#### **UNITED STATES** SECURITIES AND EXCHANGE COMMISSION

WASHINGTON, D.C. 20549

#### FORM 8-K CURRENT REPORT

Pursuant to Section 13 or 15(d) of the Securities Exchange Act of 1934

Date of report (Date of earliest event reported): April 9, 2024

Verastem, Inc.

(Exact Name of Registrant as Specified in Charter)

001-35403 (Commission

Delaware (State or Other Jurisdiction of Incorporation)

File Number)

27-3269467 (IRS Employer Identification No.)

02494 (Zip Code)

117 Kendrick Street, Suite 500, Needham, MA (Address of Principal Executive Offices)

Registrant's telephone number, including area code: (781) 292-4200

(Former Name or Former Address, if Changed Since Last Report)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)

Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12) Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b)) 

Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

Securities registered pursuant to Section 12(b) of the Act:

Trading Title of each class
Common stock, \$0.0001 par value per share Symbol(s) Name of each exchange on which registered The Nasdaq Capital Market VSTM

Indicate by check mark whether the registrant is an emerging growth company as defined in Rule 405 of the Securities Act of 1933 (§230.405 of this chapter) or Rule 12b-2 of the Securities Exchange Act of 1934 (§240.12b-2 of this chapter).

Emerging growth company  $\Box$ 

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act.  $\Box$ 

#### Item 7.01. Regulation FD Disclosure

On April 9, 2024, Verastem, Inc. posted its corporate presentation on its website, a copy of which is furnished hereto as Exhibit 99.1 to this Current Report on Form 8-K.

#### Item 9.01. Financial Statements and Exhibits

Exhibit No.		Description	
99.1	Corporate Presentation, dated April 9, 2024		

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Cover Page Interactive Data File (formatted in Inline XBRL)

#### SIGNATURES

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

#### VERASTEM, INC.

By: /s/ Daniel W. Paterson Daniel W. Paterson President and Chief Executive Officer

Dated: April 9, 2024

Exhibit 99.1



# Delivering Novel Therapies in RAS/MAPK Pathway Driven Cancers

April 2024 Corporate Presentation



#### Disclaimers

#### Forward-Looking

This presentation includes forward-looking statements about Verastem Oncology's programs and product candidates, strategy, future plans and prospects, including statements related to the expected outcome and benefit GenFleet Therapeutics (Shanghai), Inc. ("GenFleet"), the potential clinical value of various of its clinical trials, the timing of commencing and completing trials, including topline data reports, interactions with regulators, the commercialization of product candidates and potential for additional development programs involving Verastem Oncology's lead compound. The words "anticipate," "believe," "estimate," "expect," "intend," "may," "plan," "p "potential," "will," "would," "could," "could," "continue," 'can," "promising" and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying statement is subject to risks and uncertainties that could cause actual results to differ materially from those expressed or implied in such statement.

Applicable risks and uncertainties include the risks and uncertainties, among other things, regarding: the success in the development and potential commercialization of our product candidates, including avutometinib in con including defactinib, LUMAKRAS<sup>TM</sup> and others; the uncertainties inherent in research and development, such as negative or unexpected results of clinical trials, the occurrence or timing of applications for our product candidates; in any jurisdictions; whether and when regulatory authorities in any jurisdictions; whether and when regulatory authorities in any jurisdictions; whether and when regulatory authorities in any jurisdictions ary such applications that may be filed for our product candidates; and, if approved, whether our product candidates; the scope, timing, and outcome of any legal proceedings; decisions by trial design, labeling and other matters that could affect the timing, scope and rate of reimbursement for our product candidates; whether preclinical testing of our product candidates and preliminary or interim data from the results or success of ongoing or later clinical trials; that the timing, scope and rate of reimbursement for our product candidates may take longer than expected; that any not be available when expected; that enrollment of clinical trials may take longer than expected; that our product candidates will cause adverse safety events a arise from additional data or analysis, or result in unmanageable safety profiles as compared to their levels of efficacy; that our product candidates; that we do which could result in reduced market share or market potential for our product candidates; that we for onducting additional studies; that we are not here sults of our product candidates; that we will be unable to successfully initiate or complete the clinical development and eventua candidates; that we or by which could result in reduced market share or market potential for our product candidates; that we will be unable to successfully initiate or complete the clinical d

Other risks and uncertainties include those identified under the heading "Risk Factors" in the Company's Annual Report on Form 10-K for the year ended December 31, 2023, as filed with the Securities and Exchange Cor and in any subsequent filings with the SEC, which are available at www.sec.gov and www.verastem.com.

The forward-looking statements in this presentation speak only as of the original date of this presentation, and we undertake no obligation to update or revise any of these statements.

#### Use of Non-GAAP Financial Measures

This presentation contains references to our non-GAAP operating expense, a financial measure that is not calculated in accordance with generally accepted accounting principles in the US ("GAAP"). This non-GAAP financial measures from the corresponding financial measures determined in accordance with GAAP. Management believes this non-GAAP information is useful for investors, taken in conjunction with the Company's operating performance and enhance investors' ability to identify operating trends in the Company's builts. Management believes this non-GAAP information is useful for investors, taken in conjunction with the Company's operating performance and can enhance investors' ability to identify operating trends in the Company's builts. Management believes this non-GAAP information is not prepared under a comprehensive set of accounting rules and should only be understanding of the Company's operating results as reported under GAAP, not in isolation or as a substitute for, or superior to, financial information prepared and presented in accordance with GAAP. In addition, this ne unlikely to be comparable with non-GAAP information provided by other companies. The determination of the amounts that are excluded from non-GAAP financial measures is a matter of management judgment and depe the nature of the underlying expense or income amounts. Reconciliations between this non-GAAP financial measure and the most comparable GAAP financial measure are included in the footnotes to the slides in this pres GAAP number appears.

#### **Third-Party Sources**

Certain information contained in this presentation relates to or is based on studies, publications, surveys and other data obtained from third-party sources and Verastem Oncology's own internal estimates and research. We these third-party sources to be reliable as of the date of this presentation, it has not independently verified, and makes no representation as to the adequacy, fairness, accuracy or completeness of, any information obtained addition, all of the market data included in this presentation involves a number of assumptions and limitations, and there can be no guarantee as to the accuracy or reliability of such assumptions.



## Verastem Oncology

Strong progress in 2023 sets up multiple valuecreation opportunities

### Well-Positioned To Deliver on 2024 Catal

#### > On track to deliver the first approved therapy in LGSOC

- Data at ASCO 2023 of avutometinib, a RAF/MEK Clamp in combination with defact demonstrated robust responses in patients with recurrent low-grade serous ovariai
- Phase 3 confirmatory study underway with plans to report updated topline data fro H1 2024
- Commence rolling NDA for Accelerated Approval in H1 2024
- > Ongoing studies in additional indications including Pancreatic Cance
  - Report initial safety and efficacy results from RAMP 205 trial of avutometinib + gem paclitaxel + defactinib in first-line metastatic pancreatic cancer in H1 2024
  - Report updated data from both non-small cell lung cancer (NSCLC) trials RAMP 2 Amgen) and RAMP 204 (adagrasib-Mirati) trials in Mid-2024

#### > GenFleet collaboration furthers pipeline potential in RAS/MAPK dri

- GenFleet's IND application for GFH375/VS-7375, a potential best-in-class oral KRA (ON/OFF) inhibitor, was filed in China and accepted for review; upon clearance exp Phase I trial in China in H2 2024
- Expect to initiate Phase I trial for GFH375/VS-7375 in China in H2 2024
- · Ongoing discovery/lead optimization for second and third programs
- > Strong balance sheet to support ongoing programs and operations
  - Company ended Q4 2023 with \$137.1M in cash and investments and \$31.1 million expenses (\$29.5 million non-GAAP operating expenses\*)

\* Q4 2023 GAAP operating expenses - \$31.14M less Q4 2023 stock compensation of \$1.60M = \$29.54M Q4 2023 non-GAAP oper IND: investigational new drug application; NDA: new drug application

