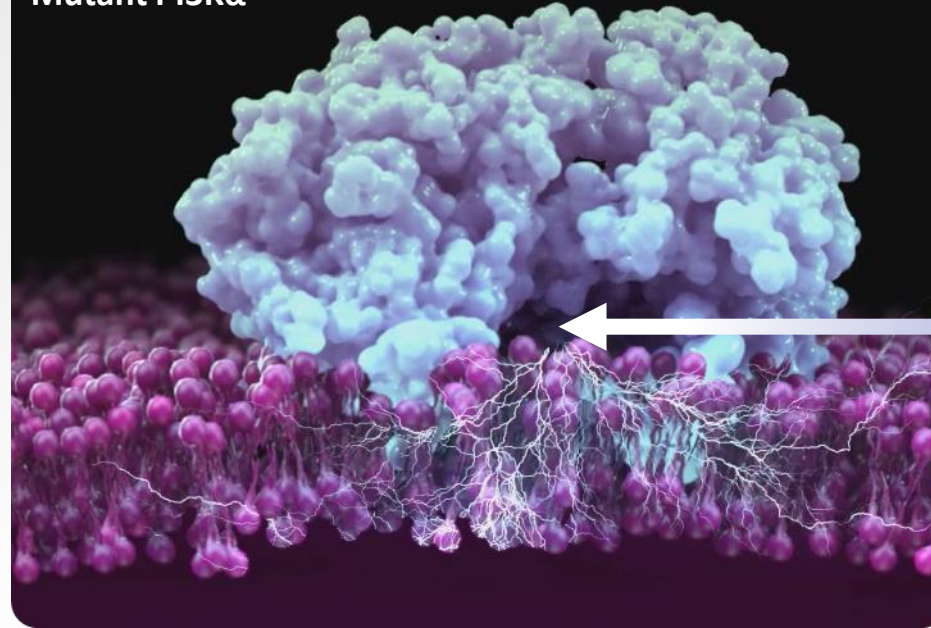
Discovered novel  
allosteric pocket favored  
in mutant protein



Designed pan-mutant  
selective PI3K $\alpha$   
inhibitor (PI3K $\alpha$ <sup>PAN</sup>)



Mutant PI3K $\alpha$



Orthosteric Site

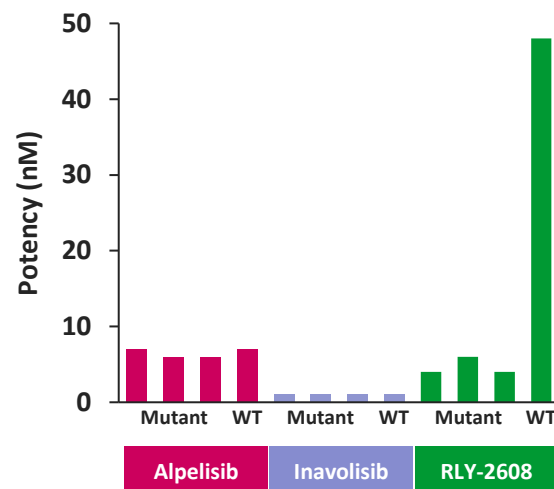
A differentiated understanding of the structure of PI3K $\alpha$  and its relationship to function  
equips Relay Tx to design optimal mutant-selective inhibitors of PI3K $\alpha$

# RLY-2608 – First Mutant Selective Inhibitor to Enter the Clinic

All Data Shown is Preclinical

## Favorable Selectivity

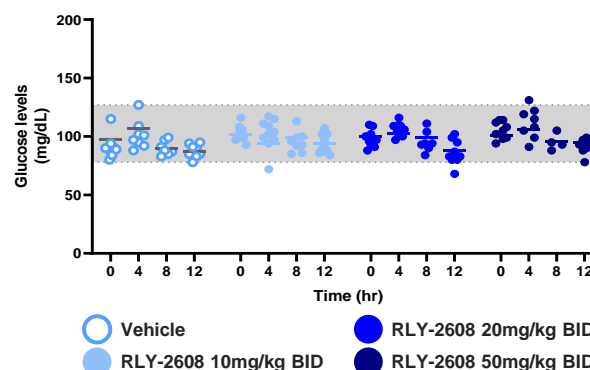
Limited potency against WT PI3K $\alpha$  and other PI3K isoforms



## Favorable Tolerability

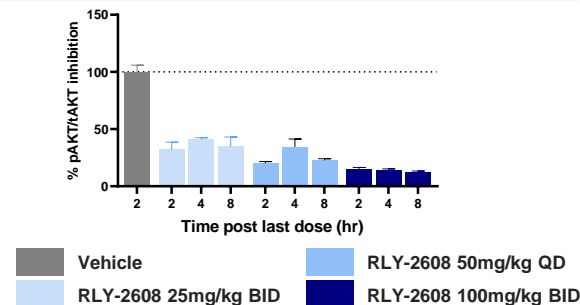
Manageable key toxicities, especially hyperglycemia shown in dog study

28-Day Repeat Dose Dog Study



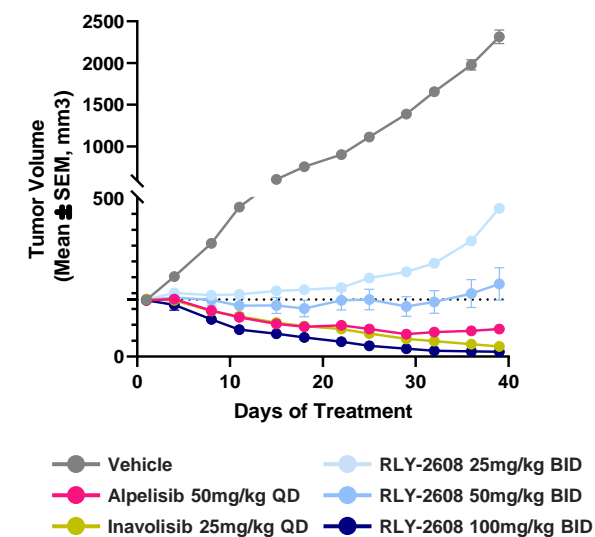
## Favorable Target Inhibition

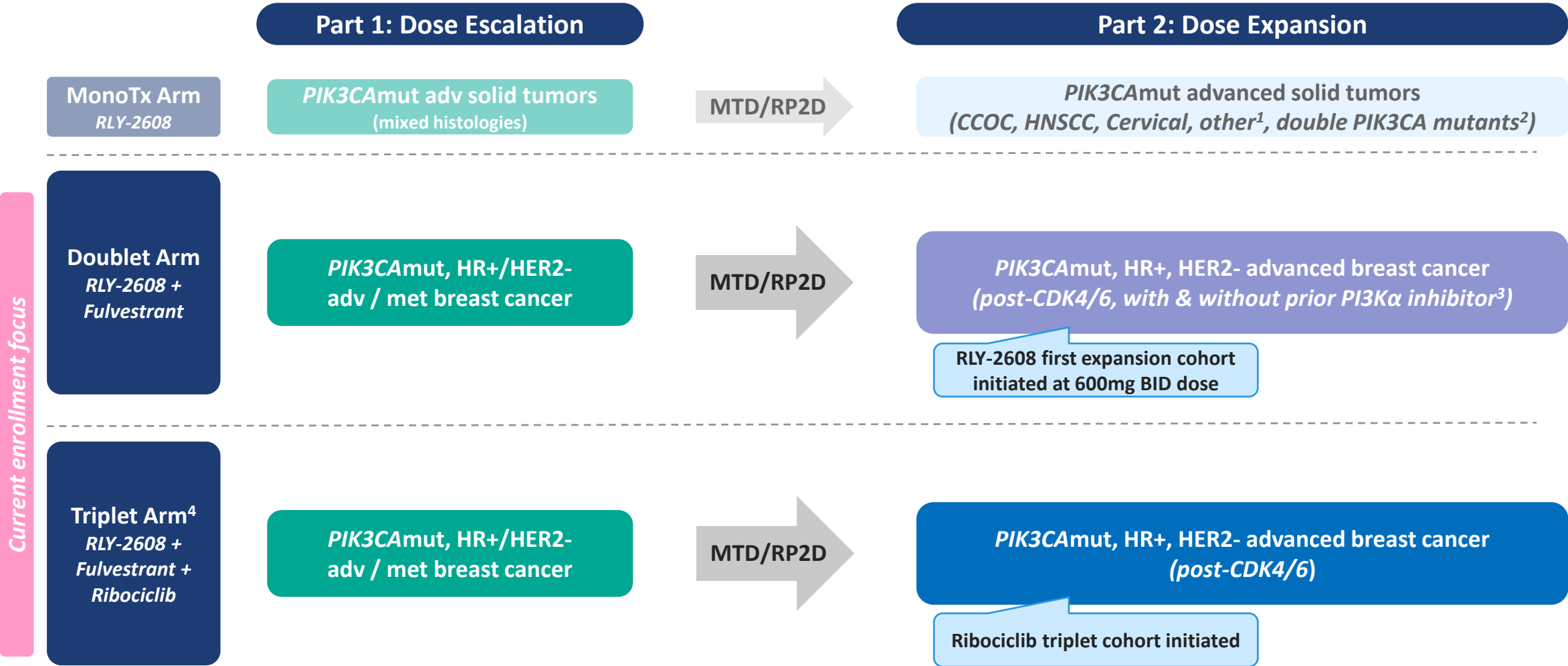
Maintains approx. 80% mutant PI3K $\alpha$  inhibition in mouse model



