

AlphaTAU

(NASDAQ:DRTS)

Revolutionary Alpha-Emitters Radiotherapy

KOL Investor Event

July 18, 2022

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Today's Presenters



Raphi Levy
CFO
Alpha Tau



Dr. Robert Den
Radiation Oncologist
*Alpha Tau Chief Medical
Officer*



Prof. Michael J. Zelefsky
Radiation Oncologist
*Memorial Sloan Kettering
Cancer Center
Alpha Tau Scientific
Advisory Board*



Dr. Marc D'Andrea
Radiation Oncologist
*University Cancer
Centers*

AGENDA

1

Intro to Alpha Tau
Raphi Levy, CFO

11:00 – 11:15

Systemic Effect

2

Overview of Potential Systemic Effect
Dr. Robert Den, CMO

11:15 – 11:25

Local Effect

3

Results from Previous Clinical Studies
Dr. Michael Zelefsky, Memorial Sloan Kettering Cancer Center

11:25 – 11:40

4

User Experience & Selected Case Studies
Dr. Mark D'Andrea, University Cancer Centers

11:40 – 11:55

5

Upcoming Clinical Studies, Including US Pivotal Study
Dr. Robert Den, CMO

11:55 – 12:15

Closing Remarks

6

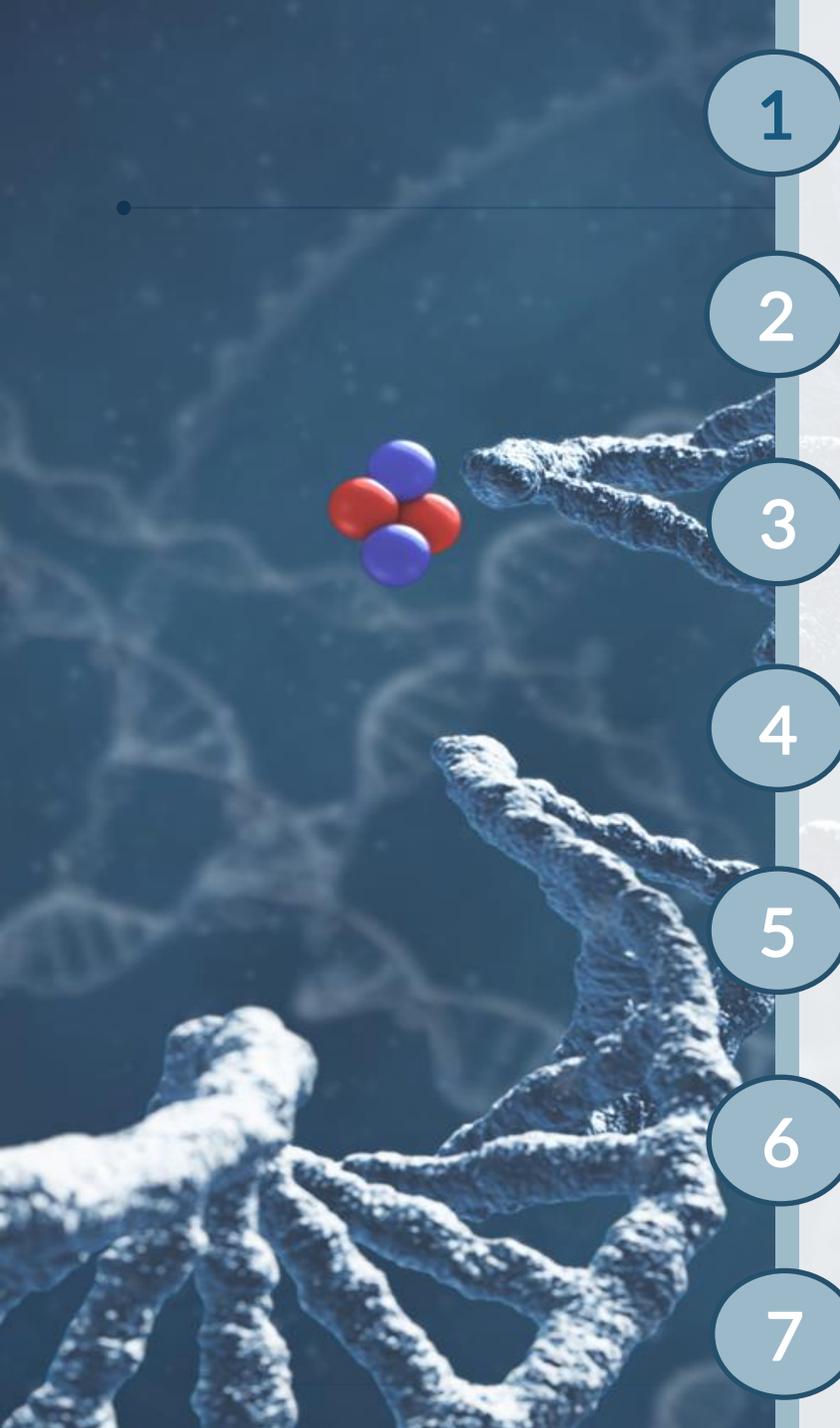
Future Outlook
Raphi Levy, CFO

12:15 – 12:20

7

Q & A

12:20 +



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Intro to Alpha Tau

Raphi Levy, CFO

Alpha Tau – Key Investment Highlights

1

Proprietary Alpha DaRT designed to safely deliver alpha radiation with localized precision in solid tumors, sparing surrounding healthy tissue

2

Broad potential and preclinical evidence supporting evaluation across various solid tumors (skin, pancreas, breast, GBM, etc.) with 18 peer-reviewed pre-clinical papers

3

Compelling potential immuno-stimulatory effect and synergetic combination with other therapies

4

Exhibited 100% ORR and ~78% CR in first-in-human clinical trial in 28 SCC tumors. Over 100 superficial tumors treated to date, with a similar profile observed. 100% CR seen at 12 weeks in 1st US study

5

Favorable safety profile observed, no systemic toxicities

6

Robust clinical-trial strategy with leading global centers, with U.S. pivotal study forthcoming in recurrent cutaneous SCC. Two FDA Breakthrough Device Designations (skin & GBM)

7

Solid logistics based on purpose-built manufacturing facilities, built or in planning, in the US, Israel and Asia, with a highly scalable and optimized proprietary production process

8

Strong intellectual property (method and device) with over 160 issued and pending patents worldwide

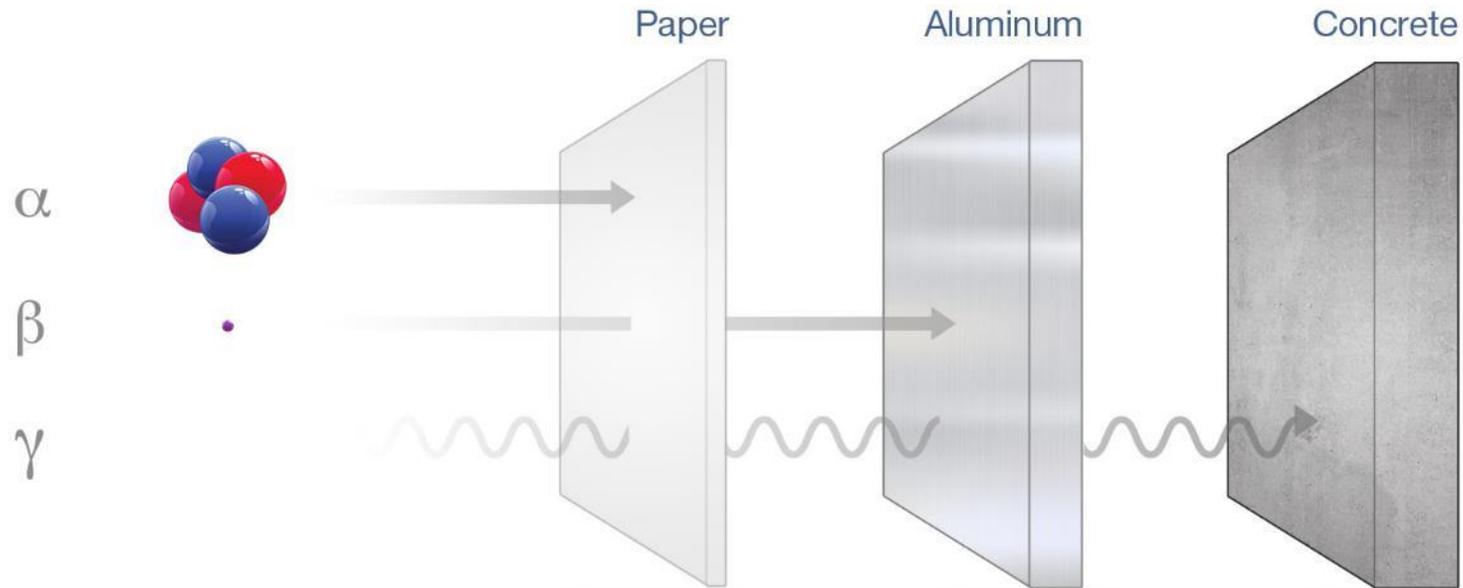
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Experienced management team, including Alpha DaRT's co-inventors, with expertise in oncology development, manufacturing scale up and commercialization

AlphaTAU

Types of Radioactive Decay

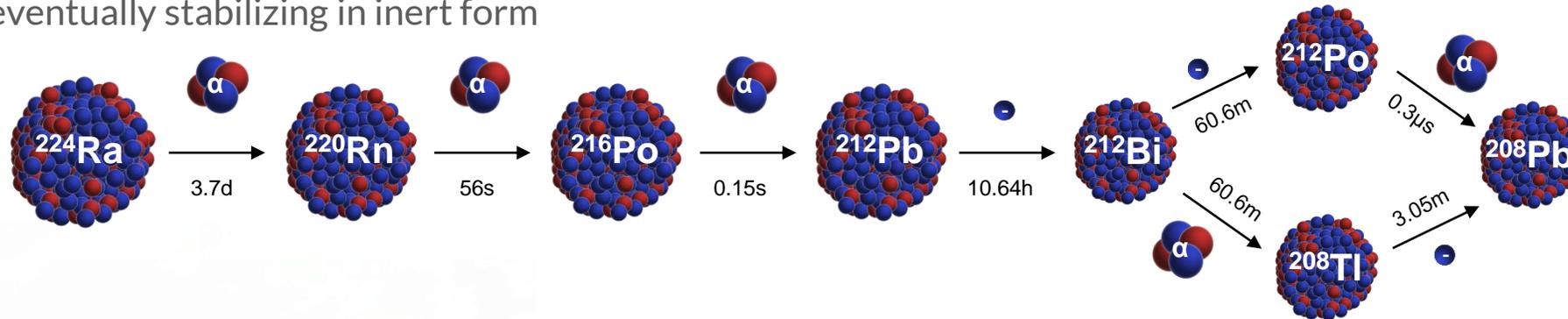
Due to the mass of the alpha particle, in comparison to beta particle, alpha has a low penetration power. This means that the outside layer of the human skin, for example, can block these particles.



Mechanism of Action of the Alpha DaRT Technology

²²⁴Ra Decay Chain

- Alpha DaRT leverages the innate decay chain of Radium-224
- The decay chain of Radium-224 includes four alpha particles
- Radium-224 has a half-life of ~3.7 days, while the remaining decay chain has a total half-life of approximately 12 hours, before eventually stabilizing in inert form



Alpha DaRT

- The Alpha DaRT utilizes stainless steel sources that are impregnated with Radium-224
- The source is designed such that when injected into the tumor, the radium remains attached to the source while its daughter atoms detach, emitting cytotoxic alpha particle payloads as they move deeper into the tumor until stabilizing

Alpha DaRT is designed to overcome the range limitations of alpha particles through precise release of alpha emitters into the tumor, generating a potent and tight distribution of alpha radiation

Alpha DaRT - Diffusing Alpha-emitters Radiation Therapy

<https://www.youtube.com/watch?v=nwfzJHm0fTQ>

Intra-tumoral Delivery Methods

We Have a Total of Seven Applicators Which Have Been Developed for a Range of Potential Uses to Accommodate for:

Treatment Delivery Method

Duration of Implantation

Tumor Location

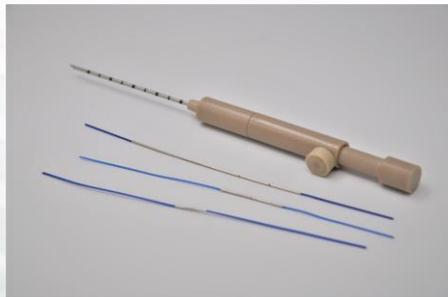
Our Applicators Allow Us Flexibility to Deliver Alpha DaRTs Into Both Superficial and Internal Tumors

Temporary Implants (Superficial Tumors)

Applicators are supplied preloaded, sealed and designed for immediate use in the procedure room

Sources are hollow and strung onto a surgical suture, allowing the clinician to insert the sources into the tumor and leave the suture in place

Alpha DaRT Needle Applicator



Needle Applicator in Action



Example Indication: Superficial Tumors.
sources are affixed to a biocompatible suture and loaded inside the needle

Permanent Implants (Internal Tumors)

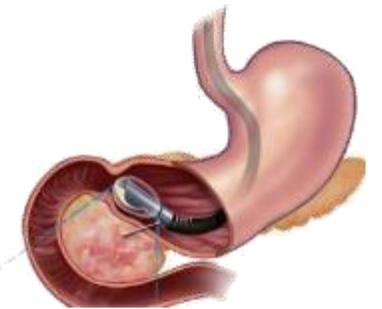
Applicators are supplied preloaded or unloaded, and are designed to allow clinicians flexibility to load the sources in the course of treatment and to select how many sources to deliver

Loading Device



Example Indication: Pancreatic Tumors.
Device is designed to be fitted to existing needles such as standard Fine Needle Aspirator (FNA) to ultimately deliver sources into the tumor

