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T H E R A P E U T I C S

Vascular Anomalies Update

May 2026

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Zovegalisib – Potential to Address 3 Large Commercial Opportunities



2L Breast Cancer

1L Breast Cancer

Vascular Anomalies

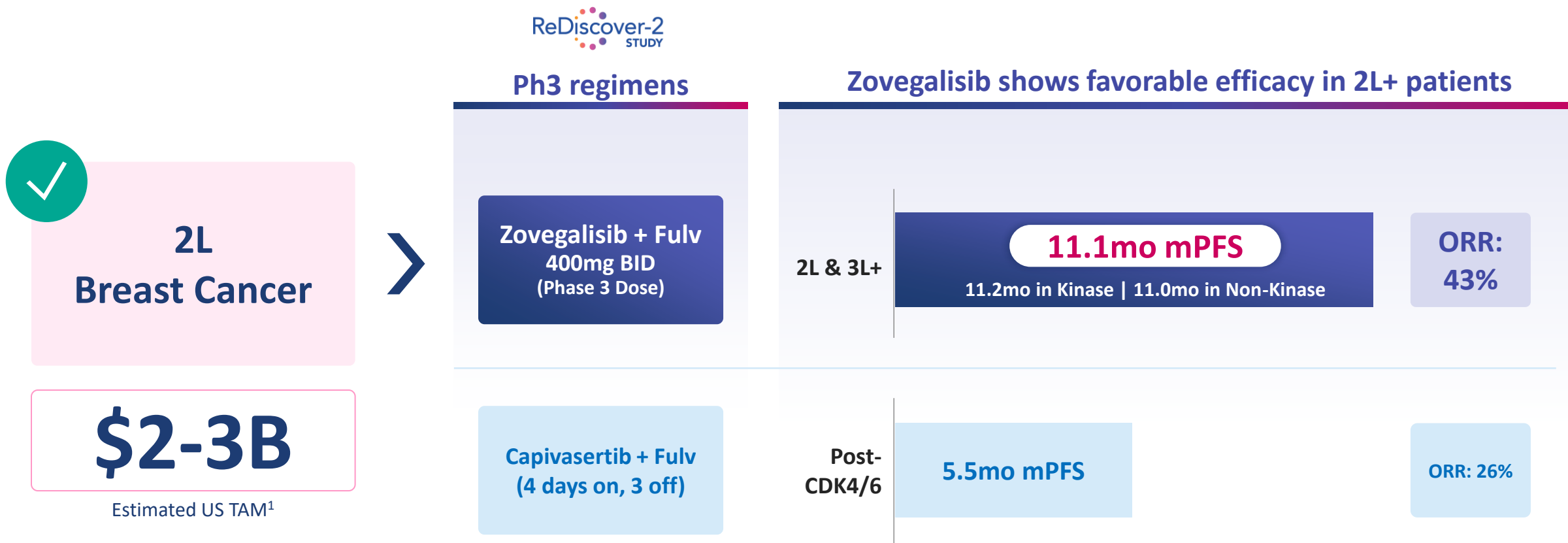
Estimated
US TAM

\$2-3B

\$7-8B

\$6-8B

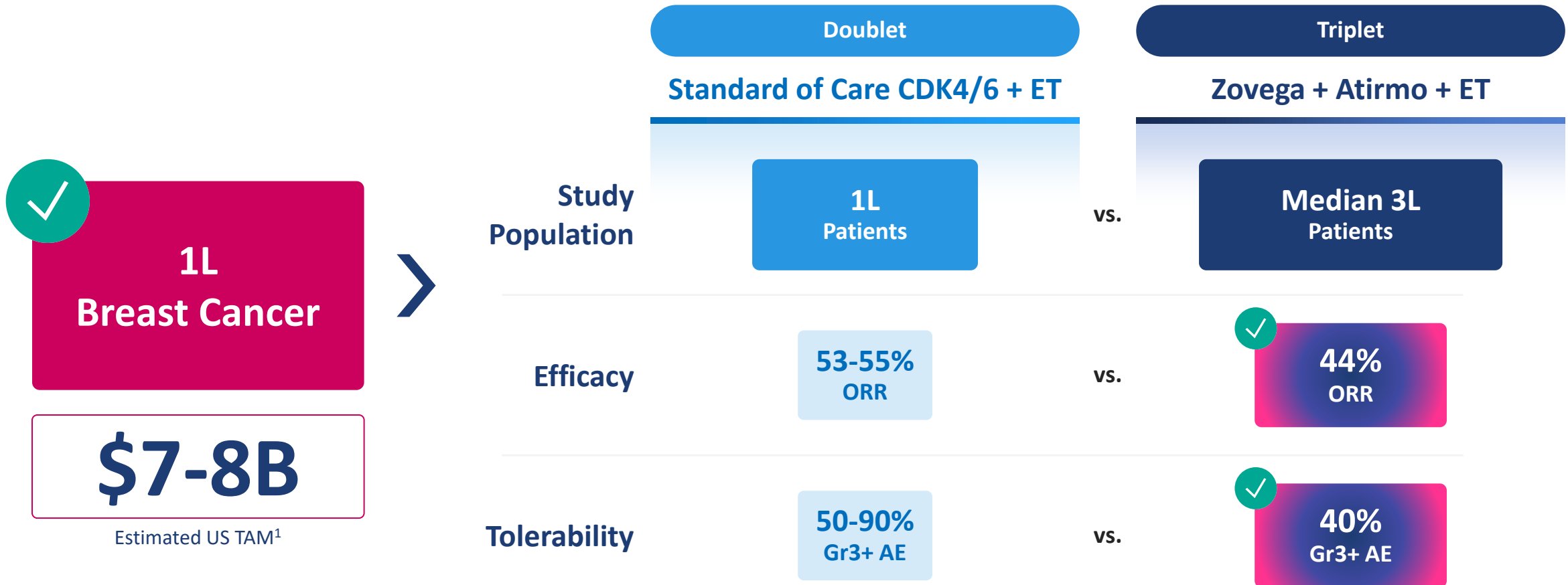
2L Breast Cancer – Ph1/2 Data at Ph3 Dose Showed Clinically Meaningful PFS



Interim zovegalisib data support ongoing Phase 3 trial against capivasertib

1. TAM calculated based on market benchmarks and internal analysis; Sources: ReDiscover Ph1/2 preliminary data as of 1/13/2026; Capi + fulv Ph3 data from CAPItello-291, Turner N Engl J Med 2023; 388:2058-2070. Note: These data are derived from different clinical trials at different points in time, with differences in molecule composition, trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.

1L Breast Cancer – Establishing Zovega + Atirmo + AI Combinability for 1L Use



Zovega + Atirmo + AI selected as go-forward 1L regimen; Supply agreement signed with Pfizer for atirmo; Trial intended to initiate in early 2027

