

### **Presenters**



Göran Forsberg CEO



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VP Cancer Research



Lars Thorsson
VP Clinical Development

### Safe Harbour Statement

The following presentation may include predictions, estimates or other information that might be considered forward-looking. The statements regarding the surrounding world and future circumstances in this presentation reflect Cantargia's current thinking with respect to future events and financial performance. Prospective statements only express the assessments and assumptions the company makes at the time of the presentation. These statements are well-considered, but the audience should note that, as with all prospective assessments, they are associated with risks and uncertainties.

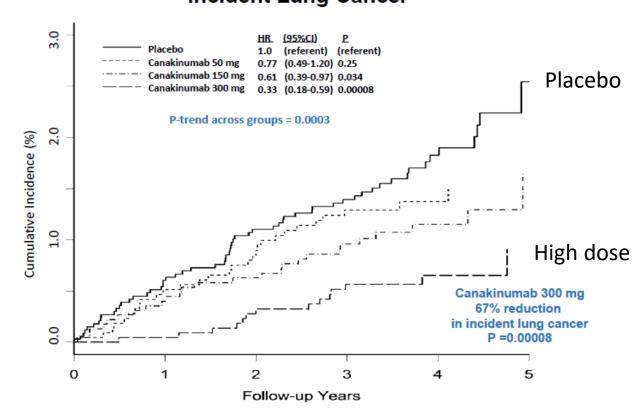


### CANTOS IL-1ß blockade - Recent strong clinical data

#### **CANTOS** trial

- Canakinumab (Novartis)
- 10 061 patients
- Designed to reduce cardiovascular events in patients with previous myocardial infarction
- Reduced lung cancer incidence by 67 % and death by 77 %.
- Novartis has clinically validated the IL-1 pathway
- Cantargia's CAN04 has higher potential than Canakinumab

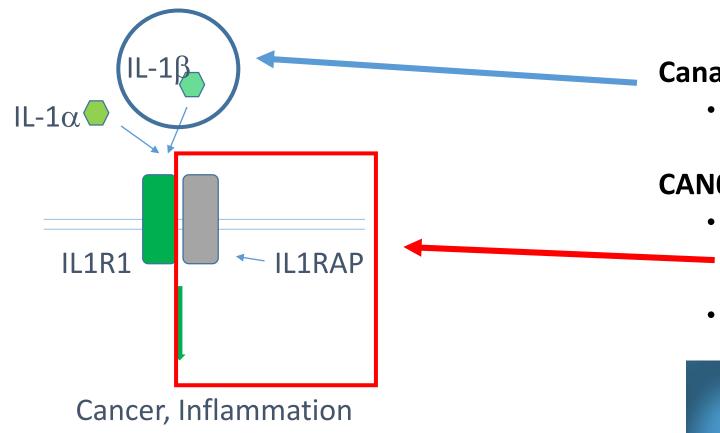
#### CANTOS: Additional Non-Cardiovascular Clinical Benefits Incident Lung Cancer





### CAN04 vs Canakinumab

(IL-6 & IL-8)

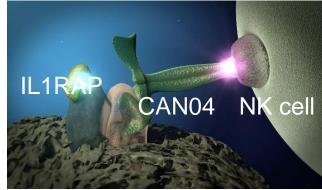


#### Canakinumab

Antibody directed against one of the two IL-1 ligands, IL-1β

#### **CAN04:**

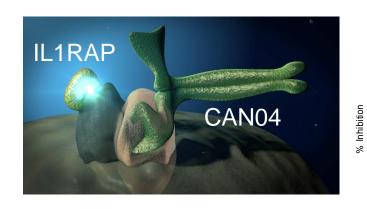
- Binds the common signaling receptor and counteracts both ligands
- Induce killing via the immune system (ADCC)



