(NASDAQ:DRTS)

Investor Presentation

April 2025

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The Alpha Tau Mission

Alpha DaRT

A novel approach using localized alpha particle radiotherapy designed to precisely destroy solid tumors while sparing surrounding healthy tissue

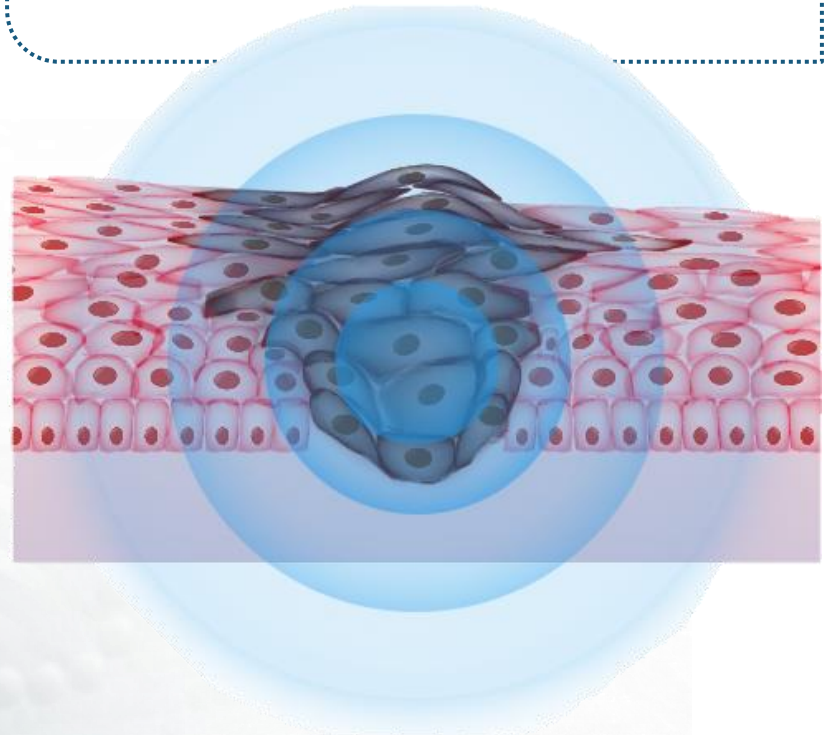
- Broad potential applicability for local tumor control, together with signs of compelling immuno-stimulatory activity
- Platform technology has the potential to be utilized alone or synergistically with other cancer treatment modalities
- Milestones and data from multiple clinical trials in various phases in different indications expected in 2025 and 2026
- 1st potential U.S. marketing authorization in 2026, with significant market opportunity across multiple tumor types

Alpha Radiation is Focal - Short Range Limits Clinical Use

Whereas beta and gamma radiation can penetrate tissue with sufficient range to facilitate tumor coverage (while risking damage to healthy tissue), alpha radiation has short range in tissue (< 100 µm), which limits its clinical usefulness in local delivery

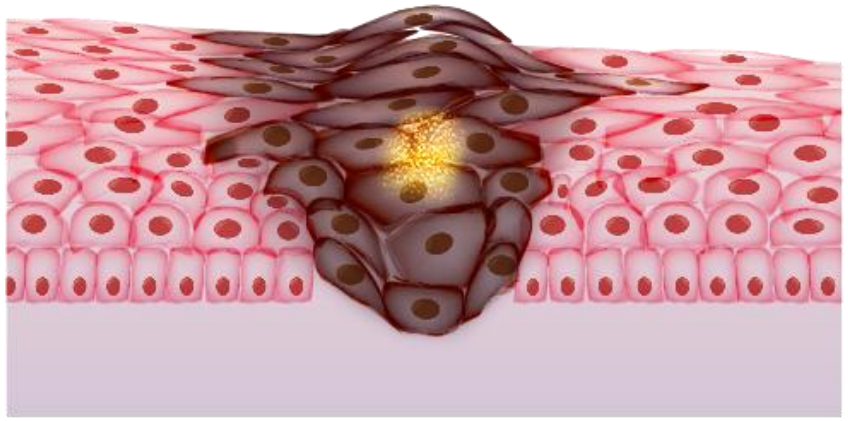
Beta/Gamma Radiation

Long therapeutic range with risk to surrounding organs



Alpha Radiation

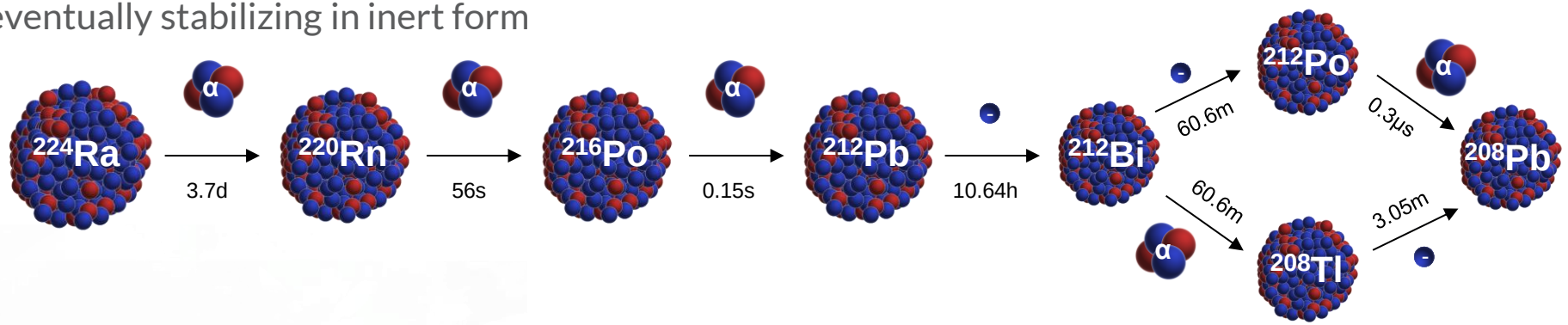
Short range in tissue limits damage to surrounding organs but also limits coverage



Alpha DaRT Technology is Designed to Overcome These Limitations

^{224}Ra 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 or titanium sources that are impregnated with Radium-224
- When the Alpha DaRT source is 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 eventually 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

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 we are targeting include SCC, H&N SCC and prostate

Metastatic

- Alpha DaRT being evaluated for its potential to induce an immune response in metastatic tumors
- Alpha DaRT being evaluated in combination with checkpoint inhibitors as an adjuvant therapy
- Tumor types we are targeting include liver, breast and H&N (which includes lip, oral cavity, salivary glands, oropharynx & pharynx) cancers

High Unmet Need

- Solid tumors that have limited treatment options with limited standard of care offering
- Alpha DaRT could potentially target broad patient populations
- Tumor types we are targeting include GBM and pancreatic cancer

Initial Foray into Superficial Tumors

Alpha DaRT first tested in superficial tumors – tumors of the skin or head & neck, due to:

- Ease of access
- Straightforward control
- Ongoing monitoring
- Strong initial preclinical data in Squamous Cell Carcinoma (SCC)

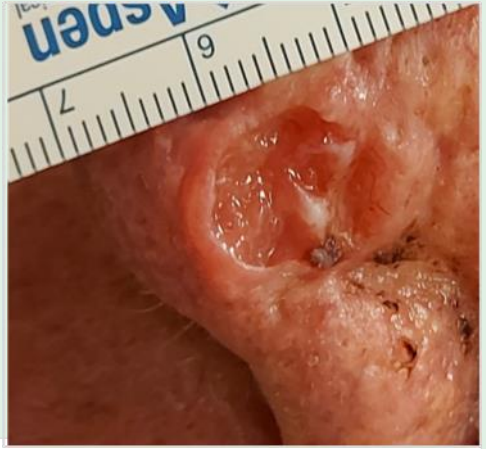
Treatment of hundreds of tumors to date:

- Indicated a mild safety profile
- Generated marketing authorization in Israel to treat SCC of the skin or oral cavity
- Allowed us to submit to PMDA in Japan for marketing authorization to treat recurrent head & neck cancer

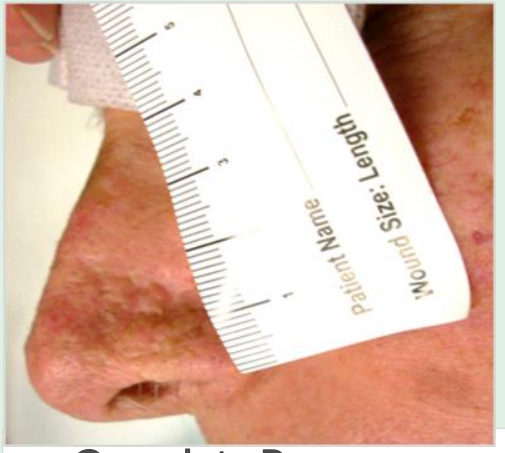
Pivotal trial ("ReSTART") underway in the U.S. for recurrent cutaneous SCC

U.S. Skin Cancer Pilot Study Leading to Pivotal Study

U.S. Pilot Feasibility Study	
Locations	5 centers – led by Memorial Sloan Kettering Cancer Center
# of Patients Treated	10
Adverse Events	**22 reported AE's, most were mild or moderate** **No treatment-related serious AEs**
Response Rate	**100% Complete Response Rate**