# Driving Momentum in 2024: Recap of Recent Key Achiever

	NSCLC	Oral G12D Inhib
<ul> <li>Received FDA Orphan Drug Designation</li> <li>Initiated Phase 3 confirmatory study</li> <li>Presented planned subgroup analysis of Part A RAMP 201 trial</li> <li>RAMP 201 FDA meeting – combination selected as go- forward regimen</li> <li>Initiated RAMP 205 combo avutometinib + gemcitabine/nab-paclitaxel + defactinib</li> </ul>	<ul> <li>Received FDA Fast Track Designation for avutometinib in combo with Amgen's G12C inhibitor sotorasib</li> <li>Presented initial results from Phase 1/2 RAMP 203 trial of avutometinib + sotorasib in KRAS G12C mutant NSCLC</li> <li>Added defactinib to avutometinib and sotorasib combination in the RAMP 203 trial</li> </ul>	<ul> <li>✓ Established dis development of with GenFleet</li> <li>✓ Selected GFH potential best- KRAS G12D ( inhibitor</li> <li>✓ IND application China and accord</li> </ul>

# Clinical Program Designed for Success in LGSOC, Signal Generation

Regimen	IND-Enabling/ Preclinical	Phase 1	Phase 2	Phase 3	Anticipated Milestones
Avutometinib + De	efactinib: Recurren	t LGSOC			
RAF/MEK Clamp + FAKi					RAMP 301 Ongoing Enrollment
RAF/MEK Clamp + FAKi					RAMP 201 Topline Data; Rolling NDA Submission Seeking Accelerated Approval: H12024
Avutometinib + KF	RAS G12C Inhibitor	s: NSCLC			
RAF/MEK Clamp + KRAS G12Ci (sotorasib) + FAKi					RAMP 203 Updated Data Mid-2024
RAF/MEK Clamp + KRAS G12Ci (adagrasib)					RAMP 204 Updated Data Mid-2024
Avutometinib + De	efactinib: Metastati	c Pancreatic Ca	ncer		
RAF/MEK Clamp + FAKi + gemcitabine, nab-paclitaxel					RAMP 205 Initial Safety/Efficacy H12024
GFH375/VS-7375					
G12D (ON/OFF) inhibitor					IND filed in China and accepted for review; upon clearance expect to initiate Phase 1 in China in H22024
ONCOLOGY					

# Avutometinib RAF/MEK Clamp Program Overview

### Avutometinib is a Differentiated Agent with the Potential to Serve a Backbone for Combinations Across RAS Pathway-Driven Cancers

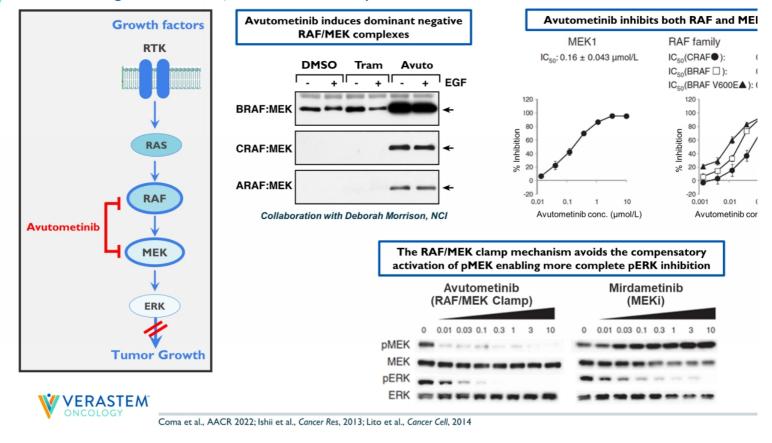
- Unique RAF/MEK clamp mechanism of action
- Novel intermittent dosing schedule; convenient oral regimen
- Orphan Drug Designation for avutometinib alone or in combination with defactinib in recur
- Breakthrough Therapy Designation for combination of avutometinib and defactinib for treatr recurrent LGSOC after one or more prior lines of therapy including platinum-based chemot
- Received FDA Fast Track Designation for avutometinib in combination with Amgen's G12C in sotorasib in KRAS G12C-mutant NSCLC
- Potential best-in-class safety & tolerability profile relative to marketed MEK inhibitors, with combinability with agents from multiple target classes
- Promising signals of clinical activity in various RAS pathway-driven cancers, including in patien tumors previously progressed on other MEK inhibitors



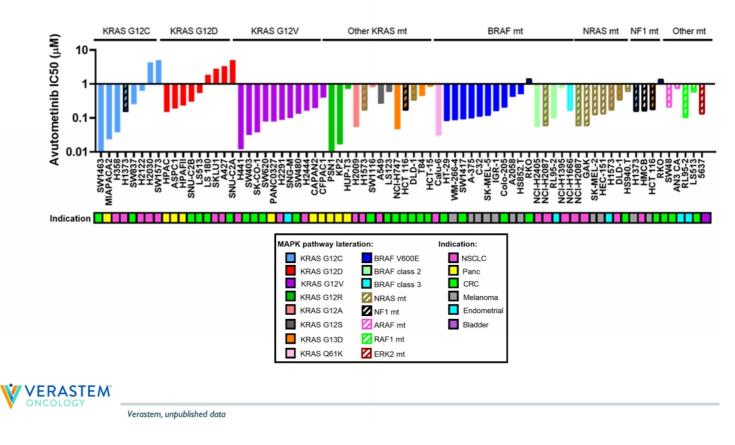
RAF-Rapidly accelerated fibrosarcoma, MEK-Mitogen-activated protein kinase kinase, RAS-Rat sarcoma virus MAPK-Mitogen-activated protein kinase KRAS-NRAS-Neuroblastoma RAS viral oncogene homolog, BRAF-v-raf murine sarcoma viral oncogene homolog B1, NF1-Neurofibromatosis type 1

## Avutometinib is a Unique Small Molecule RAF/MEK Clamp

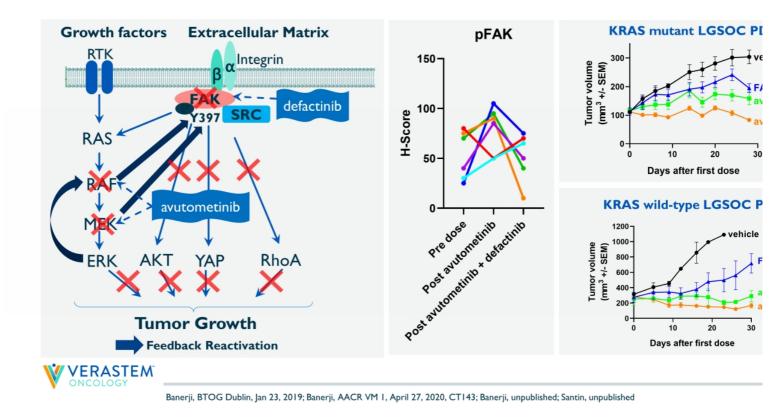
Contrasting Mechanism of Action vs. MEK-Only Inhibitors



## Avutometinib Inhibits Cell Proliferation Across Multiple RAS/MAPK Alterations and Multiple Solid Tumor Histologies



#### Strong Scientific Rationale for Avutometinib and FAK Inhibitor Con Anti-Tumor Efficacy in KRAS Mutant and Wild-Type LGSOC models



## Optimized Dosing Schedule Defined: Favorable Tolerability Profile with Novel Intermittent Dosing Regim

Summary of Adverse Events Grade  $\geq$  3 Occurring in  $\geq$  5% of patients

	Avutometinib monotherapy Daily at MTD N=6 28-day cycle	RP2D Avutometinib monotherapy 4mg twice weekly N=26 28-day cycle	RP (Avutometini weekly + defi twice N= 21 days of 2
Treatment Related Adverse Event	Grade ≥3	Grade ≥3	Grad
Rash	3 (50%)	5 (19%)	2 (!
CK elevation (Creatine phosphokinase)	I (17%)	2 (8%)	2 (!



