



Targeting IL1RAP to address unmet needs in severe cancer and autoimmune diseases

*Corporate Presentation  
June 2023*

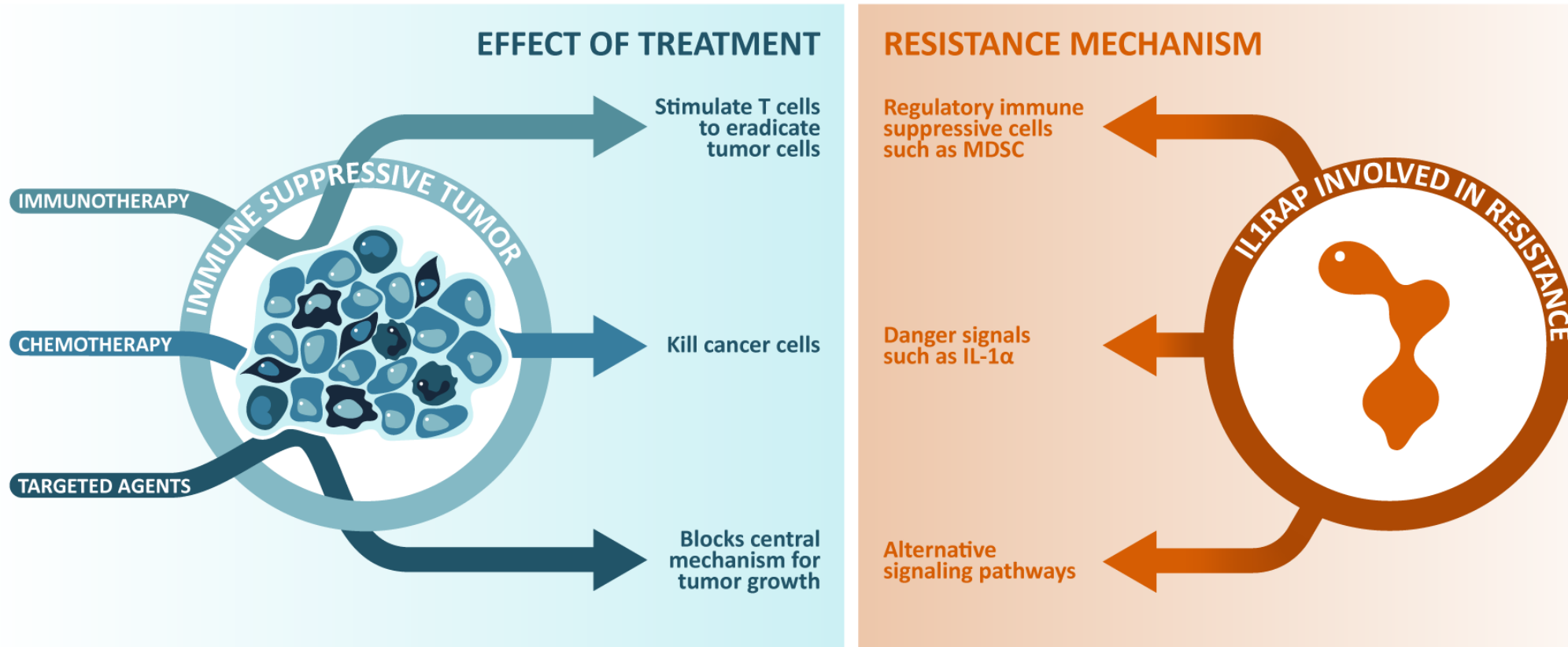
**NASDAQ STOCKHOLM MAIN LIST (CANTA.ST)**

# Safe Harbor Statement



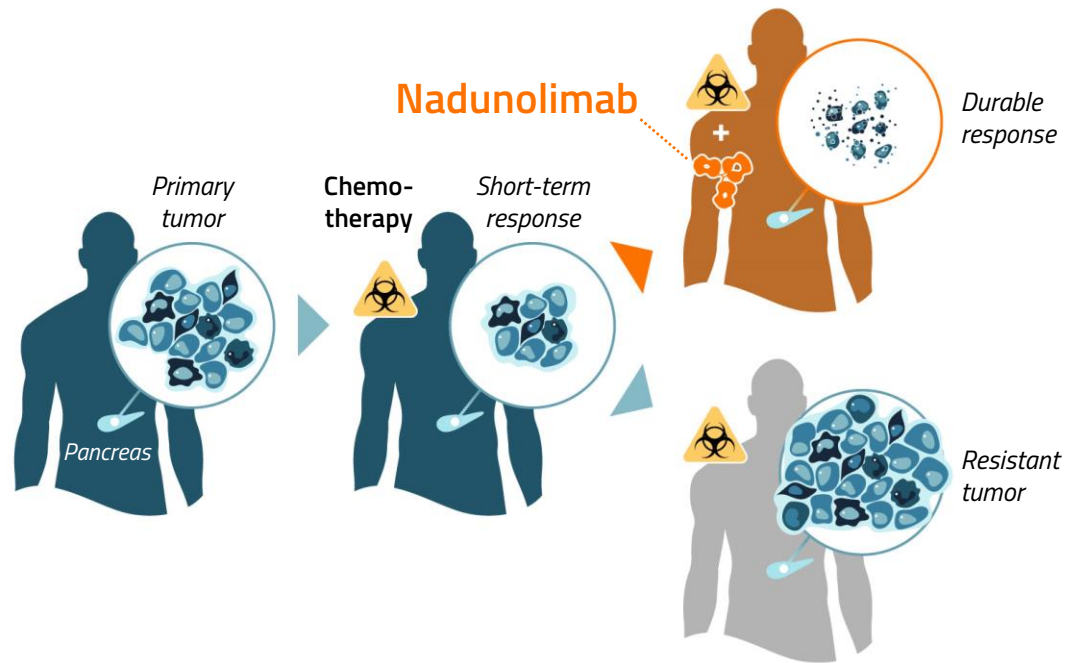
*Statements in the Investor Presentation, including those regarding the possible or assumed future or other performance of the Company or its industry or other trend projections, constitute forward-looking statements. By their nature, forward-looking statements involve known and unknown risks, uncertainties, assumptions and other factors as they relate to events and depend on circumstances that will or may occur in the future, whether or not outside the control of the Company. No assurance is given that such forward-looking statements will prove to be correct. Prospective investors should not place undue reliance on forward-looking statements. They speak only as at the date of this Investor Presentation and the Company undertakes no obligation to update these forward-looking statements. Past performance does not guarantee or predict future performance. Moreover, the Company undertakes no obligation to review, update or confirm expectations or estimates or to release any revisions to any forward-looking statements to reflect events that occur or circumstances that arise in relation to the content of the Investor Presentation.*

# Cantargia – Strategy to improve current cancer therapies



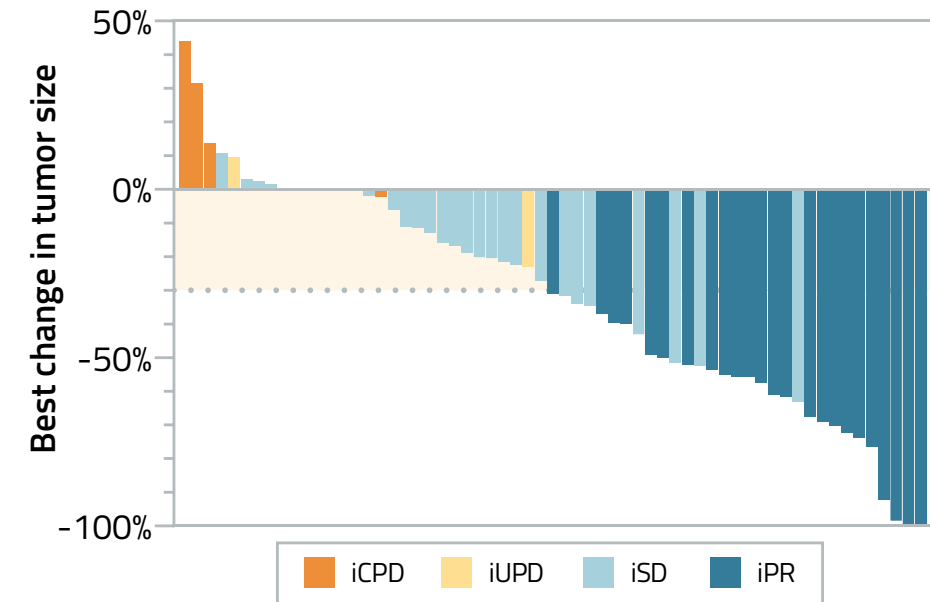
**IL1RAP – A NOVEL TARGET WITH SEVERAL OPPORTUNITIES; CURRENT FOCUS ON SYNERGISTIC COMBINATIONS**

# New strategy to treat cancer supported by clinical results



## PROMISING DATA IN PANCREATIC CANCER

- Stronger efficacy than expected from chemotherapy alone
- Patients with higher IL1RAP level benefit more



SEVERAL LINES OF EVIDENCE SUGGEST NADUNOLIMAB COUNTERACTS CHEMORESISTANCE

# IL1RAP – Broad application in cancer and autoimmune disease

Project	Disease	Type of treatment	Discovery phase	Preclinical phase	Clinical phase I	Clinical phase II	Clinical phase III
Nadunolimab	PDAC	1 <sup>st</sup> line	Gemcitabine/nab-paclitaxel				
	TNBC	1 <sup>st</sup> /2 <sup>nd</sup> line	Carboplatin/gemcitabine				
	NSCLC/ non-squamous NSCLC	1 <sup>st</sup> /2 <sup>nd</sup> line	Platinum doublets				
CAN10	Myocarditis, Systemic sclerosis						
CANxx	New opportunities within IL1RAP platform						

PDAC – pancreatic cancer; TNBC – triple-negative breast cancer; NSCLC – non-small cell lung cancer