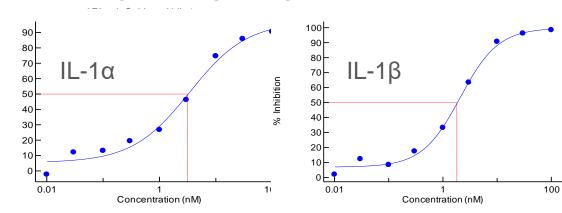
# **CANTOS** additional findings

CANCER decreased risk of death with treatment (high dose)				
Lung cancer	77 %	P=0.0002		
Non-lung cancer	37 %	P=0.06		
Decreased incidence of inflammatory disease (all doses)				
Arthritis	32%	p<0.0001		
Ostheoartritis	28%	P=0.0005		
Gout	53%	p<0.0001		
Biomarker levels (reduction)				
CRP	26-41%	P<0.0001		
IL-6	25-43%	P<0.001		

### CAN04 Dual mode of action

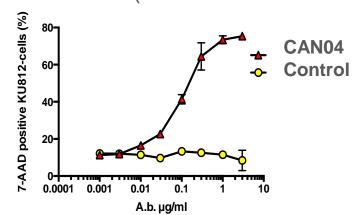


Blocking IL-1 signaling





ADCC (stimulate immune cells to kill cancer cells)





## Non-small cell lung cancer

#### **USA** estimates:

Lung cancer: 222500 new cases, 155870 deaths.

NSCLC, ~85 % of lung cancers

Subgroups (adenocarcinoma, squamous, large cell etc)

#### Inflammation of importance of tumor progression

IL-1 pathway externally validated

#### Standard therapies:

Keytruda (antibody against PD-1)

Chemotherapy

Targeted therapy (genetic form)

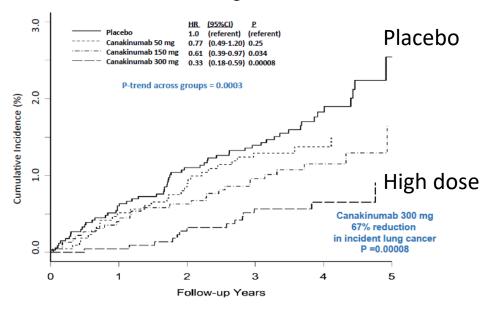
#### Canakinumab phase III trials in preparation (ref Novartis)

Adjuvant (after surgery)

First line with PD-1 antibody

Second line

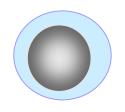
#### CANTOS: Additional Non-Cardiovascular Clinical Benefits Incident Lung Cancer



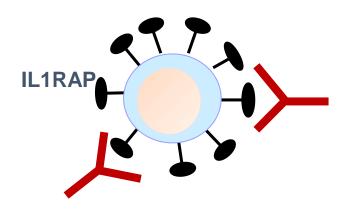


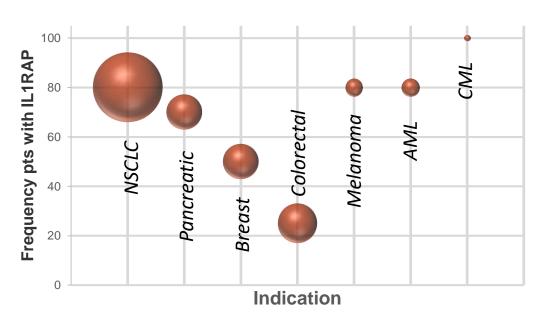
### Medical need and IL1RAP

#### Normal cell



#### Cancer cell





- Based on in house data, external data, medical need and market size, NSCLC and pancreatic cancer are primary indications.
- Biomarker studies ongoing, to identify patients most likely to respond
- Low levels of IL1RAP in normal tissue (analyzed following FDA and EMA guidelines)



### Cantargia at a glance

- Specialized in antibody therapy/immunology, with initial focus on oncology
- Granted IP around therapeutic target and drug candidates
- Lead antibody CAN04 in clinical development
- Strong management team with proven track record in clinical development and business development
- IPO March 2015 (Nasdaq First North, Stockholm), preparations for listing on main market ongoing
- More than 3000 shareholders
- Based in Lund, Sweden
- New share issue of 232 MSEK Dec 15 2017

