FivepHusion

Investor Presentation
December 2025

OPTIMISED CANCER THERAPEUTICS

A DE-RISKED DRUG DEVELOPMENT OPPORTUNITY TARGETING GLOBAL MARKETS

Dr Christian Toouli, CEO & Managing Director
c.toouli@fivephusion.com

DISCLAIMER

By accepting this presentation or attending a presentation (whether in person or by video conference), the recipient (**you**) agrees to be bound by the following terms and conditions. The purpose of this presentation is to provide general information about Detsamma Investments Pty Ltd (ABN:85 630 579 547) (trading as **"FivepHusion"**) and its business.

This presentation does not constitute an offer, invitation or recommendation to subscribe for, or purchase, any security or financial product or an advertisement for the same. This document is not a prospectus, disclosure document, product disclosure statement or other offer or disclosure document under Australian law or the laws of any other jurisdiction. This presentation does not constitute the provision of financial product or investment advice. You should seek independent legal, financial, regulatory and taxation advice before making any decision in respect of this presentation. You represent and warrant that you fall within one or more of the categories of "sophisticated" or "professional" investors as described in sections 708(8) and 708(11) of the Corporations Act 2001 (Cth), or that you are otherwise a person to whom this presentation may be provided under the securities laws of any applicable jurisdiction.

FivepHusion and its officers, employees, consultants and advisors do not make any guarantee, representation or warranty, either expressed or implied concerning the accuracy, completeness or reliability of the information contained in this presentation or accept any responsibility for errors or omissions in any information contained in this presentation or the future performance of FivepHusion; and disclaim and exclude all liability for the accuracy, completeness or reliability of the information contained in this presentation (or any error or omission thereof), and exclude all liability whatsoever for any loss, damage or costs (whether foreseeable or not and whether direct, indirect or consequential) which may be suffered by any person as a consequence of any information in this presentation or any error in or omission from it, whether the loss or damage arises in tort (including negligence), contract, statute or otherwise. Specifically, FivepHusion does not warrant or represent that the information contained in this presentation has been audited, except where specifically stated otherwise.

This presentation may include forward-looking statements, which reflect various assumptions that may or may not prove to be correct. Actual results may be materially affected by known and unknown risks including changes in economic conditions and other circumstances which may be outside the control of FivepHusion. The reliance that recipients place upon such information is a matter for their own commercial judgement however you are cautioned not to place undue reliance on forward-looking statements and to seek professional advice if in doubt.

All warranties, conditions, liabilities or representations in relation to information or advice contained in this presentation are expressly negated and excluded to the maximum extent permitted by law. The recipient agrees, to the fullest extent permitted by law that he/she shall not seek to sue or hold FivepHusion or its officers, employees, consultants and advisors liable in any respect by reason of provision of this presentation.

This information in this presentation is being furnished to you solely for your information and may not be reproduced, in whole or in part, or distributed to any other person, without the consent of FivepHusion.

Millions of cancer patients are treated with chemotherapy **unchanged since last century**

FivepHusion **is optimising treatment** safety and efficacy, & **unlocking multi-billion-dollar** commercial opportunities

Executive Summary

Optimising the standard of care, backbone of cancer treatment

FivepHusion

Deflexifol®: A next-generation, best-in-class treatment

A new & optimised standard of care therapy

- Co-formulation of **5-fluorouracil (5-FU)** & its biomodulator **leucovorin (LV)**
- Positioned to **replace standard therapy** in solid tumours
- Primary indication of **1st line metastatic colorectal cancer**
- Sales revenue potential **≥US$1B**

Broad therapeutic utility & market opportunities

- High need indications such as **paediatric brain cancers**
- **Significant upside** potential in other solid tumours
  - pancreatic, gastric, breast, head & neck cancers

Technically low-risk & clinically advanced

- **3 clinical studies successfully completed**
- **5x surrogate pII trials** support increased survival benefit
- **Fast-tracked, low-risk 505(b)(2) regulatory pathway –** market launch as early as 2029
- **Low-cost, scalable manufacture with Pfizer** in Melbourne, accessing global expertise + supply chains
- **Endorsed by leading oncologists**
- **Granted composition of matter IP** + patent pipeline
- Prospective opportunity to leverage an invited **$5M non-dilutive** Industry Growth Partnership grant proposal

An IPO will position FivepHusion to reach two registration trials with strong partnering appeal

Experienced Leadership & Strategic Partners

Established highly experienced Board, Management and Advisory Teams

Board

David Ranson
Executive Chairman
BEng(ElecEng)

Dr. Christian Toouli
CEO & Managing Director
Btech Hons; PhD; GAICD

Dr. Bill Ketelbey
Executive Director
MBBCh; FFPM; MBA; GAICD

Iain Ross
Non-Executive Director
BSc Hons; CDir (IoD)

Strategic Collaborations

Syneos Health · Pfizer CentreOne · allarity THERAPEUTICS · University of Wollongong Australia · University of South Australia · Monash University · Medicines Manufacturing Innovation Centre · FB Rice · The IP Navigators · Children's Cancer Institute

Independent Clinical Advisory Board

Advising on the clinical strategy and trial design for Deflexifol® registration for use in adult cancers

Prof. Stephen Clarke OAM
Chairman
GenesisCare · University of Sydney

Prof. John Simes AO
University of Sydney · NHMRC Clinical Trials Centre

Prof. Andrew McLachlan AM
University of Sydney · ANZAC Research Institute

Prof. John Zalcberg AO
ICON · Monash University · All.Can

Founder Advisory Board

Inventors of Deflexifol® contributing expertise to ongoing development

Prof. Philip Clingan OAM

Senior Prof. Marie Ranson

Emeritus Prof. John Bremner AM

Multiple Blockbuster Opportunities

Deflexifol® & pipeline opportunity ResectAssist™ have broad applications in solid tumours

Deflexifol®

Metastatic Colorectal Cancer

Entering Phase Ib/IIa
Primary endpoints: safety & maximum tolerated dose
Secondary Endpoints: Efficacy

Thesis

Deflexifol® to replace backbone 1st line therapy: 5-FU & LV

Blockbuster Global Market

1.9m
cases per annum
(20-30% metastatic[1])

930k
deaths per annum[2]

Paediatric Ependymoma

Entering Phase II
Primary endpoint: Efficacy (response rate)
Secondary Endpoints: Survival

Thesis

Deflexifol® to become the first approved therapy

Orphan Disease

3rd Most Common
Brain cancer in children

23 - 45%
5-year progression-free[3]

Potential Indications

Pancreatic Cancer
Gastric Cancers
Breast Cancer
Head & Neck Cancers

Thesis

Optimise treatment across other 5-FU & LV indicated solid tumours.

All present
Blockbuster
Markets

ResectAssist™

Solid Tumours

Initial Focus

Downstaging pancreatic cancer tumours to resectable with curative intent

Thesis

A novel drug delivery technology platform:

facilitating intra-tumoral delivery of approved (FOLFIRINOX) and innovative drugs

Lead indication:
Pancreatic Cancer
>$7.0B
market opportunity[4]

1. Global Cancer Observatory 2020, Cancer Today; GLOBOCAN 2020
2. https://www.who.int/news-room/fact-sheets/detail/colorectal-cancer
3. https://pmc.ncbi.nlm.nih.gov/articles/PMC10036929/
4. Polaris Market Research 2022 – pancreatic cancer market in 2030

DEFLEXIFOL®

Combining and Optimising the Current Standard of Care

FivepHusion

Metastatic colorectal cancer (mCRC)

Treated palliatively, with up to only ~55% response rate & ~30-month survival

5-fluorouracil (5-FU) + leucovorin (LV)
are the **"backbone"** of mCRC therapy

~95% of patients receive 5-FU/LV

The treatment backbone for the foreseeable future[1]

X The Problem with 5-FU + LV

5-FU + LV is synergistic, but chemically incompatible

- **Synergy:** LV enhances the efficacy of 5-FU
- **Chemically Incompatible**: Cannot be co-administered to maximise efficacy (crystallises and blocks the infusion line)

Sequential administration (current workaround) provides:

- **limited co-exposure and**
- **sub-optimal efficacy**

✓ The Solution: Deflexifol®

FivepHusion's Breakthrough: Deflexifol®

- **Deflexifol® successfully combines 5-FU + LV**
- **Overcomes chemical incompatibility**
- **Increases co-exposure from 3 hours → 47 hours**
- **Delivers new highly valuable composition of matter IP**

↑ **Enhanced Efficacy**

↓ **Reduced Toxicity**

↑ **Higher Tolerated Dose**

[1] According to KOL opinion & competitive landscape analysis, and as reviewed by Glimelius *et al.* 2021, *Cancer Treat Rev* 98:102218.

