

Oncoinvent

Innovative alpha-emitting therapy

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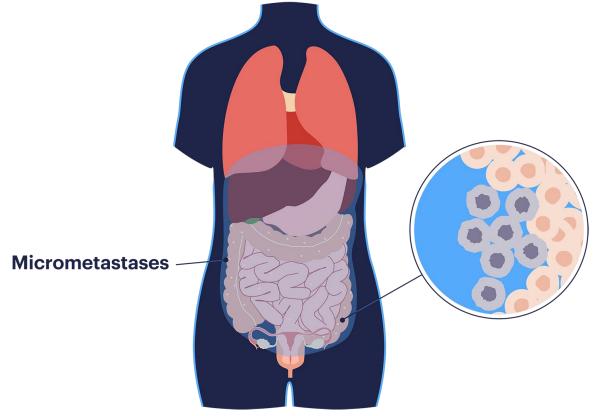
Oncoinvent at a glance





Peritoneal metastases - urgent need for novel treatments





- Peritoneal metastases are confined to the peritoneum

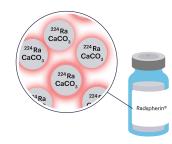
 the membrane covering the abdominal cavity and
 organs creating a 'closed compartment'
- Peritoneal metastases arise from many different primary cancers, amongst others ovarian and colorectal
- The only treatment option with curative intent is **surgery**, effect of systemic therapy limited
- Surgery is known to leave behind micro-metastases giving rise to new metastases and disease progression

Radspherin[®] - innovative alpha emitting therapy targeted to and retained in the peritoneum



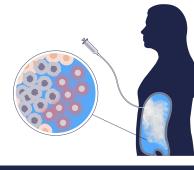
Radspherin®

- Novel formulation combining alphaemitting ²²⁴Ra with CaCO₃ microparticles
- Localized administration ensures radiation remains concentrated at the target site, minimizing off-target effects elsewhere in the body
- Good radioisotope availability and simple manufacturing
- Half-life (²²⁴Ra t_{1/2} = 3.6 days) allowing for centralized manufacturing



How does it work?

- Delivering high dose of lethal radioactivity to cancer cells in the peritoneum
- Administration 1-3 days post-operative through a catheter placed at surgery
- 75% of radiation dose delivered the first week optimal match with the post-operative treatment window
- The combination of high energy and short radiation range enables effective killing of the targeted metastases while sparing the surrounding normal tissue



Clinical development

Ovarian cancer

Phase 1: after surgical treatment of platinum-sensitive recurrence

Phase 2: after neoadjuvant chemotherapy and surgery in primary advanced ovarian cancer patients with HRD negative tumors

Colorectal cancer

Phase 1/2a: after surgery and HIPEC in patients with synchronous or metachronous peritoneal metastases

Phase 2: as above

Peritoneal metastases represent a significant market opportunity



Adds perfectly to existing patient flow

- Surgery is and will remain the cornerstone of treatment
- Treatment given 1-3 days post-operative while the patient is still hospitalized
- Simple and quick bedside administration
- Single and localized administration sustained therapeutic efficacy and decreased risk for off-target effects

Limited competition

- Distinguished by its unique mechanism of action
- Strategic advantage: complementing cytoreductive surgery, reducing threats from new therapies
- Untapped market no modern therapies and limited industry development in the specific area of peritoneal metastases

High addressable patient number

- Total treatments per year targeted more than 65,000 (ovarian and colorectal cancer) in US and Europe
- Treatment is receptor- and targetindependent –effective for peritoneal cancers regardless of origin – i.e., gastric cancer; orphan indication in the US, highly frequent in Asia, and prophylactic in highrisk patients
 - Significant potential for label expansion
- Future opportunities for tailoring to treatment of cancers in other body cavities

Potential for Radspherin® to emerge as a leading treatment option for patients with resectable peritoneal metastases

Radiopharmaceuticals have significant potential to become next generation of cancer treatments



Emergence of modern radiotherapy

- Simple, proven MOA¹) to kill cancer cells directly through DNA damage
- Traditional therapy with external beams have been used to treat cancer for **100+ years**
- Approximately 50% of all cancer patients receive radiation therapy during their course of illness²⁾
- Acceleration of field due to **novel approaches to deliver radiation directly** to tumor site

