



# PMV Pharmaceuticals

May 2026

# PMV Pharma is Harnessing the Power of p53 to Treat Cancer



PMV's lead candidate is rezatapopt, a first-in-class, investigational p53 Y220C reactivator

The p53 Y220C mutation, a previously undruggable target, is found in 3.1% of ovarian cancer and 1% of all solid tumors

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Phase 1 PYNNACLE study has achieved proof of concept data for rezatapopt and was recently published in The New England Journal of Medicine

Pivotal Phase 2 PYNNACLE study interim clinical data demonstrates favorable efficacy and safety across multiple tumor types

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NDA submission planned in 1Q2027 in platinum-resistant/refractory ovarian cancer patients

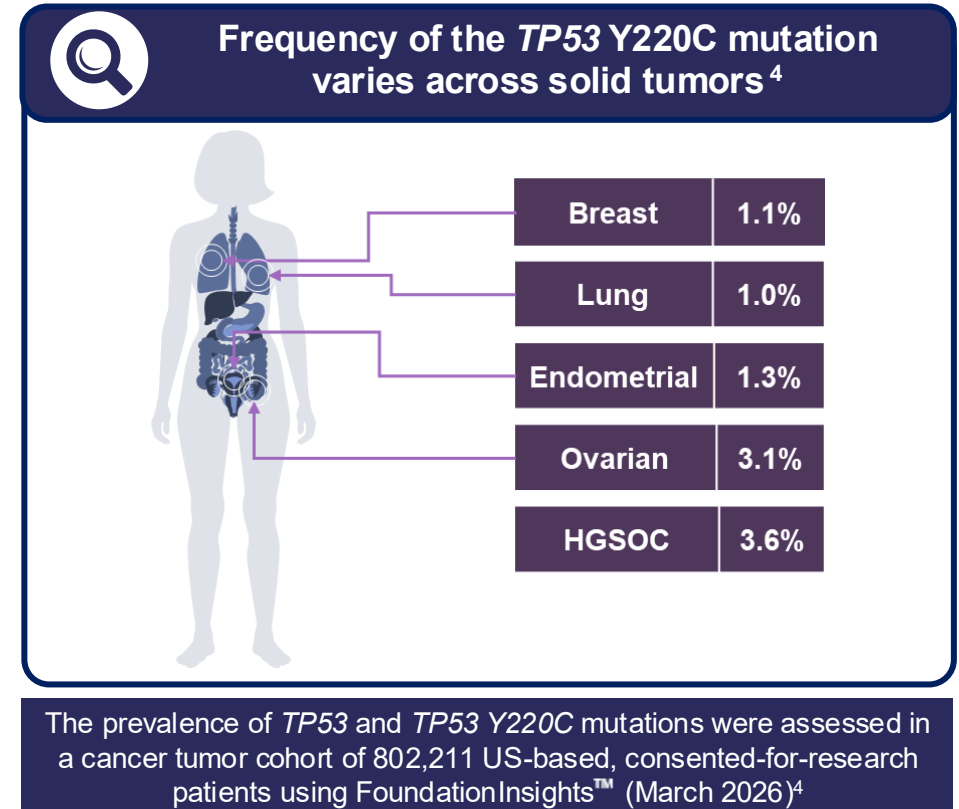
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Strong balance sheet with \$93.5M as of March 31, 2026, with cash runway through 2Q2027

# Rezatapopt Targets TP53 Y220C Hotspot Mutation Detected Across Solid Tumors

- *TP53* mutations are the most common genomic alterations across all human cancers<sup>1</sup>
- Most *TP53* mutations occur in the central DNA-binding domain and ten of them are referred to as 'hot-spot' mutations, accounting for ~30% of the *TP53* mutations observed in human cancer<sup>1-2</sup>
- p53 Y220C is a key hot-spot *TP53* missense mutation that destabilizes p53<sup>1,3</sup>
- **p53 Y220C present in ~1% of solid tumors<sup>4</sup>**
- **Addressable 2L+ U.S. & EU4/UK patients ~12K<sup>4,5</sup>**

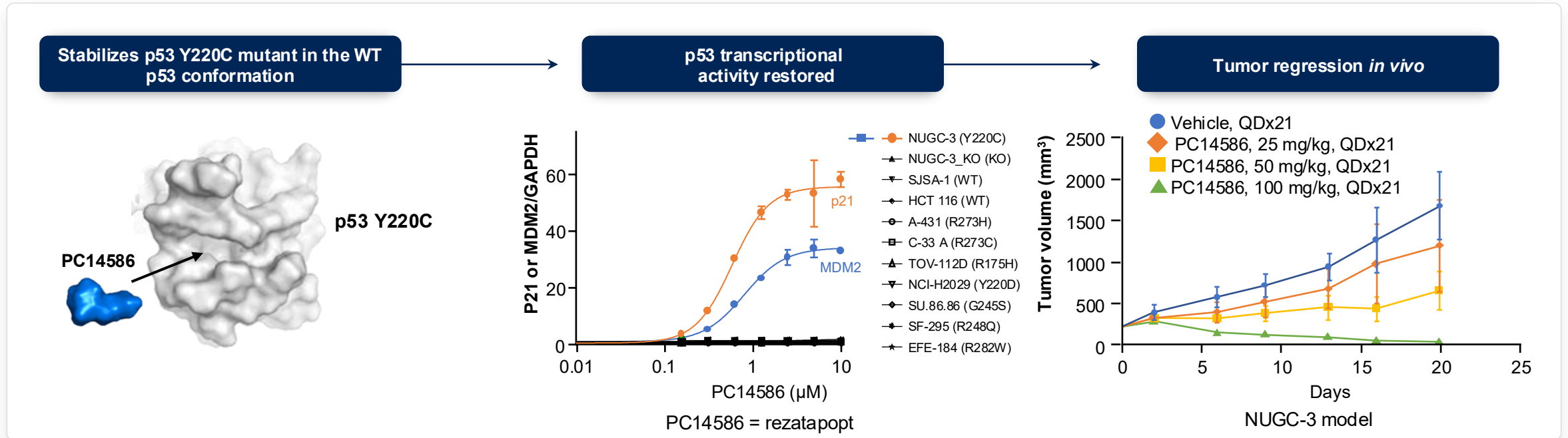


Deoxyribonucleic acid. <sup>1</sup> Baugh EH, et al. *Cell Death Differ.* 2018;25:154–160. <sup>2</sup> Roszkowska KA, et al. *Int J Mol Sci.* 2020;21:1334. <sup>3</sup> Bouaoun L, et al. *Hum Mutat.* 2016;37:865–876.