## Favorable Efficacy

Robust tumor regression at tolerable doses in mouse model



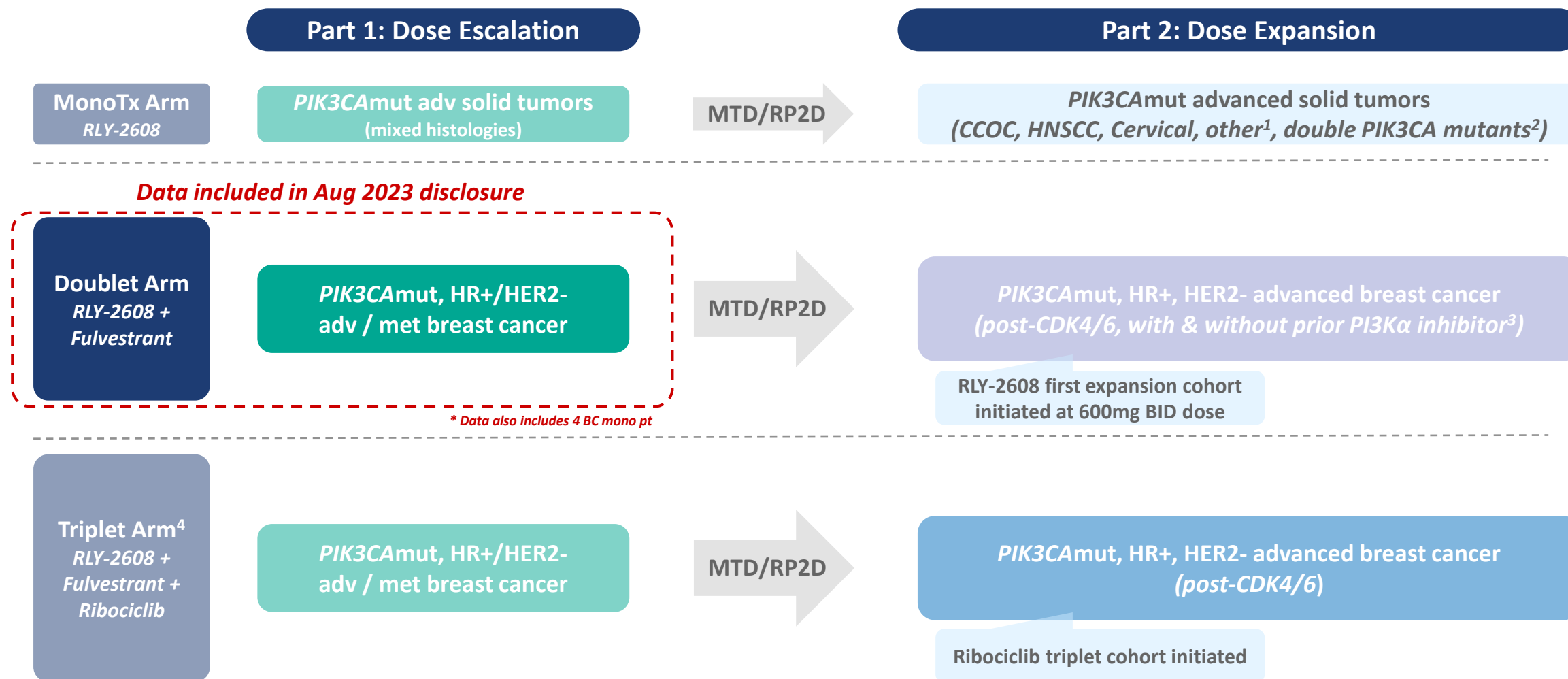


1. Excludes *PIK3CA*mut clear cell OvCA, HNSCC, Cervical cancer, and colorectal patients; 2. Double mutation defined as one major *PIK3CA* mutation (E542X, E545X, H1047X) + ≥1 additional *PIK3CA* mutation per local assessment; 3. Patients with previous *PI3Kα* inhibitor include those with intolerance to *PI3Kα*i defined as treatment discontinuation due to treatment-related AE (e.g., hyperglycemia, rash, diarrhea, stomatitis) other than severe hypersensitivity reaction and/or life-threatening reactions, such as anaphylaxis and Stevens-Johnson syndrome; 4. RLY-2608 + fulvestrant + CDK4/6 arm expected to be added in a protocol amendment

© 2024 Relay Therapeutics



# RLY-2608 – ReDiscover Trial Interim Part 1 Results Disclosed in August 2023



1. Excludes *PIK3CA*mut clear cell OvCA, HNSCC, Cervical cancer, and colorectal patients; 2. Double mutation defined as one major *PIK3CA* mutation (E542X, E545X, H1047X) + ≥1 additional *PIK3CA* mutation per local assessment; 3. Patients with previous *PI3Kα* inhibitor include those with intolerance to *PI3Kα*i defined as treatment discontinuation due to treatment-related AE (e.g., hyperglycemia, rash, diarrhea, stomatitis) other than severe hypersensitivity reaction and/or life-threatening reactions, such as anaphylaxis and Stevens-Johnson syndrome; 4. RLY-2608 + fulvestrant + CDK4/6 arm expected to be added in a protocol amendment

# RLY-2608 – Initial Data Support Selective Targeting of Mutant PI3K $\alpha$

*Summary of initial data in 43 Breast Cancer patients<sup>1</sup> as first disclosed in August 2023*

## Initial Clinical Proof of Mechanism



**Initial anti-tumor activity observed across range of doses**

- At 600mg BID dose in combination with fulvestrant:
  - 86% interim CBR (6 of 7 patients with CR, PR, or SD for  $\geq 6$ mo)
  - 1 cPR out of 5 evaluable<sup>2</sup> patients with measurable disease
- Overall, 4 PRs (of 24 evaluable<sup>2</sup> breast cancer pts) observed across mono and fulvestrant combo, dose levels and PI3K $\alpha$  genotypes



**Favorable safety profile at therapeutically active doses with evidence of selective target inhibition**

- Low rates of hyperglycemia, rash and diarrhea compared to non-selective PI3K inhibitors
- Limited observed impact on glucose homeostasis
- Continuous PK exposure above IC<sub>80</sub> achieved at  $\geq 400$ mg BID
- Safety profile at 600mg BID compelling for use in mBC combinations

Expansion cohorts at 400mg and 600mg BID underway

**Goal for Expansion Cohorts**

**Interpretable Efficacy (CBR, ORR)**

**Longer-Term Tolerability**

DLTs = dose limiting toxicities; CBR: Clinical Benefit defined as all patients with confirmed complete response or partial response or stable disease  $\geq 24$  weeks; evaluable patients started treatment  $\geq 24$  weeks prior to the data cutoff

1. N=43 Breast Cancer patients: 39 fulvestrant combo (17 at 600 mg BID), 4 monotherapy; 2. Efficacy analysis includes patients with measurable disease who had opportunity for  $\geq 1$  tumor assessment or discontinued treatment with  $< 1$  tumor assessment;

3. per CTCAE v5.0

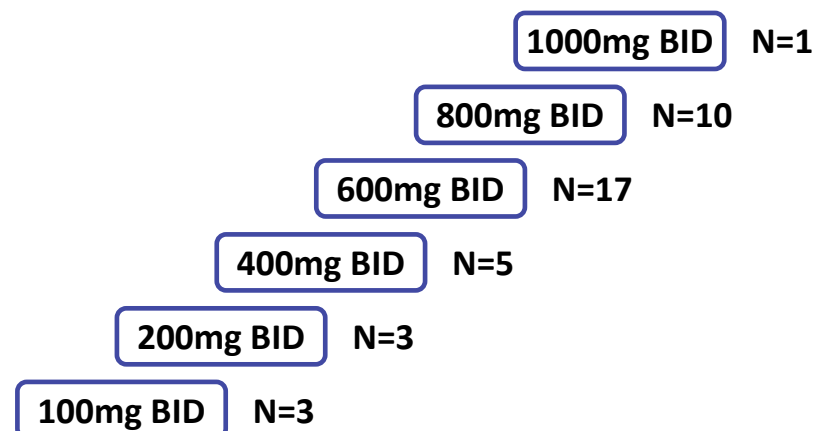
© 2024 Relay Therapeutics

Preliminary data as of 07/24/2023

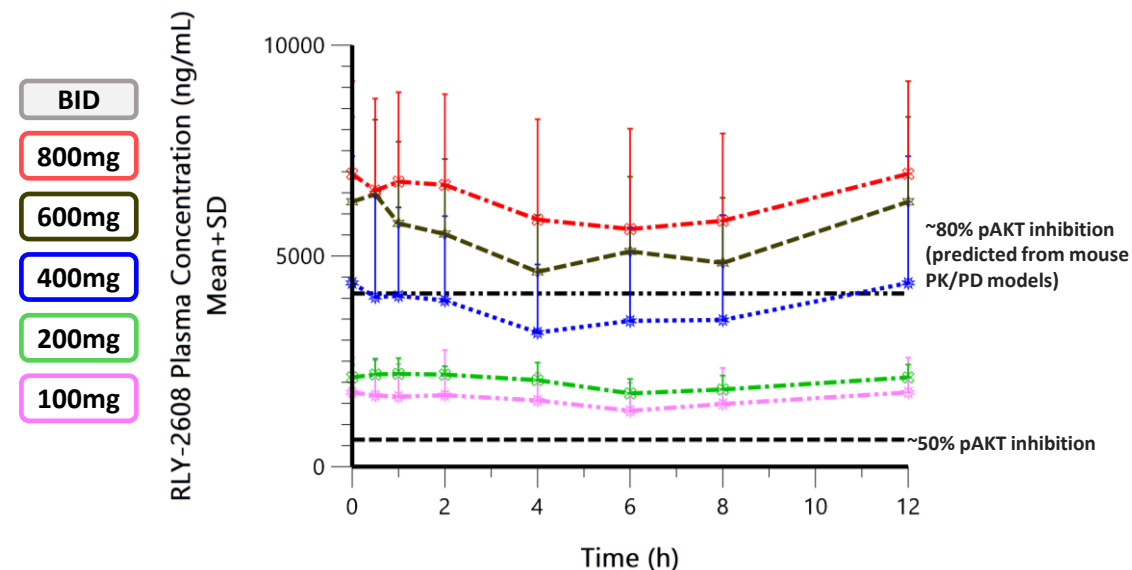
# RLY-2608 – ReDiscover Trial Interim Part 1 Results

## RLY-2608 + fulvestrant

### Dose Escalation



### Favorable PK Profile Across Dose Levels



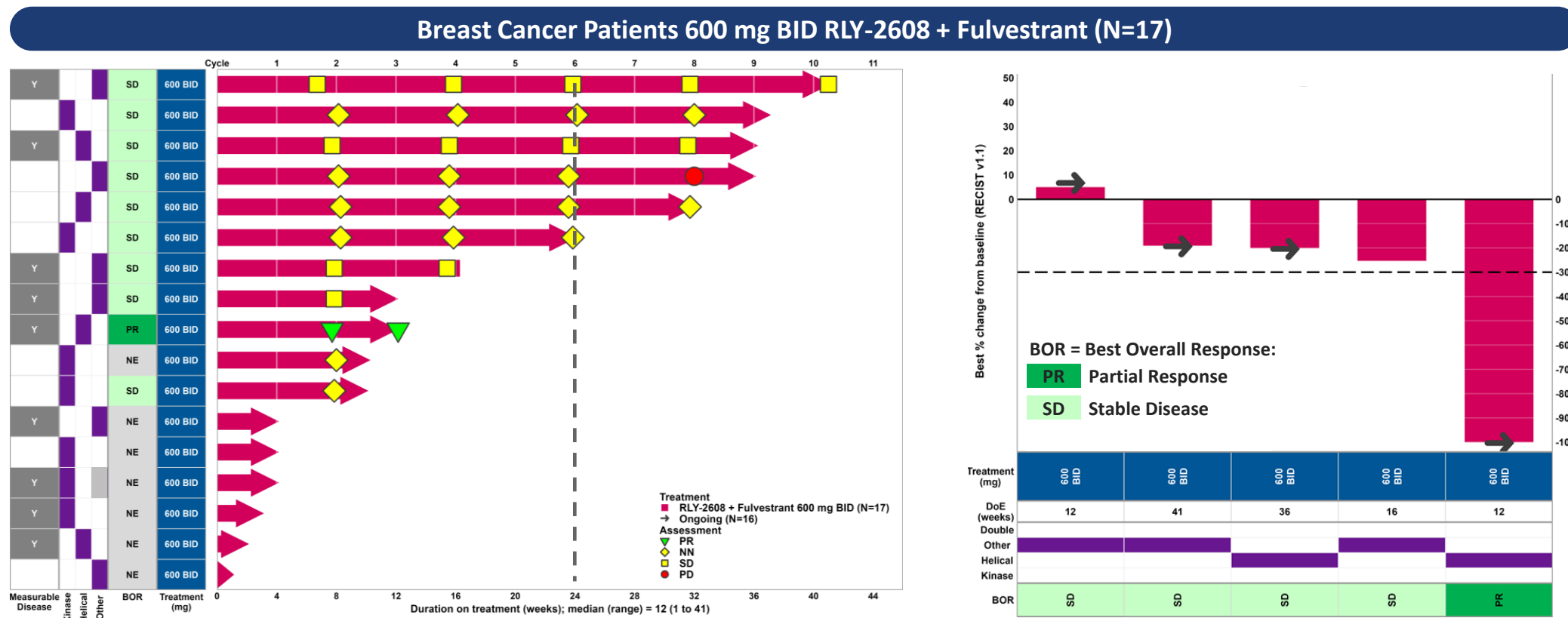
No DLTs and MTD has yet to be defined  
Dose-dependent increase in exposure and low peak to trough fluctuations across dose levels  
Continuous coverage at ~IC80+ across dosing interval at 400mg BID combo and above