The promise of radiopharmaceuticals

- Radiopharmaceuticals use a radioactive element to deliver radiation from within or very close to the tumor, in contrast to external beam radiotherapy which directs beams from outside the body
- The internal approach with radiopharmaceuticals allows for more precise treatment, often enabling higher doses to the tumor while minimizing exposure to surrounding healthy tissue

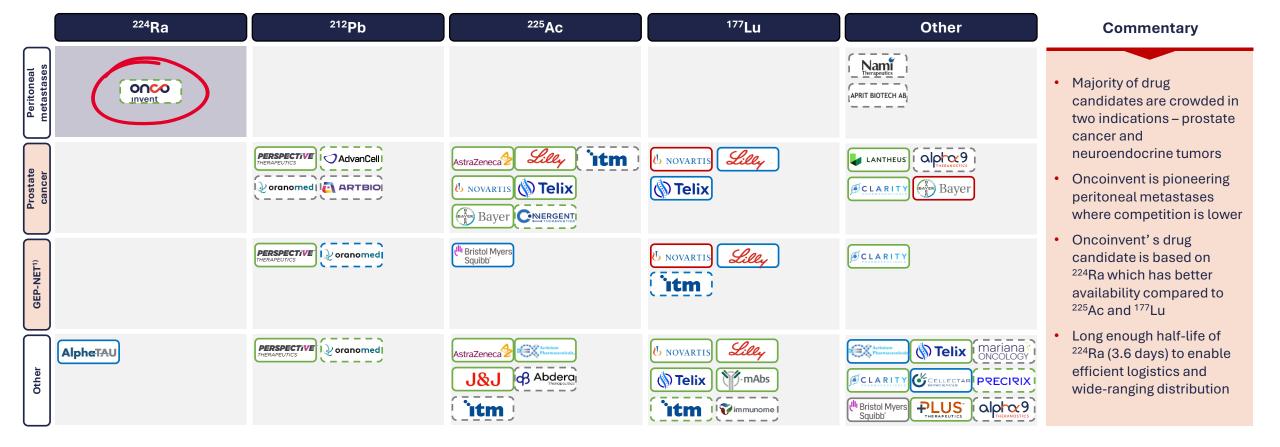
Clinical and technologic advancements have led to surge in investments, M&A and products

2) Ostuni and Taylor, Front. Med. 9:1070497

While the radiopharma sector is largely concentrated in two indications, Oncoinvent pursues peritoneal metastases



Snapshot of the Radiopharma Landscape



Notes: 1) Gastroenteropancreatic neuroendocrine tumors Source: Carnegie and DNB analysis, Guggenheim, Company websites and presentations
 Development stage
 Company type

 Preclinical
 Public

 Early Clinical
 Commercial



Radiopharmaceutical expertise at all levels



Management

Board of

Directors



Scientific founders







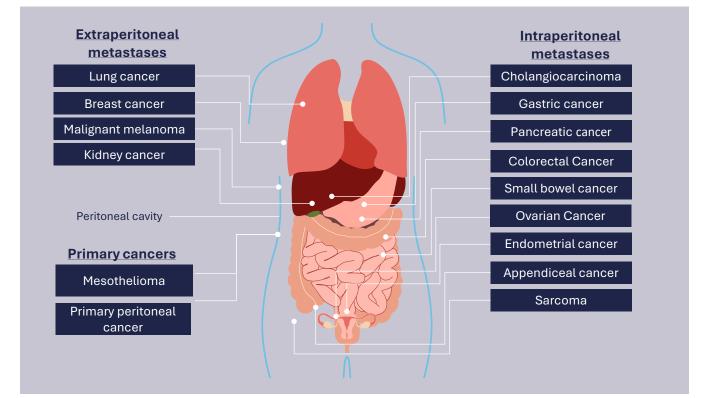


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Pipeline in one product - broad clinical application



- Peritoneal metastases arise from many different cancers
- Radspherin[®] is a receptor-independent treatment and may be effective regardless of the origin of the primary malignancy