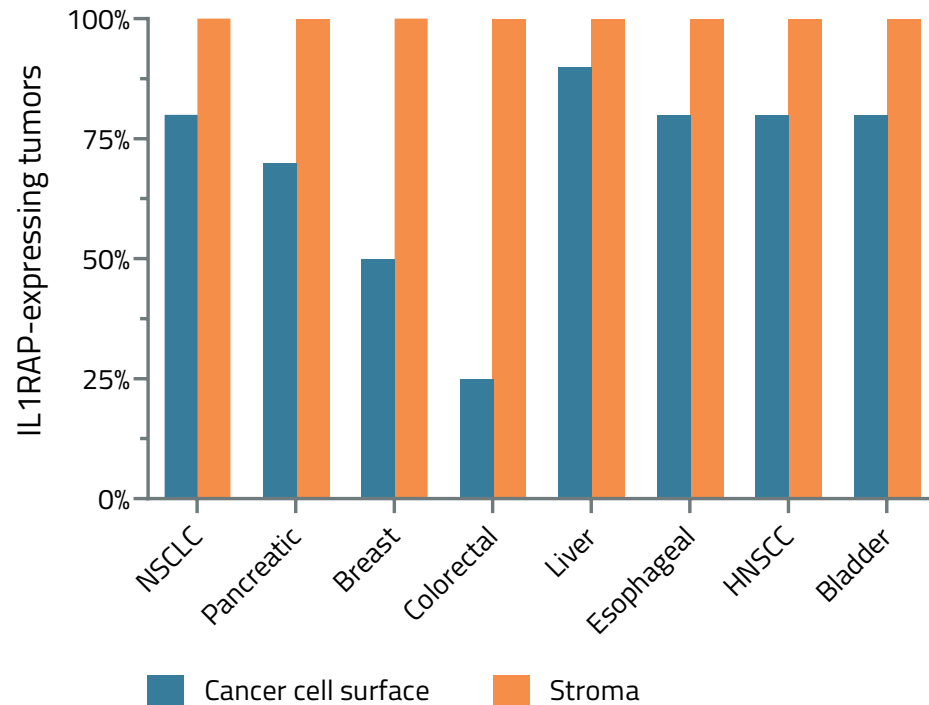


The background of the slide is a microscopic image showing several cells. Two cells are in sharp focus in the upper half, showing a granular, textured surface. The rest of the image is blurred, showing other cells in the background. A semi-transparent dark blue horizontal band runs across the middle of the image, containing the title text in white.

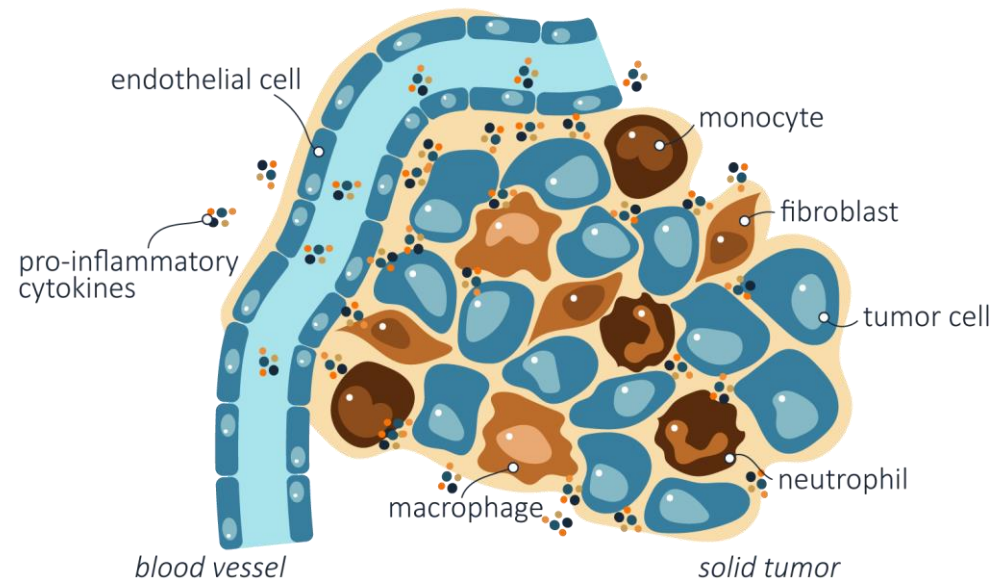
## NADUNOLIMAB (CAN04) OVERVIEW

# IL1RAP overexpressed in most solid tumors

IL1RAP EXPRESSION IN SOLID TUMOR TYPES

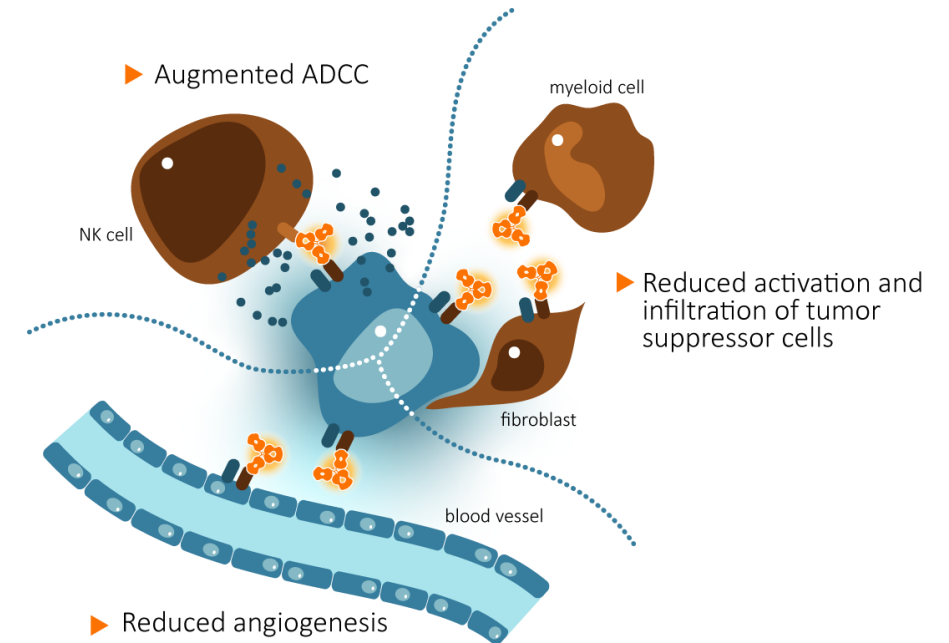
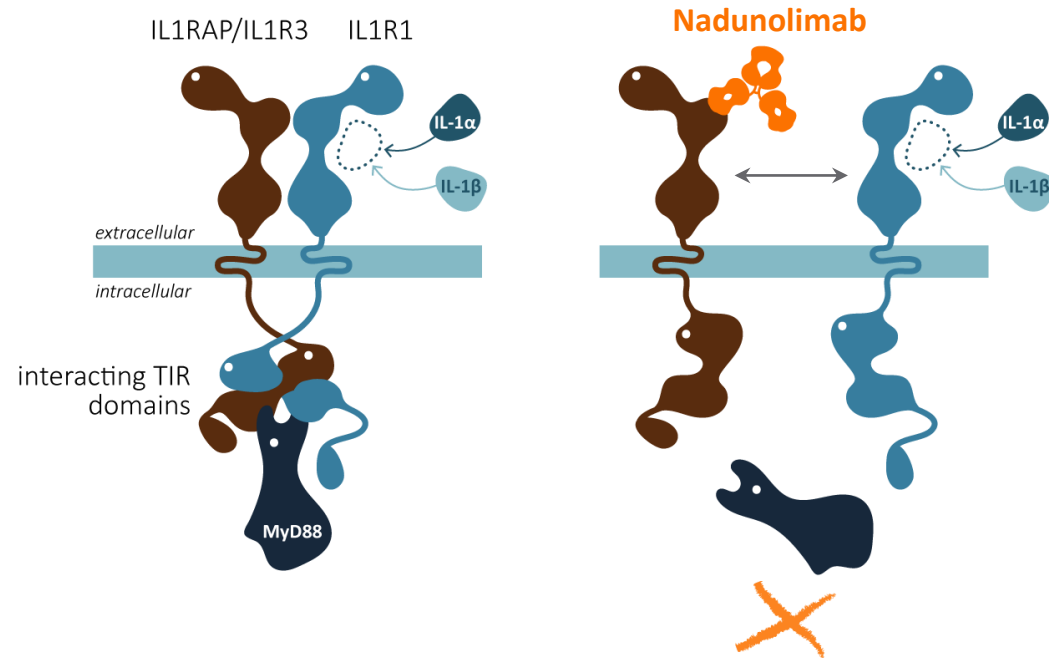


SEVERAL TUMOR-PROMOTING CELLS EXPRESSING IL1RAP IN THE TUMOR MICROENVIRONMENT



IL1RAP – DISTINCTLY OVEREXPRESSED IN TUMORS; LOW EXPRESSION IN NORMAL TISSUE

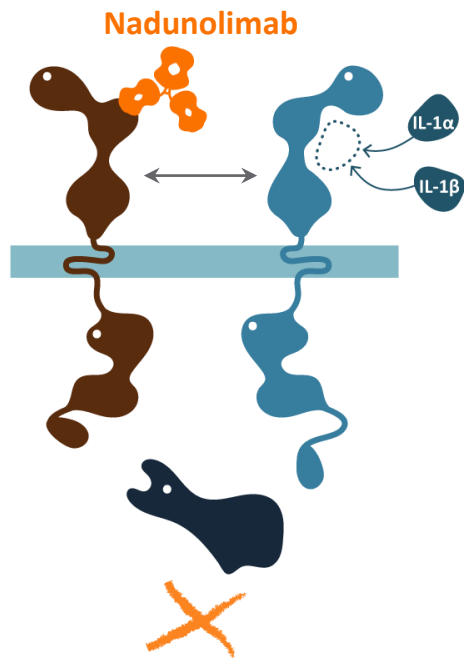
# Targeting IL1RAP provides unique opportunities to treat cancer by IL-1 $\alpha$ / $\beta$ blockade and ADCC



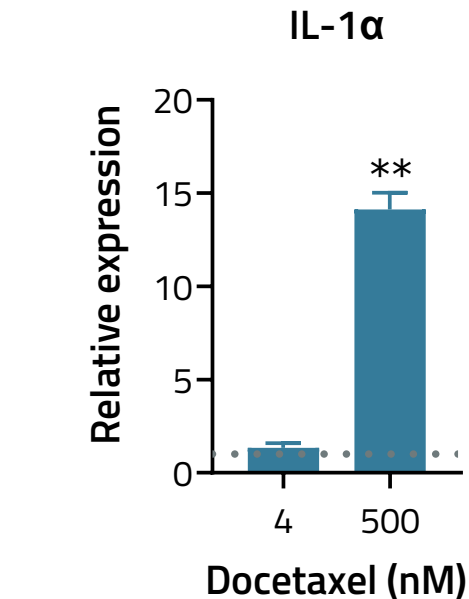
NADUNOLIMAB COUNTERACTS IMMUNE SUPPRESSION AND POTENTIATES THERAPY



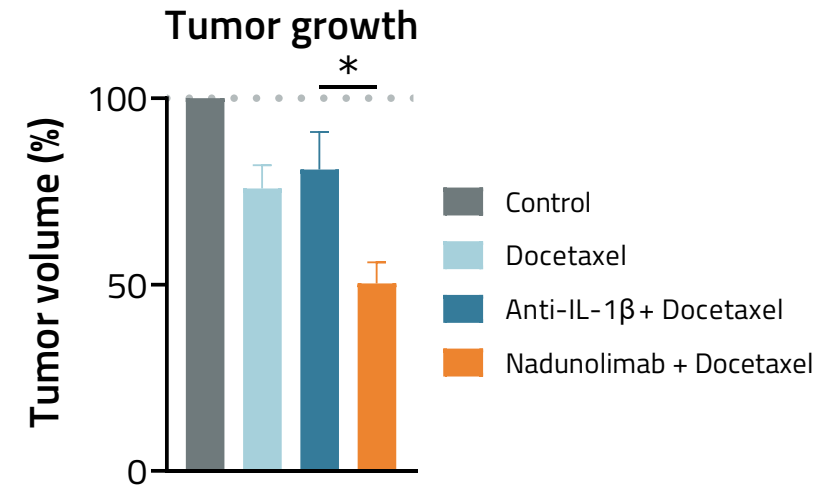
# Nadunolimab potentiates antitumor activity of chemotherapy