# RLY-2608 – ReDiscover Trial Breast Cancer Baseline Demographics and Genotype

	RLY-2608 + fulvestrant (N=39)	RLY-2608 + fulvestrant 600 mg BID (N=17)	RLY-2608 Monotherapy (N=4)
Age, median (range), years	59 (40-82)	60 (49-80)	64 (58, 85)
Female, n (%)	39 (100%)	17 (100%)	4 (100%)
Ethnicity, %			
White / Asian / American Indian / Black / Unknown	67% / 3% / 3% / 3% / 23%	59% / 0% / 0% / 0% / 41%	100% / 0% / 0% / 0% / 0%
ECOG, n (%)			
0	21 (54%)	8 (47%)	2 (50%)
1	18 (46%)	9 (53%)	2 (50%)
BMI, kg/m <sup>2</sup> , median (range)	25 (18-41)	23 (19-36)	26 (18, 44)
<30, n (%)	29 (74%)	14 (82%)	3 (75%)
≥30, n (%)	10 (26%)	3 (18%)	1 (25%)
Prior regimens of therapy in metastatic setting, median (range)	1 (1,6)	2 (1,6)	5 (1, 12)
<i>Pending data entry</i>	2 (5%)	1 (6%)	0 (0%)
1	19 (49%)	6 (35%)	1 (25%)
2	10 (26%)	6 (35%)	0 (0%)
3+	8 (21%)	4 (24%)	3 (75%)

# RLY-2608 – 600 mg BID Dose Selected for Expansion Cohort

## 17 Breast Cancer Patients Treated with RLY-2608 600 mg BID Dose + Fulvestrant



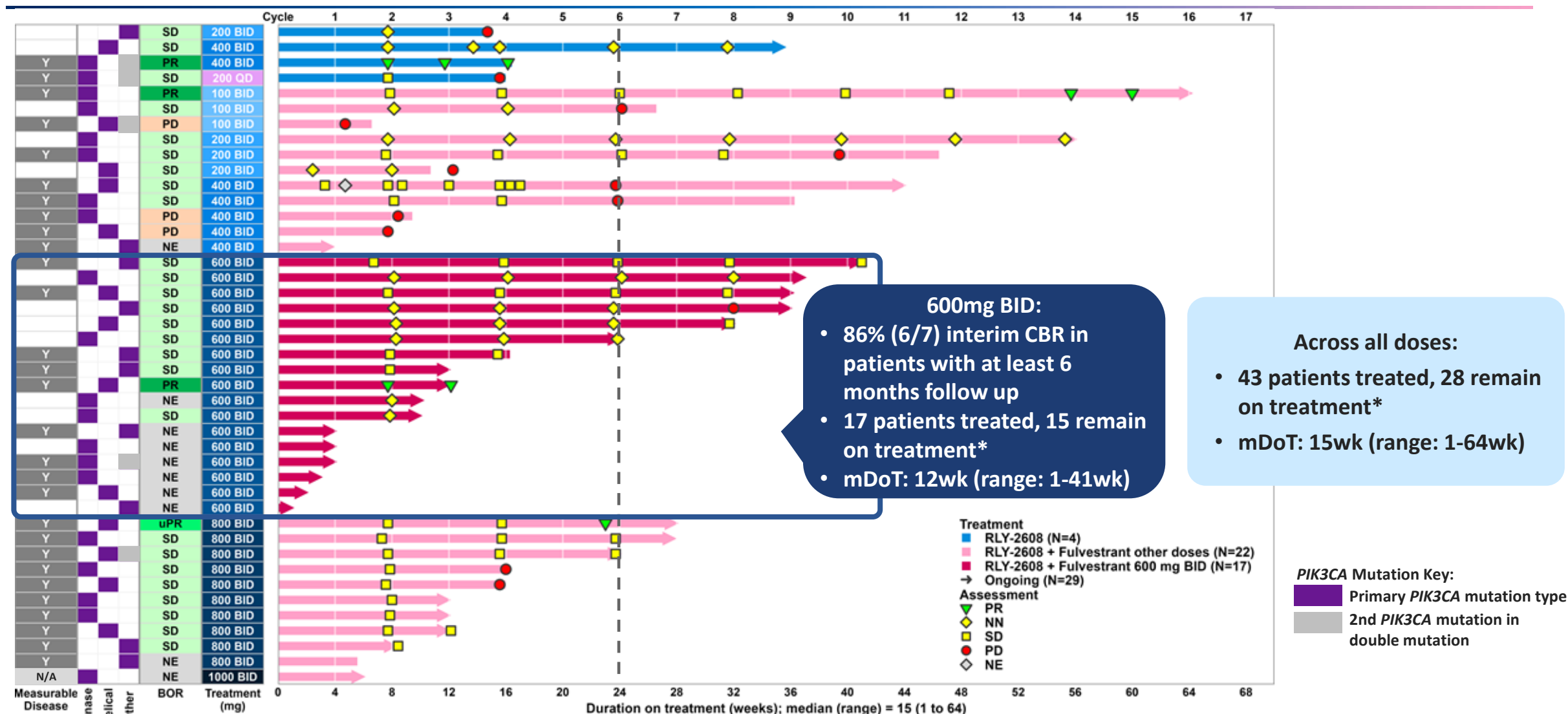
- RLY 2608 + Fulvestrant 600mg BID:**
- 86% (6/7) CBR in patients with at least 6 months follow up
  - Confirmed PR achieved in 1 of 5 efficacy evaluable<sup>1</sup> patients with measurable disease
  - 17 patients treated, 15 remain on treatment\*
  - mDoT: 12wk (range: 1-41wk)

CBR: Clinical Benefit defined as all patients with confirmed complete response or partial response or stable disease ≥24 weeks; evaluable patients started treatment ≥24 weeks prior to the data cutoff

\* Note: one additional pt at 600mg BID dose remains on treatment after PD assessment; 1. Efficacy analysis includes patients with measurable disease who had opportunity for ≥1 tumor assessment or discontinued treatment with <1 tumor assessment

# RLY-2608 – Breast Cancer Disease Control Across Dose Levels

43 Breast Cancer Patients – Measurable and Non-Measurable Disease



CBR: Clinical Benefit defined as all patients with confirmed complete response or partial response or stable disease ≥24 weeks; evaluable patients started treatment ≥24 weeks prior to the data cutoff; N/A: not available as of data cut off, pending data entry

\* Note: one additional pt at 600mg BID dose remains on treatment after PD assessment

© 2024 Relay Therapeutics

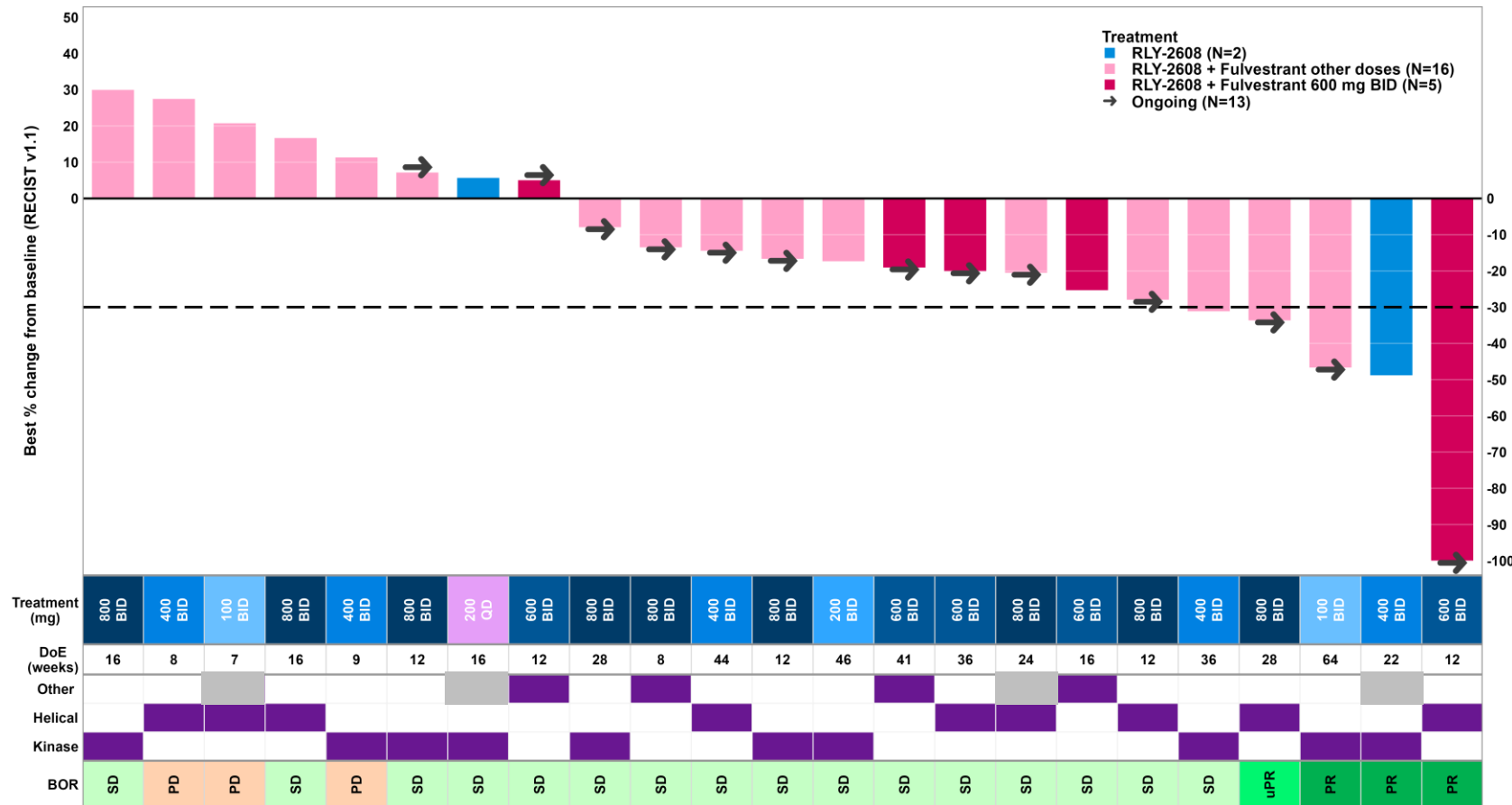
Preliminary data as of 07/24/2023

22

# RLY-2608 – Evidence of Anti-Tumor Activity Supports Selective Target Engagement

## 24 Breast Cancer Patients\* – Measurable Disease Only

### Breast Cancer Patients (RECIST Measurable Disease) N=24\*



- At 600mg BID combo, 80% of patients (4/5) exhibited radiographic tumor reductions
  - 1 pt experienced a partial response and remains on treatment
- Overall, 63% of patients (15/24) exhibited radiographic tumor reductions; 13/24 patients ongoing
- 4 partial responses observed across mono and combo, dose levels and PI3Kα genotypes