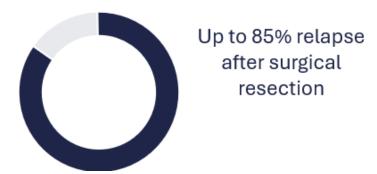


The main cause of death in ovarian cancer





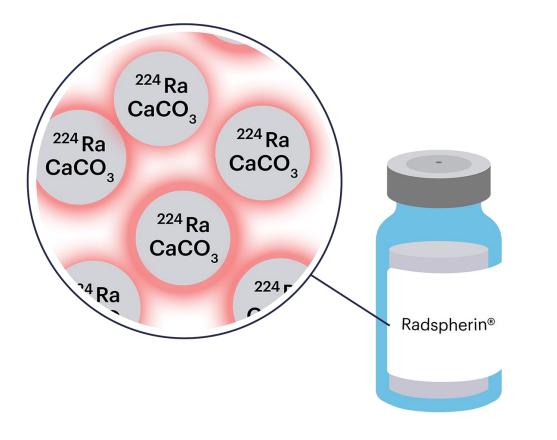
70% of all ovarian cancer patients have peritoneal metastasis at diagnosis



- Despite a comprehensive treatment approach, the majority of patients experience disease recurrence
- Ovarian cancer rarely metastasize hematogenously, recurrences almost exclusively confined to the peritoneum
- Need for improved first-line treatments that keep patients in remission – local control in the peritoneum is key to improving life expectancy

Radspherin[®] - innovative alpha emitting therapy targeted to and retained in the peritoneum



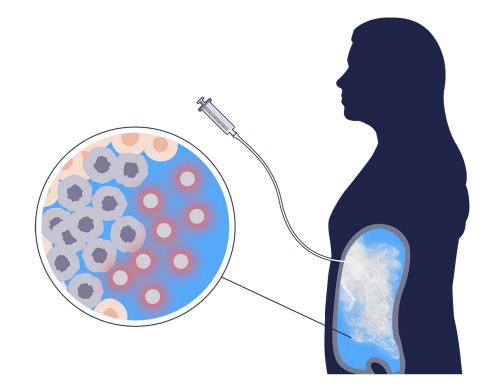


- Each vial consists of >1 billion CaCO₃ microparticles labelled with alpha-emitting ²²⁴Ra
- Ready-to-use, **single dose** delivered directly to the peritoneum
- Shelf life up to 8 days shipped as **finished goods** worldwide

Eliminating residual micro metastases after surgery

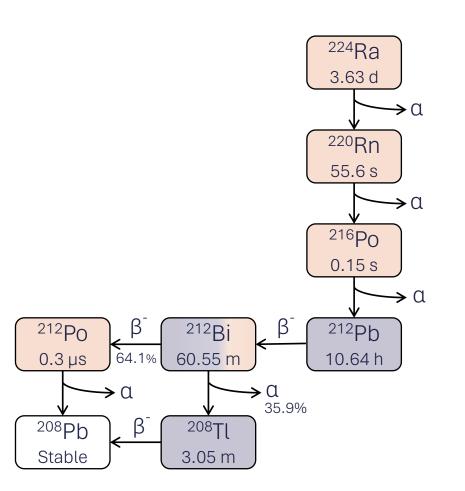


- Administration **1-3 days** post-operative through a catheter placed at surgery
- The microparticles **retain** radiation in the abdominal cavity, minimizing off-target exposure
- Alpha radiation has a very short penetration depth (<0.1 mm)
 - effectively killing cancer cells with double stranded DNA breaks
 - **limiting the risk** of harming healthy tissue protected by the peritoneal lining



Potent alpha-radiation dose delivered in the immediate post-operative invent treatment window

- Radium-224 half life 3.6 days
- Decay of radium-224 to stable lead-208 via shorterlived daughters
- The amount of lead resulting from a dose of Radspherin[®] is well below toxic levels
- **93%** of the total decay energy emitted as alpha particles



Maximized dose to the target intraperitoneal tumors versus non-target invent sites



Targeting by proximity – brilliant in its simplicity





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Radspherin[®] - in development for two lead indications



Patients undergoing surgical resection for peritoneal metastases

Ovarian cancer

Phase 1: after surgical treatment of platinum-sensitive recurrence

Phase 2: after neoadjuvant chemotherapy and surgery in primary advanced ovarian cancer patients with HRD negative tumors

Colorectal cancer

Phase 1/2a: after surgery and HIPEC in patients with synchronous or metachronous peritoneal metastases