<sup>1</sup> Chenard-Poirier, et al. ASCO 2017; References: Banerji, Q4 2020 report; Data on file; RP2D: recommended phase 2 dosing

# RAS Pathway-Driven Cancers and Rational Avutometinib Combinations

## Ongoing Comprehensive Approach to Establish More Complete Ble RAS Pathway & Resistance Pathways

	Indication	Incidence/ Prevalence	Biomarker	% Regimen	Setting	P
Incidence <sup>2,3</sup> : I I 4K	RAMP301 LGSOC	Prevalence <sup>1</sup> : 6K	70%	Avutometinib + defactinib	Relapsed Refractory molecularly profiled LGSOC	Phase 3 Confirmat
Gynecologic	RAMP201 LGSOC	Prevalence <sup>1</sup> : 6K	70%	Avutometinib + defactinib	Relapsed Refractory molecularly profiled LGSOC	Phase 2 Registratio AA cohor
Gynecologic — Cancers	Gynecologic Basket*	Incidence <sup>6-10</sup> : 85K	25%	Avutometinib + defactinib	Recurrent RAS Pathway-driven (RAS/RAF/NF1) endometrioid cancer, mucinous ovarian cancer, high-grade serous ovarian cancer or cervical cancer	Phase 2
ĺ	Mesonephric	Incidence: <sup>11</sup> ~680	96%	Avutometinib + defactinib	Advanced or recurrent mesonephric gynecologic cancer	Phase 2
NSCLC Adenocarcino	RAMP203 KRAS GI2C	Incidence <sup>2,3</sup> :		Avutometinib + sotorasib ± defactinib	Recurrent KRAS G12C with prior KRAS G12C inhibitor(i) treatment or KRAS G12Ci naïve	Phase
ma	RAMP204 KRAS GI2C	THE	$\bigcirc$	Avutometinib + adagrasib	Recurrent KRAS G12C with prior KRAS G12Ci treatment that progressed	Phase
Pancreatic	RAMP205 PDAC	Incidence⁴: 58K		Avutometinib + defactinib + gemcitabine/nab-paclitaxel	Previously untreated (front-line) metastatic pancreatic ductal adenocarcinoma (PDAC)	Phase
CRC	KRAS mt*	Incidence <sup>5</sup> : I 48K	45%	Avutometinib + cetuximab	Recurrent metastatic KRAS mt	Phase
Breast Cancer	ER+*	Incidence <sup>5</sup> : 279K		Avutometinib + abemaciclib + fulvestrant	Recurrent ER+/HER2- breast cancer following progression on CDK4/6i + aromatase inhibitor	Phase
Thyroid	MAPK alterations*+	Incidence⁴: 44K	35%	Avutometinib + defactinib	Differentiated & anaplastic thyroid cancer	Phase

<sup>1</sup> Plonk, Bandli, Grisham, The Enving Landscape of Chemothemary in Newly Dapposed Advanced Epithelial Ovarian Cancer, An Soc Cin Drowal, Base Book, 2019, Carley, Carry, Majka, Shih, Huanama, Fider, Grisham et al., Low-Grinde serves varian cancer: State of the Skinger, Gynosed Done; 2020 Grisham, Plance Participant, Shih, Huanama, Fider, Grisham et al., Low-Grinde serves varian cancer: State of the Skinger, Gynosed Done; 2020 Grisham, Plance Done

# Avutometinib ± Defactinib in Low-Grade Serous Ovarian Cancer

## LGSOC Unmet Need & Opportunity

- LGSOC is a less common type of ovarian cancer that is often diagnosed in younger women
  - LGSOC is a unique disease that is distinct from high-grade serous ovarian cancer (HGSOC) in its pathology, protracted clin response to chemotherapy and thus requires a more tailored therapeutic approach
  - An estimated 1,000-2,000 patients are diagnosed with LGSOC per year in the U.S., with prevalence of ~6,000
  - There are currently no approved therapies specifically indicated for recurrent LGSOC
    - o Recent clinical trials in recurrent LGSOC showed that standard-of-care chemo and hormonal therapy are relatively ineffectiv
    - LGSOC has a chemo-resistant nature and optimal treatment has not yet been defined. NCCN guidelines include clinical tria highlighting the lack of approved & effective therapies
- LGSOC is known to be driven by the MAPK (RAS) pathway in ≥70% of patients
- A phase I/II study in the UK (FRAME) evaluated the combination of avutometinib and defactinib
  - Results in recurrent LGSOC showed a <u>42%</u> confirmed ORR with durable responses and favorable safety/tolerability
- RAMP 201:A registration-directed Phase 2 trial of avutometinib and avutometinib + defactinib in recurrent LGSO(
  - Updated data from ASCO 2023 showed a <u>45%</u> confirmed ORR in the combination arm with tumor shrinkage in 86% of evaluation are shrinkage in 86% of evaluati
- RAMP 301:A confirmatory Phase 3 trial evaluating the combination of avutometinib and defactinib versus standarc hormonal therapy for the treatment of recurrent LGSOC

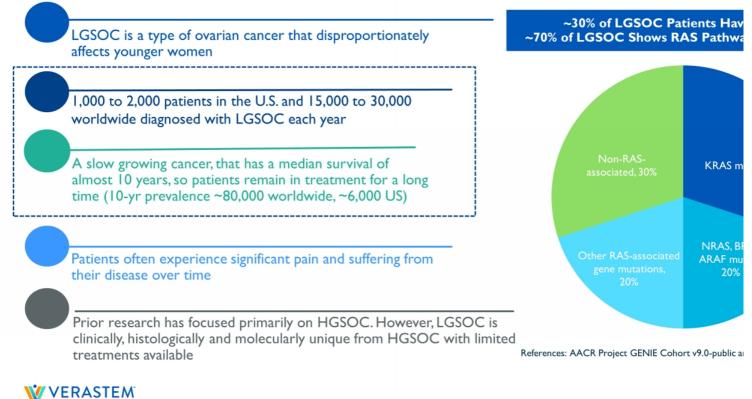
Orphan Drug Designation for avutometinib alone or in combination with defactinib in recurrent LGSOC

> Breakthrough Therapy Designation granted for avutometinib and defactinib in recurrent LGSOC after one or more



Monk et al., The Evolving Landscape of Chemotherapy in Newly Diagnosed Advanced Epithelial Ovarian Cancer, 2019; Slomovitz et al., Low-Grade serous ovarian cancer: State of the Science, 202 Grade Serous Ovarian Cancer: Current Treatment Paradigms and Future Directions, 2018; AACR. Project GENIE Cohort v9.0-public and Verastem unpublished analysis; Banerjee et al., Phase 1 stu the dual RAF/MEK inhibitor VS-6766 and the FAK inhibitor defactinib: Results of efficacy in low grade serous ovarian cancer, ESMO 2021; Malpica et al., Interobserver and intraobserver variability grading ovarian serous carcinoma, 2007; NCCN guidelines v1.2023; Zwimpfer et al. Cancer treatment Reviews 112 (2023).

