



Kazia Corporate Overview

November 2024



NASDAQ: KZIA | X: @KaziaTx

Forward Looking Statements

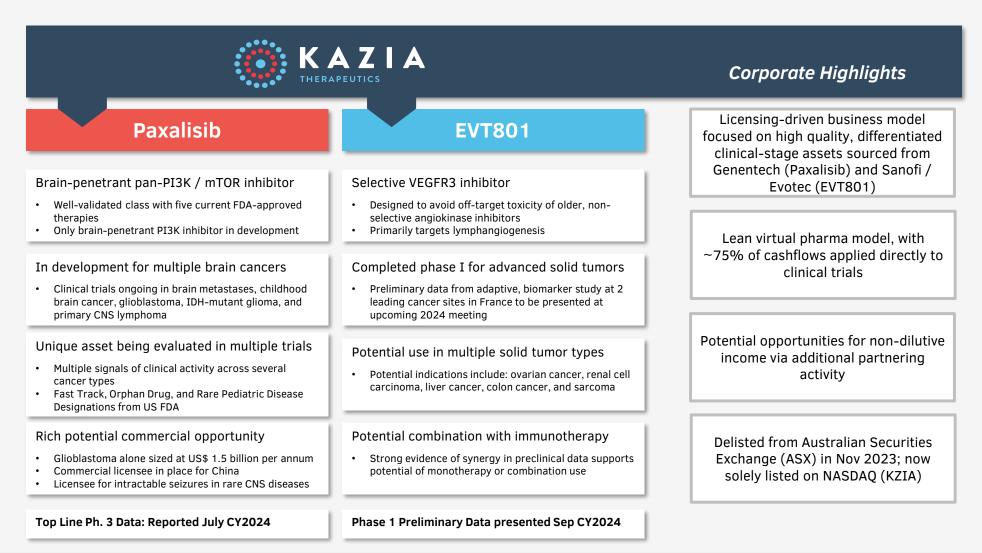
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Company Overview

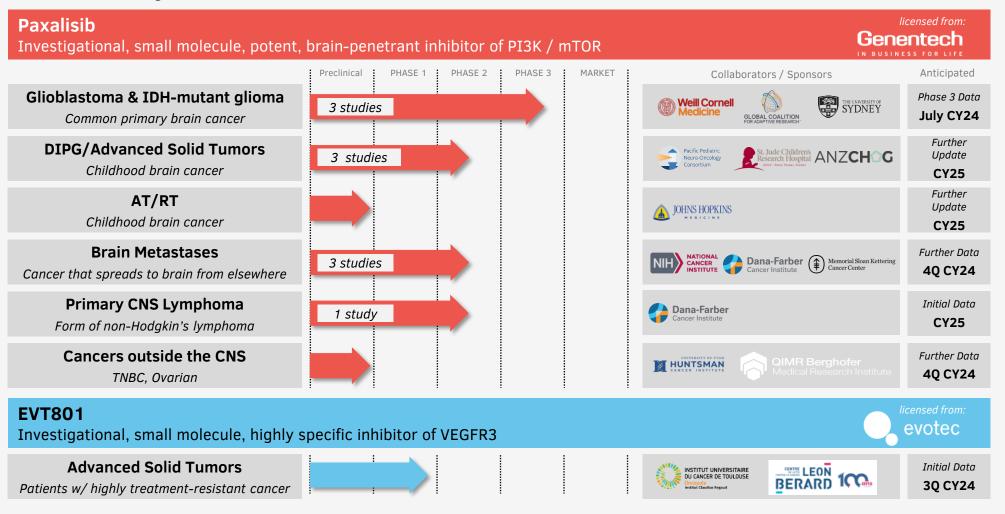
A late-clinical-stage oncology drug development company





Pipeline – Two Differentiated Assets

CY2024 positive clinical data updates driving strong interest in oncology community