Phase 2: as above

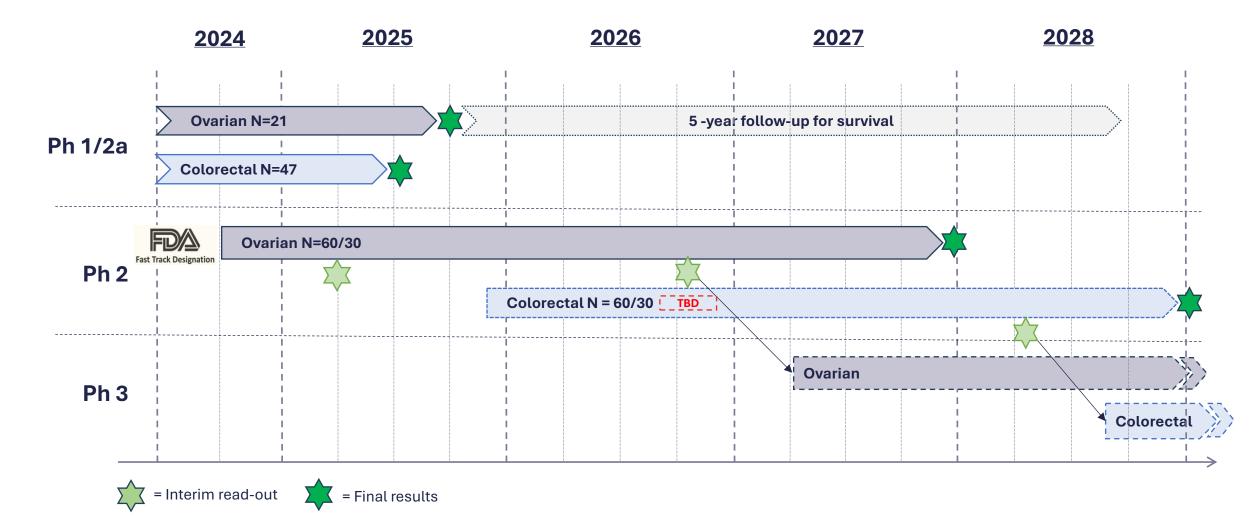
HRD negative tumors: limited/no effect of PARP inhibitors, 'the new platinum-resistant population'

HIPEC: hyperthermic intraperitoneal chemotherapy Surgery + HIPEC considered SoC in most centers and included in guidelines

Colorectal cancer



Clinical development plan







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PHASE 1/2A RESULTS

invent

Safety profile validated in two phase 1/2a studies treating 68 patients

invent	
 No dose limiting toxicity Only two out of 40 serious adverse events reported as possibly related to Radspherin* 	

\checkmark	No evidence of s	systemic	radiation	toxicity

Well tolerated and safe to use

Good safety profile for hospital staff

80% of radiation dose retained in the peritoneal cavity

Absorbed doses to other organs well below toxicity levels

- Low radioactivity dose in blood and urine
- No precautions related to external exposure required

Strong safety profile demonstrated in the completed phase 1/2a studies ovarian and colorectal cancer

• *Per cut-off date of annual DSUR 3 March 2024

-one event of small bowel perforation, 72 days after Radspherin administration - one event of procedural complication during Radspherin administration (disconnection syringe-catheter)

