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) January 9, 2024

ORIC Pharmaceuticals, Inc.

(Exact name of registrant as specified in its charter)

Delaware (State or other jurisdiction of incorporation) 001-39269 (Commission File Number) 47-1787157 (IRS Employer Identification No.)

240 E. Grand Ave, 2nd Floor South San Francisco, CA 94080 (Address of principal executive offices, including zip code)

(650) 388-5600 (Registrant's telephone number, including area code)

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Not Applicable (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)

Dere-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))

Dere-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	Name of each exchange
Title of each class	Symbol(s)	on which registered
Common stock, par value \$0.0001 per share	ORIC	The Nasdaq Global Select 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 2.02 Results of Operations and Financial Condition.

The information set forth in Item 7.01 is hereby incorporated by reference into this Item 2.02.

Item 7.01 Regulation FD Disclosure.

ORIC Pharmaceuticals, Inc. (the "Company") intends to present an updated corporate presentation (the "Corporate Presentation") at the 42nd Annual J.P. Morgan Healthcare Conference on January 9, 2024. A copy of the Corporate Presentation is furnished as Exhibit 99.1 hereto and is incorporated herein by reference.

All of the information furnished in this Item 7.01 and Items 2.02 and 9.01 (including Exhibit 99.1) shall not be deemed to be "filed" for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), or otherwise subject to the liabilities of that section, nor shall it be deemed incorporated by reference into any filing under the Securities Act of 1933, as amended, or the Exchange Act, except as shall be expressly set forth by specific reference in such a filing.

Description

Item 9.01 Financial Statements and Exhibits.

(d) Exhibits.

Exhibit Number

99.1 <u>Corporate Presentation</u>

104 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.

ORIC PHARMACEUTICALS, INC.

Date: January 9, 2024

By: /s/ Christian Kuhlen Christian Kuhlen, M.D., J.D. General Counsel





OVERCOMING RESISTANCE IN CANCER

Company Overview

January 2024

Forward-Looking Statements

This presentation contains forward-looking statements that involve substantial risks and uncertainties. All statements other than statements of historical facts contained in this presentation, including statements regarding ORIC Pharmaceuticals, Inc.'s ("ORIC", "we", "us" or "our") future financial condition, results of operations, business strategy and plans, and objectives of management for future operations, as well as statements regarding industry trends, are forward-looking statements. In some cases, you can identify forward-looking statements by terminology such as "anticipate," "believe," "continue," "could, "estimate," "expect," "intend," "may," "plan," "potentially," "predict," "should," "will" or the negative of these terms or other similar expressions. Forward-looking statements contained in this presentation also include, but are not limited to, statements regarding; our development plans and timelines; the potential advantages of our product candidates and programs; plans for the clinical trials and development of ORIC-114, ORIC-944 and ORIC-533; the expected timing of reporting data from our clinical trials; our anticipated milestones and clinical updates; and the period over which we estimate our existing cash and investments will be sufficient to fund our current operating plan.

We have based these forward-looking statements largely on our current expectations and projections about future events and trends that we believe may affect our financial condition, results of operations, business strategy and financial needs. These forward-looking statements are subject to a number of risks, uncertainties and assumptions, including, among other things: the timing of the initiation, progress and results of our preclinical studies and clinical trials; tisks associated with the process of discovering, developing and commercializing drugs that are safe and effective for use in humans and operating as an early clinical stage company; negative impacts of the COVID-19 pandemic on our operations, including clinical trials; the potential for current or future clinical trials for product candidates to differ from preclinical, initial, interim, preliminary or expected results; our ability to advance product candidates into, and successfully complete, clinical trials; the timing or likelihood of regulatory flings and approvals; changes in our plans to develop and commercialize our product candidates, if approved; our ability to advance product candidates in developing and the number of patients that may enroll in our clinical trials; the commercialize of the occurrence of any event, change or other circumstance that could give rise to the termination of our license or collaboration agreements; our ability to retain the continued service of our key personnel and to identify, hire and retain additional qualified professionals; the implementation of our business model and strategic plans for urb business and product candidates, if approved; developments relating to our competitors and our industry, including complete; cuincal trials; the scope of protection we are able to establish and maintain for intellectual property rights, product candidates, if approved; developments relating to our competitors and our industry, including completing product candidates and our pipeline; our ability to contract research

This presentation also contains estimates and other statistical data made by independent parties and by us relating to market size and other data about our industry. This data involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such data and estimates. In addition, projections, assumptions and estimates of our future performance and the future performance of the markets in which we operate are necessarily subject to a high degree of uncertainty and risk.

Except as required by law, we undertake no obligation to update any statements in this presentation for any reason after the date of this presentation.

We have filed Current Reports on Form 8-K, Quarterly Reports on Form 10-Q, Annual Reports on Form 10-K, and other documents with the SEC. You should read these documents for more complete information about us. You may obtain these documents for free by visiting EDGAR on the SEC website at www.sec.gov.

This presentation discusses our product candidates that are under preclinical or clinical study, and which have not yet been approved for marketing by the U.S. Food and Drug Administration. No representation is made as to the safety or effectiveness of our product candidates for the therapeutic use for which they are being studied.

ORIC Pharmaceuticals: Dedicated to Overcoming Resistance In Cancer

Broad Pipeline of Potential First-in-Class and Best-in-Class Programs	 Two potential best-in-class programs advancing towards pivotal studies Additional preclinical programs targeting novel and validated targets
Precision Oncology Expertise Enables Accelerated Clinical Timelines	 Rapid timelines enabled by biomarker-driven, patient-selected clinical trials and translational expertise
Dual Engines for Pipeline Expansion	 Track record of building pipeline via internal R&D and business development Targeting one new IND candidate every 18 months
Experienced Management Team	 Heritage of discovering and developing multiple approved oncology medicines at Ignyta, Medivation, Aragon and Genentech
Strong Financial Position	Cash and investments of \$235 million expected to fund company into 2026 ⁽¹⁾
Anticipated Milestones	IC-944 initiation of combination study with AR inhibitor(s): 1H 2024 IC-944 program update: mid-2024 IC-114 initiation of dose expansion in multiple cohorts: 1H 2024 IC-114 updated Phase 1b data: 1H 2025
(1) Approximate unaudited balance as of December 31, 2023.	3

