

October 2023

Developing a New Standard Therapy for Peritoneal Carcinomatosis (PC)

with local short-range alpha-radiation from Radspherin[®] micro-particles, with non-systemic administration.

Key Take Home Messages

Radiopharmaceuticals is a hot-spot

A rapidly growing development area in oncology, and as such it has substantial Investor and Big Pharma focus.

Oncoinvent Competency is solid

Founders and key personnel at Oncoinvent have a proven track record of developing a radiopharmaceutical that successfully made it to market (Xofigo[®]).

Peritoneal Carcinomatosis has a high unmet medical need

A disease area with a large unmet medical need. Peritoneal metastases can originate from several cancer types, and patient numbers are significant.

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Radspherin[®] is a Pipeline in a Product

There is significant potential for this non-systemic, receptor-independent, alpha-radiation therapy, applicable in many different cancer types.

Standard of Care status is attainable

Radspherin[®] has the possibility to become Standard of Care for peritoneal metastases from an array of different cancer types – Blockbuster Potential!

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Radiopharmaceuticals is an Investment hot-spot in Oncology



"**RayzeBio** stock opens with a big gain, to push valuation up to \$1.4 billion"

Marketwatch 15 Sep 2023

"Convergent Therapeutics Announces \$90 Million Series A Financing to Advance the Clinical Development of Radiopharmaceuticals for the Treatment of Prostate Cancer and Other Solid Tumors"

Cision 3 May 2023

"Startup **Mariana** raises \$175M for radiopharmaceutical drug research"

Biopharma Dive 7 Sep 2023

"Eli Lilly Jumps Into Radiopharmaceuticals via \$1.4B **Point Biopharma** Acquisition"

MedCity News 3 Oct 2023



Oncoinvent has a solid foundation for the creation of next generation radiopharmaceuticals

Radiochemistry and preclinical expertise

The company features **solid competency** in radiopharmaceutical development with founders and key staff having a solid track-record in developing alpha-emitting therapy that made it successfully to market



Robust isotope supply chain

Multiple supply sources of Thorium-228 raw material ensure a continuous and uninterrupted supply of Radium-224



Innovative alpha-particle product

A suspension of $CaCO_3$ micro-particles labeled with the alpha-emitting radioisotope Ra-224 uniquely delivers non-systemic, receptor-independent, radio-pharmaceutical therapy



GMP manufacturing & clinical operations

Internal GMP radiopharmaceutical manufacturing & supply capabilities. Pre-clinical and clinical studies demonstrate excellent safety and very promising long-term efficacy data

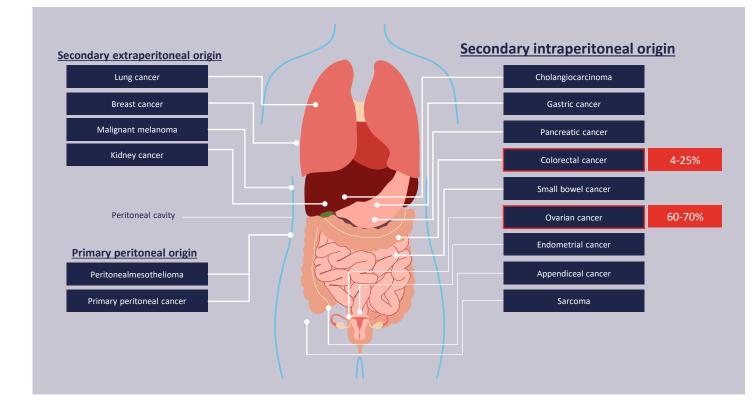


Peritoneal Carcinomatosis

- A multiple cancer type disease area with a significant unmet medical need



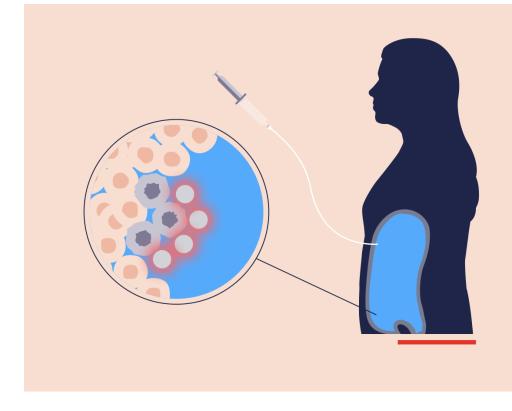
- Tumors of the peritoneum are most frequently caused by metastases from other primary cancers: ovarian and colorectal cancers are the commonest causes
- High patient burden and a dramatic impact on quality of life
- Poor prognosis and limited treatment options
- Considerable patient numbers and a significant unmet medical need!