## LGSOC is a Unique RAS Pathway-Driven Cancer with a High Unm



Monk, Randall, Grisham, The Evolving Landscape of Chemotherapy in Newly Diagnosed Advanced Epithelial Ovarian Cancer, Am Soc Clin Oncol Educ Book; 2019; Slomovitz, Gourley, Carey, Malpi Low-Grade serous ovarian cancer: State of the Science; Gynecol Oncol; 2020. Grisham, Iyer, Low-Grade Serous Ovarian Cancer: Current Treatment Paradigms and Future Directions; Curr Treat ( Surg Pathol 2007

# Low-Grade and High-Grade Serous Ovarian Cancer Are Different I

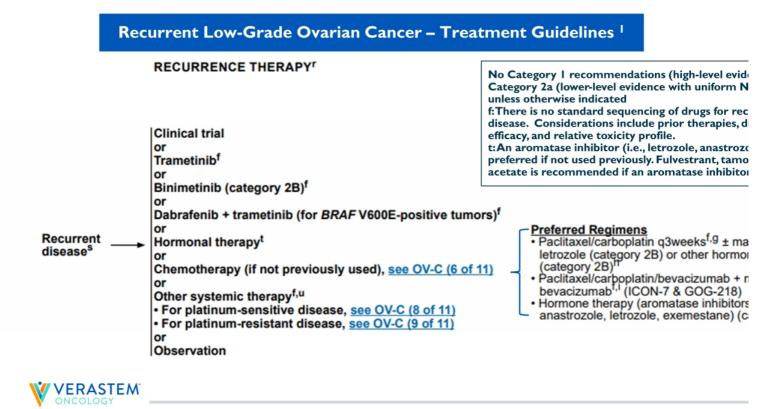
LGSOC

HGSOC

Variable	LGSOC	HGSOC
Nuclear atypia	Uniform round to oval with little variation	+++ Marked variation
Mitotic Index	<12 mitoses per 10 hpf	>l2 mitoses per l0 hpf
Chromatin and variation in size of nucleus	Little	Marked (nuclear size ratio ≥3)
Mutation	KRAS ++ BRAF + ER/PR +++	P53 +++ BRCA1/2 +
Precursor	Serous borderline tumor	Tubal intraepithelial neoplasia

Malpica et al., Am J. Surg Pathol 2007

### Recurrent LGSOC: High Medical Need No Approved Treatment Options – Limited Benefit from Available Therapic



<sup>1</sup> NCCN guidelines v1.2023

# Recent LGSOC Trials Highlight High Unmet Need

Trial	Number of Prior Systemic Therapies Median (range)	Prior MEK allowed	Prior Bevacizumab	Therapy	Response Rate ORR	Image Assessment	Median PFS Months (95% Cl												
GOG	2	No	* Low %	SoC (n=I30)	6% 95% CI: (3%, 12%)	INV	7.2 (5.6-9.9)												
2811	2811 (1-10)		* Low /⁄o	Trametinib (n=130)	26% 95% CI: (19%, 35%)	INV	13.0 (9.9-15.0)												
MIL O2	2							N	NL	NL	NL	NL		No	* 1 9/	SoC (n=101)	13% 95% CI: (7%, 21%)	BICR	10.6 (9.2 - 14.5)
MILO <sup>2</sup> (1-8)	INO	* Low %	Binimetinib <sup>2</sup> (n=198)	16% 95% CI: (11%, 22%)	BICR	9.1 (7.3-11.3)													

<sup>1</sup> Study GOG 281 trial Gershenson et al., Lancet 2022

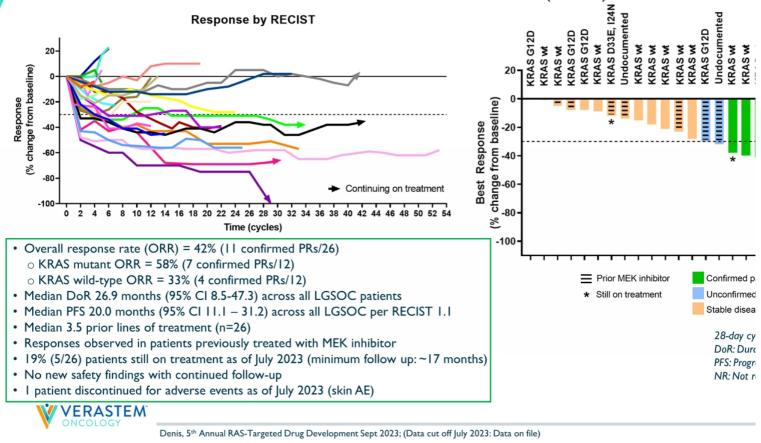
<sup>2</sup> MILO Study Monk et al., J Clin Oncol 2020.

\* Low historical use of bevacizumab during trial conduct. % not reported MILO: no more than 3 lines of prior chemotherapy

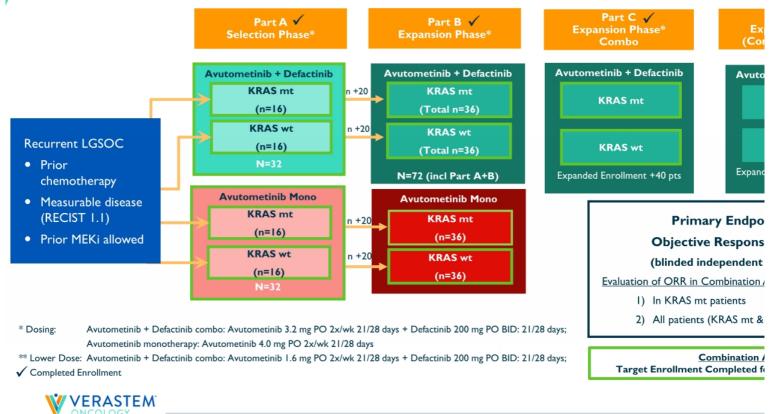
SoC = Standaı (endocrine / cł INV = Investig BICR = Blindea PFS = Progres CI = confidenc

NR = Not read

# FRAME Study: High Rate of Durable Responses with the Combination of Avutometinib and Defactinib in Recurrent LGSOC (n=26)



# RAMP 201 (ENGOTov60/GOG3052): Registration-Directed Phase Avutometinib ± Defactinib in Patients with Recurrent LGSOC



#### ASCO 2023 data Updated Data from Part A of RAMP 201

"These results demonstrate avutometinib in combination with defactinib can deliver high response rates for patients with recurrent LGSOC with a promising safety profile to date. It is particularly encouraging to see extensive tumor shrinkage in women who have had several treatment lines, including prior MEK inhibitors. These latest findings suggest the combination may offer a new treatment option for women with this hard-to-treat cancer, and we are hopeful it will become the new standard of care."

#### -Dr. Susana Banerjee, MBBS, MA PhD, FRCP, global

and lead investigator of the study, Consultant Medical Oncologist at The Royal Marsden NHS Foundation Trust and Team Leader in Women's Cancers at The Institute of Cancer Research, London

	Avutometinib + Defactinib				
	Total (n=29)				
	<b>45% (13)</b> 95% Cl: (26%, 6	64%)			
ORR, % (n)	KRAS mt 60% (9/15)				
Tumor shrinkage, % (n)	86% (25)				
Median Time to Response	5.5 months (range 1.6-14.7 months)				
Relative avutometinib Dose Intensity	83% ± 20%				

- 29 patients were evaluable for efficacy with a minimum follow-up of 12 months and 13 remain on study treatment
- Patients were heavily pretreated with a median of 4 prior systemic regimens (up to 11
  3 out of 4 patients who received prior MEK inhibitors responded to the combir
- Median duration of response and median progression free survival have not been reac
- Safety and tolerability continued to be favorable and consistent with previously report
  - The discontinuation rate due to ≥ I adverse event was 12% in the combination due to elevated blood CPK)

### Recent LGSOC Trials with Standard of Care Highlight High Unmet Need Current Trials with Avutometinib + Defactinib Show Overall Response R

Trial	Median Number of Prior lines of Therapy	Prior MEK Allowed	Prior Bevacizu mab	Therapy	Response Rate ORR	Image Assessment	Median PFS Months (95% CI)
GOG 2811	2	No	* Low %	Standard of Care	6% ^ 95% CI: (3%, 12%)	INV	7.2 (5.6-9.9)
(1-10)	(1-10)	NO	LOW /6	Trametinib	26%^ 95% CI: (19%, 35%)	INV	13.0 (9.9-15.0)
MIL O2	2		No * Low %	Standard of Care	13% 95% CI: (7%, 21%)	BICR	10.6 (9.2 to 14.5)
MILO <sup>2</sup> (1-8)	NO	Binimetinib		16% 95% CI: (11%, 22%)	BICR	9.1 (7.3-11.3)	

FRAME <sup>3</sup>	3.5	Yes	19 %	Avutometinib + Defactinib	<b>42%^</b> 95% Cl: (23%, 63%)	INV	<b>20</b> (   - 3 )
RAMP 201 Part A (ASCO 2023 data) <sup>4</sup>	4	Yes	65%	Avutometinib + Defactinib	45% 95% CI: (26%, 64%) 52%***	BICR	Not Yet Reached

Study GOG 281 trial Gershenson et al., Lancet 2022

<sup>2</sup>MILO Study Monk et al., J Clin Oncol 2020.