PHASE 1/2A RESULTS

Microparticle retention limits off-target organ exposure



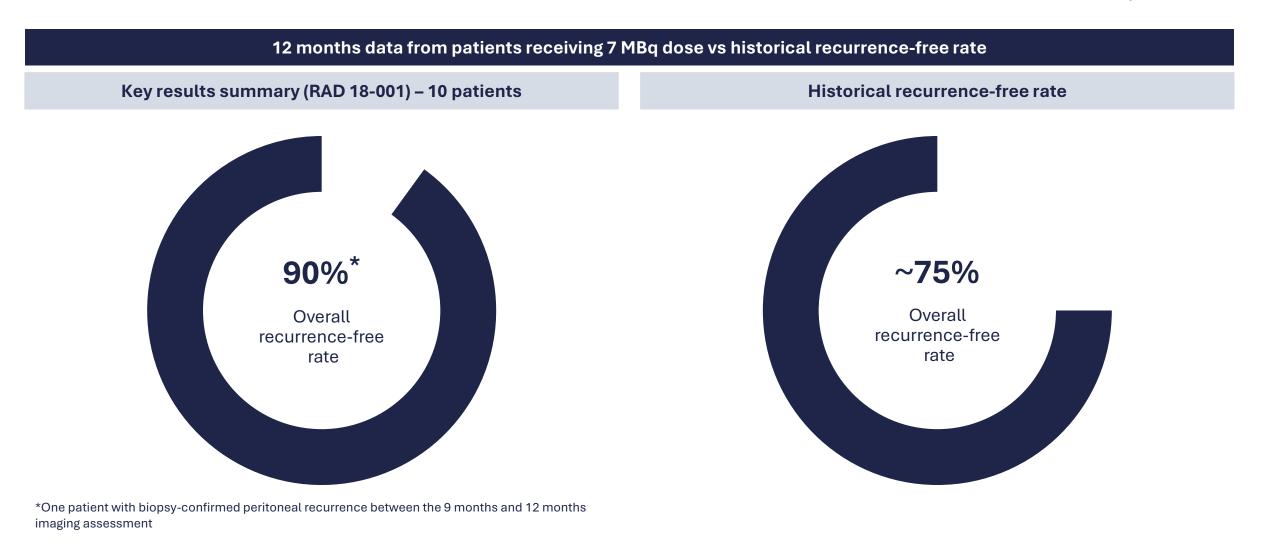
- Absorbed doses **below 1 Gy*** for all organs measured
 - Highest absorbed doses to organs at risk for osteogenic cells, followed by liver, bone marrow and kidneys
- No signs of hematological, kidney or liver toxicity observed in clinical studies

Tissue	Tolerance levels for 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

Encouraging signal of efficacy in phase 1 in ovarian cancer

Ovarian cancer



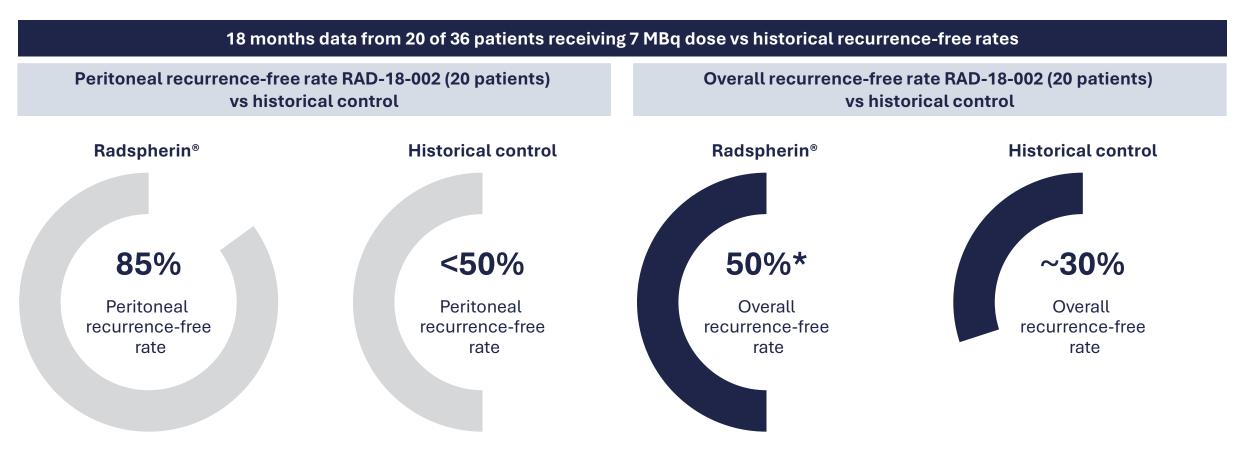


Coleman et al. N Engl J Med. 2019 Nov 14;381(20):1929-1939 Harter et al. N Engl J Med. 2021 Dec 2;385(23):2123-2131 Shi et al. Lancet Oncol. 2021 Apr;22(4):439-449

Peritoneal control in colorectal cancer

Peritoneal recurrence after surgery halves life expectancy vs non-peritoneal recurrence (21 vs 43 months)





*One pts recorded with progression after 4 weeks (liver met.). At 3 months, no radiological peritoneal recurrence noted, but peritoneal recurrence confirmed histologically after surgery 2 weeks later. The patient withdraw from the study after surgery



