



Oncoinvent

Innovative locally acting alpha-emitter technology

**Lead Product Candidate: Radspherin[®]
for Peritoneal Carcinomatosis (PC)**

Backed by Hadean Ventures & several high-value Family Offices in Norway, Europe

Oncoinvent: Differentiated, clinical stage radiopharma in multi billion market opportunity



- 1 Clinical stage opportunity in a rising market segment**
Oncoinvent is currently amongst most clinically advanced radiopharma company after several big pharma M&As
- 2 Unique radiopharmaceutical expertise at all levels of the company**
Founders, board members and management of Oncoinvent have a proven track record of developing a radiopharmaceutical asset and bringing it to market (Xofigo®).
- 3 Multi billion market opportunity in high unmet medical need indication**
The target area of Peritoneal Carcinomatosis can originate from several cancer types, and there exists a large patient population with a high unmet need.
- 4 Pipeline in a product**
Significant potential for this locally acting, receptor-independent, alpha-emitter therapy, applicable in many different cancer types.
- 5 Alpha-emitter with potential for Standard of Care status**
Radspherin® has the possibility to become standard of care for Peritoneal Carcinomatosis stemming from an array of different cancer types.

Radiopharmaceutical Expertise at all Levels

Management					
					
Anders Månsson CEO	Gro Hjellum COO	Anne-Kirsti Aksnes CCO	Kari Myren CMO	Tore Kvam CFO	
 	 	 	 	 	
					
Gillies O'Bryan-Tear Chair	Ingrid Teigland Akay Board Member	Hilde Steineger Board Member	Kari Grønås Board Member	Orlando Oliveira Board Member	Anne Cecilie Alvik Employee Rep.
Board					

	
Roy Larsen Scientific Founder & Advisor	Øyvind Bruland Scientific Founder & Advisor
Scientific Founders	