→ Nadunolimab blocks both IL-1α and IL-1β signaling



→ Docetaxel induces IL-1α release by tumor cells in vitro



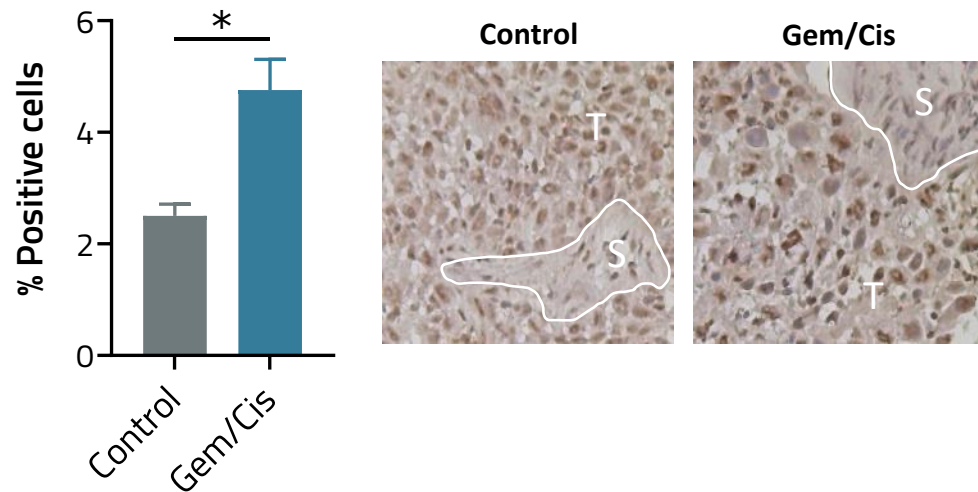
→ Nadunolimab + docetaxel reduces in vivo tumor growth more potently than anti-IL-1β + docetaxel

NADUNOLIMAB INCREASES DOCETAXEL EFFICACY IN CONTRAST TO IL-1β BLOCKADE

# Chemotherapy induces IL-1 $\alpha$ and IL-1 $\beta$ in the tumor

## IL-1 $\alpha$

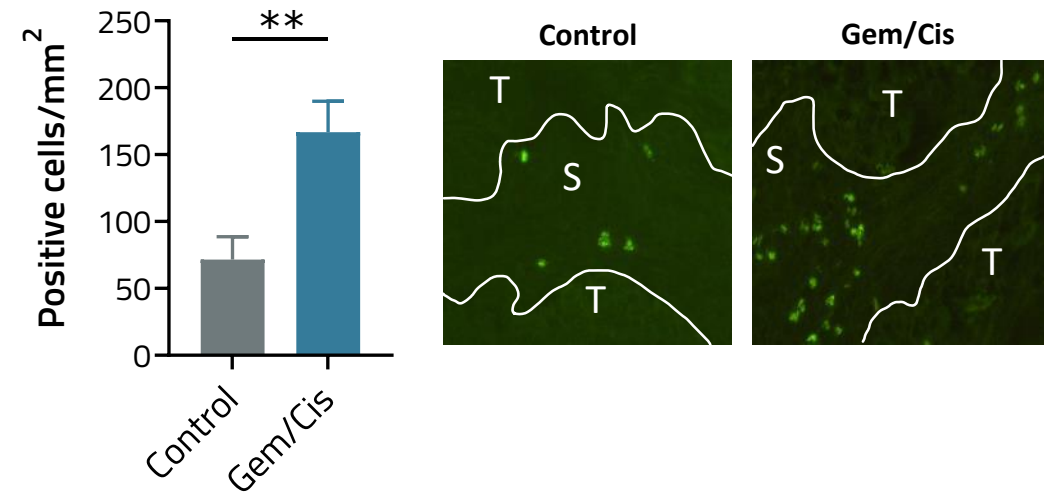
Tumor cell region (T)



→ Gem/Cis induces release of IL-1 $\alpha$  by tumor cells in tumors grown in vivo

## IL-1 $\beta$ -converting enzyme

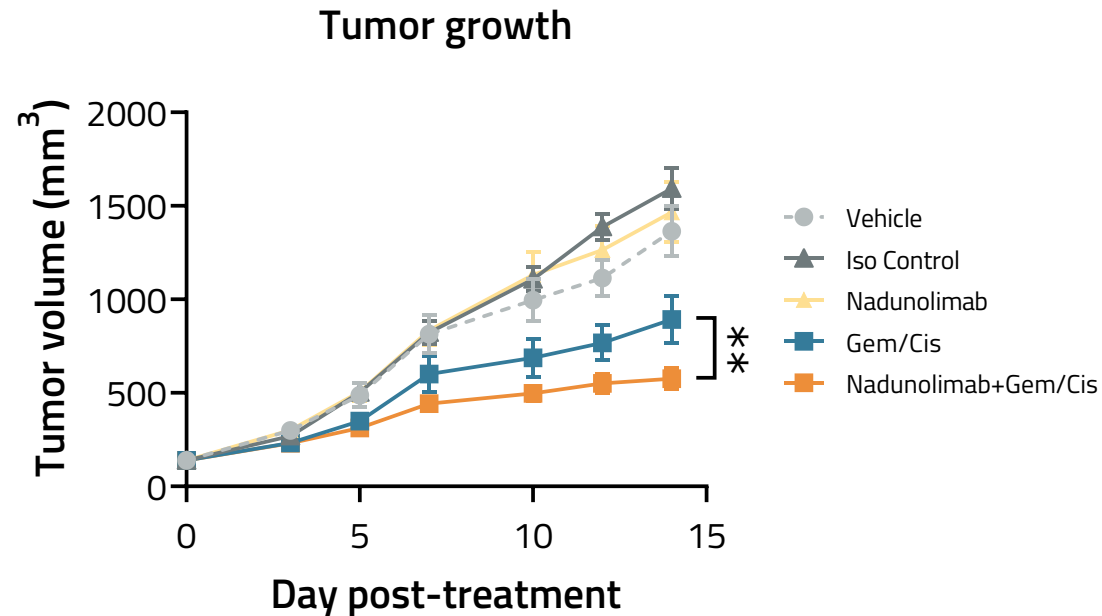
Stromal cell region (S)



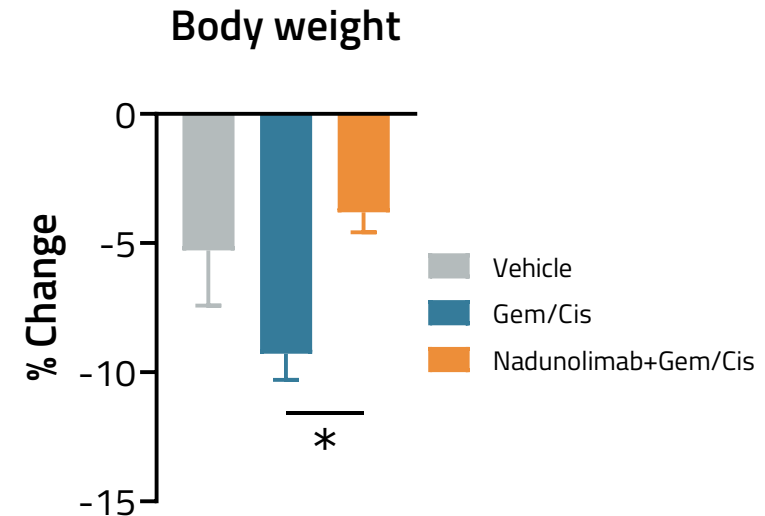
→ Gem/Cis also induces release of IL-1 $\beta$ -converting enzyme (ICE) by stromal cells