Executive Team with Expertise in Building Leading Oncology Companies

Jacob Chacko, MD Chief Executive Officer	 Previously CFO of Ignyta (acquired by Roche), raised over \$500mm in capital TPG Capital (completed \$10bn of aggregate acquisitions) and McKinsey Board member of 4D Molecular Therapeutics and Board chair of Bright Peak Therapeutics; previously Turning Point, Bonti, RentPath, EnvisionRx, Par Pharma, IMS and Quintiles 	Ignyta TPG McKinsey&Company
Lori Friedman, PhD Chief Scientific Officer	 Previously Head of Translational Oncology at Genentech; advanced over 20 drug candidates into development, two approvals to date Director of Signal Transduction at Exelixis; led new target discovery collaboration with BMS Inventor on 28 issued patents and author on 99 peer-reviewed publications Board member of NextRNA Therapeutics 	Genentech EXELI <mark>X</mark> IS
Pratik Multani, MD Chief Medical Officer	 Previously CMO of Ignyta, led development and regulatory for entrectinib CMO of Fate Therapeutics; contributed to development of Rituxan and Zevalin at Idec, and Treanda at Salmedix; earlier at Dana Farber and MGH Board member of Erasca and Chimerix 	ignyta biogenidec DANA-FARBER
Matt Panuwat Chief Business Officer	 Previously SVP of Business Development at Prothena, established Celgene collaboration for up to \$2.2bn Head of BD at Medivation (acquired by Pfizer), led M&A including the acquisition of talazoparib Global Healthcare Investment Banking at Merrill Lynch 	Prothena MEDIVATION Bankof Americ Merrill Lynch
Dominic Piscitelli Chief Financial Officer	 Previously CFO of AnaptysBio, raised over \$500mm in capital (IPO and follow-on financing) VP of Finance, Strategy and Investor Relations at Medivation VP of Treasury and Finance at OSI Pharmaceuticals (acquired by Astellas) Board member of Celyad Oncology 	AnaptysBio X MEDIVATION (051) pharmaceuticals
Christian Kuhlen, MD General Counsel	 Previously General Counsel at Synthorx (acquired by Sanofi), completed \$151 million IPO General Counsel at Ignyta and Genoptix (acquired by Novartis), executed multiple financings and M&A Attorney at Cooley LLP 	synth@rx Ignyta GENOPTIX
Edna Chow Maneval, PhD SVP Clinical Development	 Previously SVP at Ignyta; clinical lead for entrectinib, led transition team through global filings VP of Clinical Development at Seragon and Aragon, clinical lead for apalutamide Led pivotal Phase 3 study in RCC for Sutent at Pfizer 	

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Clinical Pipeline Focused on Advancement of ORIC-114 and ORIC-944

Program	Indication	Lead Lead Identification Optimization IND Enabling Phase 1 Phase 2 Phase 3	Key Differentiation
PRODUCT CANDIDATE	s		
ORIC-114 EGFR/HER2 exon 20 inhibitor	NSCLC, Breast & Tumor agnostic	Phase 1b: ORIC-114 single agent	✓ CNS active✓ Well tolerated
ORIC-944 PRC2 inhibitor	Prostate Cancer	Phase 1b: ORIC-944 single agent	✓ Potential best-in-class drug properties
OUT-LICENSING CAND	IDATE		
ORIC-533 CD73 inhibitor	Multiple Myeloma	Phase 1b: ORIC-533 combination ready	 ✓ Single agent activity ✓ Clean safety profile ✓ Immune activation
DISCOVERY RESEARC	H PROGRAMS		
ORIC-613 PLK4 inhibitor	Breast cancer		\checkmark First-in-class potential
Multiple programs	Solid tumors		
targeting resistance mechanisms	Solid tumors		
ORIC			5

Substantial Progress in 2023: Well Positioned to Build Value in 2024 and Beyond

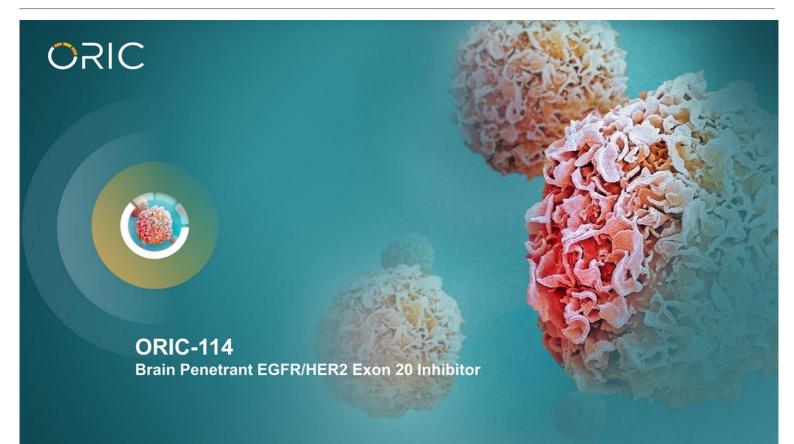
2023 Accomplishments and Next Steps

ORIC-114 EGFR/HER2 exon 20 inhibitor	 Phase 1b dose escalation data presented at ESMO 2023 demonstrated potential best-in-class profile, with favorable safety and both systemic and CNS activity in heavily pre-treated NSCLC patients Initiating multiple dose expansion cohorts in 1H 2024
ORIC-944 PRC2 inhibitor	 Phase 1b dose escalation data demonstrated potential best-in-class drug properties and favorable safety, supporting advancement into combination development in prostate cancer Initiating combination development with AR inhibitor(s) in 1H 2024
ORIC-533 CD73 Inhibitor	 Phase 1b dose escalation data presented at ASH 2023 demonstrated favorable safety and clinical activity in heavily pre-treated multiple myeloma patients Pursuing strategic partnership for combination studies
Discovery Research	 Presented preclinical data confirming therapeutic potential of highly selective PLK4 inhibitors as synthetic lethal for TRIM37 amplified breast cancer Advanced ORIC-613, a novel, highly selective PLK4 inhibitor, through IND enabling studies
Corporate	 ✓ Strengthened balance sheet with \$85 million financing from healthcare specialist funds ✓ Extended cash runway into 2026

ORIC-114 and 944 rapidly advancing, with potential registrational studies for both programs expected to initiate in 2025

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ORIC-114 Is a Promising Candidate for Patients with Tumors Harboring EGFR and HER2 Exon 20 Insertion Mutations, Including Those with Brain Metastases

ORIC-114 Target Product Profile

Selective and Potent	 Selectively targets EGFR and HER2 with high potency against exon 20 insertion mutations Exquisite kinome selectivity with limited potential for off-target activity 	
Robust In Vivo Efficacy	 Significant tumor regression in multiple exon 20 insertion models Superior therapeutic index in vivo with improved efficacy and tolerability than competitor molecules 	Promising Phase 1b Results Well tolerated safety profile Curtering estimity meet emission of the set of the
Highly Brain Penetrant	 High unbound (free) brain exposures in vivo Substantial tumor regression in intracranial efficacy studies 	 ✓ Systemic activity post-amivantamab ✓ CNS activity

ORIC-114 is a potentially best-in-class EGFR and HER2 exon 20 inhibitor with excellent selectivity and brain penetrance

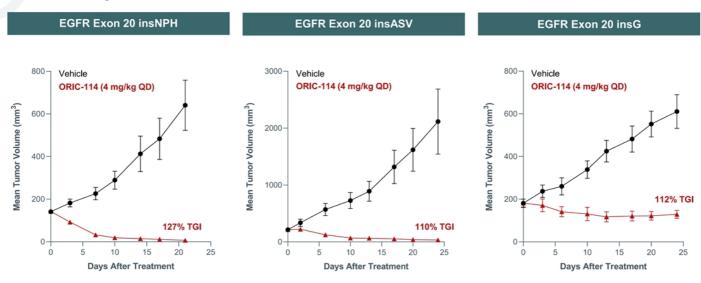
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ORIC-114 Was Designed to Selectively Target EGFR and HER2 with High Potency Against Exon 20 Insertion Mutations

Kinome Selectivity	Comparison							
ORIC-114	CLN-081	Furmonertinib	BLU-451*	Mobocertinib				
	Off-target Wild	Itype (WT) Kinases Inhibited 80	0-100% at 1 μM					
ORIC-114	CLN-081	Furmonertinib	BLU-451	Mobocertinib				
0	7	4	7*	7				
ORIC-114 has de	ORIC-114 has demonstrated an exquisitely clean kinome panel, which is especially important for covalent inhibitors							
Source: Juntila et al. ESMO Note: ORIC-114, mobocertin	Poster (2023) and Murray et al. AACR Poster (2022). b, CLN-081, data conducted head-to-head in 468 kinases at	1uM. Top 10% shown. *BLU-451 data not conducted h	ead-to-bead in 409 kinases at 1uM.					

