

Developing high value drugs for challenging diseases

Tim Oldham PhD, CEO and Managing Director July 2022



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AdAlta at a glance



Building out pipeline

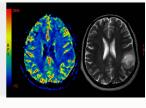
Targeting 10 programs by end 2023



Immuno-oncology: two co-development collaborations



- 1. i-CAR-T with Carina Biotech: US\$20b market by 2028
- 2. GZMB i-PET imaging agent with GE Healthcare: US\$6.4b market





Fibrosis/inflammation: wholly owned pipeline

- 1. AD-214 first in class anti-fibrotic preparing for Phase II clinical trials
- 2. Second target in discovery





i-body platform

Powerful drug discovery tool for creating drugs against diseases underserved by traditional antibodies



Four human health needs AdAlta is addressing today



Antibodies cannot do everything!

AdAlta's i-bodies are a differentiated drug discovery platform for difficult diseases



Idiopathic Pulmonary Fibrosis: degenerative, fatal

AdAlta's AD-214 could meet a desperate need for new approaches for a debilitating disease



CAR-T cell therapy providing new hope ... for blood cancer patients

AdAlta and Carina's iCAR-T cells could offer same hope for patients with solid tumours



Immuno-oncology drugs revolutionising cancer treatment ... for some

AdAlta and GE Healthcare's GZMB iPET imaging agent could identify responders early



Platform: i-bodies



i-bodies: sdAB-like molecules with engineered binding loops conferring unique binding properties



AdAlta i-bodies are combination of a human protein with unique long loop binding sites that mimic the structural features of the shark single domain antibody system







Human protein scaffold

Engineered target specific binding loops



AdAlta i-body library contains 10¹⁰ unique i-bodies. Each unique i-body has different binding loops



12-15kDa protein 90% smaller than MAb 50% smaller than scFv

Naked i-body



molecules

i-bodies allow for high affinity, high specificity binding to targets that are intractable for traditional antibodies

hindered cell membrane

receptors

Small Antibodies i-bodies Flexible, modular formats Molecules CAR cell therapy ADC/ radiotherapeutic Bi-specific Fc-fusion i-bodies are ~10% the i-bodies have high The i-body CDR specificity, size of human structure confers **PEGylation** avoiding off-target antibodies, capable of unique binding issues of small capabilities, enabling engaging sterically

unique epitope engagement and

tunable pharmacology



Lead program: AD-214



About | Idiopathic Pulmonary Fibrosis (IPF)

Scaring and stiffening of the lungs progressively and irreversibly reduces lung function

>300,000 people living with IPF; 40,000 people die from IPF every year

Only 3.8 years median survival after diagnosis

Two current therapies sell for \$3b per year ...

... despite having limited effectiveness and serious side effects

Burden of fibrotic lung disease following COVID-19 likely to be high

"Long COVID" is a developing issue – further increasing the need for better anti-fibrotic drugs.*

^{*} PM George, et al, "Pulmonary fibrosis and COVID-19: the potential role for antifibrotic therapy", Lancet published online May 15, 2020.



AD-214 | Completed Phase I, multiple indication options

AD-214 is a first in class anti-fibrotic

- i-body-Fc fusion targeting CXCR4
- Pre-clinical efficacy in lung, kidney, eye fibrosis

Phase I intravenous (iv) clinical study successfully completed¹

- AD-214 (iv) is well tolerated, binds CXCR4 tightly
- Preclinical animal data supports potential iv efficacy

Drug substance manufacturing secured for next clinical studies²

- Delivery late 2023
- Next clinical studies to commence first half of 2024³



Multiple indication and partner potential

- US\$ billion need in many fibrosis indications
- Unique formulations expands partnership options



Lung
IPF/ILD
>US\$3b
Inhalation



Eye Wet-AMD >US\$15b Intravitreal



Kidney
RENAL FIBROSIS
>US\$10b
Intravenous

Preclinical development of inhaled formulation for IPF well advanced

Route of administration, indication priorities for clinical program to be finalized in 2H 2022



An inhaled formulation of AD-214 would be a superior option for IPF







Improved bioavailability

- AD-214 delivered direct to fibrotic areas
- First pass liver clearance¹ avoided
- Dosing schedule flexibility to optimise receptor coverage