Procedure: FNA in Conjunction with Endoscopic Ultrasound



Therapeutic Focus

We are focused on delivering solutions to three markets that we believe would be best served by the unique characteristics of the Alpha DaRT

Localized & Unresectable

- Localized tumors that are not surgical candidates and tumors that recur after surgery and are **resistant to other** therapies, specifically radiotherapy
- Alpha DaRT to be evaluated as a **later line therapy**
- Tumor types include **Skin SCC, H&N SCC and prostate**



High Unmet Need

- Solid tumors that have **standard of care options**
- Alpha DaRT could potentially target **broad patient populations**
- Tumor types include **GBM and pancreatic cancer**



Metastatic

- Alpha DaRT would be evaluated for its potential to induce an **immune response** in **metastatic** tumors
- Alpha DaRT would be evaluated **in combination with check point inhibitors**
- Tumor types include **liver, breast and H&N (Head & Neck) cancers**

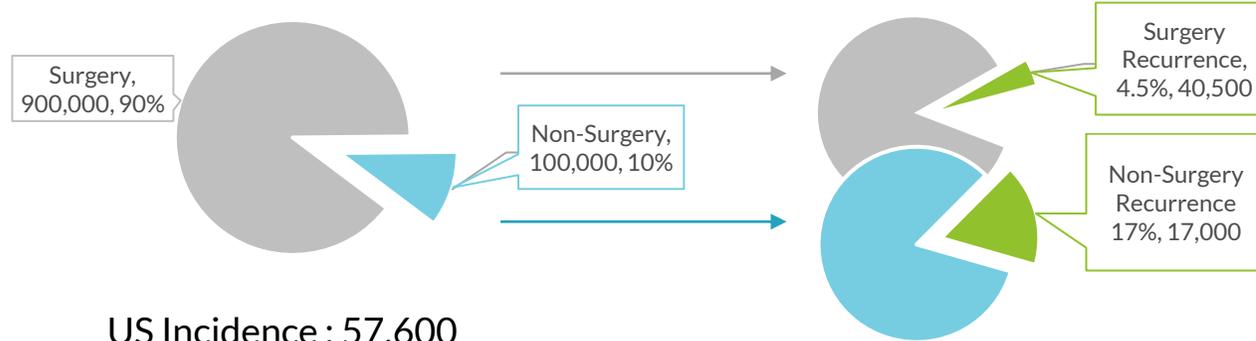
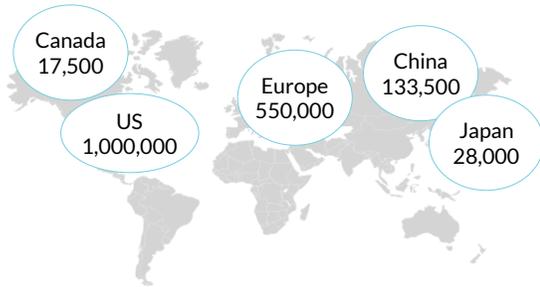


Applicable Market Size – Estimates of Annual Incidence Data

SCC / H&N Annual Incidence

US Incidence: ~1 million

US Applicable Mkt. Size

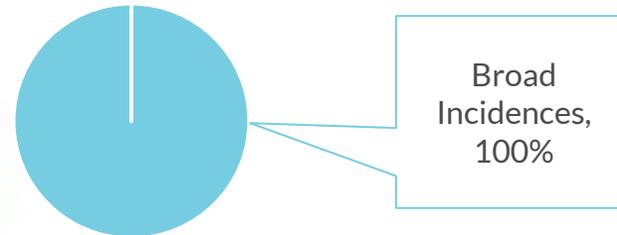
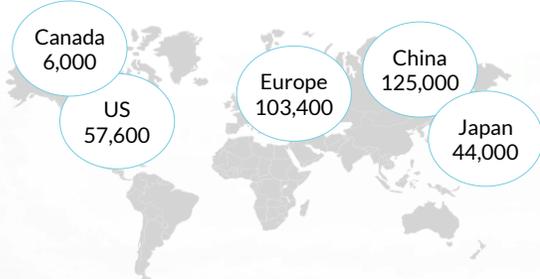


1L: 100,000
2L: 57,500

Pancreas Annual Incidence

US Incidence : 57,600

US Applicable Mkt. Size

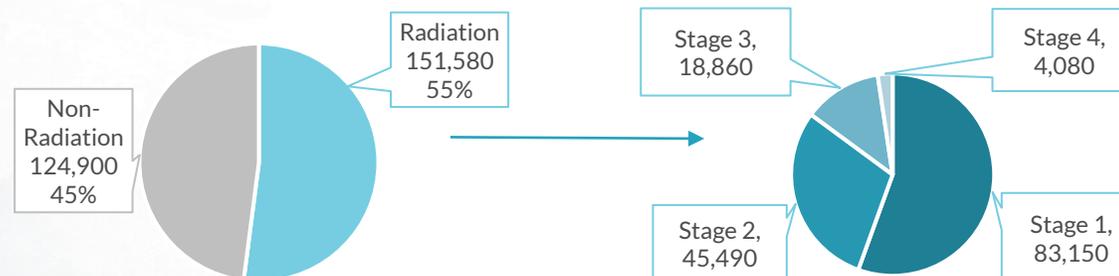
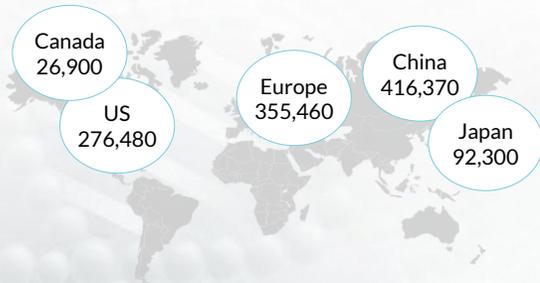


57,600

Breast Annual Incidence

US Incidence : 276,480

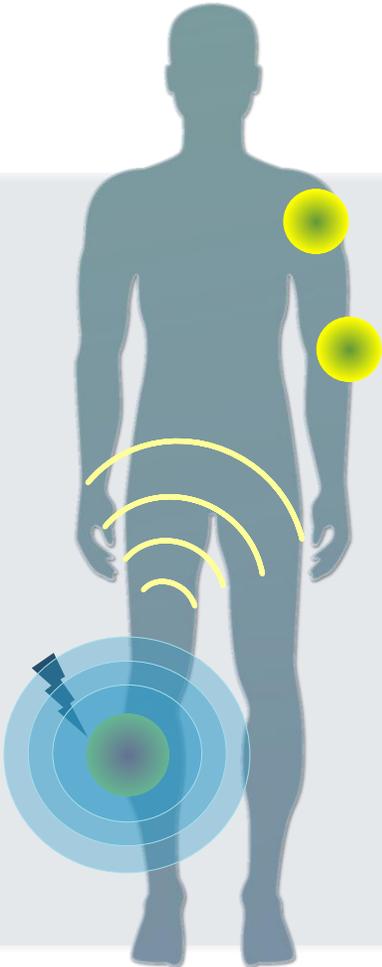
US Applicable Mkt. Size



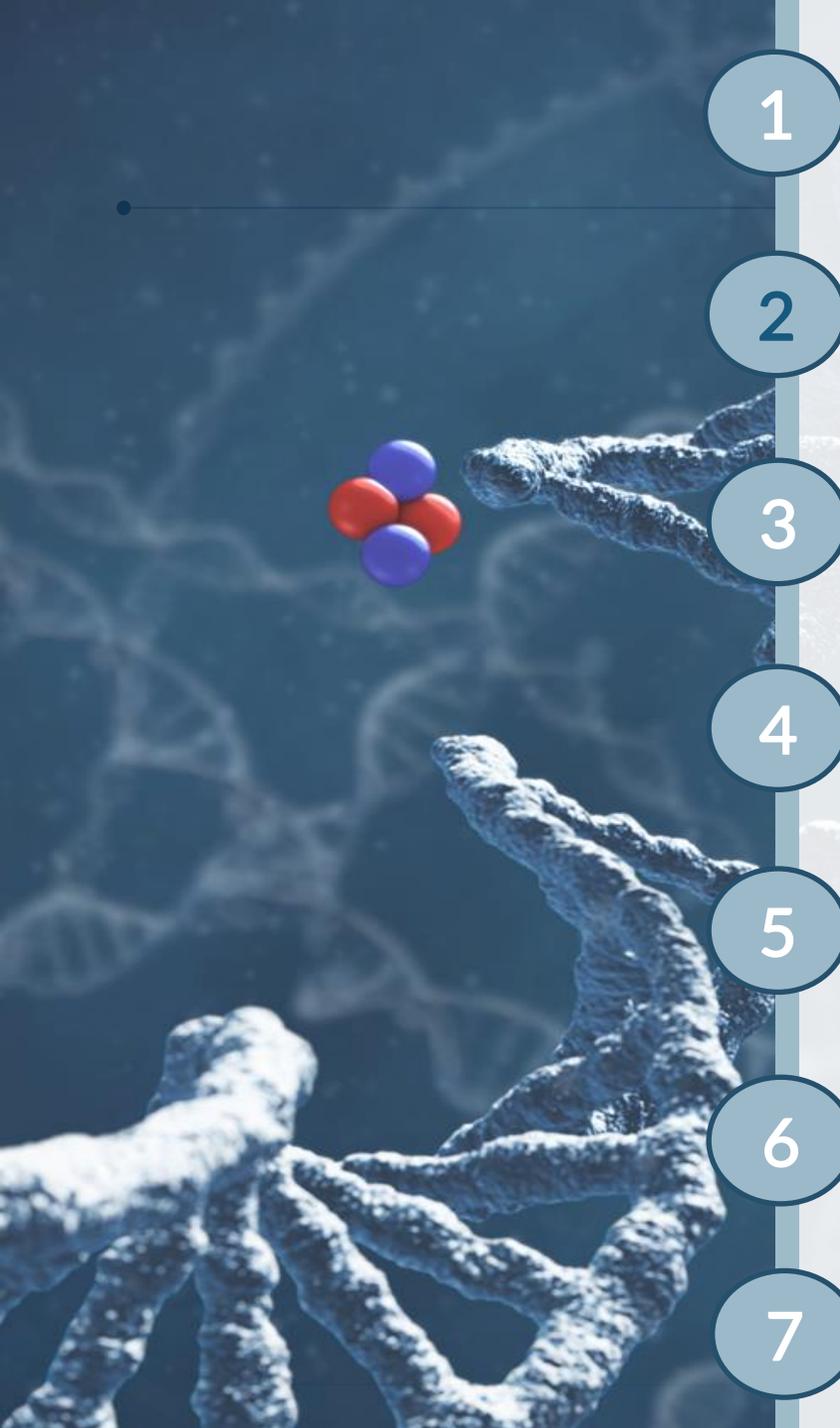
151,580

Local Control & Immune/Abscopal Effect

LOCAL CONTROL
*Dr. Marc D'Andrea &
Prof. Michael Zelefsky*



SYSTEMIC EFFECT
Dr. Robert Den



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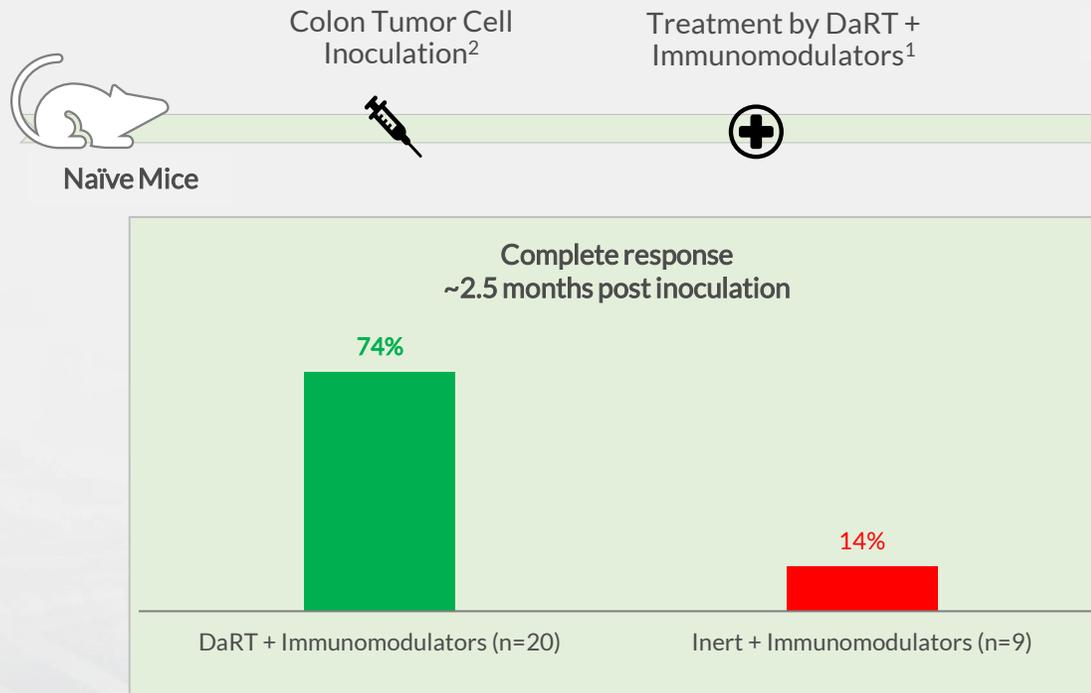
Overview of Potential Systemic Effect