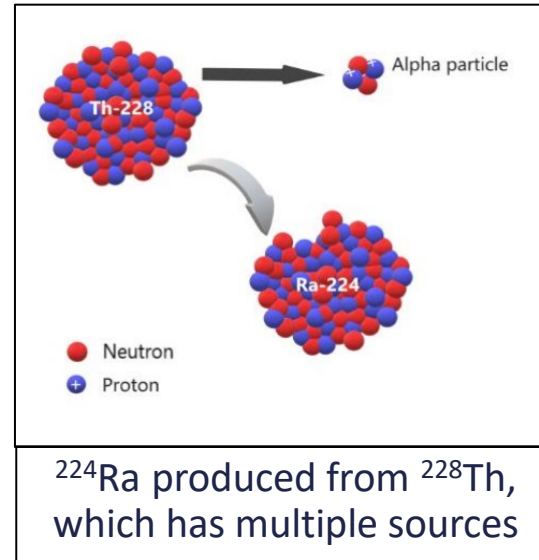


Full BIOS available at:
www.oncoinvent.com

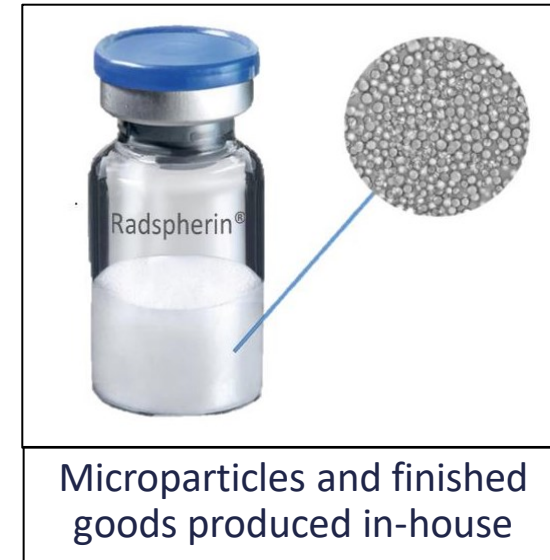
In-house Production at GMP Facility



Oncoinvent has in-house GMP Production Capability



^{224}Ra produced from ^{228}Th , which has multiple sources

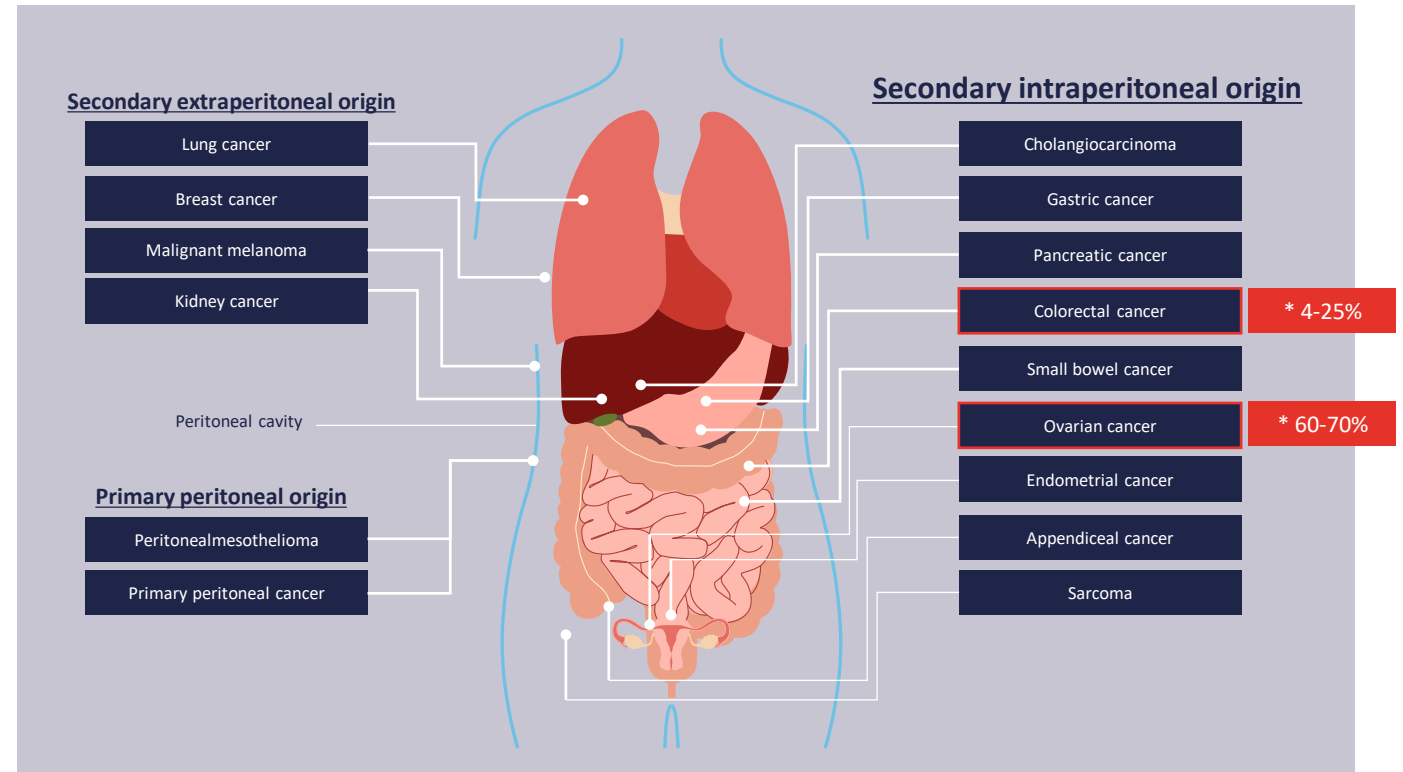


Microparticles and finished goods produced in-house

Current GMP facility can support the Clinical Phase 2b program. Outsourcing and scale-up required for Phase 3.