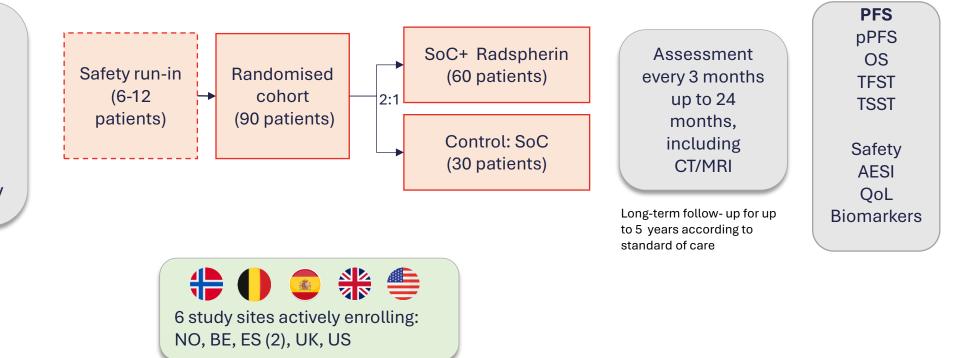


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Phase 2 study in ovarian cancer – enrollment started



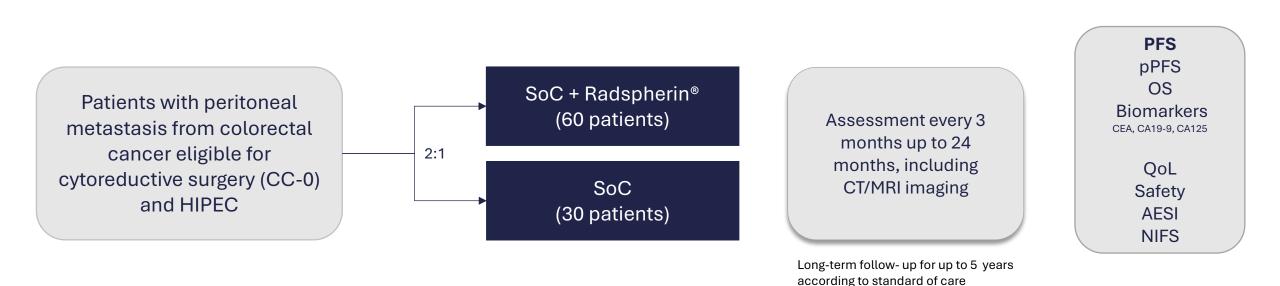
Patients with primary advanced HRD negative ovarian cancer with peritoneal metastases eligible for complete resection (R0) after neoadjuvant chemotherapy



PFS: Progression Free Survival pPFS: peritoneal Progression Free Survival OS: Overall Survival AESI: Adverse Event of Specialized interest TFST: time to first subsequent anticancer therapy or death TSST: time to second subsequent anticancer therapy or death

HRD: Homologous Recombination Deficiency IDS: Interval Debulking Surgery

Phase 2 study in colorectal cancer – IND and CTAs in place



PFS: Progression Free Survival pPFS: peritoneal Progression Free Survival OS: Overall Survival AESI: Adverse Event of Specialized interest

NIFS: Next Intervention Free Survival CRS: cytoreductive surgery HIPEC: heated intraperitoneal chemotherapy







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High addressable patient numbers with unmet need



Ovarian cancer	USA	Europe	Total
Patient diagnosed (100%)	22,000	63,000	85,000
Peritoneal metastasis (70%)	15,000	44,000	59,000
Eligible for surgery (80%)	12,000	35,000	47,000
Achieve complete resection (75%)	9,000	26,000	35,000

Colorectal cancer	USA	Europe	Total
Patient diagnosed stage IV (100%)	39,000	113,000	152,000
Peritoneal metastasis (25%)	10,000	28,000	38,000
Eligible for surgery (90%)	9,000	25,000	34,000
Achieve complete resection (90%)	8,000	22,000	30,000

Total treatments per year targeted – ca. 65,000

(in peritoneal metastasis from ovarian and colorectal cancers only, and in the US and Europe only)