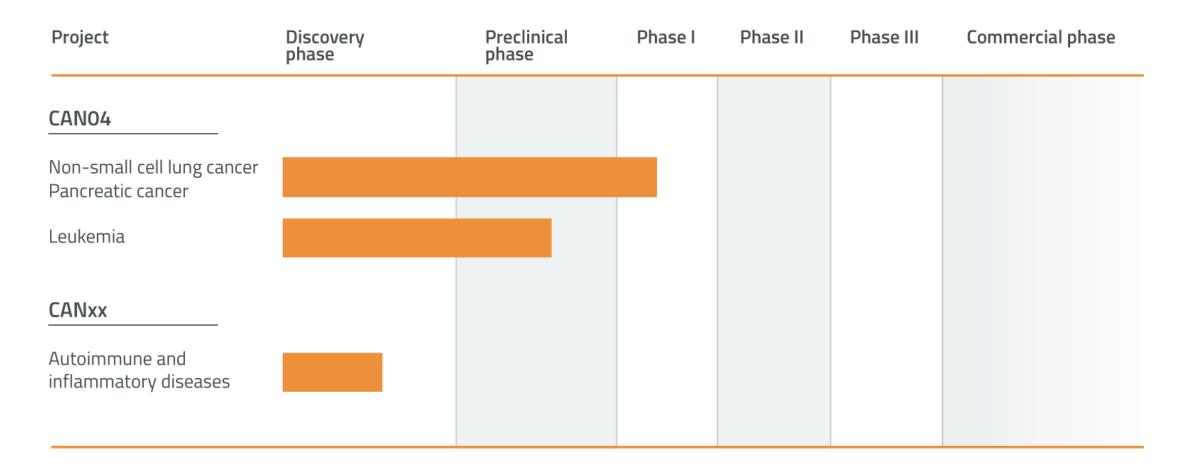
#### Financial highlights

- Share price: 6.80 SEK (0.84 USD), Feb 9, 2018
- Market cap: 450 MSEK (55.6 MUSD), Feb 9, 2018
- Cash: 52.4 MSEK (6.7 MUSD), Sep 30 2017 (excl new financing)

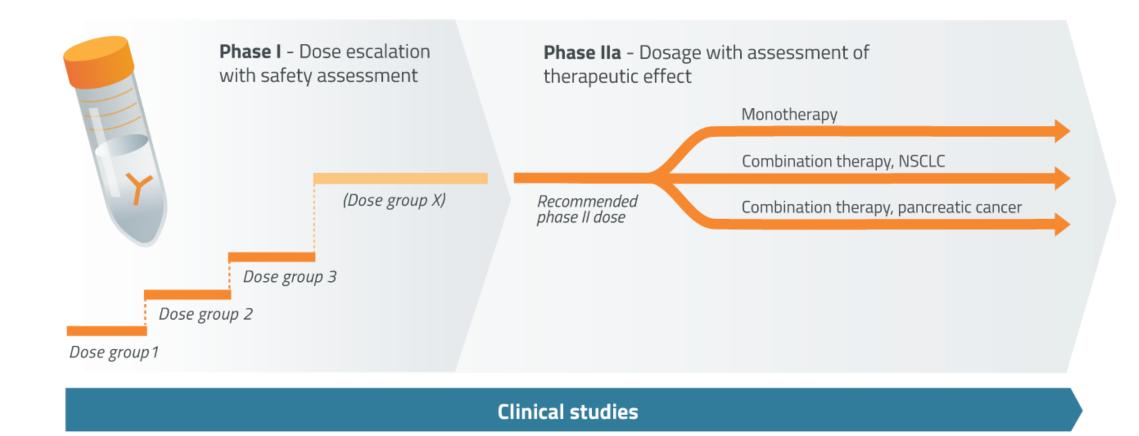
Current owners (Jan 19, 2018)			
Sunstone	9.0%		
1st AP fund	6.9%		
LU Bio	6.1%		
Avanza Pension	5.6%		
4th AP fund	4.2%		
2nd AP fund	3.3%		
SEB S.A. clients	3.0%		
Mats Invest AB	1.8%		
Tibia konsult	1.7%		
Brushamn Invest	1.6%		
SHB Pharm Fund	1.5%		
Others	55.2%		