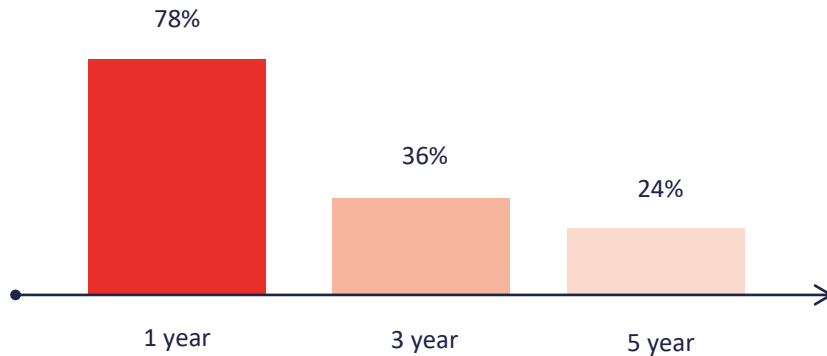
Target Area - Peritoneal Carcinomatosis (PC)

- Many underlying cancers types
- Peritoneal Metastases common at diagnosis*
- Cytoreductive Surgery is Standard of Care
- Considerable Patient Population
- No approved specific treatment
- High Unmet Medical Need
- **Significant Market Potential**

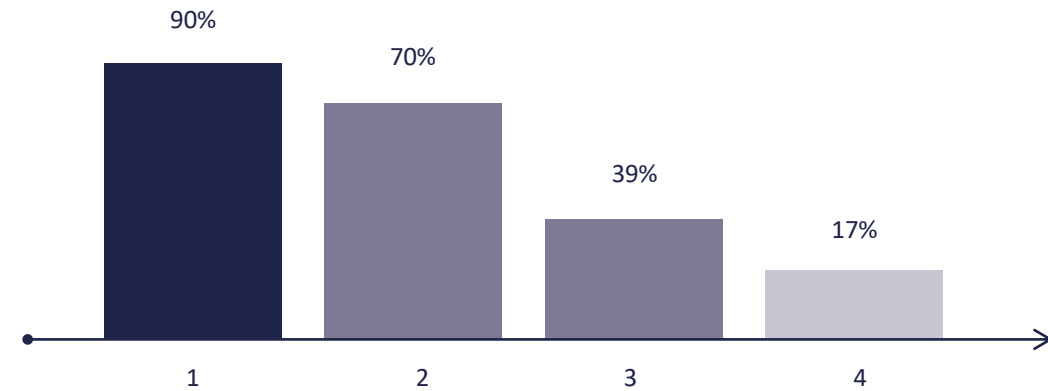


High Unmet Medical Need

Long-term survival rates of patients with PC of colorectal origin¹



Five-year survival rates in ovarian cancer² by disease stage



“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

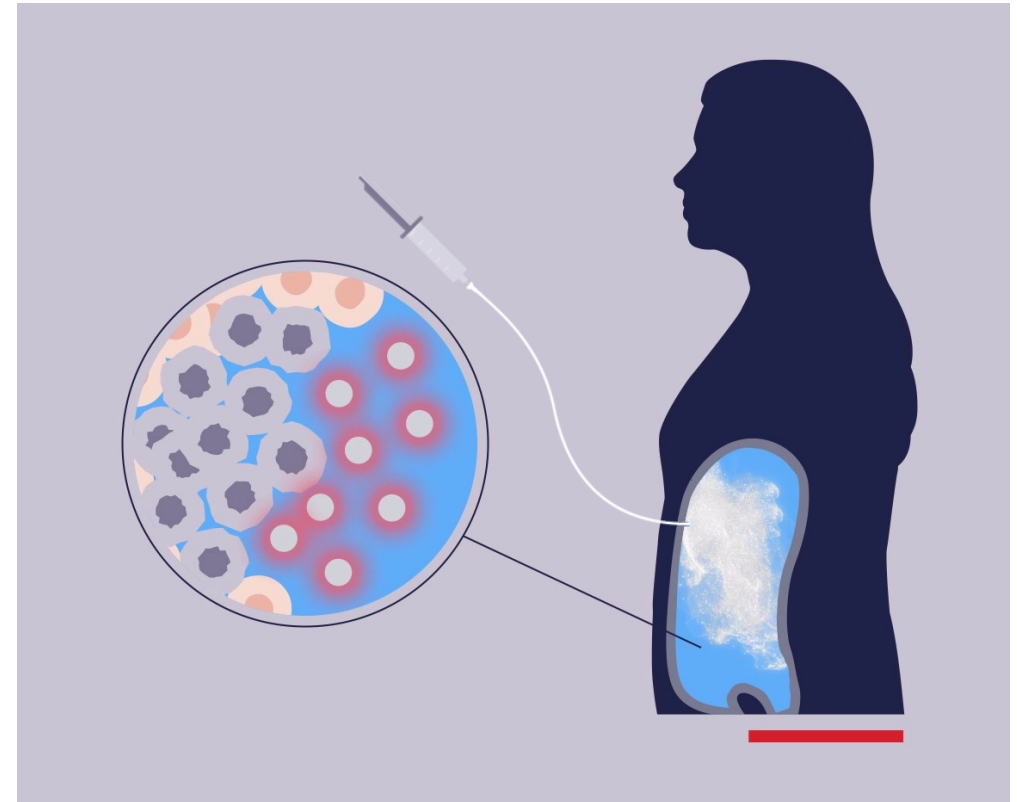
¹ source: <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4655111/>

