



# PMV Pharmaceuticals

March 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 2.9% of ovarian cancer and 1% of all solid tumors

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

Pivotal Phase 2 PYNNAACLE 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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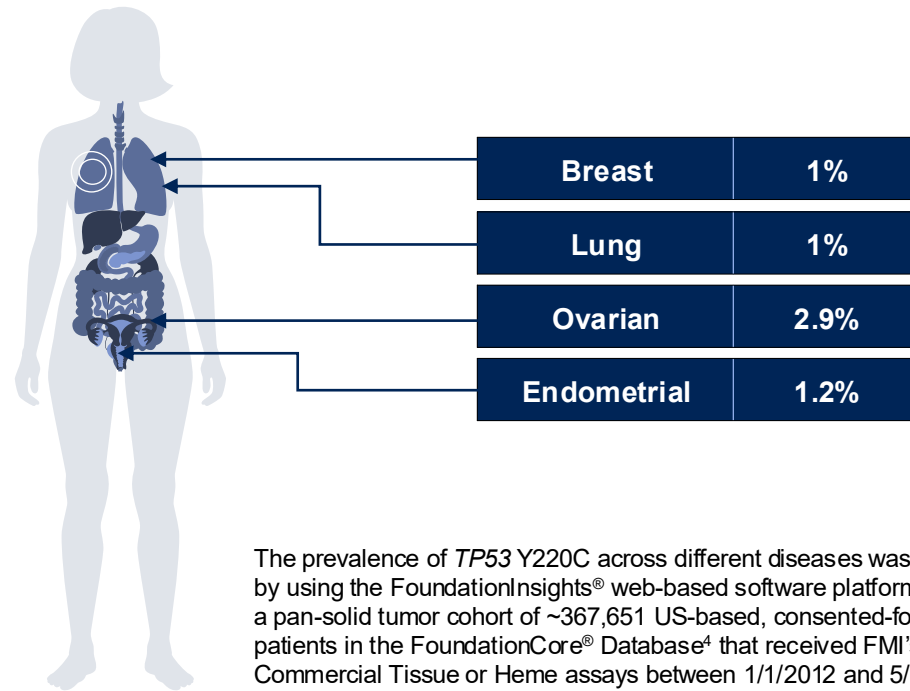
Strong balance sheet with \$112.9 M as of December 31, 2025, with cash runway through 2Q2027

# 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>**

## Frequency of *TP53* Y220C Across Common Solid Tumors

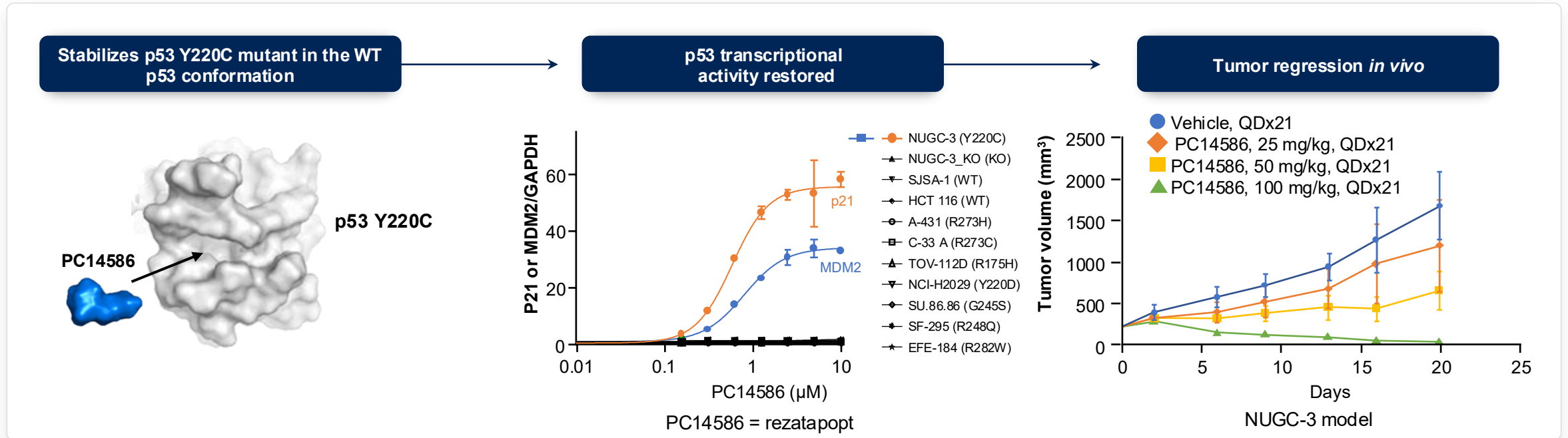
Foundation Medicine Tissue and Heme assay test results collected between 1/1/2012 and 5/31/2024