How Radspherin[®] is administered in the Clinic

- Delivering highly effective receptor-independent alpha-therapy directly to target area

- A catheter is left after Cytoreductive Surgery (CRS)
- Through that catheter a single dose of Radspherin[®] microparticles, dispersed in a solution, are injected directly into the peritoneum, avoiding the need for systemic administration
- The alpha-radiating microparticles spread throughout the cavity eradicating micro-metastases
- After microparticles have lost their capacity for radiation, the calcium-carbonate based carrier material is naturally dissolved, absorbed, and excreted by the body
- Easy-to-use, non-invasive radiotherapy with non-systemic administration.



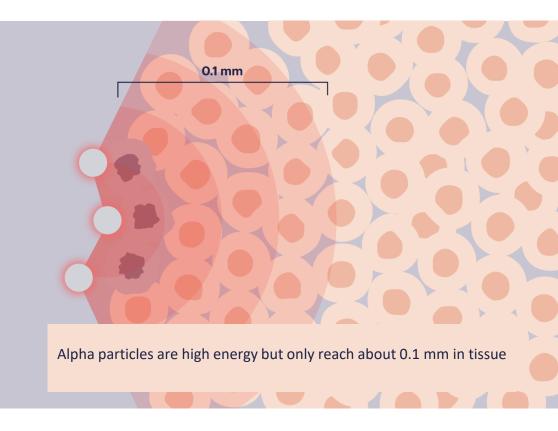
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Radspherin® in the treatment of Peritoneal Metastasis

- Delivering highly effective receptor independent alpha-therapy directly to target area

- Non-systemic, single-dose, intraperitoneal Radspherin[®] aims to eliminate residual micro-metastases after cytoreductive surgery and thus prolong disease-free period and survival
- Effective independently of cancer cell type and cellular resistance mechanisms it is a physical treatment
- The size and slow degradation of the microparticles combined with the half-life of 3.6 days of radium-224, provides sustained treatment effect over days
- Alpha radiation is highly effective, yet short-range in tissue. Ideal for large body cavity surfaces!



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Radspherin® - Clinical Development

- Clinical dose established, enrollment to phase 2a in completion



 Two Phase 1/2a studies - assessing dose, safety and tolerability, dosimetry and signal of efficacy of intraperitoneal Radspherin

RAD-18-001: Ovarian/fallopian tube cancer	 Oslo/Norway(PI: Yun Wang) Leuven, Belgium (PI: Els van Nieuwenhuysen/Ignace Vergote) Madrid/Pamplona, Spain (PI: Luis Chiva)
RAD-18-002: Colorectal carcinoma	 Oslo, Norway (PI: Stein Larsen) Uppsala, Sweden (PI: Wilhelm Graf)

For both studies, dose escalation is completed and the highest dose of 7 MBq selected, recruitment to expansion cohorts ongoing – more than 50 patients treated in total. Continued stream of follow-up data



Radspherin® - robust safety profile

- Well tolerated and minimal organ toxicity

Well tolerated and considered safe to use	 No dose limiting toxicities observed at any dose level Well tolerated with only grade 1-2 events reported as possibly related to Radspherin[®]
Clinically relevant dose determined	 7 MBq dose determined to be safe. Single-dosing!
Biodistribution measured	 80% of radioactivity dose remains in the peritoneal cavity Absorbed doses to other organs way below those associated with any toxicity
Good safety profile for hospital staff	 Low amount of activity in blood and urine No precautions related to external exposure required

Clinical Phase 1/2a

- No peritoneal recurrences in recommended dose group

- Expected median time to progression after CRS-HIPEC is around 12 months
 - Meaning that 50 % of patients are expected to have recurred at 12 months

15-months safety and efficacy presented at ASCO 2023, June

 At the 15-month time point of follow-up, only 25% of patients that received the recommended clinical dose of Radspherin[®] had experienced recurrences overall, and no peritoneal recurrences had occurred in the recommended dose group

Frøysnes et al. J Surg Oncol. 2016 Aug;114(2):222-7 Quenet et al. Lancet Oncol. 2021 Feb;22(2):256-266 ASCO 2023 poster presentation, abstract number 3518

CRS: cytoreductive surgery HIPEC: heated intraperitoneal chemotherapy



Clinical Phase 1/2a

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15-months safety and efficacy presented at ASCO 2023, June

 At the 15-month time point of follow-up, only 25% of patients that received the recommended clinical dose of Radspherin[®] had experienced recurrences overall, and no peritoneal recurrences had occurred in the recommended dose group **18-months** safety and efficacy presented at PSOGI 2023, October

 At the 18-month time point of follow-up, only 33% of patients that received the recommended clinical dose of Radspherin[®] had experienced recurrences overall, and still no peritoneal recurrences had occurred in the recommended dose group

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Phase 2b Target Populations in Peritoneal Carcinomatosis

