

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): September 8, 2025

Verastem, Inc.

(Exact Name of Registrant as Specified in Charter)

Delaware
(State or Other Jurisdiction
of Incorporation)

001-35403
(Commission
File Number)

27-3269467
(IRS Employer
Identification No.)

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

02494
(Zip Code)

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:

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

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

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.

Item 7.01 Regulation FD Disclosure

On September 8, 2025, Verastem, Inc. posted its updated 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
99.1 104	Corporate Presentation dated September 8, 2025 Cover Page Interactive Data File (embedded within the Inline XBRL document)

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.

Dated: September 8, 2025

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

Delivering Novel Therapies for RAS/MAPK Pathway Driven Cancers

Corporate Presentation | September 2025



Commercial Product and Pipeline Positioned to Deliver Long-Term Shareholder Value

- **Strong early execution of commercial launch**
 - May 2025 approval of AVMAPKI™ FAKZYNJA™ CO-PACK, six weeks ahead of PDUFA action date
 - First-ever FDA-approved treatment specifically for KRAS-mutated recurrent low-grade serous ovarian cancer (LGSOC) with listing as Category 2A in NCCN Guidelines
- **Advancing our novel, early-stage pipeline in RAS/MAPK pathway-driven cancers**
 - VS-7375, an oral KRAS G12D (ON/OFF) inhibitor: initiated Phase 1/2a trial in patients with advanced KRAS G12D mutant solid tumors outside of China; encouraging initial safety and efficacy results from partner in China
- **Maximizing the synergistic potential of avutometinib plus defactinib combination in other advanced solid tumors**
 - RAMP 205: front-line metastatic pancreatic cancer; DL1 achieved confirmed ORR of 83% (10/12)
 - RAMP 203: advanced KRAS G12C mutant non-small cell lung cancer; no DLTs were observed in the triplet combination cohort
- **Existing cash, investments, and anticipated future product sales give us a strong financial position and runway into the second half of 2026**



PDUFA: Prescription Drug User Fee Act; KRAS: Kirsten Rat Sarcoma Virus; MAPK: Mitogen-activated Protein Kinases; FDA: Food and Drug Administration; NCCN: National Comprehensive Cancer Network; DL: dose-level; ORR: objective response rate; DLT: dose-limiting toxicity

AVMAPKI FAKZYNJA CO-PACK Commercial Launch is Off to a Str

Achieved net product revenue of
\$2.1 million
In the first six weeks
on the market



AVMAPKI™
FAKZYNJA™ CO-PACK
(avutometinib capsules; defactinib tablets) 0.8 mg/200 mg

FDA Approved on May 8, 2025, Nearly Two Months Ahead of date

RAS/MAPK Pathway Directed MOA

- AVMAPKI offers dual inhibition of RAF and MEK
- FAKZYNJA mediates drug resistance of activated RAF/MEK
- Together, they offer a more complete blockade of the signal growth and drug resistance in the RAS/MAPK pathway

Clinically Meaningful Response Rates and Long Duration of T

- 44% ORR, 3.3 to 31.1 months mDOR

Manageable Safety Allows for Treatment Until Progression fo

Convenient, Two Orally Dosed Treatments

- Novel intermittent dosing schedules



Please see the full [Prescribing Information](#) for more information

MOA: Mechanism of Action; RAF: Rapidly Accelerated Fibrosarcoma; MEK, Mitogen-Activated Extracellular Signal-regulated Kinase; mDOR: median Duration of Response

Our Pipeline: Addressing RAS/MAPK-Driven Cancers

Asset	Trial Name/ Therapeutic Area	Regimen	Phase 1	Phase 2	Phase 3	FDA Approved	Ant
Avutometinib + Defactinib	RAMP 201, recurrent LGSOC	RAF/MEK Clamp + FAKi				 <small>AVMAPKI™ FAKZYNJA™ CO-PACK defactinib 100mg/100mg + avutometinib 400mg/200mg</small>	
Avutometinib + Defactinib	RAMP 301, recurrent LGSOC	RAF/MEK Clamp + FAKi vs ICT					Complete RAMP 301 Report ID 2025
VS-7375*	VS-7375-101, Advanced solid tumors	KRAS G12D (ON/OFF) inhibitor					Expect to update or monother Q4 2025
Avutometinib + Defactinib	RAMP 205, 1L mPDAC	RAF/MEK Clamp + FAKi + gemcitabine, nab- paclitaxel <i>PanCAN Collaboration</i>					Expect to the expan
Avutometinib ± Defactinib	RAMP 203, advanced KRAS G12C NSCLC	RAF/MEK Clamp ± FAKi + KRAS G12Ci (sotorasib) <i>Amgen Collaboration</i>					Report an 2025

Not shown:

*GenFleet Therapeutics has an ongoing Phase 1/2 clinical trial in China with VS-7375, known as GFH375 in China. GenFleet retains greater China right. Verastem has two undisclosed assets at discovery phase targeting the RAS/MAPK-pathway as part of the GenFleet collaboration.



FAKi: focal adhesion kinase inhibitor; ICT: investigator choice of treatment; NSCLC: non-small cell lung cancer; mPDAC: metastatic Pancreatic Ductal Adenocarcinoma;

Delivering on the Key R&D Milestones Set for 2025

Maximize the synergistic potential of avutometinib + defactinib in other advanced solid

RAMP 205: Avutometinib + Defactinib + SOC Chemotherapy in 1L mPDAC

- ✓ Reported updated data from the ongoing RAMP 205 trial selected dose level 1 cohort for recommended phase 2 cohort
- ✓ Close to completing enrollment in the expansion cohort patients at the RP2D

RAMP 203: Avutometinib + Sotorasib ± Defactinib in KRAS G12C in advanced NSCLC

- ✓ Completed enrollment to the KRAS G12C inhibitor, prior stage 1 part B doublet cohort
- ✓ Completed enrollment in the planned dose level evaluation for the triplet combination

Advance novel, early-stage pipeline

VS-7375, Oral KRAS G12D (ON/OFF) Inhibitor

- ✓ U.S. IND filed and cleared
- ✓ Dosed first patient in Phase 1/2a trial in U.S.
- ✓ GenFleet reported updated Phase 1 clinical data from C
- ✓ Granted FDA Fast Track Designation in 1L and 2L mPDA



R&D: Research and Development. IND: Investigational New Drug

Achieved All Key Milestones in 1H25, On Track To Deliver in 2H25

PRODUCT LAUNCH

- Effectively **reach HCPs**
- **Engage** with **patients**
- Ensure **seamless access**

VS-7375-101

Expect to report preliminary update on Phase 1 monotherapy dose escalation in Q4 2025. Initiate various dose escalation combination cohorts in Q4 2025.

RAMP 301

Complete plan enrollment in the confirmatory study end of 2025. Report recommendation in

RAMP 205

Complete enrollment in the expansion cohort in Q3 2025.

RAMP 203

Report an interim data update on both doublet and triplet combinations in Q4 2025.

RAMP 201J

Report initial data **Phase 2** clinical trial conducted in Japan in



IDMC: Independent Data Monitoring Committee; HCPs: health care professionals



AVMAPKI™ FAKZYNJA™ CO-PACK

(avutometinib capsules; defactinib tablets) 0.8 mg; 200 mg

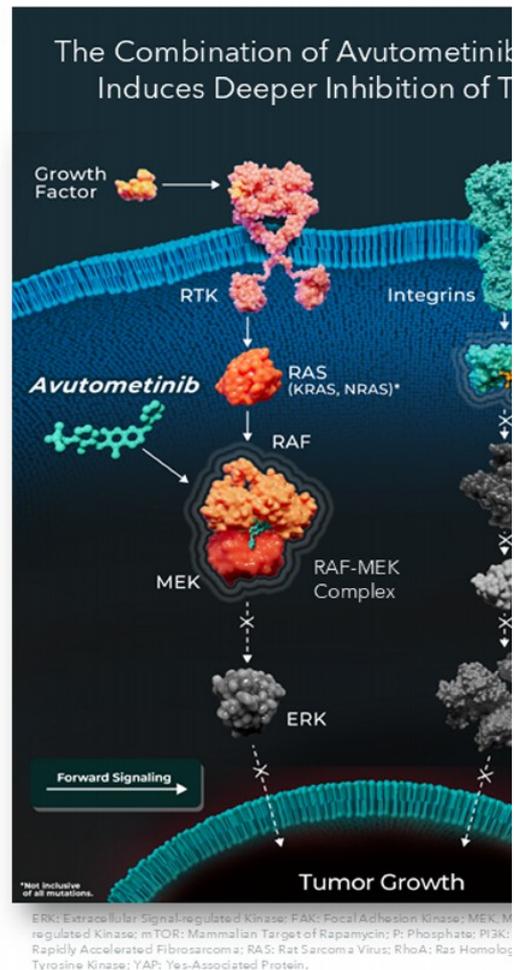
**Commercially launched in the U.S. for
KRAS-mutated Recurrent LGSOC**

**FDA Approval Date:
May 8, 2025**



Providing More Complete Blockade of the Signaling that Drives Growth and Resistance of RAS/MAPK Pathway-Dependent Tumors

- 70% of LGSOC tumors are driven by the RAS/MAPK pathway and about 30% of these have a KRAS mutation^{1,2,3,4}
- Avutometinib inhibits MEK kinase activity while blocking the compensatory reactivation of MEK by upstream RAF^{5,6,7}
- Blocking RAF and/or MEK activates FAK, a key mediator of drug resistance^{8,9}
- Defactinib, a FAK inhibitor, inhibits parallel pathway signaling^{10,11,12}
- Together, avutometinib plus defactinib offer more complete blockade of the signaling that drives the growth of RAS/MAPK pathway-dependent tumors



1. AACR Genie v16, 1:2. Chesley et al., J Pathol 2021; 2. Thomson et al., Gynecol Oncol 2023; 3. Gershenson et al., Gynecol Oncol 2022; 4. Coma et al., AACR 2022; 5. Ishii et al., Cancer Res, 2013; 6. Lito et al., Cancer Cell, 2014; 7. Lubrano et al., AACR 2024; 8. Banerjee et al., AACR 2020; 9. Jones et al., Invest New Drugs 2015; 10. McNamara et al., Gynecol Oncol 2024; 11. Banerjee et al., ASCO 2023; 12. Banerjee et al., AACR 2022

High Unmet Need for an Effective and Tolerable Therapy in Recurrent LGSOC

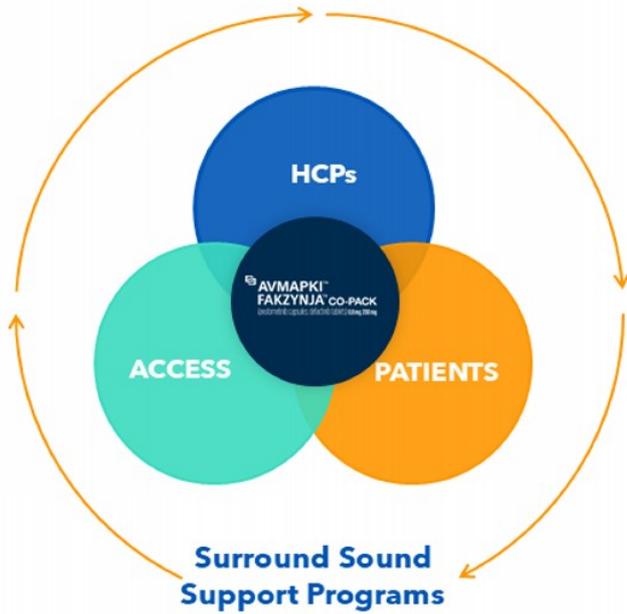
- **U.S. incidence / prevalence:** 1k-2k¹ / 6k-8k²
- **LGSOC affects younger women** with bimodal peaks of diagnosis at ages between 20-30 and 50-60; disproportionately **impacts health, fertility, and long-term quality of life**^{3,4}
- **80-90% of patients will experience a recurrence**⁵
- **Current standard of care offers low to moderate response rates** (6-13%)^{6,7,8}
- **Median overall survival (OS) of ~10 years** from time of diagnosis⁹: KRAS-mutated - ~12 years¹⁰ and KRAS wild-type: ~7 years¹⁰

1. Verastem DOF; 2. US Cancer Statistics. Accessed 2024; 3. Slomovitz *Gynecol Oncol* 2020; 4. Manning-Geist B et al. *Clin Cancer Res* 2022;28(20):4456-4465; 5. Babaier 2022/p1/para1/ln6,7; 6. Gershenson *Gynecol Oncol* 2022; 7. Slomovitz *Gynecol Oncol* 2020; 8. Monk 2020/p3758/table2/footnote-b; 9. Banerjee SN. *J Clin Oncol*. 41. No 16_suppl (June 1, 2023) 5515-5515; 10. Manning-Geist B et al. *Clin Cancer Res* 2022;28(20):4456-4465; Calculated using figures in Gershenson *Gynecol Oncol* 2022.