BOR = Best Overall Response:

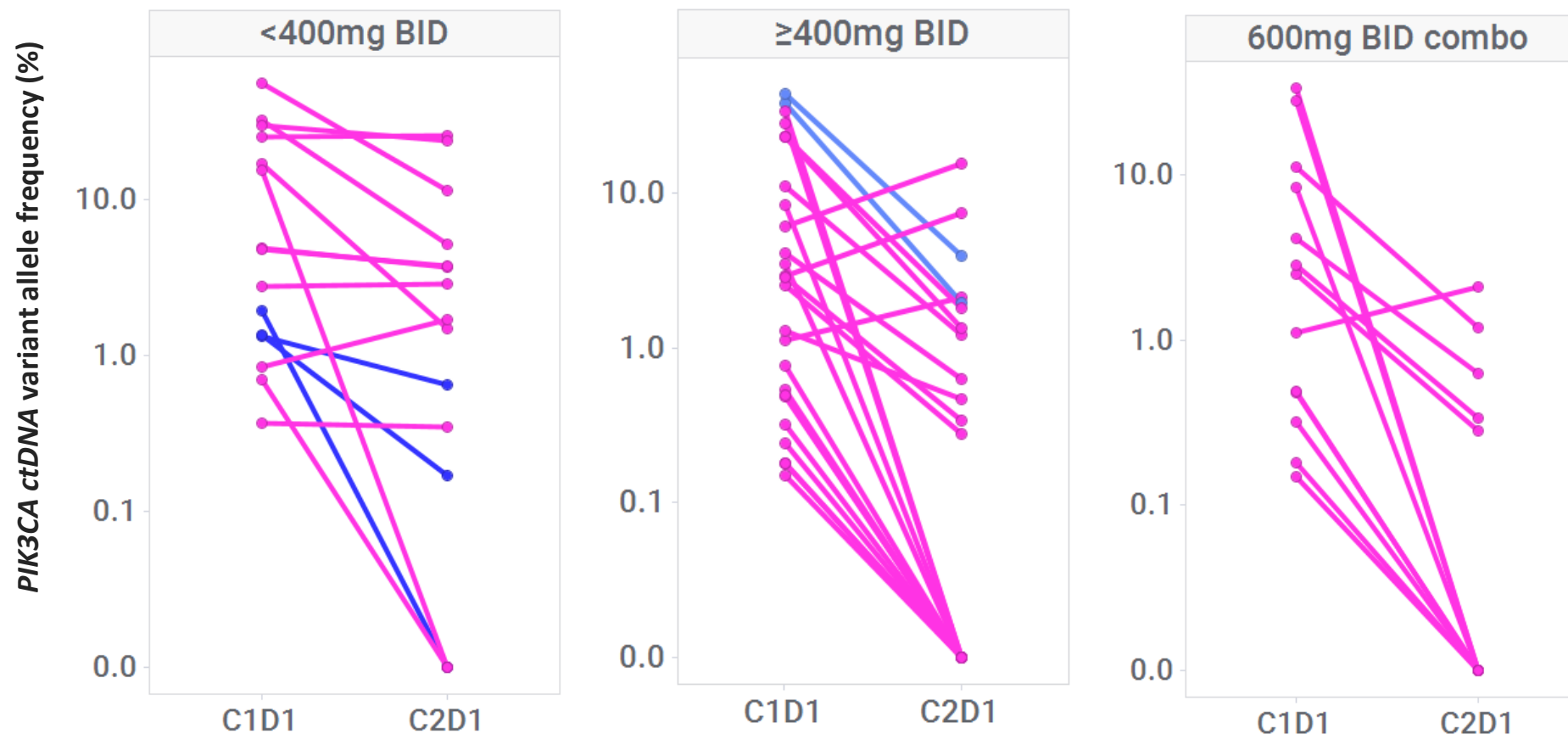
- PR Partial Response
- uPR Unconfirmed Partial Response
- SD Stable Disease
- PD Progressive Disease

PIK3CA Mutation Key:

- Primary PIK3CA mutation type
- 2nd PIK3CA mutation in double mutation

\* one patient discontinued prior to first scan and is not shown on waterfall plot

# RLY-2608 – Mutant PIK3CA Decline Supports Dose Dependent Target Inhibition



- 30 Breast cancer patients with evaluable paired C1D1-C2D1 ctDNA Sample
  - Mono: 3 pt; combo: 27 pt
- 6 patients have  $\geq 2$  PIK3CA mutation
- 26 patients had decline in PIK3CA ctDNA
- 13 patients completely cleared PIK3CA ctDNA by C2D1

**Patients with paired evaluable ctDNA**

**Mono: n=2**

**Combo: n=10**

**Mono: n= 1**

**Combo: n=17**

**Mono: n= 0**

**Combo: n=8**

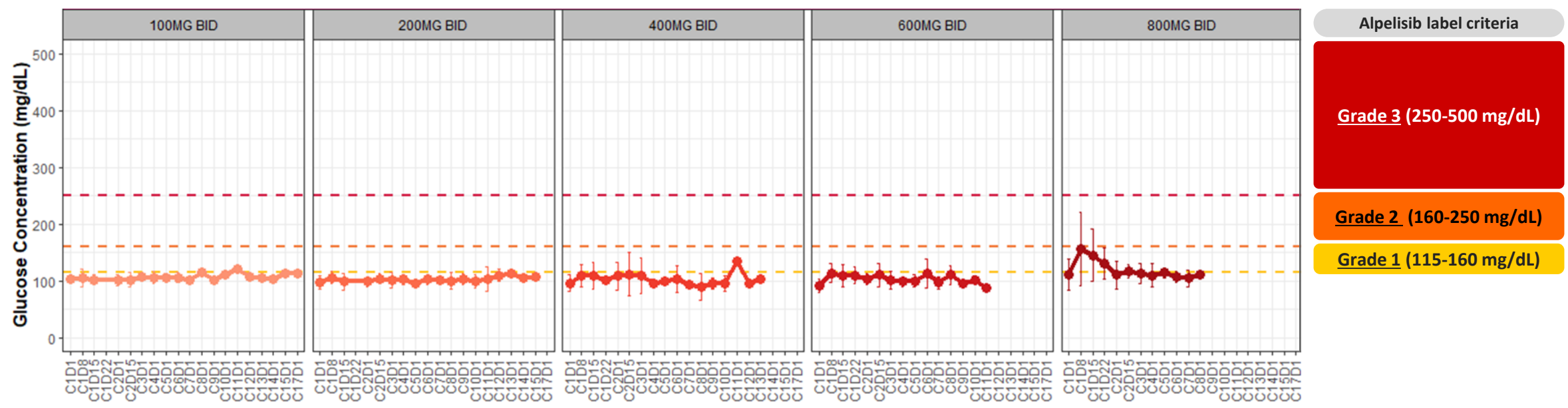
Note: data points at zero are below limits of detection  
Source: Central lab analysis



# RLY-2608 – Limited Observed Impact on Glucose Homeostasis Supports Selectivity

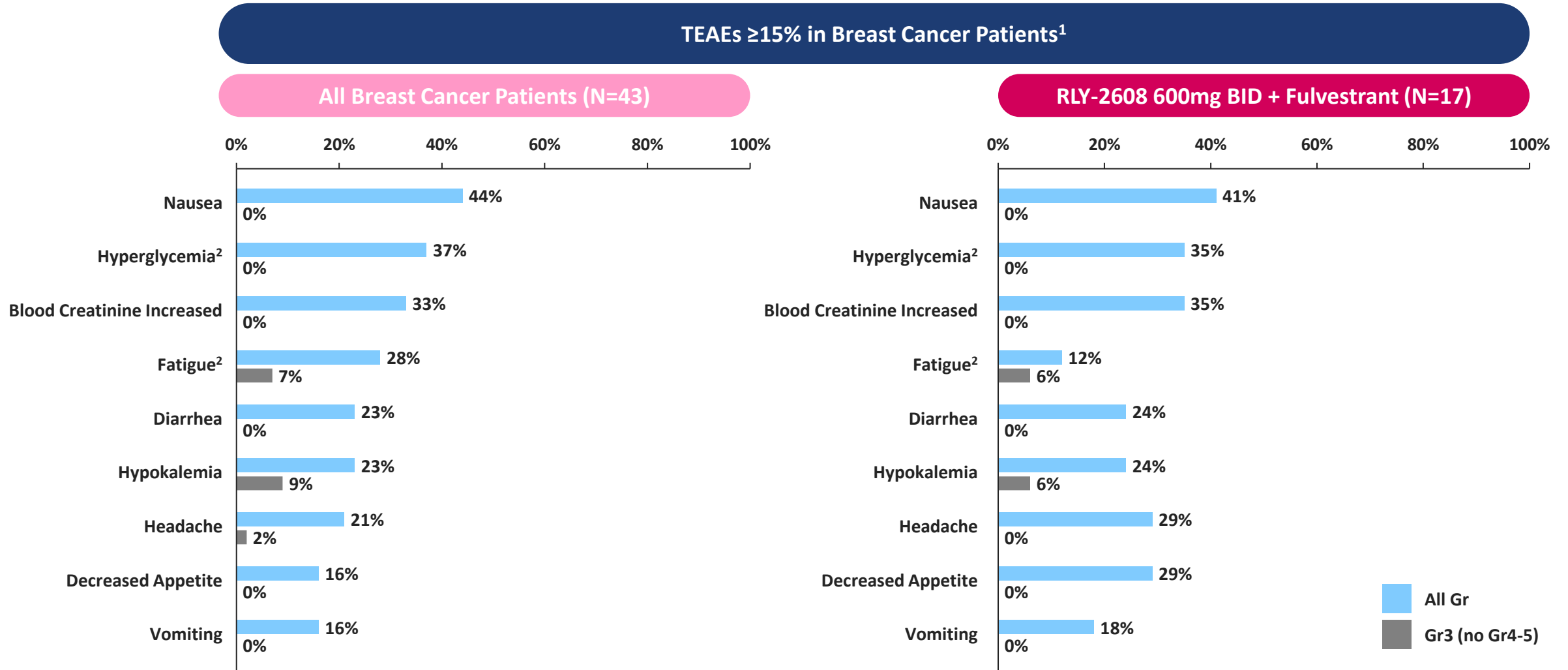


## RLY-2608 + Fulvestrant Combination



Note: one 1000mg BID combo pt not shown; pt had Gr2 glucose elevation per alpelisib label criteria; Data represent mean per cohort +/- standard deviation  
Source: Central lab analysis