ORIC-114 Demonstrates Potent In Vivo Activity in EGFR Exon 20 Insertion Models



In Vivo Efficacy – NSCLC EGFR Exon 20 Insertion Models

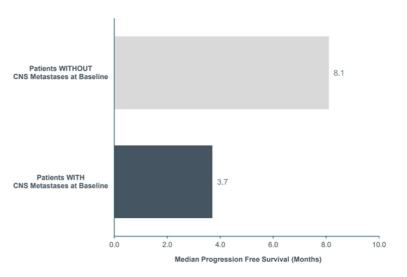




Drugs Lacking CNS Activity Often Have Worse Clinical Outcomes in NSCLC

Case Study on NSCLC Targeted Therapy without CNS Activity: Mobocertinib

- In mobocertinib's phase 1/2 trial, ~35% of patients had CNS metastases at baseline
- Patients with CNS metastases at baseline had markedly worse outcomes
 - Brain was the first site of progression in 68% of patients with CNS metastases at baseline and in 38% of all patients
 - ORR was 25% in patients with CNS metastases at baseline compared to 43% in all patients



Approximately 35% of EGFR exon 20 NSCLC patients have CNS metastases at baseline and the brain is a frequent site of progression in patients with and without CNS metastases at baseline, leading to shorter PFS with therapies lacking CNS activity

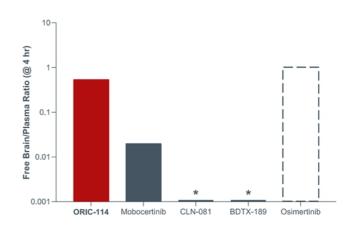
ORIC Source: Janne et al. ASCO Presentation (2019) and Ramalingam et al. ASCO Poster (2021).

Superior Brain Penetration of ORIC-114 Differentiates from Comparator Exon 20 Targeted Agents

ORIC-114 Properties Allowed Optimization of Brain Exposure

- Minimal pump engagement
 - Key pumps that limit brain penetration, PGP and BCRP drug transporters, have minimal impact on ORIC-114 in cell assays
- Suitable physicochemical properties
 - LogP, LogD, TPSA, MW, HBD/HBA, pKa
- · High free unbound exposure in brain tissue
 - Mouse Kp,uu 0.5
 - Dog Kp,uu 1.5

ORIC-114 Exhibits High Ratio of Free (Unbound) Brain/Plasma Exposure in Mice

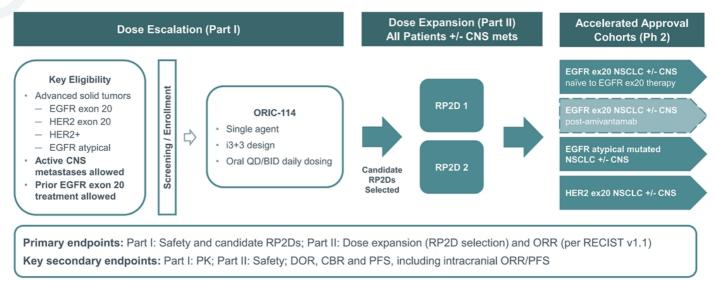


Extensive preclinical profiling demonstrates superior CNS properties of ORIC-114 versus competitors; Excellent free brain exposure across species for ORIC-114 as exhibited by Kp,uu

ORIC Source: Juntila et al. AACR Poster (2021), Juntila et al. AACR-NCI-EORTC Presentation (2021) and ORIC data on file. * Brain exposures were below quantification limit.

First-In-Human Phase 1b Study of ORIC-114

Phase 1b, Multicenter, Open-Label Study



Initial safety, PK/PD, and preliminary antitumor data from dose escalation (part I) presented at ESMO 2023

ORIC Note: ClinicalTrials.gov identifier: NCT05315700. Dose expansion may include QD and BID dosing, fed/fasted dosing. RP2D = recommended Phase 2 dose

ORIC-114 Phase 1 Patient Disposition and Baseline Characteristics

Patient Disposition and Baseline Characteristics

		EGFR Ex20 (n=21)	HER2 Ex20 (n=24)	HER2+ (n=5)	Total (N=50)
50 patients were treated with	Age, years, median (range)	63 (31,80)	63 (25,86)	66 (48,68)	63 (25,86)
increasing doses of ORIC-114	Females, n (%)	10 (48)	11 (46)	3 (60)	24 (48)
	ECOG performance score, n (%)				
Of the NSCLC patients with	0	1 (5)	10 (42)	3 (60)	14 (28)
EGFR exon 20	1	20 (95)	14 (58)	2 (40)	36 (72)
— ≥1 prior EGFR ex20: 81%	Non-smoker, n (%)	12 (57)	16 (68)	3 (60)	31 (62)
	Prior lines of therapies, median (min, max)	2 (1,6)	2 (0,7)	4 (1,7)	2 (0,7)
— ≥2 prior EGFR ex20: 19%	Prior therapies, n (%)				
 CNS mets at baseline: 86% 	Chemotherapy	21 (100)	23 (96)	5 (100)	49 (98)
	EGFR targeted agents	18 (86)	1 (4)	-	19 (38)
Of the NSCLC patients with	EGFR exon 20 targeted agents	17 (81)	-	-	17 (34)
HER2 exon 20	Amivantamab	15 (71)	-	-	15 (30)
— ≥1 prior HER2 agent: 30%	Mobocertinib	4 (19)	-	-	4 (8)
CNC moto at baseline: 200/	Other (CLN-081, BLU-451)	2 (10)	-	-	2 (4)
 CNS mets at baseline: 38% 	HER2 targeted agents	-	7 (30)	3 (60)	10 (20)
	CNS metastases at baseline, n (%)	18 (86)	9 (38)	1 (20)	28 (56)

Phase 1b enrolled heavily pretreated patients with exceptionally high rates of prior exon 20 targeted therapy and CNS metastases at baseline



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ORIC-114 Has Been Generally Well Tolerated Despite More Heavily-Pretreated Patients and Less Stringent Enrollment Criteria for Prior Therapy and CNS Disease