✓

Vascular Anomalies

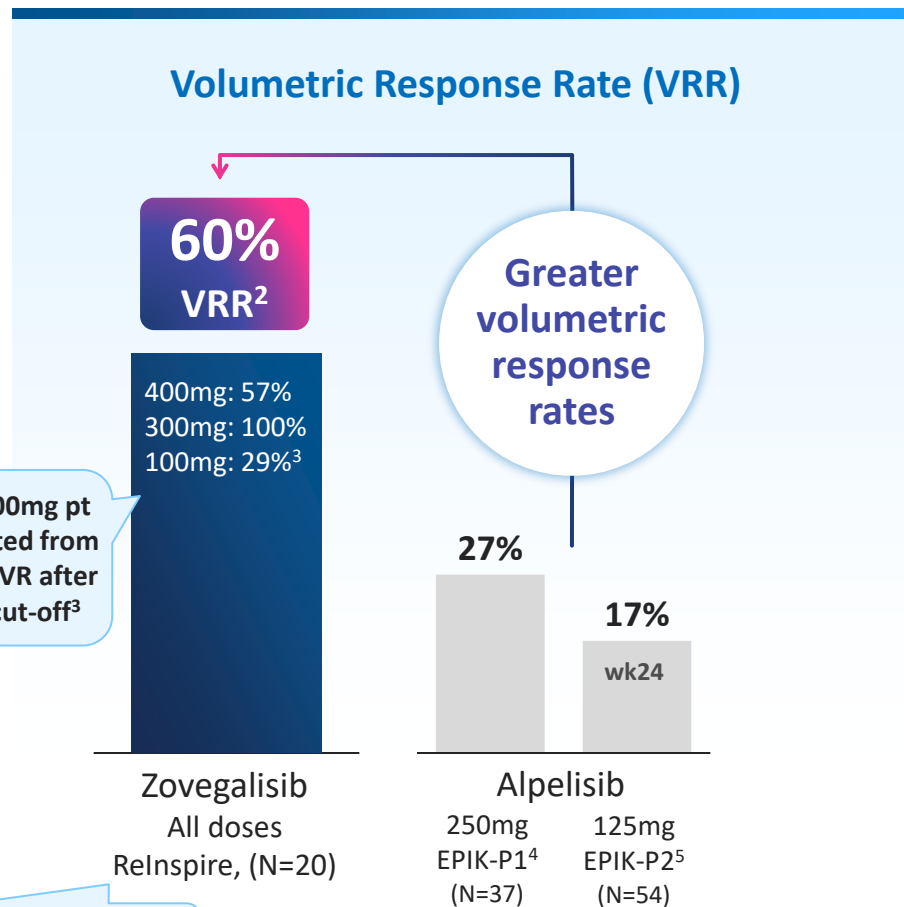
\$6-8B

Estimated US TAM¹

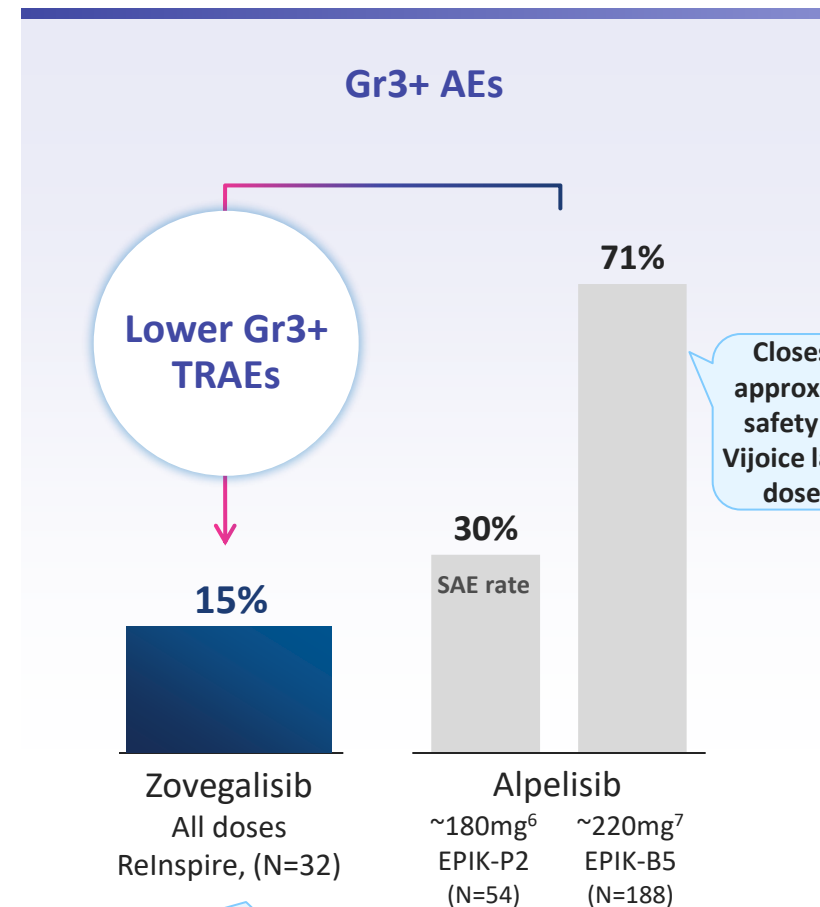
One 100mg pt converted from SD to uVR after data cut-off³

89% of patients with investigator-reported clinical improvement (IGIC at week 12)

Initial Efficacy Data



Initial Tolerability Data



Gr3 hyperglycemia: 1 pt (3%) (pre-diabetic pt at 400mg)

ReInspire median follow-up: 14 weeks

ReInspire preliminary data as of 04/15/2026 ⁶

1. TAM calculated based on market benchmarks and internal analysis; 2. Includes both confirmed and unconfirmed responses. 3. After the data cut-off date, one 100mg BID patient that did not have a volumetric response as of the data cut-off date has converted to an unconfirmed response, resulting in a 100mg BID volumetric response rate of 43% (3/7), a volumetric response rate of 69% (9/13) for patients treated at 300mg BID or 100mg BID, and a volumetric response rate of 65% (13/20) across doses. None of the other response-evaluable patients' response statuses have changed since the data cut-off date; 4. EPIK-P1 as cited in Vioice FDA label, label dose is 250mg QD; 5. EPIK-P2: Canaud 2024 Blood 144:5512 and results from clintrials.gov listing, 125mg QD was starting dose; 6. 180mg dose approximated from rates of dose escalation after week 26 listed on clintrials.gov listing; 7. EPIK-B5: SABCS 2025 #RF7-02, 220mg dose approximated from dose modification data; Gr3 TRAEs = Grade 3+ Treatment-Related Adverse Events, IGIC = Investigator Global Impression of Change scale, SD = Stable Disease, uVR = unconfirmed volumetric response. Note: These data are derived from different clinical trials at different points in time, with differences in molecule composition, trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.

Vascular Anomalies Overview

~170k US patients with PIK3CA-driven Vascular Anomalies

Somatic PIK3CA mutation



drives malformed vasculature



leading to vascular anomalies



Zovegalisib is uniquely positioned to address driver of disease

First mutant-selective PI3K α inhibitor

Initial clinical data showing:

- ✓ Selectivity
- ✓ Tolerability
- ✓ Efficacy

Potential for chronic systemic treatment option

Current treatment options are limited

Local Treatments: temporary, only treat symptoms

Systemic Treatments: non-selective, limited toxicity/efficacy

Large unmet medical need

Vascular Anomalies – Patient Numbers

PIK3CA-driven Vascular Anomalies (VAs)

~170k US patients



Vascular Anomaly Subtypes

Initial clinical focus: ~25k US patients seeking systemic therapy

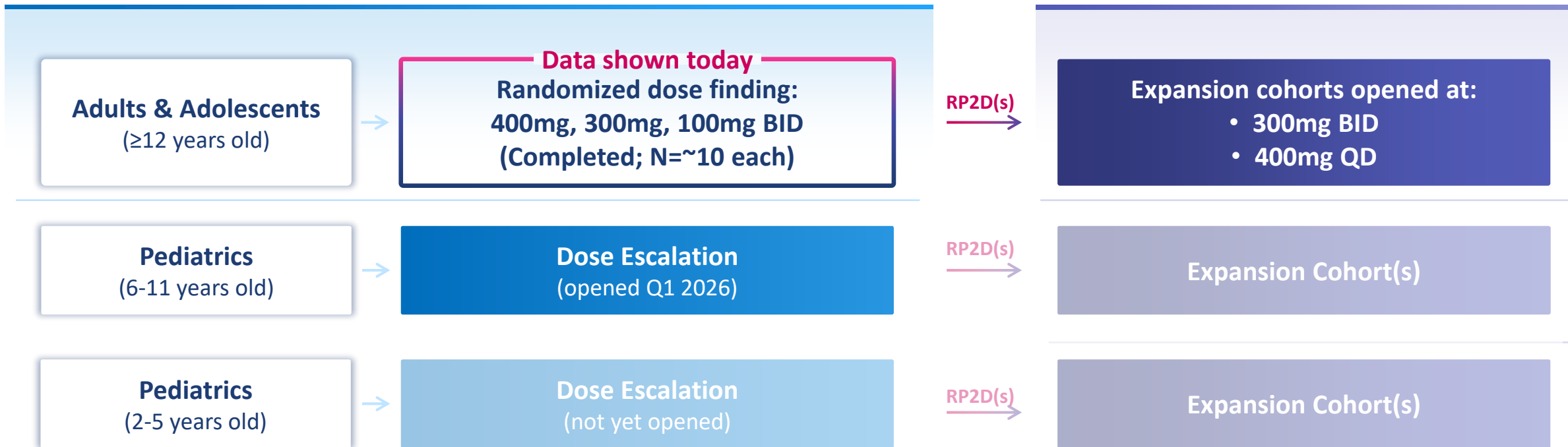
	PIK3CA-Related Overgrowth Spectrum (PROS)	PIK3CA-driven Lymphatic Malformations (LM)	PIK3CA-driven Venous Malformations (VeM)	PIK3CA-driven Cerebral Cavernous Malformations (CCM)
	~5-10k US patients	~60-65k US patients	~20-25k US patients	~50-70k US patients
	25-30% seek systemic tx	20-25% seek systemic tx	15-20% seek systemic tx	25-30% seek systemic tx

Sources: ISSVA classification, NORD, Mayo Clinic, Novartis, Penington et al 2023, Gallagher et al 2022, Luks et al 2015, Limaye et al 2015, Peyre et al 2021, Hong et al 2021, and company primary market research © 2026 Relay Therapeutics

Zovegalisib – Study Design: ReInspire

Part 1: Dose Selection

Part 2: Dose Expansion



Enrollment open in adults & adolescents in part 2 and pediatrics (6-11 y/o) in part 1