A portrait of a woman with long, wavy brown hair, wearing a black top with a white pattern. She is looking directly at the camera with a slight smile.

“When you get told that you have a recurrence, the mental load is a lot. You think, ‘What did I have to do for treatment?’ Now I have to repeat that. And I wonder, ‘Is there something available for me to prevent a second or a third recurrence?’”

Laser-Focused on Three Strategic Imperatives to Drive a Successful Commercial Launch



Effectively reach all healthcare providers	Top 100 commercial health organizations contribute to patient claims ¹
Engage and support patients	Patients likely to have prior experience through other therapies, and need to be ready for a new treatment
Ensure seamless access	Support the patient to ensure there are no barriers to reimbursement and access



1VSTM DOF - Claims LGSOC Proxy

Encouraging Launch Momentum Reinforces Strategic Imperatives Success

Achieved net product revenue of

\$2.1 million

in the first six weeks
on the market

First patient received product:

May 22, 2025

- ✓ First-ever approved treatment for people living with KRAS-mut recurrent LGSOC nearly two months in advance of PDUFA date
- ✓ Urgent, unmet patient need and significant market opportunity
- ✓ Experienced and energized field
- ✓ Two years of launch preparedness
- ✓ High physician enthusiasm and engagement
- ✓ Effective digital surround sound website traffic
- ✓ Extensive payer education



Early Launch Performance Tracking as Expected



- ✓ Prescriptions coming from a mix of academic and community physicians
- ✓ Seeing both repeat prescriptions and refills
- ✓ Payer mix is a combination of commercial and Medicare

Effectively reach all healthcare providers

- ✓ Field team engaged with 93% of top 100 organizations
- ✓ Medical Science Liaisons held 100+ meetings and hosted >30 educational forums
- ✓ Branded healthcare professional website is seeing high traffic and downloads of enrollment forms

Engage and support patients

- ✓ Branded patient website is seeing high traffic with significant patient brochure downloads
- ✓ Prelaunch disease education website netted engagement with ~2,500 potential patients

Ensure seamless access

- ✓ NCCN listing as a Category 1 recommendation for appropriate indication
- ✓ Payer coverage has been fast
- ✓ Verastem Cares® Program supporting patient needs



NCCN Submission for Treatment Guideline Inclusion Under Review Entire Population Enrolled in RAMP 201 Study

	NCCN Category 1	NCCN Category 2a	NCCN Category 2b
General % Commercial Payer Coverage			
Examples of Clinical Data in LGSOC and Current NCCN Guideline Category	No category 1 recommendation	<p>Current Listing: Avutometinib + Defactinib Combination Therapy</p> <ul style="list-style-type: none"> • KRAS mt recurrent LGSOC <p>Hormonal therapy (e.g., Anastrozole, Letrozole) & chemotherapy</p> <ul style="list-style-type: none"> • 6-13% ORR⁴ • 17-30% discontinuation rate due to AEs⁴ <p>Trametinib (2-4% US utilization rate¹)</p> <ul style="list-style-type: none"> • 26% ORR by INV assessment, no BICR⁵ • 36% discontinuation rate due to AEs⁵ 	<p>Binimetinib</p> <ul style="list-style-type: none"> • Study stopped early due to futility • 16% ORR by BICR • 31% discontinuation rate due to AEs • Supported by MILO study³

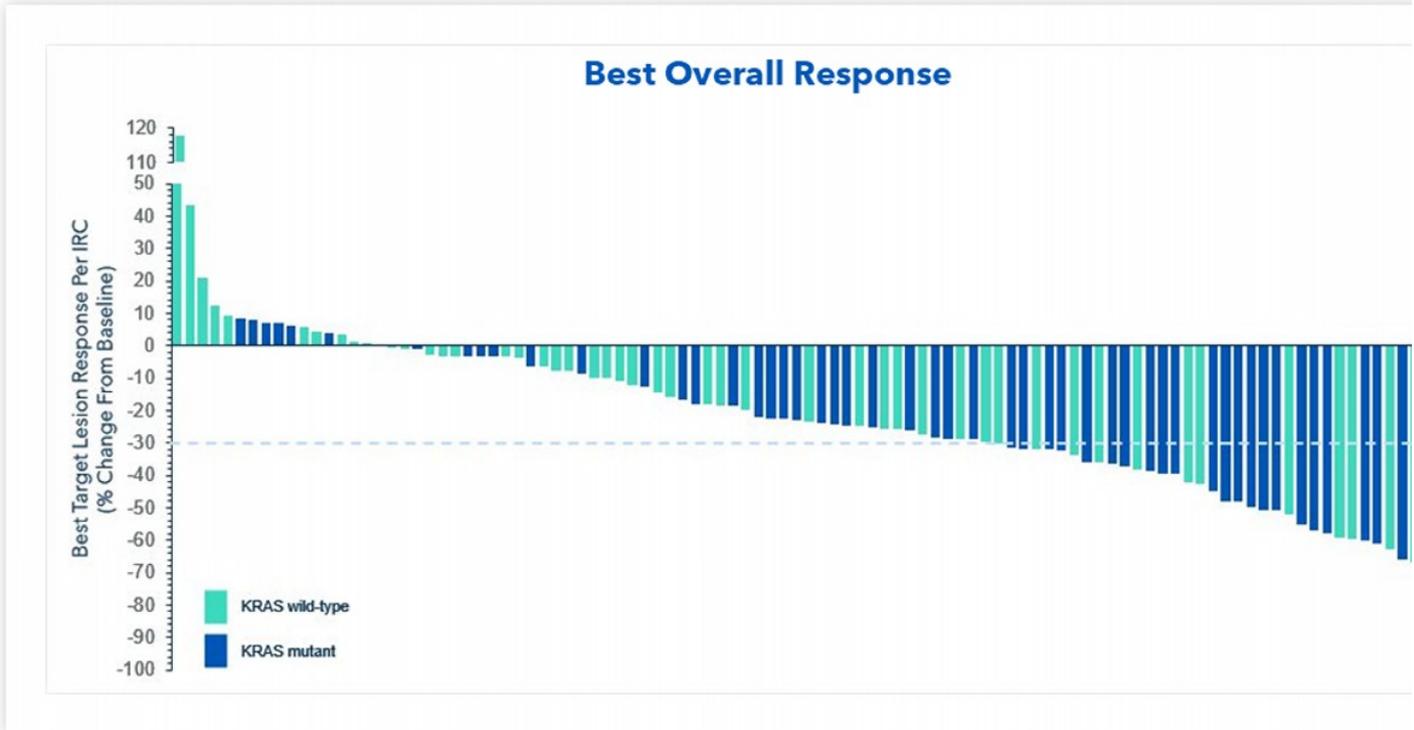


General source: NCCN; McGivney Global Advisory research and analysis; L.E.K. research and analysis. NCCN categories of preference: Preferred intervention, Other recommended intervention, High-level of evidence generally means large randomized controlled Phase 3 trials; Pie charts represent coverage by all major commercial payers; 1. Data on File; 2. GOG 281 trial Gershenson, Study Monk et al., J Clin Oncol 2020; 4. Supported by GOG 281 and MILO studies^{2,3}; 5. Supported by GOG 281⁴

Continuing Medical Progress in LGSOC



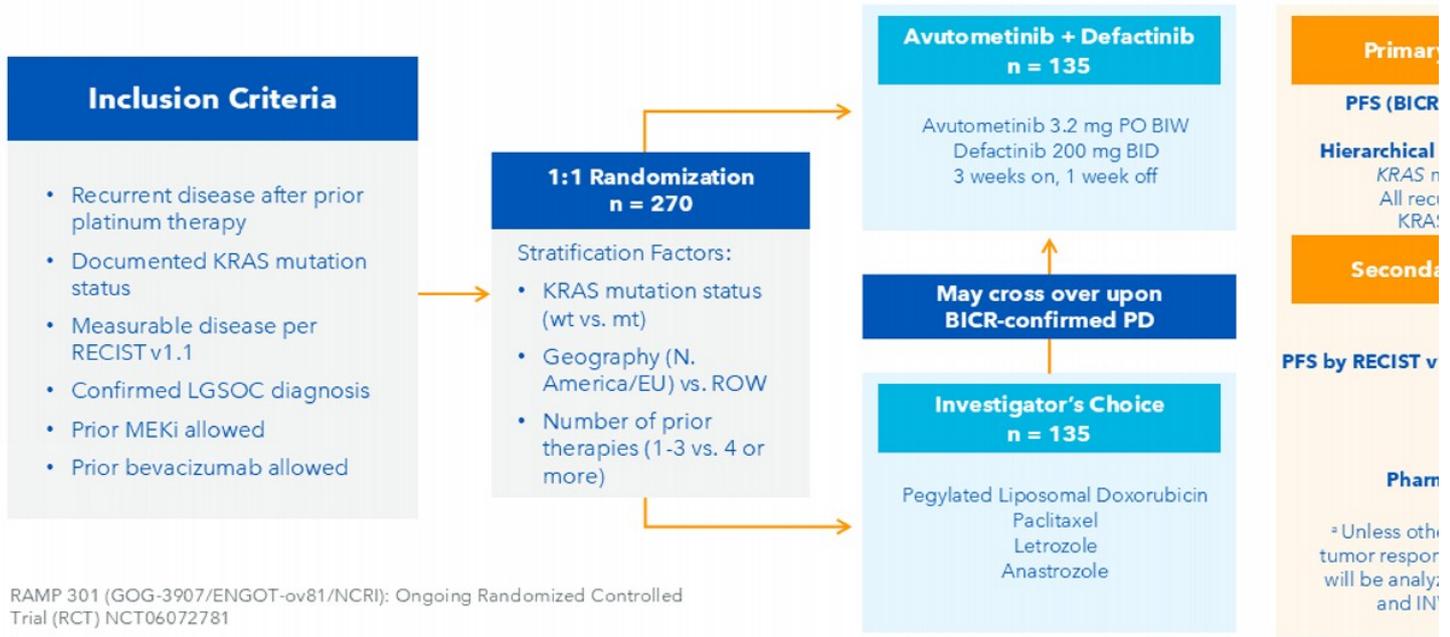
Across Patients With and Without KRAS Mutations in RAMP 201, 8 Reduction in Target Lesions while Receiving Avutometinib and De



Source for all data: RAMP 201 data cut off as of June 30, 2024; Responses for 3 patients (KRAS wild type, n=1; KRAS mutant, n=2) were unknown

RAMP 301: International Phase 3 Confirmatory Trial of Avutometinib + Defactinib in Recurrent LGSOC on Track for Full Enrollment by YE2024

- Entry criteria similar to RAMP 201 patient population, KRAS mt and KRAS wt recurrent LGSOC; prior MEKi and bevacizumab allowed and post at least one line of platinum chemotherapy
- Study sites include the U.S., Canada, UK, Europe, Australia, New Zealand, Japan and South Korea



RAMP 301 (GOG-3907/ENGOT-ov81/NCRI): Ongoing Randomized Controlled Trial (RCT) NCT06072781



*US FDA analysis plan will evaluate PFS independently in KRAS-mt and KRAS wt LGSOC. BICR: blinded independent central review; BID: twice a day; BW: twice a week; DCR: disease control rate; INV: investigator; KRAS: kirsten rat sarcoma virus; MEKi: MEK inhibitor; mt: mutant; PO: per oral; pts, patients; ORR: objective response rate; OS: overall survival; PD: progressive disease; PFS: pre patient-reported outcomes; RECIST: response evaluation criteria in solid tumors; wt: wild type.