INCREASED LEVELS OF IL-1 $\alpha$  AND IL-1 $\beta$  RESULTS IN CHEMORESISTANCE

# Targeting IL1RAP uniquely synergizes with chemotherapy



→ Nadunolimab increases efficacy of platinum-based chemotherapy in vivo



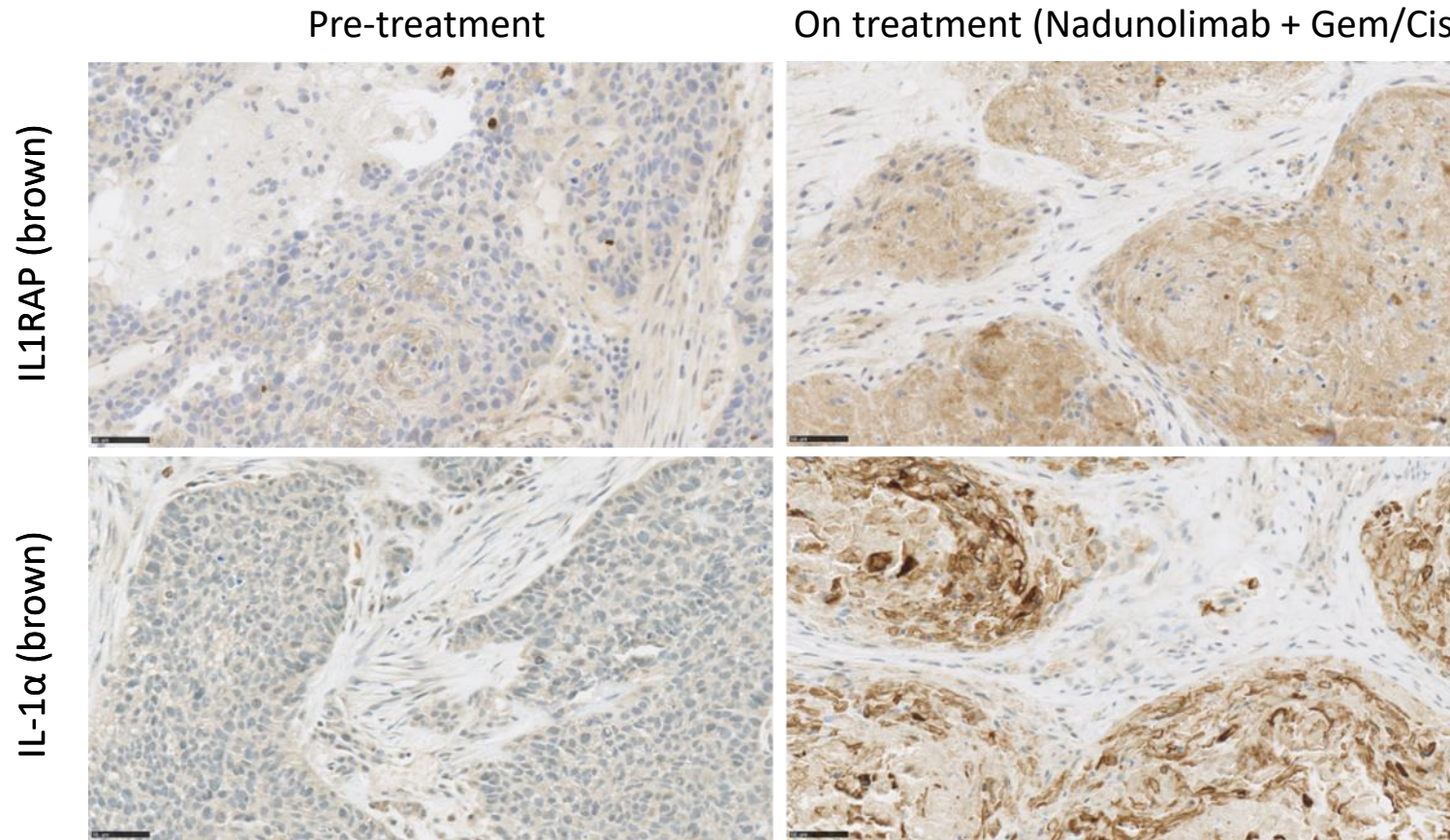
→ Nadunolimab also counteracts weight loss after chemotherapy

**NADUNOLIMAB HAS POTENTIAL TO IMPROVE CHEMOTHERAPY EFFICACY AND TOLERABILITY**

A microscopic image of cells, likely cancer cells, with a blue overlay. The cells are spherical and have a textured, fibrous surface. The background is a soft, out-of-focus blue.

## NADUNOLIMAB (CAN04) CLINICAL RESULTS

# NSCLC – Induction of IL1RAP and IL-1 $\alpha$ with therapy

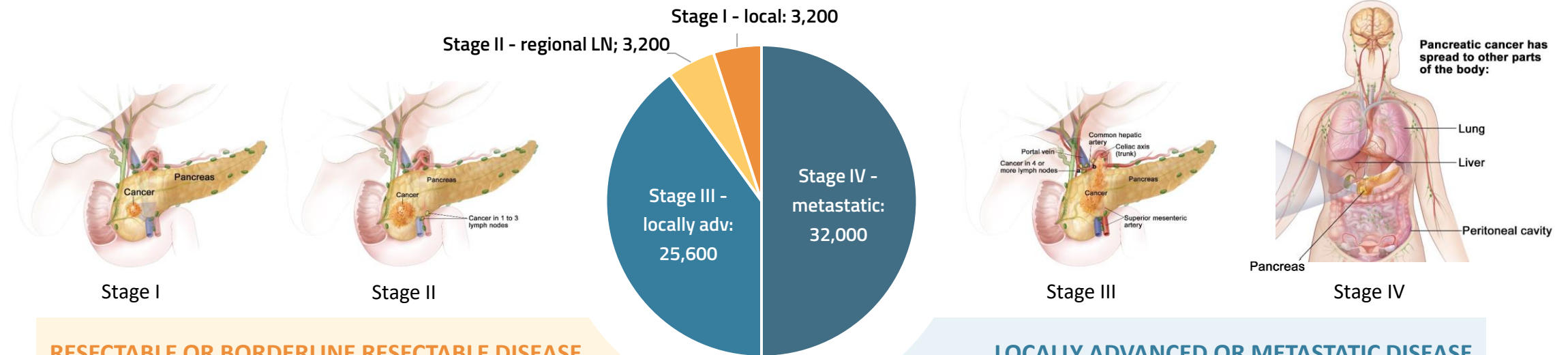


IL-1 $\alpha$  INDUCED BY CHEMOTHERAPY IN LINE WITH PRECLINICAL FINDINGS;  
WELL ESTABLISHED DANGER SIGNAL – ACTIVITY BLOCKED BY NADUNOLIMAB



# PDAC – Staging and treatment

Expected number of cases US 2023: 64,000



## RESECTABLE OR BORDERLINE RESECTABLE DISEASE

### Survival:

→ 15.7 – 28.0 mo

### Treatment:

- Neoadjuvant: gemcitabine+radiotherapy or mFOLFIRINOX
- Radical pancreatic resection
- Post-op chemo (gemcitabine+capecitabine or mFOLFIRINOX) or chemo+radiotherapy

## LOCALLY ADVANCED OR METASTATIC DISEASE

### Survival:

→ 8.5 – 11.1 mo

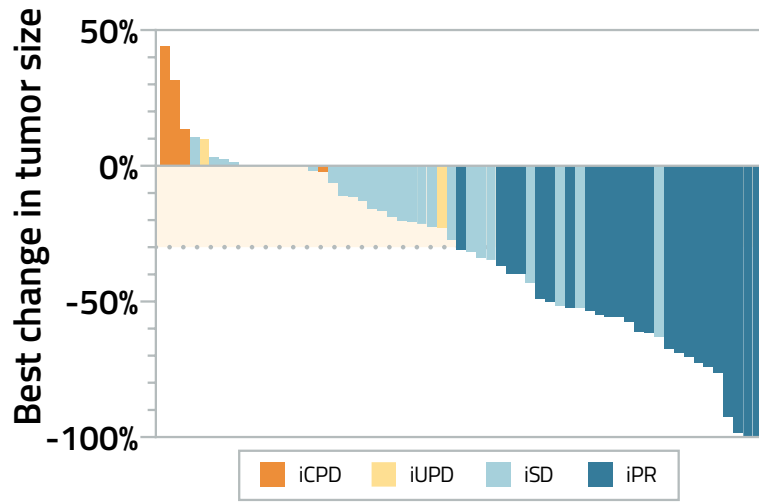
### Treatment:

- FOLFIRINOX if good performance status
- Gemcitabine/nab-paclitaxel
- Gemcitabine if poorer performance
- Jan 2023: 1<sup>st</sup> line NALIRIFOX (OS 11.1 mo)

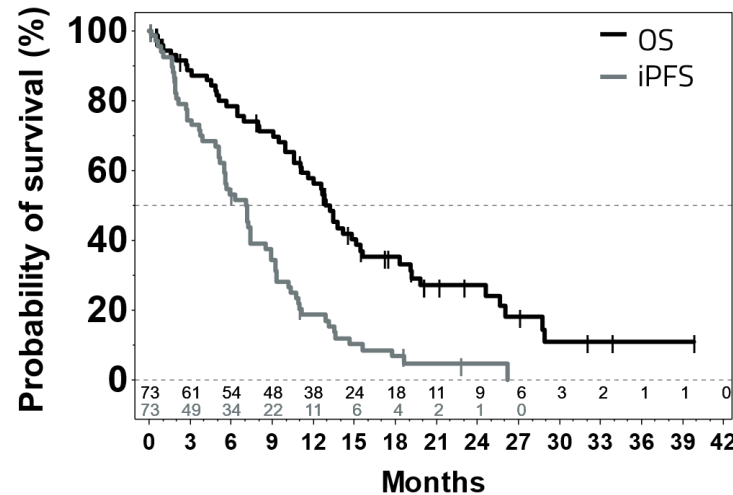
CURRENT DEVELOPMENT FOCUSES ON FIRST-LINE METASTATIC DISEASE WITH POTENTIAL TO MOVE TO EARLIER TREATMENT SETTINGS

# PDAC – Positive interim data in 1<sup>st</sup> line patients

Best responses according to iRECIST



OS and iPFS for mITT patients



## Nadunolimab combination with Gem/Abraxane in 1<sup>st</sup> line PDAC (n=73):

- 33% response rate with long PFS and OS
  - Additional 5 (7%) patients had on-treatment benefit beyond progression
- Promising OS (12.9 mo), PFS (7.2 mo) and DCR (71%)
- 2 patients still on treatment

### Benchmark efficacy Gem/Abraxane:

ORR 23%; DCR 48%; PFS 5.3 mo; OS 8.5 mo (Von Hoff et al, N Engl J Med 2013)

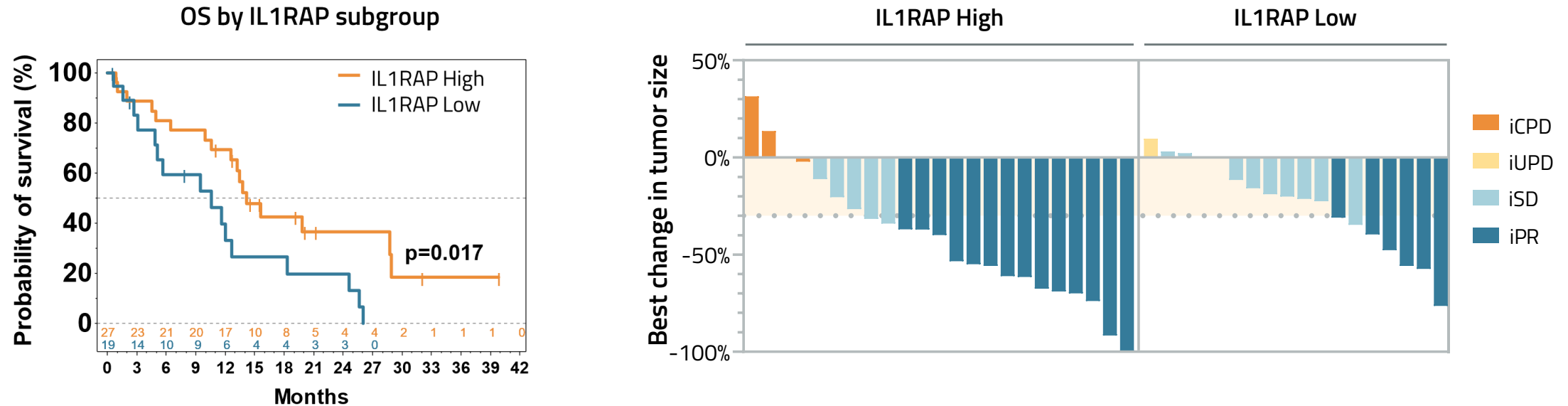
ORR 36%; DCR 62%; PFS 5.6 mo; OS 9.2 mo (NAPOLI 3 trial, ASCO GI 2023)