Greater patient convenience

- Self administration (no scheduled clinic visits; freedom of movement)
- Less invasive

Enhanced cost effectiveness

- Lower drug dose means lower cost of goods
- Lower healthcare costs for administration

^{1.} ASX Release 19 July 2021; not observed for i-bodies to other targets; these studies were part supported by a Biomedical Translational Bridge grant, a program of Australia's Medical Research Future Fund administered by MTPConnect and supported by UniQuest

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Inhaled AD-214 | Milestones and opportunities

September 2022 quarter

- Antifibrotic effects in cultured human lung tissue
- Imaging distribution, retention in sheep
- Efficacy in bleomycin mouse model*

December 2022 quarter

- Inhaled AD-214 formulation selected
- Preparation for inhalation toxicology studies

First half 2023

- Manufacturing AD-214 for toxicology studies
- cGMP manufacturing commences for Phase lb/lla clinical trials

Aim of pre-clinical studies

- 1. Demonstrate nebulised AD-214 can reach lower airways of sheep lungs (similar to human)
- 2. Demonstrate that AD-214 reaching the lower airways is retained in fibrotic tissue (bleomycin mice, sheep, cultured lung tissue)
- 3. Demonstrate AD-214 delivered to fibrotic tissue can moderate disease progression (bleomycin mice, cultured lung tissues and cells)

Pre-clinical success anticipated to accelerate existing partnering discussions



AD-214 | Valuable IPF partnering options as early as Phase I

Date	Licensee	Licensor	Transaction Terms	Clinical Phase	
Nov-21	BLADE O THERAPEUTICS	BIOTECH ACQUISITION COMPANY	US\$254m Upfront	2 (Ready)	
Nov-21	OncoArendi Therapeutics	Galápa gos	€320m Milestones	2 (Ready)	
Sep-21	Syndaxૐ	Incyte	US\$152m Upfront +US\$602m Milestones	2 (Ready)	
Nov-19	Promedior	Roche	US\$390m Upfront +US\$1b Milestones	2	
Feb-21	東德制药 TIDE PHARMACEUTICAL	G R AVIT Q N	US\$517.5m Milestones 1		
Jul-19	bridgebio	Boehringer Ingelheim	€45m Upfront +€1.1b Milestones	1	



Co-developed immuno-oncology assets



About | CAR-T therapies

CAR-T therapies are providing new hope for patients with cancer who have failed all other options

Therapy involves removing immune cells from blood and re-engineering them so they "see" cancer as a pathogen

Already 6 FDA-approved CAR-T therapies ... but so far only for blood cancers

>\$US1 billion earned by CAR-T therapy products in 2020

\$US20.3 billion¹ revenue forecast for 2028 as more products are commercialised, science evolves

Solid tumours to account for >50% of CAR-T revenues by 20302

^{1.} Grandview Research, "T-cell Therapy Market Size, Share & Trends Analysis" Feb 2021 2. Polaris Market Research, "CAR-T Cell Therapy Market Share, Size Trends, Industry Analysis Report", June 2021

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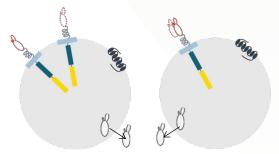


i-bodies enable optimized CAR constructs (i-CARs)

Feature	Benefit		
Small size	Increased CAR gene cassette/vector capacity, efficient multi-functional CAR cell creation		
Long CDR3 binding domain	Access to unique tumor antigens/epitopes and TME modulating proteins in cancer tissue		
Tunable binding	Control of immune synapse (length + strength)		
Robust conformation	Natural stability delivers robust CAR binding domain and stable secreted molecules		

Superior i-CAR products

- CARs against novel tumor antigens
- Dual and bi-specific CARs for enhanced specificity, reduced tumor escape and logic gated CARs
- · Secreted antibodies to modulate TME



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i-CAR-T assets | Carina co-development collaboration

AdAlta and Carina are combining i-bodies and a world class CAR-T platform to create iCAR-Ts that could offer improved precision, performance and persistence

Further expands AdAlta's pipeline in an attractive deal space

- Collaborating on up to five tumour targets
- Sharing costs to pre-clinical proof of concept (in mice)
- Jointly own resulting products: ready for partnering or further development

Current status

- i-body enabled CAR-T (i-CAR-T) cells have been successfully generated by Carina and demonstrate in vitro cell killing (lysis)¹
- First target A selected, A-i-CAR-T cells incorporating i-bodies against Target A being built



immuno-oncology asset

About | Immuno-oncology (I/O) PET imaging

Immuno-oncology (I/O) drug market is worth US\$95 billion¹...