<sup>3</sup> Banerjee et al., ESMO Sept 2021

<sup>4</sup> Banerjee et al., ASCO June 2023 \* Low historical use of bevacizumab during trial conduct. % not reported MILO: no more than 3 lines of prior chemotherapy

🖉 VERASTEM



SoC = Standard of Care

GOG 281: (chemotherapy / endocrine therapy)

PLD (liposomal doxorubicin), paclitaxel, topotecan, letrozole or tamoxifen

MILO: (chemotherapy only)

PLD (liposomal doxorubicin), paclitaxel or topotecan

INV = Investigator BICR = Blinded indet PFS = Progression fr

\*\*\* Confirmed + Ui

CI = confidence inter AE = adverse event

\*\*12% discontinuation in all combination pts (Part A + B (n=81): 4

#### ASCO 2023 data

#### RAMP 201 Part A: Heavily Pre-Treated Patient Population

Prior Platinum-Based Chemotherapy, Endocrine Therapy, Bevacizumab in Most Patients; Prior MEK Inhibitor Therapy was Permitted

	Avutometinib Monotherapy			Avutor	metinib +
	KRAS mt	KRAS wt	Total	KRAS mt	KRAS
	(n=16)	(n=17)	(n=33)	(n=16)	(n=15
Age (yrs), median (min, max)	58 (27, 72)	48 (27, 74)	51 (27,74)	61 (29,71)	50 (30,
ECOG PS, n (%)					
0	8 (50)	15 (88)	23 (70)	11 (69)	9 (60)
1	8 (50)	2 (12)	10 (30)	5 (31)	6 (40)
Number of Prior Systemic Regimens, median (min, max)	4 (1,10)	3 (1,9)	3 (1,10)	4 (1,8)	5 (2, 1
Prior platinum-based chemotherapy, n (%)	15 (94)	17 (100)	32 (97)	16 (100)	15 (100
Prior MEK inhibitor therapy, n (%)	5 (31)	5 (29)	10 (30)	2 (13)	2 (13)
Prior Bevacizumab, n (%)	8 (50)	8 (47)	16 (48)	7 (44)	13 (87
Prior Hormonal therapy, n (%)	(69)	13 (76)	24 (73)	15 (94)	13 (87



#### ASCO 2023 data

#### RAMP 201 Part A: Evaluable Patient Population\*

Positive ORR Confirmed by Blinded Independent Central Review (BICR) Support Avutometini Defactinib as Go Forward Regimen in LGSOC - Regardless of KRAS Status

		Avutometinib	A	vutometinib + Defa	
	KRAS mt	KRAS wt	Total	KRAS mt	KRAS wt
	(n=15)	(n=15)	(n=30)	(n=15)	(n=14)
Confirmed ORR, n (%)	2 (13)	I (6)	3 (10) 95% CI (2%, 24%)	9 (60)	4 (29)
CR, n (%)	I (7)	0	I (3)	0	0
PR, n (%)	I (7)	I (6)	2 (7)	9** (60)	4 (29)
SD, n (%)	12 (80)	3 (81)	25 (83)	6 (40)	7 (50)
Disease control rate <sup>***</sup> , n (%)	14 (93)	14 (88)	28 (93)	15 (100)	(79)
PD, n (%)	I (7)	2 (13)	3 (10)	0	3 (21)
Confirmed + unconfirmed ORR, n (%)	2 (13)	I (6)	3 (10)	(73)	4 (29)

\* Evaluable for Efficacy: At least one blinded imaging assessment in 31 of 33 and 29 of 31 patients enrolled in respective treatment arms

\*\* Includes patient deepened to CR at last assessment; CR not yet confirmed

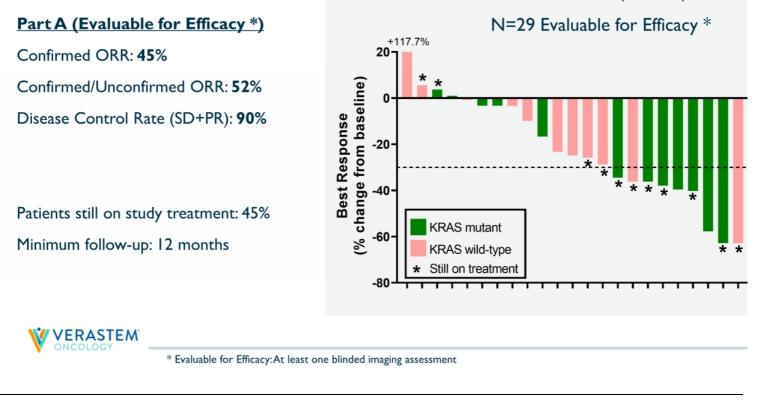
\*\*\*Disease control rate (SD + PR + CR) at 8 weeks.



BICR, blinded independent central review; mt, mutant; ORR, objective response rate; PD, progressive disease; PR, partial response; SD, stable disease; wt, wi

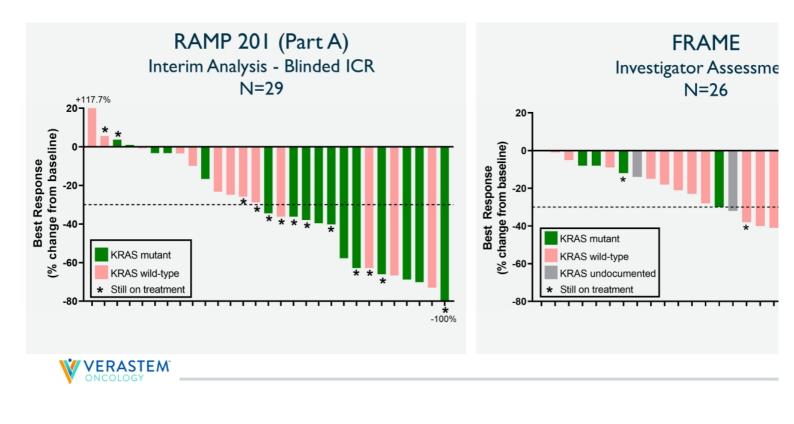
#### ASCO 2023 data Combination of Avutometinib and Defactinib High Disease Control Rate + Tumor Reduction in Recurrent LGSOC

RAMP 201 (Part A)



#### ASCO 2023 data

## Combination of Avutometinib and Defactinib Initial Data from RAMP 201 Trial Reinforce Findings from FRAME Trial



#### ASCO 2023 data

RAMP 201: Safety and Tolerability Profile of Avutometinib + Defacti No New Safety Signals; Few Discontinuations Due to Adverse Events

Most Common Treatment-Related Adverse Events (>20%) in All Treated Patients

- Majority of adverse events are mild to moderate and manageable/reversible<sup>1</sup>
- Few discontinuations due to adverse events (12.3% in combo due to ≥ 1 TEAE 4.9% due to elevated blood CPK\*)
   \* Ne association to data with elipically.