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> Foundation Insights, Schram *et al.* AACR-NCI-EORTC Conference 2023. <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: 46%
  - Median duration of response: 8.0 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



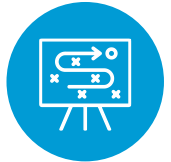
Overall results across all cohorts

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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

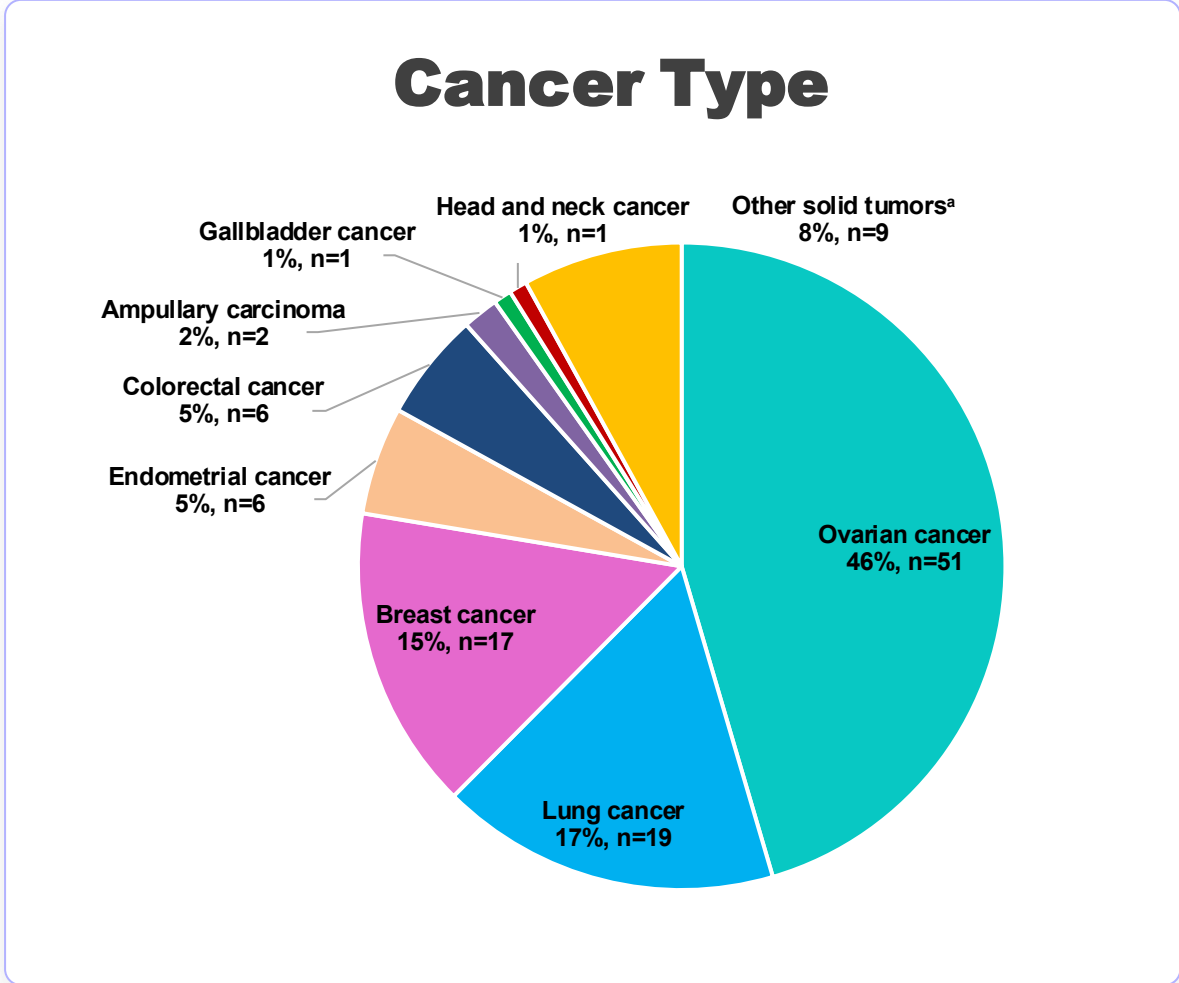
<p><b>Patient Population</b></p> <ul style="list-style-type: none"> <li>• Aged ≥ 12 years</li> <li>• Locally advanced or metastatic solid tumors, excluding primary CNS tumors</li> <li>• Documented <i>TP53</i> Y220C and <i>KRAS</i> WT only</li> <li>• Prior standard therapy or ineligible for appropriate 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%