**PFS AND OS LONGER THAN EXPECTED GIVEN HISTORICAL CONTROL IN PDAC – PHASE IIB TRIAL IN PREPARATION**

iCPD – Confirmed Progressive Disease; iUPD – Unconfirmed Progressive Disease; iSD – Stable Disease; iPR – Partial Response (all according to iRECIST)

mITT – Modified Intention to Treat

# PDAC – Strong efficacy in patients with high tumor IL1RAP level



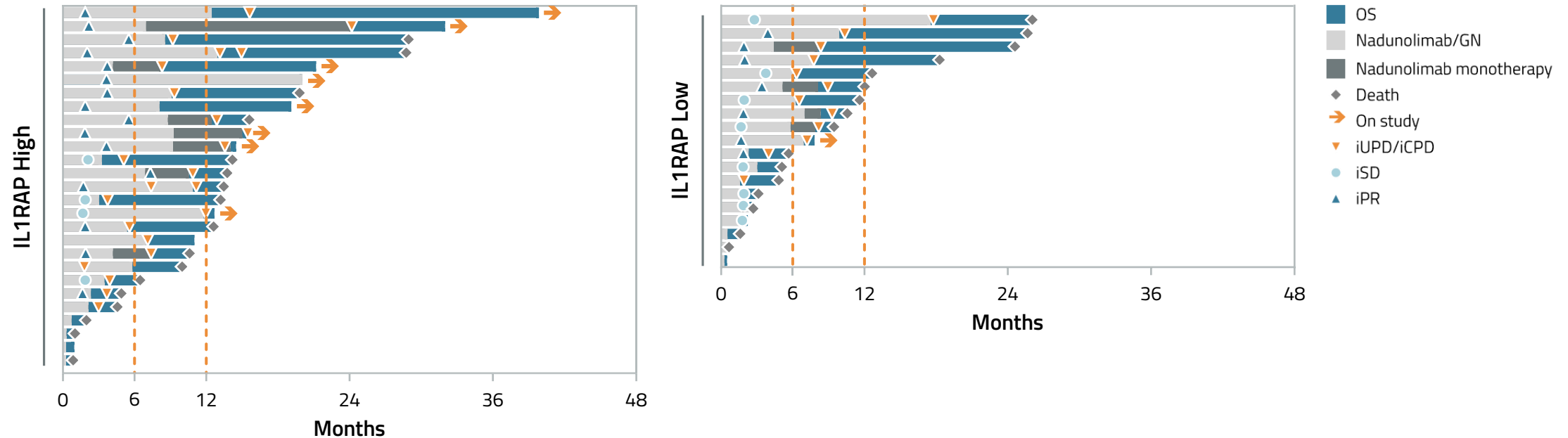
## Efficacy analysis for IL1RAP High (n=27) vs IL1RAP Low (n=19) PDAC patients:

- Significantly prolonged OS in ILRAP High vs IL1RAP Low patients (14.2 vs 10.6 mo;  $p=0.017$ )
- Deeper and more durable responses in IL1RAP High subgroup: 11 patients had 50% or more tumor size decrease

NEW DATA SUPPORT ONGOING DEVELOPMENT AND EXPLORATION OF NEW OPPORTUNITIES

# PDAC – Patients with high tumor IL1RAP level have the strongest benefit of nadunolimab combination therapy

Treatment course for each individual patient in the IL1RAP High and IL1RAP Low subgroups



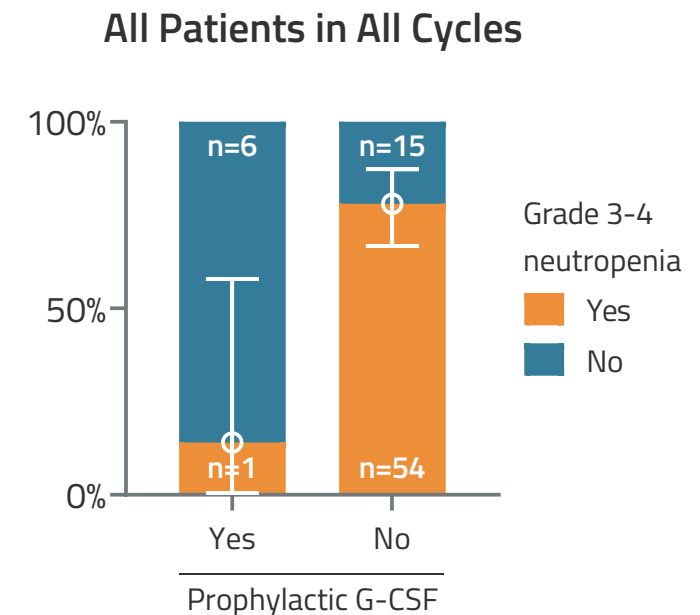
**IL1RAP HIGH PATIENTS STAY LONGER ON THERAPY AHEAD OF PROGRESSION AND DEATH**

iCPD – Confirmed Progressive Disease; iUPD – Unconfirmed Progressive Disease; iSD – Stable Disease; iPR – Partial Response (all according to iRECIST)  
GN – Gemcitabine/Nab-Paclitaxel (Abraxane)

# PDAC – Safety profile is manageable and supports MOA

- Neutropenia manageable through G-CSF prophylaxis
  - In 7 patients given G-CSF prophylaxis, only 1 developed grade 3-4 neutropenia
- Only 1 % peripheral neuropathy grade 3-4 observed (17% in historical controls)

Grade 3 or higher adverse events	Gem/Abraxane Von Hoff, 2013 (n=421)	Nadunolimab+Gem/Abraxane CANFOUR (n=76)
Neutropenia	38%	65%
Leukopenia	31%	24%
Thrombocytopenia	13%	15%
Febrile neutropenia	3%	13%
Anemia	13%	13%
Fatigue	17%	8%
Diarrhea	6%	3%
Peripheral neuropathy	17%	1%



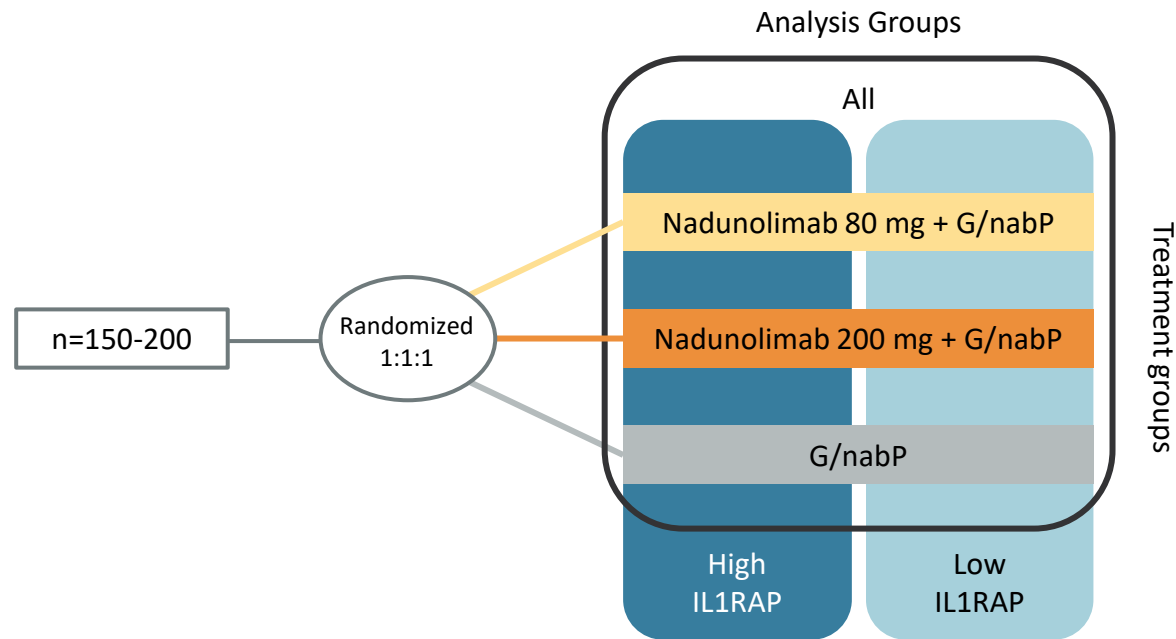
**G-CSF PROPHYLAXIS IMPLEMENTED IN FUTURE TRIALS;  
POTENTIAL REDUCTIONS OF SOME SIDE EFFECTS TO BE DOCUMENTED IN RANDOMIZED TRIALS**

Median duration of treatment 5.5 mo (ref 3.9 mo); most common reasons for termination: gastrointestinal events or general health deterioration  
No patients discontinued due to neutropenia



# PDAC – Phase IIb study design

Open-label, randomized, controlled, non-comparative, 3-arm study evaluating 2 dose levels of nadunolimab + gemcitabine/nab-paclitaxel with gemcitabine/nab-paclitaxel as control:



## Timelines:

- Regulatory submission H2 2023
- FPI early 2024; top-line results 2025

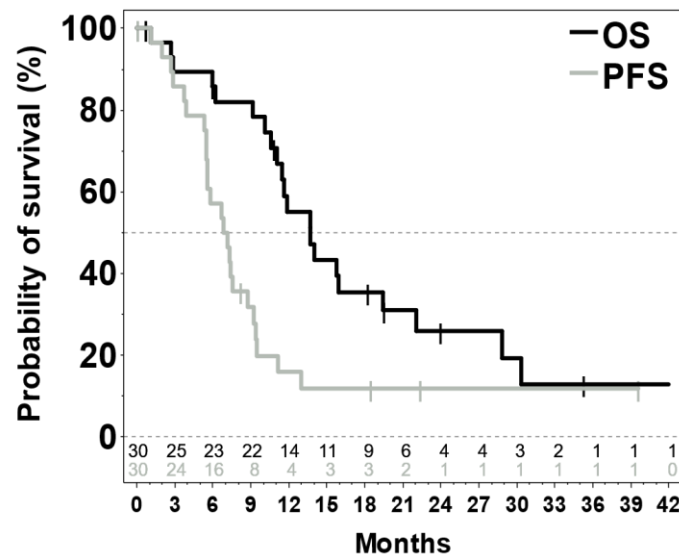
## Geography:

- USA and Europe

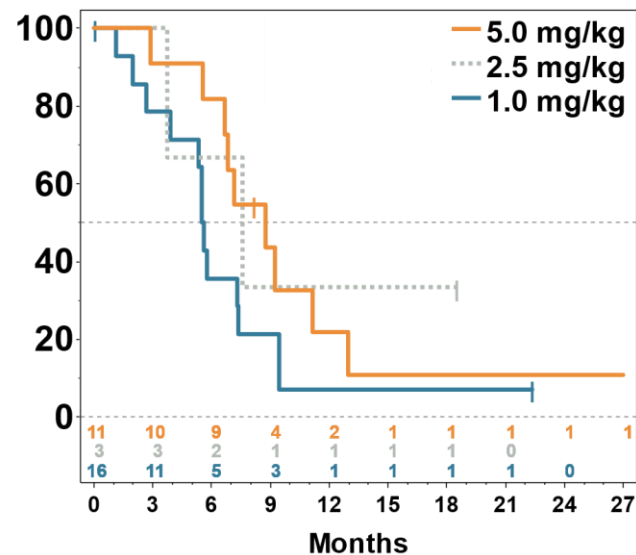
PHASE IIB TRIAL TO VALIDATE STRONG SIGNAL OF ACTIVITY IN IL1RAP HIGH PATIENTS

# NSCLC – Promising efficacy of nadunolimab combination therapy

OS and PFS for mITT patients



PFS by dose level



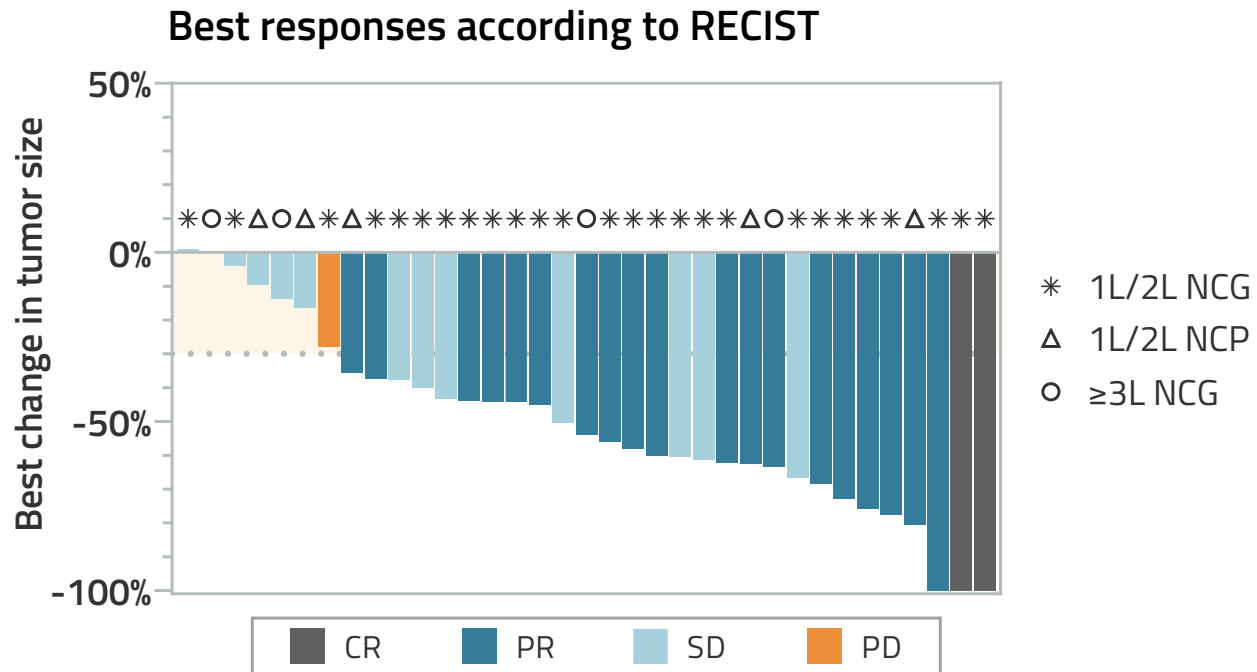
## Nadunolimab combination with Gem/Cis in 1<sup>st</sup>/2<sup>nd</sup> line NSCLC (n=30):

- ORR 53%, 13.7 mo median OS, 7.0 mo median PFS (dose-response trend for PFS)
- Generally well tolerated; neutropenia frequency higher than expected for chemo (managed by dose reductions or G-CSF)

Data presented at ASCO 2023

STRONG EFFICACY OF NADUNOLIMAB IN COMBINATION WITH GEM/CIS IN 1L/2L NSCLC

# NSCLC – Promising efficacy of nadunolimab combination therapy



## High ORR to nadunolimab and platinum doublets in different lines of therapy:

- Gem/Cis 1<sup>st</sup>/2<sup>nd</sup> line: ORR 53% (n=30)
- Carbo/Pemtrex 1<sup>st</sup>/2<sup>nd</sup> line: ORR 60% (n=5)
- Gem/Cis ≥3<sup>rd</sup> line: ORR 50% (n=4)

Data presented at ASCO 2023

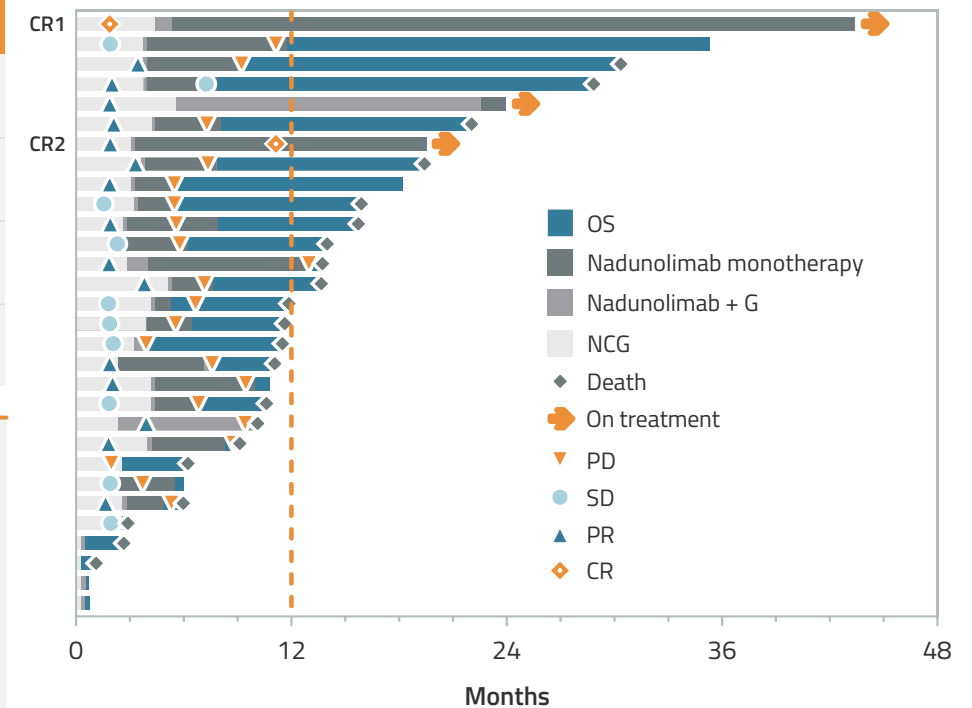
**CONSISTENTLY HIGH RESPONSE RATES WITH NADUNOLIMAB AND PLATINUM DOUBLETS**

# NSCLC – Long-term benefit with strong signal in non-squamous subtype

	All (n=30)	Historical data <sup>1,2</sup>	Non-squamous (n=16)	Non-squamous, historical data <sup>3</sup>
Median OS	<b>13.7 mo</b>	10.3 mo	<b>15.9 mo</b>	11.3 mo
Median PFS	<b>7.0 mo</b>	5.1 mo	<b>7.3 mo</b>	4.9 mo
ORR	<b>53%</b>	22-28 %	<b>56%</b>	19%
Complete response	<b>6.7% (n=2)</b>	<1%	<b>12.5% (n=2)</b>	<1%

- Strongest efficacy in 16 non-squamous patients
- Long-term benefit of nadunolimab combination therapy, including two complete responses

Treatment course for each individual patient



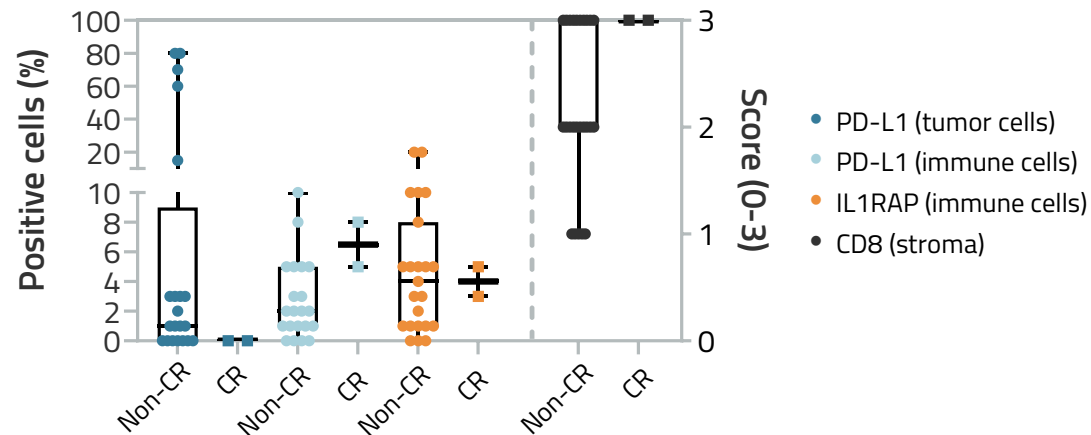
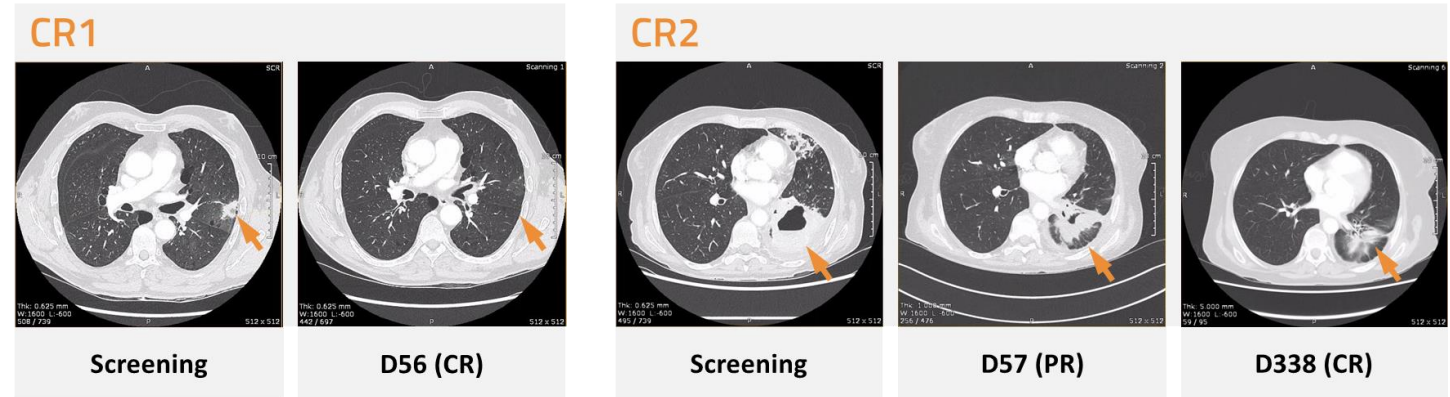
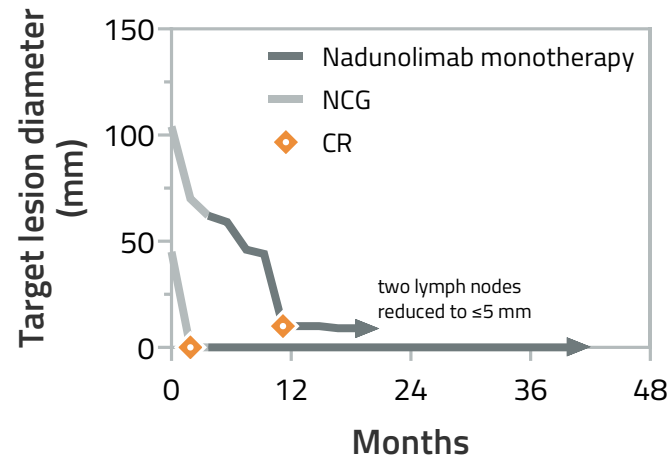
Data presented at ASCO 2023