Next Steps in LGSOC Clinical Program

Complete planned enrollment in RAMP 301 Phase 3 confirmatory study by end of 2025

Announce outcome of IDMC recommendation for RAMP 301 in Q4 2025

Report initial data from the RAMP 201J Phase 2 clinical trial being conducted in Japan in Q4 2025

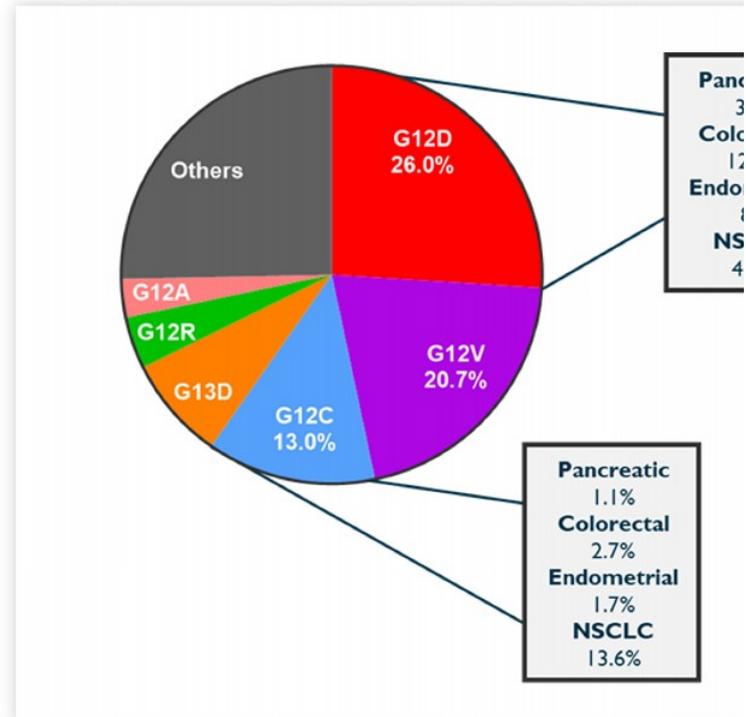
Continue to advance regulatory pathway in Japan and Europe

VS-7375, oral KRAS G12D (ON/OFF) Inhibitor



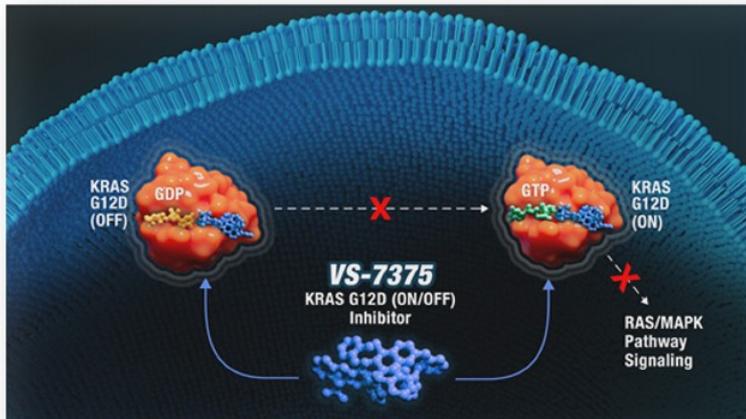
KRAS G12D is the Most Frequent KRAS Mutation in Human Cancer

- The only approved KRAS inhibitors target KRAS G12C, which is largely restricted to NSCLC
- KRAS G12D accounts for 26% of all KRAS mutations
- KRAS G12D mutations are especially prevalent in pancreatic and colorectal cancers
- Targeting KRAS G12D has historically been challenging due to the shallow pocket for drug interaction and lack of a cysteine for covalent binding



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

Non-covalent inhibitor of KRAS G12D (ON/OFF) with potent anti-tumor efficacy across preclin



VS-7375 is a dual inhibitor of ON (GTP OFF (GDP) states of KRAS G12D*

KRAS G12D State	VS-7375 IC ₅₀ (KRAS G12D)
GppNHp-bound (ON/active)	2 ± 1
GDP-bound (OFF/inactive)	6 ± 1

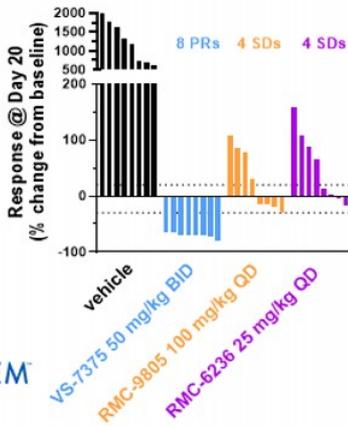
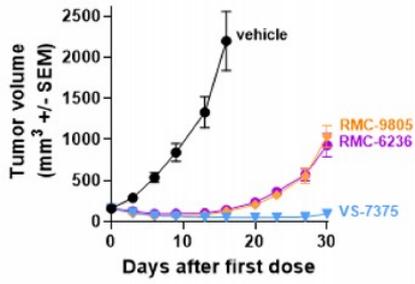
- KRAS-GTP is the active (ON) state, which drives cancer
- KRAS-GDP is the inactive (OFF) state and represents will cycle back to the active ON state
- OFF-state selective agents (e.g., approved G12C inhibitors) have sub-optimal efficacy because they do not target the ON state
- ON-state selective agents (e.g., RMC-6236) can also hydrolyze to the OFF state, which they can no longer target
- May be ideal to have an inhibitor capable of targeting both ON and OFF states of KRAS to maintain inhibition around the clock, aiming for maximum efficacy



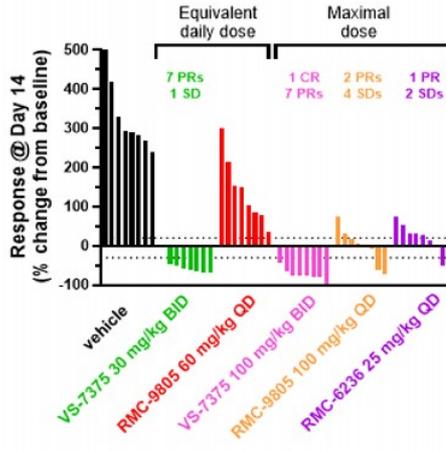
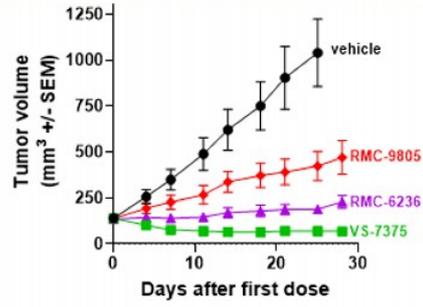
*Piro Lito, RAS Initiative Conference 2024; GEF: Guanine nucleotide exchange factor; GAP: GTPase-activating protein; *Zhou et al., AACR 2024

VS-7375 (G12D ON/OFF inhibitor) is more efficacious than G12D pan-RAS ON inhibitors in KRAS G12D mutant tumor models

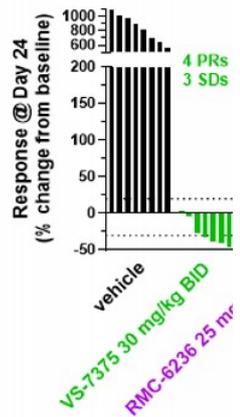
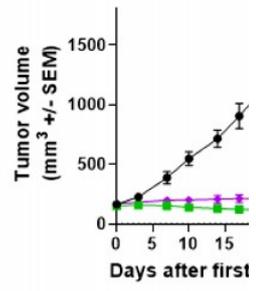
KP4 Pancreatic Cancer Model



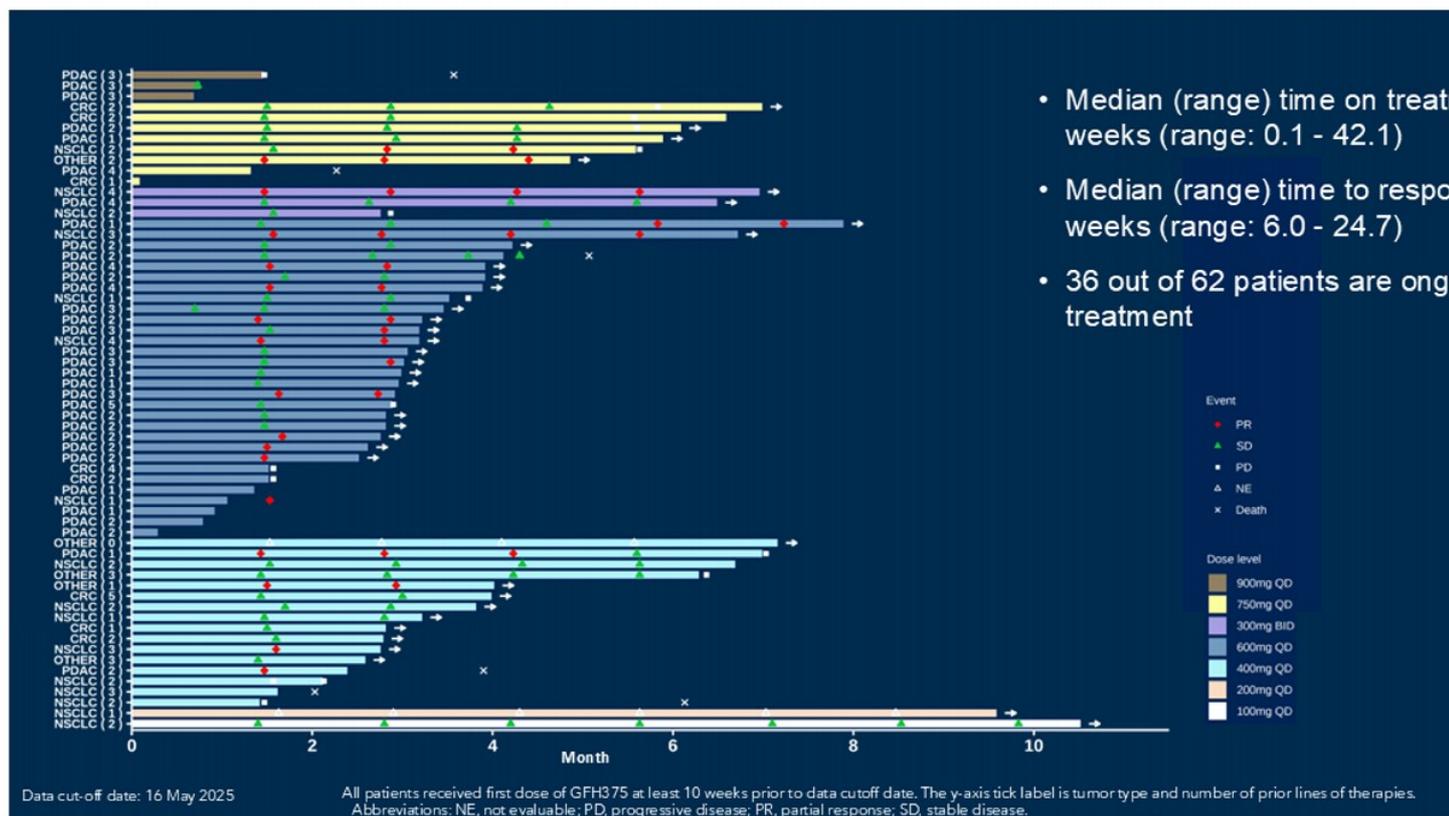
LS513 Colorectal Cancer Model



LU0876 NSCLC



Duration of Treatment and Tumor Assessment of All Patients



Data cut-off date: 16 May 2025

All patients received first dose of GFH375 at least 10 weeks prior to data cutoff date. The y-axis tick label is tumor type and number of prior lines of therapies. Abbreviations: NE, not evaluable; PD, progressive disease; PR, partial response; SD, stable disease.