Dr. Robert Den, CMO

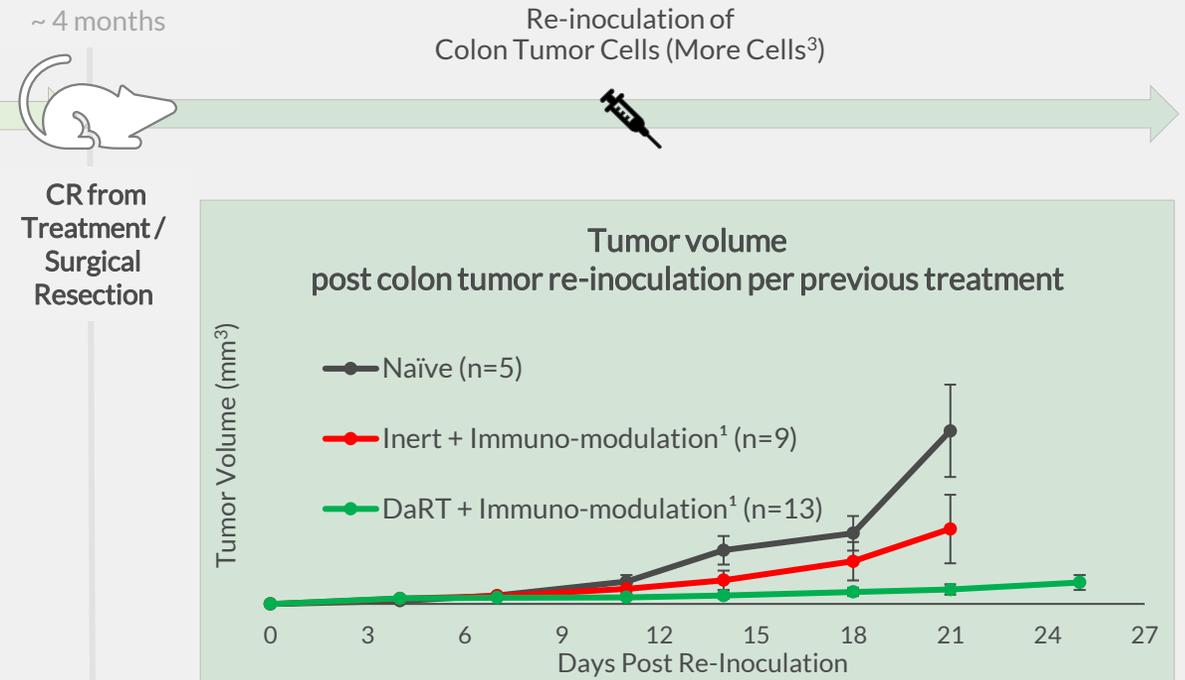
Observed Cancer-Specific Immune Protection (1/2)

In challenging mice 4 months after treatment, those previously treated by the Alpha DaRT displayed a meaningful retained protection against regrowth of the same tumor type, as compared to the two control groups

Tumor Treatment by DaRT + Immunomodulators¹



Tumor Re-Inoculation after Treatment by DaRT + Immunomodulators vs. Inert¹

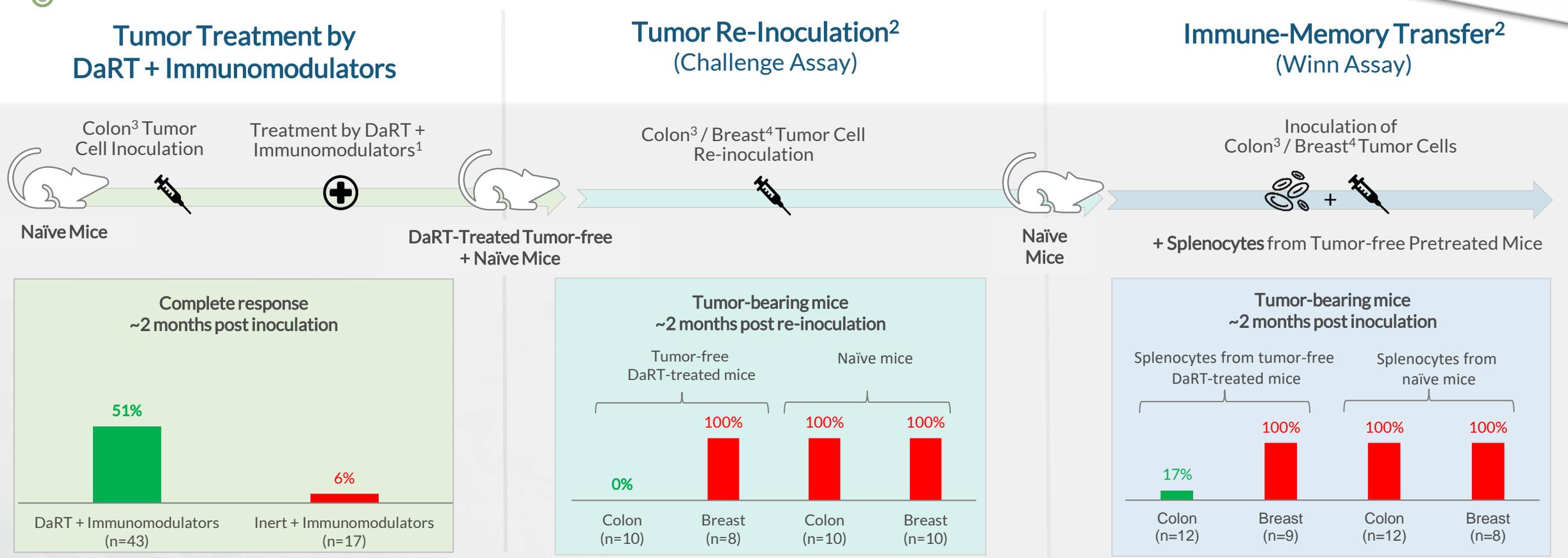


(1) Three groups of mice were inoculated with 5×10^5 CT26 tumor cells and then treated with (1) DaRT + CP, Sildenafil and 2xCpG, N=10 (2) DaRT + CP, Sildenafil and CpG, N=10 or (3) inert + CP, Sildenafil and 2xCpG, N=9. Complete responders or tumor-resected mice were re-challenged ~4 months after DaRT with 5×10^6 CT26 tumor cells.
 (2) $CT26 \ 5 \times 10^5$.
 (3) $CT26 \ 5 \times 10^6$.

Observed Cancer-Specific Immune Protection (2/2)

This activity was then shown to be tumor-specific – the challenge only resisted regrowth of the same tumor line. It was also shown to be transferrable via the transfer of splenocytes

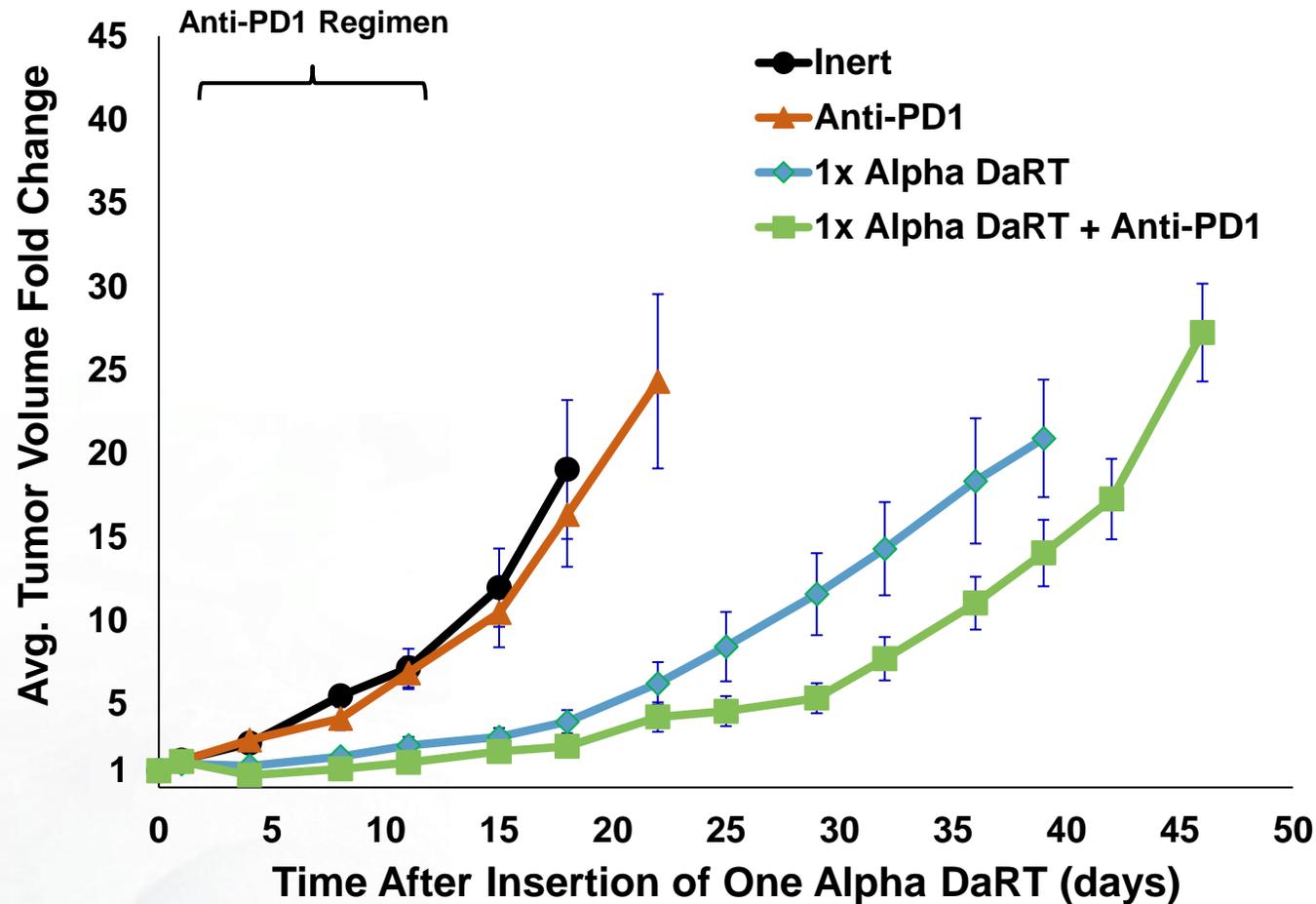
Combining alpha radiation-based brachytherapy with immunomodulators promotes complete tumor regression in mice via tumor-specific long-term immune response
 Vered Domankevich, Adi Cohen, Maroalit Efrati, Michael Schmidt, Hans-Georg Rammensee, Sujit S. Nair, Ashutosh Tewari, Itzhak Kelson & Yona Keisari



(1) Immuno-modulation refers to a combination of low dose CP, Sildenafil and CpG.
 (2) Mice with CR from DaRT + immuno-modulators (n = 18) and naive mice (n = 20) were inoculated with 5×10^5 CT26 or DA3 cells 52 days post inoculation (Challenge Assay). Naive mice were injected intradermally with splenocytes from either naive or CT26-bearing mice treated by DaRT and immunomodulators, coupled with CT26 or DA3 tumor cells (Winn assay). The presented results are based on cumulative data from two different experiments.
 (3) CT26 5×10^5 .
 (4) DA3 5×10^5 .

Alpha DaRT Elicits Effect from anti-PD1 in SCC Mouse Model (SQ2)

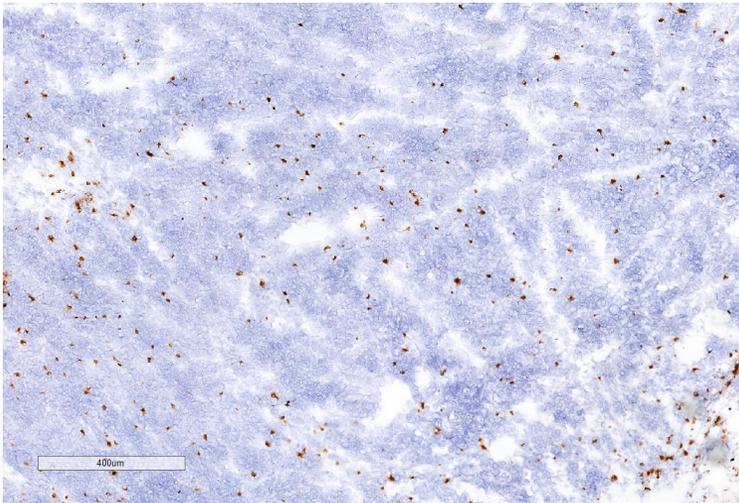
While mice with the SQ2 squamous cell carcinoma model showed little to no effect when treated with a murine anti-PD1 agent, the observed effect was larger for the combination with Alpha DaRT than for Alpha DaRT on its own



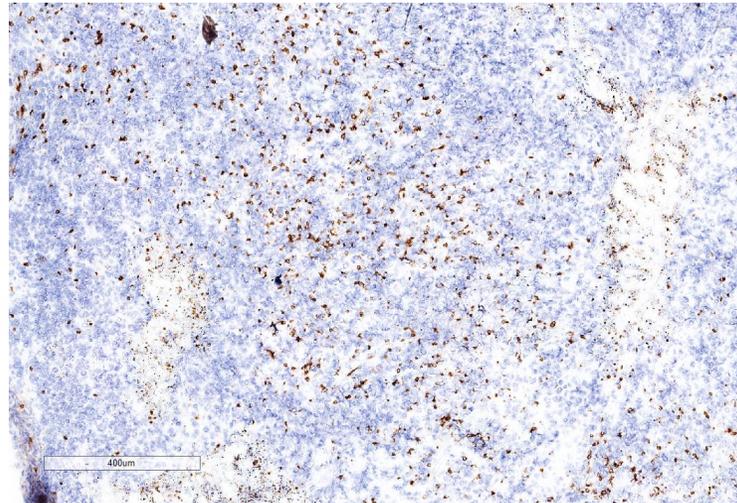
Alpha DaRT Increases Infiltration of CD3+ T-cells Into the Tumor

The combination of Alpha DaRT with anti-PD1 demonstrates the highest level of TILs in mice with SQ2 SCC tumors, potentially indicating an ability to potentiate the checkpoint blockade