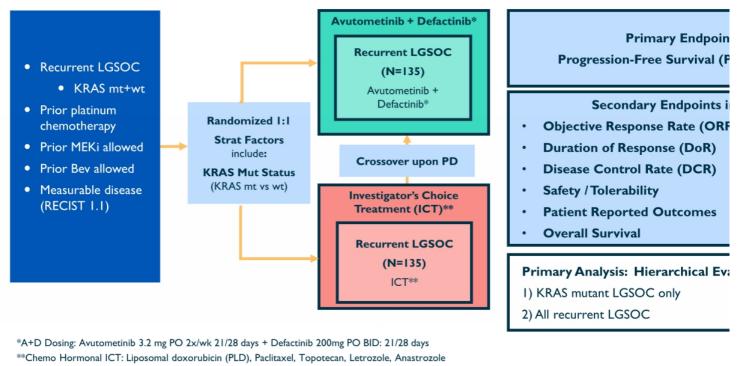
\* No association to date with clinically significant toxicities, including rhabdomyolysis

Avutometinib + Defactinib (n=81			
	Any Grade	G	
Nausea, n (%)	50 (61.7)		
Diarrhea, n (%)	40 (49.4)		
Blood CPK increased, n (%)	39 (48.1)		
Oedema peripheral, n (%)	34 (42.0)		
Vomiting, n (%)	30 (37.0)		
Vision blurred, n (%)	29 (35.8)		
Dermatitis acneiform, n (%)	28 (34.6)		
Fatigue, n (%)	27 (33.3)		
Rash, n (%)	25 (30.9)		
Dry skin, n (%)	18 (22.2)		
Anemia, n (%)	14 (17.3)		

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Banerjee et al., ASCO June 2023; <sup>1</sup> J Clin Oncol 41, 2023 (suppl 16; abstr 5515)

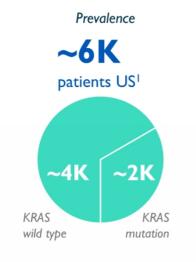
# RAMP-301: Avutometinib + Defactinib Phase 3 Confirmatory Trial – Randomized Controlled Trial (RCT)





BICR: Blinded Independent Central Review

## RAMP 201 Part A Interim Data Support Meaningful Market Potential f Recurrent LGSOC Regardless of KRAS Status with Long Duration of



~80K patients WW<sup>1</sup>

-	50,000	100,000	150
NSCLC KRAS G12C <sup>3</sup>			
Pancreatic Cancer <sup>3</sup>			
LGSOC <sup>3</sup>			
Endometrioid Cancer <sup>3</sup>			
Metastatic uveal melanoma <sup>3</sup>	I		

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<sup>1</sup> Monk, Randall, Grisham, The Evolving Landscape of Chemotherapy in Newly Diagnosed Advanced Epithelial Ovarian Cancer, Am Soc Clin Oncol Educ Boc <sup>1</sup> Monk, Kandali, Grisham, The Evolving Landscape of Chemotherapy in Newly Diagnosed Advanced Epithelial Ovarian Cancer, Am Soc Clin Oncol Educ Boc Slomovitz, Gourley, Carey, Malpica, Shih, Huntsman, Fader,, Grisham et al, Low-Grade serous ovarian cancer: Squecol Oncol; 2020. Gi Low-Grade Serous Ovarian Cancer: Current Treatment Paradigms and Future Directions; Curr Treat Options Oncology; 2018; Globocan 2020
 <sup>2</sup> Patient-months of Therapy metric calculated by multiplying relevant incidence/prevalence rate times estimated duration of therapy; represents US market c patient population estimates from Globocan 2020, American Cancer Society 2021, AACR Genie Cohort V9.0 public, and scientific publications. Duration of estimates from clinical studies and clinician experience. Patient-months on therapy is for 2<sup>nd</sup>-line+ patients
 <sup>3</sup> NSCLC KRAS G12C 2<sup>nd</sup> line patients (incidence); Pancreatic RAS/RAF mutant 2<sup>nd</sup>-line patients (incidence); Useal melanoma RAS/RAF mutant 2<sup>nd</sup>-line patients (incidence)

# Plan to File for Accelerated Approval with Mature RAMP 201 and FRAME Study Results

#### **Recent Achievements/Milestones**

- Encouraging efficacy results include independently confirmed responses (FRAME study)
- RAMP 201 Part A data at ASCO 2023 demonstrated ORR of 45% (13/29) and tumor shrinkage in 86% (25/29) of evaluable patients
- No new safety signals; few discontinuations due to adverse events
- Initiated RAMP 301, a Phase 3 confirmatory trial
- High unmet need in rare ovarian cancer with no currently FDA approved therapies specifically for recurrent LGSOC
- Received FDA Breakthrough Therapy Designation and Orphan Drug Designation for avutometinib in combination with defactinib in LGSOC



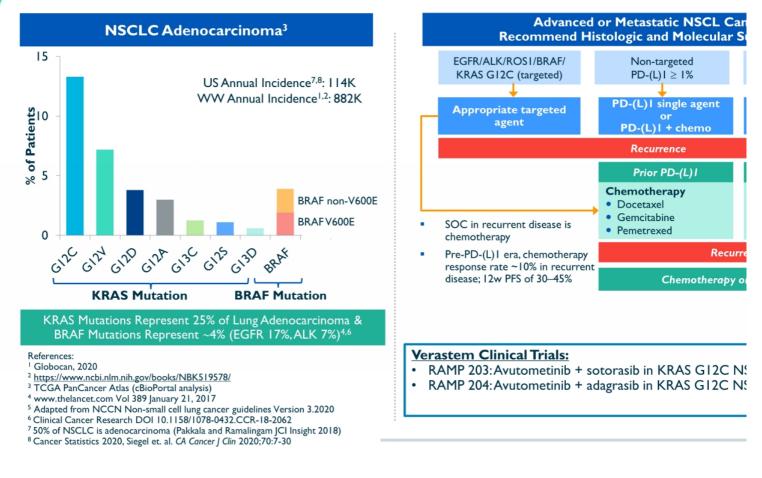
#### **Next Milestones**

- Plan to file for accelerated approval ba totality of the data from the RAMP 20 studies
- Report updated topline data from RAI H1 2024
- Continue site activation (sites current and Australia) and enrollment of RAM confirmatory study



# Avutometinib with KRAS G12C Inhibitors in Non-Small Cell Lung Cancer

### High Unmet Need in Refractory NSCLC Adenocarcinoma

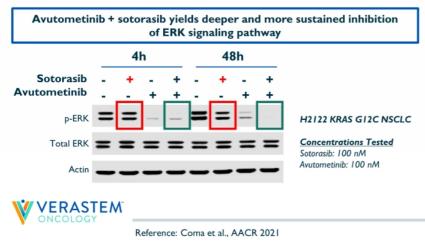


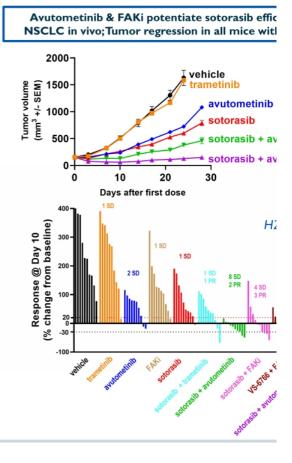
### Preclinical Synergy of Avutometinib + GI2C Inhibitors in KRAS GI2

Synergy of avutometinib + G12C inhibitors across G12C mutant NSCLC, CRC & Pancreatic cancer cell lines

		Combined S	Synergy Score
Indication	Sensitivity to G12C inhibitors	Avutometinib + sotorasib	Avutometinib + adagrasib
NSCLC	Moderately sensitive	44.7	44.6
NSCLC	Sensitive	10.0	3.4
NSCLC	Insensitive	8.6	12.0
NSCLC	Sensitive	6.9	5.4
NSCLC	Moderately sensitive	5.1	ND
CRC	Sensitive	16.1	18.5
Panc	Sensitive	2.3	5.3
	NSCLC NSCLC NSCLC NSCLC NSCLC CRC	Indication         G12C inhibitors           NSCLC         Moderately sensitive           NSCLC         Sensitive           NSCLC         Insensitive           NSCLC         Sensitive           NSCLC         Moderately sensitive           NSCLC         Sensitive           NSCLC         Sensitive           CRC         Sensitive	Sensitivity to G12C inhibitors         Avutometinib + sotorasib           NSCLC         Moderately sensitive         44.7           NSCLC         Sensitive         10.0           NSCLC         Insensitive         8.6           NSCLC         Sensitive         6.9           NSCLC         Moderately sensitive         5.1           CRC         Sensitive         16.1

ND: not determined





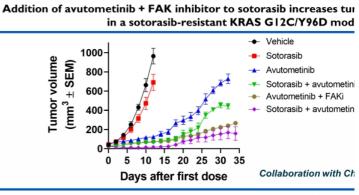
# Avutometinib $\pm$ FAKi Restores Anti-Tumor Efficacy of Sotorasib in G12Ci-Resistant KRAS G12C Models