Source: GenFleet Therapeutics ASCO 2025 Rapid Oral Presentation June 2, 2025; A First-in-human Phase I/II Study of GFH375, a Highly Selective and Potent Oral KRAS G12D Inhibitor in KRAS Advanced Solid Tumors

GFH375: NSCLC Baseline Characteristics & Patient Disposition

- 28 NSCLC patients with local tumor testing KRAS G12D+ were treated with GFH375 single agent, including 16 at 600 mg QD.
- Majority (60.7%) were never smokers.
- Among the 22 patients with baseline PD-L1 TPS data available, none had high expression (TPS \geq 50%).
- 17.9% had baseline brain metastases.
- Majority (64.3%) had received at least 2 prior lines of therapies; all received prior platinum-based chemo therapies; 96.4% received prior ICIs.
- Most patients were ongoing; the longest follow-up was 55.6 weeks.

	All NSCLC (n=28)	NSCLC at 600 mg QD (n=16)
Discontinued n(%)	12 (42.9)	4 (25.0)
Adverse event	1 (3.6)	1 (6.3)
Progress disease	8 (28.6)	2 (12.5)
Patient's decision	2 (7.1)	1 (6.3)
Physician's decision	1 (3.6)	0

	All NSCLC (n=28)
Age, median (range), years	61 (36-74)
Female, n(%)	13 (46.4)
Smoking status, n(%)	
Current or former	11 (39.3)
Never	17 (60.7)
ECOG PS, n(%)	
1	26 (92.9)
Pathology adenocarcinoma, n(%)	28 (100)
Baseline metastasis, n(%)	28 (100)
Bone	12 (42.9)
Brain	5 (17.9)
Liver	3 (10.7)
PD-L1 TPS, n(%)	
Known	22 (78.6)
< 1%	13 (59.1) ¹
1-49%	9 (40.9) ¹
\geq 50%	0
Prior lines of therapies, median (range)*	2 (1-4)
Prior ICI, n(%)	27 (96.4)
Prior platinum, n(%)	28 (100)
Prior ICI +platinum concurrent, n(%)	25 (89.3)

¹Percentage denominator is 22. ²Percentage denominator is 11.

* A neoadjuvant and adjuvant therapy would be counted as a line of system therapy during treatment or within 6 months of last dose.

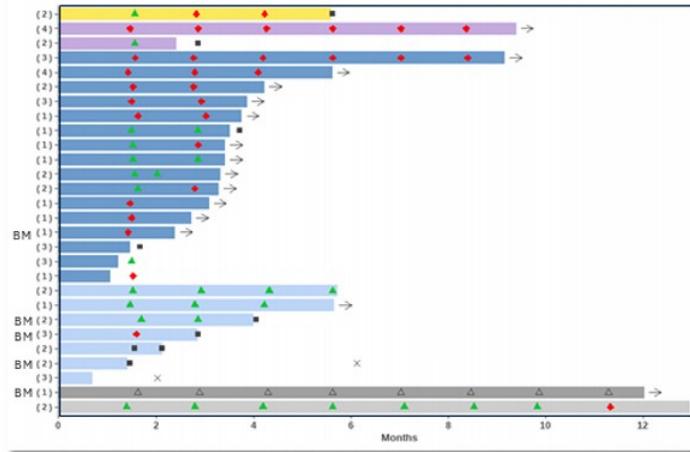
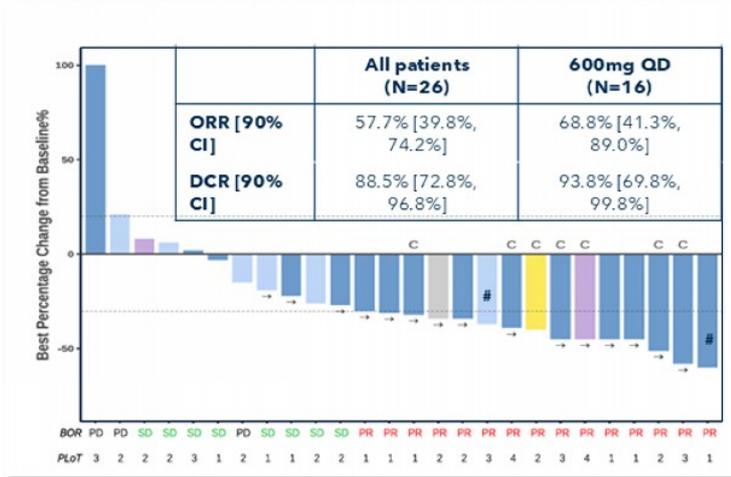


Cut-off date: 01 August 2025.

Source: GenFleet Therapeutics WCLC 2025 Mini Oral Presentation September 7, 2025; Efficacy and Safety of GFH375 in Advanced Non-Small Cell Lung Cancer Patients with KRAS G12D Mutation

GFH375: Efficacy in NSCLC

- ORR is 57.7% in 26 evaluable patients, and 68.8% in 16 patients at 600 mg QD*
- Of the 11 patients at 600mg QD with PR, 5 have confirmed and 5 have the potential to confirm the PR
- 2 additional patients at 600mg QD with 20-30% reduction are still on treatment with potential for response
- Among the 5 patients with baseline brain metastases, 2 achieved PR
- Median (range) time on treatment: 15.1 week
- Median (range) time to response: 6.3 weeks (



Data cut-off date: 01 August 2025.
 All patients received first dose of GFH375 at least 10 weeks prior to data cut-off date. Median follow-up 21.8 weeks (range: 8.3 - 55.6).
 *One patient (200 mg QD) had no target lesion at baseline but remains stable; one patient (400 mg QD) dropped out early without post-baseline tumor assessment due to patient days after last dose due to disease progression.
 # Two patients achieved PR but discontinued before response confirmation. One (400 mg QD) had PD at the 2nd assessment. The other dropped out due to G4 hepatic function at discontinuation.
 BOR, best overall response; CI, confidence interval; DCR, disease control rate; ORR, objective response rate; PD, progressive disease; PR, partial response; SD, stable disease; PLo^t therapies.



Source: GenFleet Therapeutics WCLC 2025 Mini Oral Presentation September 7, 2025; Efficacy and Safety of GFH375 in Advanced Non-Small Cell Lung Cancer Patients with KRAS G12D Mutation

GFH375: Safety in All Patients

- 142 patients, including 28 NSCLC, with advanced KRAS G12D mutation were treated with GFH375 single agent.
 - 93% had baseline ECOG PS = 1.
 - 75% had received at least ≥ 2 prior lines of therapies.
- 97 patients, including 16 NSCLC, were treated at 600 mg QD.
- No treatment related death.
- 4.2% of patients discontinued treatment due to TRAEs.
- 27.5% of patients had grade 3/4 TRAEs.
- Overall, GFH375 has manageable safety profile in previously treated advanced cancer patients.
- Grade ≥ 3 TRAEs and SAEs are more frequent in NSCLC patients; however, discontinuation rate appears to be similar.

Adverse events, n(%)	All dose levels		A
	All types ¹ (N=142)	NSCLC ² (N=28)	
TEAEs			
All grades	141 (99.3)	28 (100)	9
Grade ≥ 3	60 (42.3)	15 (53.6)	4
SAE	27 (19.0)	6 (21.4)	1
Leading to interruption	51 (35.9)	10 (35.7)	3
Leading to reduction	12 (8.5)	5 (17.9)	:
Leading to discontinuation	9 (6.3)	1 (3.6)	:
Leading to death	4 (2.8)	0	:
TRAEs			
All grades	139 (97.9)	28 (100)	9
Grade ≥ 3	39 (27.5)	13 (46.4)	2
SAE	11 (7.7)	4 (14.3)	:
Leading to interruption	33 (23.2)	9 (32.1)	1
Leading to reduction	11 (7.7)	5 (17.9)	:
Leading to discontinuation	6 (4.2)	1 (3.6)	:
Leading to death	0	0	:

Data cut-off date: 17 Jun 2025

The median exposure time to GFH375 in all patients was 9.6 (range: 0.9-49.1) weeks by the cut-off date.

¹ Including all patients treated at all dose levels: 100 mg QD (n=1), 200 mg QD (n=1), 400 mg QD (n=29), 600 mg QD (n=97), 750 mg QD (n=8), 900 mg QD (n=3), and 300 mg BID (n=2)

² Including NSCLC patients treated at all dose levels: 100 mg QD (n=1), 200 mg QD (n=1), 400 mg QD (n=7), 600 mg QD (n=16), 750 mg QD (n=1), and 300 mg BID (n=2)

SAE, serious adverse event; TEAE, treatment emergent adverse event; TRAE, treatment related adverse event.

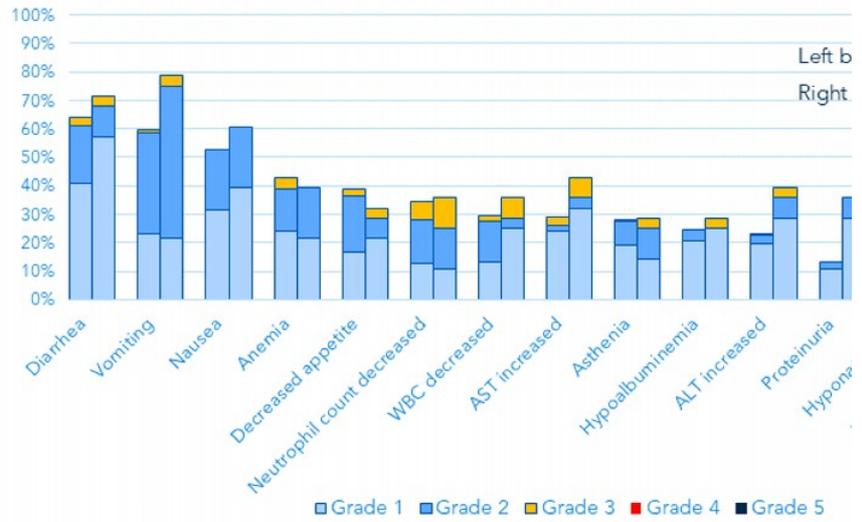
Source: GenFleet Therapeutics WCLC 2025 Mini Oral Presentation September 7, 2025: Efficacy and Safety of GFH375 in Advanced Non-Small Cell Lung Cancer Patients with KRAS G12D Mutation



GFH375: Safety in All Patients

- No new safety signals were observed.
- The most frequent TRAEs were gastrointestinal AEs (diarrhea, vomiting, and nausea) and hematological AEs (anemia and neutropenia), mainly grade 1/2 and manageable with supportive medications.
- Most frequent G3/4 TRAEs were neutropenia (6.3%) and anemia (4.2%).
- Multiple AEs had higher rates in NSCLC patients, potentially due to prior use of ICIs.
 - 96.4% pts with NSCLC had received ICIs before GFH375 treatment.
 - The median time of last dose of ICIs to first administration of GFH375 was 2.8 months (range: 1.0 - 40.0); the median time of ICI treatment was 4.3 months (range: 0.7 – 26.5).