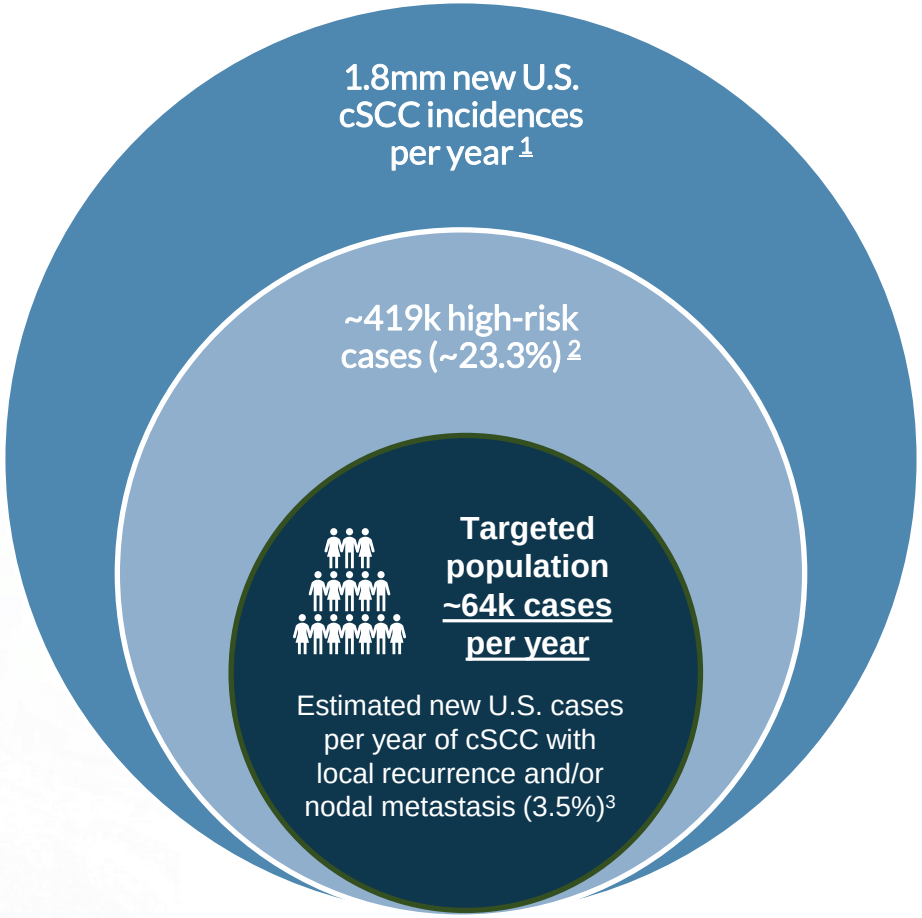
Simulation Day



Complete Response
12 weeks

Multicenter Pivotal Recurrent SCC Study	
Locations	Multiple centers, including UCLA, Emory University, Mayo Clinic, etc.
# of Patients	86
Primary Objectives	Overall Response Rate, Durability of Response @ 6 months, adverse events assessment
Targeted Completion of Recruitment	Q3 2025

Potential cSCC Patient Breakdown - Estimated U.S. Incidence



[1] https://www.skincancer.org/blog/our-new-approach-to-a-challenging-skin-cancer-statistic/

[2] Evaluation of American Joint Committee on Cancer, International Union Against Cancer, and Brigham and Women's Hospital Tumor Staging for Cutaneous Squamous Cell Carcinoma
Pritesh S. Karia, Anokhi Jambusaria-Pahlajani, David P. Harrington, George F. Murphy, Abrar A. Qureshi, and Chrysalyne D. Schmults. Journal of Clinical Oncology 2014 32:4, 327-334

[3] Factors Predictive of Recurrence and Death From Cutaneous Squamous Cell Carcinoma: A 10-Year, Single-Institution Cohort Study
Schmults CD, Karia PS, Carter JB, Han J, Qureshi AA. JAMA Dermatol. 2013;149(5):541–547. doi:10.1001/jamadermatol.2013.2139

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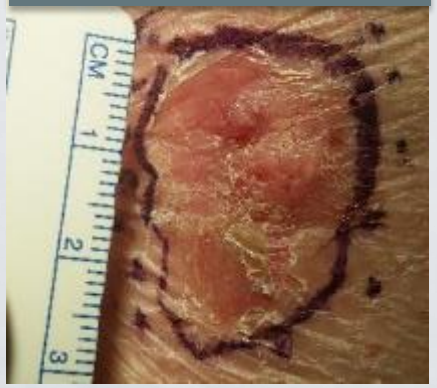
Case Study: Potential Systemic Immune Effect Observed in One cSCC Patient Where a Second, Untreated Lesion Manifested CR

Complete Response + Potential Systemic Immune Effect

Treated Tumor

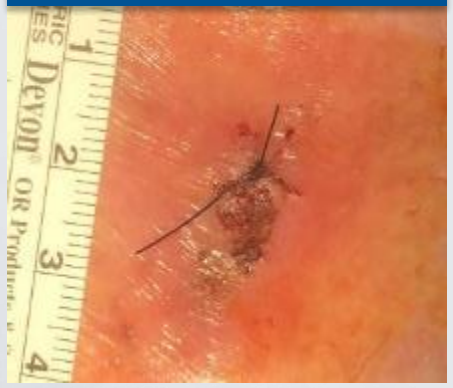
Before

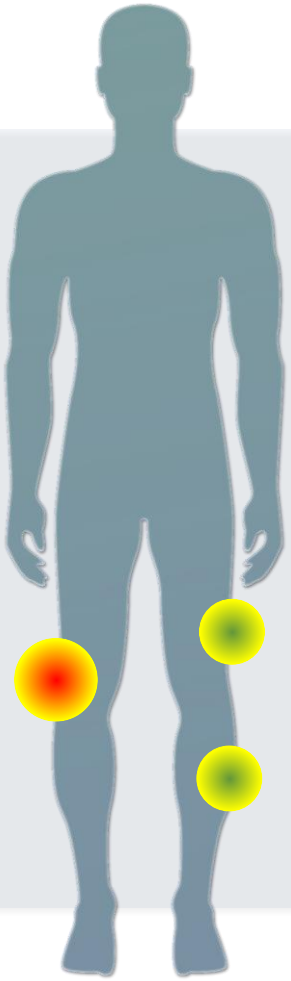
30-Nov-17



After

29-Dec-17

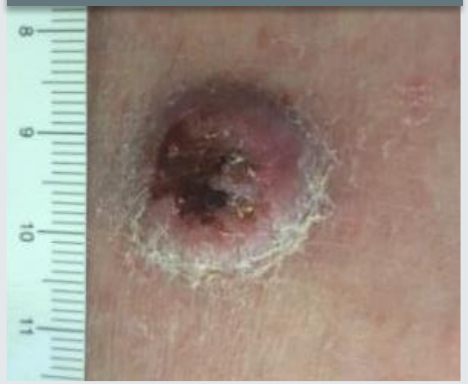




Untreated Tumors

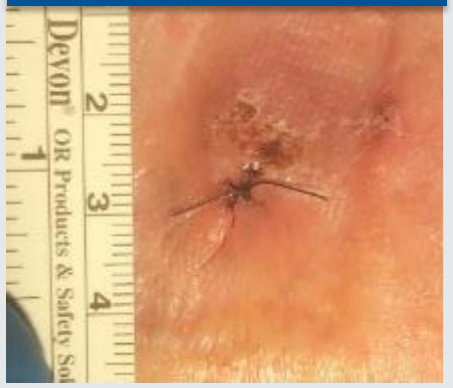
Before

30-Nov-17



After

29-Dec-17



Outline of Checkpoint Inhibitor Combination Trial – CTP-HNCPI-00

Key Eligibility Criteria

Recurrent unresectable or metastatic head and neck squamous cell carcinoma (like KEYNOTE-048)

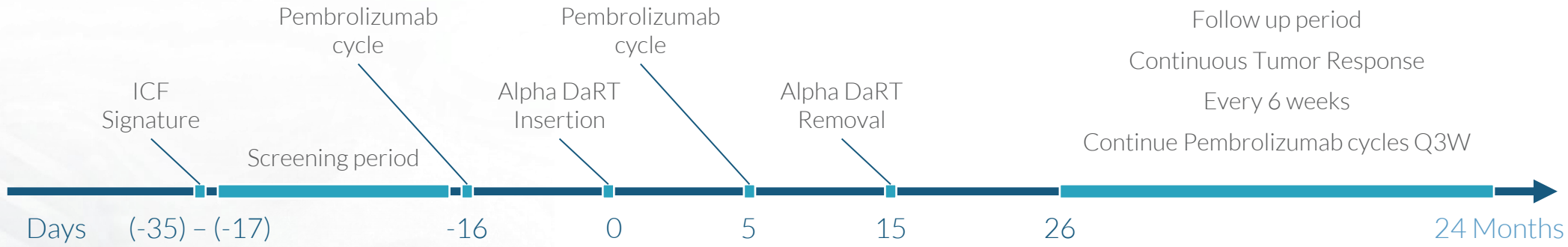
No previous treatment for metastatic disease

Benchmark Comparator

KEYNOTE-048: Benchmark comparator data for 1L Pembrolizumab in patients with recurrent or metastatic HNSCC[1]

Population	Benchmark Regimen	Systemic ORR	Systemic CR %
PD-L1 CPS ≥ 20	Pembrolizumab Alone	23%	8%
PD-L1 CPS ≥ 1	Pembrolizumab Alone	19%	5%
Total population	Pembrolizumab Alone	17%	5%

Treatment Regimen



[1]*Benchmark data provided for illustrative purposes only. Not a head-to-head trial*

Source: Burtness, B. et al (2019). Pembrolizumab alone or with chemotherapy versus cetuximab with chemotherapy for recurrent or metastatic squamous cell carcinoma of the head and neck (KEYNOTE-048): a randomised, open-label, phase 3 study. The Lancet. doi:10.1016/s0140-6736(19)32591-7

Early Interim Data Show Strong Systemic Responses

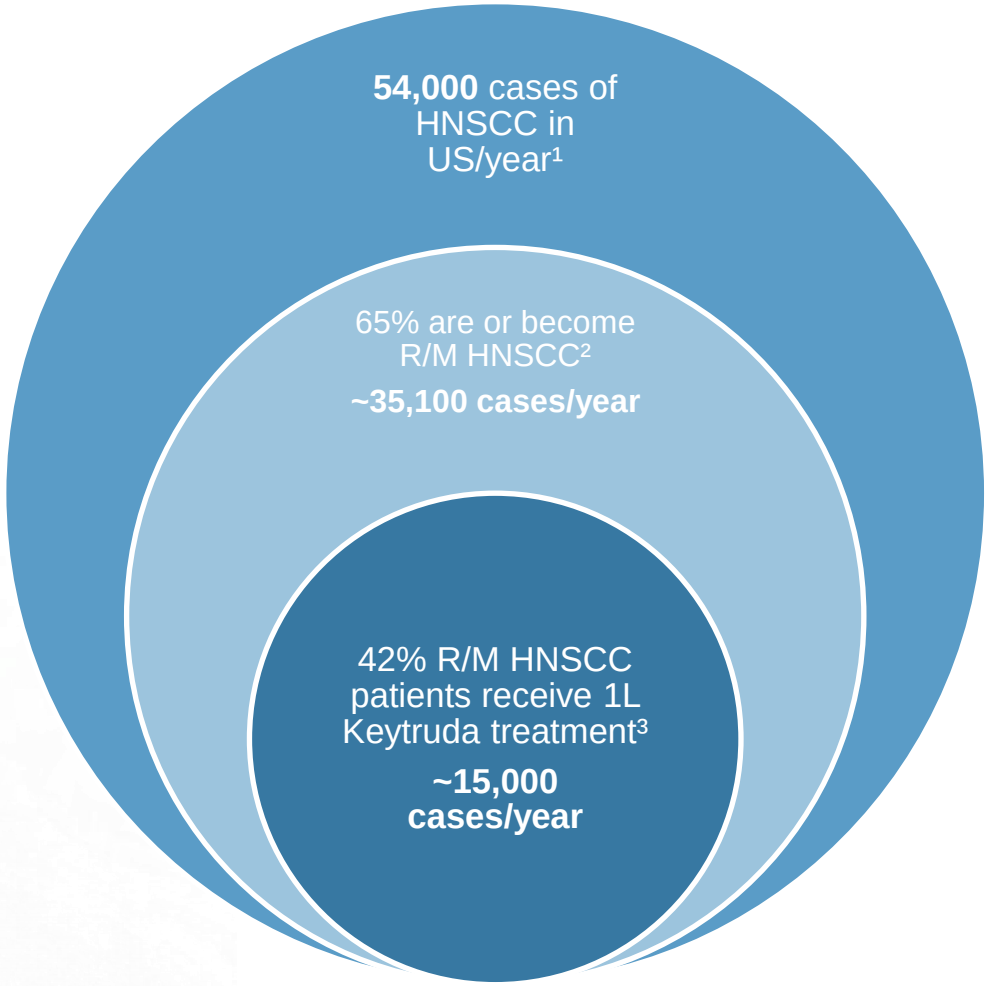
- As of January 9, 2025, eight patients were treated with Alpha DaRT and pembrolizumab in the study
- Baseline characteristics:
  - 3 female / 5 male
  - Mean age of 73 years (range 61-96)
  - 6 mHNSCC / 2 IaHNSCC
- Patients received an average of 4 cycles of pembrolizumab (range 2-9)
- Systemic responses observed:
  - Three complete responses
  - Three partial responses
  - Two patients died prior to evaluation
- Only two Alpha DaRT-related adverse events, both were Grade 1 (mild)