CONFIRMED IMPROVED SAFETY AND EFFICACY

FivepHusion's two clinical trials demonstrated safety and efficacy signals

FivepHusion has treated 59 end-stage patients with a variety of solid tumours demonstrating[1]

- Reduced toxicity and improved tolerability
- Effective disease control in the majority of patients despite failing all prior therapies (including 5-FU)

Supported by **five independent phase II studies[2]** demonstrating improved anti-tumour activity and significant survival benefits

Deflexifol® monotherapy

64-69% disease control rates

Across two dose-escalation studies in **end-stage, heavily pre-treated** patients

vs.

Approved mCRC monotherapies

41% disease control rate

for regorafenib (**US$580M** sales in 2023)

&

44% disease control rate

for Lonsurf® (**US$550M** sales in 2023)

[1,2] Deflexifol® publications, conference proceedings and supportive literature can be found at: https://fivephusion.com/publications/

DEFLEXIFOL® CASE STUDIES

Efficacious after 5-FU + LV failure in end-stage cancer patients

FivepHusion

Heavily pre-treated patients experienced benefit from optimised 5-FU/LV delivery

Activity after repeated failure of treatment with the same drugs - Indicates Deflexifol® superiority

Phase Ib/IIa trial^ Demonstrated

Disease control:
9/13 (69%) evaluable patients

Median progression free survival:
28.2 weeks

Metastatic Colorectal Cancer

Patient: male, 59 years

Failed two lines previously:

- FOLFOX
- FOLFIRI + bevacizumab

Treatment: Deflexifol®

- 525 mg/m^2 bolus + 3000 mg/m^2 infusion

Result: Stable Disease 5 months

Pancreatic Cancer

Patient: female, 75 years

Failed two lines previously:

- FOLFIRINOX
- Gemcitabine/Abraxane

Treatment: Deflexifol®

- 525 mg/m^2 bolus + 3000 mg/m^2 infusion

Result: Stable Disease 6 months

Metastatic Colorectal Cancer

Patient: male, 61 years

Failed four lines previously:

- FOLFOX + bevacizumab
- FOLFIRI
- Panitumumab
- Lonsurf®

Treatment: Deflexifol®

- 525 mg/m^2 bolus + 3800 mg/m^2 infusion

Result: Partial Response 6 months

^ Patients treated in the FP101A phase Ib/IIa trial (ACTRN12619001533189, completed May 2023). Presented at ASCO GI 2024: Link

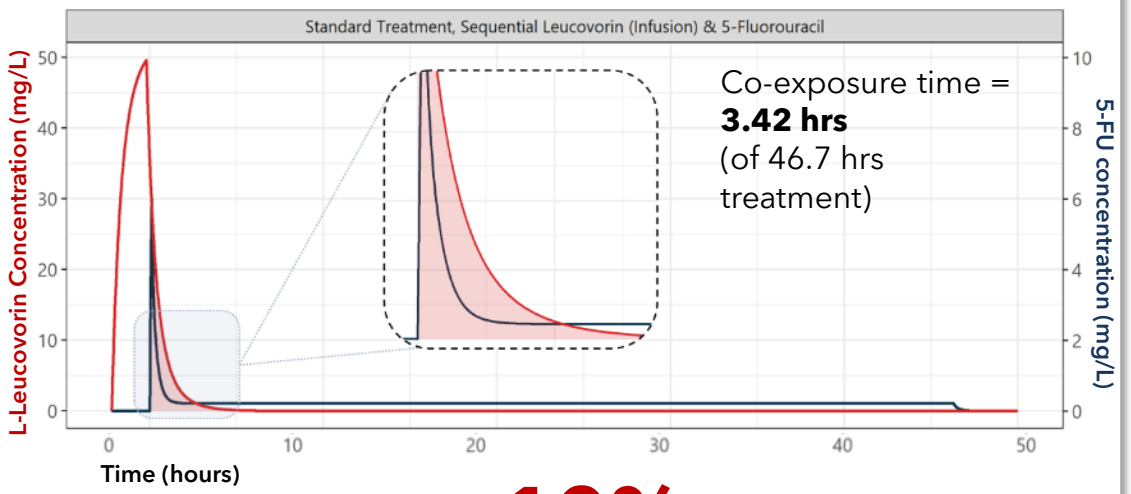
FOLFOX = 5-FU, LV & oxaliplatin; FOLFIRI = 5-FU, LV & irinotecan; FOLFIRINOX = 5-FU, LV + oxaliplatin & irinotecan
PFS = Progression Free Survival; Partial response = tumour reduced in size by ≥30%

WHY DEFLEXIFOL® ENHANCES EFFICACY

Deflexifol® increases co-exposure from 3.4 hours to 46.7 hours

Deflexifol® co-formulates 5-FU/LV safely with an FDA-approved cyclodextrin to enable maximal tumour co-exposure over the standard 46 hr infusion treatment cycle, enhancing 5-FU activity for optimal treatment efficacy

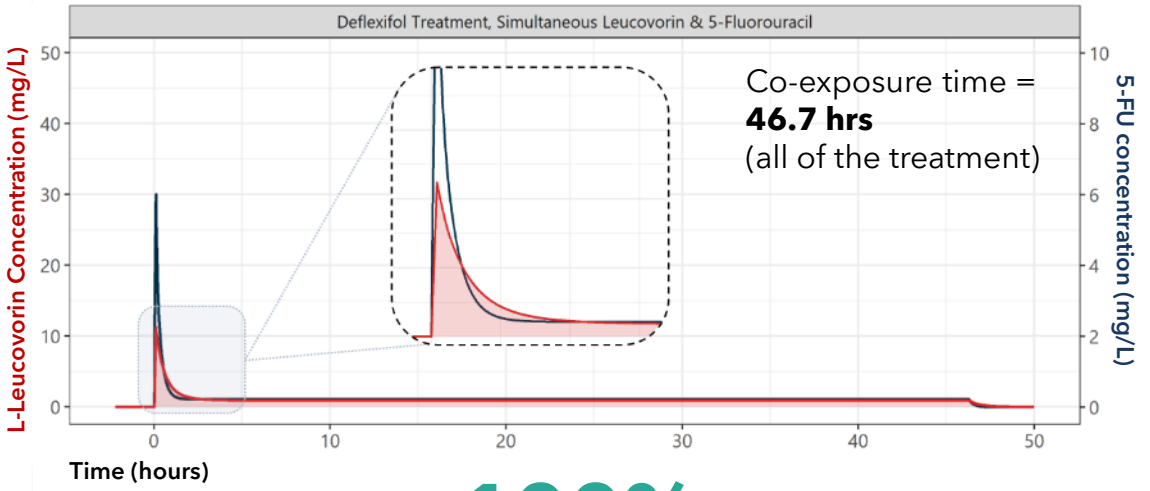
Current Standard of Care
Sub-Optimal Serial Administration



<10%

5-FU/LV co-exposure

Co-infusion via Deflexifol®
The New Gold Standard of Care™



100%

5-FU/LV co-exposure

Modelled pharmacokinetics based on published independent literature:
Maring et al. 2003, *Cancer Chemother Pharmacol.* 51:167
Straw et al. 1984, *Cancer Res.* 44:3114