Avutometinib is effective against acquired KRAS mutations that occur clinically upon progression on G12C inhibitors

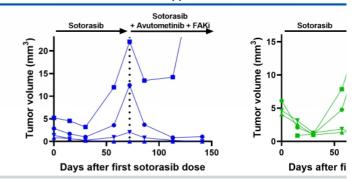
	IC50 (nM)		
Cell Line	Sotorasib	Adagrasib	Avutometinib
G12C	29	3	14
G12D	435	382	7
G12C/R68S	157	85	13
G12C/H95D	11	235	10
G12C/Y96C	438	216	4
G12C/Y96D	>5000	578	17

<30 nM 30 - 150 nM >150 nM

Collaboration with Andy Aguirre, DFCI



Addition of avutometinib + FAKi restores anti-tumor activity af sotorasib monotherapy in a KRAS GI2C NSCLC GEM

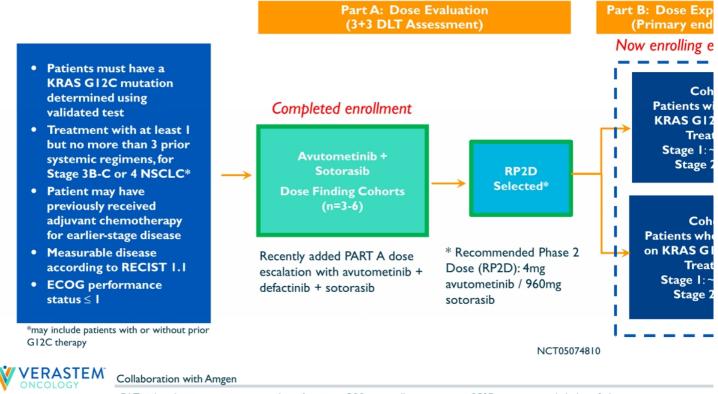


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Reference: Coma et al., AACR RAS meeting 2023

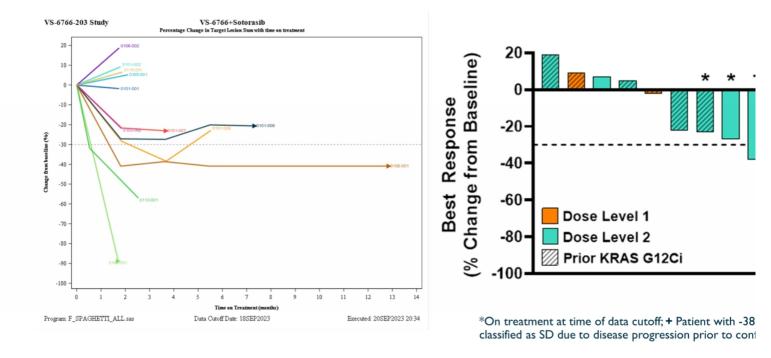
Collaboration with Mariano Barbacid, CNIO (S

## RAMP 203: Phase 1/2 Trial of Avutometinib + LUMAKRAS<sup>TM</sup> (Soto Defactinib in KRAS G12C Advanced NSCLC



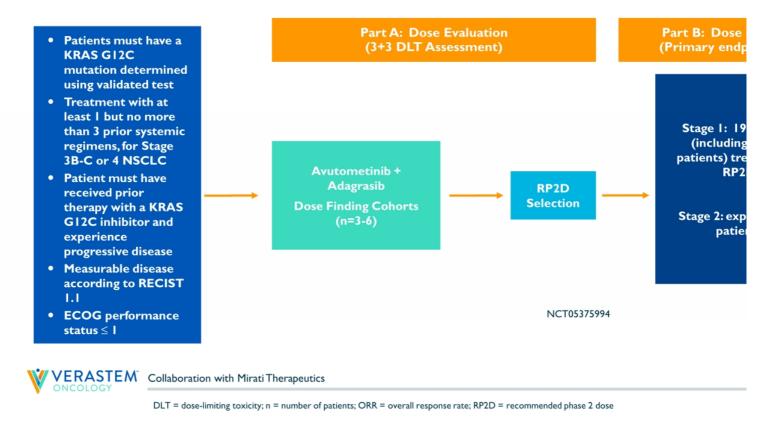
DLT = dose-limiting toxicity; n = number of patients; ORR = overall response rate; RP2D = recommended phase 2 dose

### RAMP 203: Objective Responses in KRAS GI2C NSCLC Sotorasib + Avutometinib Combination



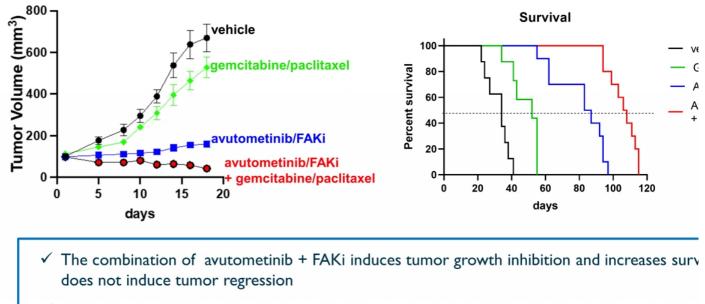
Reference: Awad et al., EORTC- NCI – AACR Conference Oct 2023

# RAMP 204: Phase 1/2 Trial of Avutometinib + KRAZATI<sup>™</sup> (Adagras KRAS G12C Advanced NSCLC



Avutometinib Combinations in Pancreatic Cancer and Colorectal Cancer

### Addition of Avutometinib + FAKi to Chemotherapy Induces Tumor and Increases Survival in a KRAS/p53 Pancreatic Cancer Mouse Mo

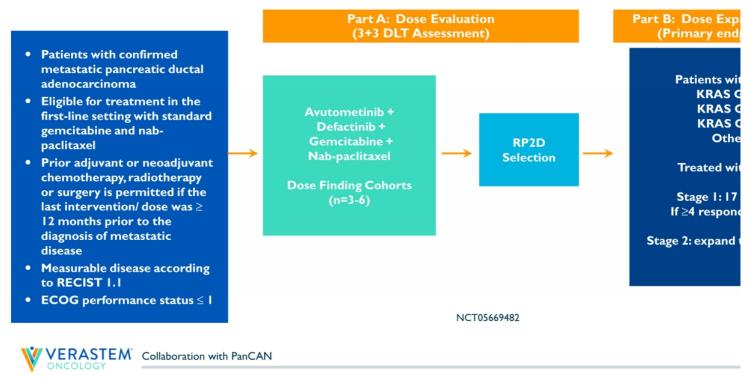


✓ Addition of chemo (gemcitabine + paclitaxel) to avutometinib/FAKi induces tumor regression



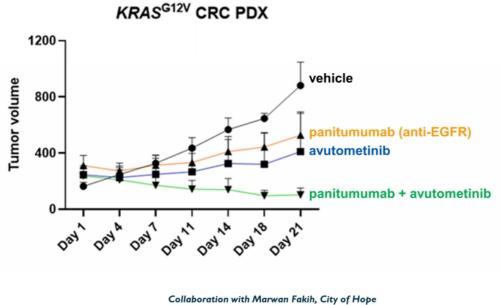
Collaboration with David DeNardo, Washington University; unpubl

# RAMP 205: Phase 1/2 Trial of Avutometinib/Defactinib + GEMZAR<sup>™</sup> (Gemcitabine)/ABRAXANE<sup>™</sup> (Nab-paclitaxel) in Front Line Metast Pancreatic Cancer



DLT = dose-limiting toxicity; n = number of patients; ORR = overall response rate; RP2D = recommended phase 2 dose

### Combination of Avutometinib with anti-EGFR mAb Induces Tumor Regression in a KRAS mt Colorectal PDX Model



- Avutometinib + anti-EGFR (pa induces tumor regression in a CRC patient-derived xenograf
- GI2Ci + anti-EGFR (sotorasib panitumumab and adagrasib + have shown partial responses i CRC (Fakih et al. ESMO 2021; ESMO 2021)
- These data support the on clinical evaluation of avuto cetuximab (anti-EGFR) for of KRAS mt CRC (NCT05