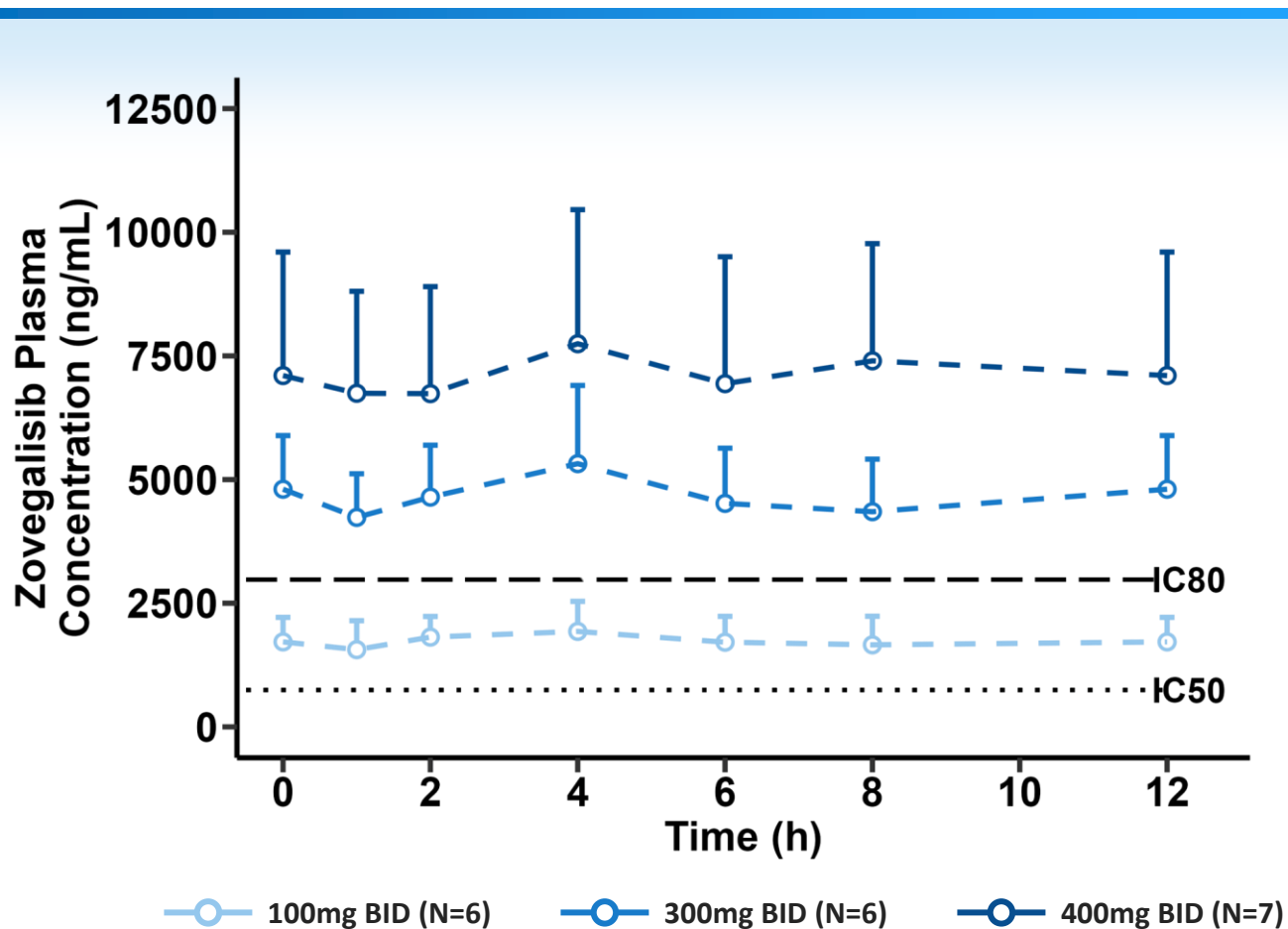
Zovegalisib – ReInspire Trial Demographics

	Total (N=32)	100mg BID (N=11)	300mg BID (N=11)	400mg BID (N=10)
Age (years), median (range)	24.5 (12, 63)	31 (13, 50)	24 (13, 54)	19.5 (12, 63)
12-17 / ≥18, n (%)	10 (31) / 22 (69)	4 (36) / 7 (64)	2 (18) / 9 (82)	4 (40) / 6 (60)
Sex, M/F, n (%)	14 (44) / 18 (56)	6 (55) / 5 (45)	5 (45) / 6 (55)	3 (30) / 7 (70)
Disease Classification, n (%)				
PROS	22 (69)	8 (73)	6 (54)	8 (80)
CLOVES	5 (16)	1 (9)	3 (27)	1 (10)
KTS	10 (31)	4 (36)	2 (18)	4 (40)
Other	7 (22)	3 (27)	1 (9)	3 (30)
LM	8 (25)	3 (27)	4 (36)	1 (10)
VeM	2 (6)	0	1 (9)	1 (10)
Performance Status at Baseline, 50-70/ ≥80¹, n (%)	5 (16) / 27 (84)	2 (18) / 9 (82)	1 (9) / 10 (91)	2 (20) / 8 (80)
Pre-diabetic², n (%)	8 (25)	1 (9)	6 (55)	1 (10)
Local PIK3CA Status at Baseline, n (%)				
Kinase mutation	10 (31)	4 (36)	4 (36)	2 (20)
Non-Kinase mutation	16 (50)	4 (36)	6 (55)	6 (60)
No mutation documented	6 (19)	3 (27)	1 (9)	2 (20)
Prior disease-related systemic treatment, median	1	1	2	1
None, n (%)	9 (28)	3 (27)	3 (27)	3 (30)
Prior alpelisib / sirolimus, n (%)	23 (72)	8 (73)	8 (73)	7 (70)
Prior disease-related surgery, n (%)	19 (59)	5 (45)	6 (55)	8 (80)
Prior catheter-based procedures, n (%)	18 (56)	6 (55)	8 (73)	4 (40)

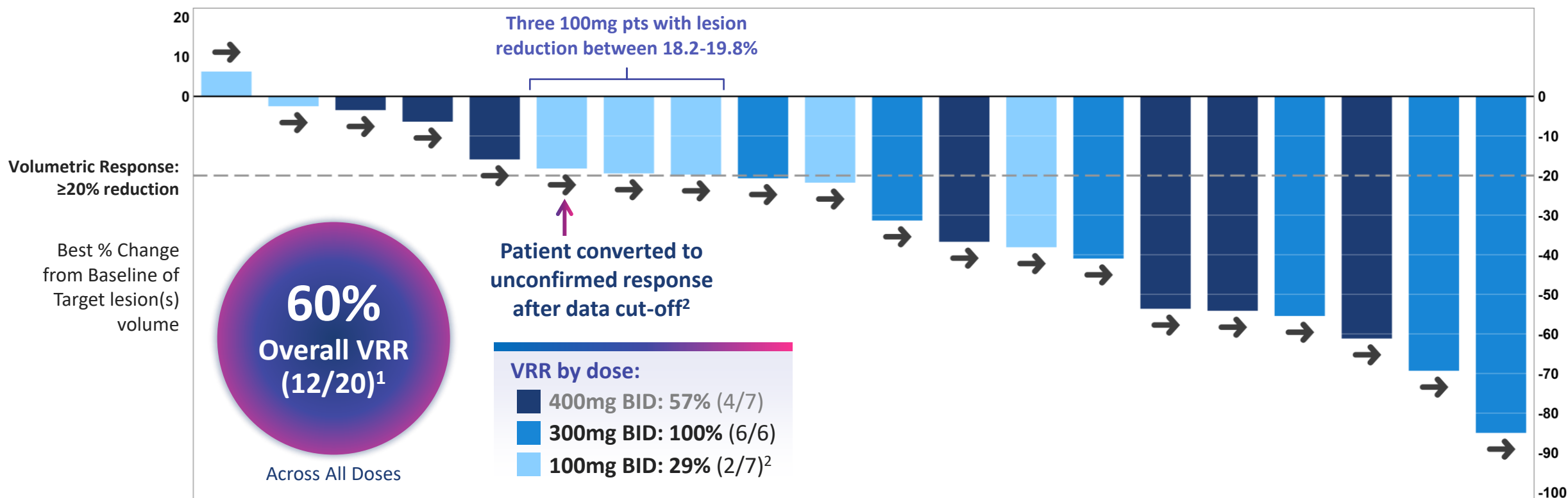
1. Lansky performance status for patients <16 years old or Karnofsky performance status for patients ≥16 years old; 2. Baseline HbA1c ≥5.7, glucose ≥100, or medical history of pre-diabetes mellitus

Zovegalisib – All Initial Doses Resulted in Exposures Projected to be Active

C1D15 Mean Concentration-Time Profiles By Dose



Zovegalisib – 60% Volumetric Response Rate by BICR Across All Doses



VRR by dose:

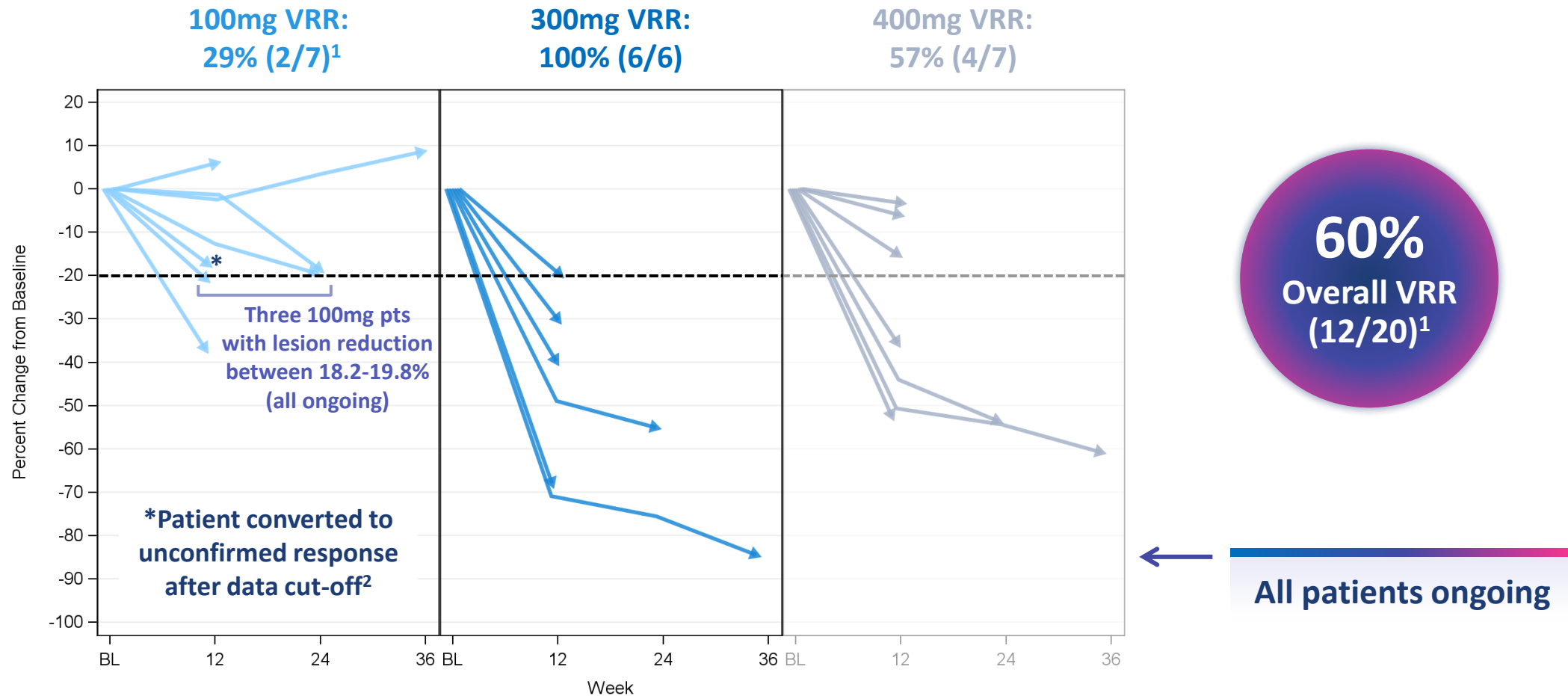
- 400mg BID: 57% (4/7)
- 300mg BID: 100% (6/6)
- 100mg BID: 29% (2/7)²

60%
Overall VRR
(12/20)¹
Across All Doses

Subtype	LM	PROS	VeM	PROS	PROS	PROS	PROS	PROS	PROS	PROS	LM	PROS	PROS	PROS	PROS	PROS	PROS	PROS	LM	LM
PROS subtype		KTS		FAO	KTS	FAO	FAVA	KTS	CLOVES	CLOVES		KTS	KTS	KTS	CLOVES	FAVA	CLOVES	KTS		
PIK3CA mutation	NK	NK	NK	NK	Unk	K	Unk	NK	NK	K	NK	Unk	K	NK	K	NK	NK	NK	K	Unk
Prior alp/siro	S	A + S	S	—	A + S	—	A + S	—	S	A	A + S	—	—	—	A + S	A + S	A + S	S	—	S
BL volume (L)	0.4	5.4	0.5	0.9	1.2	0.4	0.8	18.0	3.4	0.9	0.2	2.7	0.4	1.0	1.2	0.1	0.2	0.3	0.1	0.1
% Change from W12 Baseline	6.3	-2.5	-3.5	-6.4	-15.9	-18.2	-1.4	-12.7	-20.7	-21.8	-31.4	-36.7	-38.1	-41.0	-53.7	-44.1	-49.0	-50.7	-69.3	-70.9
W24		3.5					-19.5	-19.8								-54.2	-55.5	-54.4		-75.6
W36		8.9																-61.2		-85.0
BOR	SD	SD	SD	SD	SD	SD	SD	SD	uVR	uVR	uVR	uVR	uVR	uVR	uVR	cVR	cVR	cVR	uVR	cVR

1. Includes both confirmed and unconfirmed responses. 2. After the data cut-off date, one 100mg BID patient that did not have a volumetric response as of the data cut-off date has converted to an unconfirmed response, resulting in a 100mg BID volumetric response rate of 43% (3/7), a volumetric response rate of 69% (9/13) for patients treated at 300mg BID or 100mg BID, and a volumetric response rate of 65% (13/20) across doses. None of the other response-evaluable patients' response statuses have changed since the data cut-off date. Volumetric Response (VR) = 20% or greater reduction in target lesion volume by blinded independent central review (BICR); cVR = Confirmed Volumetric Response (VR with 2nd scan to confirm response), uVR = Unconfirmed Volumetric Response (VR without confirmatory scan), SD = Stable Disease

Zovegalisib – Volumetric Response Over Time (BICR)



Reductions generally deepened over time at all doses

1. Includes both confirmed and unconfirmed responses. 2. After the data cut-off date, one 100mg BID patient that did not have a volumetric response as of the data cut-off date has converted to an unconfirmed response, resulting in a 100mg BID volumetric response rate of 43% (3/7), a volumetric response rate of 69% (9/13) for patients treated at 300mg BID or 100mg BID, and a volumetric response rate of 65% (13/20) across doses. None of the other response-evaluable patients' response statuses have changed since the data cut-off date.

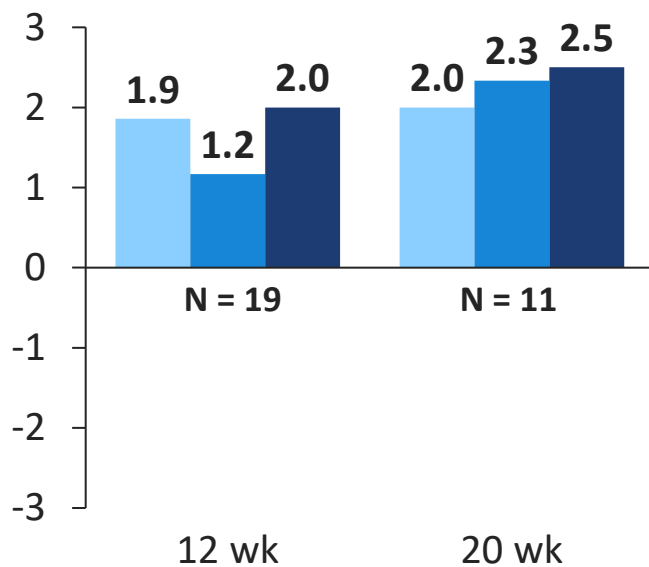
Zovegalisib – Initial Efficacy Data Supports Clear Symptomatic Benefit



Investigator-Reported (IGIC)

89%
of patients improved by week 12

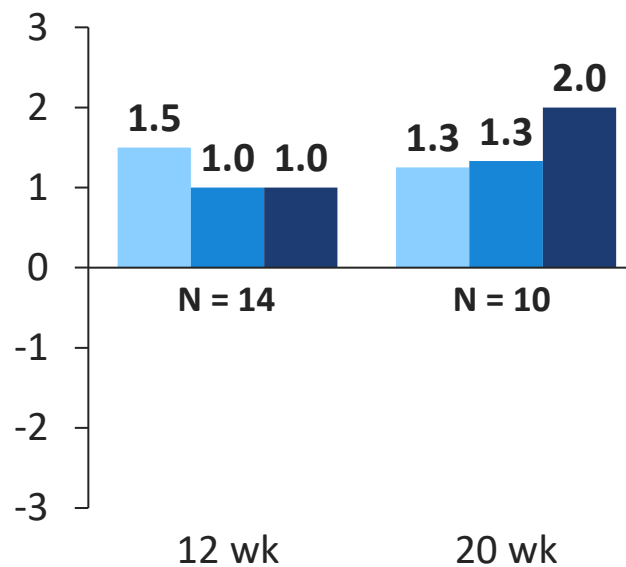
Mean Change by timepoint



Patient-Reported (PGIC)

79%
of patients improve by week 12

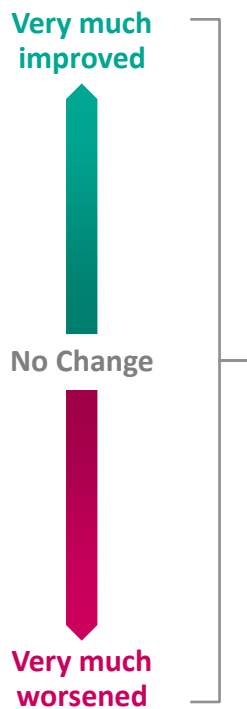
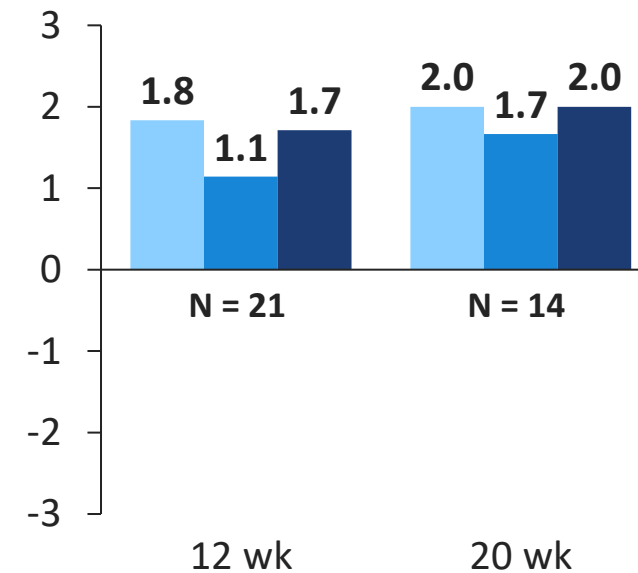
Mean Change by timepoint