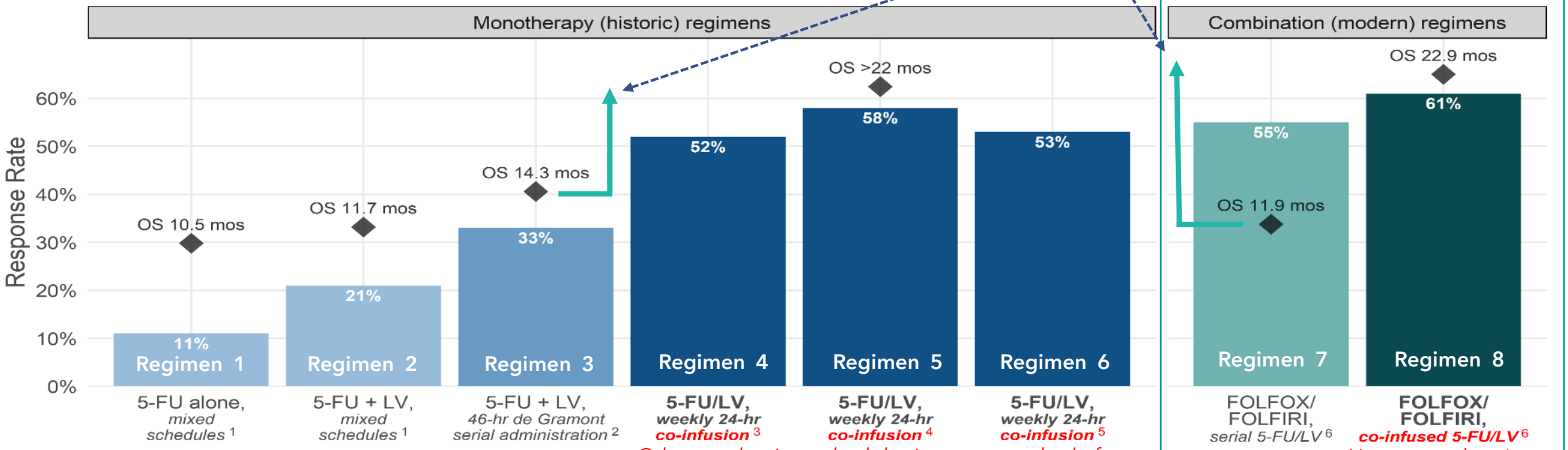
5-FU/LV Co-Infusion Improves Anti-Tumour Efficacy

FivepHusion's thesis is supported by robust third-party data

mCRC 1st line treatment has only incrementally improved over decades.

- **Independent phase II trials indicate superiority of 5-FU/LV co-infusion** (using unsafe / impractical/ unapproved methods).
- **FivepHusion's Phase III trial will aim to outperform results from Regimen 7:**
  - Results anticipated to **meet/exceed Regimen 8, resulting in successful approval**

Precedent for Deflexifol® - Designed to safely co-infuse 5-FU/LV to enhance efficacy



OS = Overall Survival

Other academic methodologies - not standard of care due to chemical incompatibility leading to safety risks

Unapproved regimen

Direct comparison in a two-arm phase II study

1. Thirion et al. 2004, *J Clin Oncol.,* 22(18):3766-75.
2. de Gramont et al. 1997, *J Clin Oncol.,* 15(2):808-15.
3. Ardalan et al. 1991, *J Clin Oncol.,* 9(4):625-30.
4. Yeh et al. 1997, *Anticancer Res.,* 17(5B):3867-71.
5. Yang et al. 1999, *Cancer*, 85(9):1925-30.
6. Bleiberg et al. 2012, *Acta Gastroenterol Belg.*, 75(1):14-21.

FOLFOX: 5-FU + LV + oxaliplatin
FOLFIRI: 5-FU + LV + irinotecan

DEFLEXIFOL® A NEW BACKBONE THERAPY

Deflexifol® aims to replace 5-FU + LV as the backbone therapy of mCRC

A **backbone therapy** is the foundation treatment in a combination regimen.

Other drugs, such as targeted therapies, immunotherapies, or supportive agents, are added to, in order to enhance effectiveness.

5-FU + LV is a First-Line Backbone Therapy:

- ✓ **Proven Efficacy**: Supported by strong clinical evidence and often forms the standard of care.
- ✓ **Benchmark in Trials**: Commonly used as the control group in clinical trials testing new treatments.
- ✓ **Combination Platform**: Designed to work alongside new or investigational drugs.
- ✓ **Persistent Role**: Remains a central part of treatment unless clearly outperformed by newer therapies.

Deflexifol® aims to outperform and replace 5-FU + LV

Backbone Therapy	Approved Combinations
5-FU + LV / Deflexifol® (if successful)	+ OXALIPLATIN (**FOLFOX**)
	+ IRINOTECAN (**FOLFIRI**)
	+ BEVACIZUMAB
	+ CETUXIMAB / PANITUMUMAB
	+ ENCORAFENIB + CETUXIMAB
	Emerging Add-Ons
	+ Checkpoint Inhibitors NIVOLUMAB & PEMBROLIZUMAB
	+ Immunotherapies
	+ Biomarker-Driven Therapies

5-FU & LV (Deflexifol®) faces limited competition risk as it will likely remain as a Backbone Therapy with new mCRC treatments utilised in combination.

FDA Confirmed Path To Market

Feedback on FivepHusion data set, clinical development, CMC and regulatory plans for Deflexifol®