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

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

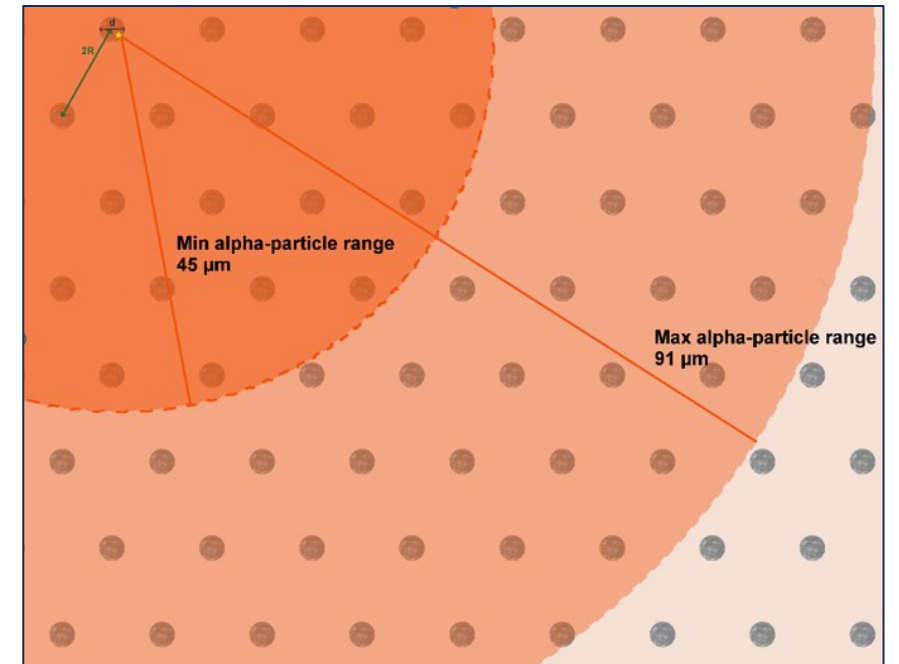
Clinical Administration of Radspherin®

- Radspherin® is shipped as dispersion of microparticles in a vial
- Single Dose Administration of 7 Mbq through standard catheter 1-3 days after surgery to eradicate micro metastases
- No incremental invasiveness or hospitalization
- Can be administered bed-side
- No special precautions
- **Easy-to-use, non-invasive, alpha-emitting radiotherapy with local administration**



^{224}Ra alpha-emitter ideal for large body cavity surfaces

- **High Radiation Energy** – Effectively treating a large body cavity with minimal exposure to other organs
- **Shallow Radiation Depth** – Limited collateral tissue damage
- **Convenient half-life** – A 3.6-day half-life means that Radspherin[®] can have ca. 10 days of clinically effective radiation and up to 8 days shelf life
- **Receptor Independence** – No cell specific target needed
- **Microparticle Depot** – Retains radiation, CaCO_3 microparticles naturally absorbed once radiation has reached sub-clinical levels
- **Alpha-radiation is highly effective, yet short-range in tissue. Ideal for large body cavity surfaces**



Alpha particles are high energy but only reach about 0.1 mm in tissue

Radspherin[®] - 68 patients treated in two indications

- Phase 1/2a fully recruited and first interim readout completed

- 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 completed Q4-23 – 68 patients treated in total. Continued stream of follow-up data.