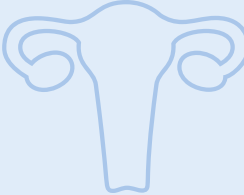


# Demographics and Baseline Characteristics (Ovarian Cancer)

*Heavily pre-treated population with poor prognostic features*

	n=51
Age (years)	Median 67
ECOG Performance Status	0: 47%, 1: 51%
Prior lines of systemic therapy	Median 4 prior lines (range 1-10) 3 or more prior lines: 73%
Prior therapies	Platinum-based tx: 100% Bevacizumab: 78% PARP inhibitors: 59%
Platinum status at study entry	Platinum-resistant: 59% Platinum-refractory: 35%* Platinum-sensitive 6%
Histology	High grade serous: 96%
Somatic BRCA1/2 mutation	BRCA1: 8%, BRCA2: 4%

**Heavily pre-treated patients:**



- **94%** platinum-resistant or refractory
- **78%** received prior bevacizumab
- **73%** with three or more prior lines of therapy

\* Including 14% (n=7) primary platinum-refractory

Data Cutoff 04Sep2025

# 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 (%)
<b>ORR per Investigator assessment</b>	<b>35 (34%)</b>
Confirmed Complete Response (CR)	1
Confirmed Partial Response (PR)	29
Unconfirmed Partial Response (uPR)	5

By Cohort	ORR n (%)
Ovarian	22/48 (46%)
Platinum Resistant/Refractory	21/45 (47%)
Platinum Sensitive	1/3 (33%)
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.
- **Ovarian cancer cohort:** ORR reached 50% (24/48) following 2 additional responses, comprising 1 confirmed CR, 22 confirmed PRs and 1 uPR pending confirmation.

Data Cutoff 04 Sep 2025

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

**Ovarian Cancer  
46% ORR  
8.0 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.

# Consistent ~40-50% Response Rates Across Key Ovarian Subgroups

	Total ovarian cancer cohort n=48	Platinum resistant n=27	Platinum refractory <sup>b</sup> n=18	Prior bevacizumab <sup>c</sup> n=37	Prior PARP inhibitor n=29	FR $\alpha$ positive <sup>d</sup> n=21	FR $\alpha$ negative <sup>d</sup> n=20
<b>ORR<sup>a</sup> %</b>	<b>46</b>	<b>48</b>	<b>44<sup>e</sup></b>	<b>46</b>	<b>52</b>	<b>48</b>	<b>40</b>
CR	1 (2%)	-	-	1 (3%)	1 (3%)	-	-
PR	18 (38%)	12 (44%)	6 (33%)	14 (38%)	13 (45%)	9 (43%)	6 (30%)
uPR	3 (6%)	1 (4%)	2 (11%)	2 (5%)	1 (3%)	1 (5%)	2 (10%)