Pain (IADRSS)

71%
of pain symptoms improved
by week 12

Mean Change by timepoint



100mg BID 300mg BID 400mg BID

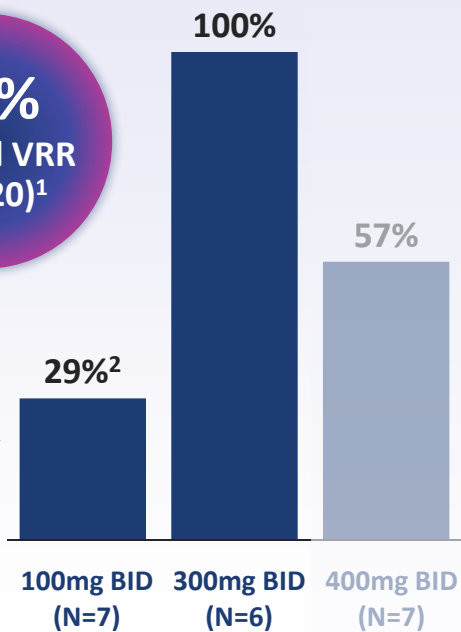
IGIC = Investigator Global Impression of Change, PGIC = Patient Global Impression of Change, IADRSS = Investigator Assessment of Disease-Related Signs and Symptoms.
Note: N for IGIC and PGIC is number of patients; N for IADRSS pain is number of most bothersome pain symptoms, where some patients may have more than one pain symptom.
Scale: +3 very much improved, +2 minimally improved, +1 minimally improved, 0 no change, -1 minimally worse, -2 much worse, -3 very much worse

Zovegalisib – Initial Efficacy Data Summary

Meaningful Efficacy Data

Volumetric Response Rate (VRR)

60%
Overall VRR
(12/20)¹

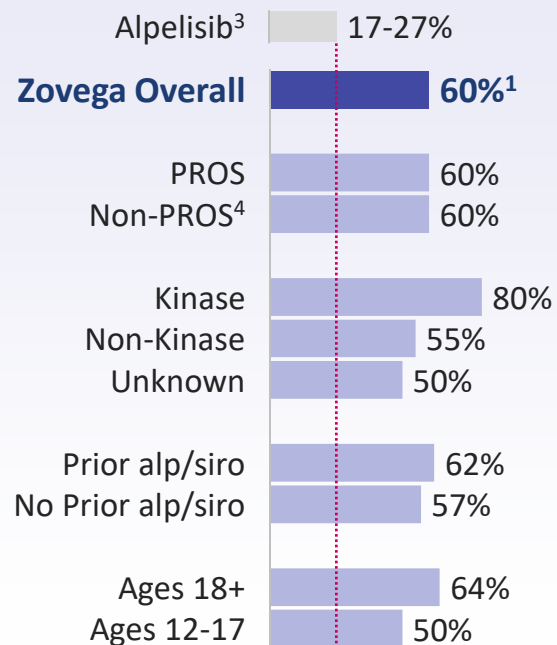


One 100mg pt converted from SD to uVR after data cut-off²

3 of 7 pts with lesion reduction 18.2-19.8%

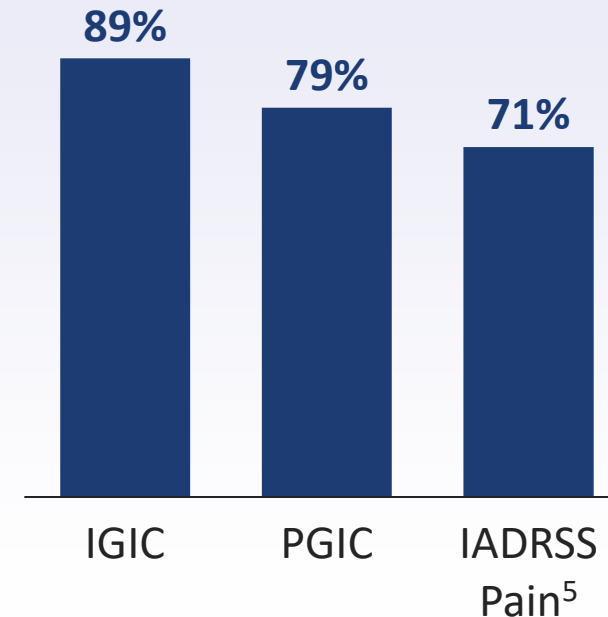
Consistent Across Subgroups

Subgroup VRR¹



Broad Symptomatic Benefit

% of Patients with Improvement at Week 12



1. Includes both confirmed and unconfirmed responses. 2. After the data cut-off date, one 100mg BID patient that did not have a volumetric response as of the data cut-off date has converted to an unconfirmed response, resulting in a 100mg BID volumetric response rate of 43% (3/7), a volumetric response rate of 69% (9/13) for patients treated at 300mg BID or 100mg BID, and a volumetric response rate of 65% (13/20) across doses. None of the other response-evaluable patients' response statuses have changed since the data cut-off date; 3. EPIK-P1: Vjoice FDA label and EPIK-P2: Canaud 2024 Blood 144:5512; 4. Non-PROS = LM and VeM; 5. IADRSS rate shown is percentage of pain symptoms improved; Alp = alpelisib, Siro = sirolimus; IGIC = Investigator Global Impression of Change, PGIC = Patient Global Impression of Change, IADRSS = Investigator Assessment of Disease-Related Signs and Symptoms, SD = Stable Disease, uVR = unconfirmed volumetric response. Note: These data are derived from different clinical trials at different points in time, with differences in molecule composition, trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.

Zovegalisib – Treatment-Related Adverse Events ≥15% of Patients

No discontinuations due to adverse events

		100mg BID (N=11)				300mg BID (N=11)				100mg+300mg BID (N=22)				400mg BID (N=10)			
		All Gr	Gr1	Gr2	Gr3+	All Gr	Gr1	Gr2	Gr3+	All Gr	Gr1	Gr2	Gr3+	All Gr	Gr1	Gr2	Gr3+
TRAE ≥15%	Any TRAE	82%	36%	45%	-	91%	55%	18%	18%	86%	45%	32%	9%	90%	20%	50%	20%
	Headache	18%	18%	-	-	73%	73%	-	-	45%	45%	-	-	50%	30%	20%	-
	Fatigue	18%	9%	9%	-	55%	36%	18%	-	36%	23%	14%	-	20%	10%	10%	-
	Nausea	27%	18%	9%	-	45%	36%	9%	-	36%	27%	9%	-	70%	40%	30%	-
	Diarrhea	27%	27%	-	-	18%	18%	-	-	23%	23%	-	-	10%	-	10%	-
	Hyperglycemia	-	-	-	-	45%	18%	27%	-	23%	9%	14%	-	40%	20%	10%	10%
Other select TRAE	Decreased appetite	18%	9%	9%	-	18%	9%	9%	-	18%	9%	9%	-	20%	10%	10%	-
	Rash	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
	Stomatitis	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

Majority of hyperglycemia observed in patients prediabetic at baseline¹

No Grade 3 hyperglycemia

Median Relative Dose Intensity	100%
Dose Reduction due to TRAE, n (%)	1 (9%) ²

Median Relative Dose Intensity	99%
Dose Reduction due to TRAE, n (%)	4 (36%)