Pachter, RAS Development Summit, 2021

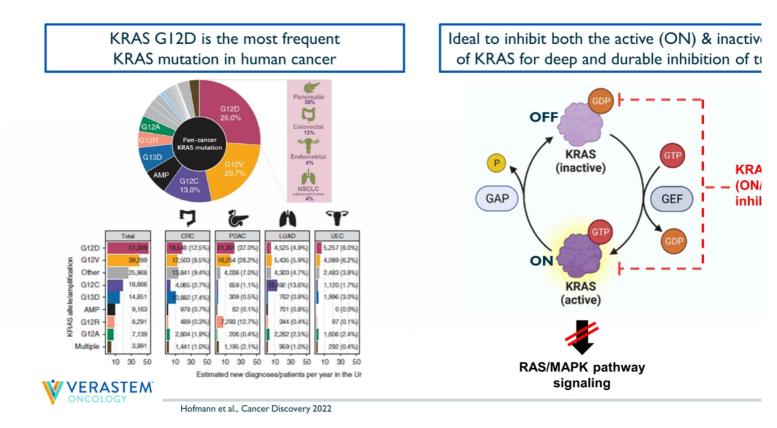
### **Discovery Efforts**

### Discovery and Development Collaboration with GenFleet Strength Pipeline Targeting RAS Pathway-Driven Cancers

- Increases the breadth of Verastem's oncology pipeline with strategically-aligned RAS pathway focus
  - Exclusive options for Verastem to exclusively license up to 3 programs with development and commercializati
    of the GenFleet markets of mainland China, Hong Kong, Macau, and Taiwan
  - $\circ\;$  Potential development in combination with Verastem's current pipeline
  - Selected GFH375 (VS-7375), an oral KRAS G12D (ON/OFF) inhibitor as lead program; programs 2 & 3 in dis
  - $_{\odot}\,$  Small molecule programs focused on anti-cancer targets related to the RAS/MAPK pathway or surrounding c
- Strategic collaboration builds on Verastem Oncology and GenFleet's experience in RAS pathway-driven cancers
  - $\circ~$  Collective worldwide strengths in RAS pathway discovery and development
  - $\circ~$  Established network of collaborators, including leading scientific and clinical experts
  - Leverages experience from GenFleet's KRAS GI2C inhibitor program and Verastem's avutometinib/defactinib
- Risk-sharing structure of the collaboration with milestone-based options provides capital efficient approach
  - At execution, Verastem paid GenFleet an upfront payment for options to obtain exclusive right to 3 programs program basis
  - Combined with the upfront amount, payments for future annual R&D support, development milestones and o first program through completion of Phase I trial could equal up to \$11.5 million
  - o Potential total deal size across all 3 programs up to \$625.5 million excluding royalties if Verastem exercises its
  - Includes exclusive rights for Verastem to obtain a license to each of the compounds after successful completic determined milestones in Phase I trials



### Rationale for Designing a Potent and Selective Orally Bioavailable Inhit KRAS G12D (ON/OFF) for the Treatment of Patients with KRAS G12



### GFH375 (VS-7375) is an Oral KRAS G12D (ON/OFF) Inhibitor

First program from the GenFleet collaboration

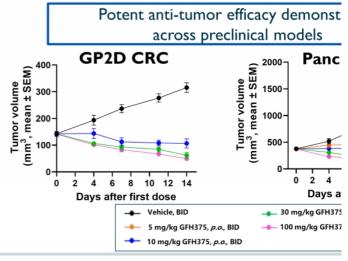
- GFH375 (VS-7375) is a potent and selective orally bioavailable inhibitor of KRAS G12D (ON/OFF) with potent anti-tumor efficacy demonstrated across preclinical models
- Dual inhibitor of ON (GTP) and OFF (GDP) states of KRAS G12D
- · Orally bioavailable across preclinical species
- Potent against intracranial tumor models suggesting potential to treat brain metastases
- Avutometinib enhances anti-tumor efficacy of GFH375 (VS-7375) in preclinical models
- IND-enabling GLP toxicology studies complete
- IND application filed in China and accepted for review; upon clearance expect to initiate Phase I trial in China in H2 2024



Zhou et al., AACR 2024

states of KRAS G12D		
KRAS GI2D State	GFH375 IC (KRAS G12D	
GppNp-bound (ON/active)	2 ± 1	
GDP-bound (OFF/inactive)	6 ± 1	

Dual inhibitor of ON (GTP) and O



### Financials

### **Key Financial Statistics**

### As of and for the quarter ended December 31, 2023

Cash, cash equivalents & investments	\$137.IM
GAAP Operating Expenses	\$31.IM
Non-GAAP Operating Expenses*	\$29.5M
Shares Outstanding	25.3M**

### Sources of Non-Dilutive Capital

#### **Oxford Finance LLC Credit Facility**

- Up to \$150M available in a series of term loans •
  - \$40M term loans outstanding •
  - Remaining \$110M available upon achievement of pre-defined milestones or at lender's discretion
- Floating interest rate, subject to a floor and a cap; 5% final payment charge, and loan subject to 1-3% early pa
- Interest only payments through April 2025
- No financial covenants

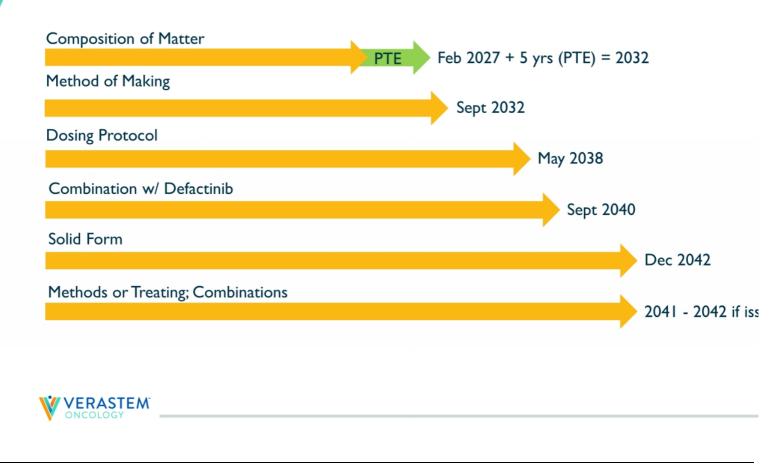
#### Secura Bio, Inc. (Secura) Asset Purchase Agreement – COPIKTRA

- Regulatory and commercial milestone payments up to \$95M
- Entitled to receive 50% of royalties, milestones, and sublicensee revenue payments made to Secura related tc •
- Low double-digit royalties on annual net sales over \$100M in US, EU, and UK •



VERASTEM \* Q4 2023 GAAP operating expenses - \$31.14M less Q4 2023 stock compensation of \$1.60M = \$29.54M Q4 2023 non-GAAP operating expenses \*\*Excludes Series A Preferred (0.8M Shares on as-converted basis), Series B Preferred (4.2M Shares on as-converted basis), and outstanding un warrants (1.5M Shares).

### Avutometinib Patent Exclusivity



### Experienced Senior Management Team



#### **Daniel Paterson** President and Chief Executive Officer

- CEO The DNA Repair Co. (now On-Q-ity)
- PharMetrics (now IMS), Axion



#### Mike Crowther Chief Commercial and Business Strategy Officer

- CBO, Minerva Biotechnologies
- Interim US lead and VP of US
- Marketing, Kite Pharma
- Celgene



**Dan Calkins** Chief Financial Officer

- Technical Accounting Consultant-CFGI
- PwC LLP



- Head of Cancer Biology OSI (now Astellas)
- Schering-Plough



Cathy C Chief Organ Effectiveness

Principal –
Ironwood,/ Tufts Healt

Hagop Y MSc, M.I Head of Me • CMO, BINI

Progenics,CMO & EV

 CMO & EV SVP, Imclon