Data Cutoff 04 Sep 2025

**Post-data cutoff: Total ovarian cancer population:** ORR reached 50% (24/48) following 2 additional responses, comprising 1 confirmed CR, 22 confirmed PRs and 1 uPR pending confirmation.

- **46% ORR (95% CI: 31%-61%) 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**

a 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.

Investigator assessed ORR is calculated from confirmed CR, confirmed PR, and uPR.

b Platinum refractory: Defined as relapse during platinum therapy or within 1 month of the last dose of a platinum agent.

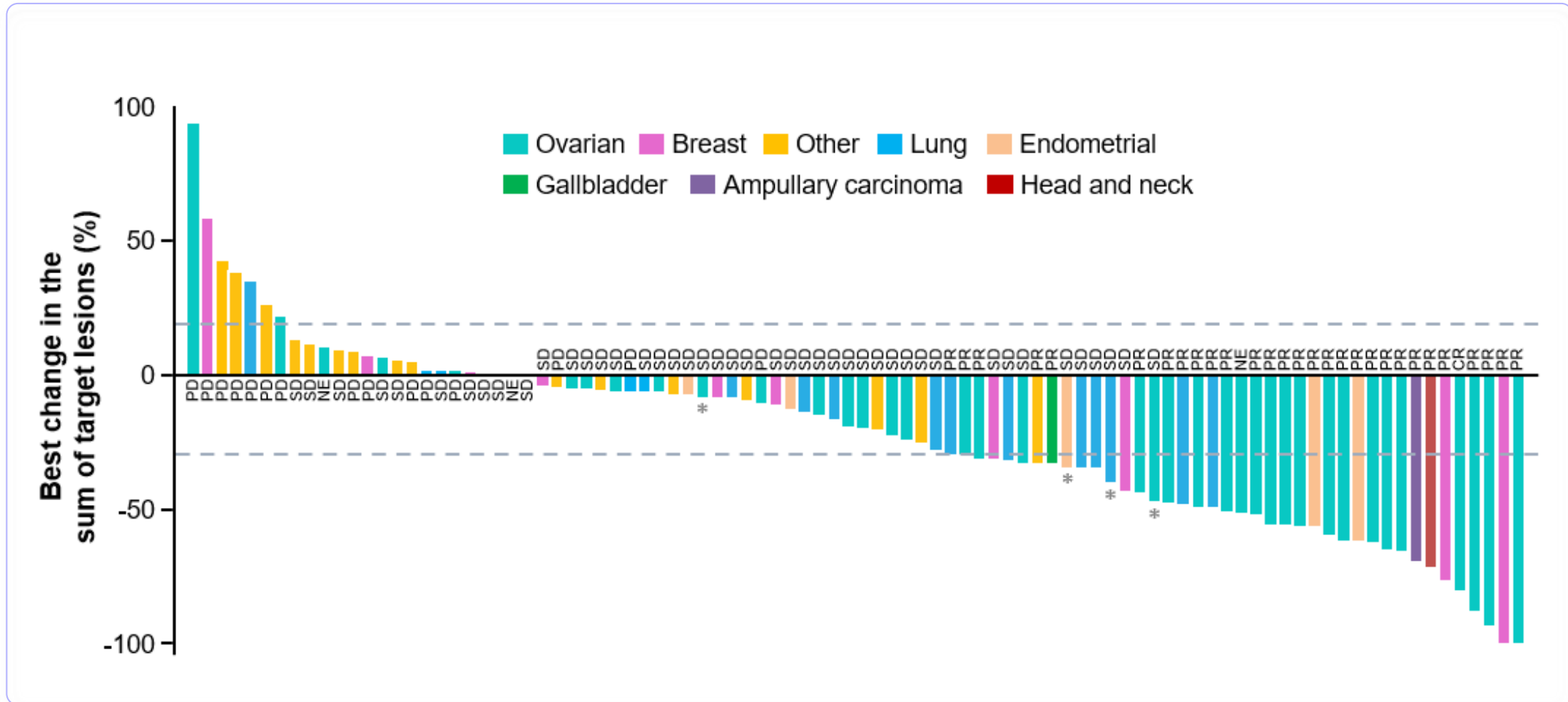
c Prior bevacizumab unless medically contraindicated.