Treatment Related Adverse Events Occurring in ≥10% of Patients

		2	<45 mg (n=1			4	5 – 60 n (n=2	•)		≥75 mg (n=			Total (N=50)
Well tolerated safety profile	Preferred Term, n (%)	Gr1	Gr2	Gr3	≥Gr4	Gr1	Gr2	Gr3	≥Gr4	Gr1	Gr2	Gr3	≥Gr4	All Grades
with mostly Grade 1-2 TRAEs	Rash*	6 (33)	4 (22)	-	- 1	6 (26)	6 (26)	-	-	4 (44)	1 (11)	-	-	27 (54)
 Minimal EGFR-wt related or 	Diarrhea	2 (11)	2 (11)	-	-	7 (30)	2 (9)	2 (9)	-	2 (22)	2 (22)	1 (11)	-	20 (40)
other toxicities	Stomatitis	4 (22)	2 (11)	-	-	2 (9)	2 (9)	1 (4)	-	2 (22)	2 (22)	-	-	15 (30)
 Low rates and severity of 	Paronychia	1 (6)	2 (11)	-	-	4 (17)	4 (17)	-	-	2 (22)	1 (11)	-	-	14 (28)
rash and diarrhea	Pruritis	2 (11)	-	-	-	4 (17)	2 (9)	1 (4)	-	1 (11)	1 (11)	-	-	11 (22)
— No Grade ≥3 rash	Nausea	1 (6)	-	_	_	2 (9)	2 (9)	-	-	1 (11)	1 (11)	1 (11)	_	8 (16)
- Low rate of Grade 3	Decreased appetite	-	1 (6)	_	-	5 (22)	1 (4)	-	-	-	-	-	-	7 (14)
diarrhea (6%)	Vomiting	2 (11)	-	-	-	2 (9)	-	-	-	1 (11)	1 (11)	1 (11)	-	7 (14)
 Infrequent dose reduction and discontinuations 	Dose Reductions		2 (1	8)			3 (1	13)			3 (3	33)		8 (16)
	Dose Discontinuations		1 (9	9)			1 (4	4)			-			2 (4)

ORIC-114 was well tolerated with mainly Grade 1 and 2 adverse events and little evidence of off-target toxicities

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Note: All data as of the data cut-off on September 26, 2023. * Rash includes the following terms: acne, dermatitis, dermatitis, acentiform, eczema, hand dermatitis, and rash. TDD, total daily dose. TRAE, treatment related adverse ex

EGFR Exon 20: Tumor Regression Observed Across All Active ORIC-114 Doses





Systemic and CNS activity observed in heavily pretreated patients, including prior EGFR exon 20 therapy & active brain metastases

Note: All data as of the data cut-off on September 26, 2023. ND: not done, SD/PR: Non-CR/Non-PD (RECIST 1.1), Response-evaluable: Dose level ≥45 mg TDD and at least one post-baseline tumor assessment

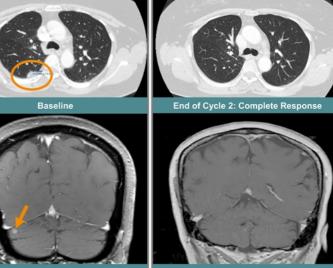
Confirmed Complete Intracranial and Systemic Response in Patient with EGFR Exon 20 Mutated NSCLC and Active CNS Metastases Progressed on Prior EGFR Exon 20 Targeted Therapy

- · Patient: 55F with EGFR exon 20 mutated NSCLC
- Prior therapy: Pemetrexed/cisplatin and amivantamab · Metastases at baseline: Four active CNS non-target
 - lesions
 - Previously untreated
 - No prior surgery
- No prior radiation
- ORIC-114 dose: 75 mg QD
- · Systemic response: Partial response after Cycle 1 (60% reduction in all target and non-target lesions) followed by complete response at the end of Cycle 2 (100% reduction of all target and non-target lesions), subsequently confirmed
- CNS response: Complete response after Cycle 1 (100% reduction of all 4 CNS lesions) confirmed after Cycle 2
- Grade ≥2 treatment-related AEs: Grade 2 mucositis and paronychia
- Duration of treatment: Cycle 9 (ongoing)

Baseline (1 out of 4 lesions shown) End of Cycle 1: Complete Response

ORIC-114 demonstrated single agent clinical activity in a key cancer and molecular subtype (i.e., NSCLC and EGFR exon 20) in a patient typically excluded from clinical trials (i.e., active brain metastases) and previously treated with EGFR exon 20 therapy

ORIC Note: All data as of the data cut-off on September 26, 2023



ORIC-114 Is the Only EGFR Exon 20 Inhibitor to Demonstrate a Systemic Complete Response and CNS Complete Response, Despite More Challenging Patients

Comparison of Selected Eligibility Criteria, Baseline Characteristics, and Clinical Activity

- EGFR exon 20 inhibitor clinical studies typically EXCLUDE:
 - Prior EGFR exon 20 treatment
 - Untreated CNS metastases
- ORIC-114 trial enrolled significantly higher percentage of patients with prior EGFR exon 20 treatment and baseline CNS metastases
- Despite more challenging patients, ORIC-114 demonstrated:
 - Systemic complete response
 - CNS complete response in untreated CNS metastases
 - Responses post-amivantamab

	Amivantamab	CLN-081	Sunvozertinib	Furmonertinib	BLU-451	ORIC-114
Trial	Phase 1	Phase 1	Phase 2	Phase 1	Phase 1	Phase 1
ENROLLMENT						
Prior EGFR ex20i Allowed (1)	No	No	No	No	Yes	Yes
% Prior EGFR ex20i	1%	4%	3%	NR	75%	81%
Untreated CNS Mets Allowed	No	No	No	No	Yes	Yes
% Baseline CNS Mets	22%	38%	32%	34%	58%	86%
CLINICAL ACTIVITY						
Systemic Complete Response	Yes	No	No	No	No	Yes
CNS Complete Response in Untreated CNS Mets ⁽²⁾	No	No	No	No	No	Yes
ORR in EGFR ex20i Naive	~40%	~41%	~61%	42%	TBD	TBD
Post-Amivantamab Response	NA	No	Yes	No	No	33% confirme ORR (at 75 m

Even while allowing patients with prior exon 20 treatment and untreated brain metastases, ORIC-114 is the only EGFR exon 20 inhibitor to demonstrate a confirmed complete systemic response and confirmed complete CNS response

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Note: All data as of the data cut-off on September 26, 2023. Source: Park et al. J Clin Oncol (2021), Zhou et al. JANA Oncology (2021), Piotrowska et al. J Clin Oncol (2023), Han et al. WCLC Presentation (2023), Wang et al. ASCO Presentation (2023), and Nguyen et al. ASCO Poster (2023). (1) Anivnatamab prohibited priving CFGR exon 20 treatment in dose expansion. CLN-081 allowed prior EGR exon 20 treatment selectively during accelerated titration dose escalation only. (2) Treatment history for brain metastases not disclosed for BLU-451. 18

HER2 Exon 20: Tumor Regression Observed Across All Active ORIC-114 Doses





ORIC-114 demonstrated clinical activity in patients with HER2 exon 20 NSCLC, including in patients with baseline brain metastases

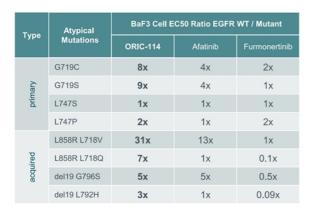
Note: All data as of the data cut-off on September 26, 2023. ND: not done, SD/PR: Non-CR/Non-PD (RECIST 1.1), Response-evaluable: Dose level ≥45 mg TDD and at least one post-baseline tumor assessment