# Cantargia pipeline



### CANO4 – CANFOUR clinical trial



@antargia

Details on www.clinicaltrials.gov

# Preclinical development



# Tumor inflammation – key to cancer progression

**Enablers** 

Genomic instability and mutation (2000)



Tumor-promoting inflammation (2011)

Deregulating cellular energetics

Sustaining proliferative signaling

**Evading growth suppressors** 

Resisting cell death

**Enabling replicative immortality** 

**Inducing angiogenesis** 

Activating invasion and metastasis

**Avoiding immune destruction** 

The inflammatory cytokine IL-1 – Well established role in cancer progression:

Cancer hallmarks

#### **Tumor cells**

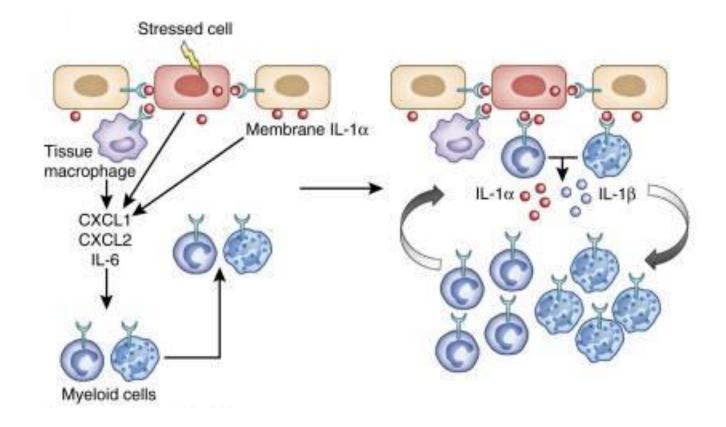
- Signaling/proliferation of cancer cells
- Chemoresistance

#### **Tumor microenvironment**

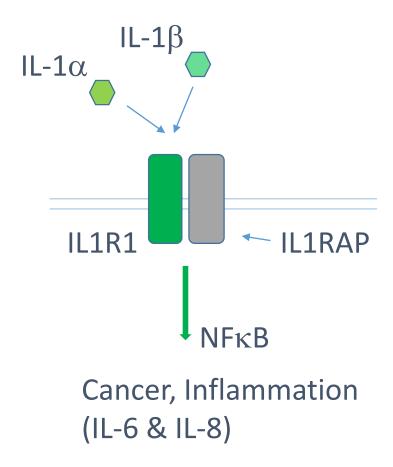
- Metastasis
- Crosstalk between tumor cells and stroma
- Inflammation and local suppression of the immune system



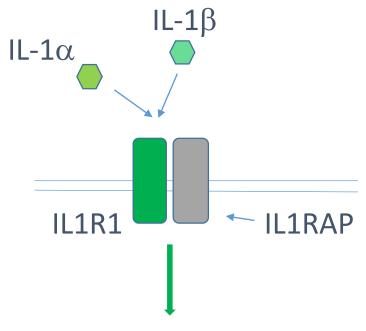
# In sterile inflammation, IL-1 $\alpha$ drives an inflammatory loop that is amplified by IL-1 $\beta$



# IL1RAP is required for Interleukin-1 signaling

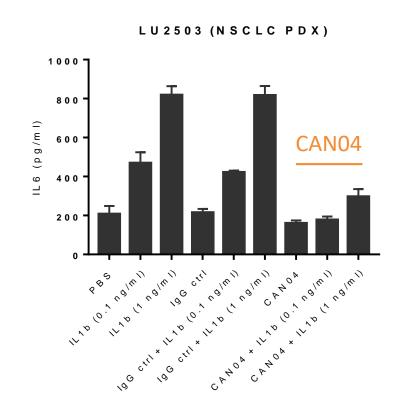


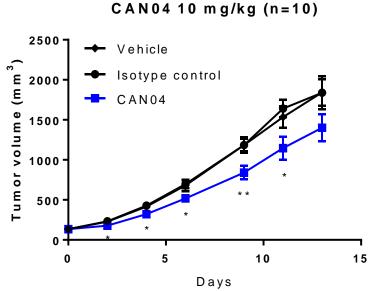
## CANO4 targets IL1RAP to inhibit tumor growth