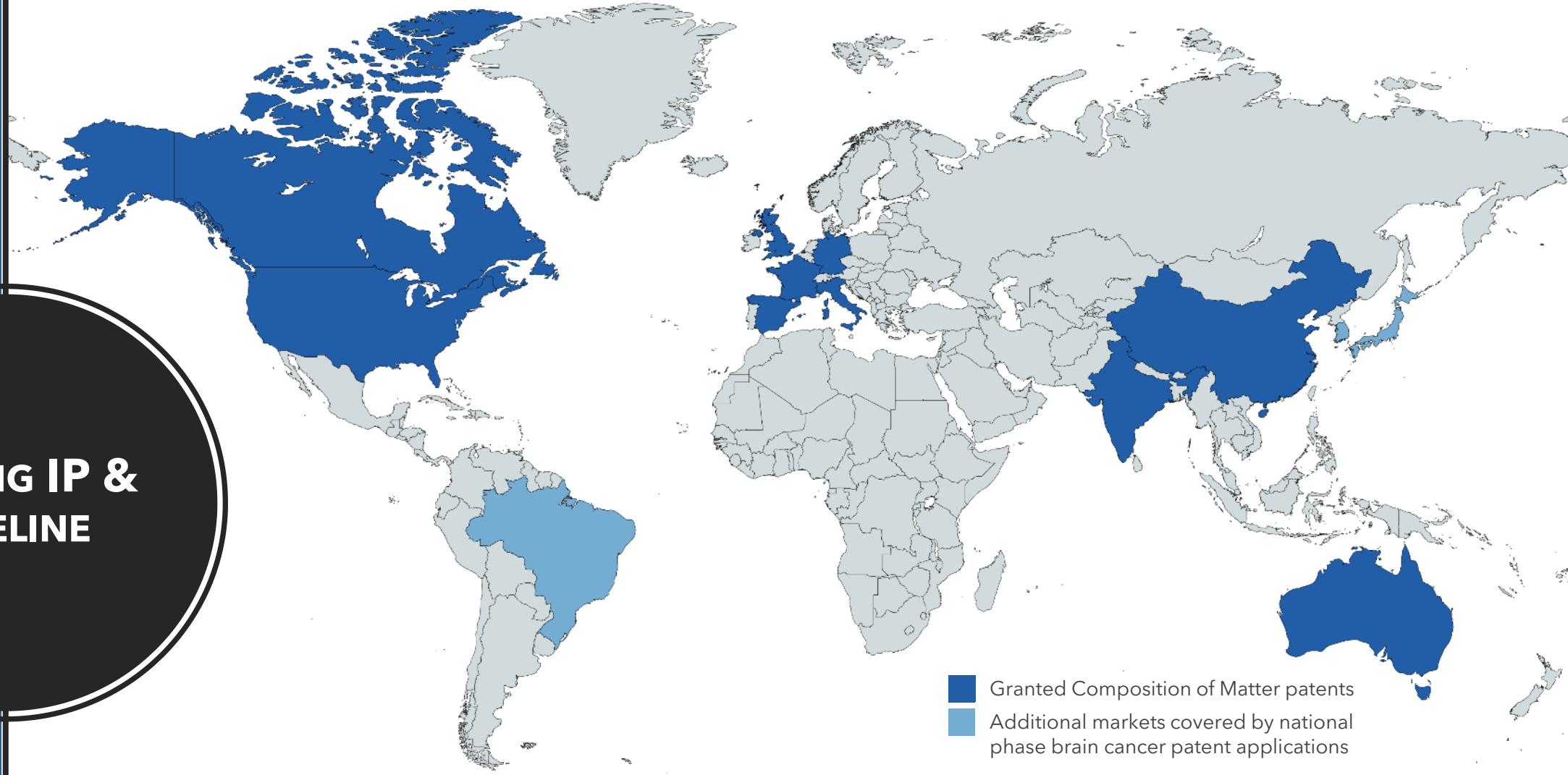
Key FDA Feedback

1. **Deflexifol® can be immediately developed for 1st line mCRC patients**
   - **No need for phase II** due to approved drugs with established safety and tolerability
   - **No need to first seek registration in later lines of therapy**
2. **Advice on design of planned phase Ib/IIa ("combo") trial** confirming Deflexifol® dose when combined with oxaliplatin and bevacizumab (HREC approved – ready to initiate)
3. **Only one successfully conducted phase III** pivotal trial required to support registration
4. **Accelerated regulatory path** for registration in mCRC (FDA 505(b)(2))

Endorses our plan to accelerate towards phase III development and registration for Deflexifol®

*FDA Type C meeting feedback received H1 2023. Press release can be found here: Link

Strong IP & Pipeline



- **Granted composition of matter**
- Patents in prosecution
- **New composition** filed in 2025
- IP pipeline 2026

expected exclusivity to

>2045

DEFLEXIFOL®: PAEDIATRIC EPENDYMOMA

Aiming to be the first approved drug for Paediatric Ependymoma

PAEDIATRIC EPENDYMOMA	• **The third most common brain cancer in children** • Peak incidence <4 years of age
CURRENT TREATMENT	• Surgical resection and adjuvant radiotherapy • There are no approved drug therapies
RATIONALE	• **US trial[1]: 5-FU activity in children that had failed prior therapy** • **Deflexifol® is safer and more efficacious than 5-FU alone**
DEFLEXIFOL® AT RELAPSE TRIAL (DART)	• **National,** investigating safety and tolerability in children with brain cancer • **A safe & tolerable dose confirmed, encouraging reports of extended treatment durations. Oncologist enthusiasm to commence phase II**

Orphan indication with a fast path to approval

[1] Wright et al. 2015, *Neuro Oncol.*, 17(12):1620-27

SIGNIFICANT COMMERCIAL OPPORTUNITIES

FivepHusion's conservative modelling indicates blockbuster status for Deflexifol®

Deflexifol® addresses global markets

Global annual colorectal cancer incidence: 1.9M

- **20-30%** diagnosed metastatic[1]
  - **380K - 570K new cases are metastatic (mCRC)**
- **~50% of patients with earlier-stage CRC** will eventually develop metastases[2]
- **US$13B** mCRC market[3], **majority receive 5-FU + LV**[4]
- FDA confirmed immediate **path to 1st line treatment**
- **Strong pharmacoeconomic value** / basis for **premium pricing**
- **Limited competition** – other drugs typically combine with 5-FU + LV

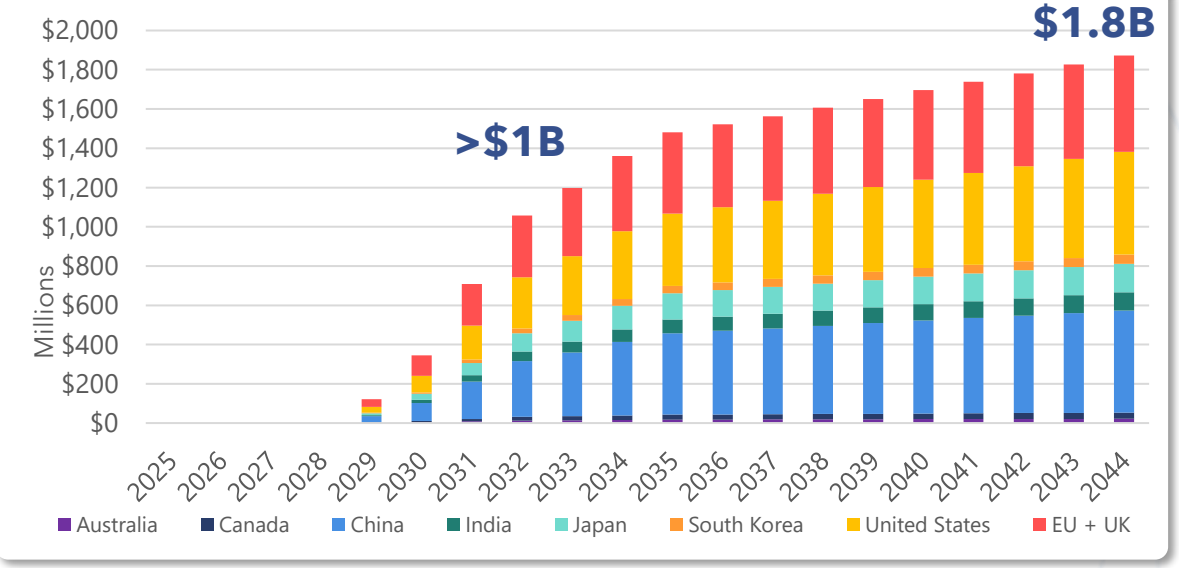
Pipeline Upside:

+ Paediatric brain cancer: **US$1.84B**[5] → Adult brain cancer
+ Replace 5-FU+LV across solid tumour indications = **>5M patients**

On mCRC approval:

Deflexifol® may also receive an **FDA registration label enabling** physician use **across all other solid tumour indications for which 5-FU + LV are currently utilised.**

Deflexifol® Projected Sales[6]



Path to Substantial Value

- **De-risked & accelerated regulatory pathways** to market
- Commercial launch: As early as **2029**
- **Strong KOL** interest to switch to a superior co-formulation
- Projected global peak sales: **US$1.8B**

[1] Global Cancer Observatory 2020, Cancer Today; GLOBOCAN 2020
[2] Metastatic colorectal cancer: ESMO Clinical Practice Guidelines for diagnosis, treatment and follow-up † Van Cutsem, E. et al.Annals of Oncology, Volume 25, iii1 - iii9

[3] 2025 Colorectal Cancer Market Insight, Epidemiology And Market Forecast - 2034
[4] Glimelius et al., 2021, *Cancer Treatment Reviews* 98:102218
[5] Market Research Future 2023
[6] Indications: drug sales for the treatment of mCRC, ependymoma, CRC, breast, gastric, pancreatic

RESECTASSIST™: BIODEGRADABLE DRUG-ELUTING IMPLANT

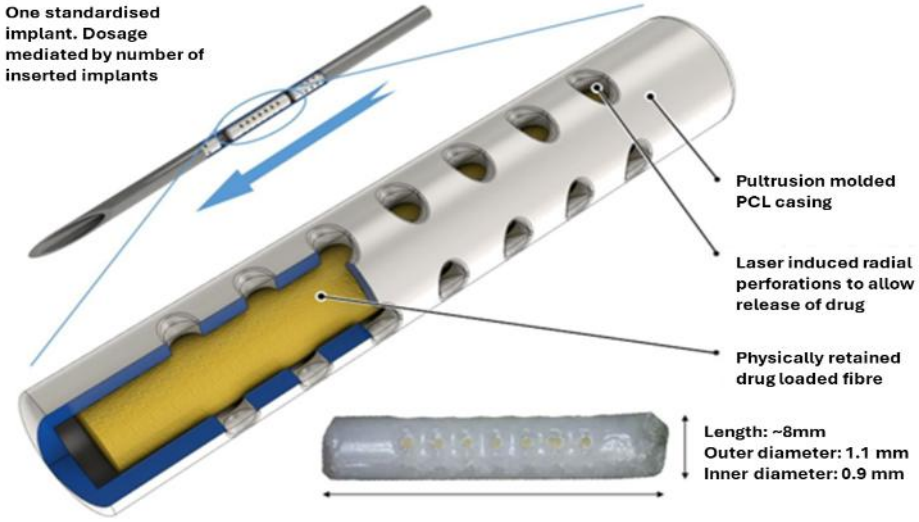
Exclusive option over ResectAssist™ significantly bolsters FivepHusion's pipeline