<sup>4</sup> FoundationInsights™. A proprietary database used under license with review and approval from Foundation Medicine®. Available at: <https://www.foundationmedicine.com/service/genomic-data-solutions>. Accessed March 2026, <sup>5</sup> DRG Epidemiology Estimates 2028.

# Rezatapopt is a p53 Y220C-Selective First-in-Class p53 Reactivator

- Orally available small molecule designed to selectively bind to the pocket contained in the p53 Y220C mutant protein<sup>1</sup>
- Stabilizes the p53 Y220C mutant protein in the wild-type p53 conformation, thereby restoring transcription and tumor-suppressor function<sup>1</sup>
- Inhibits proliferation across all Y220C-expressing cell lines



# Compelling Rezatapopt Monotherapy Phase 2 Interim Data

## Across All Cohorts:

- Encouraging efficacy in heavily pre-treated patients with a *TP53* Y220C mutation with poor prognoses<sup>1</sup>
- Promising rate of tumor responses observed across multiple tumor types
  - ORR: 34%
  - Median duration of response: 7.6 months
- Differentiated safety and tolerability profile compared to standard of care

## Ovarian Cancer:

- Significant unmet medical need
- Strong response rate and benefit
  - ORR: 44%
  - Median duration of response: 8.2 months
- Initial registrational opportunity in platinum-resistant or refractory ovarian cancer (PROC) informed by FDA feedback

- *TP53* Y220C mutation leads to a worse prognosis<sup>1</sup>
- Emerging clinical data supports rezatapopt as an effective, well-tolerated, oral option
- Opportunity to deliver a novel, biomarker-selected chemo-alternative

# Overview of PYNNACLE Phase 2 Interim Data



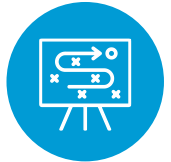
Overall results across all cohorts

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Updated Ovarian cohort results

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NDA submission strategy

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Looking ahead

# PYNNACLE Phase 2 Study Design

Ongoing Phase 2 study actively enrolling patients across ~70 sites globally

## Cohorts

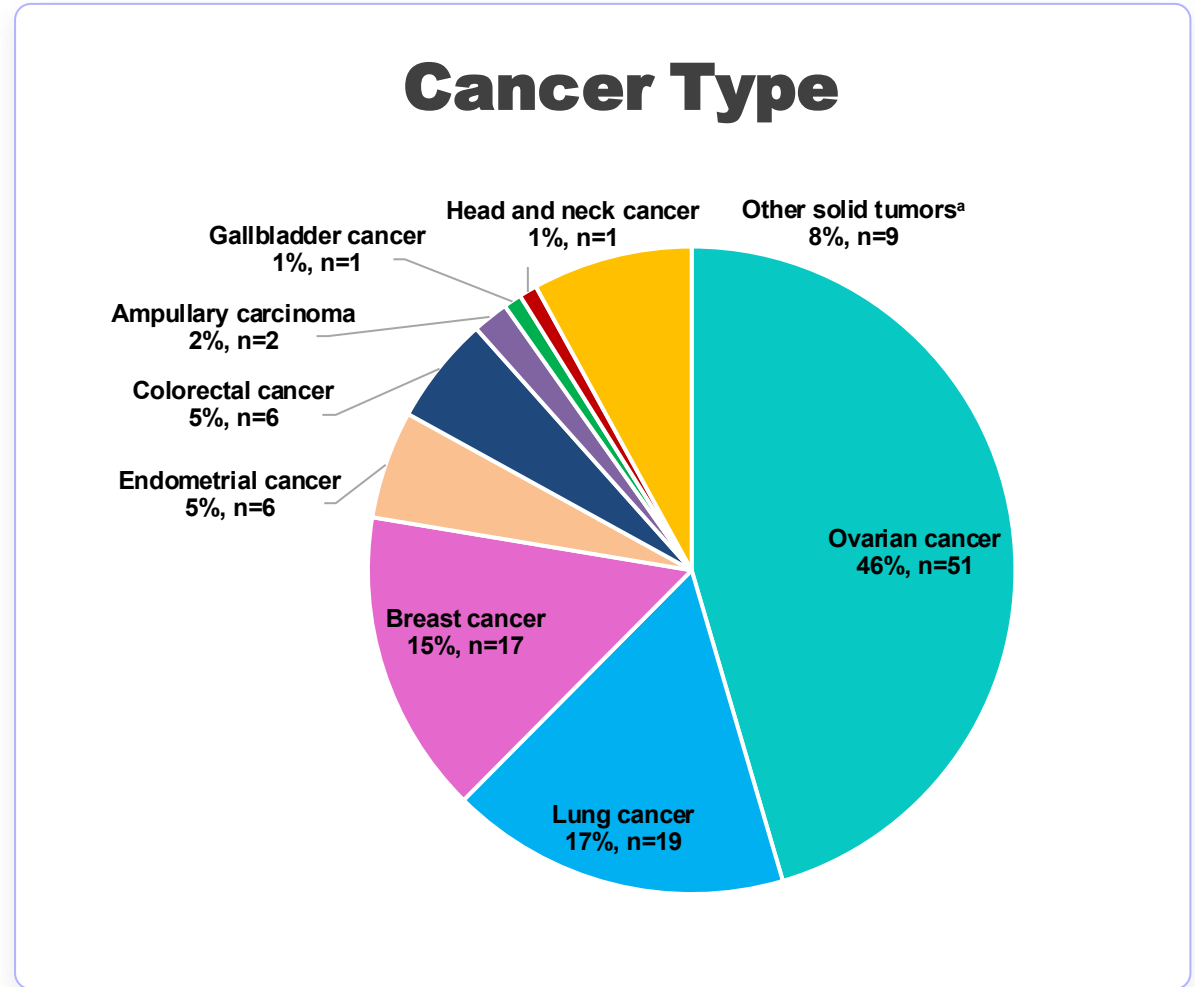
<p><b>Patient Population</b></p> <ul style="list-style-type: none"> <li>• Aged ≥ 12 years</li> <li>• Locally advanced or metastatic solid tumors, excluding primary CNS tumors</li> <li>• Documented <i>TP53</i> Y220C and <i>KRAS</i> WT only</li> <li>• Prior standard therapy or ineligible for appropriate standard of care therapy</li> </ul>	<p><b>Basket</b> <b>N = ~200</b></p> <p>Rezatapopt at 2000mg QD</p>	<p>Cohort 1: Ovarian cancer</p> <p>Cohort 2: Lung cancer</p> <p>Cohort 3: Breast cancer</p> <p>Cohort 4: Endometrial cancer</p> <p>Cohort 5: All other solid tumors</p>	<p><b>Primary endpoint:</b> ORR per BICR</p> <ul style="list-style-type: none"> <li>- Across all cohorts</li> <li>- Ovarian cancer cohort</li> </ul>
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Accelerating development in key tumor types via a streamlined single-arm pivotal study design