37.5%
Systemic Complete Responses

75%
Systemic Objective Response Rate (CR + PR)

No Related SAEs

HNSCC Patient Breakdown



[1]*Epidemiology, Risk Factors, and Prevention of Head and Neck Squamous Cell Carcinoma* Adam Barsouk, John Sukumar Aluru, Prashanth Rawla, Kalyan Saginala, Alexander Barsouk. *Med. Sci.* 2023, *11*(2), 42; https://doi.org/10.3390/medsci11020042

[2]*Recent Advances and Future Directions in Clinical Management of Head and Neck Squamous Cell Carcinoma* Jameel Muzaffar, Shahla Bari, Kedar Kirtane, Christine H. Chung *Cancers* 2021, *13*(2), 338; https://doi.org/10.3390/cancers13020338

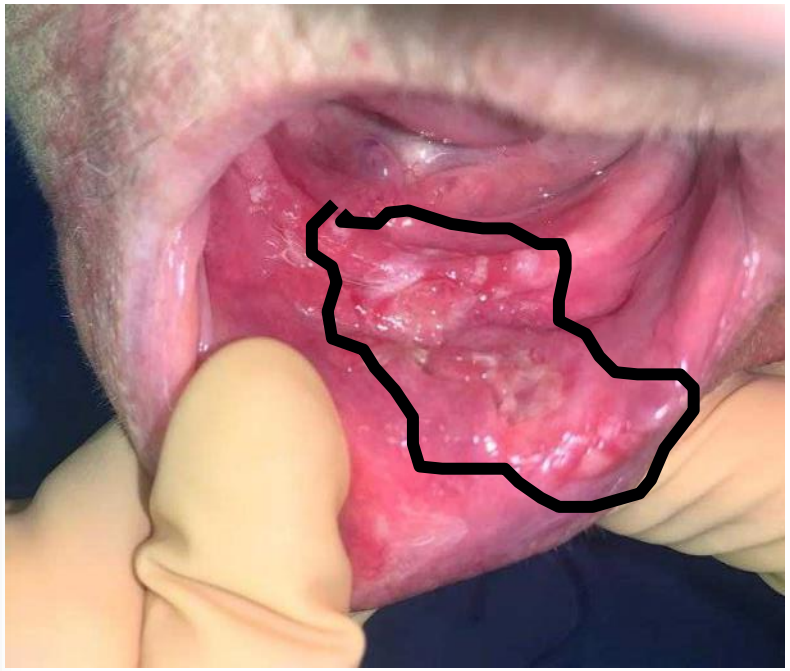
[3]*Real-world treatment patterns and outcomes among individuals receiving first-line pembrolizumab therapy for recurrent/metastatic head and neck squamous cell carcinoma* Christopher M Black, Glenn J Hanna, Liya Wang, Karthik Ramakrishnan, Daisuke Goto, Vladimir Turzhitsky, Gleicy M Hair Front Oncol. 2023 May 22;13:1160144. https://doi.org/10.3389/fonc.2023.1160144

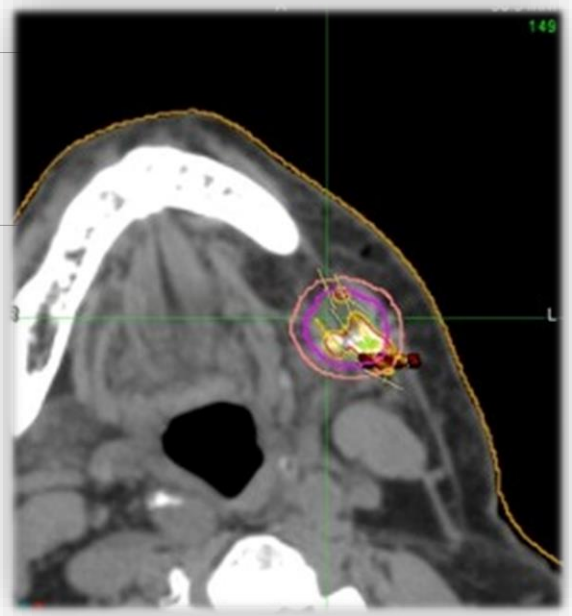
HNCPI-00-01-003

Pembrolizumab Combination Case Study

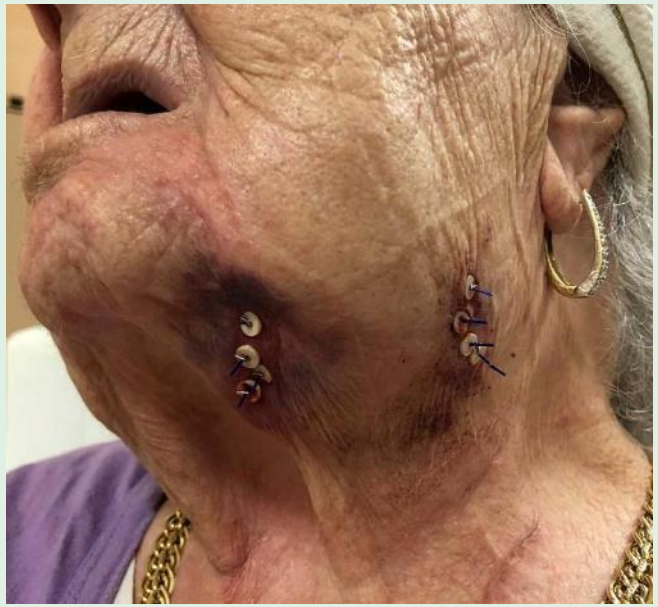
Case Background – HNCPI-00-01-003

Age	96
Sex	Female
Tumor Type	SCC
Date of First Diagnosis	Jul-2022
Location	Alveolar ridge & lip plus dermal involvement
Prior Treatments	None
Medical Background	• Cardio • Dementia • ECOG3
Cancer Stage	• Stage IV • T2N1M1