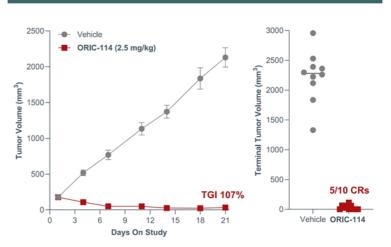
In Addition to EGFR Exon 20, ORIC-114 Also Demonstrates Excellent Preclinical Activity Against Atypical Mutations in EGFR, Revealing an Additional Opportunity

ORIC-114 In Vitro and In Vivo Activity in Atypical EGFR Mutations

ORIC-114 Demonstrates Superior In Vitro Potency

ORIC-114 Induces Complete Tumor Regressions In Vivo



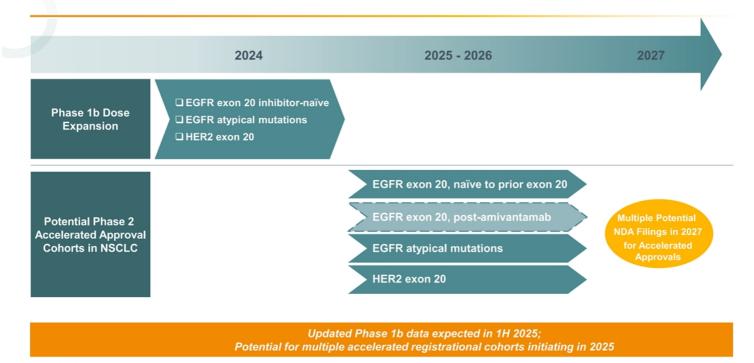


ORIC-114 is a promising therapy for NSCLC patients with atypical mutations in EGFR, based on promising preclinical activity and the emerging clinical profile in patients with EGFR and HER2 exon 20 insertion mutations

Source: Junitia et al. ESMO Poster (2023). Note: Left graph, Bal/F3 cells stably expressing EGFR wild-type or EGFR carrying classical or atypical mutations. Middle graph, EGFR G719S a terminal tumor volumes at day 21. Mean is indicated by a line. No body weight loss observed. Complete response, CR, defined as tumor of undetectable size.

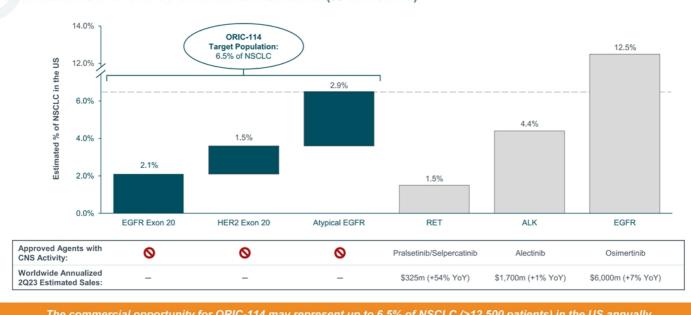
ograft model. Right graph: Individual

ORIC-114 Planned Next Steps and Potential Registrational Path(s)



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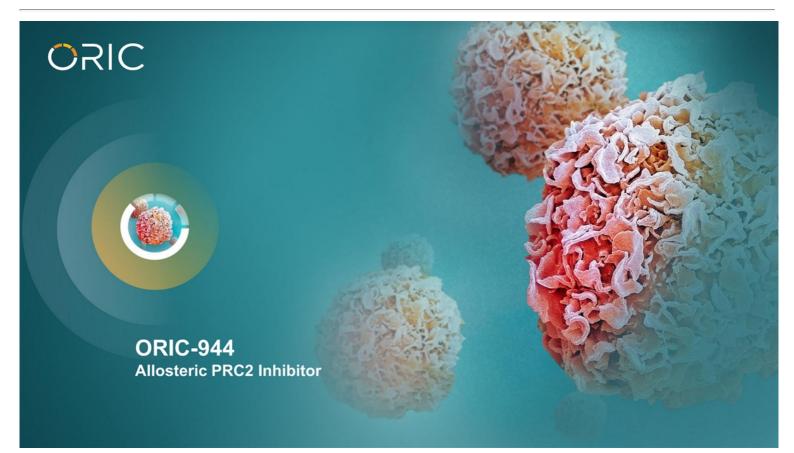
ORIC-114 Is Pursuing a Significant Commercial Opportunity Across Multiple Patient Populations that Do Not Have CNS Active Agents Approved or in Late-Stage Development



Estimated US Prevalence of Exon 20 Insertions (% of NSCLC)

Source: American Cancer Society Cancer Facts & Figures 2023, Gainor et al. Clin Cancer Res (2013), Hirsch et al. Lancet (2017), Rosen et al. Clin Cancer Res (2021), Heymach et al. WCLC Presentation (2018) and Robichaux et al. Nature (2021). AstraZeneca, Roche and Lilly corporate filings. Note: Worldwide annualized 2023 estimated sales calculated using reported 2023 sales and adjusted for an annual run rate.

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ORIC-944 Is a Promising Next-Generation PRC2 Inhibitor Focused on Patients with Prostate Cancer

ORIC-944 Target Product Profile

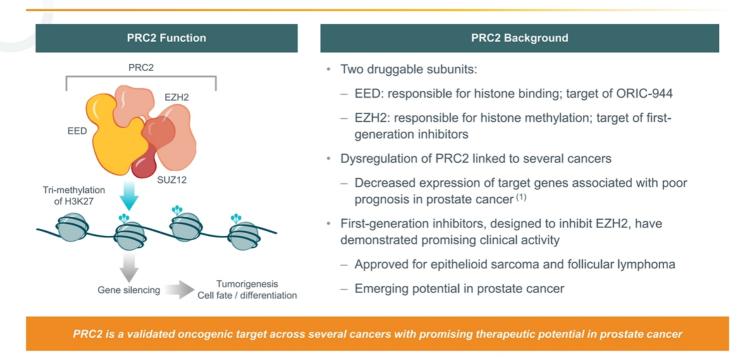
Selective and Potent	Allosteric inhibitor of PRC2 by selectively targeting EEDPicomolar biochemical potency	
Robust In Vivo Efficacy	 Significant single agent activity demonstrated in treatment resistant prostate cancer models ORIC-944 appears more effective than EZH2 inhibitors in preclinical models 	Promising Phase 1b Results ✓ Best-in-class drug properties ✓ Robust target engagement
Best-in-Class Drug Properties	 Approved and other PRC2 inhibitors in development appear to be limited by poor drug properties ORIC-944 designed for improved drug properties over other PRC2 inhibitors 	 ✓ Well tolerated safety profile

ORIC-944 is a potential best-in-class PRC2 inhibitor with superior drug properties suitable for combination development in prostate cancer

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PRC2 Plays Pivotal Role in Transcriptional Regulation and Cancer

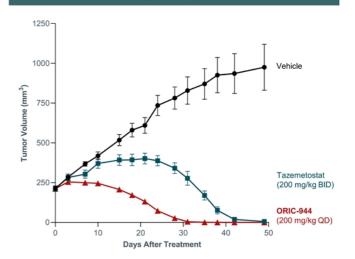


Note: EZH1, enhance of zeste homolog 1. EZH2, enhance of zeste homolog 2. EED, embryonic ectoderm development. SUZ12, suppressor of zeste 12. H3K27, histone H3 at lysine 27. (1) Yu et al. Cancer Res. (2007).