# Demographics and Baseline Characteristics (All Cohorts)

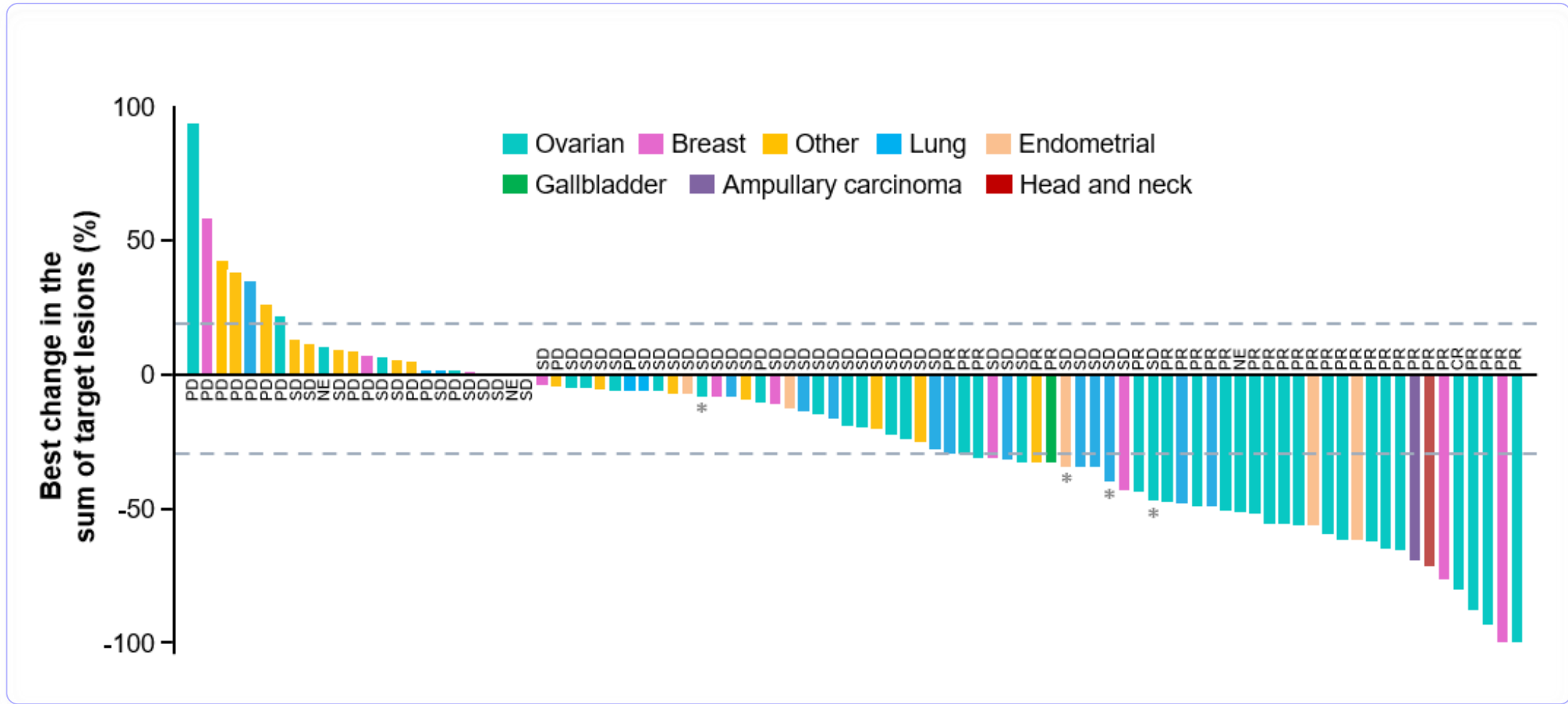
*Heavily pre-treated patients across broad spectrum of tumor types*

	Total N=112
Age (years)	Median 65
Sex	Female 73%, Male 27%
ECOG PS	0: 44%, 1: 56%
Prior line of systemic therapy	Median of 3 prior lines (range 1-10) 3 or more prior lines 64%
TP53 Y220C mutation status	100%
KRAS status	Wild type 100%



Data Cutoff 04Sep2025

# Target Lesion Reduction Observed in the Majority of Patients



Data Cutoff 04 Sep 2025

# Responses Observed Across All Cohorts in Eight Tumor Types

## TP53 Y220C / KRAS WT Efficacy Population <sup>a</sup> (n=103)

Across All Cohorts	ORR n (%)
ORR per Investigator assessment	35 (34%)
Confirmed Complete Response (CR)	1
Confirmed Partial Response (PR)	29
Unconfirmed Partial Response (uPR)	5

By Cohort	ORR n (%)
Ovarian	22/48 (46%)
Breast	2/12 (17%)
Lung	4/19 (21%)
Endometrial	3/5 (60%)
Other Solid Tumors	4/19 (21%)

**Post-data cutoff:**

- **Across all cohorts:** All 5 uPR patients converted to confirmed PRs, including 1 lung cancer and 4 ovarian cancer patients.