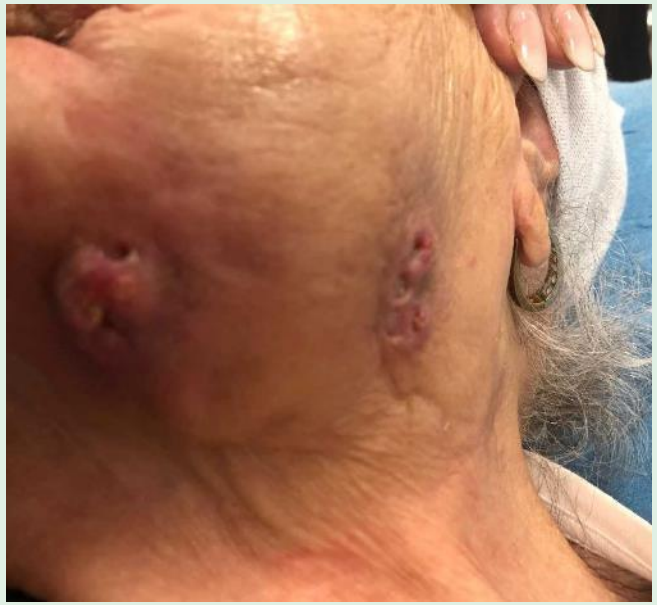




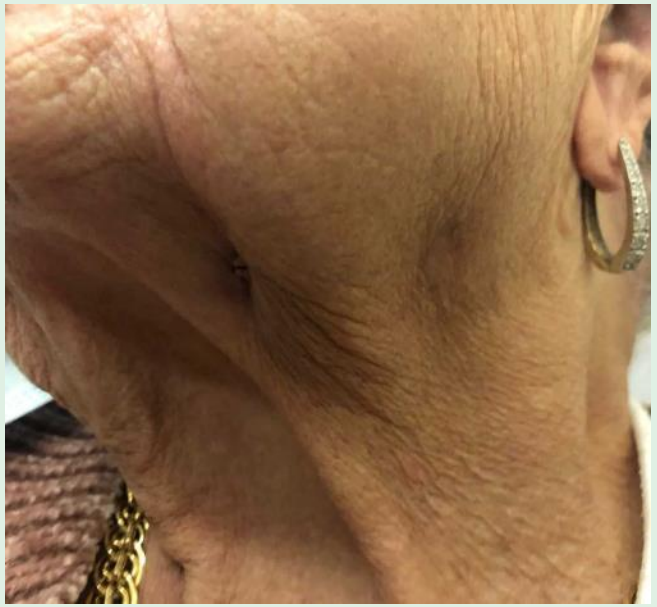
Alpha DaRT Treatment



Alpha DaRT Insertion
Sept-2022



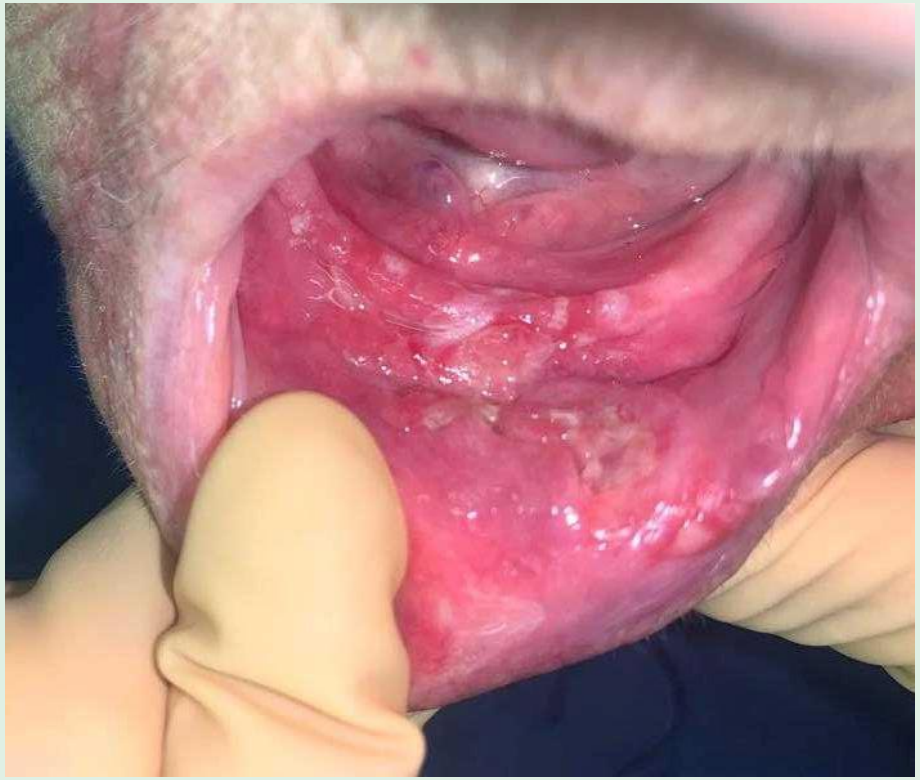
After Alpha DaRT Removal
Oct-2022



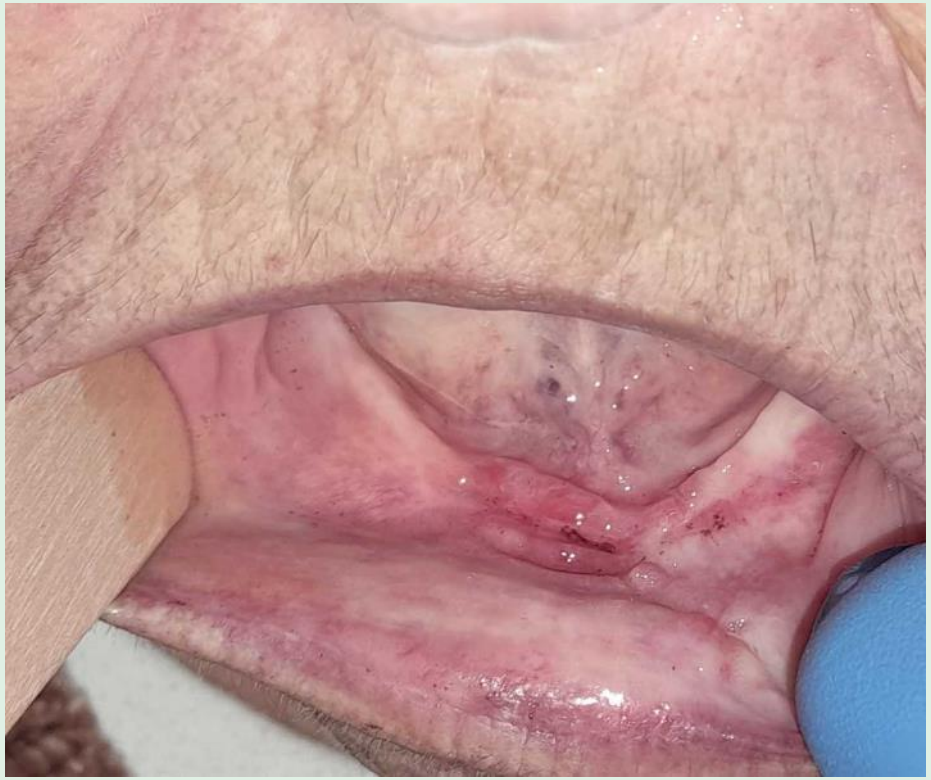
Follow-Up

Jan-2023

Clinical Follow-Up



Pre-Treatment



Nine Weeks Post Treatment

Patient Status

- Patient stopped Pembrolizumab after 12 months
- Patient still alive with no evidence of disease at October 2024 followup

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Focus on Internal Organ Treatments

We continue to make progress across internal organ programs, with trials underway in multiple targeted indications and others in various stages of planning and start-up

Internal Organs in Focus

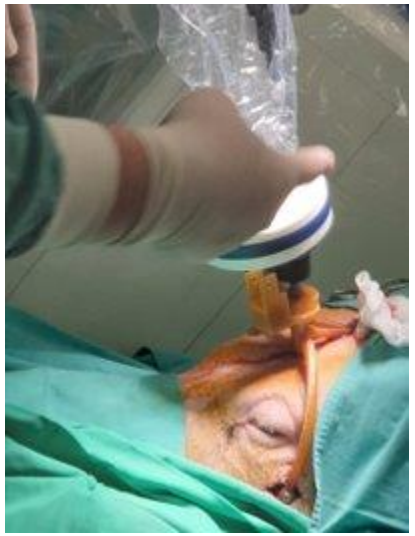
- Pancreas – clinical trial underway
- Liver – clinical trial underway
- Lung – clinical trial underway
- Prostate – clinical trial underway
- Brain – GBM + Brain Mets
- Breast
- Rectum

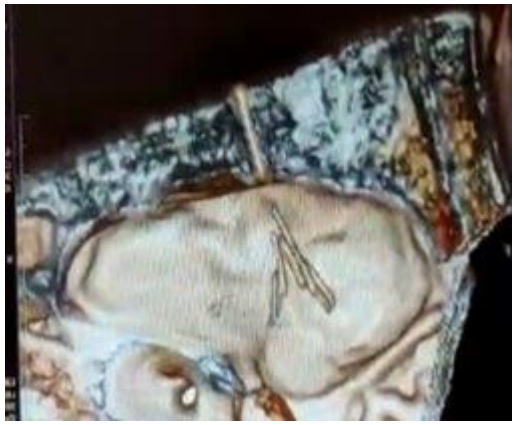
CHUM
Centre hospitalier de l'Université de Montréal

HADASSAH UNIVERSITY MEDICAL CENTER
Founded by Hadassah, the Women's Zionist Organization of America

HGJ | JGH

RAMBAM Health Care Campus





Interim Pancreatic Cancer Results - Overview of Trial Design

Three trials treating pancreatic cancer patients in parallel:

- CTP-PANC-101 monotherapy treatment at 2 sites in Montreal, Canada – up to 37 patients total
- CTP-PANC-02 monotherapy treatment at 1 site in Jerusalem, Israel – up to 15 patients total
- CTP-ALL-00 flexible basket trial at 1 site in Jerusalem, Israel – no specified limit on number of patients

Following initial results, there are some situations where chemotherapy has been used in the first two trials

- CTP-PANC-101 allows chemotherapy 30 days after Alpha DaRT treatment
- CTP-PANC-02 was modified to allow concomitant chemotherapy

Therefore, after initially embarking on monotherapy exploration, **a small number of patients from all three trials have received chemotherapy treatment alongside or following Alpha DaRT treatment**

Due to the exploratory nature of the trials, they do not focus on a specific patient sub-population but rather a broad mix of patients with non-resectable pancreatic cancer

High Disease Control Rate Observed

Among the 41 patients treated, 33 had a measured objective response, with 5 patients awaiting response evaluation and 3 who discontinued prior to evaluation. Results are presented below using Best Overall Response (BOR) for those with a measured response.

Including first two patients (heavily underdosed / feasibility only)	**18%** Objective Response Rate (CR + PR)	**91%** Disease Control Rate (CR + PR + SD)
Excluding first two patients (heavily underdosed / feasibility only)	**19%** Objective Response Rate (CR + PR)	**97%** Disease Control Rate (CR + PR + SD)

Note: Results as of January 8, 2025

Highlights of Overall Survival (OS) Data

Key Caveats:

- The data are still **relatively immature, but ongoing**
- Trial designs were **focused on feasibility and safety**, without the frequent monitoring visits common in studies focused on precise measurement of survival
- Five patients treated since Nov 25, 2024, and three patients who exited the study very shortly after treatment, in all cases with insufficient time to reach objective response measurement, were excluded from OS analysis for lack of data maturity
  - Therefore, a total of n = 33 patients are evaluated for OS using Kaplan-Meier analysis

Population	OS Since Diagnosis / Initiation of Last Chemotherapy (mo)	OS Since Alpha DaRT Treatment (mo)
Overall Population (n=33)	18.6	10.9

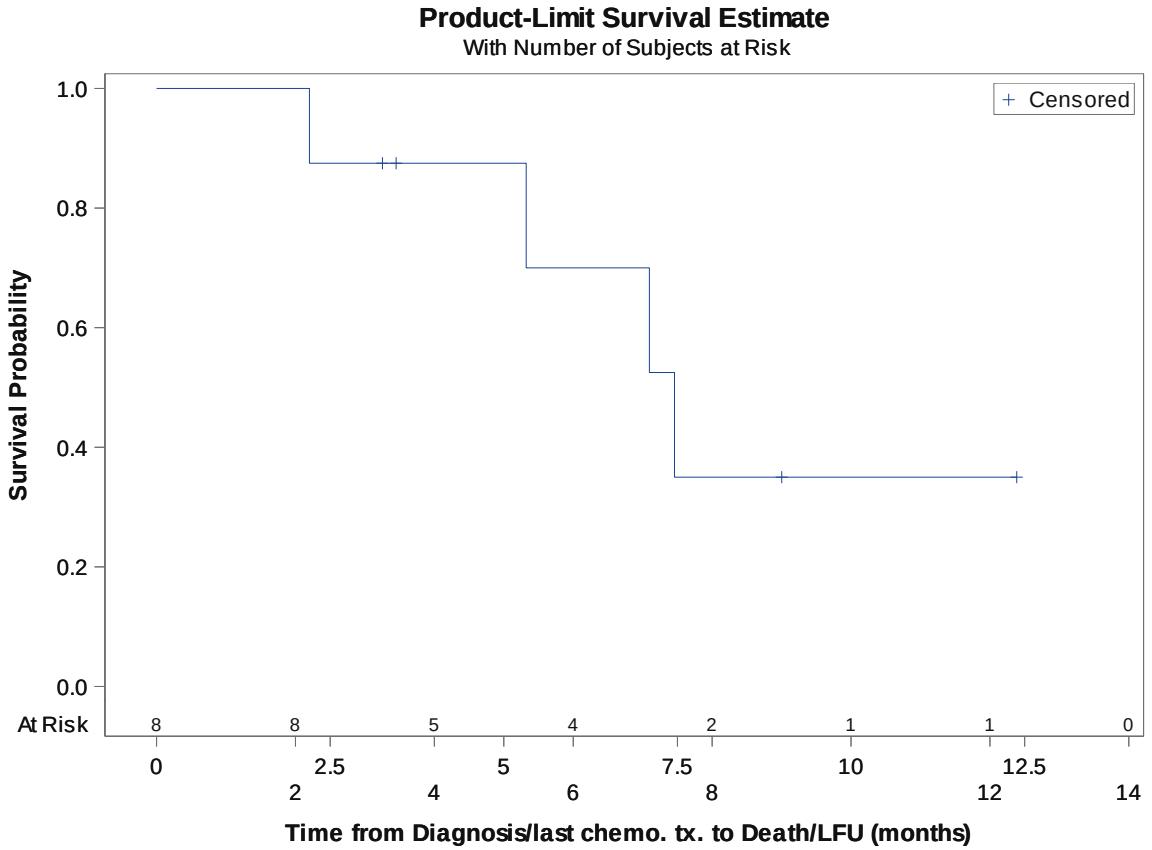
Of n=33 patients analyzed, 13 have died
The remaining 20 (and the five newer patients) remain alive