**NADUNOLIMAB COMBINATION THERAPY COMPARES VERY FAVORABLY TO HISTORICAL DATA FOR CHEMOTHERAPY ALONE**

<sup>1</sup> Schiller et al, N Engl J Med 2002; <sup>2</sup> Scagliotti et al, J Clin Oncol 2008; <sup>3</sup> Gandhi et al, N Engl J Med 2018

PD – Progressive Disease; SD – Stable Disease; PR – Partial Response; CR – Complete Response; NCG – Nadunolimab/Cisplatin/Gemcitabine

# NSCLC – Complete responders with distinct biomarker profile



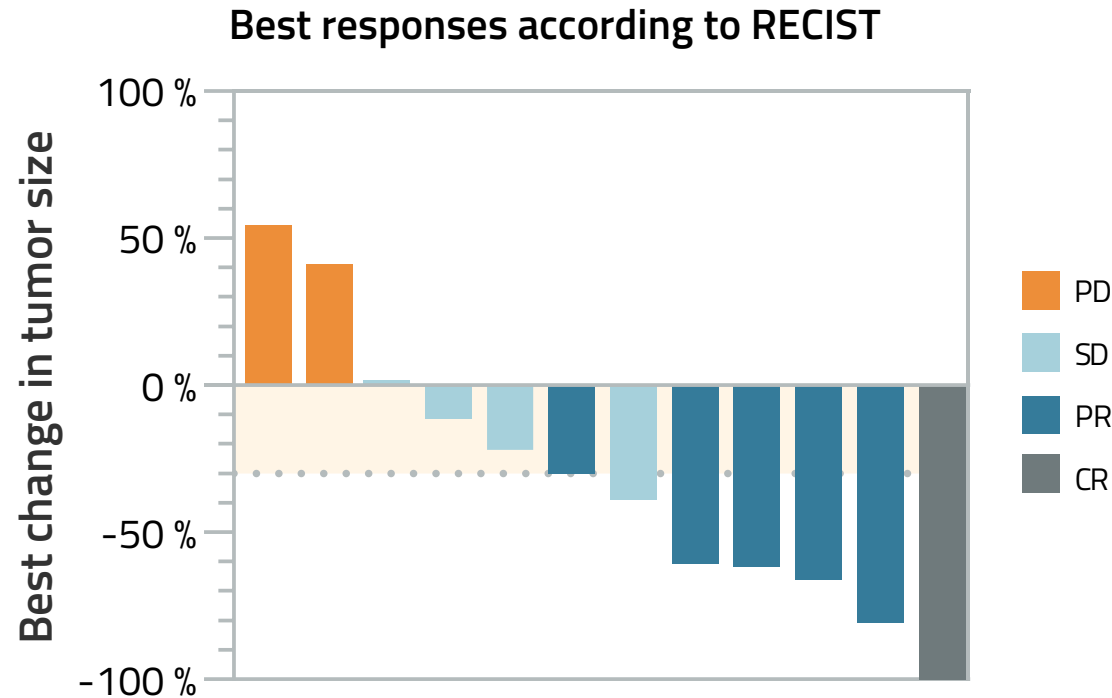
- One CR maintained over 3 yrs (CR1), second CR achieved on nadunolimab monotherapy approx. 9 mo post-chemo (CR2)
- Both non-squamous, progressed on pembro, with PD-L1<sup>-</sup> tumor cells, PD-L1<sup>+</sup> immune cells in tumor

Data presented at ASCO 2023

**SIGNAL OF NADUNOLIMAB MONOTHERAPY ACTIVITY RESULTING IN COMPLETE RESPONSE**



# TNBC – Promising early safety and efficacy



## Nadunolimab combination with Gem/Carbo in 1<sup>st</sup>/2<sup>nd</sup> line metastatic TNBC:

### 15 patients enrolled in the dose-escalation phase

- Acceptable safety profile (G-CSF given prophylactically to control neutropenia)
- 12 patients treated long enough for initial efficacy evaluation:
  - **Preliminary ORR: 50%** (1 CR, 5 PR, 4 SD, 2 PD)
- Proceeds to randomized phase including up to 98 additional patients (n=49 per arm)
- Interim futility analysis planned for Q4 2023

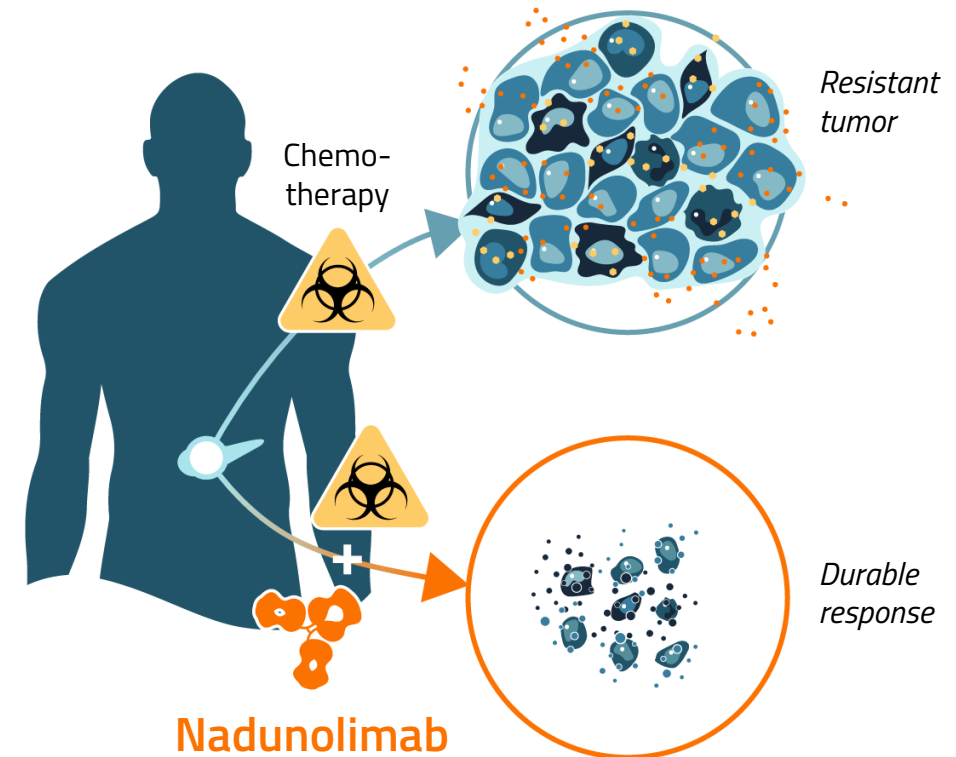
RESPONSE RATE OF NADUNOLIMAB COMBINATION THERAPY WELL ABOVE HISTORICAL DATA FOR CHEMOTHERAPY ONLY<sup>1</sup>

<sup>1</sup> O'Shaughnessy et al, J Clin Oncol 2014

PD – Progressive Disease; SD – Stable Disease; PR – Partial Response; CR – Complete Response

# Key messages

- Most chemotherapies induce chemoresistance already after a few months of therapy. Chemotherapy can upregulate both IL-1 $\alpha$  and IL-1 $\beta$ .
- Unlike other IL-1 blocking compounds, nadunolimab blocks both IL-1 $\alpha$  and IL-1 $\beta$  signalling and improves chemotherapy efficacy and tolerability in preclinical models.
- Current results are in sharp contrast to canakinumab data.
- Clinical results strongly support potential unique first-in-class opportunities in PDAC, NSCLC and TNBC.