# RLY-2608 – TEAEs Generally Consistent with Mutant-Selective Inhibition

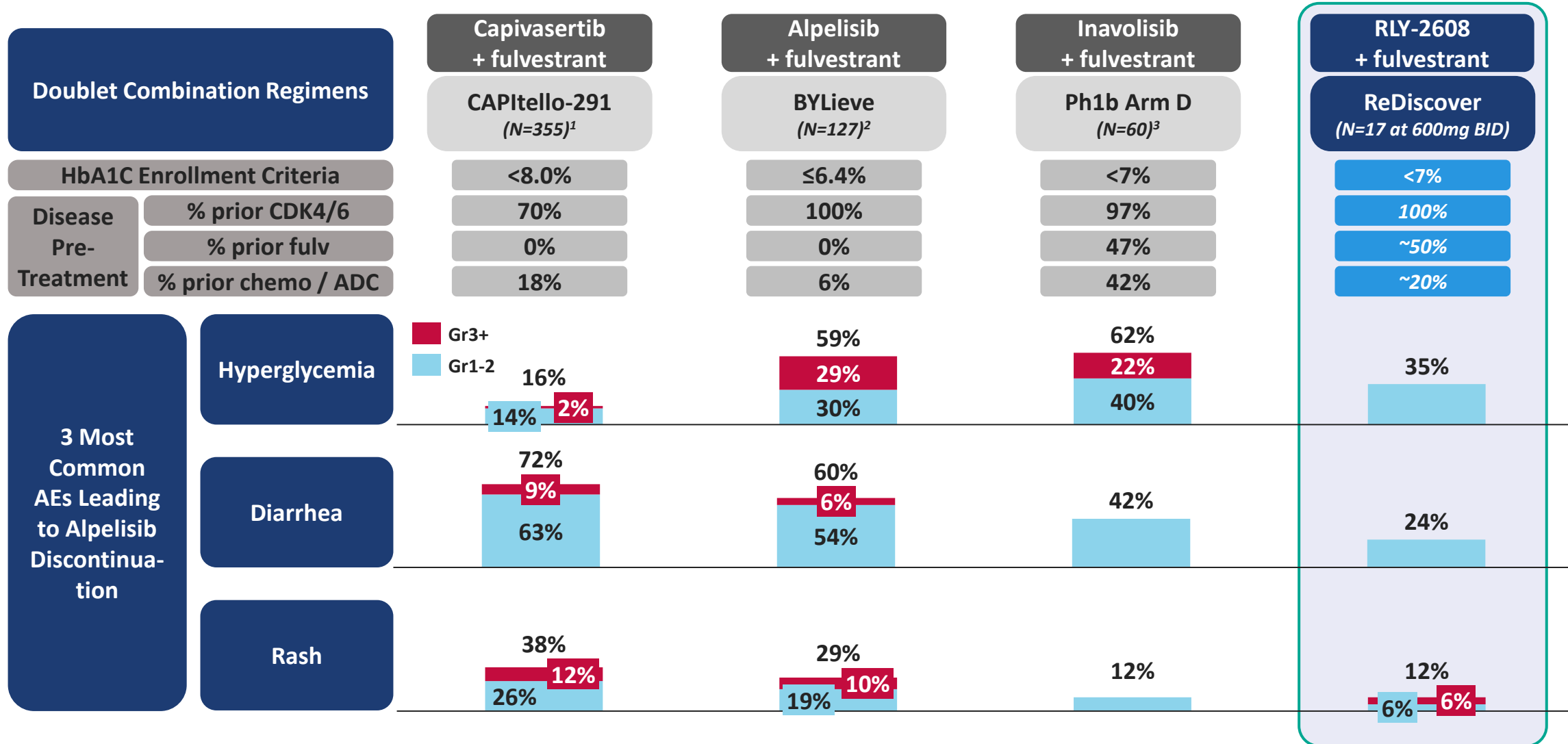


1. TEAEs that occurred in  $\geq 15\%$  of the Breast Cancer Safety Set (N=43) are shown for both populations; 2. Hyperglycemia includes the MedDRA v26.0 Preferred Terms (PT): Hyperglycemia and Blood Glucose Increased, Fatigue includes the PTs: Fatigue and Asthenia.

# RLY-2608 – Safety Profiles of Existing PI3K $\alpha$ Pathway Compounds

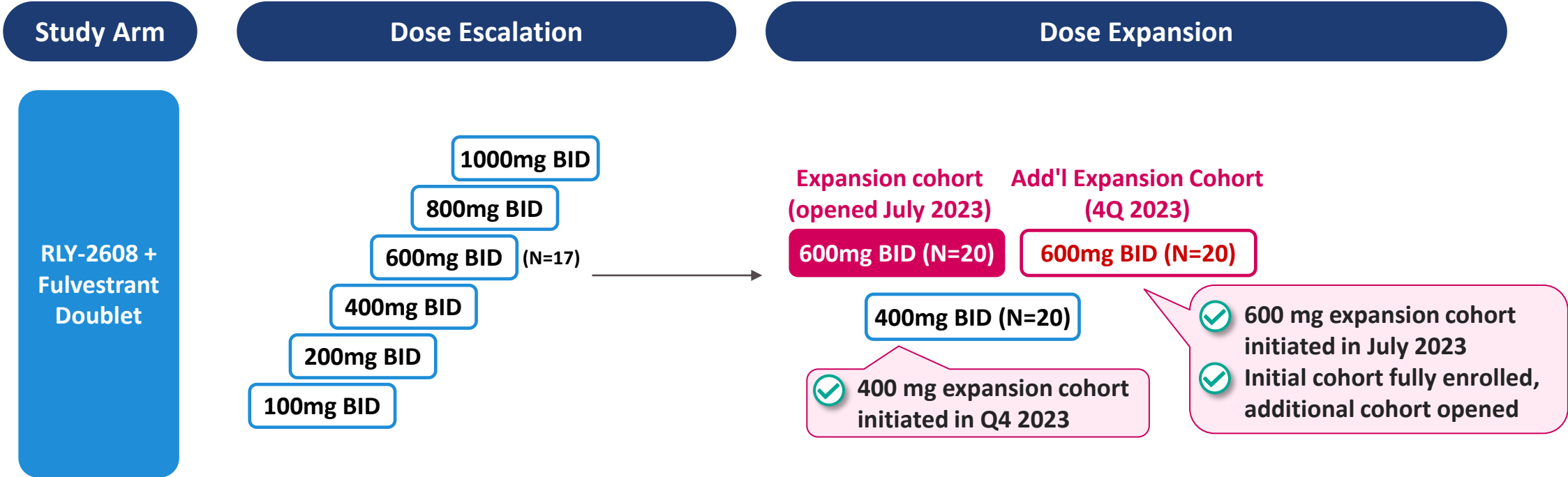
Data below are not from head-to-head studies.

Cross-trial data interpretation should be considered with caution as it is limited by differences in study population and many other factors.



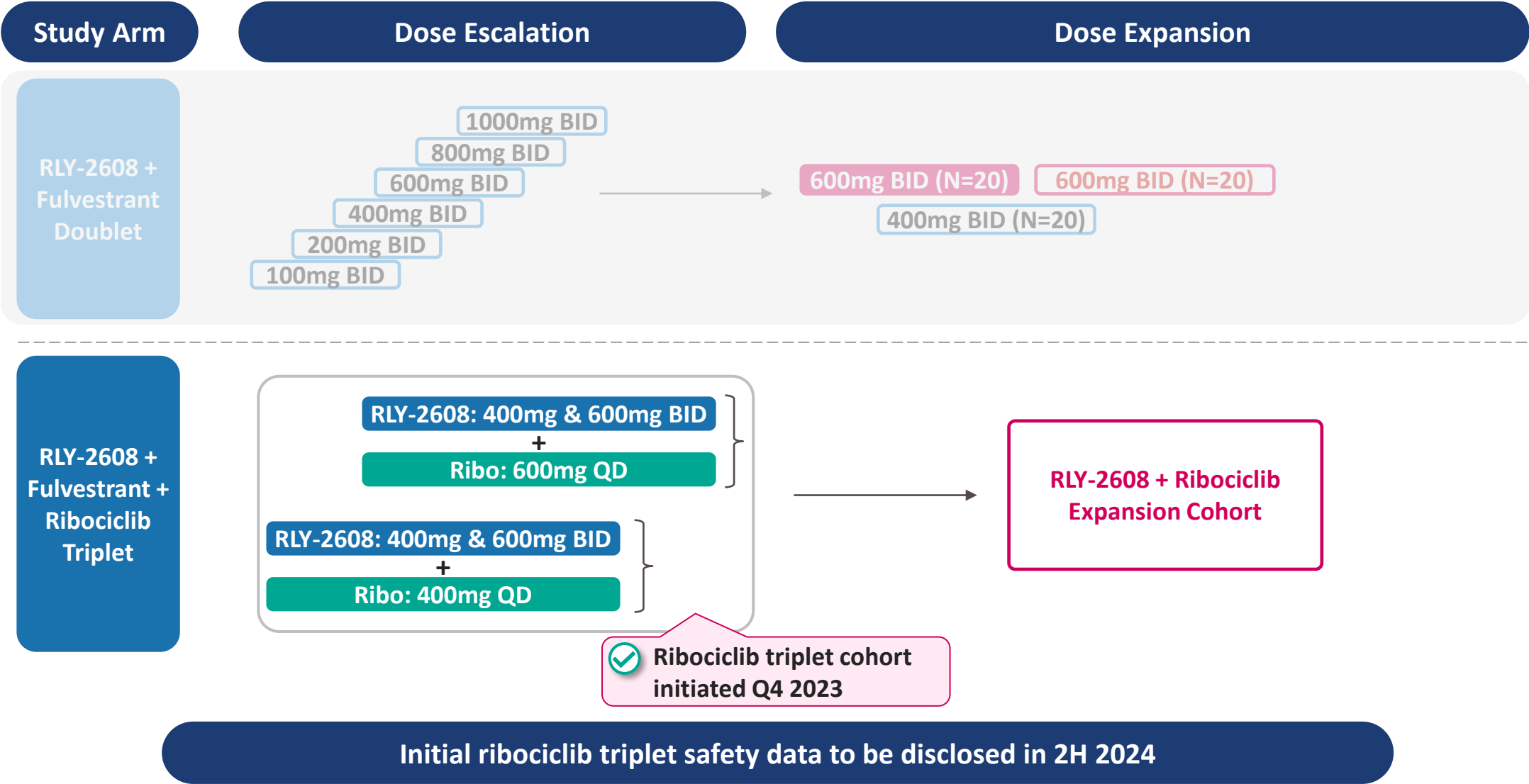
Sources: 1. Turner N Engl J Med 2023; 388:2058-2070; 2. Rugo 2021 Lancet Oncol 22:489; 3. SABCS 2021 #P5-17-05; \* For PIK3CAmut HR+/HER2- breast cancer in combination with fulvestrant; Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.

# RLY-2608 – ReDiscover Combination Trial Design

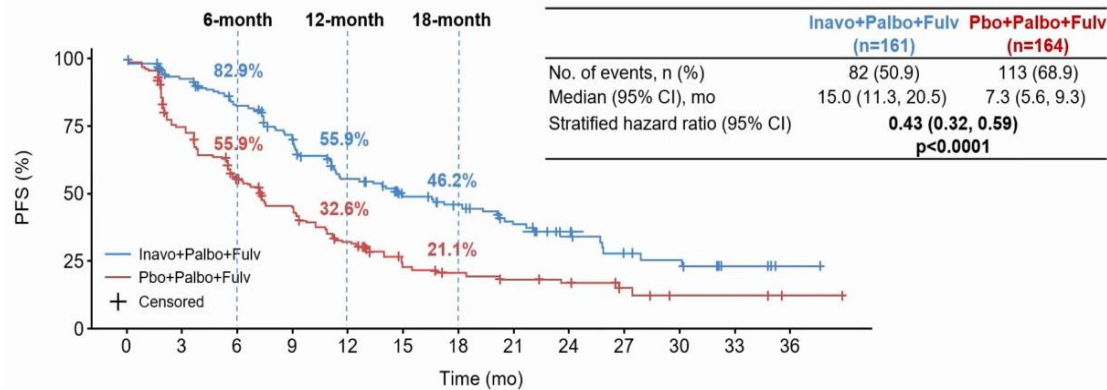


Next RLY-2608 doublet data to be disclosed in 2H 2024 after further data maturation

# RLY-2608 – ReDiscover Combination Trial Design



## Inavolisib + Palbociclib + Fulvestrant doubled PFS vs. Fulvestrant + Palbociclib Alone



**15.0mo mPFS**  
Vs. 7.3mo pbo

**HR: 0.43**

Demonstrated manageable safety in heavily selected, metabolically stable patient population

## However, INAVO120 Ph 3 Trial Included Only a Subset of 1L HR+/HER2- Breast Cancer

INAVO120  
Enrollment  
Restrictions

1 Endocrine Resistant Only

40% of 1L BC pop.