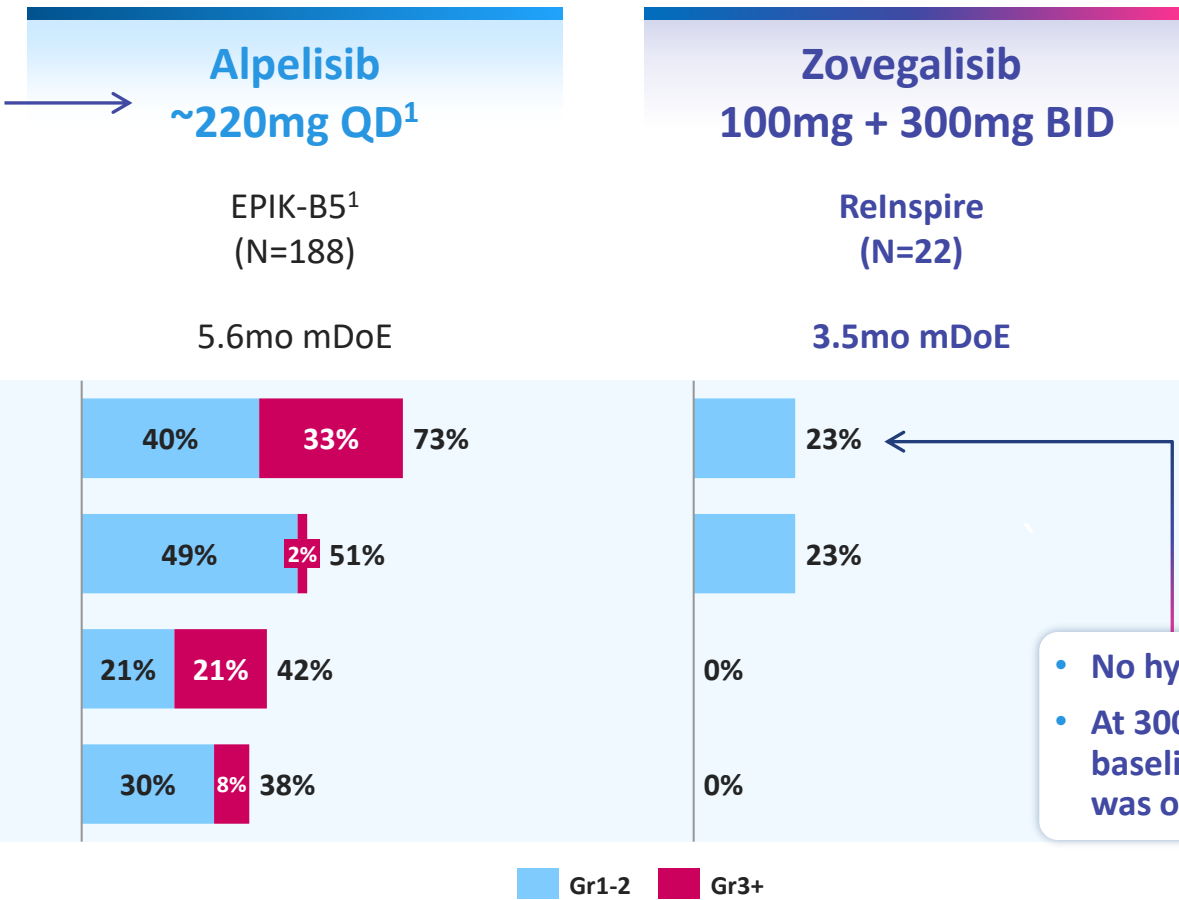
Median Relative Dose Intensity	99%
Dose Reduction due to TRAE, n (%)	5 (23%)

Median Relative Dose Intensity	77%
Dose Reduction due to TRAE, n (%)	7 (70%)

1. Baseline HbA1c ≥5.7, glucose ≥100, or medical history of pre-diabetes mellitus; 2. Patient later increased back up to original dose of 100mg BID

PI3K α Inhibitors – Tolerability Profile Across Known Key Pathway AEs

EPIK-P1 (250mg) was not a prospective study and EPIK-P2 was conducted at half the label dose⁵



Data benchmark

EPIK-B5¹
(N=188)

ReInspire
(N=22)

mDoE

5.6mo mDoE

3.5mo mDoE

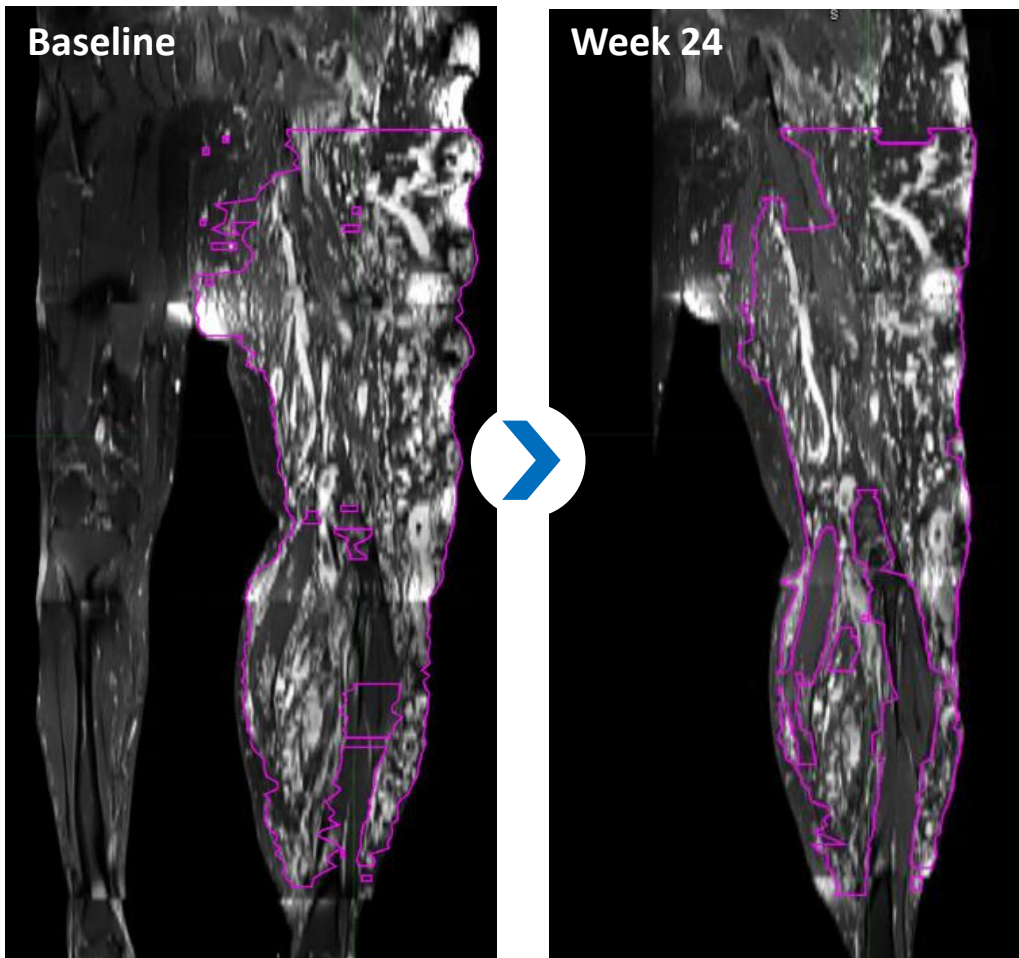
At 100mg and 300mg BID:

- No rash or stomatitis of any grade
- No Gr3 hyperglycemia or diarrhea
- No discontinuations
- Median dose intensity >99%

- No hyperglycemia at 100mg
- At 300mg, 55% of patients were pre-diabetic at baseline⁴, and the majority of hyperglycemia was observed in these patients

1. EPIK-B5, SABCS 2025 #RF7-02, 220mg dose approximated from dose modification data; 2. Rash for alpelisib references the cumulative sum of rates of rash and rash maculo-papular from the EPIK-B5 study, and may include overlap; 3. Stomatitis for alpelisib references the cumulative sum of rates of stomatitis and mucosal inflammation from the EPIK-B5 study, and may include overlap; 4. Pre-diabetic: baseline HbA1c ≥ 5.7 to < 6.5 , glucose ≥ 100 , or medical history of pre-diabetes mellitus; 5. EPIK-P1 is a retrospective study, label dose is 250mg QD, and EPIK-P2:

Patient Vignette – Adult Patient with PROS Achieved Meaningful Clinical and Radiographical Improvement with Zovega 100mg BID



18 liter lesion at baseline

3.6 liter reduction (-19.8%) at 24wk

44-year-old male with KTS (PROS)

- PIK3CA mutation: Q546K
- Prior sclerotherapy x3, no prior systemic tx
- Minimal mobility at baseline
- **Dosed with 100mg zovegalisib**

19.8% volumetric reduction at week 24
(13% reduction seen at week 12)

- 3.6 liter lesion reduction
- Deepening reduction at each scan

Dramatic and rapid clinical improvement

“Much Improved” IGIC overall status
Investigator Global Impression of Change

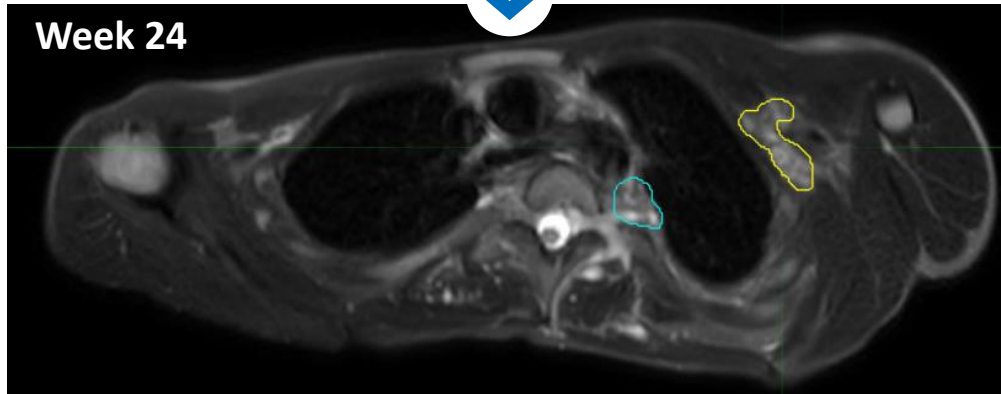
Note: Symptom improvement shown here is for Week 24