ORIC-944 Targets the EED Subunit of PRC2 and Has Demonstrated Superior Single Agent In Vivo Efficacy and Favorable Drug Properties

EED Inhibition May Improve Upon EZH2 Inhibitors

- ORIC-944 allosterically inhibits PRC2 by targeting EED
- Allosteric inhibition of PRC2 through EED may address limitations of EZH2 inhibitors
 - Active against EZH2 innate resistant PRC2 mutants⁽¹⁾
 - Prevent acquired resistance through secondary mutations in EZH2 ⁽²⁾
 - Inhibit compensatory bypass activity of EZH1 ⁽³⁾
- ORIC-944 is associated with improved drug properties over other PRC2 inhibitors ⁽⁴⁾
- ORIC-944 appears more effective than EZH2 inhibitors in preclinical models



ORIC-944 Induces Complete Regression in DLBCL Model

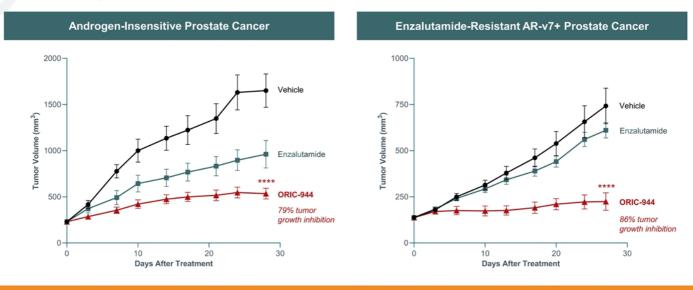
Allosteric PRC2 inhibition through EED provides many potential benefits over EZH2; Notably, ORIC-944 has improved drug properties over other PRC2 inhibitors

Source: Daemen et al. AACR Poster (2021). (1) Gi et al. Nat Chem Biol (2017). (2) Bisserier et al. Blood (2018). (3) Shen et al. Mol Cell (2008) and Honma et al. Cancer Sci (2017). (4) Italiano et al. Lancet Oncol (2018), Harb et al. TAT (2018) and Yap et al. Clin Cancer Res (2019). Note: DLBCL, diffuse large B-cell lymphoma. Right graft: KARPAS-422 xenograft model.

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ORIC-944 Demonstrated Strong Single-Agent Activity in Prostate Cancer Models

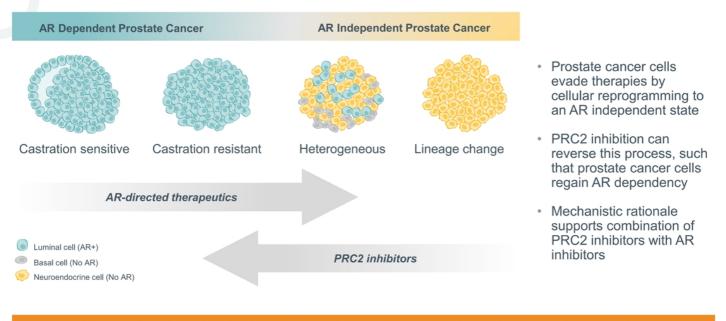
In Vivo Efficacy – Prostate Cancer Models



Based on In vivo efficacy observed in multiple prostate cancer models and improved drug properties, ORIC-944 is positioned as a potential best-in-class PRC2 inhibitor for prostate cancer

Source: ORIC data and Daemen et al. AACR Poster (2021). Note: ORIC-944 dose used was 200 mg/kg QD. Enzalutamide dose used was 30 mg/kg QD. ****p < 0.0001. Left graph: C4-2 xenograft model. Right graph: 22Rv1 xenograft model

PRC2 Epigenetic Dysregulation Plays a Key Mechanistic Role During the Progressive Reprogramming of Prostate Cancers Treated with AR Inhibitors

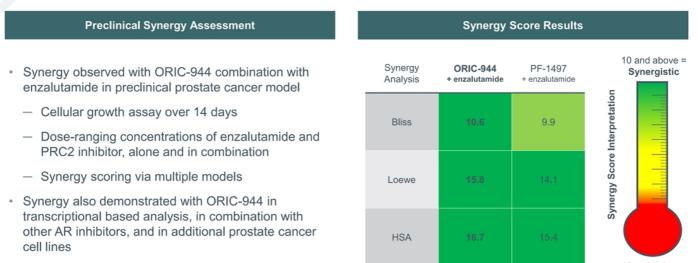


Therapeutic potential of PRC2 inhibitors in prostate cancer is maximized in combination with AR inhibitors

Source: Mu et al. Science (2017), Dardenne et al. Cancer Cell (2016), Davies et al. Nat Cell Biol (2021), Nouruzi et al. Nat Commun (2022), and Goel et al. Semin Cancer Bio (2022).

ORIC-944 Synergizes with AR Inhibitors in Preclinical Prostate Cancer Models

Combination Potential of PRC2 and AR Inhibition



-10 and below = Antagonistic

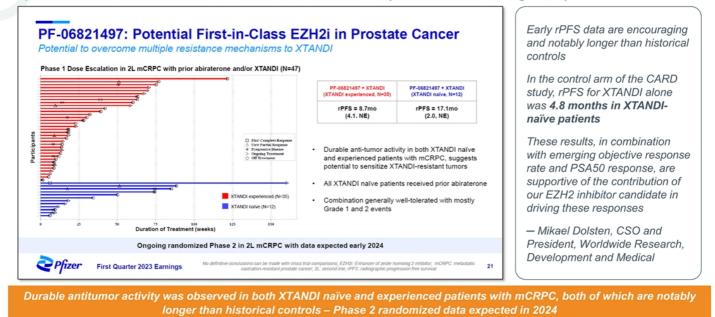
ORIC-944 synergizes with AR inhibitors in prostate cancer models, providing rationale for clinical development

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Source: Data in C4-2 prostate cancer cells shown

Pfizer Phase 1 Data Clinically Validates Synergy of PRC2 Inhibitor and AR Inhibitor in Prostate Cancer

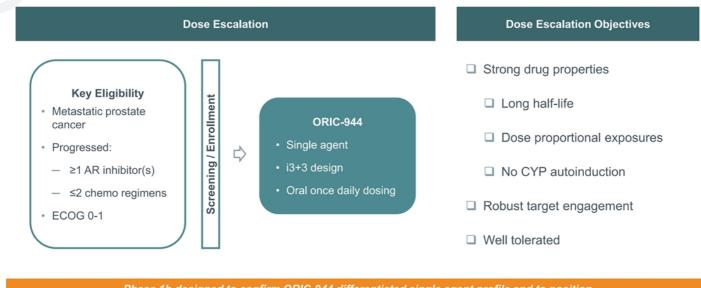
Updated PF-06821497 Phase 1 Data in Prostate Cancer (1Q23 Pfizer Earnings Call)



Source: Pfizer first quarter 2023 earning call on May 2, 2023

First-In-Human Phase 1b Study of ORIC-944 in Metastatic Prostate Cancer

Phase 1b, Multicenter, Open-Label Study



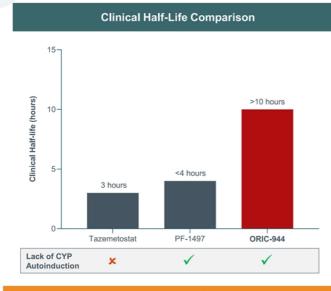
Phase 1b designed to confirm ORIC-944 differentiated single agent profile and to position



ORIC Note: All data as of December 10, 2023.