Most frequent TRAEs (≥10% of all patients)

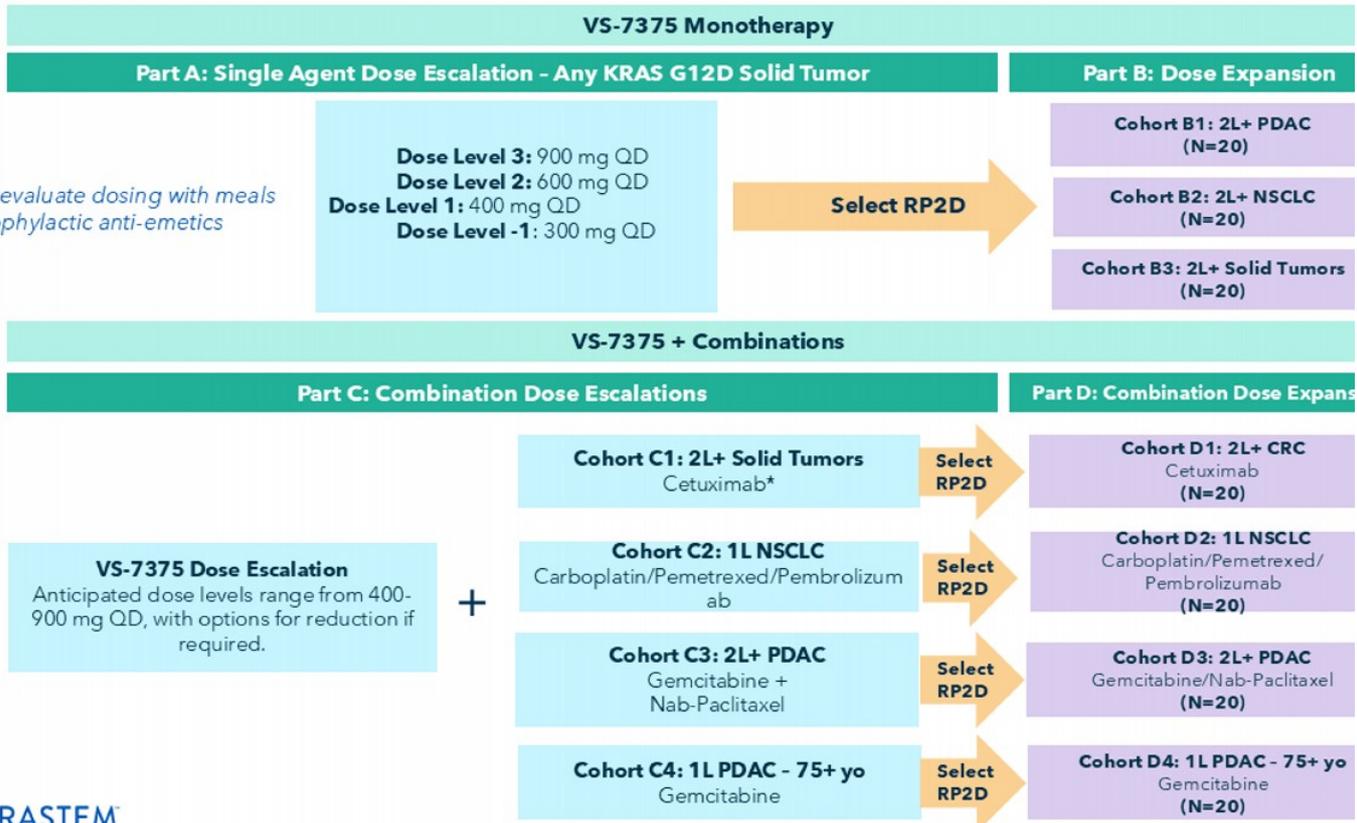


ALT, alanine transaminase; AST, aspartate transferase; TRAE, treatment related adverse event; WBC, white blood cell.

Source: GenFleet Therapeutics WCLC 2025 Mini Oral Presentation September 7, 2025; Efficacy and Safety of GFH375 in Advanced Non-Small Cell Lung Cancer Patients with KRAS G12D Mutation

VS-7375-101 Study Schema, Efficiently Testing KRAS G12D+ Indics with Monotherapy and SOC Combinations

The study will evaluate dosing with meals and utilize prophylactic anti-emetics



*Anticipated starting dose level for cetuximab combination once the monotherapy clears 600 mg dose level

Accelerated Development Approach Builds on Preliminary First-in Study Findings

Multi-Arm Clinical Study evaluating monotherapy and combinations

- **Efficacious starting dose:**
 - 400 mg starting dose supported by monotherapy efficacy/safety in FIH study
- **Evaluating GI side effect mitigations:**
 - Use prophylactic anti-emetics
 - Evaluate dosing with and without food
 - Evaluate additional formulation
- **Multiple, high-value indication strategy:**
 - Monotherapy expansion cohorts in PDAC & NSCLC
 - Combination cohorts with cetuximab for CRC, chemotherapy for PDAC, chemo plus I-O for NSCLC
- **Expand to geographies outside of U.S.**

FDA Engagement Plans

- **Granted Fast Track Designation for**
 - First-line in patients with KRAS G12E advanced or metastatic PDAC and
 - Patients with KRAS G12D locally advanced mPDAC who received at least one previous standard systemic therapy in July 2023
- **Plan to pursue Breakthrough Therapy Designation**
- **Accelerated clinical development input**

Next Steps in VS-7375 Clinical Program

Expect to report a preliminary update on the Phase 1 monotherapy dose escalation in Q4 2025.

Initiate the dose escalation cohorts in combination with cetuximab, chemotherapy, and chemotherapy with checkpoint-inhibitor for CRC, PDAC, and NSCLC, respectively, in Q4 2025.

Subject to the results of the Phase 1 monotherapy dose escalation, **initiate monotherapy expansion cohorts** in PDAC, NSCLC, and other solid tumors.

Subject to the combination dose escalation of VS-7375, **initiate combination expansion cohorts** in CRC, PDAC, and NSCLC.

Topline Data from RAMP 205: Avutometinib + Defactinib + Standard of Care in First-Line Metastatic Pancreatic Cancer



Pancreatic Ductal Adenocarcinoma (PDAC) Represents an Area of Unmet Need

Nearly 180,000 Total Incident PDAC Patients



~59,800¹

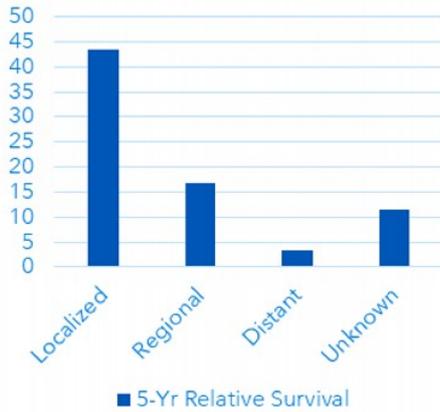


~77,500²

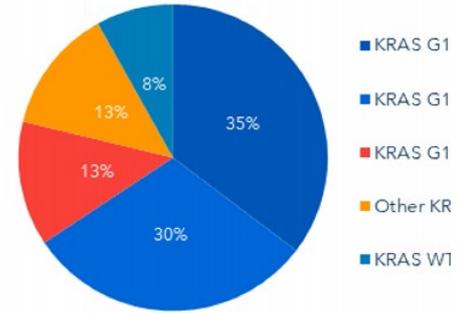


~43,200³

High Unmet Need Based on 5 Year Relative Survival of 13.3%



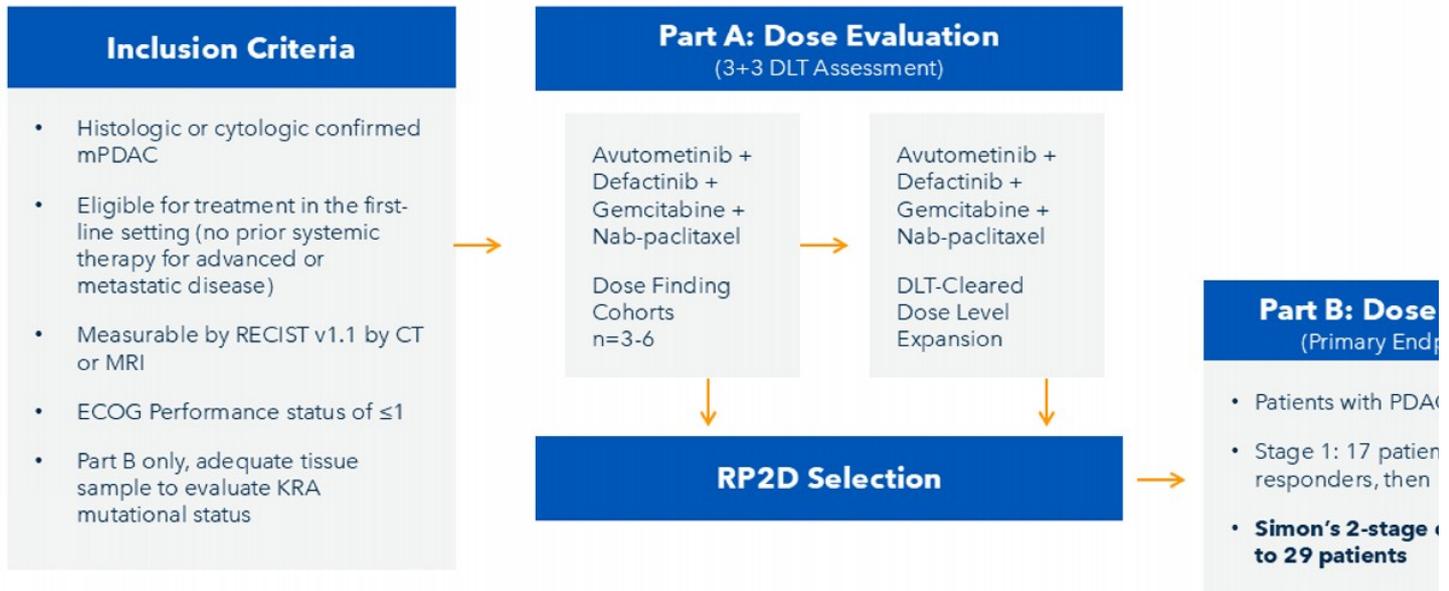
KRAS Mutations Are Present in ~90-95% of PDA



¹SEER 12; ²worldometers; ³Globocan/Japan/factsheet; ⁴McIntyre, Cancer Cell, Sept 2023; ⁵Yu, Frontiers in Oncology, Jul 2021

RAMP 205: Designed to Identify and Evaluate RP2D in Combination Chemotherapy for Treatment of Newly Diagnosed mPDAC

RAMP 205: Ongoing Phase 1/2 Evaluating Avutometinib + Defactinib with Gemcitabine and Nab-paclitaxel



Collaboration with PanCAN, NCT05669482



DLT: dose-limiting toxicity; RP2D: recommended phase 2 dose; CT: computed tomography; ECOG: European Cooperative Oncology Group; MRI: magnetic resonance imaging