Tolerable profile allowing for prolonged dosing

- No dose modifications
- Patient remains on therapy at 100mg BID

“The participant used to not be able to walk further than front door to car. Within 2 weeks, he walked around the block.”
- ReInspire Investigator

Patient Vignette – Previously Systemically Treated Pediatric Patient with PROS Achieved Meaningful Radiographic & Clinical Improvement with Zovega 300mg BID



12-year-old male with CLOVES (PROS)

- PIK3CA mutation: E542K
- Prior surgery x6, laser therapy x12, sirolimus (no response) and alpelisib
- Painful chest lesion, with lymphatic leakage
- **Dosed with 300mg zovegalisib**

55% volumetric reduction at week 24

(49% reduction seen at week 12)

- Deepening reduction at each scan

Dramatic and rapid clinical improvement

“Much Improved” IGIC overall status

Investigator Global Impression of Change

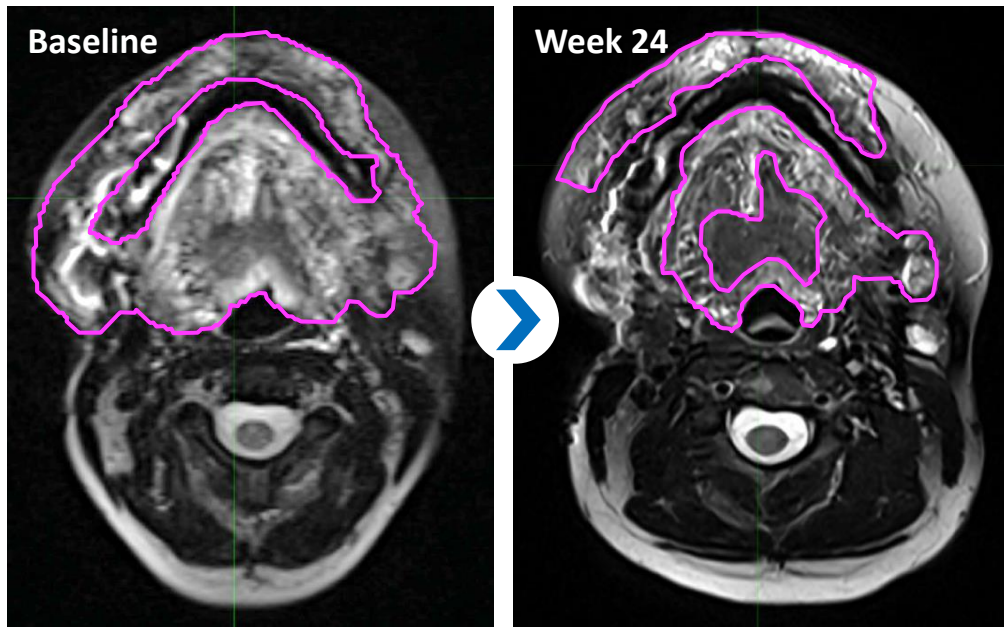
Note: Symptom improvement shown here is for Week 24

Tolerable profile allowing for prolonged dosing

- No dose modifications
- Patient remains on therapy at 300mg BID

*"After initiation of zovegalisib, the patient has experienced a marked reduction in overgrowth size, improved clothing fit, and decreased sensitivity of affected areas. Episodes of drainage and cellulitis have resolved, and previously painful stimuli such as ECG sticker placement are now well tolerated."
- ReInspire Investigator*

Patient Vignette – Adult patient with Facial LM Experienced Meaningful Improvement in Pain and Radiographic Response with Zovega 300 mg BID



↑
“She had a meaningful decrease in pain and fullness”
 - ReInspire Investigator

42-year-old female with facial LM

- PIK3CA mutation: E545K
- Prior sclerotherapy x3, embolization x2, surgery x4, sirolimus (no response, dc for AEs) & alpelisib (improvement, but dc for AEs)
- **Dosed with 300mg zovegalisib**

53% volumetric reduction at week 24*

(31% reduction seen at week 12)

- Deepening reduction at each scan

Dramatic and rapid clinical improvement

“Much Improved” IGIC overall status

Investigator Global Impression of Change

Most bothersome symptoms:

(Investigator Assessment of Disease-Related Signs and Symptoms)

Ear pain → Much improved

Oral pain → Minimally improved

Jaw pain → No change

Tolerable profile allowing for prolonged dosing

- No dose modifications
- Patient remains on therapy at 300mg BID

Note: Symptom improvement shown here is for Week 24

ReInspire preliminary data as of 04/15/2026

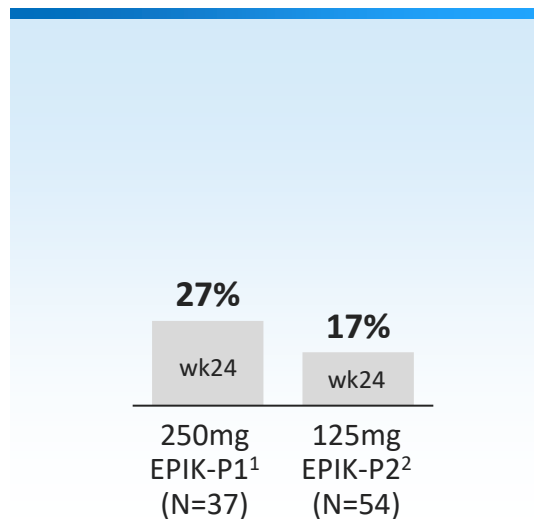
*Week 24 scan provided after data cutoff 20

Zovegalisib - Clear Path to Finding Dose to Evaluate Potential for Differentiated Profile