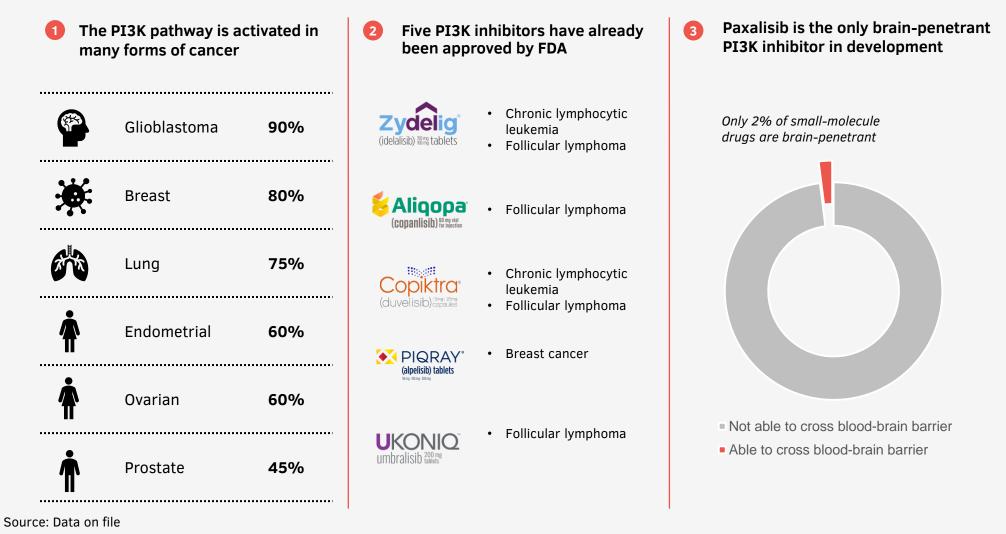


IDH: Isocitrate dehydrogenase, DIPG: Diffuse Intrinsic Pontine Glioma, AT/RT: Atypical Teratoid Rhabdoid Tumor, CNS: central nervous system, TNBC: triple negative breast cancer, VEGFR3: vascular endothelial growth factor receptor 3

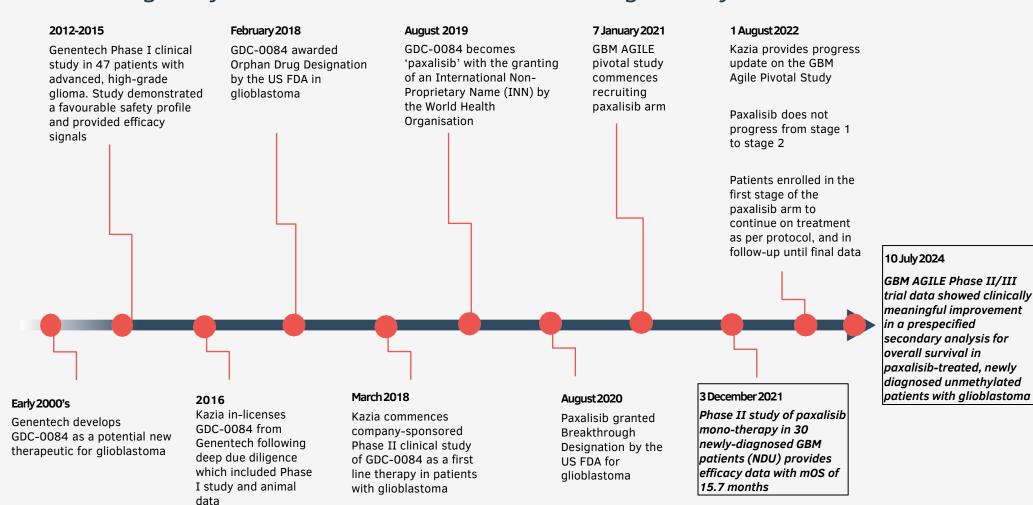


Paxalisib Mechanism of Action

Only brain-penetrant drug in development within the PI3K inhibitor class



Paxalisib – Development History Growing Body of Clinical Evidence Demonstrating Activity in GBM





Glioblastoma Background & Market potential



Glioblastoma Overview

The most aggressive malignant brain cancer





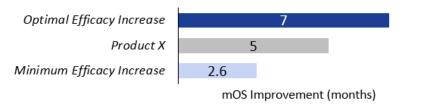
Primary Market Research Outcomes

Physicians indicated a 2-month minimum and 12-month optimum increase in efficacy for newly diagnosed unmethylated GBM treatments, but adoption would be high regardless

Physician receptivity to optimal and minimal mOS efficacy for ND* GBM

(N=15 Physicians)

Optimal and minimal mOS increase in newly diagnosed GBM patients



Adoption rates of Product X if optimal mOS improvement achieved



ND Unmethylated ND Methylated Recurrent 💧 Base Case

Key Takeaway

Due to the high unmet need for a more efficacious therapy for Newly
Diagnosed Unmethylated (NDU)
GBM patients, physicians indicated high adoption rates if Product X
(paxalisib)* is approved by the FDA and achieved their suggested minimum mOS improvement of 2-3 months for newly diagnosed unmethylated GBM patients

Source: Data on file. Company-sponsored market research performed in 2021 * There is no guarantee that the Paxalisib data generated to date will support an FDA approval for commercial use