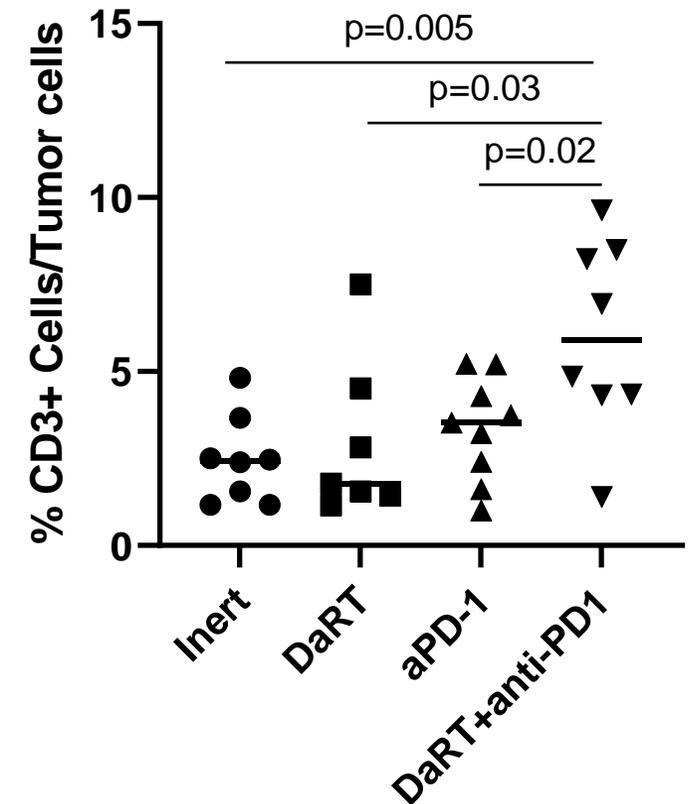
anti PD-1



Alpha DaRT + anti PD-1



TILs in SQ2 tumors



Potential Systemic Immune Effect Observed in One Patient Where a Second, Untreated Lesion Manifested CR

✔ Complete Response + Potential Systemic Immune Effect



Treated Tumor

Before

30-Nov-17



After

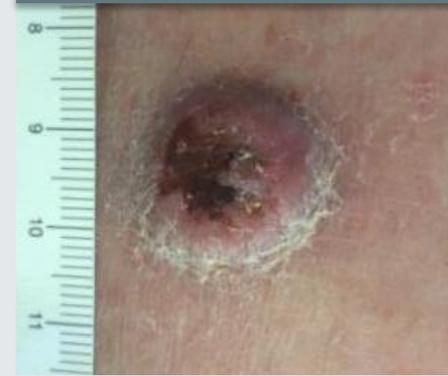
29-Dec-17



Untreated Tumors

Before

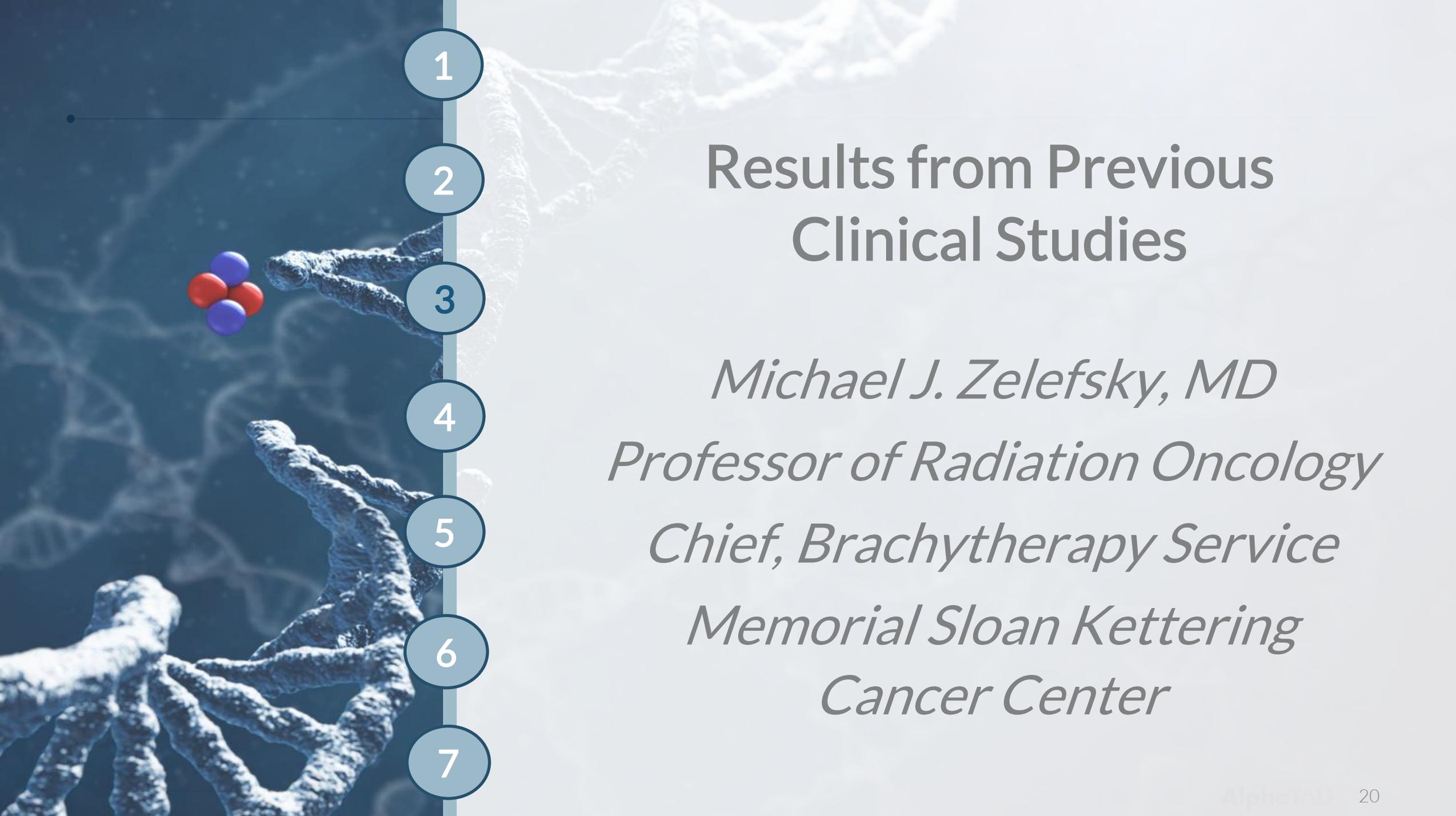
30-Nov-17



After

29-Dec-17





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Results from Previous Clinical Studies

Michael J. Zelefsky, MD

Professor of Radiation Oncology

Chief, Brachytherapy Service

Memorial Sloan Kettering

Cancer Center

Results from
Previous Clinical
Studies

CTP-SCC-00:
First Clinical Trial

Outline of Our First Clinical Study: Skin / Head & Neck SCC

Trial Sites: Israel, Italy

 **Objectives:** Evaluate feasibility, safety & effectiveness

Key Eligibility Criteria



SCC histopathologically confirmed

Lesions ≤ 5 cm*

Age ≥ 18

ECOG performance scale ≤ 2

Patients W/O immunosuppression

De-novo or **recurrent** disease

*in the longest diameter (without nodal spread).

Treatment and Procedure



Treatment plan based on CT-simulation

Sources 1cm length, 0.7mm diam.

Activity per source 2 μ Ci

Outpatient setting

Local anesthesia

Number of sources inserted: min 3, max 169

Timeline and Follow-Up



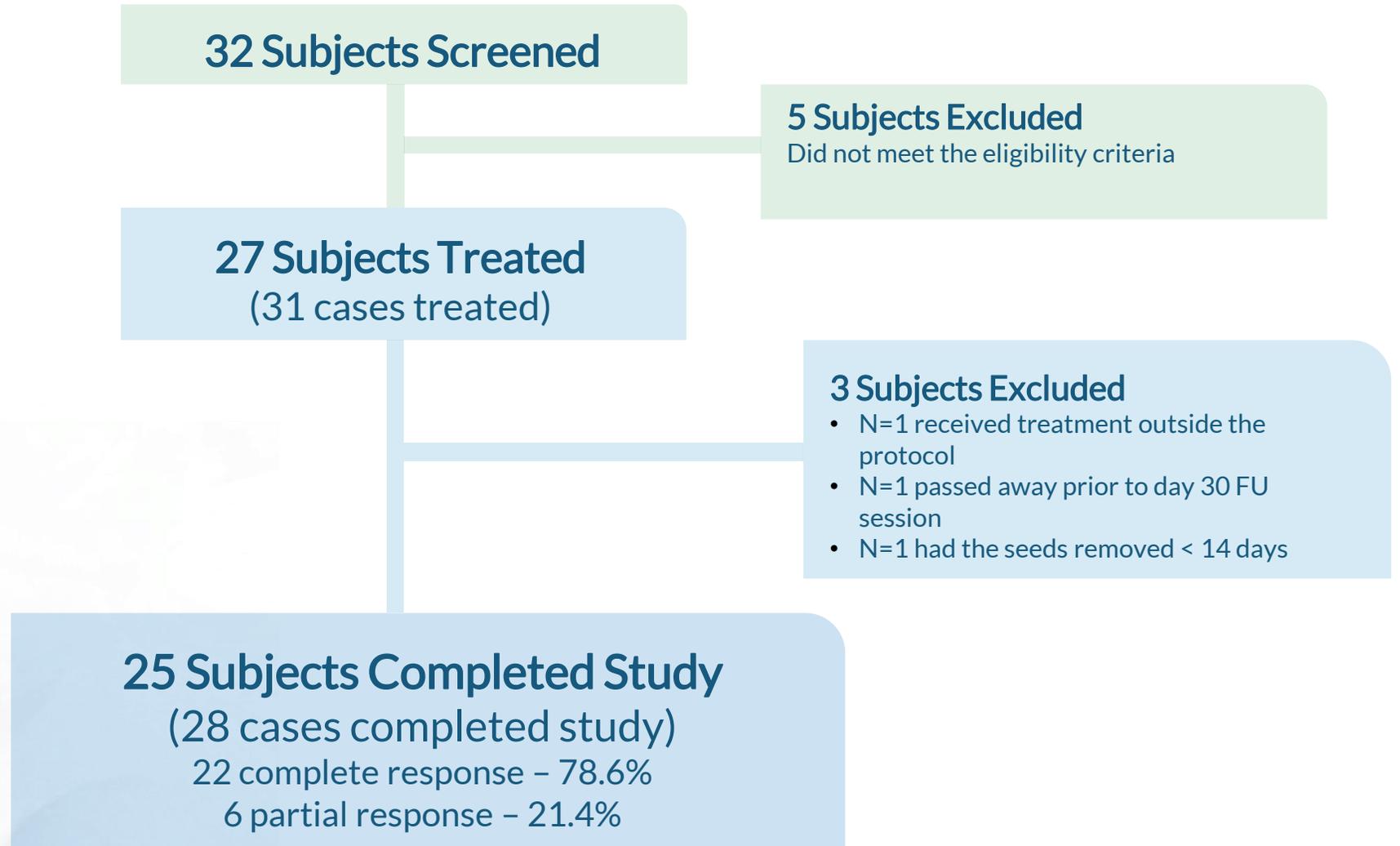
Alpha DaRT sources insertion

Removal after 15 days

Check-up on days 4, 9 and 30 after insertion

Long term follow up based on standard of care

CTP-SCC-00: Diagram of Study Subjects & Cases



Skin / Head & Neck SCC Study Results



100% overall response rate



Durable responses observed



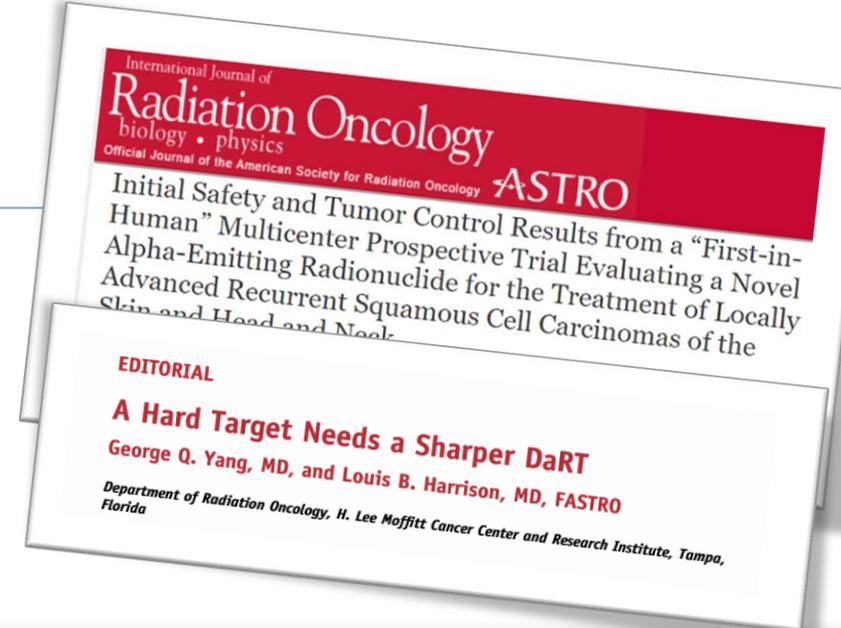
Responses observed within days



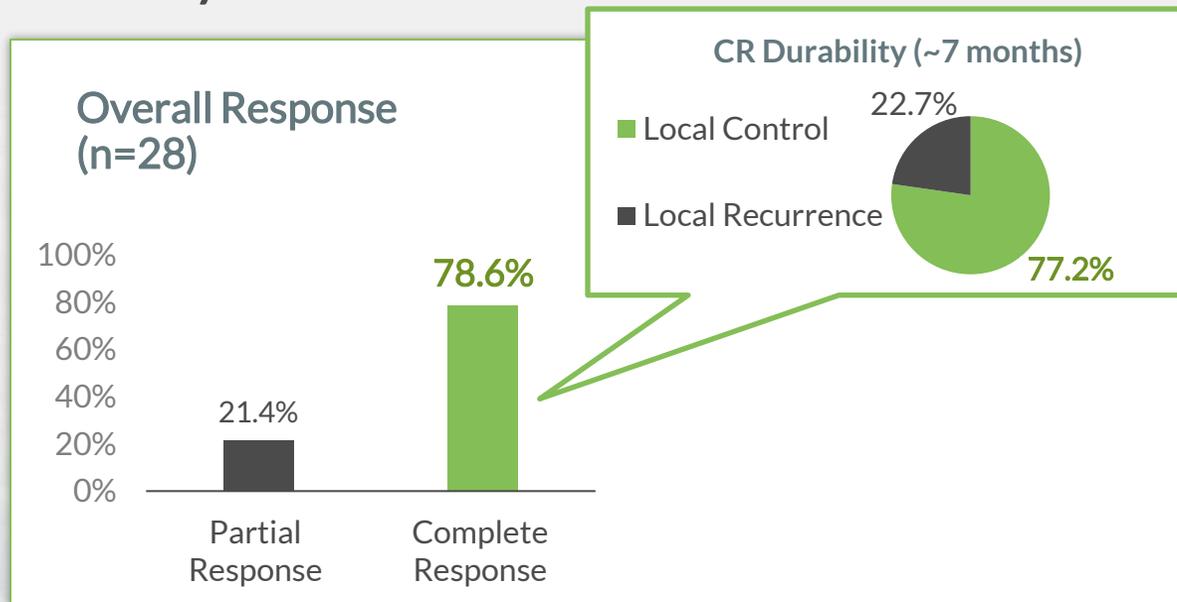
Well tolerated; no systemic toxicity observed