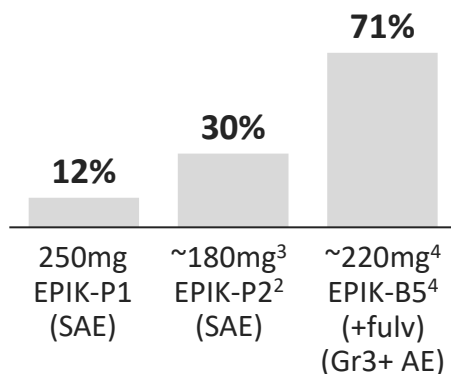


Alpelisib

Efficacy Data:
Volumetric response rate⁶

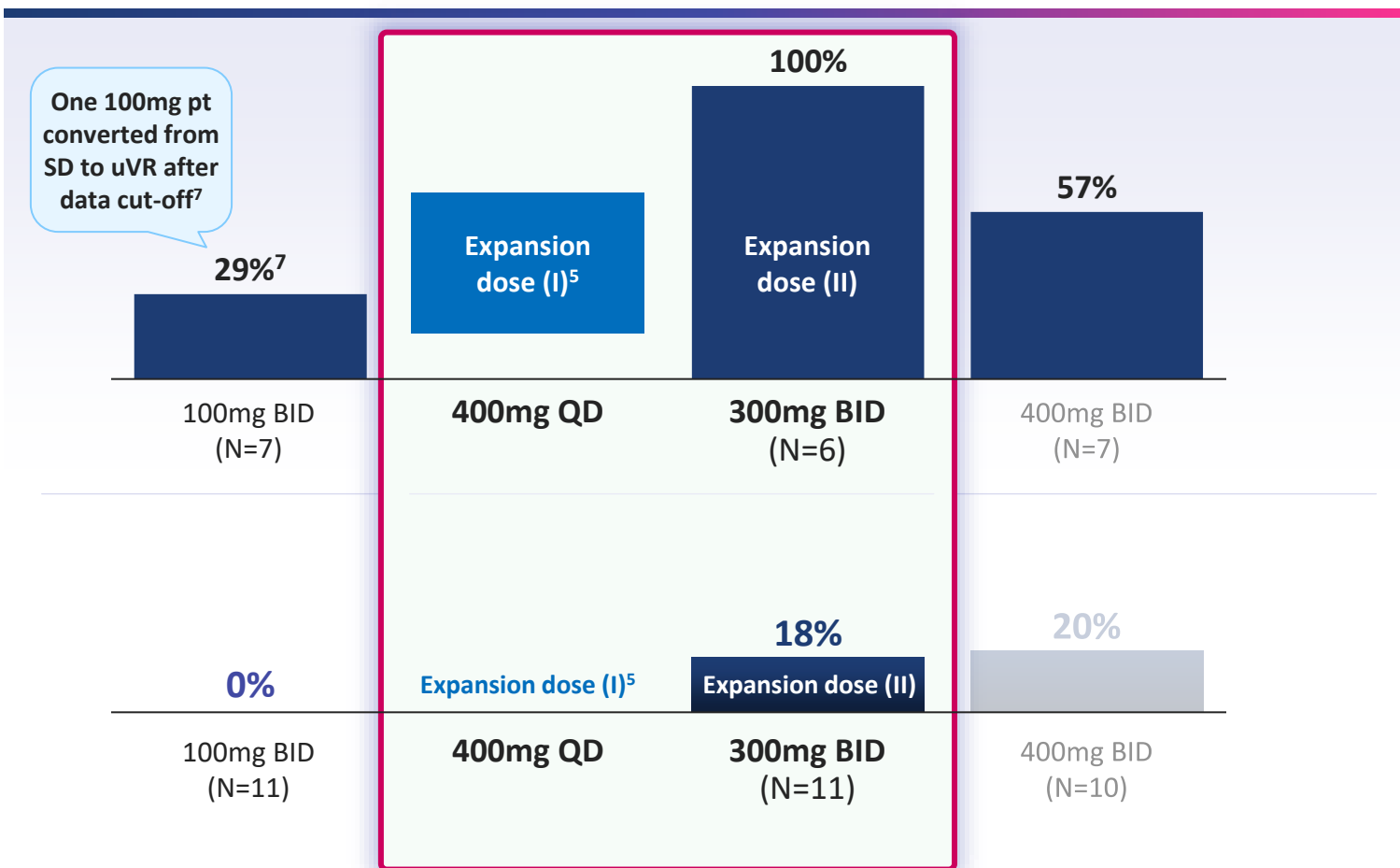


Safety Data:
Gr3+ TRAE



Zovegalisib

One 100mg pt converted from SD to uVR after data cut-off⁷

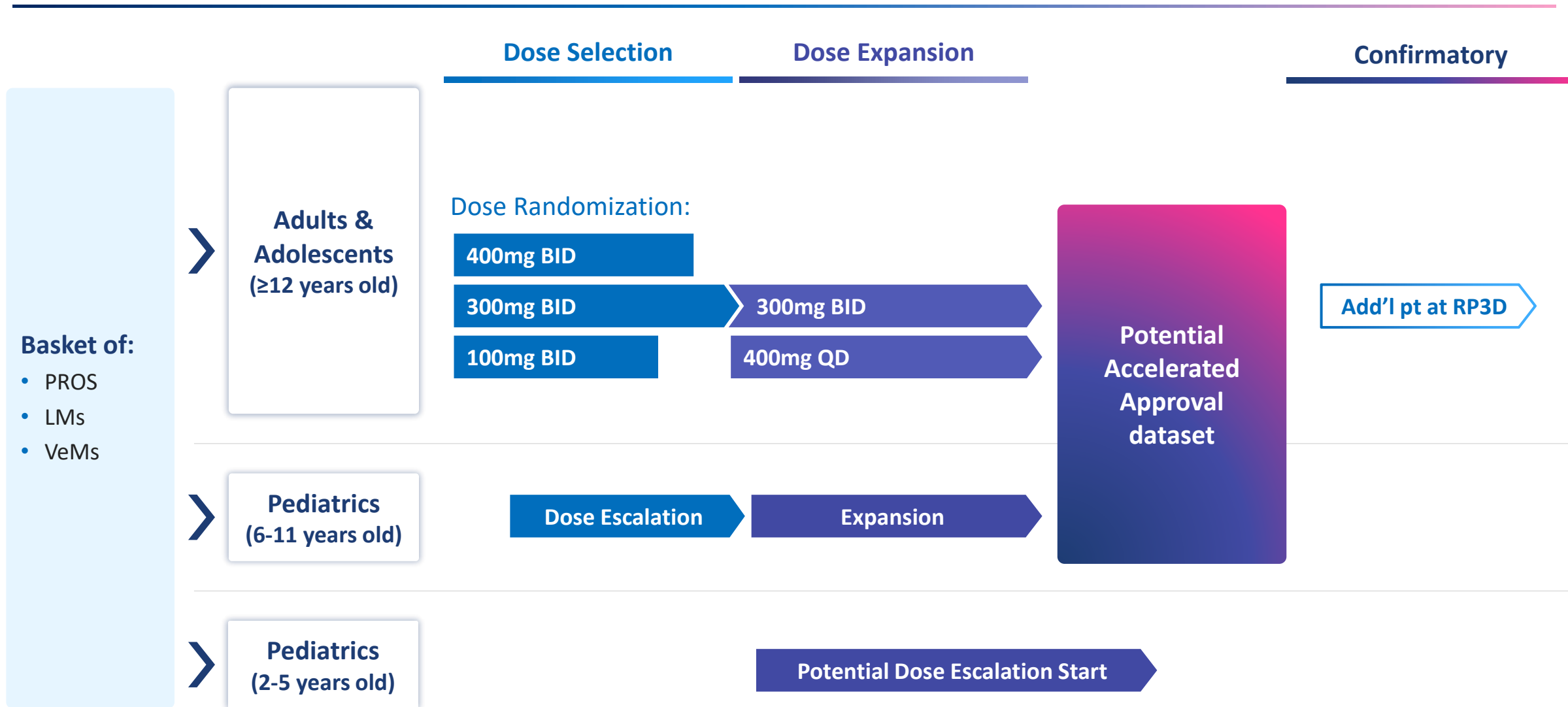


Selected as Expansion Doses

Reinspire preliminary data as of 04/15/2026

1. EPIK-P1: FDA review document; 2. EPIK-P2: Canaud 2024 Blood 144:5512 and results from clintrials.gov listing, 125mg QD was starting dose; 3. 180mg dose approximated from rates of dose escalation after week 26 listed on clintrials.gov listing; 4. EPIK-B5: SABCS 2025 #RF7-02, 220mg dose approximated from dose modification data; 5. 400mg QD expansion cohort yet to be initiated; 6. Reinspire VRR Includes both confirmed and unconfirmed responses. 7. After the data cut-off date, one 100mg BID patient that did not have a volumetric response as of the data cut-off date has converted to an unconfirmed response, resulting in a 100mg BID volumetric response rate of 43% (3/7), a volumetric response rate of 69% (9/13) for patients treated at 300mg BID or 100mg BID, and a volumetric response rate of 65% (13/20) across doses. None of the other response-evaluable patients' response statuses have changed since the data cut-off date. SD = Stable Disease, uVR = unconfirmed volumetric response. Note: These data are derived from different clinical trials at different points in time, with differences in molecule composition, trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.

Potential for Accelerated Approval Pathway*



*Illustrative and subject to discussion with the U.S. Food and Drug Administration

Vascular Anomalies – Mutant-Selective Approach Has Potential to Unlock Significant Market



Target Profile of Zovegalisib

Superior
Efficacy

Superior
Tolerability

Superior
Activity across VAs

Potential Commercial Implications

Greater
Penetration

Greater
Chronicity

Greater
Breadth of use

Significant
market opportunity

\$6-8B

Estimated US TAM¹

Relay Tx – Clear Path to Addressing Large Commercial Opportunities



Zovegalisib granted BTB

Zovegalisib Program
Anticipated 2026 disclosures
→ key value drivers

Clinical Benchmark Hurdle¹

Anticipated Next Steps

2L Breast Cancer
~\$2-3B TAM²

✓ **11.1mo mPFS at pivotal dose**
→ Rapid execution of ongoing 2L pivotal trial

Capi+fulv in 2L:
5.5mo mPFS

Phase 3 enrollment update by YE2026

1L Breast Cancer
~\$7-8B TAM

✓ **44% ORR in median 3L patients for Zovega + Atirmo + fulv triplet**
→ Aim to initiate 1L pivotal trial in early 2027

CDK+ET in 2L+
14-32% ORR

Regulatory update by YE2026, Phase 1/2 data in 1H 2027

Vascular Anomalies
~\$6-8B TAM

✓ **60% VRR across all doses⁴**
→ Enrolling adult expansion; pediatric cohort open

Alpelisib & KP-001
11-16% VRR³
at week 12 & 16

Data and regulatory update by YE2026

~\$642M | Cash as of end 1Q 2026

1. Clinical benchmark references: 2L breast cancer: capivasertib + fulvestrant (CDK4/6-experienced patient sub-population of CAPitello-291, Turner N Engl J Med 2023; 388:2058-2070) ; 1L breast cancer: CDK+ET in 2L+ (PACE Ph2: SABCS 2022 #GS3-06; postMONARCH Ph3: ASCO 2024 #1001; MAINTAIN Ph2: ASCO 2022 #LBA1004); atirmociclib Ph1: Pfizer R&D Oncology Day Feb 2024; vascular anomalies: alpelisib (EPIK-P2, Canaud 2024 Blood 144:5512) and KP-001 (Ozeki 2025, Orphanet Journal of Rare Diseases 20:64); 2. TAM calculated based on market benchmarks and internal analysis; 3. these benchmarks represent the earliest volumetric response evaluation timepoint; 4. Includes both confirmed and unconfirmed responses. After the data cut-off date, one 100mg BID patient that did not have a volumetric response as of the data cut-off date has converted to an unconfirmed response, resulting in a 100mg BID volumetric response rate of 43% (3/7), a volumetric response rate of 69% (9/13) for patients treated at 300mg BID or 100mg BID, and a volumetric response rate of 65% (13/20) across doses. None of the other response-evaluable patients' response statuses have changed since the data cut-off date. VRR = Volumetric Response Rate. Note: These data are derived from different clinical trials at different points in time, with differences in molecule composition, trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.



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THERAPEUTICS