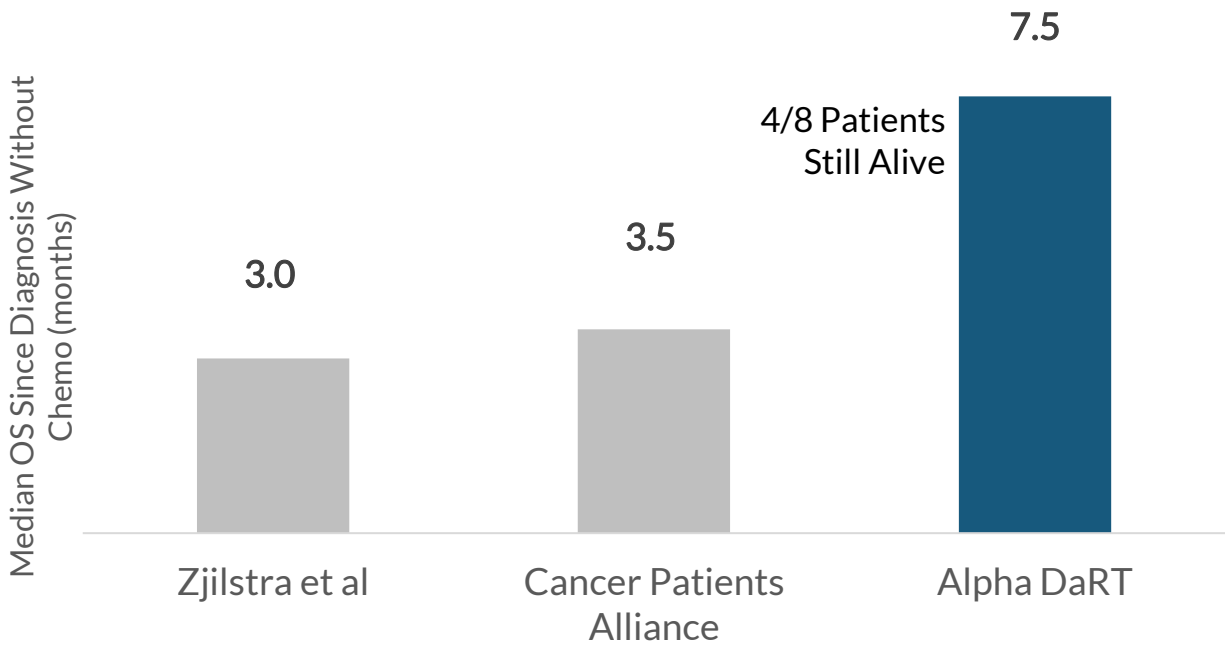
In light of the **heterogeneity of the population**, we conducted ad-hoc analyses **of key sub-groups** to offer context vs. expected OS for each group

Note: Results as of January 8, 2025

Analysis of Overall Survival in Key Sub-Populations (1/3)

Newly Diagnosed / Not Eligible for Chemotherapy (n=8)

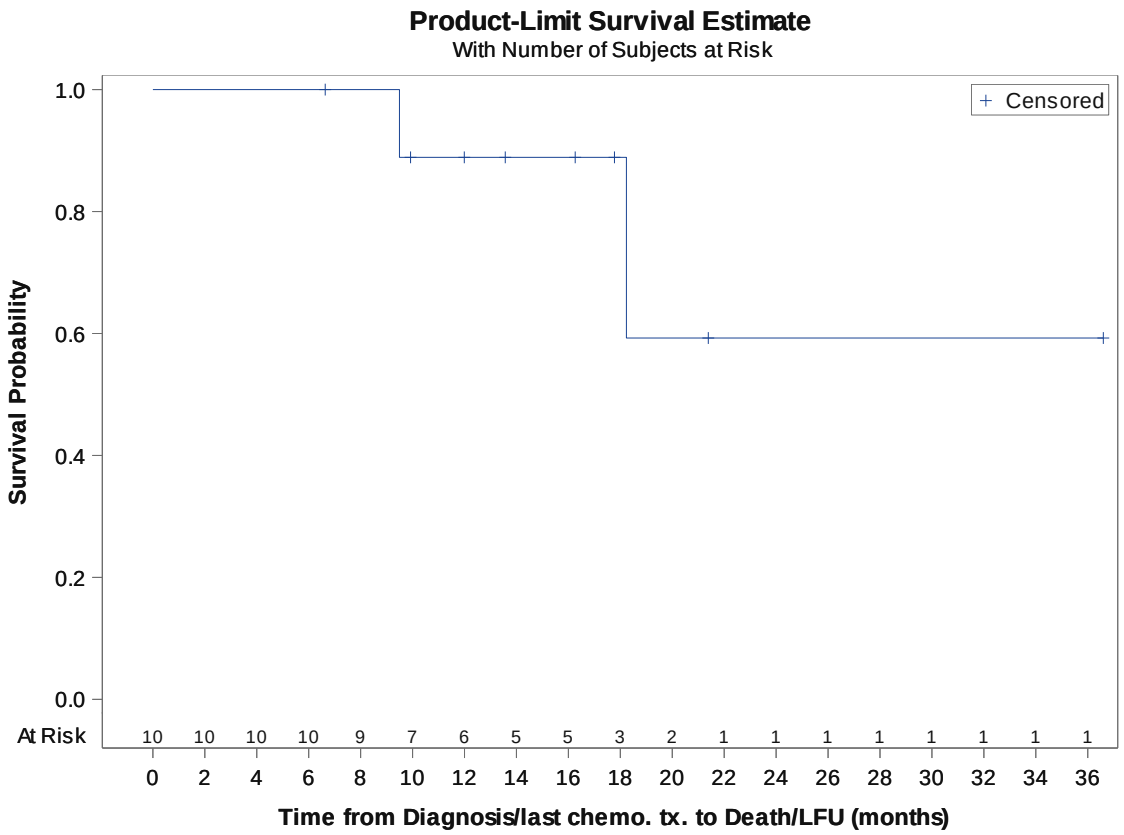


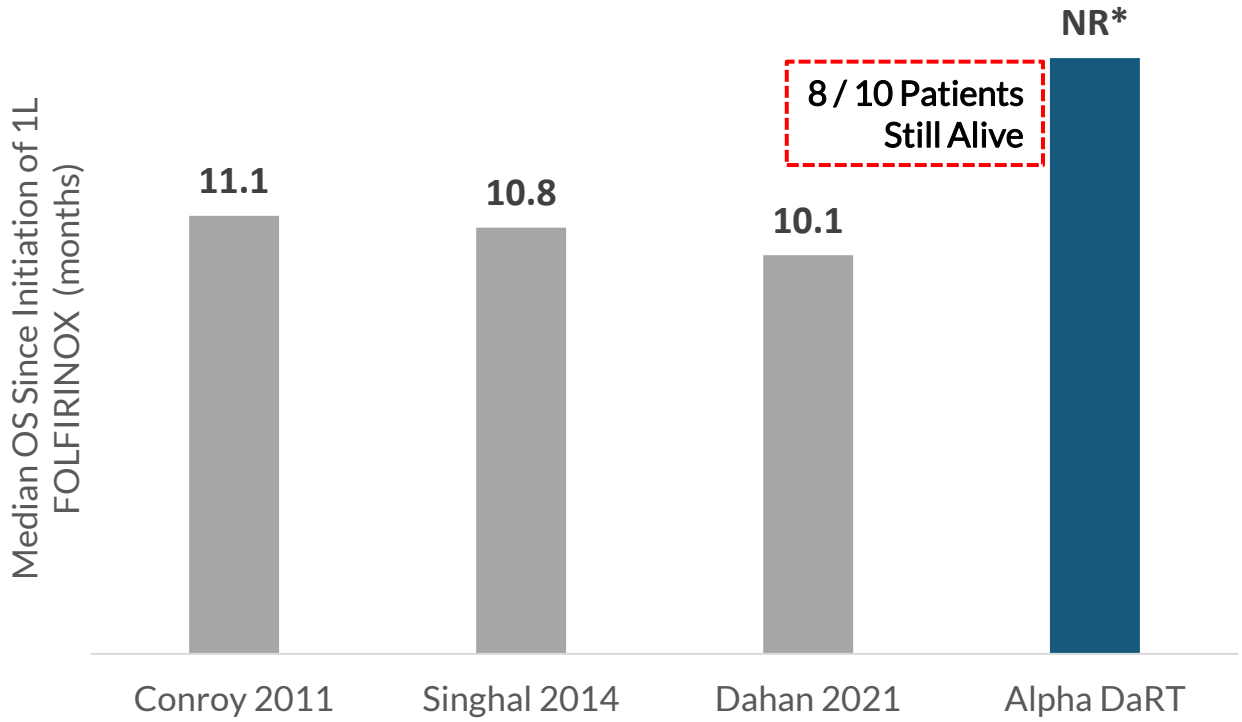


Note: Median follow-up in Alpha DaRT group of 6.3 months
Results as of January 8, 2025
For Illustrative Purposes Only - no head-to-head clinical study has been conducted comparing Alpha DaRT to other products or candidates. Differences exist between trial designs, subject characteristics and other factors, and caution should be exercised when comparing data across unrelated studies
Sources:
Zijlstra, M. et al (2018). Patient characteristics and treatment considerations in pancreatic cancer: a population based study in the Netherlands. https://doi.org/10.1080/0284186X.2018.1470330
https://pancreatica.org/pancreatic-cancer/pancreatic-cancer-prognosis/

Analysis of Overall Survival in Key Sub-Populations (2/3)

Metastatic (Stage IV) Patients After 1L FOLFIRINOX (n=10)