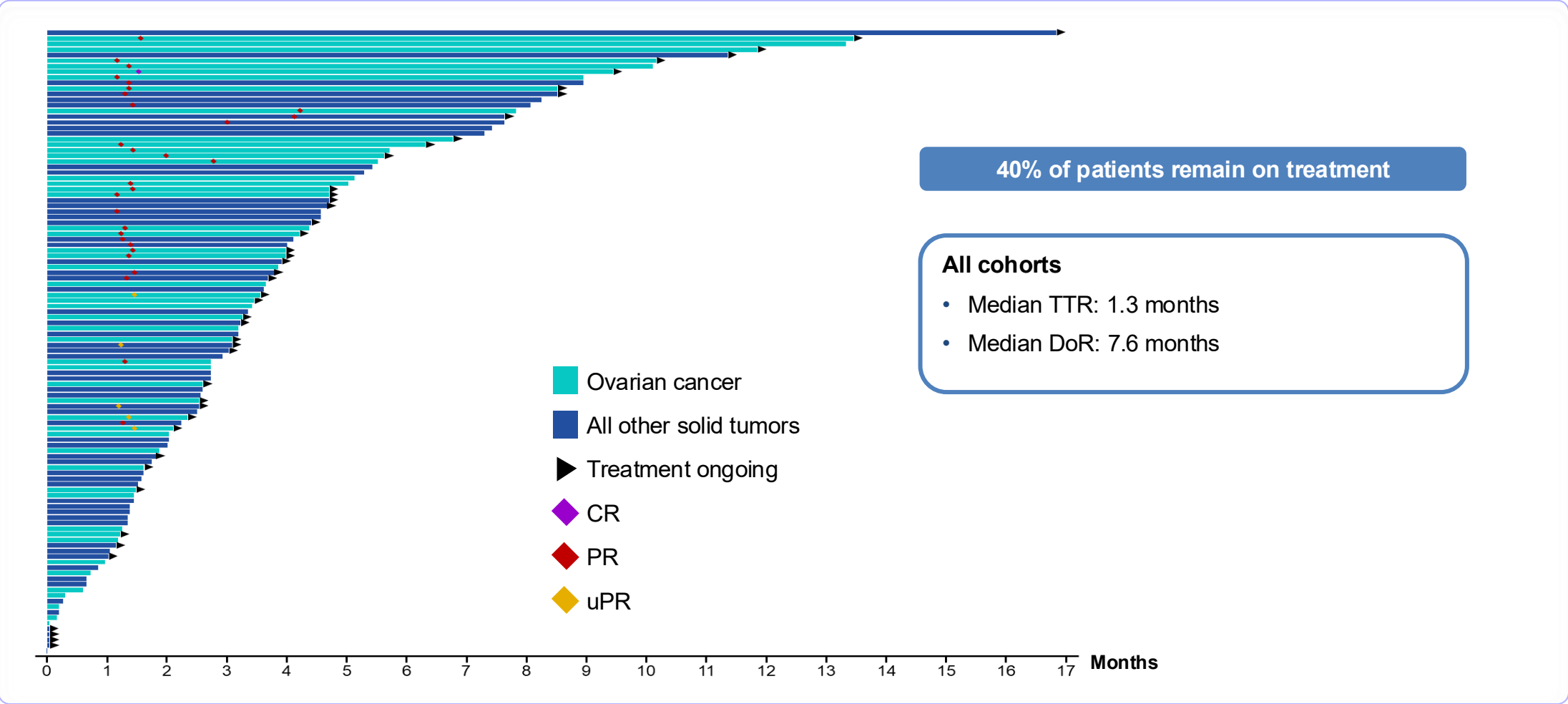
Data Cutoff 04Sep2025

*Overall*  
**34% Overall ORR**  
**7.6 months median Duration of Response <sup>b</sup>**

<sup>a</sup> Patients with the opportunity to reach first post-baseline scan. Patients discontinuing before the first post-baseline scan are included in the efficacy population.

<sup>b</sup> Duration of Response accounts only for confirmed responses.

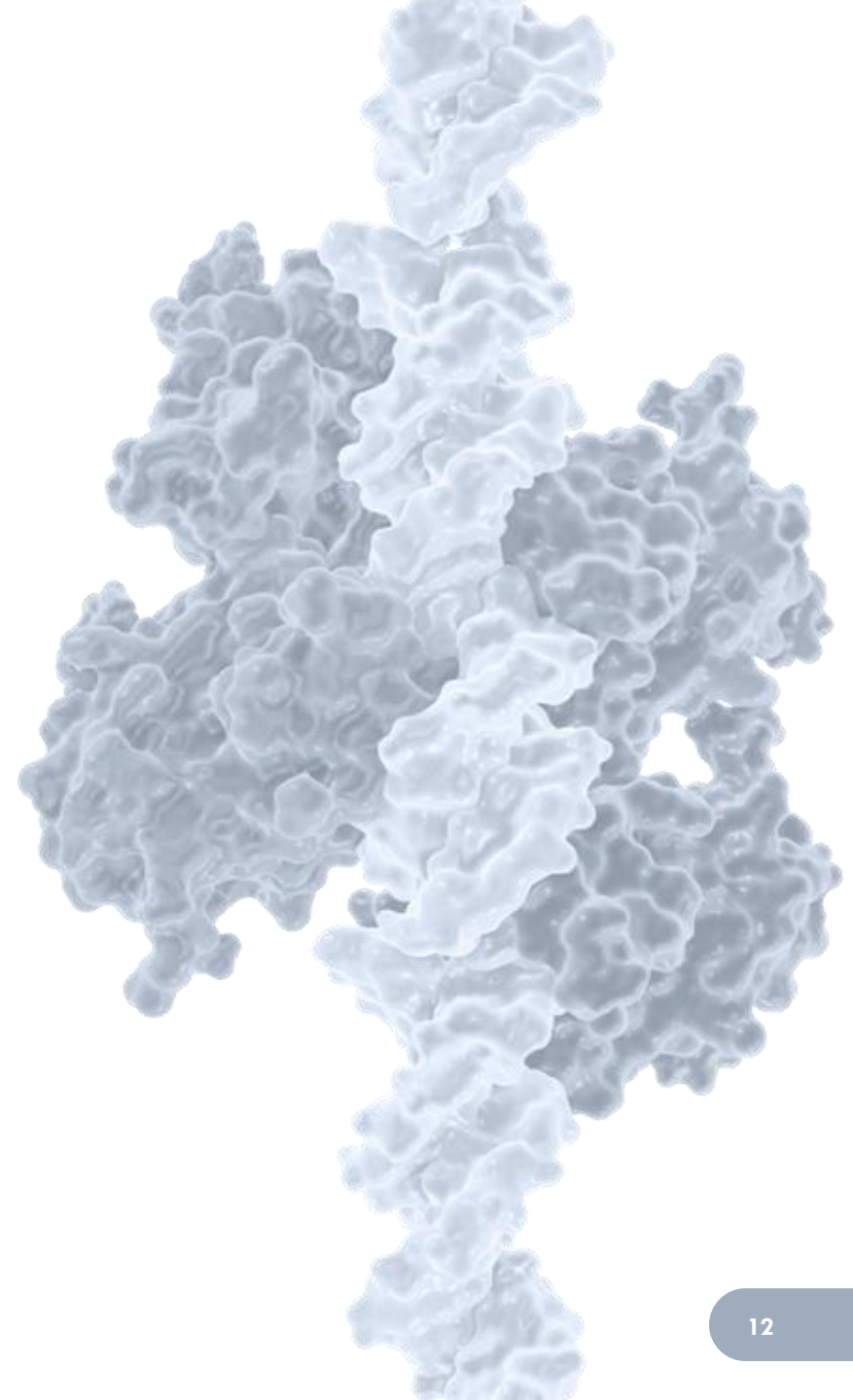
# Rapid Time to Response and Long Duration of Treatment