Peritoneal Carcinomatosis

Ovarian cancer	Colorectal cancer	
~4 000 pat/year	~20 000 pat/year	Populations in Ph 2b
HRD-negative subgroup with pre-operative chemotherapy Median PFS ~12 months	Adjuvant to CRS/HIPEC Median PFS ~12 months	



Total Target Populations

Peritoneal Carcinomatosis

Ovarian cancer	Colorectal cancer	Gastric cancer	Other peritoneal and other body cavity lesions (Bladder, Pleura)	
~4 000 pat/year	~20 000 pat/year			Populations in Ph 2b
~ 4 000 - 8 000 pat/year	~20 000 pat/year			Populations in Ph 3 / MA
Same as Ph2b or possibly extended	Same as Ph2b as it is extended to Ph2b/3			
~28 000 pat/year	~90 000 pat/year	~25 000 pat/year	Significant potential	Potential Future Expansion
HRD independent Primary & Secondary	Prophylactic after surgery for high-risk primary tumors			

Radspherin® Radiopharmaceutical Pricing benchmarks

Product	PFS Benefit	OS Benefit	Price
Xofigo	N/A	3.6 m	USD 69.000
Lutathera	8.5 m	N/A	USD 190.000
Pluvicto	N/A	4.0 m	USD 255.000

PFS = Progression Free Survival, OS = Overall Survival

 Radspherin[®] has a targeted PFS improvement of 12 months and a minimum acceptable profile of 6 months.

Key tenets of the Oncoinvent Business Case

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- Targeted Patient Population: 24.000 eligible patients per year (US & Europe)
 - CRC: 20.000 eligible patients (based on Phase 3 target only)
 - Ovarian Cancer: 4.000 eligible Patients (based on conservative, no extension, Phase 3 target only)
- Peak Penetration Estimate (assuming Standard of Care): 70% and rapid uptake
- Average Pricing Estimate given PFS Target: 50.000 USD per treatment

20% of US & Europe sales estimate added for RoW

Peak Sales Estimate (*Phase 3 target only*): 24.000 patients x 70% x 50.000 USD/pat x 120% ~ <u>1 Bill. USD</u>

Implied future Enterprise Value: 3-5 x Peak Sales Estimate ~ <u>3-5 Billion USD</u>

Peritoneal Carcinomatosis – room for further expansion

- Addressable market, beyond ph3 target, with significant unmet medical need

Long-term survival rates of patients with PC Five-year survival rates in ovarian cancer² by disease stage of colorectal origin¹ 90% 70% 78% 39% 36% 24% 17% 2 3 1 year 3 year 5 year 1 4

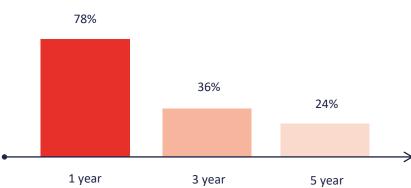
"Historically, the survival rate for gastric carcinoma patients with peritoneal carcinomatosis has been poor, ranging from 2.2 to 8.8 months and no survival at 5 years."³

1 source: <u>https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4655111/</u>

2 source: <u>http://www.cancer.org/cancer/ovariancancer/detailedguide/ovarian-cancer-survival-rates</u>

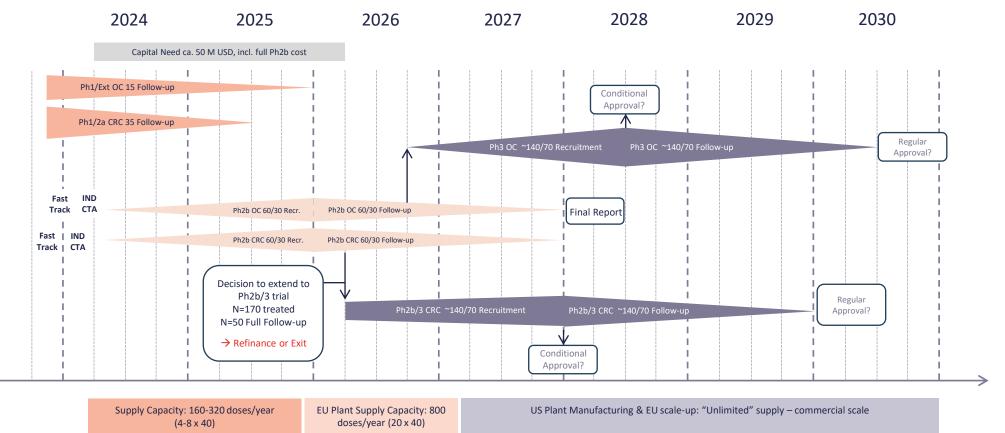
3 source: Cytoreductive Surgery and Hyperthermic Intraperitoneal Chemotherapy for Gastric Cancer - PMC (nih.gov)





Radspherin® Clinical Development Plan





17

Identified Partner Candidates



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Thank You!

Questions are more than Welcome!

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