A multi billion USD market opportunity



- Targeted patient population: Up to 65,000 eligible patients per year
- Average pricing assumption: \$100,000 per treatment (conservative vs. benchmarks)
- The total addressable market in the US and Europe, for two cancer types in US and EU only, is more than \$6 billion

At 15% penetration, sales surpasses \$1bn

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





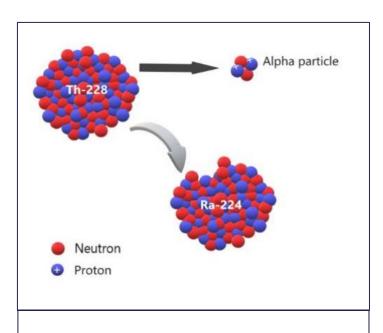
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In-house production at GMP facility

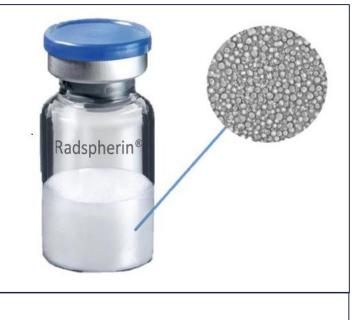




Oncoinvent has in-house GMP production capability



²²⁴Ra produced from ²²⁸Th, which has multiple sources

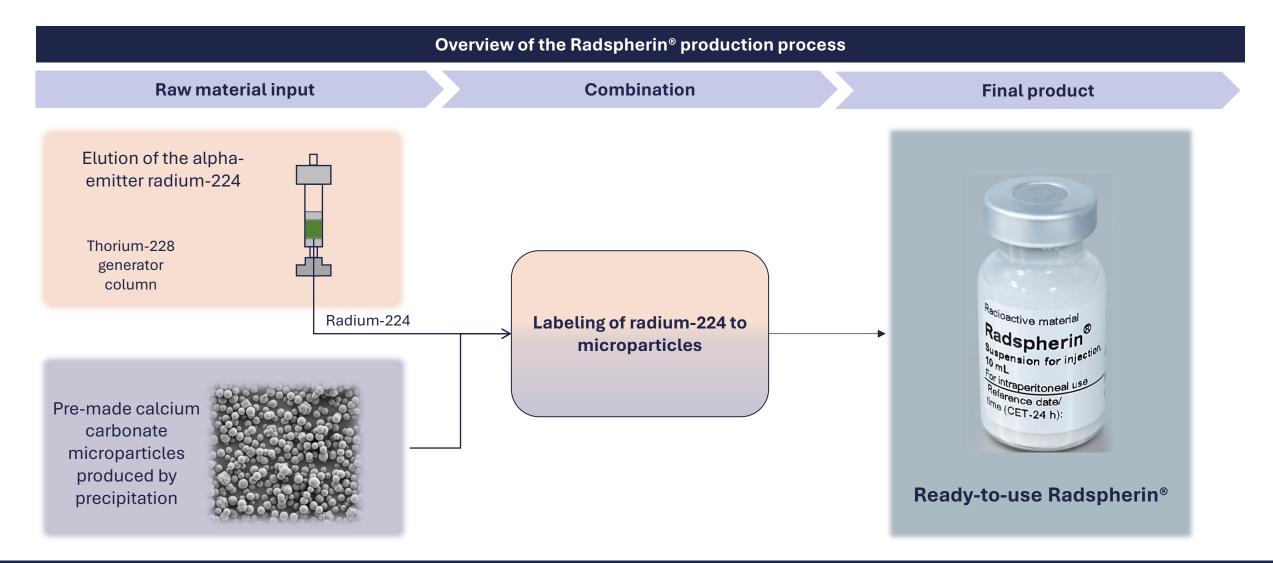


Microparticles and finished goods produced in-house

Current GMP facility supplies phase 2, outsourcing and scale-up required for phase 3

Radspherin® production process









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Radspherin® - solid multilayer intellectual property protection



Radspherin[®] composition of matter & use

- Granted in US, EU, China, Japan and additional countries
- Patent expiry 2035 (2036 in some countries) with an option for 5 years extension

Radspherin® formulation

- Filed in 2021 in: USA, Europe, Japan, China, Canada, India, Mexico, Hong Kong
- Patent expiry 2041 with an option for 5 years extension