2 Non-Pre-Diabetic or Diabetic

50% of US Pop.

US 1L BC Population which  
meets enrollment criteria

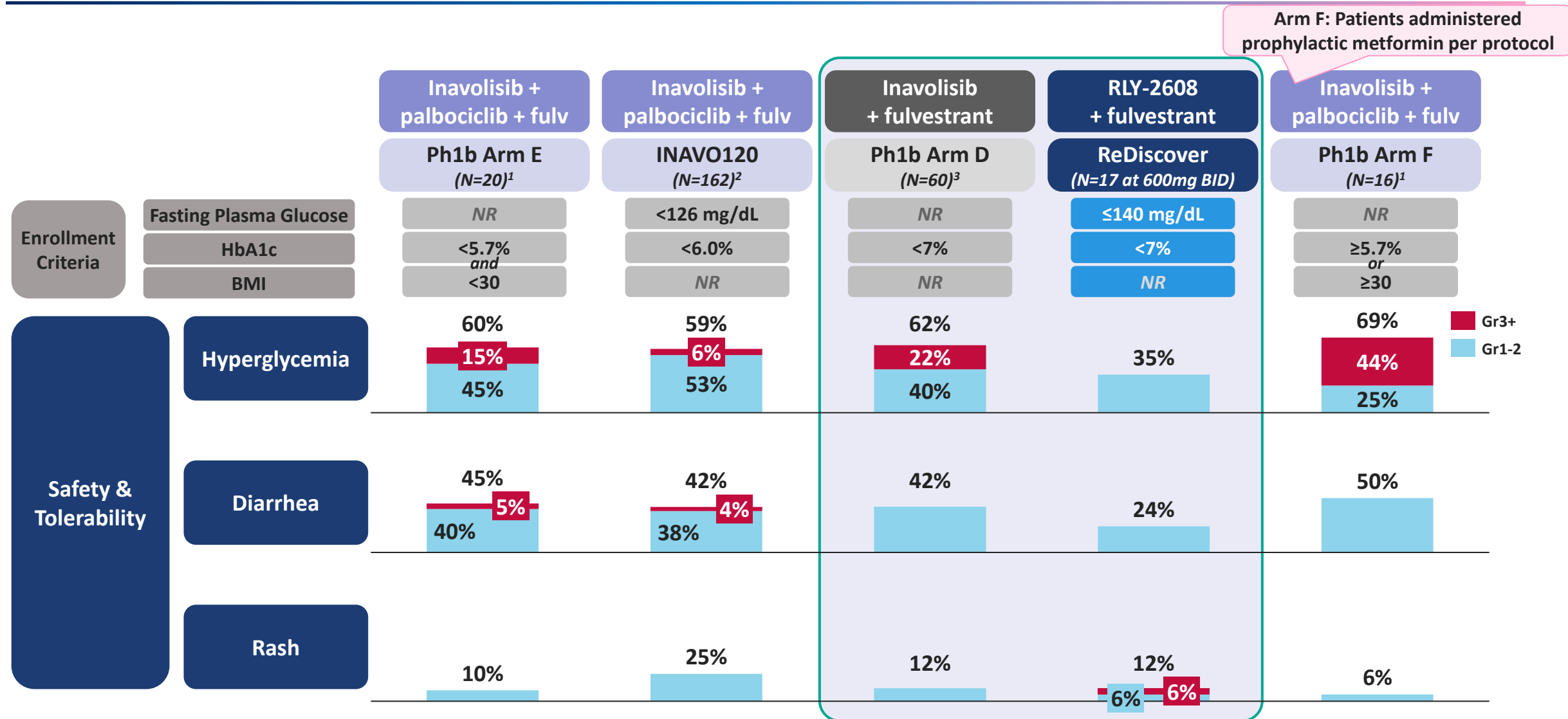
~20%

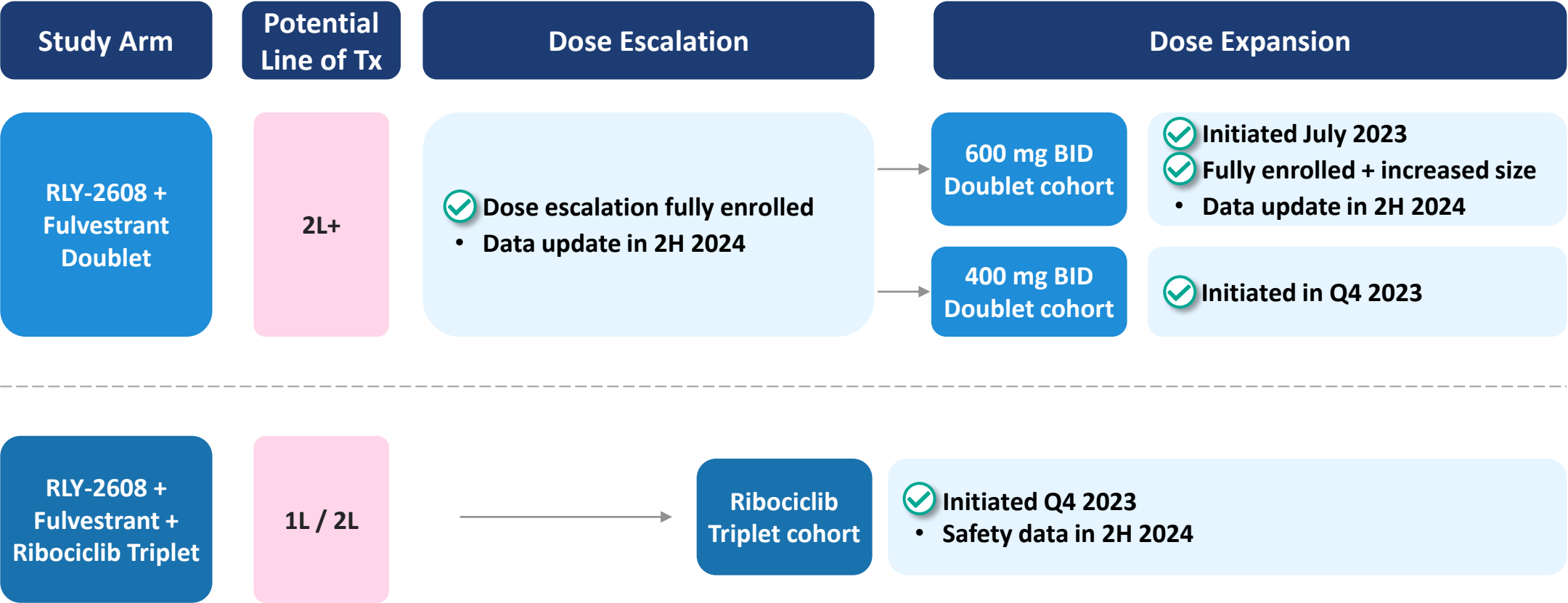
Metabolically selected  
patients limit market size

# RLY-2608 – Safety Profiles of Existing PI3K $\alpha$ Pathway Compounds

Data below are not from head-to-head studies.

Cross-trial data interpretation should be considered with caution as it is limited by differences in study population and many other factors.





Next RLY-2608 data update in 2H 2024



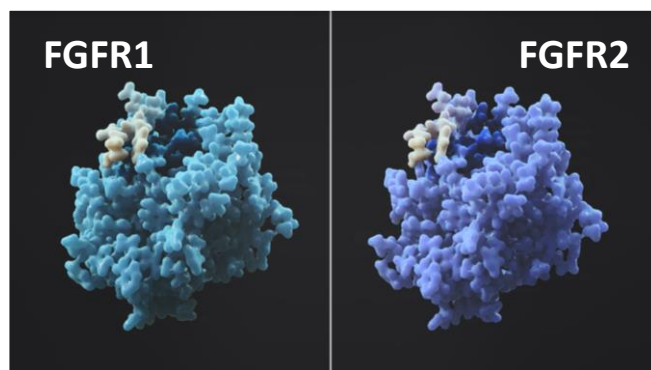
# Relay Tx – Broad Precision Medicine Pipeline



Target	Program	Preclinical	Early Clinical	Late Clinical
PI3Kα franchise	Monotherapy			
	RLY-2608 PI3Kα <sup>PAN</sup> Endocrine Tx (ET) doublet			
	CDK4/6i + ET triplet			
	RLY-5836 (PI3Kα <sup>PAN</sup> )Dose Escalation	Deprioritized		
	PI3Kα <sup>H1047R</sup>			
FGFR2	Lirafugratinib (RLY-4008)			
Solid Tumor	2 programs			
Genetic Disease	2 programs			
CDK2	RLY-2139	Paused; IND ready		
ERα	ERα Degradar	Paused at DC		
SHP2	Migoprotafib (GDC-1971) Genentech <small>A Member of the Roche Group</small>	3 ongoing combo studies		

# FGFR2 – Limitations of Current CCA and Non-CCA Treatment Options

## FGFR1-4 static structures look the same



## No FGFR2-targeted therapy available

Pan-FGFRi's lead to high rates of off-target toxicity, esp. for FGFR1,4

FDA Approved Compound <sup>1</sup>	% of Patients with Hyperphosphatemia	% of Patients with Diarrhea
Pemigatinib	93%	39%
Futibatinib	88%	33%
Erdafitinib	71%	59%

Chemo and other late line therapies also have high rates of AEs and dose modifications

## Efficacy limited by off-target tox

### CCA

**36-42%** ORR in currently approved tx<sup>1</sup>  
(in fusion+ CCA, FGFRi-naïve pt)

### Non-CCA Solid Tumors

**0-15%** ORR in approved late-line tx<sup>2</sup>  
(based on NCCN guidelines)

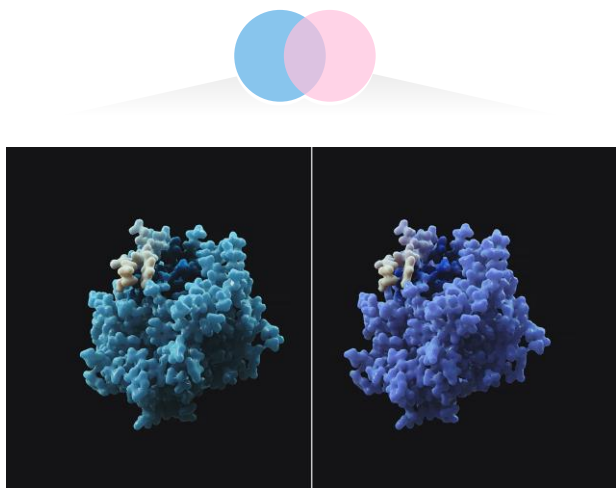
mPFS 1-5mo in  
non-CCA solid tumors

1. Sources: Pemigatinib – prescribing information; futibatinib – prescribing Information; erdafitinib – prescribing information; (note: AEs are reflective of respective label indications); 2. Reflects reported ORRs in key randomized studies evaluating NCCN recommended regimens for recurrent/metastatic patients (second/third line or later) for the following tumor types: HR+ breast cancer, gastric cancer, pancreatic cancer, NSCLC, ovarian cancer, and head and neck