Novel Drug Delivery Technology Platform[1,2]

- Intra-tumoural drug delivery
- **Manufactured using FDA-approved biomaterials**
- **Delivers diverse drug payloads:** Approved medicines and in-development drugs (small molecule, biologics, antibody-drugs, mRNA and others)

Higher
Focused Dose

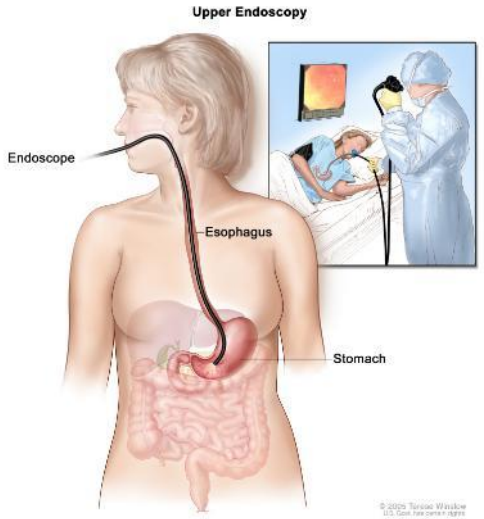
Lower
Systemic Toxicity

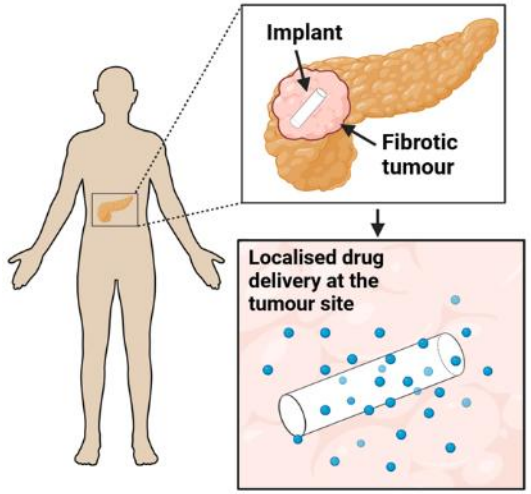


Lead Program: Pancreatic Cancer

ResectAssist™-FOLFIRINOX: Downstaging tumours to resectable with curative intent

- ✓ **Unmet Market: >$7B market opportunity[3]**
- ✓ **Strong IP: Composition of matter patents & IP pipeline, including novel drug payload - device combinations**
- ✓ **Govt Grant: $500K Federal AEA Ignite grant**



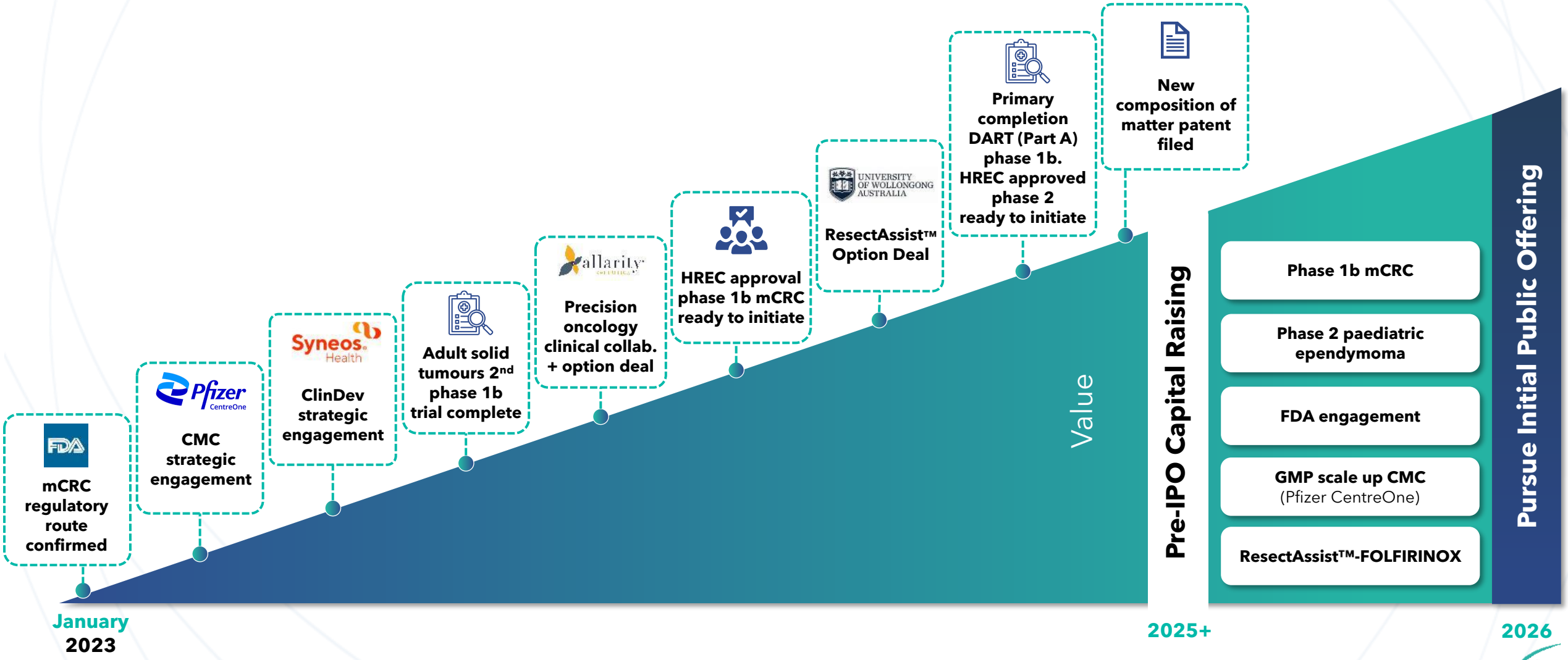


UNIVERSITY OF WOLLONGONG AUSTRALIA

[1] Wade et al., 2020, Adv. HealthcareMater. :Nov;9(21):e2001115; [2] Minaei et al., 2024, J Immunother Cancer 2024;12(Suppl 2):A1–A1683;
[3] Polaris Market Research 2022 – pancreatic cancer market in 2030