** Median Kaplan-Meier estimate was not reached (NR); median follow-up time was 15.1 months*

For Illustrative Purposes Only - no head-to-head clinical study has been conducted comparing Alpha DaRT to other products or candidates. Differences exist between trial designs, subject characteristics and other factors, and caution should be exercised when comparing data across unrelated studies

Note: Results as of January 8, 2025

Sources:

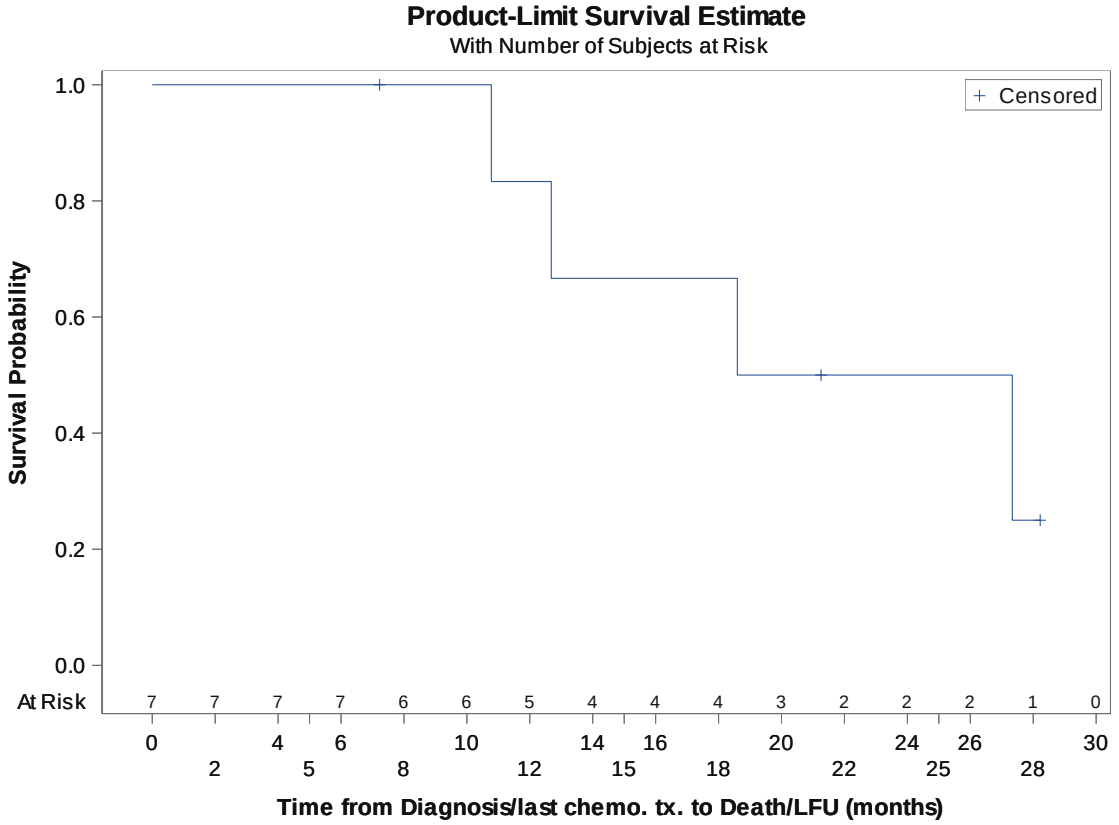
Thierry Conroy et al., FOLFIRINOX versus Gemcitabine for Metastatic Pancreatic Cancer. New England Journal of Medicine (2011). DOI: 10.1056/NEJMoa1011923

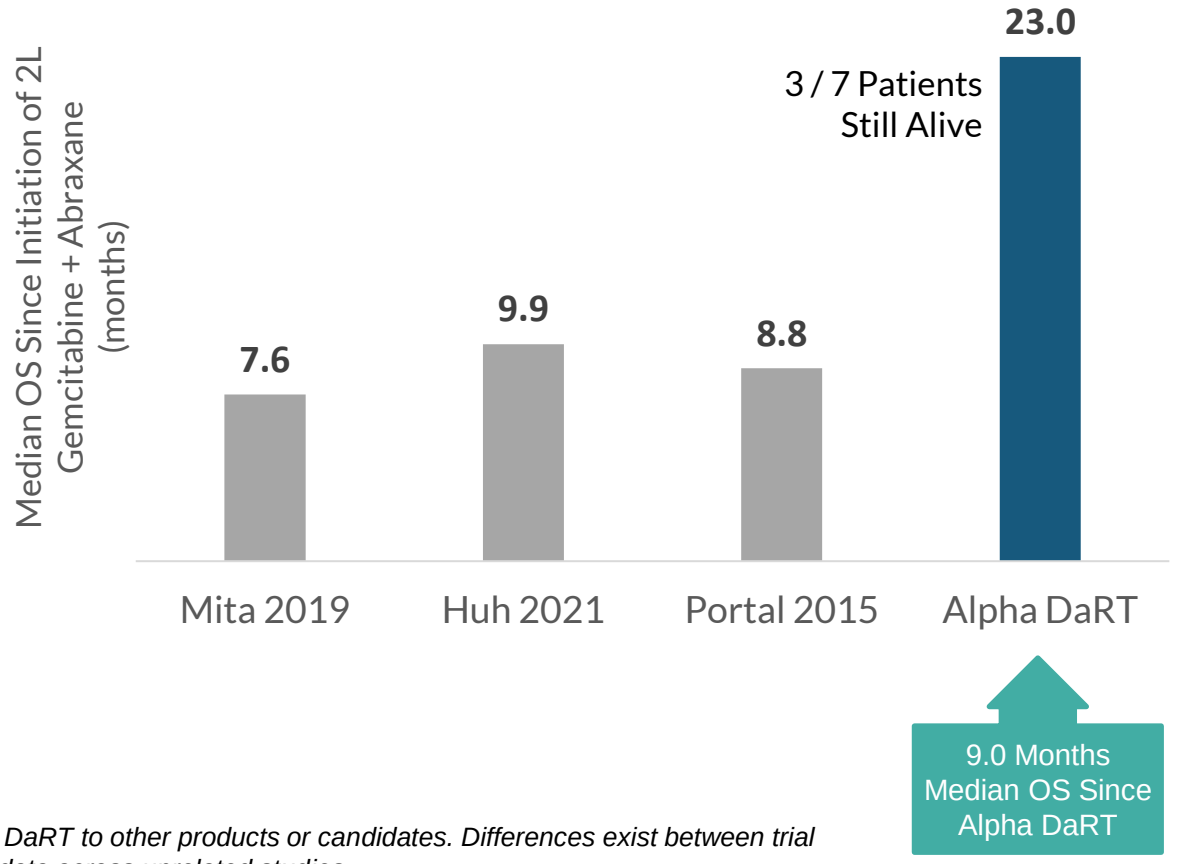
Singhal MK, et al. A phase III trial comparing FOLFIRINOX versus gemcitabine for metastatic pancreatic cancer. Ann Oncol. 2014;25(suppl 4):iv210–53.

Laetitia Dahan et al., Randomized Phase II Trial Evaluating Two Sequential Treatments in First Line of Metastatic Pancreatic Cancer: Results of the PANOPTIMOX-PRODIGE 35 Trial. JCO 39, 3242-3250(2021). DOI:10.1200/JCO.20.03329

Analysis of Overall Survival in Key Sub-Populations (3/3)

Progressed After 2L Gemcitabine-Abraxane (n=7)





Note: Median follow-up in Alpha DaRT group of 18.9 months

For Illustrative Purposes Only - no head-to-head clinical study has been conducted comparing Alpha DaRT to other products or candidates. Differences exist between trial designs, subject characteristics and other factors, and caution should be exercised when comparing data across unrelated studies

Note: Results as of January 8, 2025

Source:

Mita N, Iwashita T, Uemura S, Yoshida K, Iwasa Y, Ando N, Iwata K, Okuno M, Mukai T, Shimizu M. Second-Line Gemcitabine Plus Nab-Paclitaxel for Patients with Unresectable Advanced Pancreatic Cancer after First-Line FOLFIRINOX Failure. J Clin Med. 2019 May 29;8(6):761. doi: 10.3390/jcm8060761. PMID: 31146420; PMCID: PMC6616879

Huh G, Lee HS, Choi JH, Lee SH, Paik WH, Ryu JK, Kim YT, Bang S, Lee ES. Gemcitabine plus Nab-paclitaxel as a second-line treatment following FOLFIRINOX failure in advanced pancreatic cancer: a multicenter, single-arm, open-label, phase 2 trial. Ther Adv Med Oncol. 2021 Nov 10;13:17588359211056179. doi: 10.1177/17588359211056179. PMID: 34790261; PMCID: PMC8591648.

Portal A et al. Nab-paclitaxel plus gemcitabine for metastatic pancreatic adenocarcinoma after Folfirinox failure: an AGEO prospective multicentre cohort. Br J Cancer. 2015 Sep 29;113(7):989-95. doi: 10.1038/bjc.2015.328. Epub 2015 Sep 15. PMID: 26372701; PMCID: PMC4651133.

Pancreatic Cancer Clinical Trial: FDA Approval of IDE for U.S. Pilot

N=15

Newly Diagnosed
Locally Advanced Pancreatic Cancer

N=15

Newly Diagnosed
Metastatic Pancreatic Cancer

AlphaDaRT
Insertion
(During Cycles 1-4)