RAMP 205: 12 Patients Enrolled in Each Dose Cohort

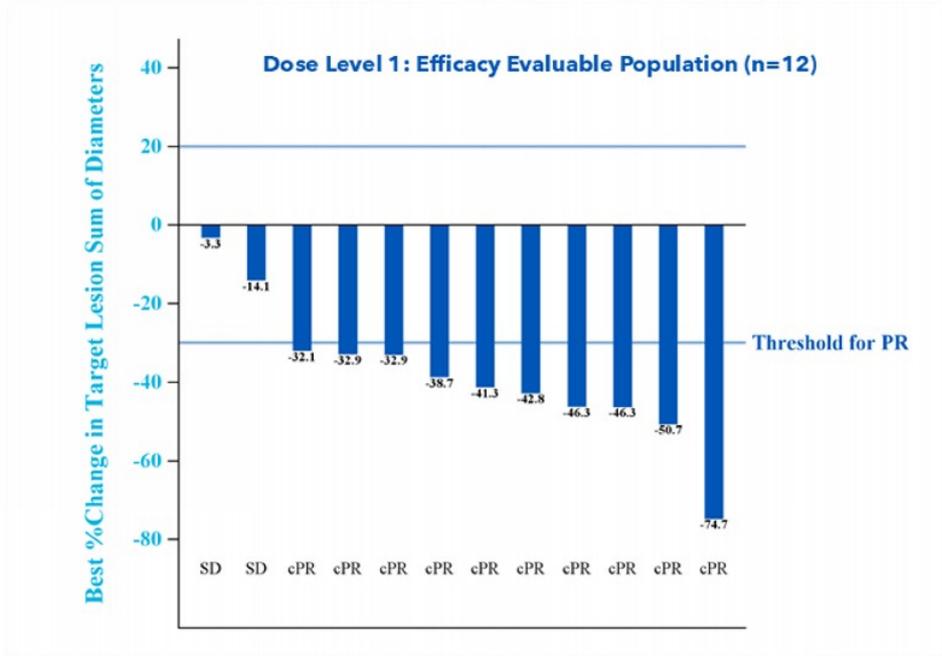
Dose level 1 selected as the RP2D

PART A Enrollment Summary Dose Levels & Administration Schedule (28-day) Cycle

- **Enrolled:** (n=60)
- **Enrolled Study:** (n=30)
- **Treatment Ongoing:** (n=19)
- **Survival Follow Up**

Dose Level	Avutometinib	Defactinib	Gemcitabine	Nab-Paclitaxel	Days Chemo Dosing
1	2.4	200	800	125	1-8-15
0	3.2	200	800	100	1-8-15
-1	2.4	200	800	100	1-8-15
1a	3.2	200	800	125	1-15
2a	3.2	200	1000	125	1-15

Dose Level 1 Demonstrated a Confirmed 83% ORR with Tumor Shrinkage Observed in All Patients

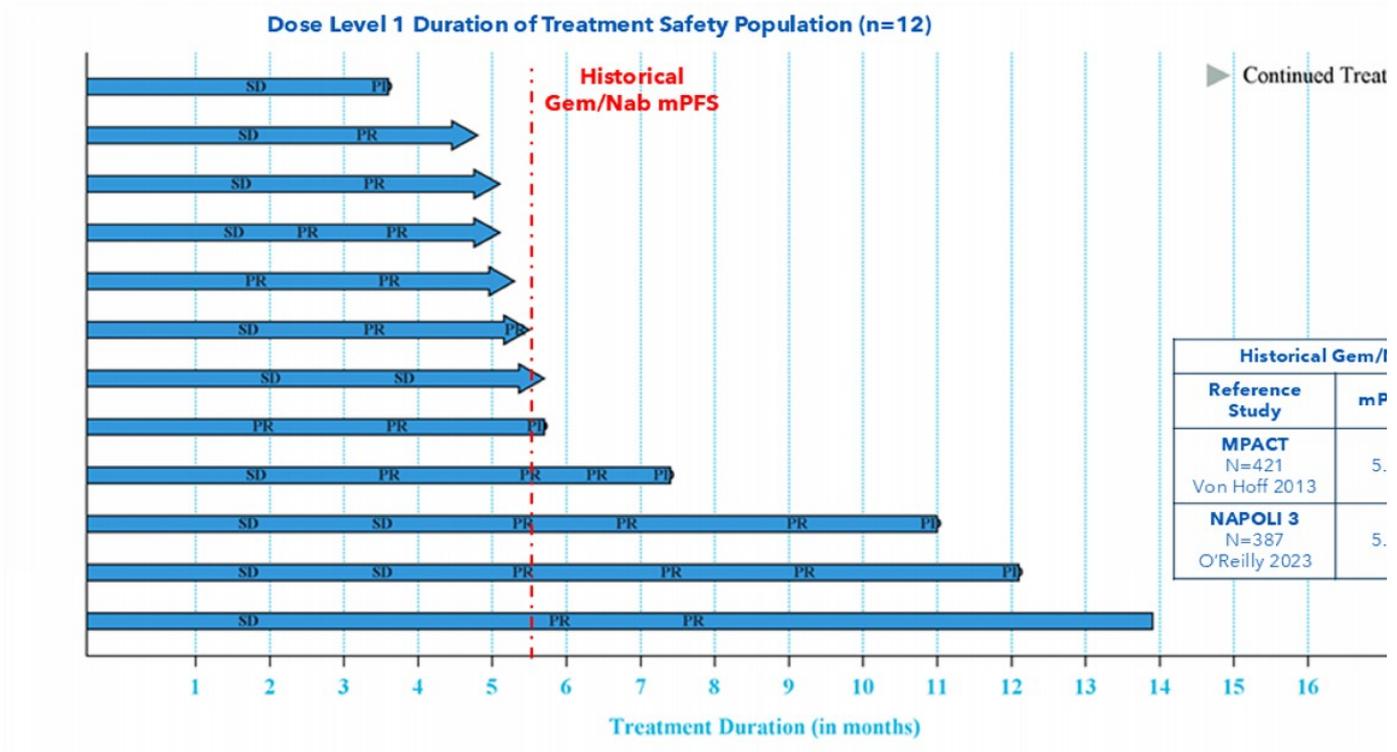


Dose Level 1: Response & Disease Control as of August 1, 2025	
Confirmed ORR, n (%)	
PR, n (%)	
SD, n (%)	
PD, n (%)	
DCR, n (%) ≥ 4 cycles	



Source: F_TR_WATERFALL.sas Data Cut: 01 AUG 25

Encouraging Duration of Treatment Observed to Date for Dose Level



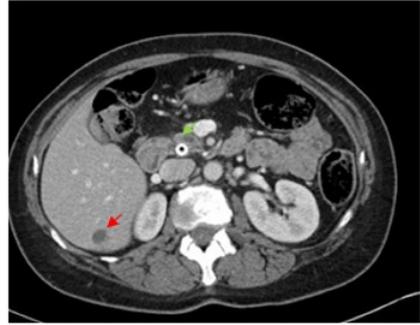
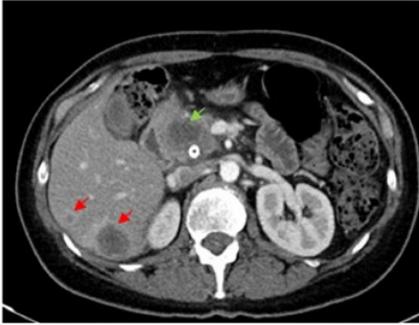
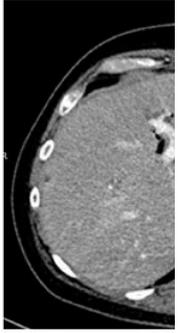
Source: Program:F_TX_R5_SWIMMER_DL1.sas Data Cut: 25 APR 25

Patient with mPDAC Showed Substantial Tumor Shrinkage in Dose

Before C1

After C2

After C4



▲ metastasis ▲ pi

AEs were Generally Manageable, Allowing Patients to Remain on

DL1 Treatment Emergent AE All Grades ≥ 25% / Non-laboratory AEs

TEAEs	Dose Level 1 (n=12)		NAPOLI 3* (N=379)	
	All Grades n (%)	Grade ≥3 n (%)	All Grades n (%)	Grade ≥3 n (%)
Nausea	10 (83)	0	162 (43)	10 (3)
Diarrhoea	9 (75)	0	139 (37)	17 (4)
Fatigue	9 (75)	0	143 (38)	20 (5)
Alopecia	8 (67)	0	119 (31)	Not Listed**
Constipation	8 (67)	0	113 (30)	8 (2)
Oedema peripheral	7 (58)	2 (17)	108 (29)	Not Listed**
Rash maculo-papular	7 (58)	0	Not Listed**	Not Listed**
Stomatitis	7 (58)	1 (8)	45 (12)	Not Listed**
Vomiting	7 (58)	1 (8)	100 (26)	8 (2)
Dysgeusia	6 (50)	0	58 (15)	Not Listed**
Hypotension	6 (50)	0	Not Listed**	Not Listed**
Pyrexia	6 (50)	1 (8)	87 (23)	Not Listed**
Decreased appetite	5 (42)	0	106 (28)	10 (3)
Hypertension	5 (42)	1 (8)	Not Listed**	8 (2)
Neuropathy peripheral	5 (42)	1 (8)	66 (17)	22 (6)
Cough	4 (33)	0	Not Listed**	Not Listed**
Dyspepsia	4 (33)	0	Not Listed**	Not Listed**
Dyspnoea	4 (33)	0	47 (12)	8 (2)
Retinopathy	4 (33)	1 (8)	Not Listed**	Not Listed**
Vision blurred	4 (33)	1 (8)	Not Listed**	Not Listed**
Abdominal Distension	3 (25)	0	Not Listed**	Not Listed**
Abdominal Pain	3 (25)	0	77 (20)	14 (4)
Depression	3 (25)	0	Not Listed**	Not Listed**
Epistaxis	3 (25)	0	43 (11)	Not Listed**
Febrile Neutropenia	3 (25)	3 (25)	Not Listed**	9 (2)
Rash	3 (25)	0	Not Listed**	Not Listed**

- No new or unexpected AEs observed
- Most non-laboratory AEs were grade 1 or 2
- AEs were generally manageable, allowing patients to remain on treatment
- The rates and severities of most AEs are comparable to the individual rates reported for Gem/Nab and Avutometinib/Defactinib
- Nausea, diarrhea, constipation, febrile neutropenia, and anemia may be increased in comparison to treatment with Gem/Nab

DL1 Treatment Emergent AE All Grades ≥ 25% / Laboratory AEs

TEAEs	Dose Level (n=12)	
	All Grades n (%)	Grade ≥3 n (%)
Anaemia	8 (67)	5 (42)
Neutropenia***	8 (67)	6 (50)
Hyperbilirubinaemia***	7 (58)	2 (17)
Thrombocytopenia***	5 (42)	1 (8)
Aspartate aminotransferase increased	3 (25)	1 (8)
Hypokalemia	3 (25)	0



RAMP 205 data source: T_AE_GRADE_FT.sas; Data Cut: 11 APR 25

**Supplement Appendix: (NAPOLI 3), Wainberg et al. The Lancet, Volume 402, Issue 10409, 1272 - 1281

***"Not Listed" when an adverse event is either not reported or did not meet threshold of <10% all grade TEAE or <2% G3 or higher TEAE.

***"Neutropenia" and "Neutrophil count decreased" are grouped as "Neutropenia", "Thrombocytopenia" and "Platelet count decreased" are grouped as "Thrombocytopenia", "Hyperbilirubinaemia" and "Blood bilirubin increased" are grouped as "Hyperbilirubinaemia".