Radspherin[®] is safe and well tolerated – reduced radiation exposure to organs vs systemic treatment

✓ Well tolerated and considered safe to use

- So far, in patients that have finished the follow up period, no dose limiting toxicities were observed, no deaths, **no serious adverse events related to Radspherin[®]**, and no discontinuations due to adverse events were reported

✓ 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, advantage over systemic treatment**

✓ Good safety profile for hospital staff

- Low amount of activity in blood and urine
- **No precautions related to external exposure required**

Absorbed Radiation Doses to Normal Organs are low

- Calculated doses well below known limiting thresholds

- None of the patients received absorbed doses higher than 1 Gy* for any normal organ
 - Biokinetic modelling resulted in highest uptake values for bone, blood, kidneys and liver
 - Highest absorbed doses to organs at risk for osteogenic cells (mean value 0.55 Gy*/7MBq), followed by liver, red marrow and kidneys (mean value ≤ 0.1 Gy*/7MBq)
 - No signs of hematological depression or negative effects on kidney or liver function observed in clinical studies

Tissue	Normal tissue tolerance for standardly fractionated external beam radiotherapy	Corresponding administered activity of Radspherin (MBq)
Colon	< 11 Gy‡	>3 000
Small intestine	≤ 15 Gy¶	>4 000
Stomach	≤ 45 Gy¶	>10 000
Liver	≤ 30 Gy¶	>400
Kidney	< 20 Gy¶	>300
	Threshold for possible major hematotoxicity§	
Red marrow	≤ 2 Gy§	~30

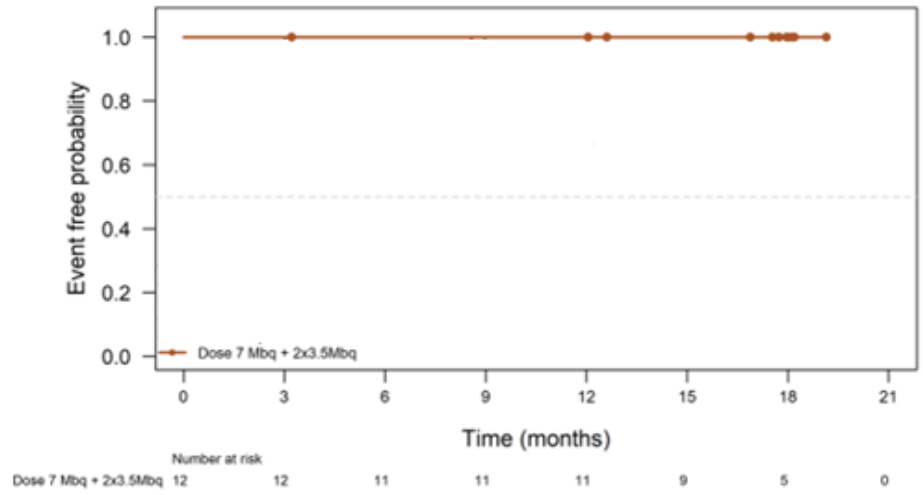
¶ From clinical experience with standardly fractionated external beam radiation therapy, Emami et al. 2013, QUANTEC group review
 ‡ From SBRT, Emami et al. 2013.
 § Hobbs, R.F., et al., A bone marrow toxicity model for ²²³Ra alpha-emitter radiopharmaceutical therapy. Phys Med Biol, 2012. 57: p. 3207-22.

*To compare doses from alpha-radiation and external beam radiotherapy or beta-radiation head-to-head, a relative biological effectiveness (RBE) factor of 5 must be used for alpha-radiation.