8-12 cycles of **mFOLFIRINOX**

1

Primary Endpoint:
Incidence of Treatment-Emergent Adverse Events

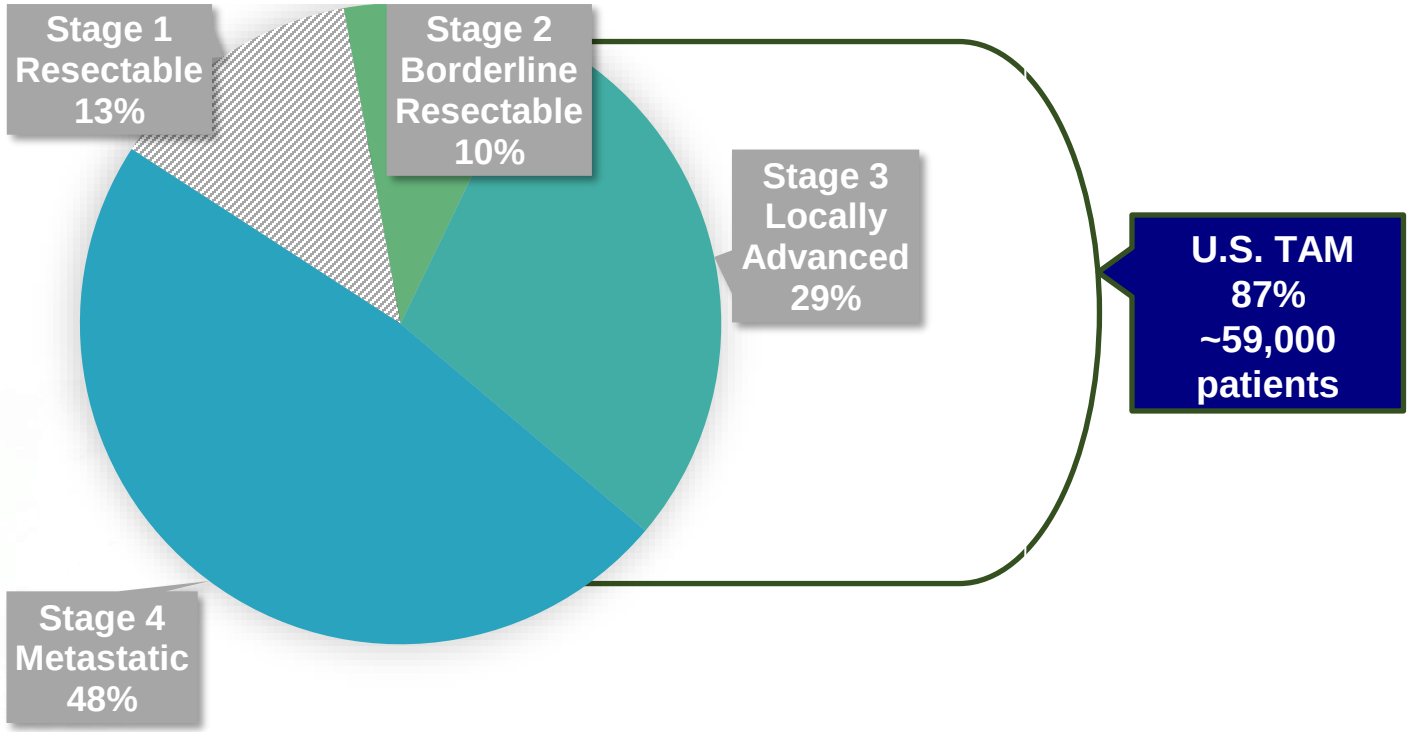
2

Secondary Endpoints:
Overall Survival
Progression-Free Survival
Pain Improvement
Surgical Resectability (LA cohort only)

Pancreatic Cancer: Massive and Growing Unmet Need

There are over half a million new cases of pancreatic cancer per year. Approx. 66k of them are in the U.S.

Incidence rate of pancreatic cancer is trending upward, especially in younger patients

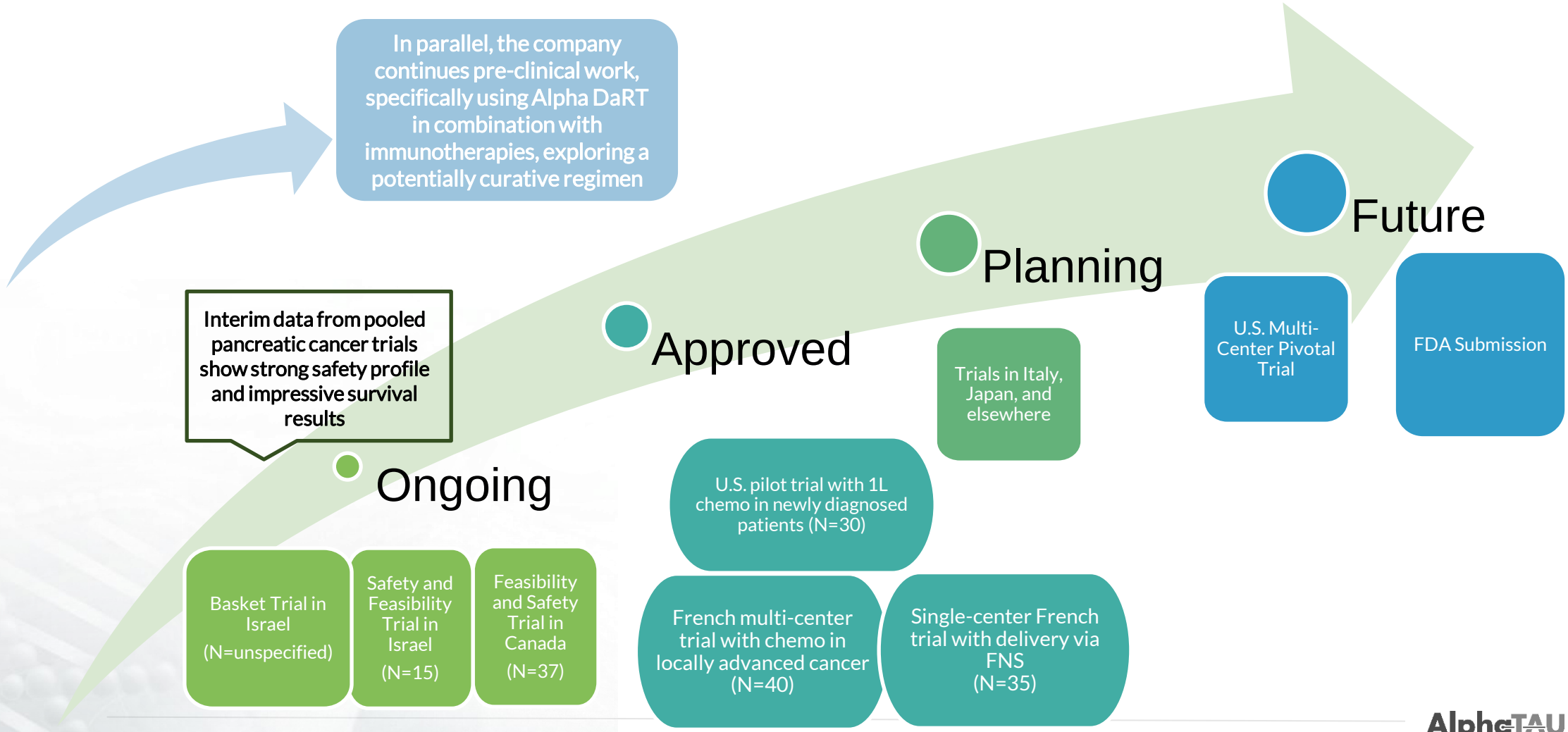


At diagnosis, 87% of pancreatic cancer cases are not eligible for surgical resection and/or have metastasized

Note: Excludes cancers of stage "unknown" or "N/A" - data from 1400 Hospitals
Source: https://www.facs.org/media/ztllhkfu/cancer-cases-reported-to-the-ncdb-by-tumor-type-and-ajcc-stage.pdf
https://gco.iarc.who.int/media/globocan/factsheets/cancers/13-pancreas-fact-sheet.pdf
https://www.cancer.org/cancer/types/pancreatic-cancer/about/key-statistics.html
https://www.thelancet.com/journals/langas/article/PIIS2468-1253%2823%2900039-0/fulltext

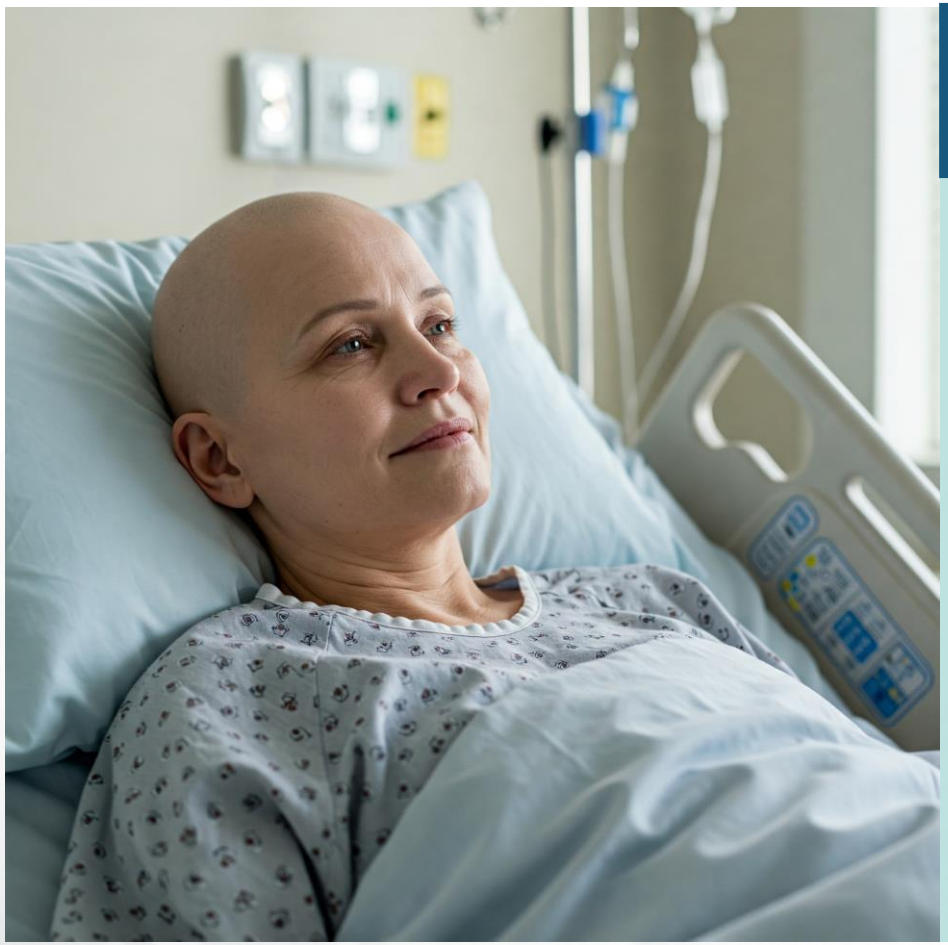
Increasing Momentum in Pancreatic Cancer "Killer Application"

The feasibility and safety of delivering Alpha DaRT sources to solid tumors in the pancreas has been the focus, but we continue to build momentum toward a pivotal clinical pathway in the U.S. with an eye toward a potential submission for FDA authorization