ORIC-944 Has Demonstrated a Potential Best-in-Class Pharmacokinetic Profile

Preliminary Phase 1b Pharmacokinetic Data



Key Takeaways

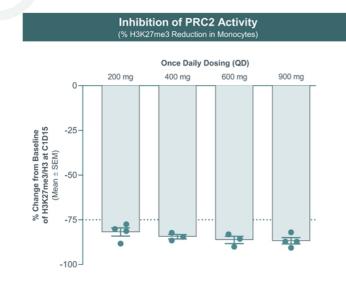
- · Excellent pharmacokinetic profile observed to date
 - Increased exposure with dose level
 - Low inter-patient variability
- No signs of CYP autoinduction that is observed with first-generation PRC2 inhibitors
- Clinical half-life consistent with preclinical prediction of >10 hours, which is superior to other PRC2 inhibitors and supports QD dosing
- Exposures at ≥600 mg QD exceed target Cmin that provides 90% TGI in preclinical in vivo prostate cancer models

Dose exploration continues with favorable plasma half-life and exposures consistent with best-in-class drug properties

ORIC Note: All data as of December 10, 2023. Source: Tazemetostat half-life from FDA label. PF-06821497 half-life estimated from Schweizer et al. ESMO Poster (2022).

ORIC-944 Has Demonstrated Robust and Consistent Target Engagement

Preliminary Phase 1b Pharmacodynamic Data



 Robust target engagement demonstrated with oncedaily monotherapy dosing

Key Takeaways

- Maximal decrease (≥75%) in H3K27me3 in monocytes from peripheral blood samples achieved across multiple dose levels, starting as low as 200 mg QD
- · Low inter-patient variability observed

ORIC-944 has demonstrated promising pharmacodynamic data, indicating strong target engagement

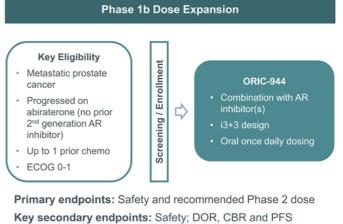
Note: All data as of December 10, 2023. Note: % H3K27me3 inhibition in monocytes from Phase 1b patients shown on plot. Samples collected at cycle 1 day 15, or at cycle 1 day 22 for 2 patients at 400 mg QD. H3K27me3, trimethylation of lysine 27 of histone H3.

ORIC-944 Is Advancing into Combination Development Based on Phase 1 Results

Phase 1b, Multicenter, Open-Label Study

Initial Phase 1b Results

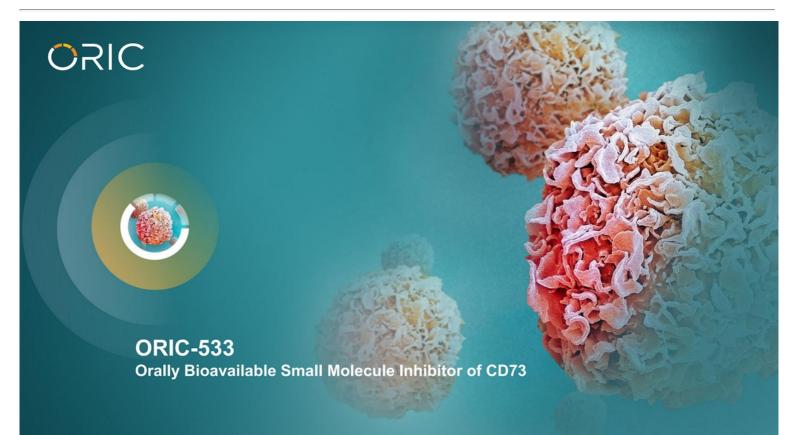
- Potential best-in-class drug properties with half-life >10 hours supporting once-daily dosing
- Robust target engagement demonstrated with maximal decrease (≥75%) in H3K27me3 in monocytes from peripheral blood samples
- Well tolerated to date
 - Only grade 1 and 2 TRAEs at dose levels less than 900 mg QD



Exploratory endpoints: H3K27 trimethylation, PRC2 target gene expression, PSA, and genomics

ORIC-944 demonstrated potential best-in-class drug properties with favorable safety and strong PK profile supporting QD dosing; Combination study with AR inhibitor(s) to be initiated in 1H 2024

Note: All data as of December 10, 2023.



ORIC-533 Is a Potential Best-in-Class Inhibitor of CD73 and First-in-Class for the Treatment of Multiple Myeloma

ORIC-533 Target Product Profile

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CD73 Has Significant Therapeutic Potential in Oncology	 Adenosine is immunosuppressive and impairs antitumor immunity CD73 reverses immunosuppression in preclinical studies CD73 inhibition has demonstrated positive randomized phase 2 data in NSCLC in combination with PD-L1 inhibition 	Promising Phase 1b
ORIC-533 Is a Potential Best-in-Class CD73 Inhibitor	 More potent than benchmark inhibitors in T cell activation assays Orally administered small molecule provides benefits over antibodies (ease of administration and increased tumor penetration) Clean safety profile observed in preclinical toxicology studies 	 ✓ Once daily oral dosing ✓ Clean safety profile ✓ Dose dependent immune activation
ORIC-533 Is First-in-Class for Multiple Myeloma	 CD73 and adenosine role in multiple myeloma supported by research of Dr. Kenneth Anderson lab at Dana Farber Cancer Institute Activity demonstrated in ex vivo bone marrow assays from patients with relapsed/refractory multiple myeloma 	 Clinical activity as single agent in heavily-pretreated multiple myeloma
ORIC-533 demonstra	ted immune activation with an exceptionally clean safety profile, which transla	ated into the first single agent

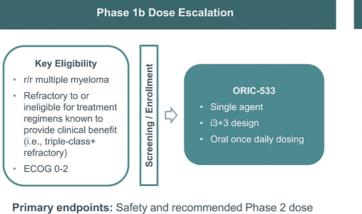


- Once daily oral dosing
- Clean safety profile Dose dependent immune activation
- Clinical activity as single agent in heavily-pretreated multiple myeloma

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Initial First-In-Human Phase 1b Results of ORIC-533 in r/r Multiple Myeloma Were Presented at ASH 2023

Phase 1b, Multicenter, Open-Label Study



Key secondary endpoints: PK

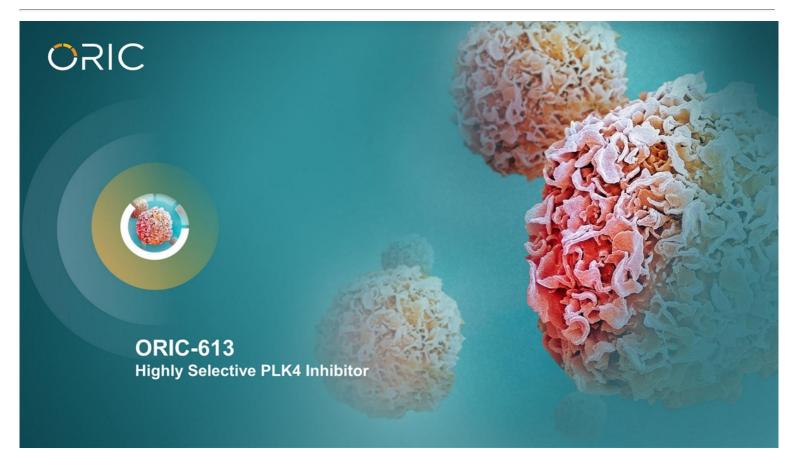
Exploratory endpoints: Exploratory biomarkers⁽¹⁾