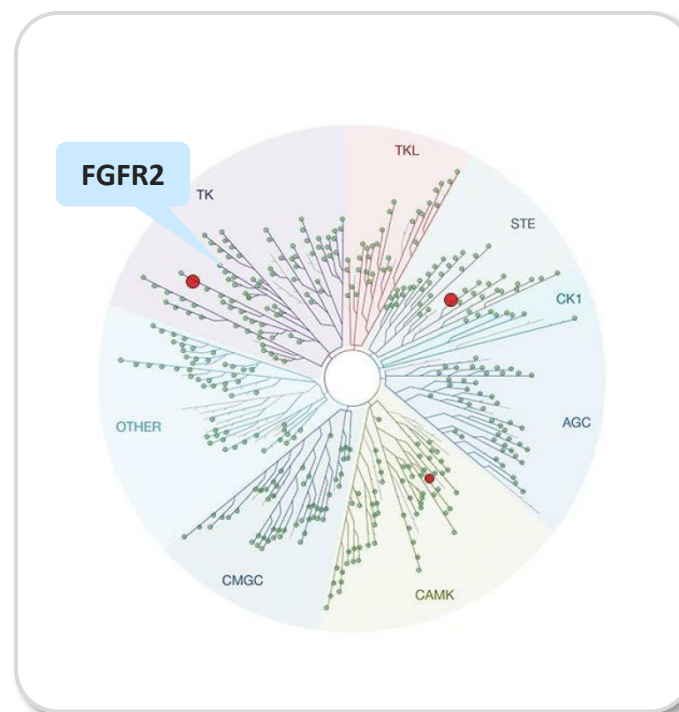
# Lirafugratinib (RLY-4008) – Embodies The Power of Our R&D Engine

## Motion Based Drug Design...

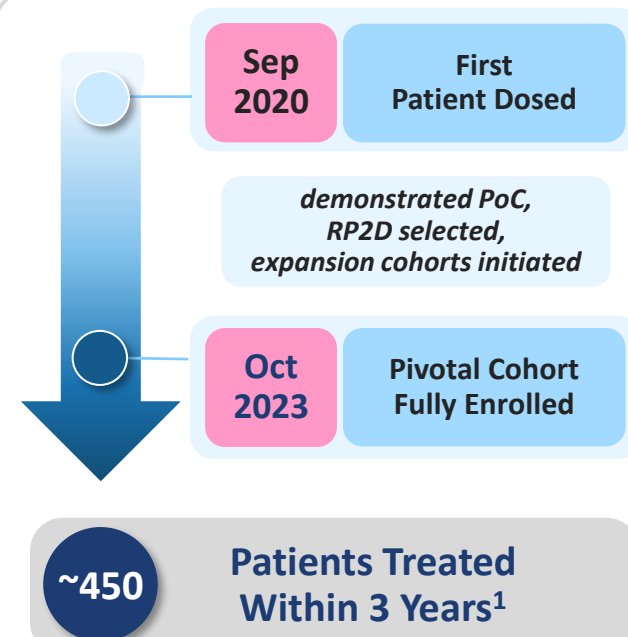
### Relay Approach



## ...Created First Known Selective FGFR2



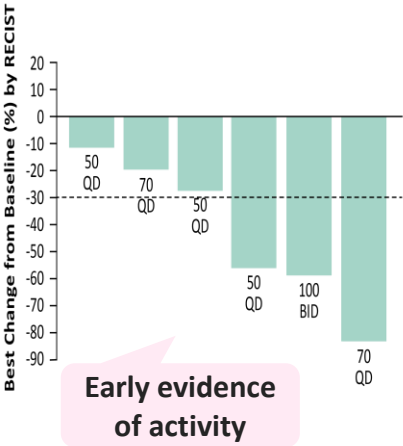
## Strong Clinical Execution Drives Rapid Pathway to Potential Registration



# Lirafugratinib (RLY-4008) – Evolution of Data Maturity

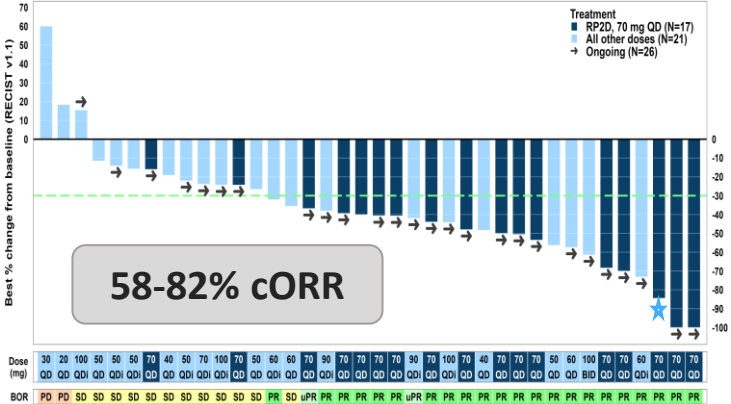


## 2021 – Initial Clinical Activity Demonstrated<sup>1</sup>



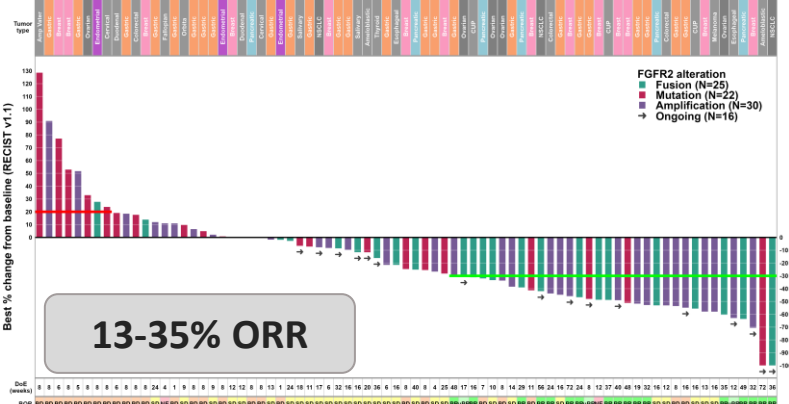
**N = 49**  
(All solid tumors)

## 2022 – Interim Efficacy in CCA Fusion<sup>2</sup>



**N = 38**  
(fusion+, FGFRi-naïve CCA)

## 2023 – Interim Tumor Agnostic Efficacy<sup>3</sup>



**N = 84**  
(non-CCA solid tumor expansion cohorts)

## Tumor agnostic data and regulatory update in 2H 2024

Data presented at: 1. 2021 ENA Meeting (data as of 09 September 2021); 2. 2022 ESMO Congress (data as of 01 August 2022); 3. 2023 Triple Meeting (data as of 27 September 2023)  
© 2024 Relay Therapeutics

# Relay Tx – Broad Precision Medicine Pipeline



Target	Program	Preclinical	Early Clinical	Late Clinical
PI3Kα franchise	Monotherapy			
	RLY-2608 PI3Kα <sup>PAN</sup> Endocrine Tx (ET) doublet			
	CDK4/6i + ET triplet			
	RLY-5836 (PI3Kα <sup>PAN</sup> )Dose Escalation	Deprioritized		
	PI3Kα <sup>H1047R</sup>			
FGFR2	Lirafugratinib (RLY-4008)			
Solid Tumor	2 programs			
Genetic Disease	2 programs			
CDK2	RLY-2139	Paused; IND ready		
ERα	ERα Degradar	Paused at DC		
SHP2	Migoprotafib (GDC-1971) Genentech <small>A Member of the Roche Group</small>	3 ongoing combo studies		

# SHP2 – Genentech Global Collaboration for Migoprotafib (GDC-1971)



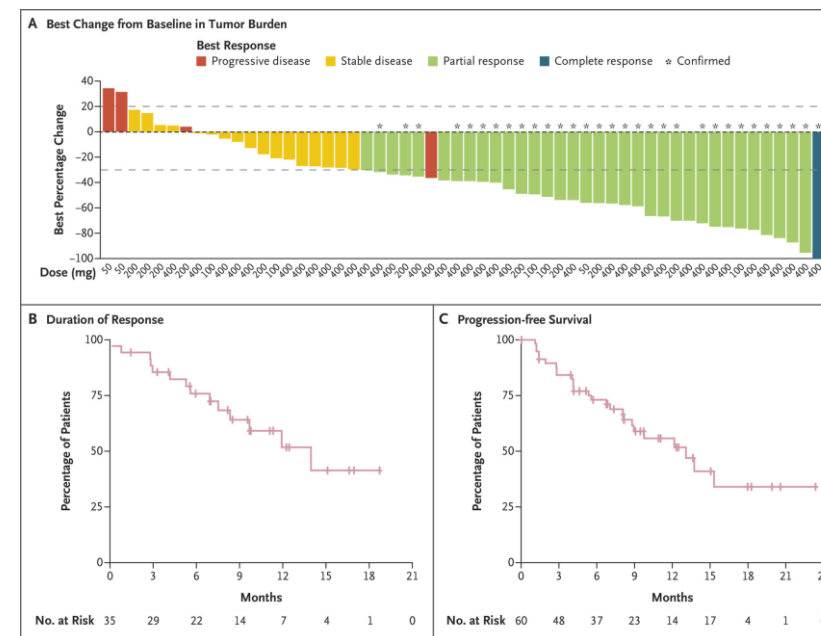
## Three ongoing trials with migoprotafib