Data Cutoff 04Sep2025

# Updated Ovarian Cancer Data

29 March 2026

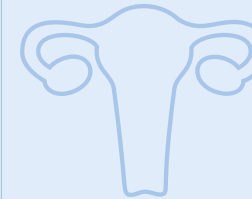


# Demographics and Baseline Characteristics

*Heavily pre-treated population with poor prognostic features*

Ovarian cancer cohort, n=76	
<b>Median age, years</b> (min–max)	66 (46–91)
<b>ECOG performance status, n (%)</b> 0 / 1	36 (47) / 40 (53)
<b>Prior lines of systemic therapy, n (%)</b> 1 / 2 3 ≥4 Median (min–max)	3 (4) / 14 (18) 16 (21) 43 (57) 4 (1–10)
<b>Prior therapies, n (%)</b> Bevacizumab	61 (80)
<b>Platinum status, n (%)</b> Sensitive Resistant Refractory Primary platinum refractory	3 (4) 46 (61) 27 (36) 10 (13)
<b>Histology, n (%)</b> High grade serous	74 (97)
<b>FRα status, n (%)</b> Positive / Negative	n=63 30 (48) / 33 (52)
<b>BRCA mutation, n (%)</b> Somatic BRCA 1/2 mutation Germline BRCA 1/2 mutation	7 (9) 3 (4) 4 (5)
<b>HRD status, n (%)</b> Positive / Negative	n=67 27 (40) / 40 (60)

## Heavily pre-treated patients:



- **96%** platinum-resistant or refractory
- **80%** received prior bevacizumab
- **78%** with three or more prior lines of therapy

# Rezatapopt Demonstrates Consistent ~42-53% ORR Across Key Ovarian Subgroups

	Ovarian cancer cohort n=72 <sup>a</sup>	Platinum resistant n=44	Platinum refractory <sup>b</sup> n=25	Prior bevacizumab n=57	Prior PARPi n=38	FR $\alpha$ positive <sup>c</sup> n=30	FR $\alpha$ negative <sup>d</sup> n=31
<b>ORR<sup>e</sup> %</b>	<b>44</b>	<b>45</b>	<b>44<sup>f</sup></b>	<b>44</b>	<b>53</b>	<b>50</b>	<b>42</b>
<b>CR</b>	1 (1.4)	0 (0.0)	0 (0.0)	1 (1.8)	1 (2.6)	0 (0.0)	0 (0.0)
<b>PR</b>	31 (43.1) <sup>g</sup>	20 (45.5)	11 (44.0)	24 (42.1)	19 (50.0)	15 (50.0)	13 (41.9)

Data Cutoff 29Mar2026

**Post-data cutoff: Ovarian cancer population:** ORR reached 46% (34/74) following 2 additional patients experiencing unconfirmed PRs

- **44% ORR (95% CI: 33%-57%) in ovarian cancer cohort exceeding ORR for single agent non-platinum based chemotherapy**
- **Compelling efficacy observed regardless of prior treatment received in a heavily pretreated population**

<sup>a</sup> Patients with the opportunity to reach first post-baseline scan. Patients discontinuing before the first post-baseline scan are included in the efficacy-evaluable population.

<sup>b</sup> Platinum refractory: defined as relapse during platinum therapy or within 1 month of the last dose of a platinum agent.