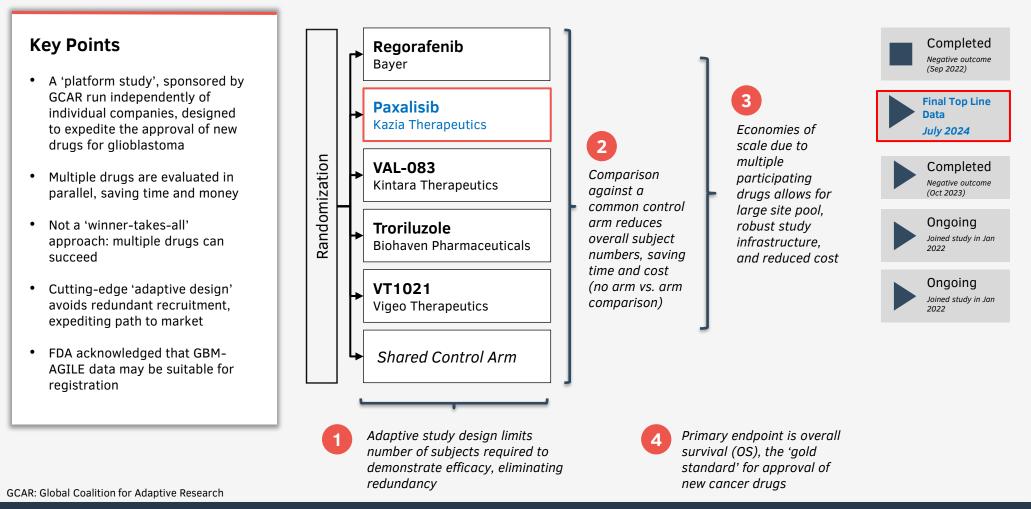


GBM AGILE study data – Primary and secondary analysis



Paxalisib and GBM-Agile

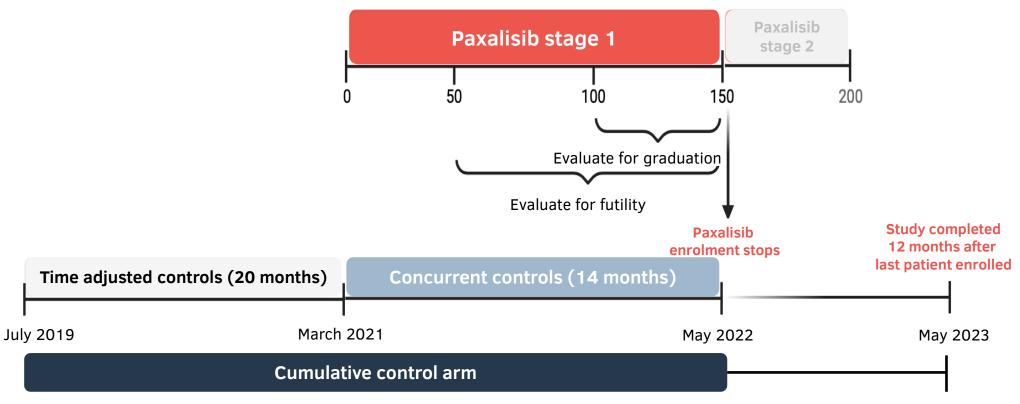
International, multi-center, adaptive, phase 2/3 study evaluating promising therapeutics in patients with glioblastoma





Paxalisib & GBM-Agile

Study schema; Paxalisib arm (n=154) enrolled Newly Diagnosed Unmethylated GBM patients (NDU) and Recurrent GBM patients



Important notes:

- The Cumulative control arm is a combination of Concurrent control patients and the "Time adjusted control" patients that were enrolled in the study before the Paxalisib arm joined the study
- Bayesian Primary Analysis uses data from the Cumulative control arm, while Prespecified Secondary Analysis uses data from the Concurrent Control Arm (i.e., Compares paxalisib data with standard of care)
- All patients (Paxalisib, concurrent control, and cumulative control) were censored on May 2023 if still alive



Paxalisib and GBM-Agile

Newly Diagnosed Unmethylated GBM patients (NDU) Primary endpoint: median Overall Survival analyses (ITT)

Bayesian Primary Analysis: Paxalisib (n=54) vs Standard of Care (n=77)¹ 14.77 months vs 13.84 months Prespecified Secondary Analysis: Paxalisib (n=54) vs Standard of Care (n=46)² 15.54 months vs 11.89 months

A prespecified sensitivity analysis in NDU patients showed a similar median OS difference between paxalisib treated patients (15.54 months) and concurrent SOC patients (11.70 months)

Important notes:

- The Cumulative control arm is a combination of Concurrent control patients and "Time adjusted control" patients that were enrolled in the study before the Paxalisib arm joined the study
- Primary Analysis comparator is the Cumulative control arm, while Prespecified Secondary Analysis comparator is the Concurrent Control Arm
- An efficacy signal was not detected in the recurrent disease population [median OS of 9.69 months for concurrent SOC (n=113) versus 8.05 months for paxalisib (n=100)]

^{1.} Cumulative controls. ^{2.} Concurrent controls

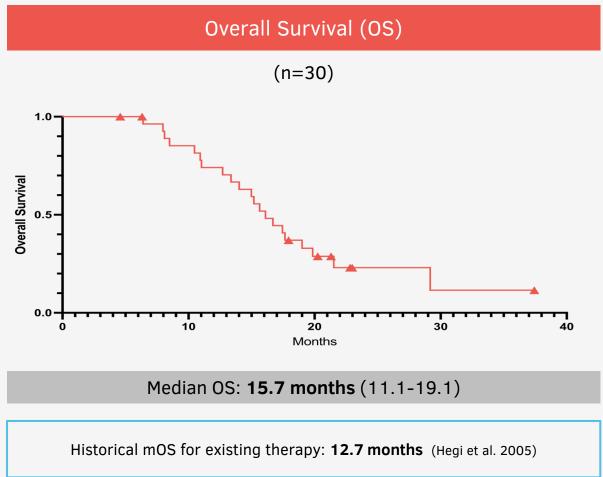


Recap and next steps for Paxalisib in glioblastoma



Paxalisib in Glioblastoma Phase II Clinical Study

Encouraging median OS (mOS) in Newly Diagnosed Unmethylated GBM patients



Note: Figures for existing therapy are for temozolomide, per Hegi et al. (2005); No head-to-head studies have been published



Paxalisib in Glioblastoma Phase II Clinical Study

Encouraging safety profile

Number of Patients at Any Dose (n=30) Experiencing AEs 'Possibly' or 'Likel	ly' Related to Paxalisib (affecting $\geq 10\%$ of patients)
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Term	Gr 1	Gr 2	Gr 3	Gr 4	Total (%)
Fatigue	3	13	2		18 (60%)
Stomatitis	4	7	3		14 (47%)
Decreased appetite	6	6	1		13 (43%)
Hyperglycemia	3	1	6	2	12 (40%)
Nausea	4	6	1		11 (37%)
Rash, maculo-popular	1	1	7		9 (30%)
Diarrhea	7	1			8 (27%)
Vomiting	4	2	1		7 (23%)
Rash	2	4	1		7 (23%)
Neutrophils decreased	3	3		1	7 (23%)
Platelets decreased	6	1			7 (23%)
Weight decreased	5	2			7 (23%)
Lymphocytes decreased	2	3			5 (17%)
Dehydration		4	1		5 (17%)
Dysgeusia		4			4 (13%)
Cholesterol increased	4				4 (13%)
ALT increased	1		2		3 (10%)
Triglycerides increased	1	2			3 (10%)
Malaise	2	1			3 (10%)



Paxalisib in Glioblastoma

Consistent median Overall Survival data in two studies of NDU glioblastoma patients

Compelling Paxalisib data in NDU patients when compared to SOC		Standard of Care data G and STUPP historical co patie	ontrols (right) in NDU
Paxalisib in GBM Agile	Paxalisib in Kazia sponsored phase II study	Concurrent SOC GBM Agile	STUPP historical control
(n=54)	(n=30)	(n=46)	(N/A)
Median OS: 15.54 months*	Median OS: 15.7 months	Median OS: 11.9 months*	Median OS: 12.7 months



*GBM Agile; Prespecified secondary analysis of median Overall Survival

Paxalisib glioblastoma program highlights

Significant body of clinical evidence from two clinical trials supporting paxalisib's activity in newly diagnosed GBM

- ✓ GBM AGILE Phase II/III trial data showed improvement in a prespecified secondary analysis for overall survival (15.5 months) in paxalisib-treated, newly diagnosed unmethylated patients with glioblastoma; presentation of data from GBM AGILE anticipated in 4Q CY 2024
- ✓ Earlier Phase II study of paxalisib as a mono-therapy in 30 newly-diagnosed GBM patients (NDU) provides additional clinical evidence of activity with mOS of 15.7 months

US FDA has granted a December 2024 Type C meeting with Kazia Therapeutics to discuss results and possible pathways to registration

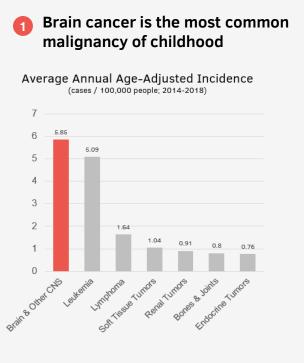


Childhood Brain Cancers



Paxalisib in Childhood Brain Cancer

High unmet need especially in patients with diffuse midline gliomas (DMGs)