**Migoprotafib**  
+  
**GDC-6036 (KRAS G12Ci)**  
*initiated July 2021*

**Migoprotafib**  
+  
**Atezolizumab (PD-L1 Ab)**  
*initiated August 2022*

**Migoprotafib**  
+  
**Osimertinib/Cetuximab (EGFRi)**  
*initiated July 2023*

## Clinical Update for GDC-6036 Monotherapy in 2L NSCLC



**ORR: 61% (35/58 patients, across doses, 53% cORR)**  
**mPFS: 13.7mo, mDoR: 11.9mo (39 pts at 400mg)**

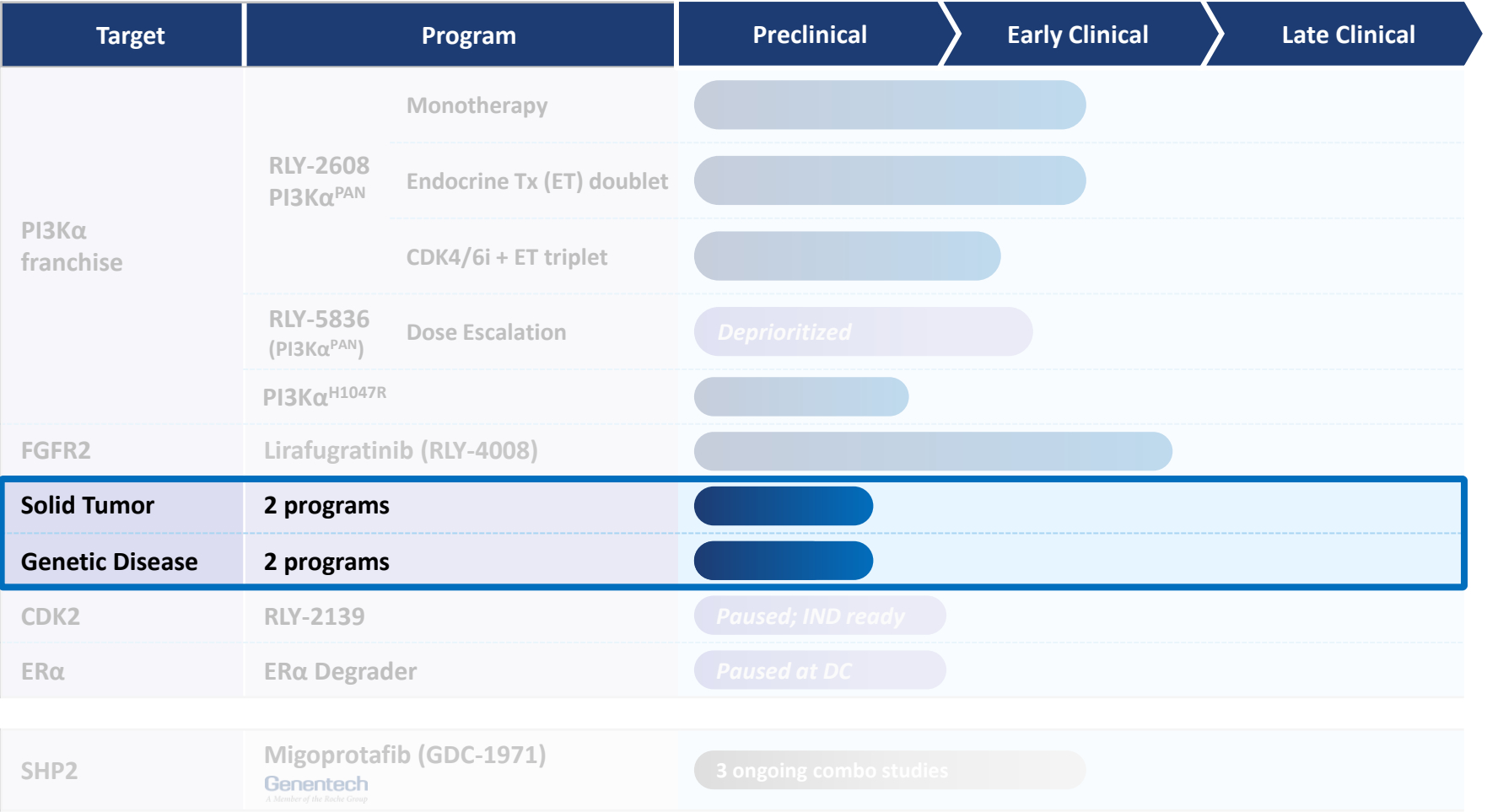
**Collaboration provides meaningful economics to Relay Tx<sup>1</sup>**

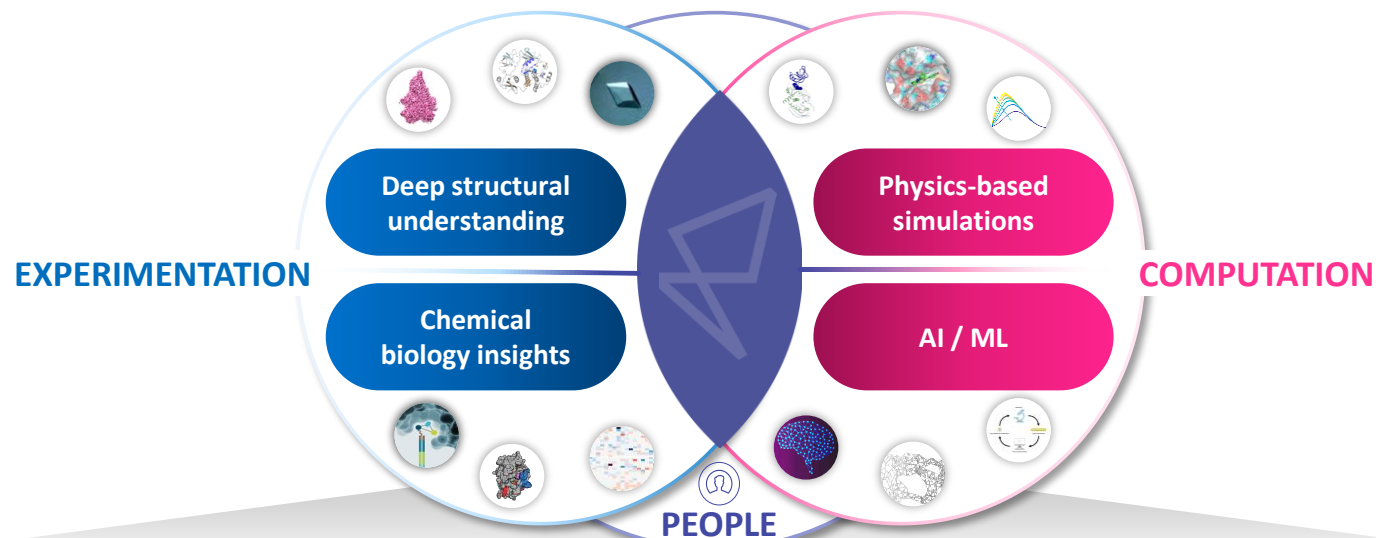
Source: Sacher 2023 N Engl J Med 389:710

1. As of the date of this presentation: \$120 million in upfront & milestone payments received, and eligible to receive up to \$675M in potential additional total milestones, low-to-mid teen royalties on global net sales plus additional royalties upon approval of GDC-1971 and GDC-6036 in combination

© 2024 Relay Therapeutics

# Relay Tx – Broad Precision Medicine Pipeline





## Already Productive Platform...

IND	Compound	Achievement
2019	Migoprotafib <sup>1</sup> (SHP2)	Partnered with GNE
2020	Lirafugratinib <sup>2</sup> (FGFR2)	Enrolled ~450+ pt
2021	RLY-2608 (PI3Kα)	Clinical POC
2022	RLY-5836 (PI3Kα)	Clinical Start
2023	RLY-2139 (CDK2)	IND Ready



## ...Potential To Generate More Assets In Future

Pipeline	7+ pre-clinical programs
TAs	Oncology and Genetic Disease
Modalities	Inhibitors, chaperones and degraders
Platform	Expansion of integrated tools & capabilities



# Relay Tx – Broad Precision Medicine Pipeline



Target	Program	Preclinical	Early Clinical	Late Clinical	Annual US Patient #
PI3Kα franchise	Monotherapy				~10-71K breast cancer ~76-243K all solid tumors
	RLY-2608 PI3Kα <sup>PAN</sup> Endocrine Tx (ET) doublet				
	CDK4/6i + ET triplet				
	RLY-5836 (PI3Kα <sup>PAN</sup> ) Dose Escalation	Deprioritized			~4-27K breast cancer ~15-50K all solid tumors
	PI3Kα <sup>H1047R</sup>				
FGFR2	Lirafugratinib (RLY-4008)				~11-35K <sup>4</sup>
Solid Tumor	2 programs				To be announced
Genetic Disease	2 programs				To be announced
CDK2	RLY-2139	Paused; IND ready			~35K <sup>2</sup>
ERα	RLY-1013 (Degradar)	Paused at DC			~30-205K <sup>3</sup>
SHP2	Migoprotafib (GDC-1971) Genentech <small>A Member of the Roche Group</small>	3 ongoing combo studies			~36-69K <sup>5</sup>

**Note:** Unless otherwise indicated, patient #'s refer to total annual number of US patients with late-line cancers compared to comprehensive annual incidence that may be amenable to treatment with our programs

1. Unless otherwise indicated, all breast cancer patient numbers refer to HR+/HER2- breast cancer tumors; 2. ~35K HR+/HER2- breast cancer patients expected to receive CDK 4/6 inhibitors in adjuvant setting, first-line setting, and second-line setting in 2024, per Decision Resources Breast Cancer Market Forecast report dated November 2023; 3. HR+/HER2- US late-line breast cancer patients compared to HR+/HER2- US incident breast cancer patients; 4. FGFR2 altered late-line solid tumors compared to comprehensive annual FGFR2 altered incident solid tumors including additional FGFR gene fusions and rearrangements resulting from truncation of the protein at exon 18 and all breast cancer patients with FGFR2 alterations; 5. SHP2 combo only includes KRAS G12C in lung and colorectal, EGFR mutations in lung, and ALK fusions in lung

# Relay Tx – Capital, Team & Execution Focus to Deliver on Milestones



## 2024 Corporate Objectives

**RLY-2608 Doublet**  
(PI3K $\alpha$ )

- Additional clinical data in 2H 2024

**RLY-2608 Triplet**  
(PI3K $\alpha$ )

- ✓ Ribociclib triplet initiation in Q4 2023
- Ribociclib triplet safety data in 2H 2024

**Lirafugratibnib (RLY-4008)**  
(FGFR2)

- Tumor agnostic data and regulatory update in 2H 2024

**Pre-clinical Pipeline**  
(Targets unnamed)

- New program(s) to be disclosed in 2024
- 7+ undisclosed programs in preclinical development and additional early-stage efforts across platform

**Migoprotafib (GDC-1971)**  
 (SHP2)

- Three ongoing combination trials  
\*Genentech controls data disclosures

Goal is a first- or best-in-class profile

## Significant Capital to Achieve Goals

~\$750M

Cash, cash equivalents and investments as of the end of 4Q 2023

Expected to be sufficient to fund current operating plan  
into 2H 2026

# Relay Tx 2022 ESG Report – Continuing Our ESG Journey



## Relay Tx's 2<sup>nd</sup> ESG Annual Report



### Patients

4 clinical programs

Committed to clinical trial patient safety

Committed to product safety and quality

*Note: Relay Tx is a development stage company*

### Community



Our patients / future patients



Our community in Cambridge and the broader Boston area



The next generation of scientists

### People

93% of employee respondents "would recommend Relay Tx as a great place to work"

Turnover below industry average rates

Diversity & inclusion advisory group

Training and development opportunities

Equitable compensation

### Environment



Responsible energy consumption\*



Reducing water consumption



Hazardous and lab waste management



Non-hazardous waste management

\*Efforts to reduce energy consumption lend to our ambitions to limit carbon emissions

### Governance

8 Directors Total\*

The Nom/Gov and Audit Committees oversee ESG efforts, with the full BOD getting ~quarterly updates

38%

Racial/Ethnic Diversity

38%

Women

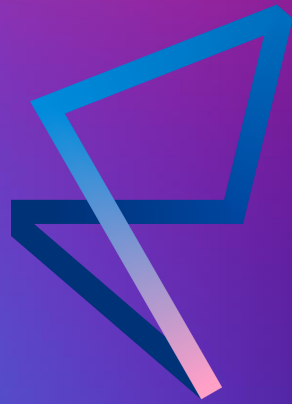
5yrs

Average Tenure

88%

Independence  
(Separate CEO and Chair Role)

\*As of December 2022



**RELAY**<sup>®</sup>  
THERAPEUTICS