Similar response from radioresistant tumors

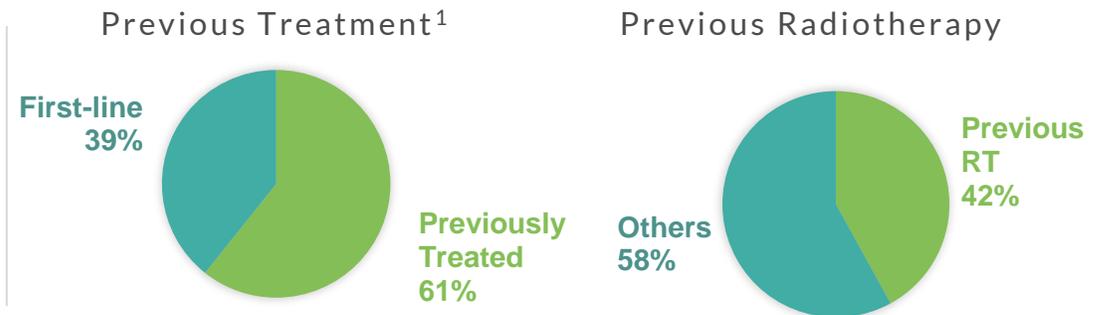


Efficacy Results



Baseline Disease Characteristics

(Patient median age = 80.5 years)



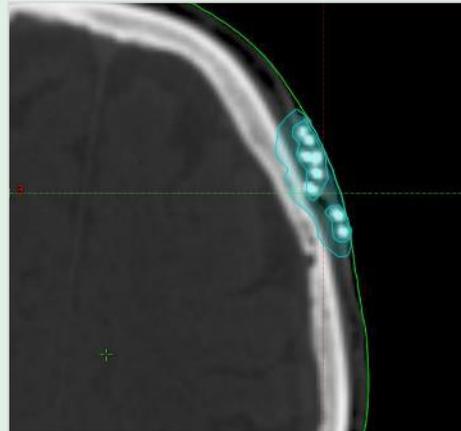
¹Most patients (60.7%) had recurrent and previously treated disease by either surgery, prior external beam radiotherapy or both; 13 of 31 (42%) had received prior RT.

AP-022 Complete Response

Age	68	Applicators used	12
Previous treatments	None	Alpha DaRT sources inserted	24
Tumor initial volume [cm ³]	2.8	Total activity [μCi]	48



Before
27-Aug-2018



During
30-Aug-2018



During
30-Aug-2018



After
30-Sep-2018

Alpha DaRT Treatment was Well Tolerated

No systemic toxicities and minimal (\leq grade 2) local toxicities observed to date



Targeted treatment

Designed to spare neighboring healthy tissue



No systemic toxicity observed

Negligible and short-term radioactivity in the patient's body



Minimal local toxicity observed

Minimal local toxicity with grade ≤ 2 resolved within a month



Safe procedure for caregivers

No special shielding required



No suppression of immune system observed

Critical in these times of pandemic

Acute Local Toxicity	Incidence (%)		
	Severity Grade		
	1	2	3
Administration site erythema	11 (41%)	9 (33%)	0 (0%)
Administration site edema	9 (33%)	10 (37%)	0 (0%)
Administration site pain	8 (30%)	11 (41%)	0 (0%)
Administration site exudate	2 (7%)	8 (30%)	0 (0%)
Administration site ulcer	4 (15%)	5 (19%)	0 (0%)
Administration site numbness	1 (4%)	0 (0%)	0 (0%)
Administration site pruritus	3 (11%)	0 (0%)	0 (0%)
Administration site bleeding	1 (4%)	0 (0%)	0 (0%)
Aural myiasis (administration site)	1 (4%)	0 (0%)	0 (0%)
Decreased appetite	1 (4%)	0 (0%)	0 (0%)

Results from
Previous Clinical
Studies

U.S. Pilot Skin / Superficial Soft Tissue Study
CTP-SCC-MSK-00

Outline of Our U.S. Pilot Skin / Superficial Soft Tissue Study

- 🎯 **Primary objective:** Demonstrate feasibility & safety malignant skin and superficial soft tissue tumors
- 🎯 **Secondary objective:** Evaluate tumor response, radiation exposure safety, stability of sources, QoL metrics

Key Eligibility Criteria

Malignant skin or superficial soft tissue

Lesions ≥ 1 to 5 cm, depth ≥ 4 mm

Suitable for **percutaneous interstitial radiotherapy**

Sample size N = 10 patients

Treatment and Procedure

Treatment plan based on CT-simulation

Sources 1cm length, 0.7mm diam.

Activity per source 2 μ Ci

Local anesthesia

Timeline and Follow-Up

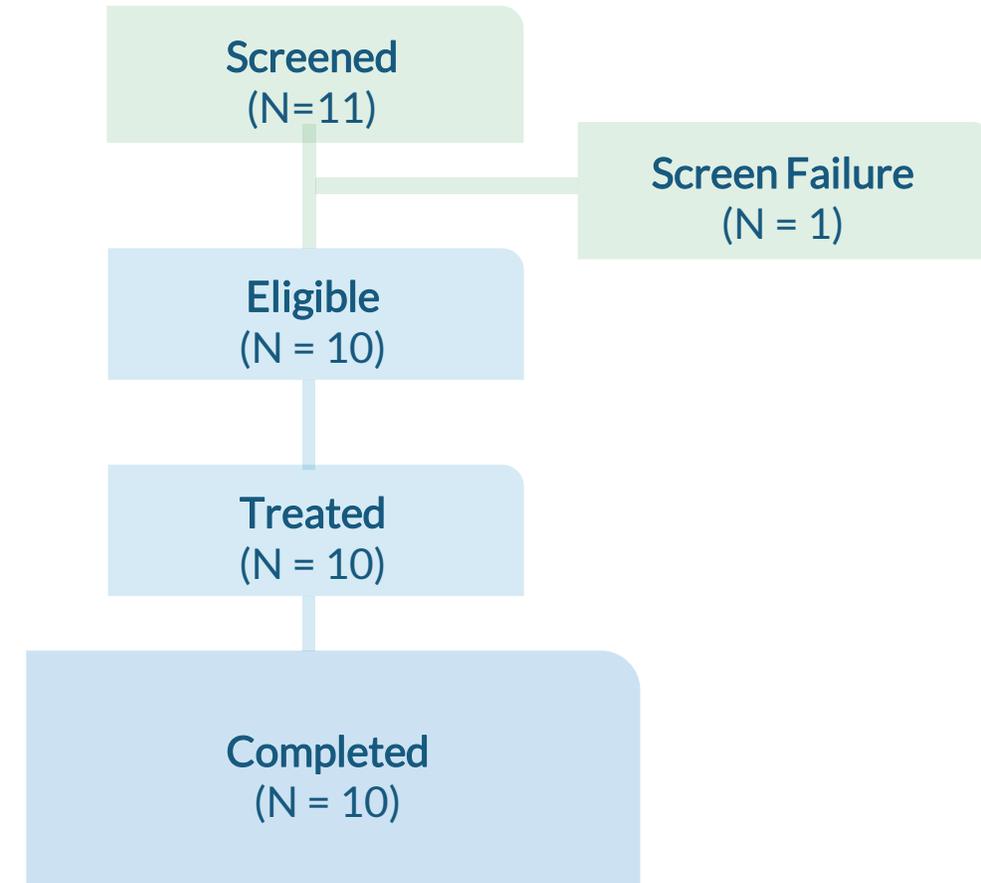
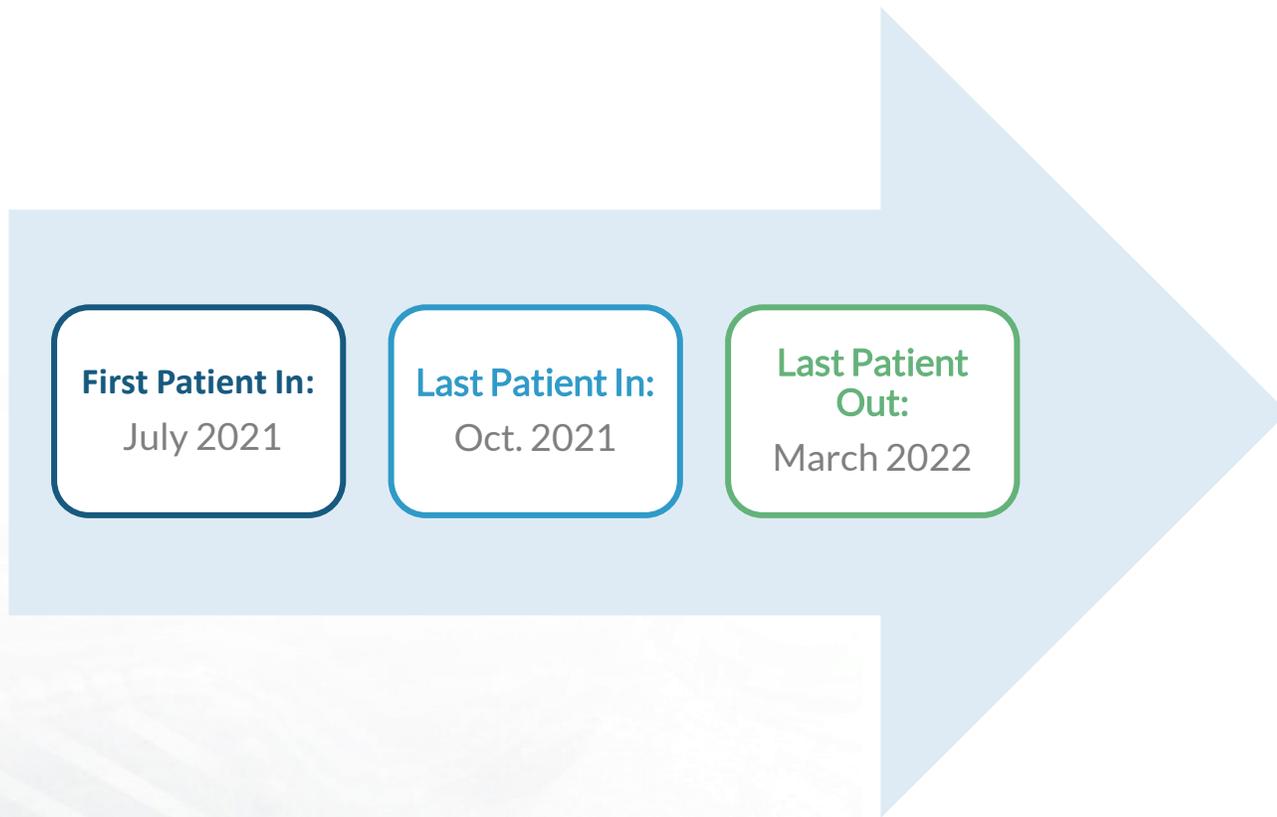
Alpha DaRT sources insertion

Removal after 14 to 21 days

Check-up on days 4, 12, 30, 38 after insertion

Long term follow up based on standard of care

Trial Overview & Subject Disposition



Patient / Tumor Characteristics

Demographics & Disease Characteristics

Treated Subjects (N = 10)

Age, years	
mean ± std	73.9 ± 10.8
Median (min, max)	71.5 (57, 92)
Sex, n (%)	
Female	4 (40%)
Male	6 (60%)
Race, n (%)	
White	10 (100%)
Histopathology, n (%)	
Basal Cell Carcinoma (BCC)	6 (60%)
Squamous Cell Carcinoma (SCC)	4 (40%)
Stage, n (%)	
Stage I	7 (70%)
Stage II	3 (30%)

Tumor Characteristics

Treated Subjects (N = 10)

Tumor Volume, cm³	
Median (min, max)	2.1 (0.7, 12.7)
Time from first diagnosis, months	
Median (min, max)	18.7 (0.3, 284.5)
Longest Diameter (LD), cm	
Median (min, max)	2.4 (1.6, 4.3)
ECOG Score, n (%)	
0	8 (80%)
1	0
2	2 (20%)
Tumor Volume, cm³	
Median (min, max)	2.1 (0.7, 12.7)

Safety Results

- Twenty-two (22) **total adverse events (AEs)** were reported in 7 subjects
- Most AEs were of **mild or moderate severity**
- Two (2) serious AEs (SAEs) in a single subject – **both not related to study device or procedure**

Number of Subjects with Procedure- or Device-Related* Adverse Events by Severity Grade

Adverse Event	Severity Grade		
	1	2	3
Dermatitis radiation	2	1	0
Localized edema	1	0	0
Joint range of motion decreased	0	1	0
Pain	0	1	0
Pruritis	2	0	0
Wound infection	0	1	0

Note: Adverse events are presented according to CTCAE V5 coded terms.