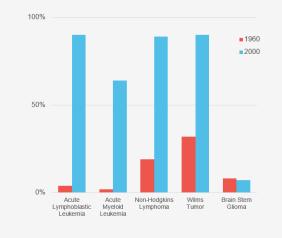


2 Brain cancer represents about one third of childhood cancer deaths

Mortality



Prognosis of childhood brain cancer, especially DMGs, has improved little in recent decades



FDA-Approved Drug The	erapies
Diffuse Midline Gliomas	Nil
Atypical Teratoid / Rhabdoid Tumors	Nil
Medulloblastoma	Nil

Source: CBTRUS; CDC; Ages 0-14 shown; Adamson PC, CA Cancer J Clin. 2015;65:212-220



Summary of Paxalisib in Childhood Brain Cancer

Kazia is actively pursuing three forms of childhood brain cancer

	Diffuse Midline Gliomas (DMG, DIPG)	Atypical Teratoid / Rhabdoid Tumors (AT/RT)	Advanced Childhood Cancer (PI3K/mTOR activated)
Preclinical Research	Positive preclinical data in combination with ONC201	Positive preclinical data as monotherapy and in combination (AACR 2022, 2023, 2024)	Research proposals under discussion
Clinical Trials	Phase I monotherapy clinical trial at St Jude Children's Research Hospital completed	Clinical trial design/execution discussions ongoing between PNOC and Kazia	Additional clinical trial opportunities under discussion for medulloblastoma and HGG
	PNOC022, Phase II clinical trial in combination with ONC201, ongoing		Phase II clinical trial in combination with chemotherapy for treatment of high-risk malignancies commenced 2023
Regulatory Interaction	Orphan Drug Designation (ODD) and Rare Pediatric Disease Designation (RPDD) granted by FDA in Aug 2020	ODD and RPDD granted by FDA in June and July 2022, respectively	Regulatory strategy under discussion

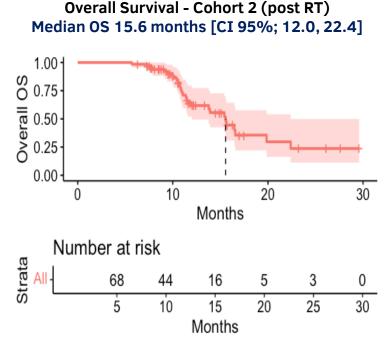


Paxalisib in Diffuse Midline Gliomas

Follow-up Phase II data presented at ISPNO 2024 Annual Meeting

In spite of research that has helped improve treatment for DIPG patients, the prognosis remains poor–with the median survival range being from 8-11 months¹

- 68 patients with biopsy-proven DMG were enrolled in the PNOC Phase II study between November 2021 and June 2023 (median age 9 years [range 3-37], n=41 female [60%])
- Updated Median OS from time of diagnosis was 15.6 months (Confidence interval (CI) 12.0, 22.4)
- Cohort 3 enrolled 30 recurrent patients (in conjunction with radiation therapy) had median OS 8.7 months [CI 95% 8.5, NA]
- Most common grade 3 and above treatment-related adverse events were decreased neutrophil count (n=4); mucositis (n=3); and colitis, drug reaction with eosinophilia and systemic symptoms, decreased lymphocyte count, hyperglycemia, and hypokalemia (n=2)
- Next Steps: Further PK and biomarker analyses ongoing for subsequent cohorts; anticipate clinical update 1HCY2025



Central imaging review analysis of PFS ongoing

1. Hargrave, D., Bartels, U. & Bouffet, E. Diffuse brainstem glioma in children: critical review of clinical trials. Lancet Oncol 7, 241-8 (2006)



Brain Metastases



Paxalisib in Brain Metastasis

MSKCC-sponsored Phase I trial's interim analysis showed encouraging clinical activity of paxalisib in combination with radiation therapy (NCT04192981)

July 2023	February 2024
Fast Track Designation granted by US FDA for paxalisib in combination with radiation therapy in patients with solid tumor brain metastases and PI3K pathway mutations	Announced early conclusion , based on Stage 2 positive safety data and promising clinical response findings observed to date.
	Fast Track Designation granted by US FDA for paxalisib in combination with radiation therapy in patients with solid tumor brain metastases and