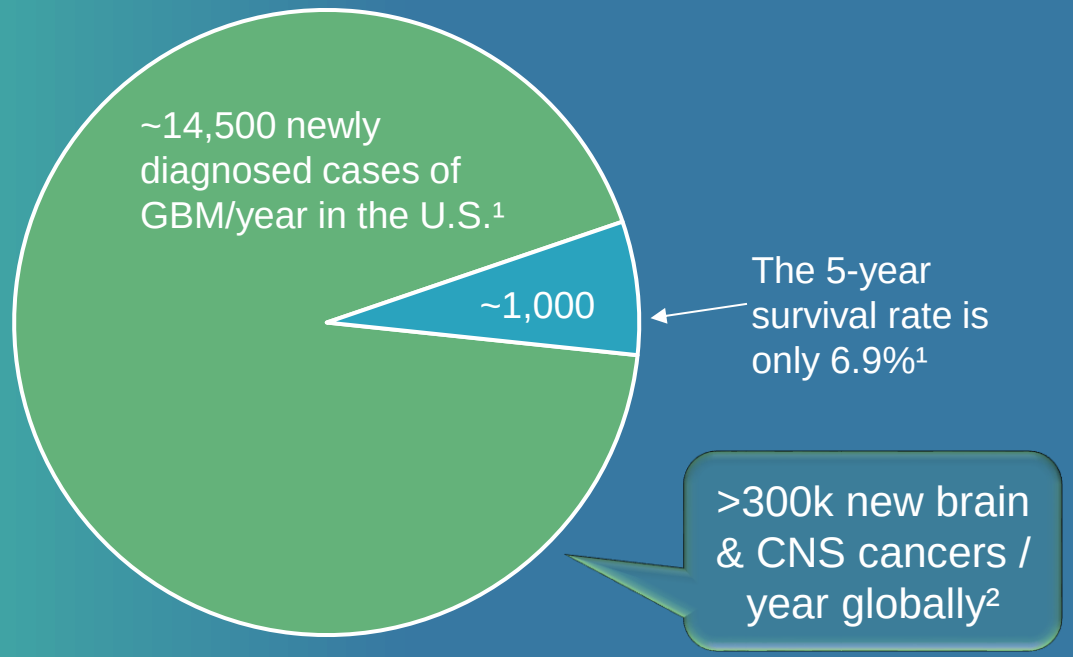


The Next Frontier: Glioblastoma Multiforme (GBM) / Brain Cancers

GBM is one of the most complex, deadly, and treatment-resistant cancers, with an average length of survival estimated at only 8 months[1]. Alpha Tau has IDE approval to initiate a U.S. pilot study in GBM.



There is a high unmet need for a life extending treatment for GBM, with a mortality rate of 93% after 5 years[1]

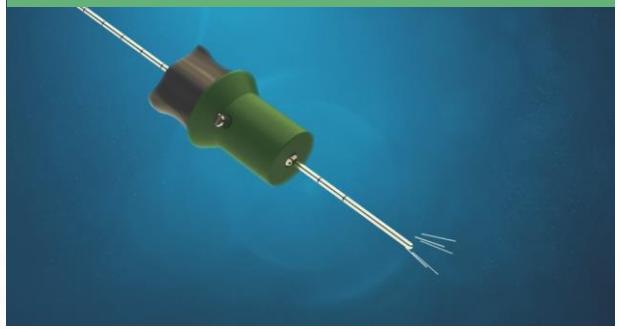


[1]*https://braintumor.org/events/glioblastoma-awareness-day/about-glioblastoma/*
[2]*https://gco.iarc.who.int/media/globocan/factsheets/cancers/31-brain-central-nervous-system-fact-sheet.pdf*

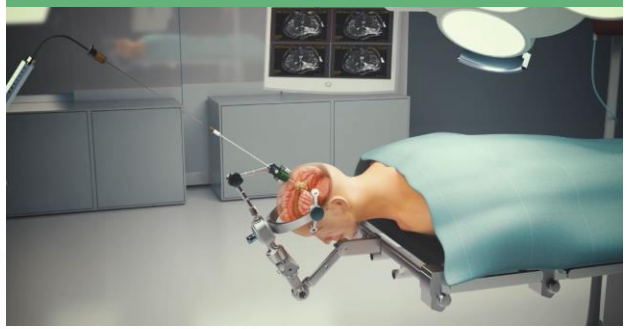
Radial Applicator & Pre-Clinical Study in Swine Brain

Alpha Radial Applicator for the Delivery of Alpha DaRT into the Brain

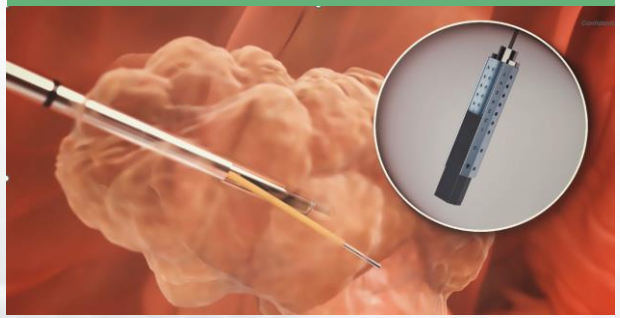
Designed to deliver sources in precise spacing while minimizing damage to the brain



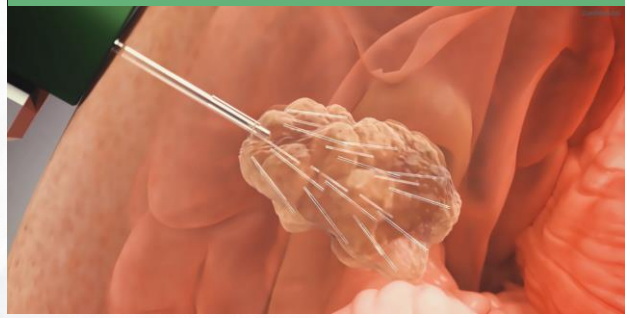
A stereotactic biopsy needle is inserted into the target area of the brain. The alpha radial applicator is affixed to the biopsy needle hub



The physician pushes the flexible applicator tube into the tumor, and once in place, pushes the stylet forward and retracts the tube



The needle is then rotated to the next position, to deliver a layer of sources without inserting the needle repeatedly



RESEARCH

Journal of Neuro-Oncology

Stereotactic implantation of diffusing alpha-emitters radiation therapy sources in the swine brain: a potential new focal therapy for brain tumors

Yigal Shoshan[1] · Moshe J. Gomori[2] · Lior Moss[3] · Saleem Eben Bari[3] · Nir Edery[3] · Robert B. Den[4] · Lior Arazi[5] · Aron Popovtzer[6] · Jon Feldman[6] · Samuel Moscovici[1]

Pre-Clinical (Swine) Study Results

- Alpha-DaRT sources were reproducibly and efficiently delivered to the brain cortex and subcortex
- No unexpected abnormalities in blood / CSF
- No evidence of major bleeding or infection
- Minimal spacial and temporal movements of sources

Conclusion: Alpha-DaRT sources can be safely delivered into a large animal brain using image-guided stereotactic implantation

Radial Applicator Overview

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

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

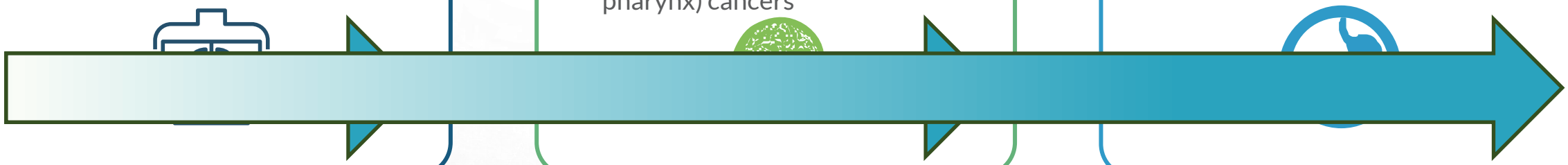
- Localized tumors that are not surgical candidates and tumors that recur after surgery and are **resistant to other** therapies, specifically radiotherapy
- Alpha DaRT to be evaluated as **a later line therapy**
- Tumor types we are targeting include **SCC, H&N SCC and prostate**

Metastatic

- Alpha DaRT being evaluated for its potential to induce an **immune response** in **metastatic** tumors
- Alpha DaRT being evaluated **in combination with checkpoint inhibitors** as an adjuvant therapy
- Tumor types we are targeting include **liver, breast and H&N** (which includes lip, oral cavity, salivary glands, oropharynx & pharynx) cancers

High Unmet Need

- Solid tumors that have **limited treatment options** with limited standard of care offering
- Alpha DaRT could potentially target **broad patient populations**
- Tumor types we are targeting include **GBM and pancreatic cancer**



Anticipated Milestones

Geography	Target Indication	H1 2025	H2 2025	H1 2026
United States	Recurrent Cutaneous SCC		Completion of multi-center pivotal trial recruitment	Data Readout + Potential FDA submission
	Pancreatic Cancer	First Patient in Pilot Study	Complete Recruitment in Pilot Study	Readout from Pilot Study
	Recurrent GBM	Early Feasibility Study IDE		Readout from Early Feasibility Study
Israel	Brain Cancer (GBM or Metastases)	Targeted first patient treated		
Europe	Pancreatic Cancer (French Multicenter)		Targeted first patient treated	
Japan	Head & Neck Cancer		PMDA Response	

Clinical | Regulatory

Development Pipeline

FDA Breakthrough Device Designation Received for certain uses in skin cancer and GBM

Indication	Geography	Pre-Clinical Research / Feasibility Trial / Pivotal Trial / Marketing Authorization	Notes / Anticipated Milestones
Skin Cancer	U.S.	Recurrent Cutaneous SCC (ReSTART)	• Target patient recruitment completion in Q3 2025
	Israel	Skin SCC	
Head & Neck Cancer	Japan		• PMDA application sent Q4 2023, awaiting reply
	Israel	Oral Cavity SCC	
	Israel - la/mHNSCC	Combo with Pembro	• Interim data released Jan 2025
Pancreatic Cancer	Israel		• Interim data released Jan 2025
	U.S.		• IDE approved by FDA
	Canada		• Interim data released Jan 2025
	Europe		• Two approved studies in France; planning in Italy
	Japan		• Trial in planning
Brain (GBM + mets)	Israel		• Targeting first treatment in Q2 2025
	U.S.		• IDE approved by FDA
Liver Metastases	Canada		• First patient treated Q2 2024
Lung Cancer	Israel		• First patient treated Q4 2024
Prostate Cancer	Israel		• Two trials underway (focal recurrent + neoadjuvant)
Vulvar Cancer	U.K.		• Trial initiated in Q2 2023

Note: ReSTART trial may also be used for potential ex-U.S. submissions. Additional superficial tumor studies not listed above

Global Manufacturing Facilities

For efficient commercial operations, we look to establish manufacturing operations in multiple regions of the world, to enable relatively short shipping times to our core markets. We are currently building our first commercial-scale facility in Hudson, NH

Hudson, New Hampshire
(First Phase Under Construction – ~400,000 local sources per year)

Jerusalem
(Operational ~200,000 local sources per year)

Lawrence, Massachusetts
(Operational – Producing Generators)

Jerusalem
(In Planning)

Togane, Japan
(In Planning)

Financial Position

- Public Since Mar-2022 (NASDAQ:DRTS)
- $62.9mm in Cash & Deposits at YE 2024
- 2+ Years of Cash Runway

AlphaTAU

Saving Lives Globally