

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

Corporate Presentation Feb 2024 NASDAQ STOCKHOLM MAIN LIST (CANTA.ST)

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## Cantargia – Investment highlights

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

- IL1RAP elevated in most solid and liquid tumors
- IL1RAP signaling drives several autoimmune and inflammatory diseases

### NADUNOLIMAB: CLEAR ACTIVITY SIGNALS IN CANCER THERAPY WITH UPCOMING CATALYSTS

- Strong clinical interim results in PDAC and NSCLC, and promising initial results in TNBC; >250 patients treated
- Randomized Phase II trial ongoing in TNBC (initial data late 2024); Phase IIb trial in preparation in PDAC (top-line data 2025)



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#### **CAN10: OPPORTUNITY IN AUTOIMMUNITY/INFLAMMATION**

- Pronounced activity in models of systemic sclerosis, myocarditis, psoriasis, atherosclerosis and inflammation
- Phase I clinical trial ongoing, initial results show good safety and receptor occupancy. New data Q2 2024



#### **CORPORATE STRENGTH DRIVING INNOVATION**

- Solid cash position with runway into 2025 (195MSEK (19 MUSD) cash & equivalents at Q4 2023)
- Robust patent portfolio: IL1RAP antibody target in oncology (2032), nadunolimab (2035) and CAN10 (2041)



### Current pipeline

Project	Disease	Type of treatment	Discovery phase	Preclinical phase	Clinical phase l	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





### 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

ADCC – Antibody-Dependent Cellular Cytotoxicity; NK – Natural Killer; TIR – Toll-Interleukin-1 Receptor



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



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

- 33% response rate with long OS and iPFS
  - Additional 5 (7%) patients had on-treatment benefit beyond progression
- Promising OS (13.2 mo), iPFS (7.2 mo) and DCR (71%); 2 patients still on treatment

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

Benchmark Gem/Abraxane: OS 8.5 mo, PFS 5.3 mo, ORR 23%, DCR 48% (Von Hoff et al, N Engl J Med 2013); OS 9.2 mo, PFS 5.6 mo, ORR 36%, DCR 62%, (NAPOLI-3, ASCO GI 2022) iCPD – Confirmed Progressive Disease; iUPD – Unconfirmed Progressive Disease; iSD – Stable Disease; iPR – Partial Response (all according to iRECIST)



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



Efficacy analysis for IL1RAP High (n=29) vs IL1RAP Low (n=20) PDAC patients (1<sup>st</sup> line, combination with Gem/Abraxane):

- → Significantly prolonged OS in ILRAP High vs IL1RAP Low patients (14.2 vs 10.6 mo; p=0.026)
- → Deeper and more durable responses in IL1RAP High subgroup: 11 patients had 50% or more tumor size decrease

NEW DATA IN IL1RAP HIGH PATIENTS SUPPORT ONGOING DEVELOPMENT AND EXPLORATION OF NEW OPPORTUNITIES



### Nadunolimab PDAC milestone targets

	mid-2024	H1 2025	H2 2025	H1 2026	H1 2027	H1 2028	H2 2028
	Start PANFOUR study	PANFOUR enrolment completed	PANFOUR study results				
		FDA meeting	FDA EOP2 meeting				
		Phase III study	preparation	Start Phase III study	Phase III enrolment completed	Phase III study results	
						Potential	Potential US
<ul> <li>Confirm CANFOUR high IL1RAP results and accelerated path to market</li> <li>Interim efficacy &amp; subgroup analysis</li> </ul>					BLA / MAA submission	market launch	
<ul> <li>Discuss and agree dose and data driven patient selection strategy for Phase III / BLA</li> <li>IL1RAP or KRAS or serum BM patient selection</li> </ul>							

PANFOUR study design address FDA Project Optimus and Frontrunner guidelines and de-risks development with interim snapshot to evaluate efficacy, safety and biomarker subgroup analysis