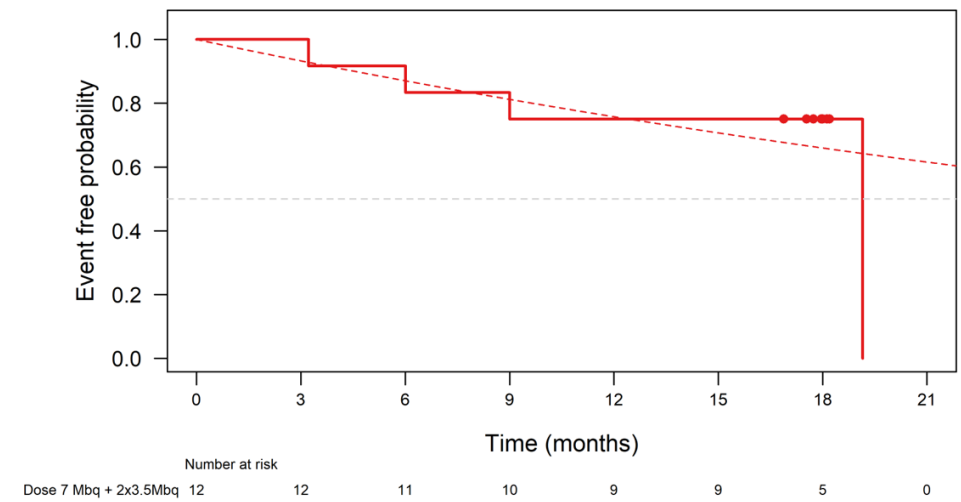
Efficacy - Data indicates substantial DFS Improvement

- Interim efficacy analysis phase 1/2a in CRC (n=22 all dose levels)

Disease-free survival at 18 months with 7 MBq Radspherin® in combination with SOC was 67 % (n=12)



Local disease-free survival: No local recurrences



Overall disease-free survival with dotted line corresponding to an expected median of 30 months (vs. 12 months expected in current SOC).

2024

2025

2026

2027

2028

2029

2030

Funding required:

\$ 10 M per year Op. Cost. Each Phase 2b trial \$ 10 M. TT cost ca. \$ 10 M → \$ 50-60 M to reach Phase 3

Milestones:

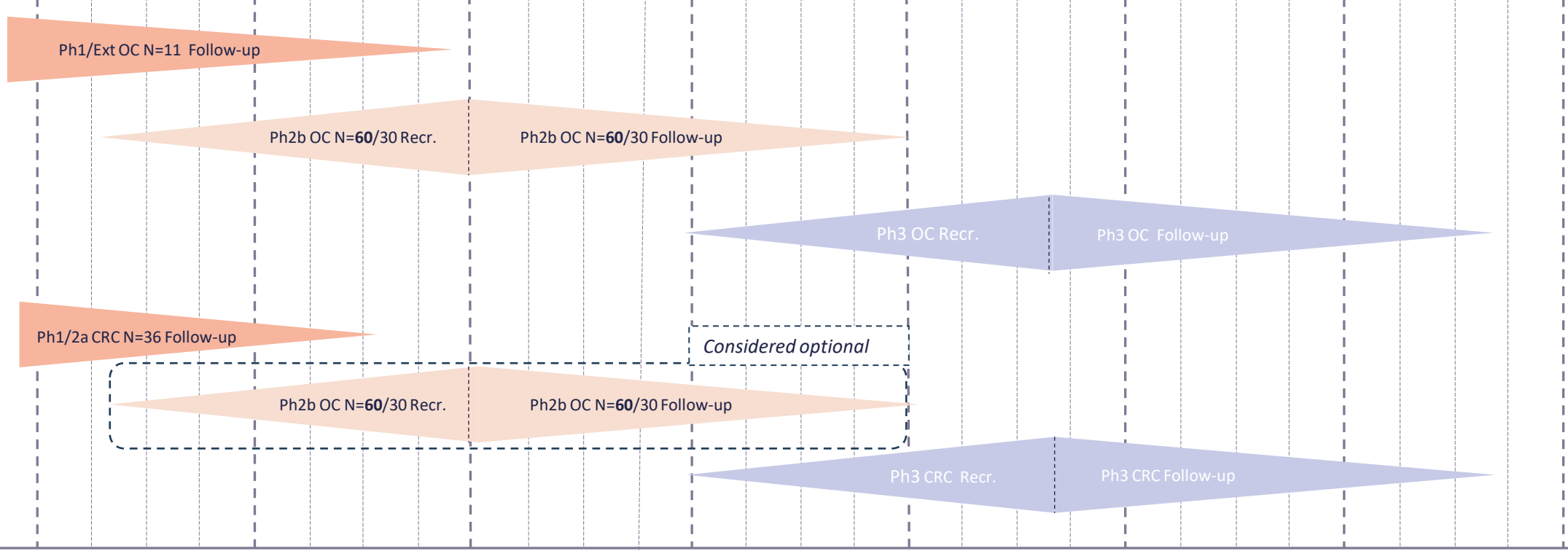


IND CTA ✓

IND CTA ✓

Ovarian Cancer (OC)