d 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.

e In the primary platinum-refractory population (n=7), three patients experienced a confirmed PR as of 4 Sept 2025.

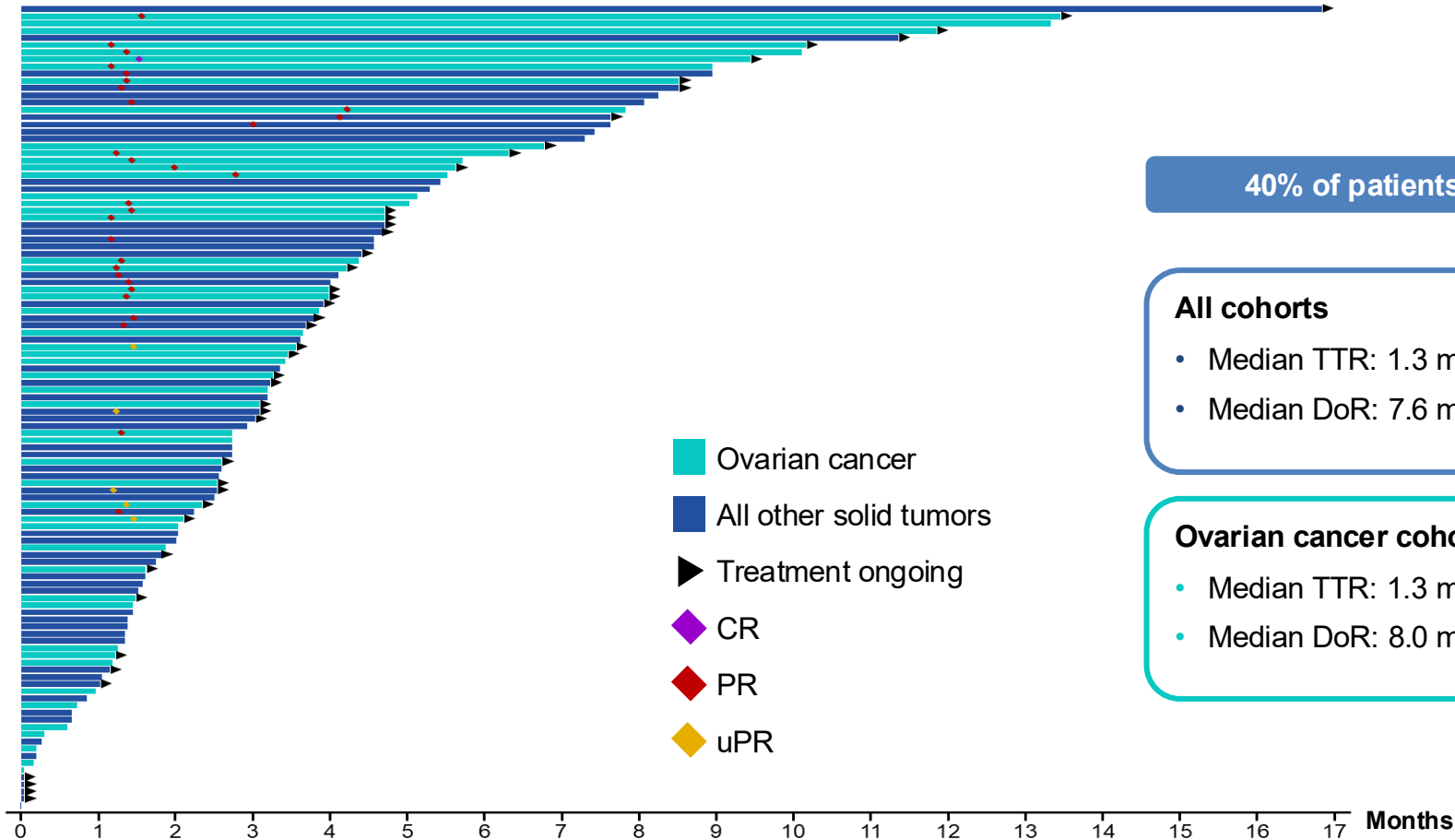
ORR: Overall Response Rate

# Target Lesion Reduction Observed in the Majority of Patients



Data Cutoff 04 Sep 2025

# Rapid Time to Response and Long Duration of Treatment



40% of patients remain on treatment

**All cohorts**

- Median TTR: 1.3 months
- Median DoR: 7.6 months

**Ovarian cancer cohort**

- Median TTR: 1.3 months
- Median DoR: 8.0 months

Data Cutoff 04Sep2025

# Favorable Safety and Tolerability

All TRAEs (≥ 10% of Patients) Preferred Term, n (%)	Overall N = 112	Grade 1	Grade 2	Grade 3	Grade 4
Nausea	38 (34)	24 (21)	13 (12)	1 (1)	-
Fatigue	26 (23)	11 (10)	13 (12)	2 (2)	-
Blood creatinine increased	22 (20)	5 (4)	16 (14)	1 (1)	-
Alanine aminotransferase increased	20 (18)	8 (7)	5 (4)	6 (5)	1 (1)
Aspartate aminotransferase increased	16 (14)	6 (5)	3 (3)	7 (6)	-
Anemia	16 (14)	5 (4)	6 (5)	5 (4)	-
Decreased appetite	14 (13)	11 (10)	3 (3)	-	-
Vomiting	13 (12)	7 (6)	6 (5)	-	-