Next Steps for RAMP 205

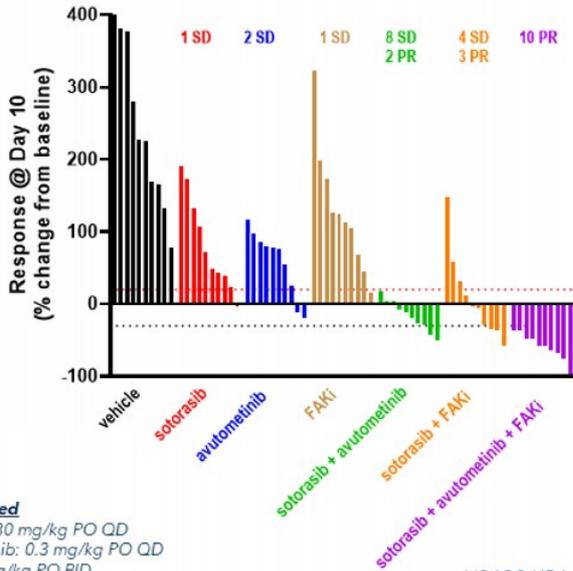
Complete enrollment up to 29 patients in RAMP 205 at the RP2D in Q3 2025

Avutometinib ± Defactinib with Sotorasib (G12Ci) in KRAS G12C mutant NSCLC



Addition of FAK inhibitor Augments the Efficacy of Sotorasib + Av and Reverses Sotorasib Resistance in KRAS G12C NSCLC Preclinical

AVUTOMETINIB ENHANCES SOTORASIB EFFICACY. ADDITION OF FAK INHIBITOR INDUCES DEEP TUMOR REGRESSIONS IN ALL TREATED MICE



Doses Tested

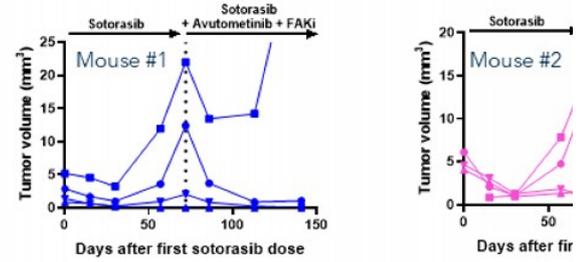
Sotorasib: 30 mg/kg PO QD
Avutometinib: 0.3 mg/kg PO QD
FAKi: 50 mg/kg PO BID

H2122 KRAS G12C NSCLC

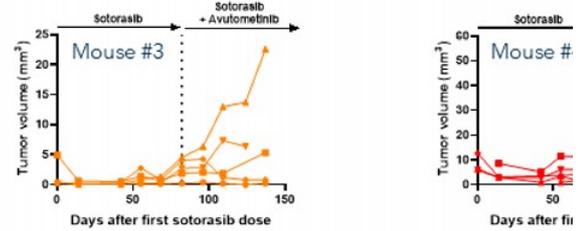


Collaboration Verastem & Mariano Barbacid, CNIO (Spain)

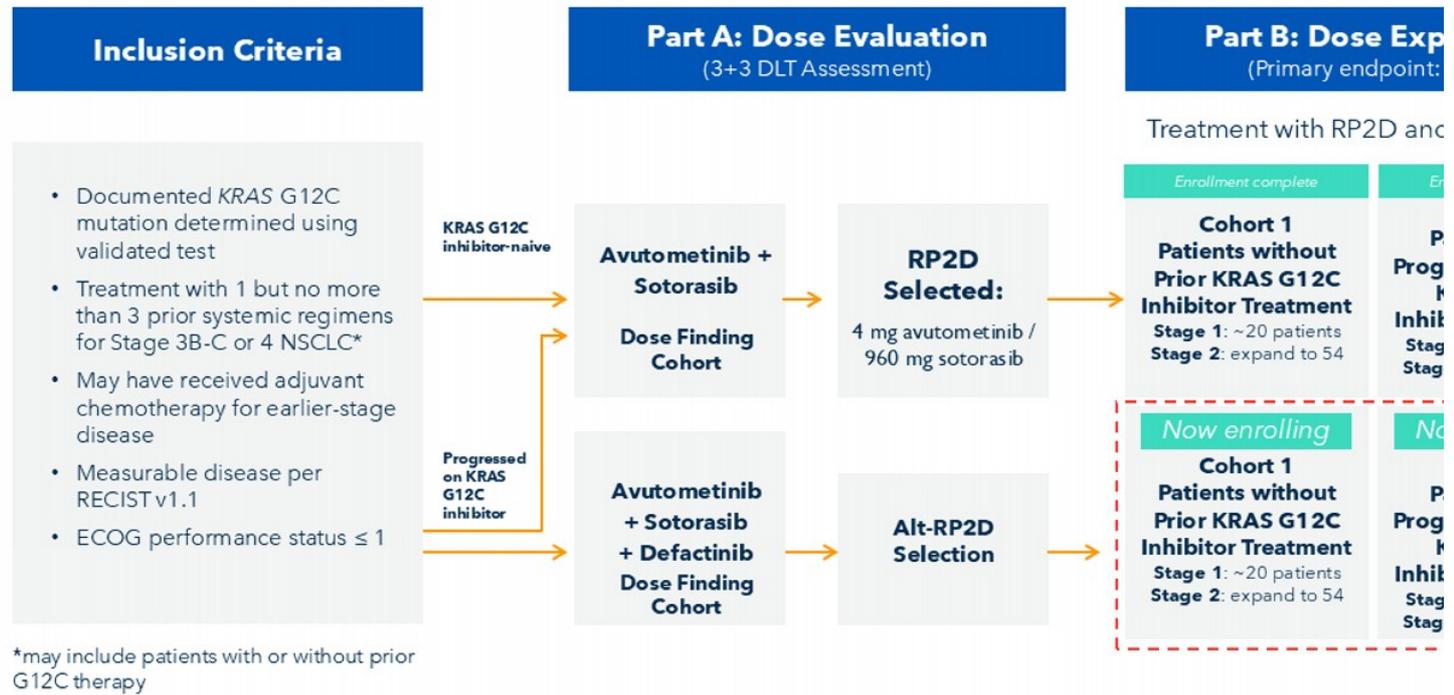
ADDITION OF FAKi + AVUTOMETINIB REVERSES SOTORASIB RESISTANCE



ADDITION OF AVUTOMETINIB IS INSUFFICIENT TO REVERSE SOTORASIB RESISTANCE



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



Collaboration with Amgen, NCT05074810

RECIST v1.1, response evaluation criteria in solid tumours version 1.1

RAMP 203: No Dose-Limiting Toxicities (DLTs) Were Observed in the Triplet Combination Cohort in Patients Previously Treated with a G12C Inhibitor

Triplet Combination Update:

- As of a November 21, 2024, data cutoff, three patients whose cancer previously progressed on a G12C inhibitor have been treated with the triplet combination of sotorasib 960 mg administered daily on a continuous schedule and avutometinib 3.2 mg twice-weekly (BIW) plus defactinib 200 mg twice-daily (BID). Avutometinib and defactinib are administered on a three out of four weeks schedule.
- 2 of the 3 patients demonstrated initial tumor reductions of at least 20% at the first scan. As of the data cutoff, all three patients remain on treatment.
- Completed enrollment of the planned dose level evaluation cohorts for the triplet combination in Q1 2025.

Doublet Combination Update:

- As previously reported, the doublet combination of avutometinib with sotorasib has completed enrollment (n=28) in the G12C inhibitor treatment-naïve Stage 1 Part B cohort.
- Completed enrollment to the KRAS G12C inhibitor treated Stage 1 Part B doublet cohort in Q1 2025.
- Patients enrolled in the doublet cohorts continue to be followed for safety and efficacy results (both treated and treatment-naïve cohorts).

Next Steps for RAMP 203

Report an interim update on the efficacy and safety results from both the doublet and triplet combinations in Q4 2025

Anticipated Milestones & Financials



Execute Successful Commercial Launch in LGSOC, Followed by Me Catalysts to Expand Into Larger, Underserved Patient Populations

Program	Anticipated Milestones & Activities in 2H 2025
 AVMAPKI™ FAKZYNJA™ CO-PACK <small>(avutometinib capsules; defactinib tablets) 0.8 mg, 200 mg</small> KRAS-mutated Recurrent LGSOC	<ul style="list-style-type: none"> <input type="checkbox"/> Complete planned enrollment in RAMP 301 by end of 2025 <input type="checkbox"/> Announce outcome of IDMC's sample-size re-estimation recommendation for RAMP 301 in Q4 2025 <input type="checkbox"/> Report initial data from RAMP 201J Phase 2 clinical trial being conducted in Japan in Q4 2025
VS-7375, oral KRAS G12D (ON/OFF) Inhibitor	<ul style="list-style-type: none"> <input type="checkbox"/> Expect to report a preliminary update on the Phase 1 monotherapy dose escalation in Q3 2025 <input type="checkbox"/> Subject to the results of the Phase 1 monotherapy dose escalation, the Company plans to initiate expansion cohorts in both advanced PDAC and NSCLC. <input type="checkbox"/> Initiate the dose escalation cohorts in combination with cetuximab, chemotherapy, and immunotherapy with checkpoint-inhibitor for CRC, PDAC, and NSCLC, respectively. <input type="checkbox"/> Subject to the results of the Phase 1 dose escalation combinations cohorts with VEGFR inhibitors, the Company plans to initiate a combination expansion cohort in CRC, PDAC, and NSCLC.
Avutometinib + Defactinib + SOC in First-Line Metastatic Pancreatic Cancer	<ul style="list-style-type: none"> <input type="checkbox"/> Complete enrollment in the expansion cohort in Q3 2025
Avutometinib ± Defactinib + Sotorasib: mKRAS G12C NSCLC	<ul style="list-style-type: none"> <input type="checkbox"/> Plan to share interim safety and efficacy results from both double and triplet combinations in Q4 2025

Key Financial Statistics

As of and for the quarter ended June 30, 2025

Cash, cash equivalents & short-term investments	\$164.3M
Net Product Revenue	\$2.1M
GAAP Operating Expenses	\$45.9M
Non-GAAP Operating Expenses	\$42.5M*
Shares Outstanding	61.5M**

Oberland Finance Credit Facility

- Up to \$150M available in a series of notes
 - \$75M principal of notes outstanding
 - Remaining \$75M available at Company's option upon achievement of pre-defined milestones
 - \$25M tranche upon FDA approval of avutometinib and defactinib for treatment of LGSOC
 - \$50M tranche upon trailing six months revenue of at least \$55M
- Floating interest rate, subject to a floor and a cap
- Interest only payments through January 2031
- No financial covenants



*Q2 2025 GAAP operating expenses of \$45.90M less Q2 2025 stock-based compensation expense of \$3.41M = \$42.49M Q1 2025 non-GAAP operating expenses.
**Excludes unexercised warrants (14.1M shares upon exercise) and unexercised pre-funded warrants (9.8M shares upon exercise).

THANK YOU!