Colo-rectal Cancer (CRC)



High addressable patient numbers and unmet medical need

<u>Ovarian Cancer</u>	<u>USA</u>	<u>Europe</u>	<u>Total</u>
Patient Diagnosed (100%)	22,000	63,000	85,000
Peritoneal Mets (75%)	17,000	47,000	64,000
Eligible for Surgery (80%)	13,000	38,000	51,000
Achieve R0 (75%)	10,000	28,000	38,000

<u>Colorectal Cancer</u>	<u>USA</u>	<u>Europe</u>	<u>Total</u>
Patient Diagnosed Stage IV (100%)	39,000	113,000	152,000
Peritoneal Mets (25%)	10,000	28,000	38,000
Eligible for Surgery (90%)	9,000	25,000	34,000
Achieve R0 (90%)	8,000	22,000	30,000

Total Treatments per Year Targeted – ca. 68,000
 (in PC from *ovarian* and *colorectal* cancers only, and in the US and Europe only)

Addressing a multi billion dollar market opportunity

- Targeted Patient Population: Up to 68,000 eligible patients per year (US & Europe)
- Average Pricing Estimate: \$100,000 per treatment (conservative vs. benchmarks)
- **The total addressable market in the US and Europe alone is up to \$7 billion**
- It takes a penetration rate of only 15% to reach a billion USD and thereby blockbuster sales levels

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 is currently trending at 18 months.

Key Intellectual Property

Patent	Priority date	Area covered	Geography
WO2017005648A1	03-July-2015	To provide particles comprising a degradable compound and an α emitting nuclide and/or a radionuclide generating an α emitting daughter nuclide, or a pharmaceutical composition comprising a suspension of the particles	DK NO RS PT PL SI EP ES HU US KR JP AU CA WO MX CN RU BR CN NZ JP
WO2015044218A1	24-Sept.-2013	The present invention relates to a novel anti-CD146 antibody and derivatives thereof. The antibody and/or derivatives can be used for therapy and/or imaging, diagnosis and/or immunostaining.	EP WO DK ES US
WO2018033630A1	19-Aug.-2016	The invention relates to chimeric antigen receptor (CAR) specific to p80 and CD146, vectors encoding the same, and recombinant T cells comprising the p80 or CD146 CAR. The invention also includes methods of administering a genetically modified T cell expressing a CAR that comprises a p80 or CD146 binding domain.	WO

Patent	Priority date	Area covered	Geography
WO2022058337A1	15-Sept.-2020	The present disclosure relates to a particle comprising a degradable compound, a radionuclide, and a phosphorus containing additive. Phosphorus containing additives, such as phosphonates, have the unique ability to control the size of particles for medical applications. The applications allow for use of the particles as medicaments and for imaging, especially within the field of cancer.	WO
WO2022058338A1	15-Sept.-2020	The present invention related to a combination of radium-224 (224Ra) and/or progeny of 224Ra, and a DNA repair inhibitor for use in the treatment of cancer. The DNA repair inhibitor can for example be a poly (ADP-ribose) polymerase inhibitor (PARPi), a MGMT inhibitor, a DNA-dependent protein kinase inhibitor (DNA-PK inhibitor), an ataxia telangiectasia and Rad3-related (ATR) kinase inhibitor, an ataxia telangiectasia mutated (ATM) kinase inhibitor, a Wee1 kinase inhibitor, or a checkpoint kinase 1 and 2 (CHK1/2) inhibitor. The radium-224 (224Ra) and/or progeny of 224Ra can be comprised in nano- and/or micro sized particles.	WO

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