Data Cutoff 04Sep2025

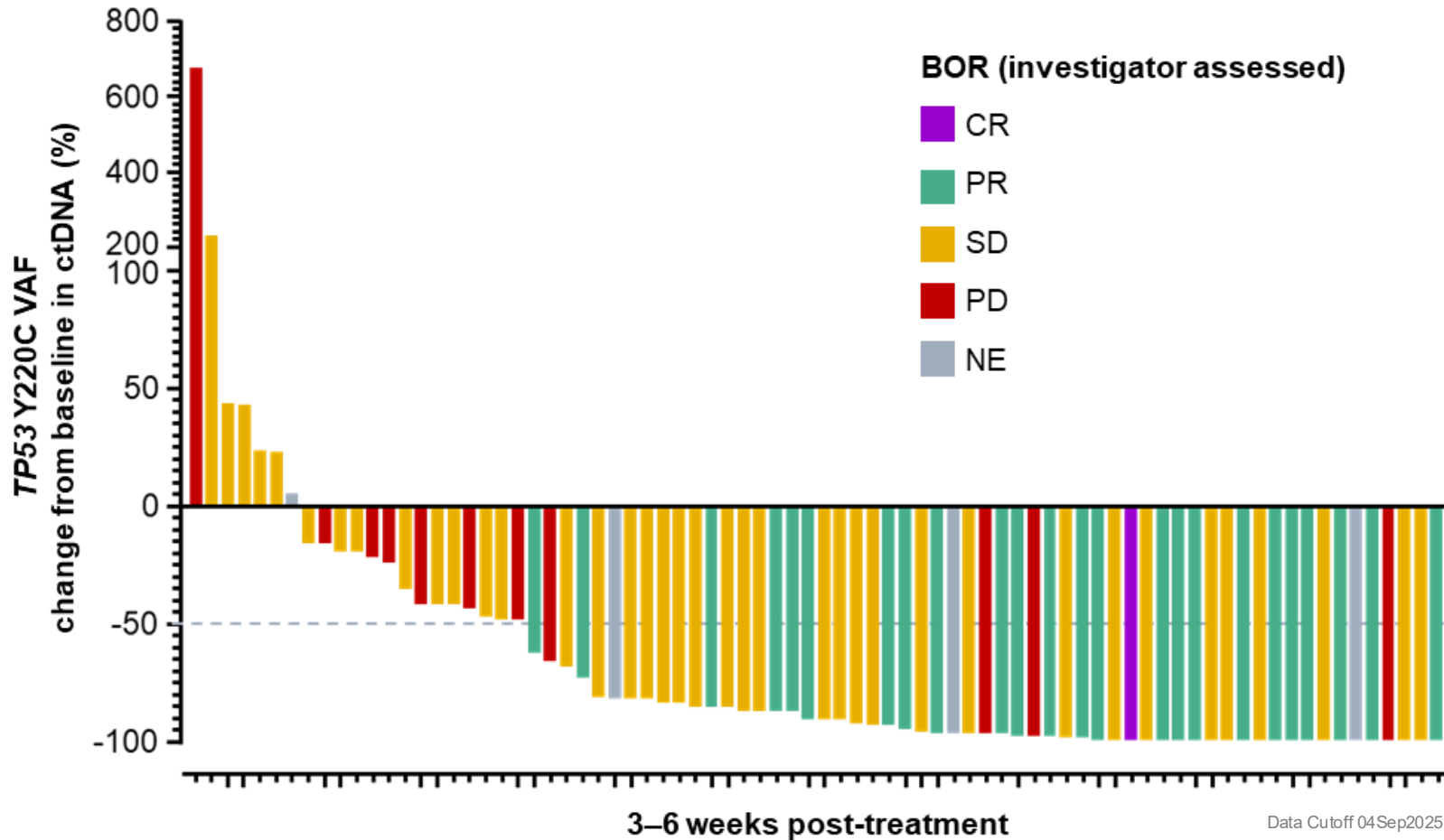
- TRAEs were mostly Grade 1/2
- Most frequent TRAEs: Nausea, fatigue, blood creatinine increased, ALT increased
- Laboratory abnormalities were manageable / monitorable with most cases being transient and reversible
- Four patients (4%) discontinued treatment due to TRAEs
- Administration of rezatapopt with food decreased incidence of gastrointestinal TRAEs compared with Phase 1<sup>1,2</sup>