... but only 20-40% of patients respond² to therapy

Granzyme B (GZMB) is produced by immune cells to kill cancer: potential biomarker of I/O drug activation of the immune system

PET imaging GZMB could help identify early who will – and won't – respond to I/O drugs

The PET imaging agent market is valued at US\$6.4billion³

Largest products >US\$400m⁴

^{1. 2026} forecast by ResearchandMarkets.com, Immuno-Oncology - Market Analysis, Trends, Opportunities and Unmet Needs - Thematic Research, March 2021 2. P Sharma, et al, Cell 168(4) 707 (2017) 3. 2027 forecast by Global Industry Analysts, Imaging Agents: Global Market Trajectory and Analytics, April 2021 4. AD Nunn, J Nucl Med (2007) 169



GZMB i-PET imaging asset | GE Healthcare co-development collaboration

AdAlta and GE are co-developing a GZMB i-body PET imaging (iPET) asset to evaluate the effectiveness of immuno-oncology drugs

Revenue generative pipeline asset

- AdAlta earns research fees, development and sales milestone payments and royalties on product sales
- A\$2.27 million revenue* earned to December 2021
- GZMB iPET asset could generate royalty revenue sooner than a therapeutic due to shorter diagnostic development timelines

Current status

- Panel of GZMB specific i-bodies identified
- Pre-clinical proof of concept studies and i-body optimization underway
- Manufacturing development underway







Co-developed iPET imaging immuno-oncology asset.

^{*} Milestones, research fees and contributions to third party costs; AdAlta Half Year Report 23 Feb 2022



I/O assets | Milestones and opportunities



September 2022 quarter

 Initial in vitro cancer cell killing screening assays for A-i-CAR-T

 i-CAR-T targets B and C selected December 2022 quarter

First half 2023

 Full in vitro cancer cell killing assays complete for A-i-CAR-T A-i-CAR-T in vivo proof of concept studies commenced



TBD

 Updates provided in consultation with GEHC and as milestones are achieved



Corporate snapshot

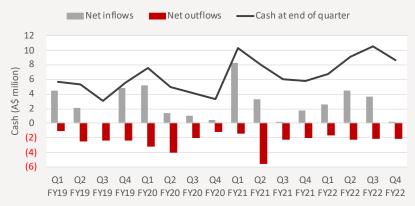
ASX code	1AD
Market capitalisation	A\$15.08m
Share price (12 month closing range)	A\$0.048 (\$0.045 - 0.114)
12 month return	(58)%
Ordinary Shares (daily volume)	314,184,746 (277,395)
Unlisted Options	14,184,060
Cash (30 Jun 2022)	A\$8.66m

Major shareholders (21 July 2022)	%
Yuuwa Capital LP	17.2
Platinum Asset Management	15.7
Meurs Holdings Pty Ltd	6.4
Radiata Super Pty Ltd	3.5
Sacavic Pty Ltd	3.1
Other (1,472 total holders)	54.1
Total	100%

Analyst Coverage				
Lodge Partners				
Louge Faithers				



Quarterly cash flows (A\$ million)





Investment proposition



i-body platform to create value



Fibrosis/inflammation
Lead asset advancing to Phase II
>\$3b market potential in first indication¹

Discovery initiated on 2nd target



Immuno-oncology
2 x co-development collaborations to leverage platform

✓ Carina Biotech: \$20b CAR-T market²

✓ GE Healthcare: \$6b PET market³



Leading expertise



Clear vision for growth through pipeline expansion



Regular near-term news flow

^{1.} GlobalData, Idiopathic Pulmonary Fibrosis Opportunity Analysis and Forecasts to 2029, November 2020 2. 2028 forecast by Grandview Research, "T-cell Therapy Market Size, Share & Trends Analysis" Feb 2021 3. 2027 forecast by Global Industry Analysts, Imaging Agents: Global Market Trajectory and Analytics, April 2021



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