Radspherin® clinical doses, application: use patent

- Filed in January 2024
- Patent expiry: 2044 with an option for 5 years extension

Radium-224 combination with PARP inhibitors

- Filed in 2020 in: USA, Europe, Japan, China, Canada, Mexico, Hong Kong
- Patent expiry 2041 with an option for 5 years extension

Composition of matter

Formulation

Clinical dosage/use

Combination



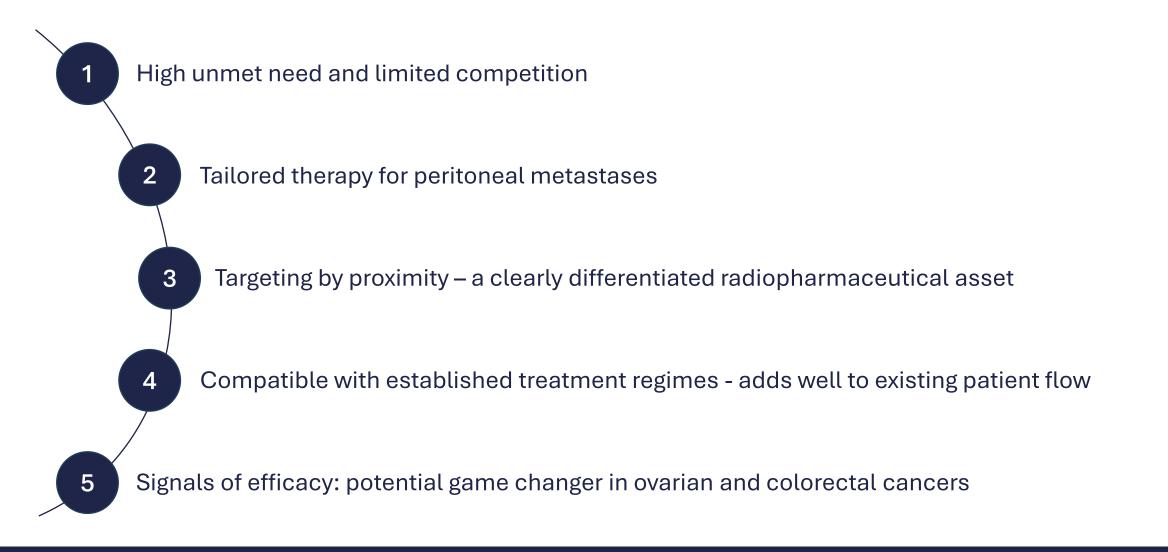


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SUMMARY

A unique radiopharmaceutical opportunity









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APPENDIX

Acquisition Price (\$ Bn)

b NOVARTIS

\$3.9

(2017)

Despite strong M&A activity within the radiopharma sector there is still significant headroom for further acquisitions



Summary of M&A Activity
Overview of Radiopharma Exposure

(* Bristol Myers Squibb"
\$4,1

Volvertial
Commercial
Late-Stage
Late-Stage
Volvertial
Late-Stage
FPI-2265, FPI-1434, FPI-2059, FPI
Late-Stage
PNT2002
PNT2002
PNT2003, PNT2004, PNT200
PNT2003, PNT2004, PNT200
PNT2002
PNT2003, PNT2004, PNT200
PNT204



(2024)

 $(2024)^{1}$

(2023)

	Commercial	Late-Stage	Early-Stage	Precunicat	
U NOVARTIS	Pluvicto, Lutathera		Lu-NeoB, ²²⁵ Ac-PSMA-617, FAP-2286	MC-339	
AstraZeneca			FPI-2265, FPI-1434, FPI-2059, FPI-2068		
Lilly		PNT2002	PNT2003, PNT2004, PNT2001		
(^{III} Bristol Myers Squibb"		RYZ101		Glypican-3	
BAYER R BAYER	Xofigo		BAY3546828, BAY3563254, BAY270439		
S MERCK			JNJ-69086420		
Johnson&Johnson	Series A investment in Aktis Oncology				
sanofi	Partnership with Orano Med and RadioMedix				
🧭 GILEAD					
abb∨ie	Key glo	bal biopharma	a companies with oncology presend	e	
P fizer		but no cur	rent radiopharma pipeline		
AMGEN					
GSK					
Datkh-Sankyo					
Roche					

(2021)