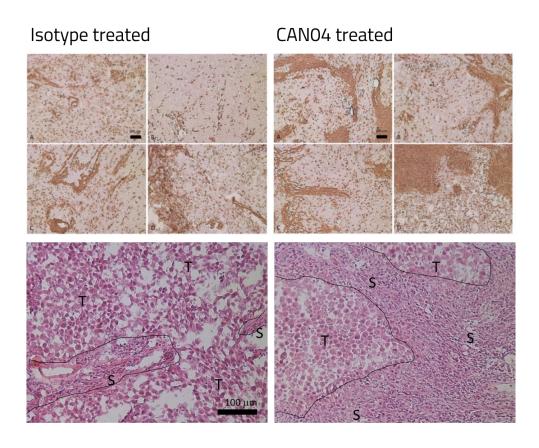
Inflammatory response e.g. IL-6

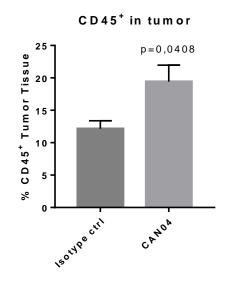
IL-6 is involved in tumor progression





## CANO4 targets IL1RAP to inhibit tumor growth





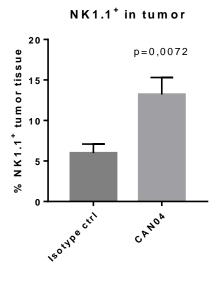


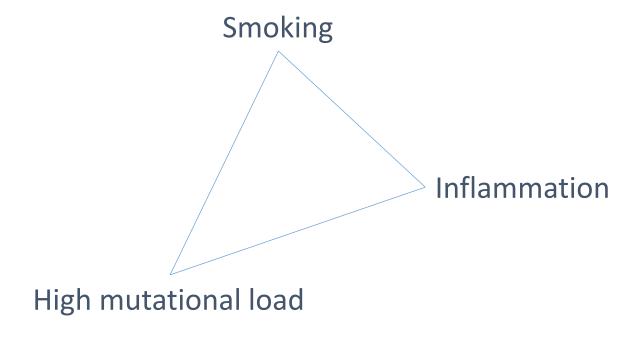
 Image analysis:
 Isotype
 CANO4

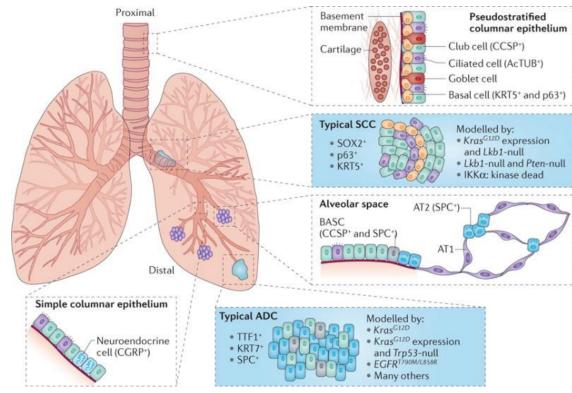
 CD45+
 14.9 %
 24.3%

 Stroma (H/E)
 12 %
 34%



# Non-small cell lung cancer (NSCLC)

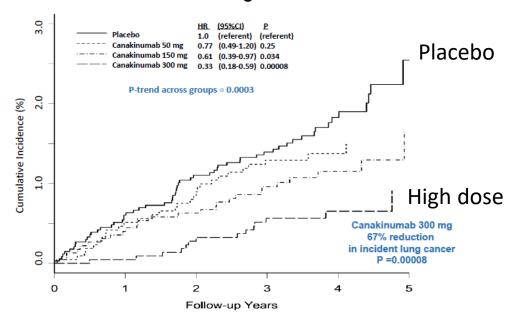


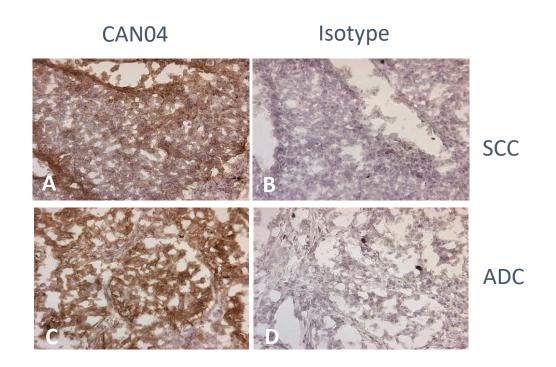


Chen et.al, Non-small-cell lung cancers: a heterogeneous set of diseases, Nat Rev Cancer 2014

# Non-small cell lung cancer (NSCLC)

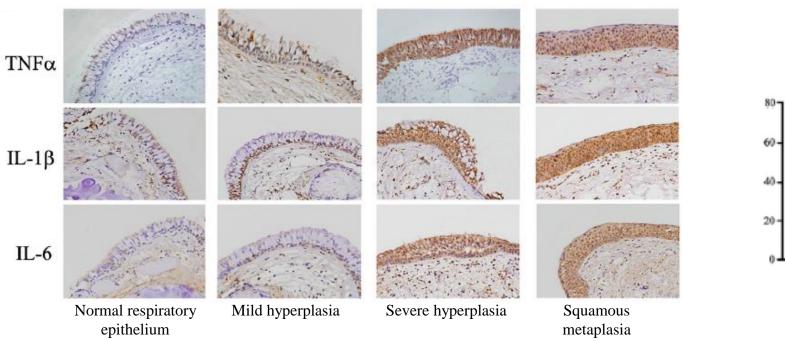