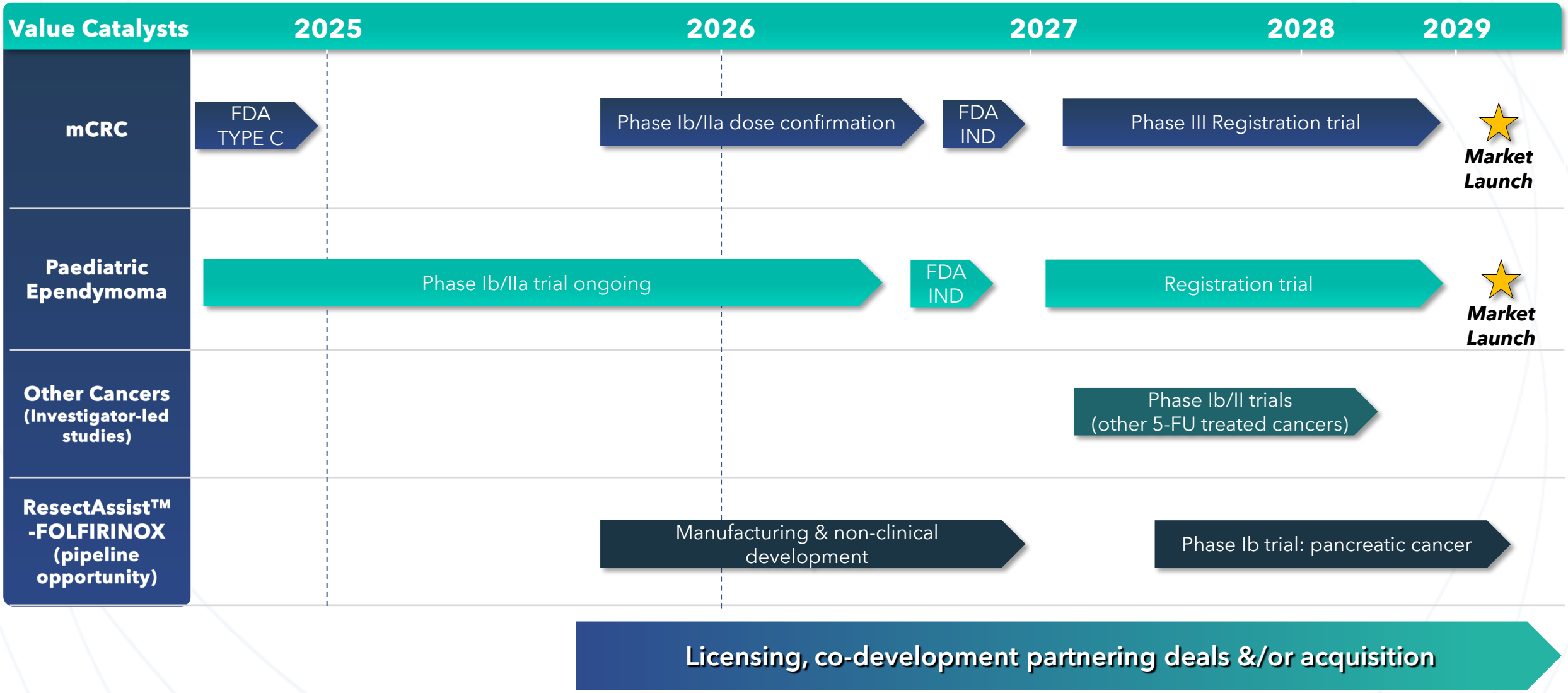
FivepHusion Timeline

Continuous value creation from 2023 → 2025 & Beyond



VALUE CREATION STRATEGY

FivepHusion's fast-tracked development strategy to global markets



All development steps and timelines are indicative

ASX Listed Oncology Peers

FivepHusion aims to bridge the valuation gap to its comparable listed peers

Peer	ASX Ticker	Description	Lead Candidate Clinical Stage	Market Cap
Telix	TLX	Telix is specialising in radiopharmaceuticals for cancer diagnosis and therapy.	Phase III Commercialisation	$4,722m
CLARITY	CU6	Clarity is targeting membrane antigen (PSMA)-expressing metastatic castration-resistant prostate cancer.	Phase III	$1,399m
RACE ONCOLOGY		Race Oncology is reformulating a version of bisantrene, a chemotherapy targeting acute myeloid leukaemia (AML) with potential applications in breast cancer and clear cell renal cell carcinoma (ccRCC).	Phase II	$539m
immutep	IMM	Immutep is developing immunotherapies targeting metastatic lung cancer.	Phase III	$383m
starpharma	SPL	Starpharma is developing its dendrimer technology for pharmaceutical applications, such as cancer	Phase II	$169m
arovella	ALA	Arovella is developing off-the-shelf cancer CAR-T Cell immunotherapies targeting CD19-positive blood cancers.	Phase I	$103m
IMUGENE Developing Cancer Immunotherapies	IMU	Imugene is developing novel therapies to activate the immune system against cancer.	Phase II	$99m
Prescient Therapeutics	PTX	Prescient is developing personalised medicine approaches to cancer, including targeted & cellular therapies	Phase II	$61m
Amplia THERAPEUTICS	ATX	Amplia is developing its Focal Adhesion Kinase (FAK) inhibitors for pancreatic cancer	Phase II	$59m

Comsec Data: Market Capitalisations as of 20th November 2025

RECENT ONCOLOGY TRANSACTIONS

FivepHusion

Date	Type of Deal	Acquirer/Licensee	Target/Licensee	Stage	Upfront (US$)	Milestones (US$)	Total Deal Value (US$)
Apr-24	Partnership	NOVARTIS	PeptiDream	Platform	$180m	$2,700m	$2,880m
Feb-25	Option Agreement	abbvie	xilio THERAPEUTICS	Multiple	$52m	$2,100m	$2,152m
Dec-24	Partnership	GSK	DualityBio 映恩生物	Multiple	$30m	$975m	$1,005m
Nov-24	Partnership	KURA ONCOLOGY	KYOWA KIRIN	Phase 3	$330m	$1,200m	$1,530m
Jun-24	Option Agreement	Takeda	Ascentage	Phase 3	$100m	$1,200m	$1,300m
Jan-23	Licensing Agreement	Takeda	HUTCHMED	Phase 3	$400m	$730m mCRC	$1,130m
Mar-25	Acquisition	Jazz Pharmaceuticals	CHIMERIX	Phase 3	$935m	Paediatric brain	$935m
Jul-24	Licensing Agreement	IPSEN	Day One BIOPHARMACEUTICALS	Phase 3	$111m	$350m	$461m
Nov-24	Acquisition	BIONTECH	BIOTHEUS 普米斯生物技术	Phase 2	$800m	$150m	$950m
Sep-24	Licensing Agreement	sanofi	oranomed	Phase 2	$110m	$250m	$360m
Jan-25	Acquisition	GSK	IDRx	Phase 1b	$1,000m	$150m	$1,150m
May-24	Licensing Agreement	NOVARTIS	ARVINAS	Phase 1	$150m	$1,000m	$1,150m
Jan-25	Licensing Agreement	MENARINI group	Insilico Medicine	Phase 1	$20m	$550m	$570m
May-24	Acquisition	NOVARTIS	mariana ONCOLOGY	Preclinical	$1,000m	$750m	$1,750m

Appendix

Potential Corporate Transactions for FivepHusion

FivepHusion is actively engaged with Global and Regional Pharma Companies

Potential US/Global Licensing Deal

- **License:** **Exclusive worldwide rights, or US +** major EU, often with option for FivepHusion to co-promote.
- **Upfront Payment:** **US$50–150m at Phase III entry.**
- **Milestones:** **>US$500m** across development, regulatory, and commercial events.
- **Royalty:** **Industry Standard.**

Phase Ib/IIa mCRC Clinical Trial › **FDA IND** › **Phase III Registration Trial** › **Regulatory Approval & Commercialisation**

Potential Regional Licensing Deal

- **License:** **A defined territory,** FivepHusion retains US/global rights.
- **Upfront Payment:** **US$10–30m** (supported by strong composition of matter IP + 505(b)(2) fast track).
- **Milestones:** **US$50–200m** tied to Phase III initiation, regulatory approvals, and first commercial sales in the region.
- **Royalty:** **Industry Standard.**

FP101B: HREC Approved Phase Ib/IIa Trial Design Study^

Dose exposure / response confirmation for Deflexifol® when combined with oxaliplatin + bevacizumab

Trial Design

- **1st line unresectable mCRC**
- **Two stage phase Ib/IIa** Trial Design
  - **40 – 50 patients**; trial duration **~12 months**
- Allarity Therapeutics collaboration: Blinded evaluation of DRP®-5-FU CDx predictive ability

Endpoints

- **Primary endpoints:** Safety and tolerability of Deflexifol® when combined with oxaliplatin and bevacizumab
- **Secondary endpoints:**
  - Pharmacokinetics of Deflexifol® when combined with oxaliplatin and bevacizumab, DRP®-5-FU evaluation
  - ORR, PFS*

PART A

Dose Escalation Cohorts (3 + 3)
(9 – 18 pts, 3 trial sites; ~6 - 8 months∅)

Standard of Care
OXALIPLATIN
85 mg/m²
BEVACIZUMAB
5 mg/kg

⊕ **DEFLEXIFOL® BOLUS#**
400 mg/m²

⊕ **DEFLEXIFOL® INFUSIONΩ**

Dose: → **2400 mg/m²**

No DLTs ↑

3000 mg/m²

No DLTs ↑

3400 mg/m²

3 patients per cohort +
an additional 3 patients at the final dose

⊕

PART B

Expansion Cohort
(~30 pts, 6 - 8 trial sites; ~6 months∅)