Initial Phase 1b Results

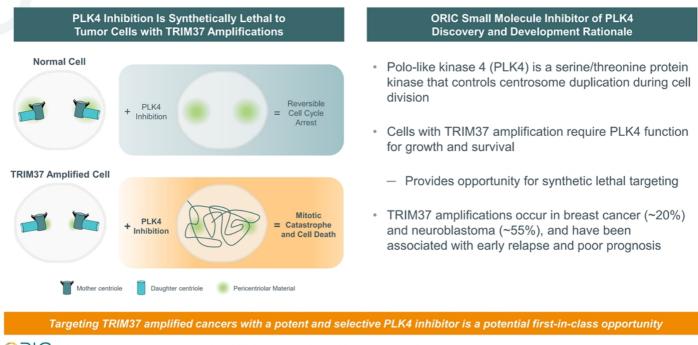
- ✓ Clinical half-life of ~24 hours supports QD dosing
- ✓ Well tolerated safety profile with no Grade ≥3 TRAEs
- Complete/substantial inhibition of CD73 activity in serum and bone marrow
- Evidence of immune modulation of CD8+ T cells and NK cells
- Meaningful reductions in sBCMA levels, suggestive of antimyeloma activity
- Preliminary evidence of clinical antimyeloma activity, including reductions in paraprotein, demonstrated in multiple patients with r/r multiple myeloma

Phase 1b remains ongoing as a monotherapy to select provisional RP2D for combination development; Strategic partnership being pursued to enable combination studies

(1) Exploratory biomarker analyses include CD73 enzymatic function, and immune cell and cytokine profiling. Note: rfr, relapsed/refractory. NK, natural killer. sBCMA, soluble B-cell maturation antigen

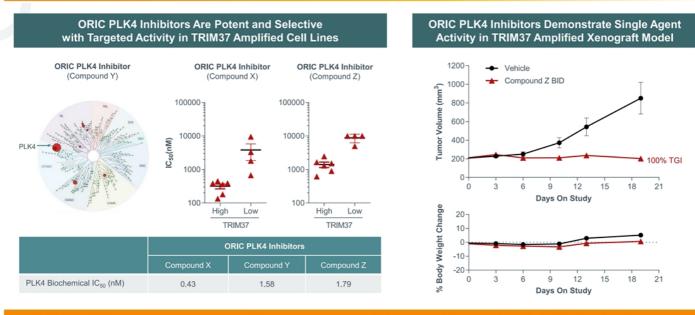


ORIC Is Developing Small Molecule Inhibitors of PLK4 Targeting TRIM37-Amplified Breast Cancer Via Synthetic Lethality



Source: Meitinger et al. Nature (2020), Yeow et al. Nature (2020), Sinclair et al. Breast Cancer Res Treat (2003) and Bown et al. N Engl J Med (1999).

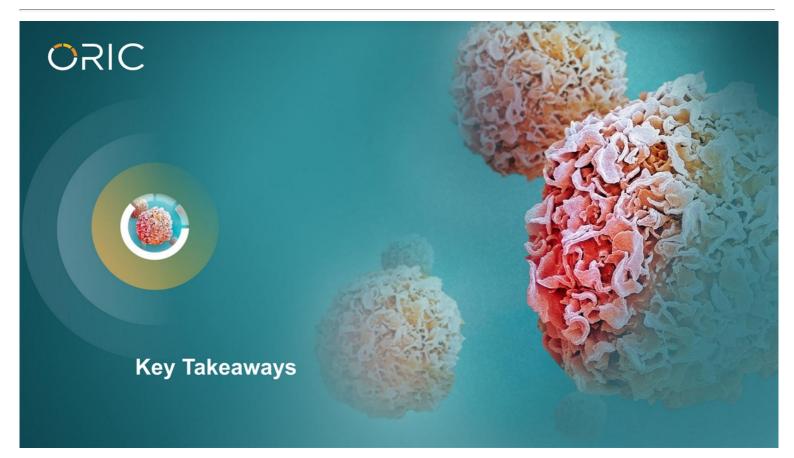
ORIC Small Molecule Inhibitors of PLK4 Are Highly Potent and Selective, and Demonstrate Single Agent Activity In Vivo



ORIC PLK4 inhibitors are highly selective and demonstrate strong single agent antitumor activity in TRIM37 amplified xenografts; Advanced novel development candidate, ORIC-613, through IND enabling studies

ORIC Source: Edgar et al. AACR Poster (2022). Left graphs: Kinome profile at 1 µM. Cell panel consists of breast cancer and neuroblastoma cell lines. Right graphs: oral dosing of ORIC PLK4 inhibitor in CHP-134 neuroblastoma xenograft model

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Clinical Pipeline Focused on Advancement of ORIC-114 and ORIC-944

Program	Indication	Lead Lead Identification Optimization IND Enabling Phase 1 Phase 2 Phase 3	3 Key Differentiation		
PRODUCT CANDIDATE	PRODUCT CANDIDATES				
ORIC-114 EGFR/HER2 exon 20 inhibitor	NSCLC, Breast & Tumor agnostic	Phase 1b: ORIC-114 single agent	✓ CNS active✓ Well tolerated		
ORIC-944 PRC2 inhibitor	Prostate Cancer	Phase 1b: ORIC-944 single agent	✓ Potential best-in-class drug properties		
OUT-LICENSING CAND	OUT-LICENSING CANDIDATE				
ORIC-533 CD73 inhibitor	Multiple Myeloma	Phase 1b: ORIC-533 combination ready	 ✓ Single agent activity ✓ Clean safety profile ✓ Immune activation 		
DISCOVERY RESEARCH PROGRAMS					
ORIC-613 PLK4 inhibitor	Breast cancer		✓ First-in-class potential		
Multiple programs targeting resistance mechanisms	Solid tumors				
	Solid tumors				
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ORIC Pharmaceuticals: Dedicated to Overcoming Resistance In Cancer

Broad Pipeline of Potential First-in-Class and Best-in-Class Programs	 Two potential best-in-class programs advancing towards pivotal studies Additional preclinical programs targeting novel and validated targets 	
Precision Oncology Expertise Enables Accelerated Clinical Timelines	 Rapid timelines enabled by biomarker-driven, patient-selected clinical trials and translational expertise 	
Dual Engines for Pipeline Expansion	 Track record of building pipeline via internal R&D and business development Targeting one new IND candidate every 18 months 	
Experienced Management Team	 Heritage of discovering and developing multiple approved oncology medicines at Ignyta, Medivation, Aragon and Genentech 	
• Cash and investments of \$235 million expected to fund company into 2026 ⁽¹⁾		
 ORIC-944 initiation of combination study with AR inhibitor(s): 1H 2024 ORIC-944 program update: mid-2024 ORIC-114 initiation of dose expansion in multiple cohorts: 1H 2024 ORIC-114 updated Phase 1b data: 1H 2025 		
ORIC (1) Approximate unaudited balance as of December 31, 2023.	43	