* Probably or possibly related

Efficacy Results



All 10 subjects achieved a **complete response (CR)** at the 12-week follow-up visit



There were **no reported relapses** of disease by the final study visit at 24 weeks

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User Experience & Selected Case Studies

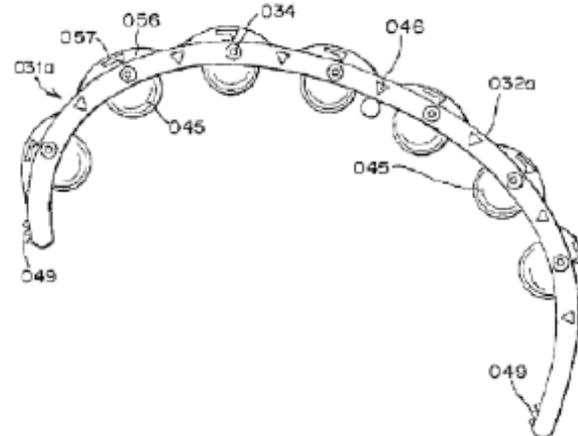
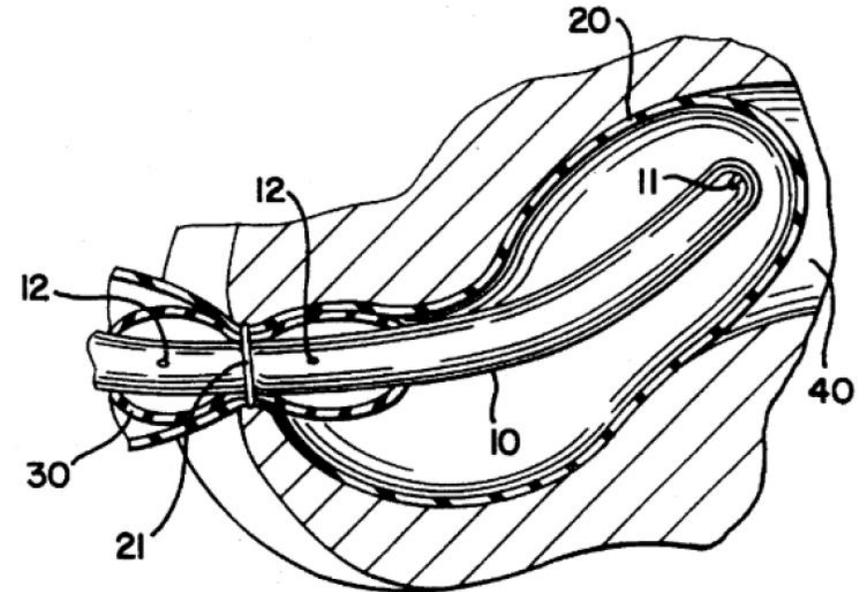
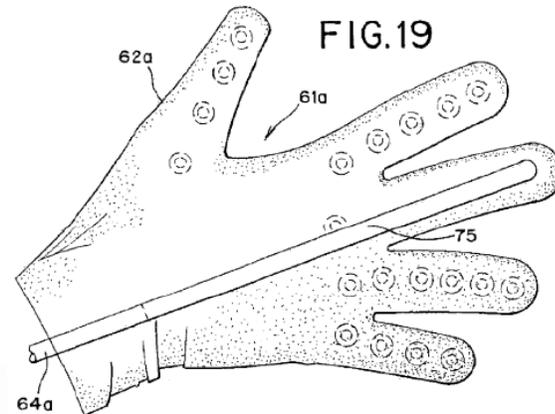
*Dr. Mark D'Andrea,
University Cancer Centers*



My Research Interests (1/2)

• **Dozens of patents** for radiotherapy / brachytherapy devices and methods of treatment for use in:

- Head & Neck
- Bladder
- Rectum
- Sarcoma
- Gynecologic
- Others



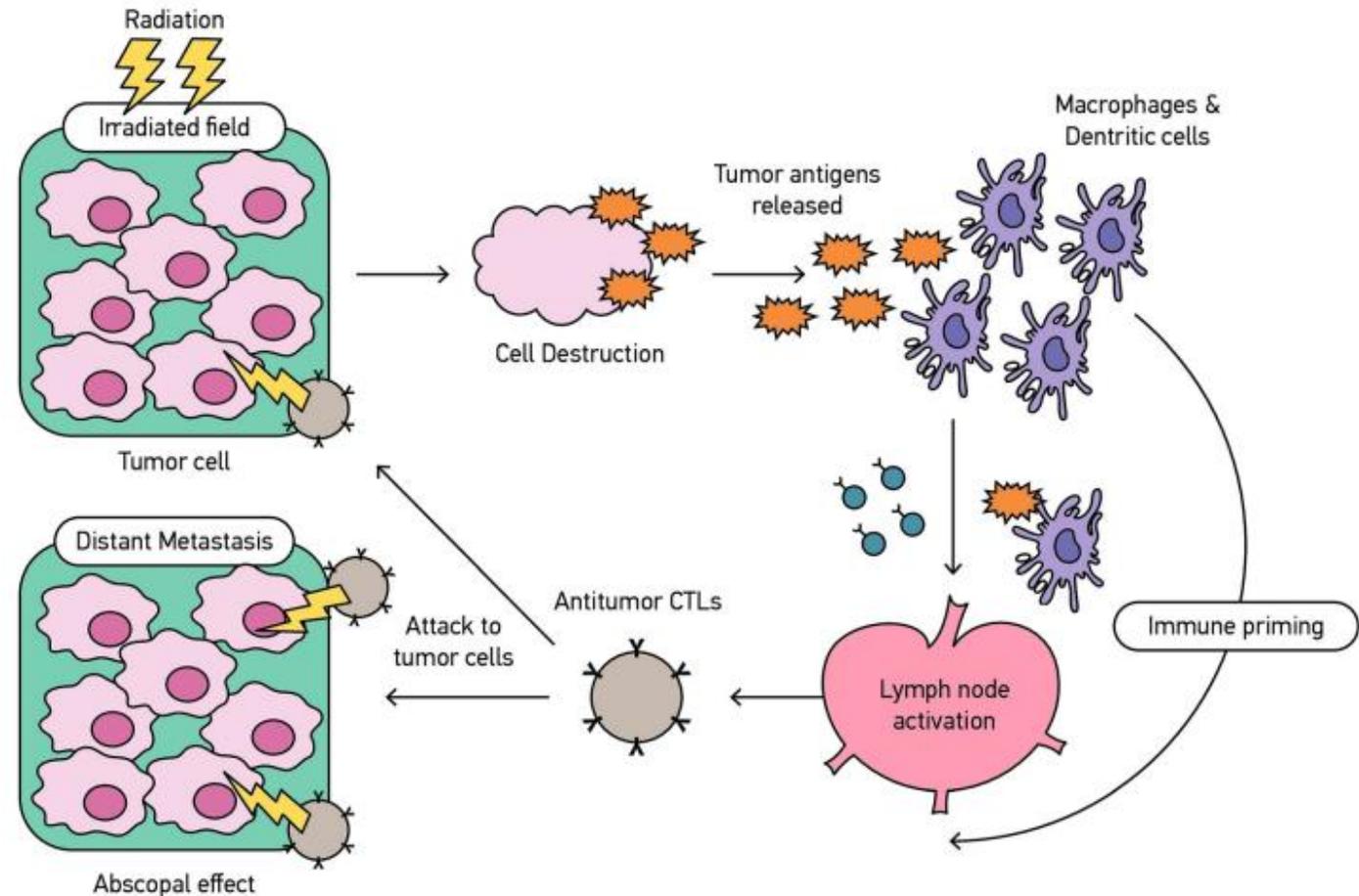
My Research Interests (2/2)

- Authored a chapter on the **abscopal effect** in radiotherapy

The systemic immunostimulatory effects of radiation therapy producing overall tumor control through the abscopal effect

Mark A. D'Andrea & G. Kesava Reddy

Journal of Radiation Oncology
ISSN 1948-7894
J Radiat Oncol
DOI 10.1007/s13566-019-00391-1



Case 1 - 77 Y/O with Recurrent BCC on the Nose



Prior treatments: Surgery (2005)

Tumor Size:

Longest diameter 1.59 cm

Depth 0.5 cm

Volume 0.65 ml

Alpha DaRT Treatment:

Applicators used 15

Alpha DaRT sources inserted 20

Total activity [μCi] 40

Case 1 - 77 Y/O with Recurrent BCC on the Nose

Results



Simulation Day



Insertion Day



Removal Day
15 days



Complete Response
12 weeks

Case 2 - 92 Y/O with BCC on the Chin

Prior treatments: None

Tumor Size:

Longest diameter 2.38 cm

Depth 0.84 cm

Volume 2.3 ml

Alpha DaRT Treatment:

Applicators used 29

Alpha DaRT sources inserted 58

Total activity [μ Ci] 116

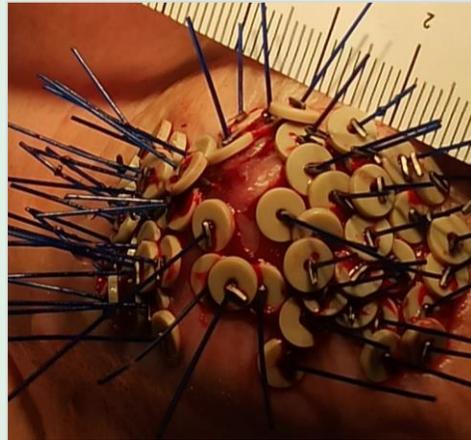


Case 2 - 92 Y/O with BCC on the Chin

Results



Simulation Day



Insertion Day



Removal Day
15 days



Complete Response
12 weeks

Case 3 - 75 Y/O with BCC on the Chin / Cheek

Prior treatments: None

Tumor Size:

Longest diameter 1.7 cm

Depth 0.9 cm

Volume 1.7 ml

Alpha DaRT Treatment:

Applicators used 23

Alpha DaRT sources inserted 35

Total activity [μ Ci] 70



Case 3 - 75 Y/O with BCC on the Chin / Cheek

Results



Simulation Day



Insertion Day



Removal Day
15 days



Complete Response
12 weeks

Case 4 - 71 Y/O with BCC on the Nose

Prior treatments: Cryotherapy (1998),
Resection (2005)

Tumor Size:

Longest diameter 2.59 cm

Depth 0.6 cm

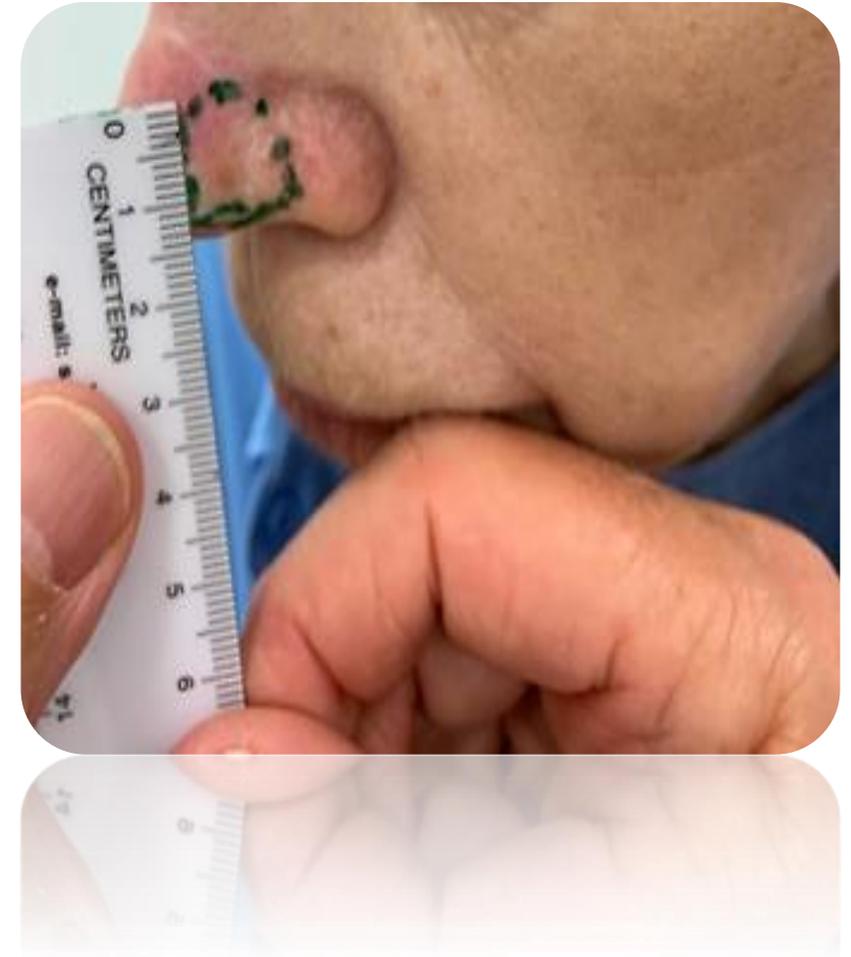
Volume 1.95 ml

Alpha DaRT Treatment:

Applicators used 15

Alpha DaRT sources inserted 23

Total activity [μ Ci] 46



Case 4 - 71 Y/O with BCC on the Nose

Results



Simulation Day



Insertion Day



Removal Day
15 days



Complete Response
12 weeks

Case 5 - 66 Y/O with Recurrent SCC on the Eye

Prior treatments: Cryotherapy (2000)

Tumor Size:

Longest diameter 1.63 cm

Depth 0.6 cm

Volume 0.98 ml

Alpha DaRT Treatment:

Applicators used 16

Alpha DaRT sources inserted 25

Total activity [μ Ci] 50



Case 5 - 66 Y/O with Recurrent SCC on the Eye

Treatment



Before



Insertion Day



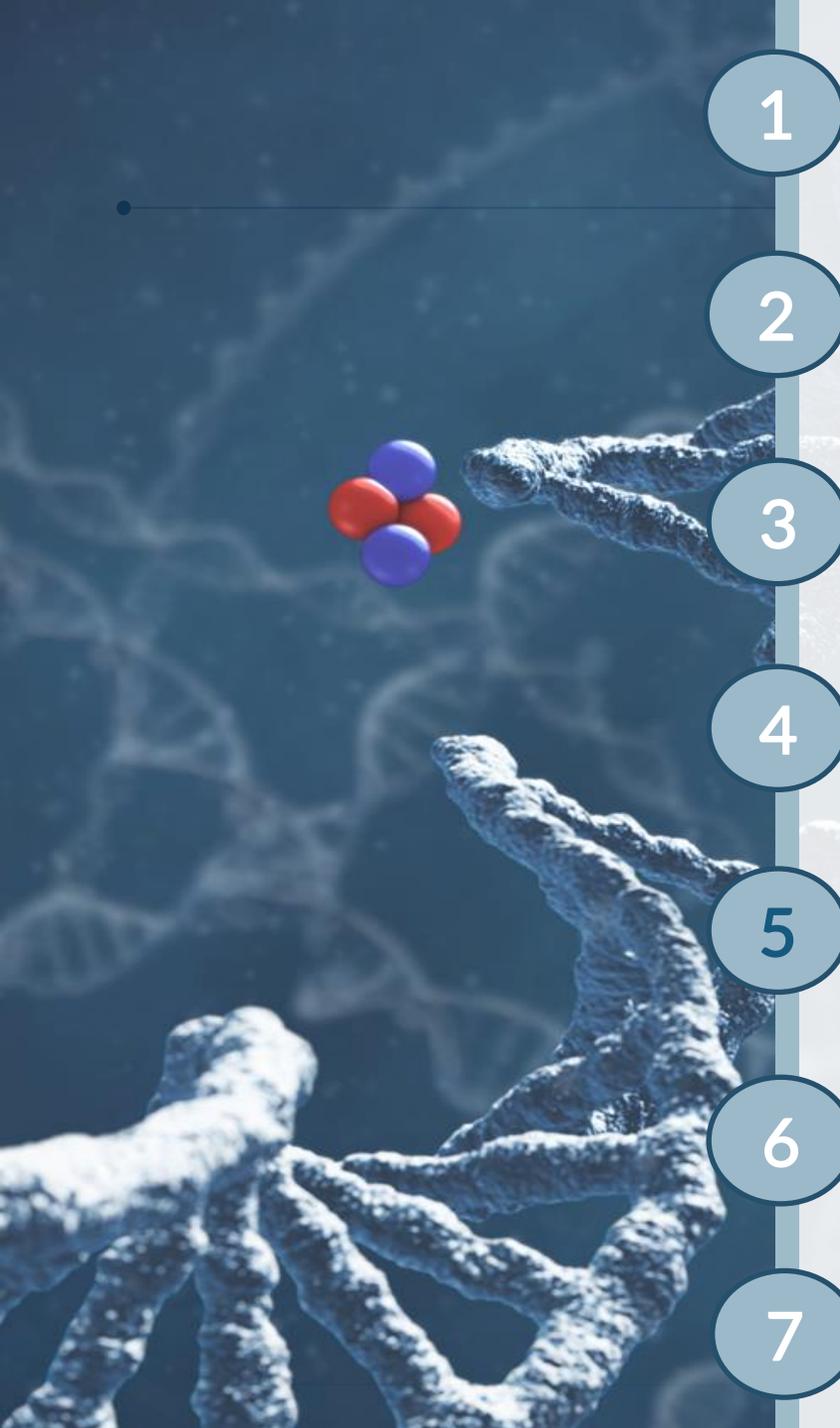
Insertion Day



Removal

Case 5 – Patient Testimonial

<https://www.youtube.com/watch?v=t2MXPoS5ql4>



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Upcoming Clinical Studies, Including US Pivotal Study

Dr. Robert Den, CMO

How Are We Moving Forward?

Our guiding principle is to have our scientific discoveries inform our clinical trials and then have our clinical trials inform our scientific research

Toward that end:

Laboratory	Clinic
Design experiments that mirror the clinic	Focused on neo-adjuvant setting
Mechanism	Built in biomarkers for immune response
Interrogate novel combinations using FDA approved agents	Interrogate novel combinations using FDA approved agents

Clinical Trial Portfolio

Superficial to Internal Organs

Temporary Implants

Japan Pivotal Trial

US Multi-center Pivotal Trial

Expanding to Other
Histologic Types: **Israel**

Pathologic Response

Neoadjuvant Prostate: Israel

Liver: Two-stage
Hepatectomy: Canada

Breast Cancer: Israel

Unmet Need / Inoperable

Pancreas: Israel/Canada

Liver: Israel/Canada

Breast: Israel/Europe

Intraluminal GI tumors: Israel

Recurrent GBM: Israel/US

Study Specifics

**A Prospective International Multicenter,
Pivotal, Single Arm, Open Label
Clinical Study to Assess the Efficacy and
Safety of Intratumoral Alpha DaRT224
for the Treatment of Patients with Recurrent
Cutaneous Squamous Cell Carcinoma**
Protocol Number: CTP-SCC-03

Outline of Our Multicenter Pivotal Recurrent SCC study

- **Primary / safety objectives:**
 - ORR based on Best Overall Response
 - DOR 6 months after initial response
 - Assess the safety based on statistics of device-related AEs (per CTCAE v5)
- **Secondary objectives:** Evaluate O-DOR, local control, PFS and OS (all up to 12 months), and QoL Metrics

Key Eligibility Criteria



Recurrent non-metastatic cutaneous **SCC**

Patient with **no curative standard-of-care options**

No **previously untreated SCC**

Sample size N = 86 patients

Treatment and Procedure



Treatment plan based on CT-simulation

Sources 1cm length, 0.7mm diam.

Activity per source 3 μCi

Local anesthesia

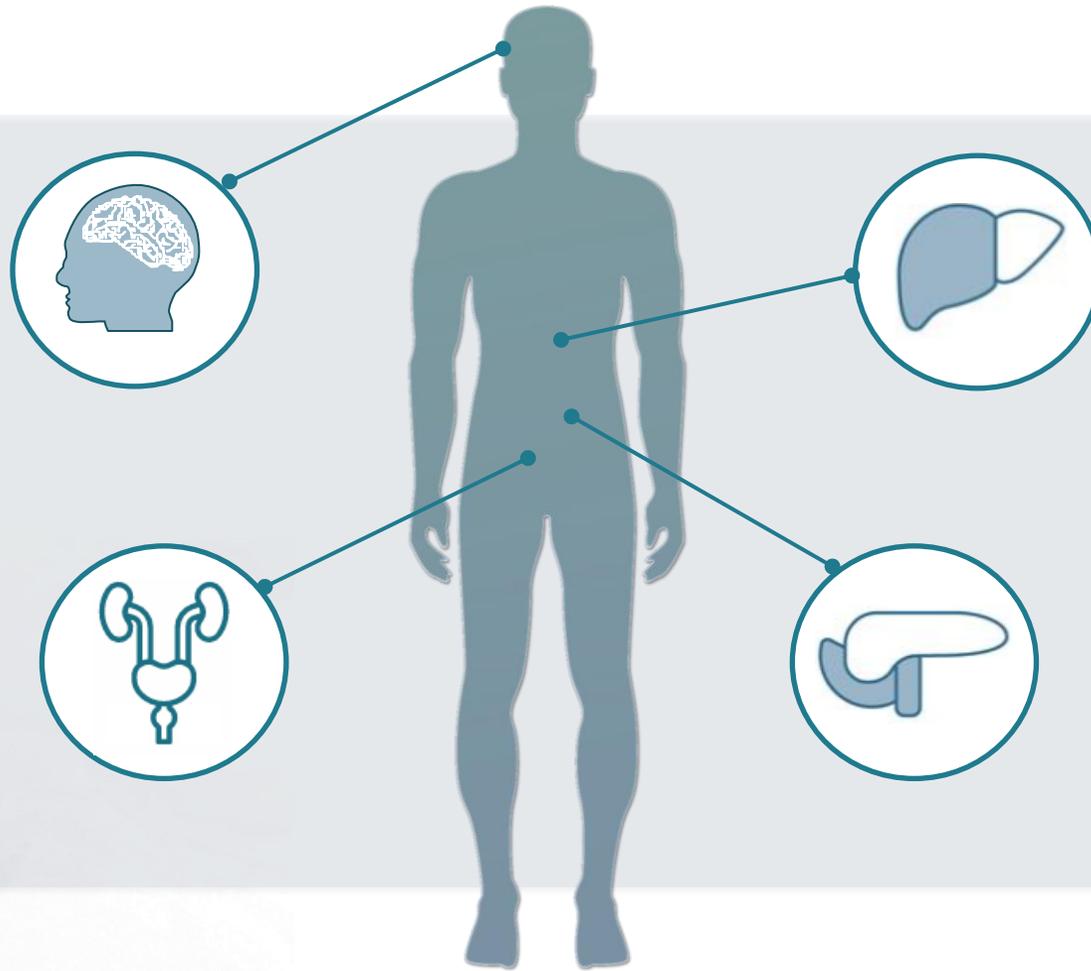
Timeline and Follow-Up



Alpha DaRT sources insertion

Removal after 14 to 21 days

Weekly **follow-up** during the treatment period



Other Indications

Other Indications

**Alpha DaRT
In Men With
Prostate Cancer**
CTP-PRST-02

Outline of our Neoadjuvant Prostate Study

- 🎯 **Primary objectives:** Feasibility & safety of intra-tumoral insertion of Alpha DaRT sources into prostate
- 🎯 **Secondary objectives:** Evaluate pathological & radiological ORR and change in QoL metrics

Key Eligibility Criteria



Resectable prostate adenocarcinoma

Non-metastatic tumor

Lesions ≤ 3 cm*

Targetable lesion

Previously **untreated**:

No prior **TURP** or **prostate surgery**

No prior **pelvic radiation**

Sample size N = 10 patients

Treatment & Procedure



Treatment plan based on PSMA

PET-CT or multiparametric MRI

Sources 0.7 mm in diameter and 1 cm in length

Activity per source 5 μ Ci

General anesthesia

Timeline and Follow-Up



Alpha DaRT sources insertion

Prostate surgery after 50 days

Check-up on days 7, 15, 22 after insertion

Follow-up duration up to 75 days

Plant Applicator Overview

<https://www.youtube.com/watch?v=hSKbj16moFQ>



Other Indications

**A Feasibility and Safety Study of Intratumoral
Diffusing Alpha Radiation Emitters on
Advanced Pancreatic Cancer**
AT-PANC-101

Outline of the Pancreas Study

- 🎯 **Primary objective:** Evaluate feasibility & safety of Alpha DaRT sources inserted into pancreas in terms of incidence of device related AEs & SAEs.
- 🎯 **Secondary objective:** Evaluate efficacy (radiological ORR and change in tumor markers), OS, stent durability, and QoL

Key Eligibility Criteria

Locally **advanced (Stage II or III)** or **metastatic (Stage IV)** pancreatic adenocarcinoma

Inoperable pancreatic cancer because:

- **Unresectable**
- **Metastatic** disease
- Medically **unfit** for surgery

No **concomitant chemotherapy** or **immunotherapy**

Sample size N = 30 patients

Treatment and Procedure

Treatment plan based on CT

Sources 0.7 mm in diameter and 10 mm in length

Activity per source 3 μ Ci

Source insertion using **endoscopic ultrasonography**

General anesthesia

Timeline and Follow-Up

Alpha DaRT sources insertion

Check-up on days 6, 15, 21, 35, 60 after insertion

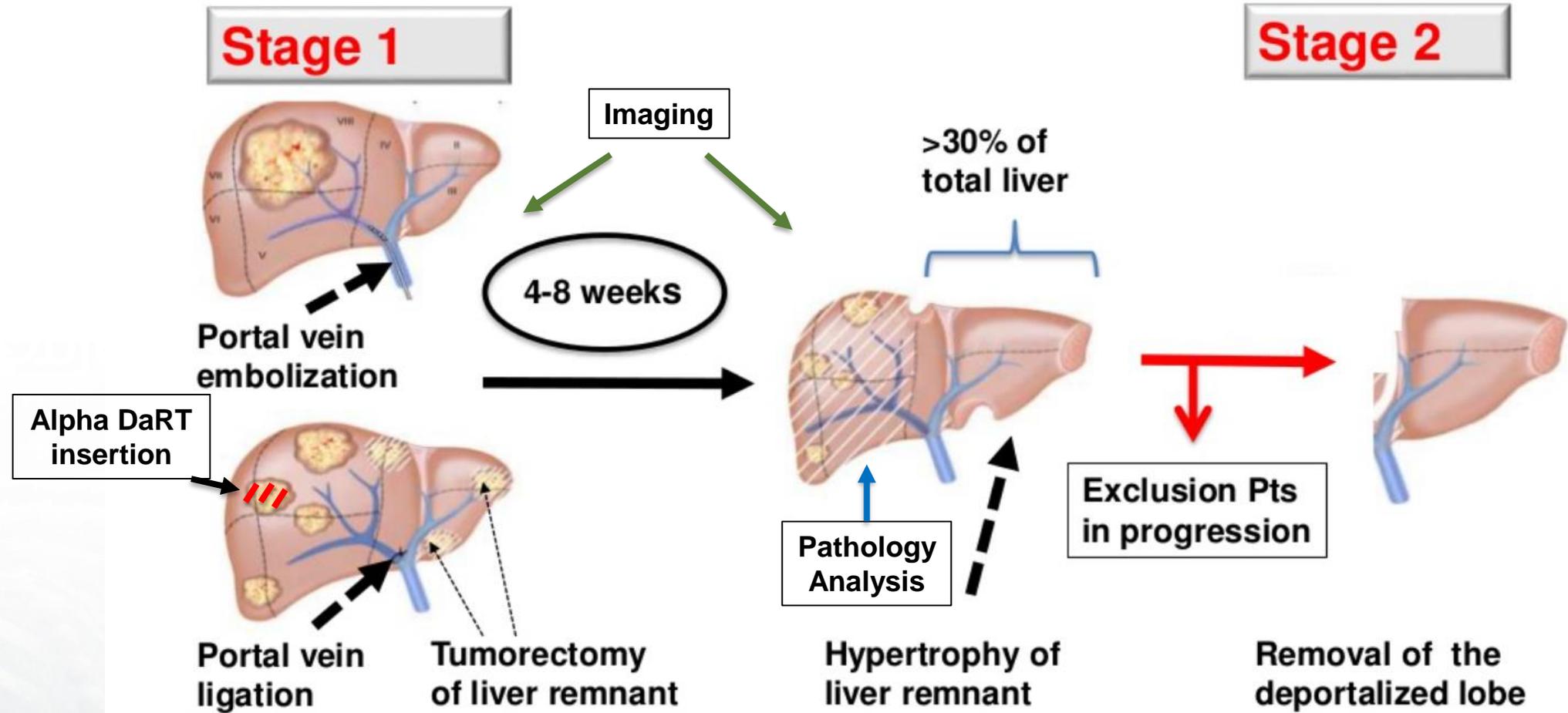
Follow-up duration up to 2 years

Other Indications

**A Feasibility and Safety
study of Intratumoral Diffusing
Alpha Radiation Emitters for the
Treatment of Liver Metastases
CTP-LIV-00**

Study Schema

Liver study



Clavien et al. Strategies for safer liver surgery. NEJM, 2017

Outline of Draft Liver Metastases Study

- Primary objectives: Evaluate feasibility & safety of Alpha DaRT implanted in liver metastases
- Secondary / exploratory objectives: Evaluate pathological and radiological response, determine immunological impact, stratify differences in response by histopath. growth patterns (vascular / immuno.)

Key Eligibility Criteria



Referred for a **two-staged hepatectomy** to resect liver metastases of colorectal cancer

No prior use of **systemic investigational agents** for primary cancer

Sample size N = 10 patients

Treatment and Procedure



Treatment plan based on CT scan or MRI

Sources 0.7 mm in diameter and 1 cm in length

Activity per source 3 μ Ci

General anesthesia

Timeline



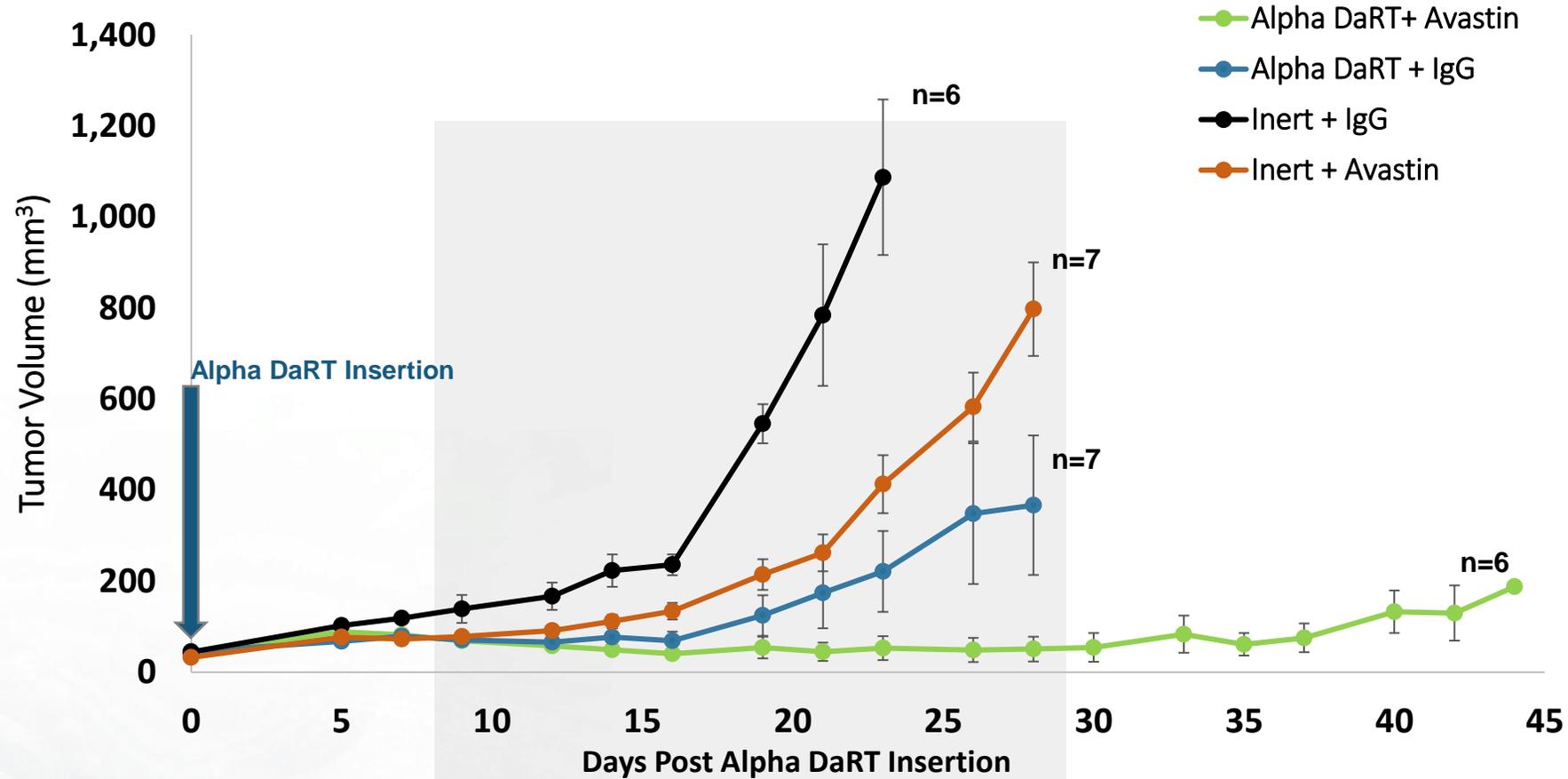
- 1st operation:** one side of the liver is cleared from its metastases & Alpha DaRT sources are implanted in the other side of the liver
- 3 - 4 cycles of **chemotherapy** (6 - 8 weeks)
- 2nd operation:** The liver lobe containing the metastasis with the sources is resected, to leave the patient with a disease-free liver

Other Indications



Early Look at Efforts in Glioblastoma Multiforme

Alpha DaRT + Avastin Combo Showed Attenuated Growth of GBM Xenografts

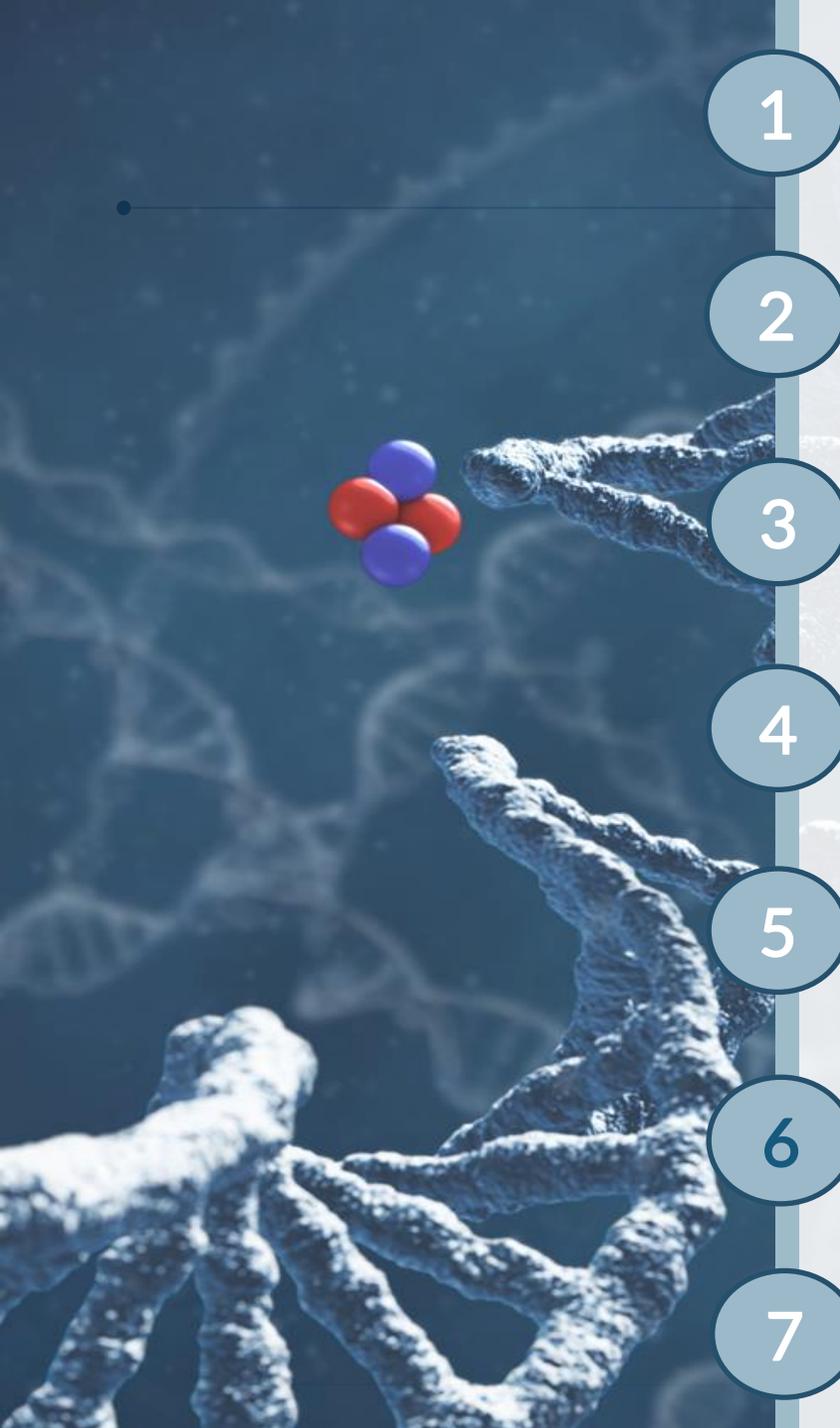


Avastin / IgG control injections 3 times per week for 3 weeks

Radial Applicator Overview

<https://www.youtube.com/watch?v=IJY965J0xMk>





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Future Outlook

Raphi Levy, CFO

Development Pipeline

- Our clinical trial strategy involves progressing our lead program (superficial tumors), particularly in the US, and conducting feasibility studies in other tumors to evaluate the Alpha DaRT in tumors of high unmet need or metastatic disease
- FDA Breakthrough Device Designation received for certain uses in skin cancer and GBM

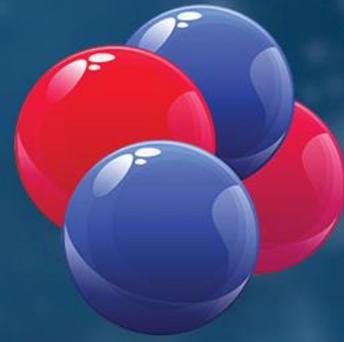
Geography	Indication	Pre-Clinical Research	Feasibility Trial	Pivotal Trial	Marketing Authorization	Anticipated Milestones
North America	Skin Cancers	U.S.				• First patient into US pivotal trial targeted for 2H 2022
	Pancreatic Cancer	Canada				• First patient in feasibility trial 2H 2022
	Liver Cancer	Canada				• Trial in planning
Israel	Skin & Oral SCC					
	All Skin & Oral Cancers					• Trial completion and submission
	la/mHNSCC (combo with pembrolizumab)					• Feasibility combination trial with Keytruda initiated 4Q 2021; awaiting interim results
	Pancreatic Cancer					• Initiate feasibility trial 4Q 2022
	Breast Cancer					• Trial in planning
	Prostate Cancer					• Trial initiated 2Q 2022 – data ~2Q 2023
Europe	Skin Cancers					• Trials underway
	Pancreatic Cancer					• Trial in planning
Japan	Head & Neck SCC					• Potential PMDA submission in 3Q 2022
	Breast Cancer					• Trial underway
Additional Tumors	Hepatic Cell Carcinoma , GBM, lung					• Development / pre-clinical trials underway

Anticipated Milestones

Geography	Indication	2H 2022	1H 2023	2H 2023
North America	Recurrent Cutaneous SCC (United States)	First patient treated in multi-center pivotal trial		Completion of multi-center pivotal trial recruitment
	Pancreatic Cancer (Canada) <i>(Tentative)</i>	First patient in feasibility trial	Interim read-out of feasibility trial	
Israel	Prostate Cancer		Read-out of prostate cancer trial data	
	Pancreatic Cancer	Initiate feasibility trial		
Japan	Head & Neck SCC	Potential submission of pivotal trial for PMDA review	Potential PMDA approval	

Clinical / Enrollment

Regulatory



AlphaTAU

Questions?