### NSCLC – Complete responders with distinct biomarker profile



SIGNAL OF NADUNOLIMAB MONOTHERAPY ACTIVITY RESULTING IN COMPLETE RESPONSE FURTHER BIOMARKER ANALYSES ONGOING FOR FUTURE DEVELOPMENT STRATEGY

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## TNBC – Promising early safety and efficacy



Benchmark Gem/Carbo: OS 11.1 mo, PFS 4.1 mo, ORR 30% (O'Shaughnessy et al, J Clin Oncol 2014)

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

#### 15 patients enrolled in the doseescalation phase:

- Preliminary ORR: 60% (1 CR, 8 PR, 4 SD, 2 PD)
- → Preliminary median OS 12.3 mo, median PFS 6.6 mo
- Acceptable safety profile
   (G-CSF given prophylactically to control neutropenia)
- → Randomized phase II ongoing

#### RESPONSE RATE OF NADUNOLIMAB COMBINATION THERAPY WELL ABOVE HISTORICAL DATA FOR CHEMOTHERAPY ONLY



## Nadunolimab and alleviation of neuropathy

- Chemotherapy induce neuropathy by several pathways including IL-1 (neuroinflammation)
- Nadunolimab, phase 2 data in PDAC with Gem/nabP
  - → lower Grade 3-4 peripheral neuropathy than expected from historical controls (1% vs 17%).
- → Further analysis from this trial on grade 1-2 neuropathy strengthens finding
- → Counteraction of chemotherapy induced neuropathy in animal models
- ADC payloads induce IL-1 system contributing to tumor progression

Detailed data to be presented at upcoming scientific conferences



IN ADDITION TO PROMISING EFFICACY NADUNOLIMAB COULD CONTRIBUTE TO SAFER COMBINATION THERAPIES





### CAN10 – OPPORTUNITY IN AUTOIMMUNE/INFLAMMATORY DISEASE

## CAN10 – New clinical asset in autoimmunity/inflammation

- → IL1RAP-binding antibody potently 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



#### **UNIQUE OPPORTUNITY FOR CAN10 IDENTIFIED IN LIFE-THREATENING DISEASES**



# IL1RAP and the IL-1/33/36 pathways are upregulated in SSc patient skin

2 publicly available human SSc cohorts show differential expression of IL1RAP and associated genes in SSc skin



14 SSc vs. 11 healthy Agilent 2-channel Microarray

Mahoney et al. 2015 GSE59787 Skaug et al. Ann Rheum Dis 2020. GSE130955



# Systemic sclerosis: mCAN10 inhibits bleomycin-induced skin fibrosis



# CAN10 first-in-human study

- Combined SAD/MAD protocol
- IV administration in healthy volunteers (SAD)
- SC administration in subjects with mild to moderate plaque psoriasis (MAD)
- Safety and PK
- Building value by adding additional PD analyses
  - Receptor occupancy
  - Ex vivo inhibition assay (proteomics)
  - Psoriasis severity scoring
  - Skin biopsy
- Phase I SAD ongoing, MAD planned to start Q3 2024
- Preparations for phase II clinical study ongoing





### MILESTONES & INVESTMENT HIGHLIGHTS

### Upcoming milestones

### Nadunolimab

PDAC	NSCLC	TNBC	CAN10	Additional milestones
<ul> <li>Start of Phase IIb trial in 150-200 patients H1 2024</li> <li>Phase IIb top-line data in 2025</li> </ul>	<ul> <li>Efficacy/biomarker data from CANFOUR during 2024</li> </ul>	<ul> <li>Full recruitment H2 2024</li> <li>Randomized Phase II top-line data in late 2024</li> </ul>	<ul> <li>Phase I data updates during 2024 (including safety and biomarkers)</li> </ul>	<ul> <li>New clinical data presented from CIRIFOUR, CAPAFOUR and CESTAFOUR trials</li> <li>New preclinical and translational results</li> </ul>

**EXTENSIVE NEWS FLOW EXPECTED DURING 2024** 



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