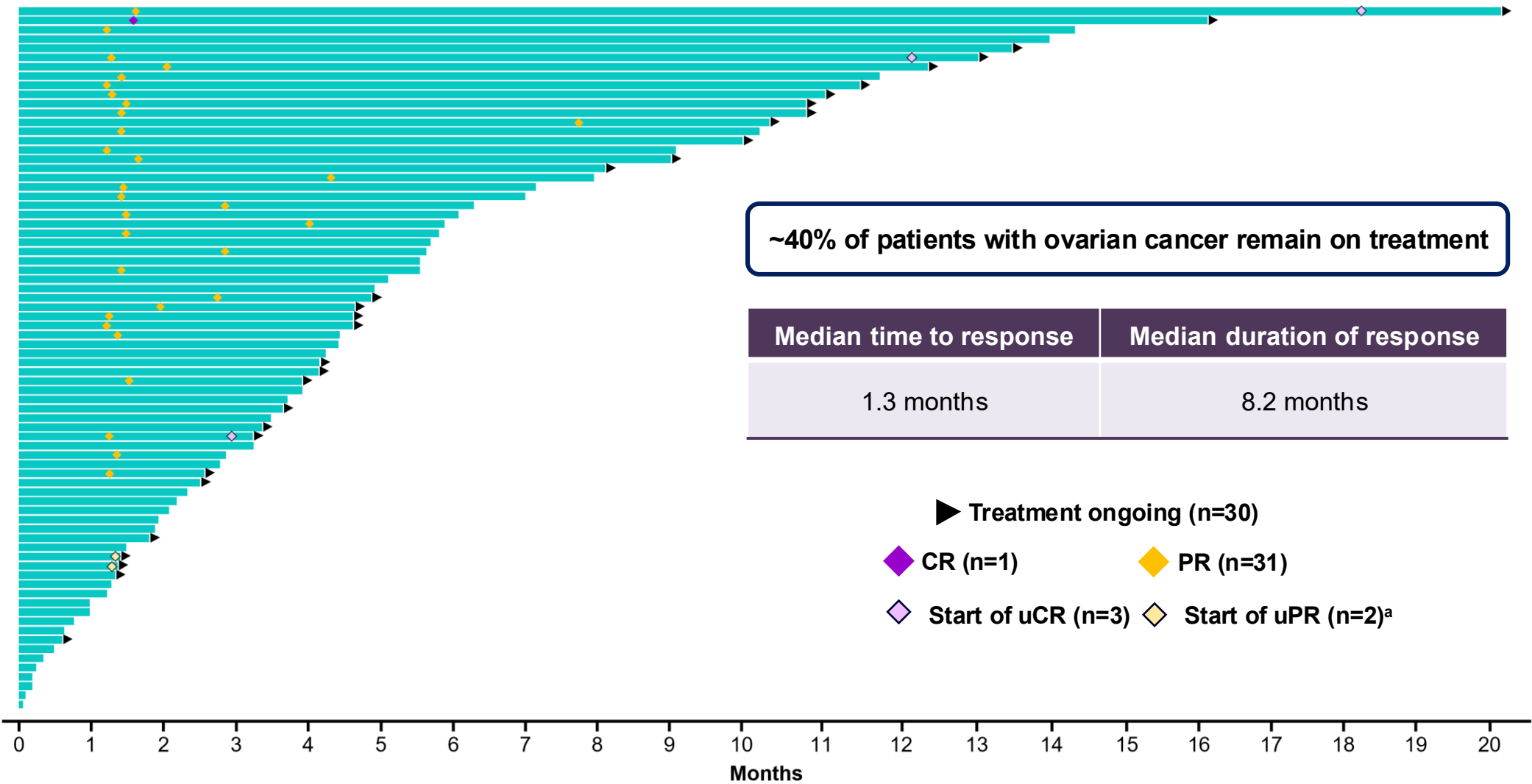
<sup>c,d</sup> FR $\alpha$  positive: FR $\alpha$  expression in  $\geq 75\%$  of viable tumor cells and FR $\alpha$  negative: FR $\alpha$  expression in  $< 75\%$  of viable tumor cells.

<sup>e</sup> ORR is calculated from confirmed CR and confirmed PR.

<sup>f</sup> In the primary platinum-refractory population (n=10), 3 patients experienced a confirmed PR as of March 29, 2026.

<sup>g</sup> 3 patients who achieved a confirmed PR as of the data cutoff experienced unconfirmed CR at last scan.

# Rapid and Durable Responses Were Observed in the Ovarian Cohort



Data cutoff: March 29, 2026. Overall duration of treatment measured in 76 patients. <sup>a</sup> Two unconfirmed PRs occurred after the data cutoff. CR, complete response; PR, partial response; uCR, unconfirmed complete response; uPR, unconfirmed partial response.

# Rezatapopt Displays Favorable Safety and Tolerability in the Overall Population

Preferred term	Maximum CTCAE grade <sup>a</sup>				
	Any grade	1	2	3	4
Nausea	59 (41.8)	41 (29.1)	18 (12.8)	0 (0.0)	0 (0.0)
Fatigue	36 (25.5)	15 (10.6)	20 (14.2)	1 (0.7)	0 (0.0)
Blood creatinine increased	31 (22.0)	10 (7.1)	19 (13.5)	2 (1.4)	0 (0.0)
ALT increased	26 (18.4)	11 (7.8)	5 (3.5)	8 (5.7)	2 (1.4)
Anemia	26 (18.4)	7 (5.0)	11 (7.8)	8 (5.7)	0 (0.0)
AST increased	26 (18.4)	12 (8.5)	5 (3.5)	8 (5.7)	1 (0.7)
Vomiting	24 (17.0)	16 (11.3)	8 (5.7)	0 (0.0)	0 (0.0)
Decreased appetite	18 (12.8)	12 (8.5)	6 (4.3)	0 (0.0)	0 (0.0)
Diarrhea	16 (11.3)	14 (9.9)	1 (0.7)	1 (0.7)	0 (0.0)
Pruritus	15 (10.6)	10 (7.1)	5 (3.5)	0 (0.0)	0 (0.0)
Asthenia	13 (9.2)	6 (4.3)	7 (5.0)	0 (0.0)	0 (0.0)
Constipation	10 (7.1)	9 (6.4)	1 (0.7)	0 (0.0)	0 (0.0)
Platelet count decreased	10 (7.1)	4 (2.8)	2 (1.4)	2 (1.4)	2 (1.4)
Headache	8 (5.7)	5 (3.5)	2 (1.4)	1 (0.7)	0 (0.0)
Rash maculopapular	8 (5.7)	1 (0.7)	3 (2.1)	4 (2.8)	0 (0.0)

<sup>a</sup> No Grade 5 treatment-related adverse events were observed.