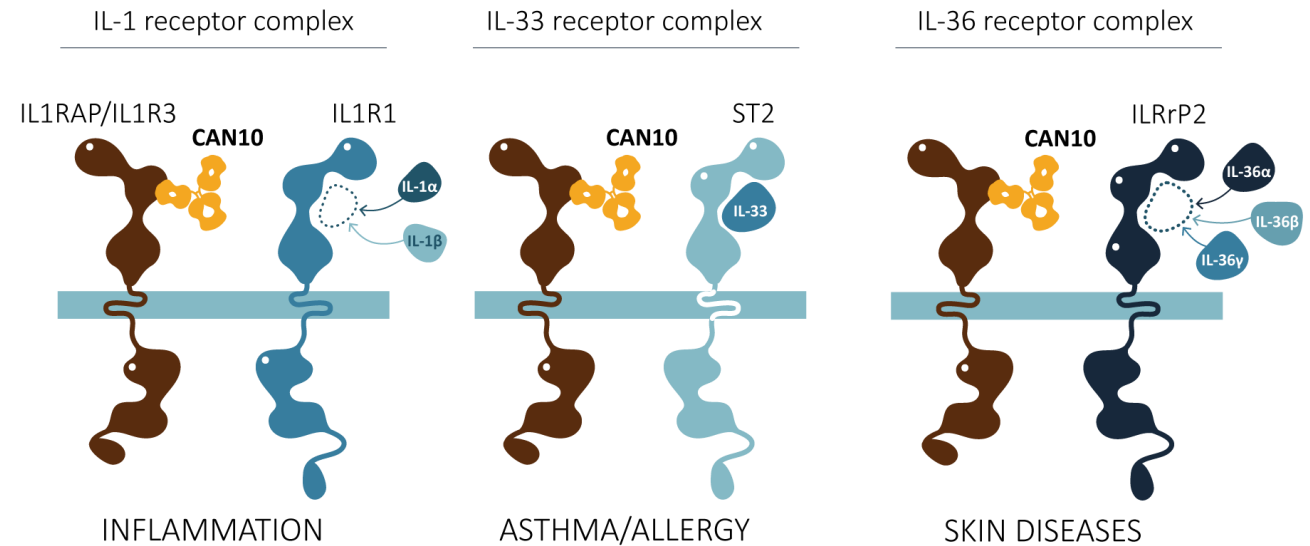
PROMISING EFFICACY OF NADUNOLIMAB WITH CHEMOTHERAPY – CURRENT FOCUS ON RANDOMIZED CLINICAL TRIALS

A microscopic image showing several cells with a complex, textured surface. The image is overlaid with a semi-transparent blue layer, which serves as a background for the text. The cells are primarily in the upper half of the frame, with one large cell on the left and another on the right. The blue overlay covers the entire image, creating a uniform background for the text.

CAN10 – OPPORTUNITY IN AUTOIMMUNE/INFLAMMATORY DISEASE

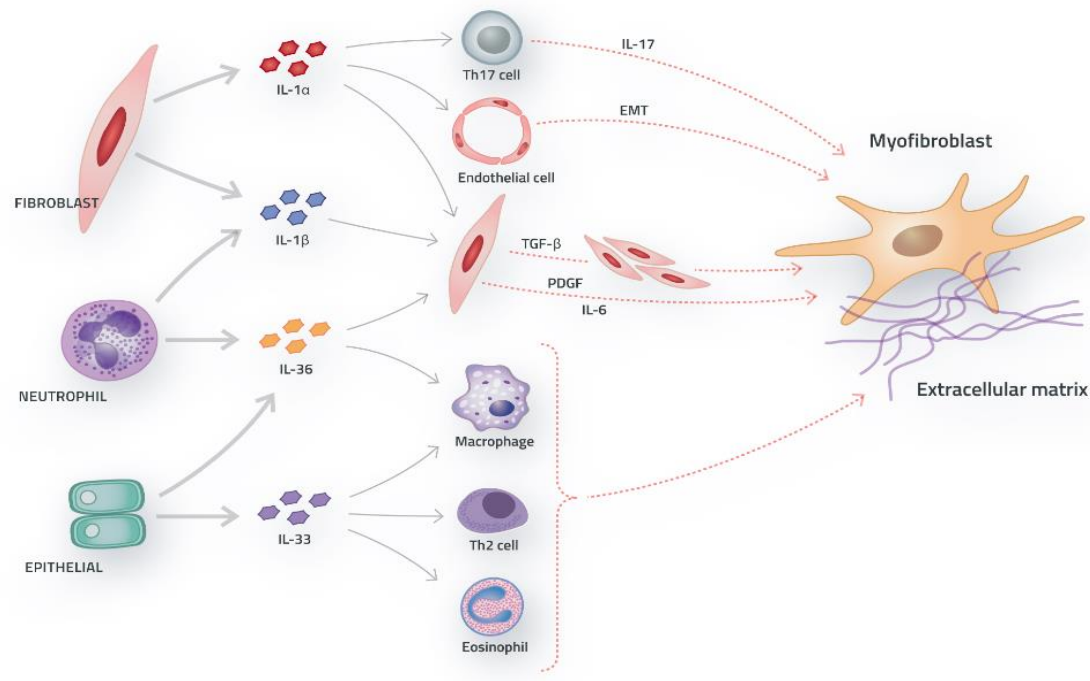
# CAN10 – New asset within autoimmunity/inflammation

- IL1RAP-binding antibody potentially blocking IL-1, IL-33 and IL-36, without ADCC
- Unique anti-inflammatory activity observed in different mouse models (myocarditis, systemic sclerosis, psoriasis, inflammation)
- Development focusing on systemic sclerosis and myocarditis, diseases involving multiple IL-1 family cytokines
- Clinical trial in healthy volunteers to start mid-2023

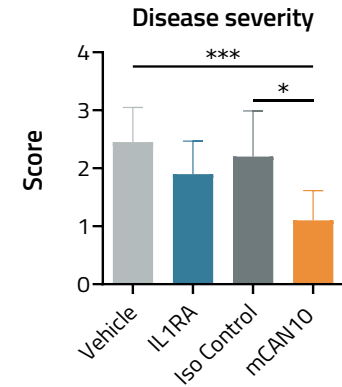


UNIQUE OPPORTUNITY FOR CAN10 IDENTIFIED IN LIFE-THREATENING DISEASES

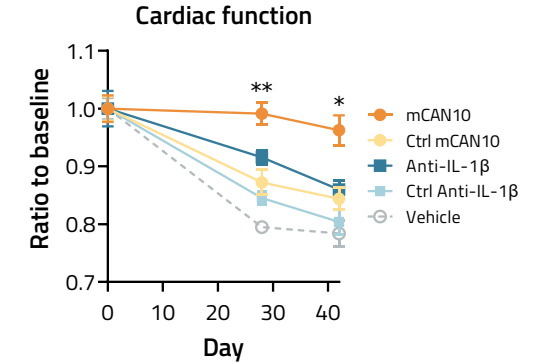
# CAN10 – Promising effects in several preclinical disease models



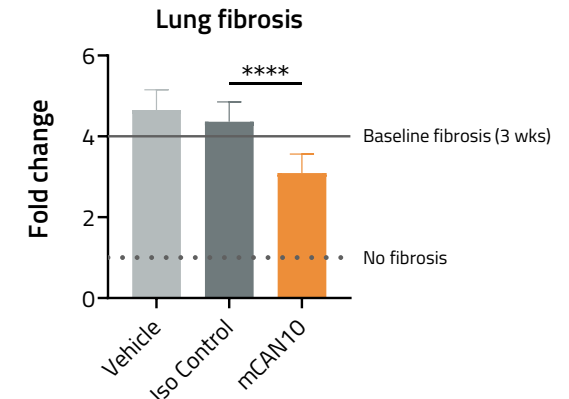
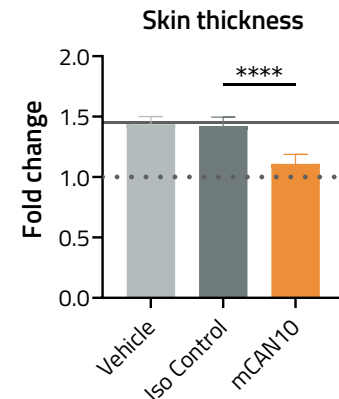
## Viral myocarditis



## Autoimmune myocarditis



## Systemic sclerosis



CAN10 SHOWS POTENTIAL IN SEVERAL AUTOIMMUNE/INFLAMMATORY DISEASES WITH HIGH MEDICAL NEED



# CAN10 – Project status

## GLP toxicity study – Completed

- CAN10 given i.v. once weekly for six weeks at doses up to 50 mg/kg or s.c. at 5 mg/kg
- No adverse findings related to CAN10 at/above clinically relevant (pharmacologically active) dose levels

## Clinical Phase I study – Study start planned for mid-2023

- Clinical trial application submitted in April 2023
- Phase I plan in healthy volunteers (SAD) followed by psoriasis patients (MAD)



## MILESTONES, FINANCIALS & SUMMARY

# Upcoming milestones

## Nadunolimab

PDAC	NSCLC	TNBC	CAN10	Additional milestones
<ul style="list-style-type: none"><li>• Start Phase IIb trial in 150-200 patients with submission in H2 2023</li><li>• Phase IIb top-line data in 2025</li></ul>	<ul style="list-style-type: none"><li>• Presented updated efficacy and biomarker data from CANFOUR at ASCO Q2 2023</li></ul>	<ul style="list-style-type: none"><li>• Randomized Phase II (TRIFOUR) – interim futility analysis in Q4 2023</li><li>• Present safety and efficacy data from Phase I in H2 2023</li></ul>	<ul style="list-style-type: none"><li>• Treatment of first subject in Phase I clinical trial mid-2023</li></ul>	<ul style="list-style-type: none"><li>• New clinical data presented from CIRIFOUR, CAPAFour and CESTAFOUR trials</li><li>• New preclinical and translational results</li></ul>

# Solid financial position with strong shareholder support

- Cash and cash equivalents 353M SEK (~\$33M) at end of Q1 2023
- Runway until mid-2024
- Market cap appr 0.8B SEK, 75M USD May 26, 2023

## Current owners (Mar 31, 2023)

4th AP fund	8.8%
Alecta	7.3%
1st AP fund	6.3%
Avanza Pension	5.1%
Six Sis AG	4.7%
Swedbank Robur Funds	3.8%
BNY Mellon	2.5%
Nordnet Pensionförs.	1.4%
Handelsbanken fonder	1.2%
Brushamn Invest	1.2%
Other	57.6%

# Cantargia – Investment highlights



## NOVEL IL1RAP ANTIBODIES, POTENTIAL TO ADDRESS CANCER & INFLAMMATORY DISEASE

- IL1RAP elevated in most solid and liquid tumors
- Potential to break down resistance to cancer treatment, enabled by unique dual action approach – nadunolimab (CAN04)
- Additional key target for inflammatory diseases – CAN10



## DEVELOPING THERAPIES IN AREAS OF HIGH UNMET NEED; WITH UPCOMING CATALYSTS

- Strong clinical interim results in PDAC and NSCLC, and promising initial results in TNBC; >200 patients treated
- Randomized trial ongoing in TNBC
- Phase IIb trial in preparation for PDAC



## CORPORATE STRENGTH DRIVING INNOVATION

- Solid cash position with runway to mid-2024 (353M SEK cash & equivalents at end of Q1 2023)
- Robust patent portfolio: antibody target in oncology (2032), nadunolimab (2035) and CAN10 (2041)