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

1. Kuo H-CD, et al. Clinical Pharmacology (ACCP) 2024; Poster presentation 044;

2. Schram AM, et al. Annual Meeting on Women’s Cancer (SGO). 2024; Oral presentation (abstract LBA 26).

# On Target Activity Supported by Significant Decreases in ctDNA TP53 Y220C Mutation VAF

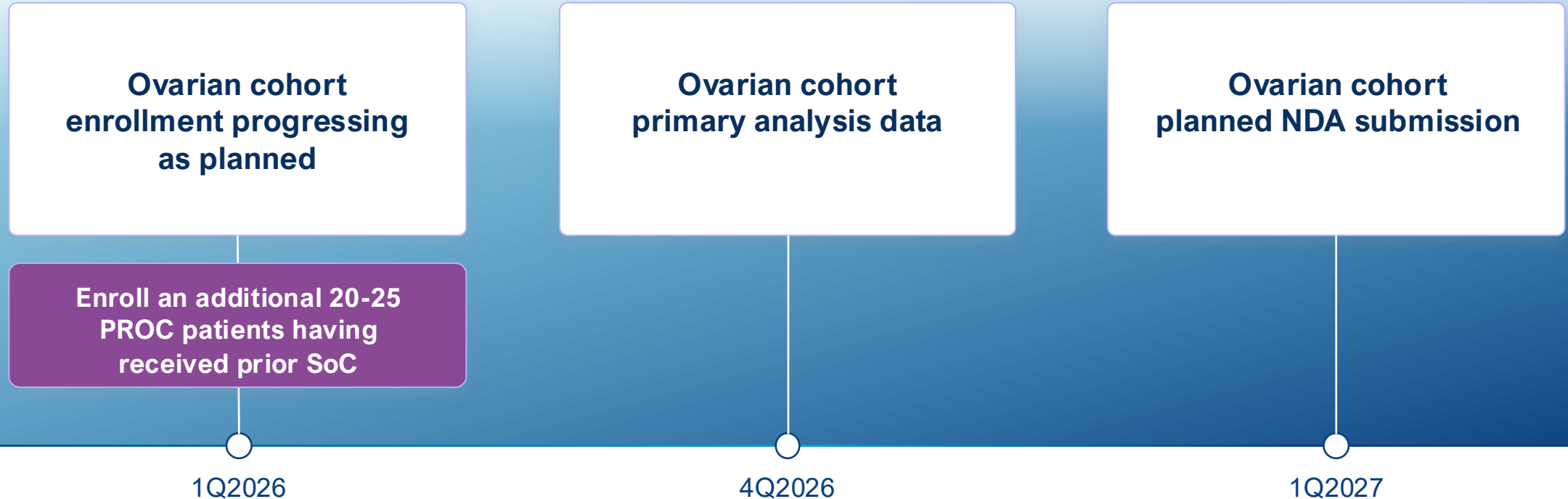


- 78 patients, including 35 ovarian cancer patients, had ctDNA TP53 Y220C VAF data available at baseline and on treatment (3-6 weeks)
- All patients experiencing a response had a reduction in TP53 Y220C VAF
- 91% had a reduction in TP53 Y220C VAF
  - 73% had a reduction of  $\geq 50\%$

# Ovarian Cancer as Lead Indication Informed by FDA Feedback

Targeting 1Q2027 NDA submission seeking accelerated approval

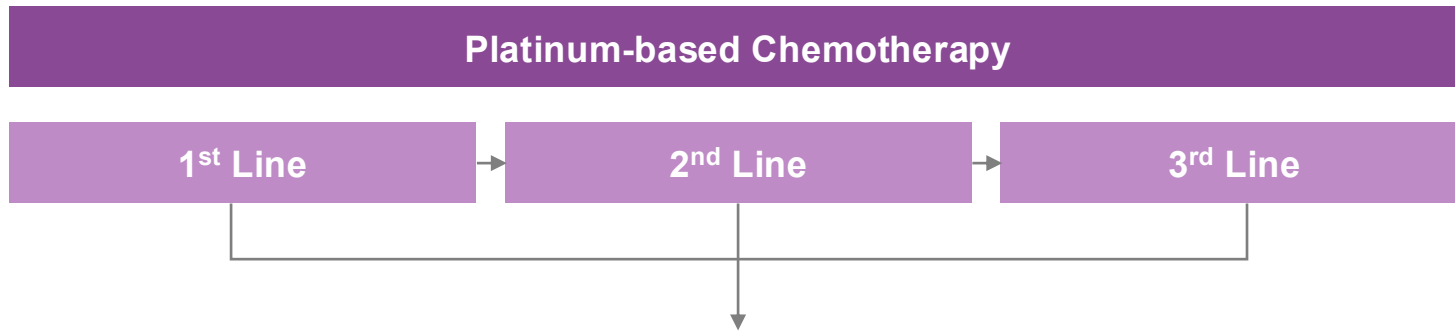
Orphan Drug Designation granted for rezatapopt in TP53 Y220C ovarian cancer



**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
- 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 46% in ovarian cancer with a median DoR of 8.0 months

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

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Strong balance sheet with \$112.9 M as of December 31, 2025, with cash runway through 2Q2027