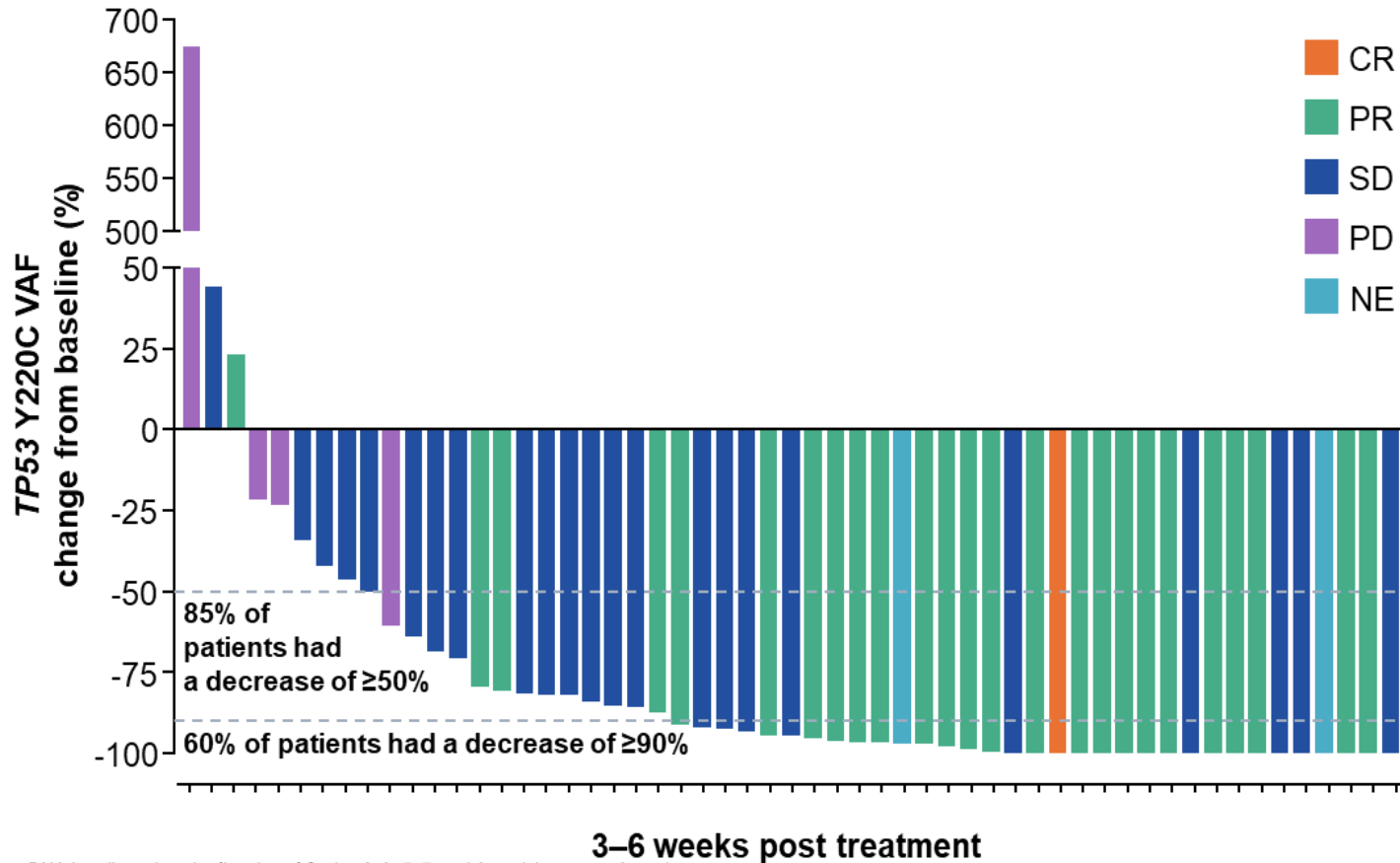
ALT, alanine aminotransferase; AST, aspartate aminotransferase; CTCAE, Common Terminology Criteria for Adverse Events; TRAE, treatment-related adverse event.

Data cutoff: March 29, 2026

- TRAEs were mostly Grade 1 or 2
- Most frequent TRAEs: Nausea, fatigue, and blood creatinine increased
- Laboratory abnormalities were manageable / monitorable
  - Most cases were transient and reversible
- A similar safety profile was observed in patients with ovarian cancer
- 7/141 patients (5.0%) discontinued treatment due to TRAEs, including 4/76 patients (5.3%) with ovarian cancer

# Rezatapopt On-target Activity Supported by Decreases in ctDNA TP53 Y220C VAF in Ovarian Cancer Patients

Reduction in *TP53* Y220C VAF was seen in most patients with ovarian cancer who had available data (n=55)



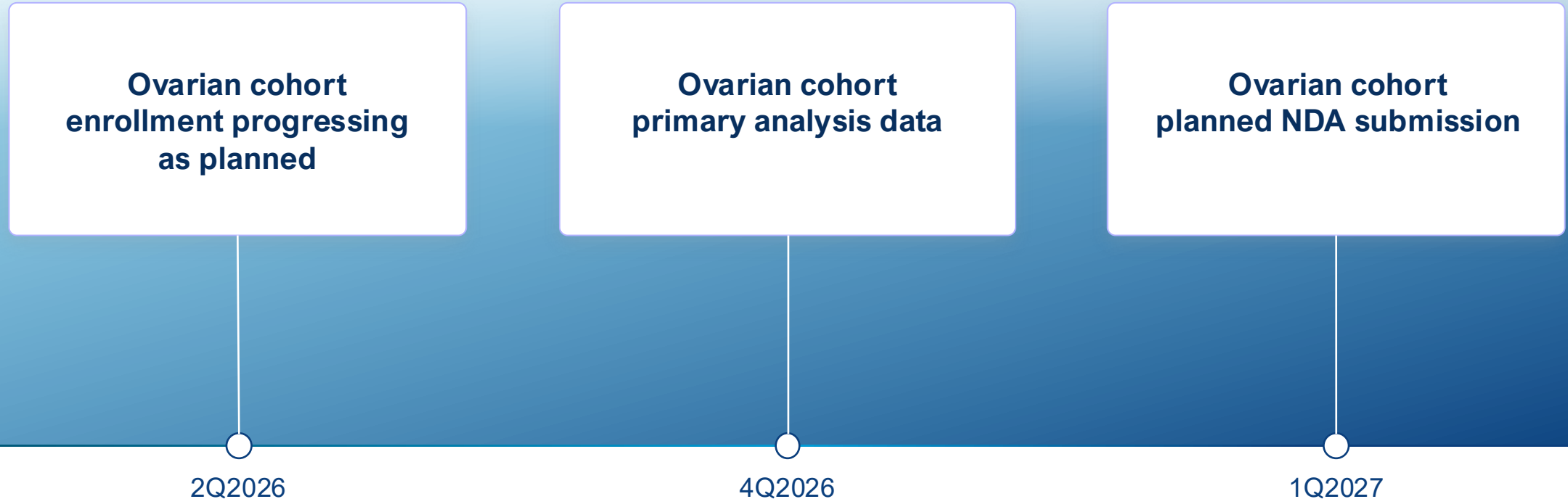
- 55 patients had ctDNA *TP53* Y220C VAF data available at baseline and on treatment (3–6 weeks)
- Nearly all patients experiencing a response had a reduction in *TP53* Y220C VAF
- 52 patients (95%) had a reduction in *TP53* Y220C VAF
  - 85% had a reduction of  $\geq 50\%$