AVMAPKI FAKZYNJA CO-PACK Patent Portfolio

OB: 7,928,109 Defactinib COM US (expiry 4/17/2028)	PTA	PTE (Aug, 2034)
OB: 7,897,792 Avutometinib COM (expiry 2/9/2027)		
OB: 8,247,411 Defactinib - genus (expiry 4/17/2028)		
OB: 11,400,090 - Avutometinib mono dosing - (expiry 10/29/2038)		PTE
OB: 11,517,573 A+D combo dosing- (expiry 9/11/2040)		
OB: 11,873,296 Avuto polymorph - (expiry 12/29/2042)		



OB: Orange Book Listed Patent

Experienced Senior Management Team



Daniel Paterson
President and CEO

John Hayslip, M.D.
Chief Medical Officer

Jonathan Pachter, Ph.D.
Chief Scientific Officer

Mike Crowther
Chief Commercial & Strategy Officer

Matt Ros
Chief Operating Officer

Dan Calkins
Chief Financial Officer

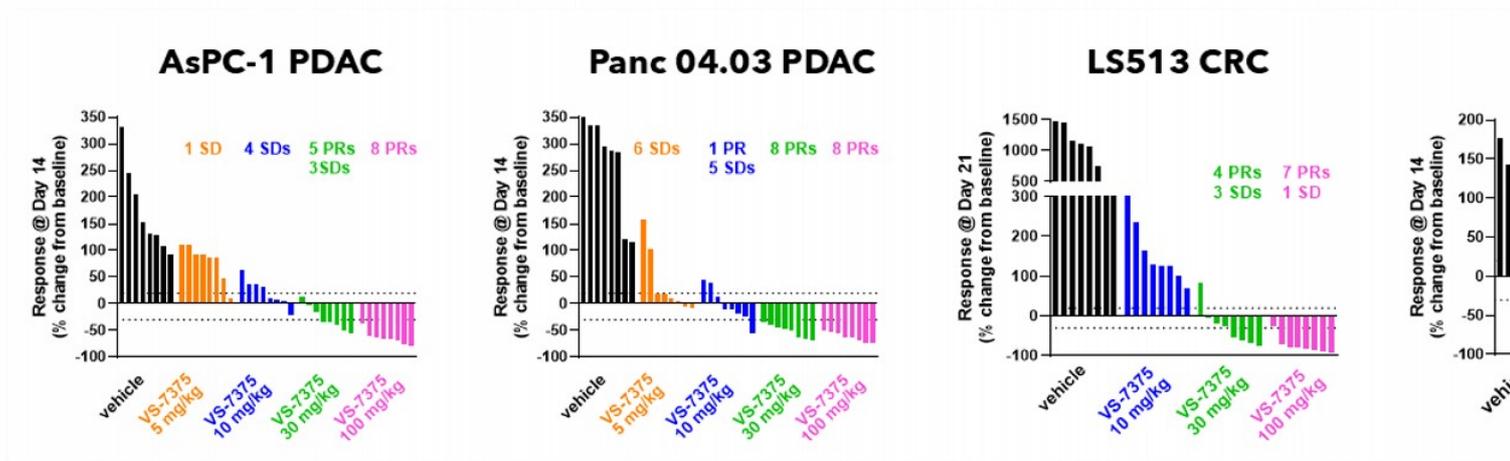
Nate Sanburn
Chief Business Officer

Colleen Mockbe
Global Head of Regulatory Affairs and Development

<p>Previous experience</p>	<ul style="list-style-type: none"> • CEO, The DNA Repair Co. (now On-Q-ity) • PharMetrics (now IMS) • Axion 	<ul style="list-style-type: none"> • CMO, I-MAB • Nektar Therapeutics, AbbVie • University of Kentucky's Markey Cancer Center 	<ul style="list-style-type: none"> • Head of Cancer Biology - OSI (now Astellas) • Schering-Plough 	<ul style="list-style-type: none"> • CBO, Minerva Biotechnologies • Interim US lead and VP of US Marketing, Kite Pharma • Celgene 	<ul style="list-style-type: none"> • CEO, FORE Biotherapeutics • EVP & CSBO, Epizyme • COO, Sanofi-Genzyme • BMS 	<ul style="list-style-type: none"> • Technical Accounting Consultant-CFGI • PwC LLP 	<ul style="list-style-type: none"> • Associate VP, Head of Collaborations & Late Phase BD, Lilly Oncology • National Gene Vector Lab, Indiana University 	<ul style="list-style-type: none"> • Chief Development Officer, SVP of Regulatory OncXerna • Head of Global Regulatory Lilly Oncology
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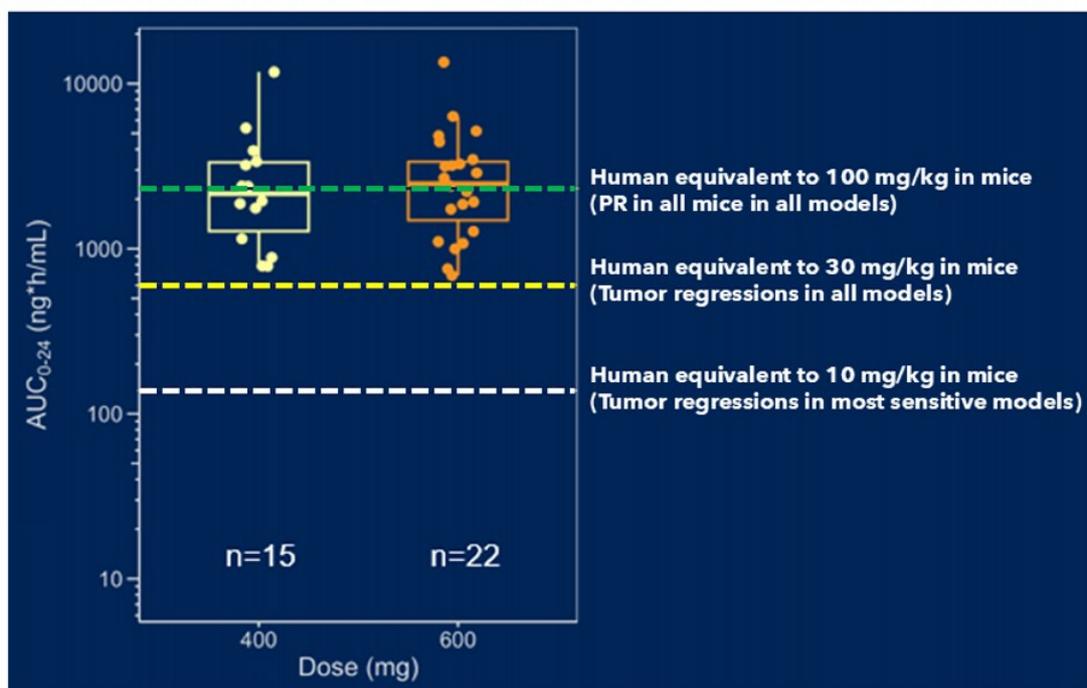


Oral Administration of VS-7375 Inhibits Tumor Growth in a Dose-Dependent Manner in KRAS G12D models

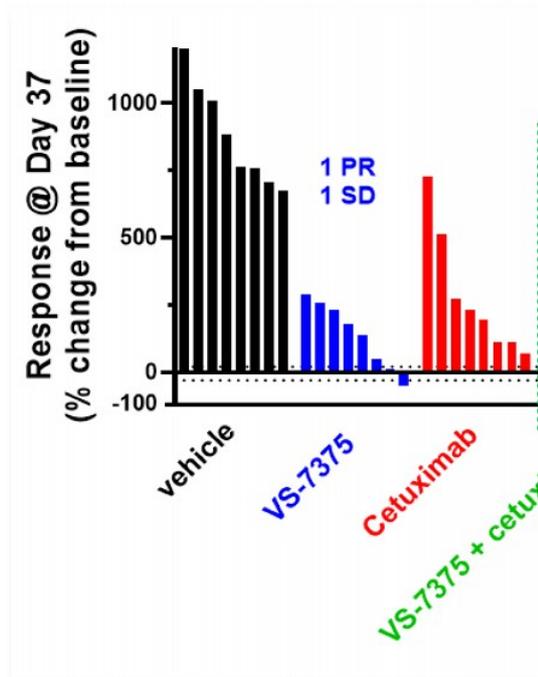
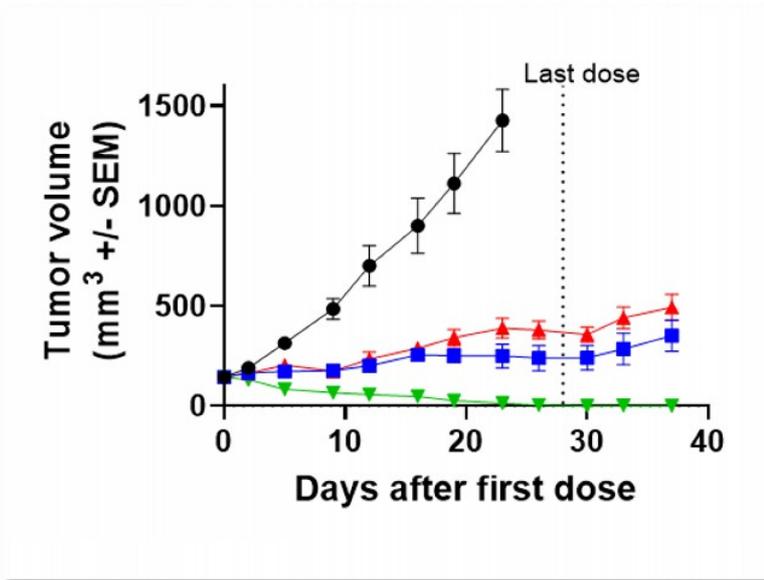


- **10 mg/kg** conferred strong tumor regressions in the most sensitive model (GP2D)
- **30 mg/kg** conferred strong tumor regressions across all models
- **100 mg/kg** conferred partial responses (>30% reduction) in >95% of all mice

Dose Levels of 400 and 600 mg QD Achieved Steady State Target Exposures in Almost all Patients



Addition of Cetuximab with VS-7375 Induces Complete Response Mice in a KRAS G12D Colorectal Cancer Model



LS513 Colorectal Cancer Model

- Vehicle p.o., BID
- VS-7375 10 mg/kg p.o., BID
- ▲ Cetuximab 0.25 mg/mouse IP Q3D
- ▼ VS-7375 + cetuximab



Presented at AACR 2025

VS-7375 Shows Potential Best-in-Class Preclinical Properties Relative to Other Known KRAS G12D Inhibitors

	Criteria	VSTM/ GenFleet VS-7375	Mirati/BMS MRTX1133	RevMed RMC-9805	Lilly LY3962673	AZ AZD0022	Incyte INCB161734	Quanta QTX3046	Tyligand TSN1611
On/Off	ON/OFF selectivity ratio	3x	0.2x	NR	0.016x	NR	1.2x	0.0003x	1.2x
Oral	Oral availability in preclinical models	Y	N	Y	Y	Y	Y	Y	Y
Potency	AsPC-1 pERK IC50	0.5 nM	NR	23 nM	NR	NR	7 nM	30 nM	NR
	Panc 04.03 pERK IC50	0.9 nM	NR	NR	NR	NR	19 nM	NR	NR
Tumor regression	GP2D Oral dose for tumor regression	10 mg/kg PO BID	30 mg/kg IP BID	100 mg/kg PO QD	30 mg/kg PO BID	NR	No regression @ 30 mg/kg PO QD	100 mg/kg PO BID	10 mg/kg PO BID
	Panc 04.03 Oral dose for tumor regression	10 mg/kg PO BID	30 mg/kg IP BID	NR	NR	Tumor stasis @ 150 mg/kg PO BID	Slight regression @ 30 mg/kg PO QD	NR	NR
BBB	Efficacy demonstrated in intracranial model	Y	NR	Y	NR	NR	NR	NR	NR



NR: not reported; IP: intraperitoneal administration; QD: once per day; QW: once per week; References: GenFleet/Verastem AACR 2024; Hallin et al (Mirati) 2022; RevMed AACR 2023; RevMed AACR 2024; AstraZeneca (AZ) AACR 2024; Incyte AACR 2024; Quanta AACR 2023; Tyligand AACR 2024; Betta AACR 2024; Zhou et al (Hangrui) 2024. The safety and efficacy of VS-7375 and the others have not been established. No conclusions on the clinical safety or efficacy of VS-7375 or any of the other G12D inhibitors can be drawn at this time.