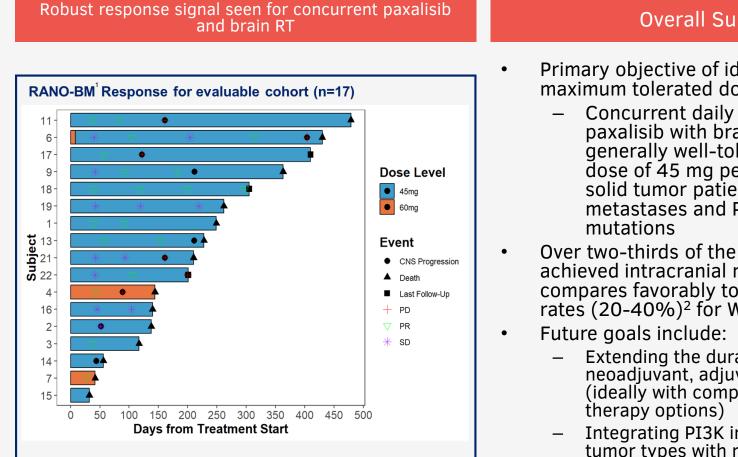
All 9 patients evaluated for efficacy exhibited a clinical response, according to RANO-BM criteria, with breast cancer representing the most common primary tumor Based on the interim Stage 1 data from the MSKCC-sponsored Phase I trial's interim analysis.

- Data presentation anticipated at upcoming scientific congress in 2H CY2024
- Coordinate and plan next clinical study in conjunction key thought leaders and FDA



Paxalisib in Brain Metastasis

MSKCC-sponsored Phase I trial's interim analysis presented at 2024 ASTRO^{*} meeting showed encouraging clinical activity of paxalisib in combination with radiation therapy (NCT04192981)



* American Society for Radiation Oncology

1. Response assessment in neuro-oncology brain metastases (RANO-BM)

2. Zhou et al. 2021, Kim et al. 2020

Overall Summary

- Primary objective of identifying the maximum tolerated dose (MTD) was met:
 - Concurrent daily administration of paxalisib with brain radiotherapy was generally well-tolerated at a maximum dose of 45 mg per day in advanced solid tumor patients with brain metastases and PI3K pathway
- Over two-thirds of the patients at MTD achieved intracranial response which compares favorably to historical response rates (20-40%)² for WBRT alone
 - Extending the duration of PI3K inhibition, neoadjuvant, adjuvant and maintenance (ideally with complementary systemic
 - Integrating PI3K inhibition with CNS tumor types with relevant pathway driver mutations and potentially SRS



Other Solid Tumors



Paxalisib in Triple Negative Breast Cancer

QIMR Berghofer Medical Institute collaboration

"In treatment-resistant pre-clinical models of breast cancer, paxalisib (4T1 mouse model, TNBC¹) has shown encouraging results in inhibiting both the primary tumor burden and metastasis by reinvigorating the immune system within the tumor microenvironment" – Professor Sudha Rao, Group Leader, QIMR Berghofer

- Leading transcriptional biology and epigenetics expert, Prof Rao identified an entirely novel effect of PI3K inhibition:
 - Immune modulator of the tumor and the surrounding microenvironment
 - Administration of PI3K inhibitors such as paxalisib, at doses and frequencies different to those conventionally used, appears to activate or reinvigorate the immune system in the tumour, making it more susceptive to immunotherapy
- The preliminary data from our collaboration will be presented at an upcoming conference in 4Q CY2024

Paxalisib Combination Combination influence on Paxalisib + Paxalisib + immune system LYNPARZA® **KEYTRUDA®** Intellectual (example, T cells, (olaparib) data in (pembrolizumab) **B** cells, NK cells) Property (IP) advanced breast data in TNBC¹ and within the update cancer preclinical tumor and its preclinical models micromodels environment

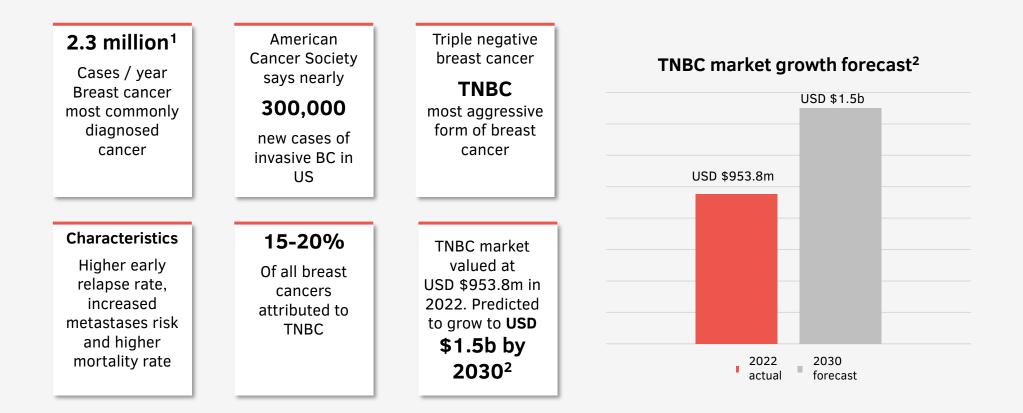


1. Triple Negative Breast Cancer



Triple Negative Breast Cancer Treatment Landscape

Projected TNBC market to exceed \$1.5 Billion by 2030



1. National Institutes of Health (NIH): Current and future burden of breast cancer: Global statistics for 2020 and 2040

2. https://www.databridgemarketresearch.com/reports/global-triple-negative-breast-cancer-market