ctDNA is collected on the first day of Cycles 2, 3, 5, 7, and 9, and then every 9 weeks.  
 Data cutoff: March 30, 2026. CR, complete response; ctDNA, circulating tumor DNA; NE, non-evaluable; PD, progressive disease; PR, partial response; SD, stable disease; VAF, variant allele frequency.

# Ovarian Cancer as Lead Indication Informed by FDA Feedback

*Targeting 1Q2027 NDA submission seeking accelerated approval*

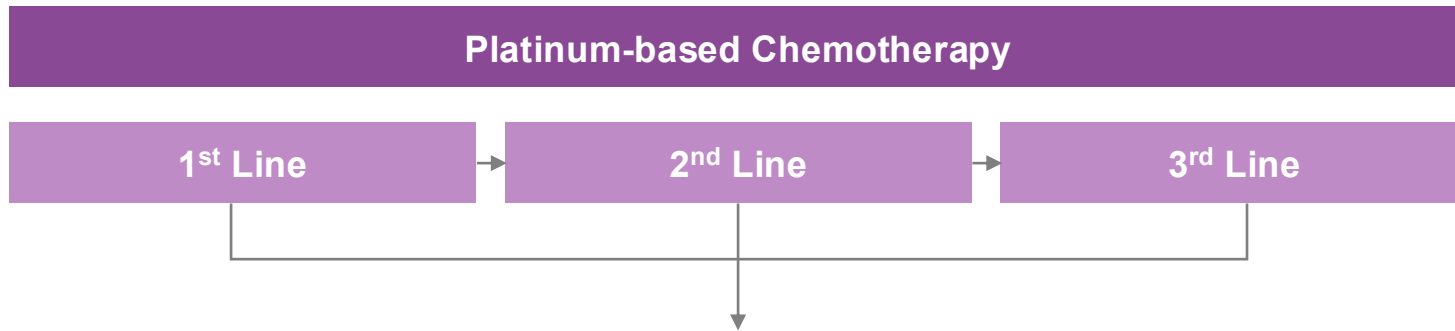
*Orphan Drug Designation granted for rezatapopt in TP53 Y220C ovarian cancer*



**Potential U.S. Launch in 2027**

# Rezatapopt Well-Positioned for Success in Ovarian Cancer and Beyond

## Platinum-Sensitive Patients



## Platinum-Resistant Patients

- Non-platinum-based Chemotherapy +/- bevacizumab
- Relicorilant + nab-paclitaxel
- Mirvetuximab (FR-alpha+)
- Trastuzumab deruxtecan (T-DXd) (HER2+)
- Pembrolizumab + paclitaxel +/- bevacizumab (PD-L1+)

### Limitation of approved options:

- Inconvenient IV administration
- AEs requiring invasive monitoring
- Chemotherapy offers limited efficacy

### Rezatapopt offers:

- Biomarker-directed approach
- Competitive and differentiated profile vs. other emerging therapies
- Convenient oral administration
- Common AEs are manageable

# TP53 Y220C Mutation is Broadly Identifiable on Existing NGS Panels

- Molecular testing is now recommended by NCCN and ESMO across many cancer types including ovarian cancer, breast cancer, NSCLC, endometrial and others
- Reimbursement of NGS testing is widely covered by Medicare and private insurance for qualifying patients



FOUNDATION  
MEDICINE

TEMPUS



Memorial Sloan Kettering  
Cancer Center

# TP53 Y220C 2L+ Ovarian Cancer Offers Meaningful Market Potential

## Total 2L+ TP53 Y220C Ovarian Cancer



**~1,700**

Addressable 2L+  
U.S. & EU4/UK Patients<sup>1</sup>



**~\$350 - 420M**

U.S. Market  
Potential<sup>2</sup>



**~\$520 - 630M**

Global Market  
Potential<sup>3</sup>

- Ovarian cancer patient population will be pursued as initial NDA submission
- Label expansion potential in other tumors

# Future Opportunities to Grow Rezatapopt Beyond Ovarian Cancer

## Monotherapy

### Endometrial

- Monotherapy data continues to be generated in Phase 2 PYNACLE
- 2L+ endometrial cancer has the potential to add ~350 patients in U.S. and EU4/UK

### Breast

- Monotherapy data continues to be generated in Phase 2 PYNACLE
- 2L+ Breast cancer has the potential to add ~2,000 patients in U.S. and EU4/UK

## Combination

### Solid Tumors

- Bevacizumab (PSOC)
- KRAS inhibitors (NSCLC, Pancreatic, CRC)

### Hematologic

- R/R AML/MDS in combination with azacitidine (ongoing IIT)
- Newly diagnosed AML/MDS in combination with azacitidine and venetoclax

# Compelling Efficacy and Defined Registrational Path for Rezatapopt



In the Phase 2 PYNNACLE trial interim data, rezatapopt demonstrated an ORR of 44% in ovarian cancer with a median DoR of 8.2 months

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NDA submission planned in 1Q2027 in platinum-resistant/refractory ovarian cancer patients

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