OXALIPLATIN
85 mg/m²
BEVACIZUMAB
5 mg/kg

⊕ **DEFLEXIFOL® BOLUS**
400 mg/m²

⊕ **DEFLEXIFOL® INFUSION**
Part A MTD

^ Trial design approved by Bellberry HREC. Trial planned to commence H1 2026, pending successful capital raising
*ORR = Objective Response Rate; PFS = Progression Free Survival, MTD = Maximum Tolerated Dose, DLT = Dose Limiting Toxicity
∅ Time frame to expected primary completion

Deflexifol® bolus = 400 mg/m² 5-FU + 27 mg/m² LV;
Ω Deflexifol® infusion dose escalation = 2400 mg/m² 5-FU + 160 mg/m² LV (equivalent to the current standard 5-FU dose) up to the currently declared MTD of 3400 mg/m² 5-FU + 227 mg/m² LV

PHASE 1/2 DEFLEXIFOL® AT RELAPSE TRIAL (DART)

Ongoing investigator-led trial; predominantly charity funded

FivepHusion

Ongoing investigator-led trial involving paediatric oncology centres across Australia[1]

Paediatric Patients with:

- **refractory/relapsed CNS tumours, including ependymoma;**
- newly diagnosed **diffuse intrinsic pontine glioma** (DIPG) / **diffuse midline glioma** (DMG) who have completed radiotherapy

Trial Design

Part A: Open-label, phase I dose escalation

- Between n= 6-24, bolus + infusional Deflexifol® commencing at the adult MTD with dose de-escalation as required

Part B: Phase II refractory or recurrent ependymoma expansion cohort^

- Up to n=10, primary endpoint of Objective Response Rate

Part A Completed - safe and tolerable dose confirmed. Encouraging treatment durations reported. Oncologist enthusiasm to commence Part B (phase II)

[1] ACTRN12623000104651 (link)

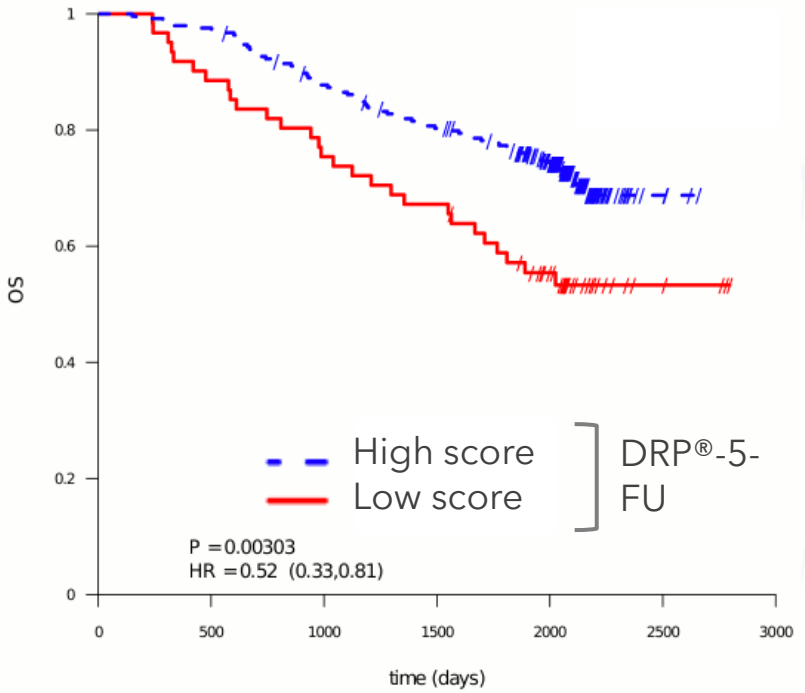
Exploring Precision-Oncology (August 2023 Deal)

Collaboration with Allarity Therapeutics to predict 5-FU responders

FivepHusion and Allarity Therapeutics Collaboration

- **Drug Response Predictor (DRP®) companion diagnostics**, highly validated via >35 clinical trials[1]
  - Proprietary DRP® algorithm applied to tumour biopsy gene expression data sets
  - Validated drug-specific response signatures, 80+% predictive response accuracy
  - 2-5 fold increase in response: *predicted* sensitive vs *predicted* resistant tumours
- **DRP®-5-FU retrospectively validated to predict response and overall survival to 5-FU** treatment in late-stage CRC and mCRC [2,3]
- **Collaboration to evaluate the DRP®-5-FU** and other DRP® companion diagnostics in the upcoming FP101B phase 1b/2a trial of Deflexifol® in 1st line mCRC
- **Option right to negotiate an exclusive license** to commercialise the DRP®-5-FU and other DRP® companion diagnostics for Deflexifol®
- **Potential to personalise cancer treatment** for patients most likely to benefit from Deflexifol®

Overall Survival of Stage III Colon Cancer Patients Treated with 5-FU + LV [2]



n = 307 stage III CRC patients from PETACC-3 trial

1. www.allarity.com 2. Buhl et al., 2016, PLoS ONE 11(5): e0155123; 3. Buhl et al., 2019, Annals of Oncology, 30(5):v242; OS = Overall survival

REGISTRATION TRIAL: DRAFT PLAN FOR PHASE III TRIAL Q4 2026

1st line treatment of unresectable mCRC

Phase III Registration Trial

- International, multi-centre registration trial (2026 – 2028)
- Designed to demonstrate that as a treatment for first-line unresectable mCRC,
  - *Deflexifol® in combination with oxaliplatin and bevacizumab (DEFLOX)*
- is superior in efficacy to*
  - *the standard of care mFOLFOX6 + bevacizumab regimen*

Rationale for superior efficacy over the standard of care

- ***Optimised 5-FU/LV co-exposure***
- ***Higher 5-FU dose***

GLOBAL PHASE III TRIAL

A ~550 patient, blinded, randomized phase III study of Deflexifol® in combination with oxaliplatin (DEFLOX) and bevacizumab vs mFOLFOX6 and bevacizumab in first-line unresectable metastatic colorectal cancer

RANDOMIZATION 1:1

DEFLOX + bevacizumab *i.v. 14-day cycle*	**mFOLFOX6 + bevacizumab** *i.v. 14-day cycle*

Radiological response every 8 weeks (RECIST 1.1)

Primary Objective: Best Objective Response Rate

Secondary Objectives: Progression Free Survival, Overall Survival, safety, Quality of Life

* Considering regulatory and commercial factors, this trial design is to be refined and confirmed based on independent expert feedback from KOL oncologists, clinical scientists and regulatory specialists, together with consultation with the FDA, EMA, NMPA and potentially other regulators

EPENDYMOMA SENSITIVITY TO 5-FU

Ependymoma (EPN) = 3rd Most Common paediatric brain tumour[1]

Ependymoma cell lines have significantly lower *thymidylate synthase* expression levels [2,3] → **increased 5-FU sensitivity**

Posterior fossa (PF)
~2/3 of childhood EPN [1]

Supratentorial
~1/3 of childhood EPN [1]

All partial responders in the St Jude 5-FU trial had primary PF-EPN [4]

PF-A = ~85-90% of PF-EPN [1]
- Predominantly younger children
- Frequent gain of chromosome arm 1q (1q+) [5]
  - ~20% at presentation
  - ~50% at first recurrence

PF-B = ~10-15% of PF-EPN [1]
- Mostly older children & adults

INCREASINGLY HIGH RISK
(Younger age, PF-A & 1q+ are negative prognostic factors)

PF-A 1q+ cell lines demonstrate:
- Repressed p53 (tumour suppressor) activity **that is restored by 5-FU**
- Significantly higher expression of *UCK2*, a 5-FU 'activating' enzyme → **increased 5-FU sensitivity**

Compared to PF-A 1q wild-type cells [6]

[1] Zaytseva *et al.* 2021, *Cancers* 13(19):4954.
[2] Atkinson *et al.* 2011, *Cancer Cell* 20(3):384-99.
[3] Donson *et al.* 2018, *Mol Cancer Ther.* 17(9):1984-94.
[4] Wright *et al.* 2015, *Neuro Oncol.* 17(12):1620-27.
[5] Donson *et al.* 2023, *Neuro Oncol.* 25(10):1854-67.
[6] Griesinger *et al.* 2024, *Clin Cancer Res.* 30(8):1544-54.