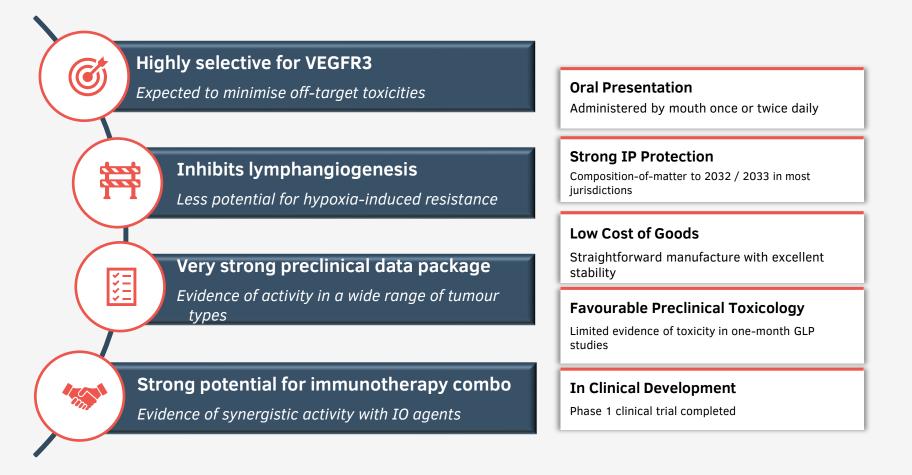


EVT801





EVT801 is a highly selective VEGFR3 inhibitor, primarily inhibiting lymphangiogenesis (formation of new lymphatic vessels)

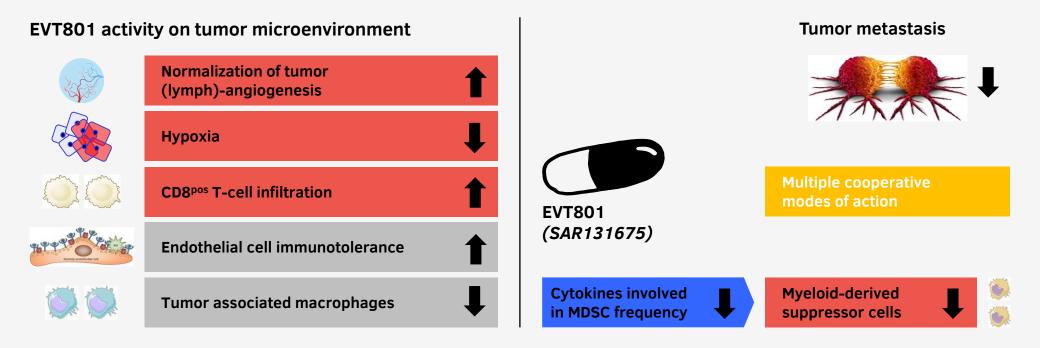




EVT801 Mechanism of Action

By targeting VEGFR3^{pos} tumor blood vessels, EVT801 may induce tumor blood vessel normalization, reduce hypoxia, and improve CD8 T-cells infiltration

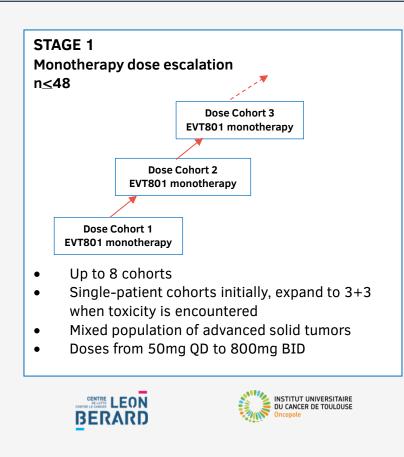
Schematic overview based on pre-clinical data





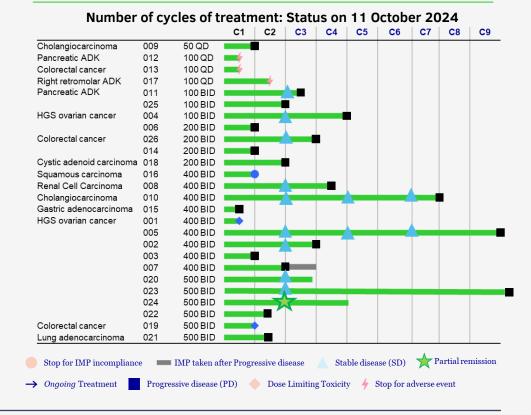
Data from Tacconi & al. with SAR131675

EVT801: Phase 1 dose-finding trial; KZA 0801-101 (NCT05114668) Staged development in patients with advanced cancer



Phase 1 study in advanced cancer patients Completed

- Primary objective of stage one of the study was successfully met:
 - MTD has been reached at 500mg BID
 - The recommended dose for phase 2 is 400 mg BID* in continuous monotherapy administration



MTD = Maximum Tolerated Dose; RP2D = Recommended Phase 2 Dose

Human active dose prediction based on predicted human clearance of 2.5 mL/min/kg: 375 mg BID $\,$



EVT801 Key Points

- 1 Well-understood mechanism (anti-angiogenesis) with unique differentiating feature (high VEGFR3 selectivity)
- 2 Strong preclinical data package, with observed activity in multiple tumours and favourable toxicology
- **3** Potential for combination use with immuno-oncology therapies
- A Phase 1 completed demonstrating encouraging safety and tolerability profile to date:
 - Clinical and biomarker data presented at AACR Ovarian Cancer Research Symposium September 2024
 - Primary and secondary objectives successfully met, with MTD and RP2D identified
 - Encouraging signal of activity observed in High Grade Serous (HGS) ovarian cancer as well as strong VEGFR3 biomarker expression
- 5 Next clinical trial under discussion with scientific thought leaders:
 - Consolidate safety data at RP2D and our hypotheses on EVT801 mode of action
 - Validate HGS ovarian cancer as indication of choice for clinical trial phase 2 as monotherapy or in combination with standard-of-care (ex. PARPi)



2024 Corporate Update



Paxalisib Licensing and Collaborations

Opportunistic partnering and strategic collaborations continue to add

Licensing			Key Collaborations
Summary	Simcere	DI Sovargen	QIMR Berghofer Medical Research Institute
Territories and responsibilities	To develop and commercialize Paxalisib in Greater China, Hong Kong, Macau, and Taiwan	To develop, manufacture and commercialize Paxalisib as a potential treatment for intractable epilepsy in focal cortical dysplasia type 2 (FCD T2) and tuberous sclerosis	Cutting edge preclinical program to evaluate Paxalisib in combination with immuno-therapies for Advanced Breast Cancer
Linforent normant	US\$11m, comprising	complex (TSC) disease	JOHNS HOPKINS
Upfront payment	US\$7m in cash and a US\$ US\$1.5 million 4m equity investment		
Milestone payments	Contingent milestone payments of up to US\$ 281 million in GBM + further milestones payable in indications beyond GBM	Potential milestone payments of up to US\$19 million upon the achievement of development and regulatory milestones	 Paxalisib alone and in combination with other targeted agents is active in preclinical models of AT/RT¹ US FDA has awarded Orphan Drug Disease and Rare Pediatric Disease Designations in AT/RT
Royalties on net sales	Mid-teen percentage royalties on commercial sales	A percentage of sub- licensing revenues and royalties on net sales of products incorporating paxalisib	 If Paxalisib were to be approved, Kazia could be entitled to receive a pediatric priority review voucher which are tradeable and have historically commanded prices in excess of USD \$100 million.



1. Atypical Teratoid Rhabdoid Tumor

Kazia Therapeutics: 2024-2025 Corporate Focus

Objectives for value creation

Progress paxalisib glioblastoma program	 Compile data from all clinical trials FDA granted Type C meeting with Kazia in December 2024 Propose potential pathways to registration
Execute paxalisib pediatric and brain metastasis programs	 PNOC team to complete PK/biomarker data analysis and provide update 1Q CY2025 Additional data presentation and advance development to evaluate Paxalisib + Radiation Therapy 4Q CY2024
Paxalisib in other key oncology indications	 Advance TNBC¹ program stemming from QIMR collaboration whereby encouraging signals of immune reinvigoration and cancer stem cell activity have been consistently observed in animal models
EVT801 program	 Complete analysis Stage one of EVT801 Phase 1 clinical study Data presented at AACR Ovarian Cancer Research Symposium, September 2024 Discuss and plan for Phase 2 study in advanced ovarian cancer patients
Corporate business development	 Continue to be opportunistic in terms of global and regional licensing for paxalisib and EVT801







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