5-FU/LV Co-administration: Precedents & Comparison

FivepHusion

Deflexifol® is clinically and commercially viable

Two separate pumps[1]	Dilution strategies[2-3]	Sodium salt LV[4-5]	Deflexifol®
✓ Considerably improved survival & response rate in untreated & previously treated mCRC patients.[1]	✓ Considerably improved response rates in untreated & previously treated mCRC patients.[2,3]	✓ Significantly improved patient survival in 1st line mCRC when used in modern infusion regimens.[4,5]	✓ **No precipitation** or catheter blockages.
× **Not clinically or commercially feasible** - requires two pumps & intravenous lines.	× **Does not prevent precipitation** and catheter blockages.	× **Not approved** for this usage.	✓ Pain-free, **physiological-pH** formulation.
	× **Not approved** for this usage.	× **Requires alkaline pH of 9.0.**	✓ Highly tolerable, with a **higher MTD than 5-FU.**
		× **Similar toxicity to standard dose administration.**	✓ **Improved safety.**
			✓ **Demonstrable efficacy** in end-stage patients following previous 5-FU failure.
			✓ **Composition of matter patent protection.**

While surrogate studies provide supporting evidence, they remain commercially unviable and impractical in real-world use – **positioning Deflexifol® as the only path forward with both safety and efficacy.**

[1] Ardalan *et al.*, 1991, J Clin Oncol. 9:625.
[2] Yeh *et al.* 1997, Anticancer Res. 17:3867.
[3] Yang *et al.* 1999, Cancer 85:1925.
[4] Bleiberg *et al.* 2012, Acta Gastroenterol Belg. 75:14.
[5] Romano *et al.* 2021, Oncotarget 12:221.

New Version Pricing Premiums

Deflexifol® has blockbuster potential at all potential pricing outcomes

New version drugs commanded between 2-175x price premiums in comparison to the originator drugs

- Most of these assets demonstrated only modest or even non-inferior improvements in safety and/or efficacy
- Isofol Medical **expected >US$4,000**/month for arfolitixorin (new version of LV) = **8x increase over LV** [1]
  - Analysts expected $3,000 – $6,600/month prior to clinical failure [2, 3]

5-FU & LV: Current Pricing ($US)	
5-FU + LV: Per month	~$180 – $800
5-FU + LV: Per course*	~$1,500 – $6,500

Deflexifol®: Potential Pricing		
Low Case (~2x)	~$1,100/month;	~$9,000/course
Mid Case (~8x)	~$4,400/month;	~$35,000/course
Courses/patient	~2-3x	
Cost/patient	**$18,000 → $70,000**	
COGS	Immaterial	
Cases/annum	mCRC = ~500,000 Other Solid Tumours = 5,000,000+	

Originator	New Drug	Region	Originator Price	New Drug Price	Price Increase
5-FU/LV	**Xeloda® (capecitabine)**	US	~$185/month	~$5,900/month	**32x**
LV calcium (folinic acid)	**Fusilev® (levo-LV)**	US	~$55/dose	~$1,500/dose	**27x**
Daunorubicin + cytarabine Price is for 2x induction cycles + 2x consolidation cycles	**Vyxeos®**	US	~$5,200	~$128,000	**24x**
		UK	~£6,886	~£82,458	**12x**
		Germany	~€2,163	~€132,056	**61x**
		Italy	~€535	~€84,474	**158x**
		Spain	~€534	~€93,600	**175x**
Paclitaxel (Taxol)	**Abraxane® (nab-paclitaxel)**	US	$150/dose generic; $1,000/dose branded	$4,200 /dose	**4-42x**
		UK	£668/3wks	£1,230/3wks	**2x**
Paclitaxel (Taxol)	**Taxotere® (docetaxel)**	US	$150 /dose generic; $1,000/dose branded	$2,500/dose	**2.5-17x**
		UK	£668/3wks	£1,232/3wks	**2x**
Doxorubicin (Adriamycin)	**Doxil® (Doxorubicin liposomal)**	US	$1,066/month	$2,311/month	**2x**
Cytarabine	**DepoCyt® (Cytarabine liposomal)**	US	$84/month	$4,762/month	**57x**

[1] Isofol Medical AB, IPO Prospectus 2017; Isofol Medical, Arfolitixorin overview, 2020
[2] DNB Markets, Isofol Medical Equity Research 15 Nov 2020; Redeye research update Isofol Medical, 15 Nov 2020
[3] Wolters Kluwer Medi-Span Price Rx Accessed May 2020
* An average course of treatment is 8 months
Pricing data sourced from published literature and national pricing & reimbursement authorities (e.g. CMS, G-BA, NICE, AIFA, etc). Prices differ across geographic regions and are subject to change over time due to changes in legislation, demand, shortages, and other factors, thus prices listed on this slide may be subject to change in the future.

FivepHusion vs. Traditional Biotech

FivepHusion's unique co-formulation strategy delivers a rare value proposition

	Cost	Timelines	Risk Profile	1st Line Therapy	Strong IP	Blockbuster
FivepHusion	✓ Low	✓ Short ~3.5 Years	✓ Low Safety & Efficacy Established	✓ Yes	✓ Yes Composition of Matter	✓ Yes
Traditional	X High	X Long 13-16 years	X High Safety & Efficacy Unproven	X Rarely	✓ Yes Composition of Matter	✓ Yes

Development Timelines

FivepHusion

FivepHusion — **Established safety and attractive trial designs result in fast and cost-effective clinical programs**

Phase Ib/IIa Clinical Trial (1 Year) → 1x Phase III Registration Trial (2.5 Years) → Regulatory Approval (<1 Year)

Traditional

Discovery & Pharmacology (2-3 Years) → Preclinical Testing → Phase 1 Clinical Trials (5-6 Years) → Phase II Clinical Trials → 2x Phase III Clinical Trials (5-6 Years) → Regulatory Approval (>1 Year)

FivepHusion Key Highlights

FivepHusion's strategy presents a unique and compelling risk-reward profile

Blockbuster Markets	• **1.9m colorectal cancers diagnosed** p.a. (≤570k metastatic) = **US$13B mCRC market** • **5.0m+ solid tumour diagnosed p.a**. (where 5-FU + LV are utilised)
First-Line Therapy	• 5-FU + LV: Established standard of care' backbone therapy for mCRC (95% of patients) • **Deflexifol®: Aims to replace current 'standard of care' (SOC) backbone therapy**
Strong economics	• **Premium pricing** potential vs. generics, **driven by superior safety and efficacy** • **Rapid uptake** - KOLs believe **Deflexifol®** would be **widely adopted within 2 years** of clinical validation and launch • Conservative Modelling suggests peak sales ~$1.8B+ (multiplies on higher pricing)
Clinically Validated	• 5x independent surrogate Phase II trials confirm rationale and therapeutic mechanism • 3x company clinical trials demonstrated **higher safety & tolerability** and **potent efficacy**

Fast-tracked and Capital-Efficient	• Fast-tracked development via 505(b)(2) pathway **enabling faster, lower cost approval**
	  ○ Phase Ib/IIa + **single pivotal Phase III** → ~3.5 years to approval
	  ○ Rare in oncology: Deflexifol® is a 'next generation' co-formulation (with composition of matter patents), differing from basic reformulations or new delivery methods.

Strong FDA Engagement	• Ongoing FDA dialogue, including Type C Meeting, guiding Phase Ib/IIa and Phase III trial designs (mCRC)
Significant Pipeline	• **Active Phase I/II paediatric brain cancer** trial moving towards Phase III. • **Broader applications where 5-FU + LV are utilised**: pancreatic, gastric, breast, head & neck cancers
IP Protection	• **Granted Composition of Matter patents**, expected **exclusivity to 2045**
Licensing Transactions	• Clear, short-term pathway to **regional and global licensing transactions** post IPO