### CANTOS: Additional Non-Cardiovascular Clinical Benefits Incident Lung Cancer





# Non-small cell lung cancer (NSCLC)

Inflammation drives metaplasia and is a hallmark of active lung cancer



IL-1β

80

60

40

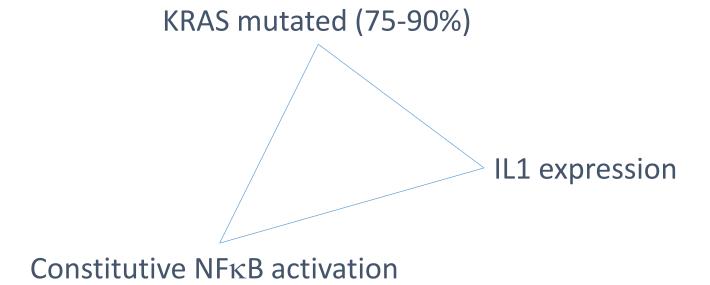
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Air Smoke Smoke Smoke Smoke Smoke Etanercept Anakinra Tocilinanah Etanercept Anakinra Tocilinanah Tocilinanah Tocilinanah Tocilinanah Tocilinanah

Herfs et.al, Proinflammatory Cytokines Induce Bronchial Hyperplasia and Squamous Metaplasia in Smokers, Am J Respir Cell Mol Biol 2012

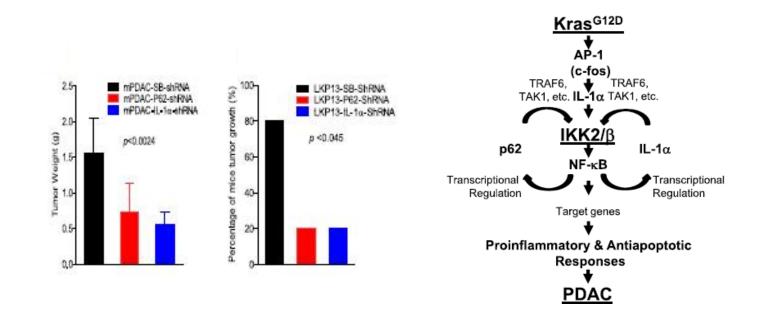
# Pancreatic ductal adenocarcinoma (PDAC)

Propensity to metastasize, resistance to chemo- and radiotherapy, 5-year survival < 6%

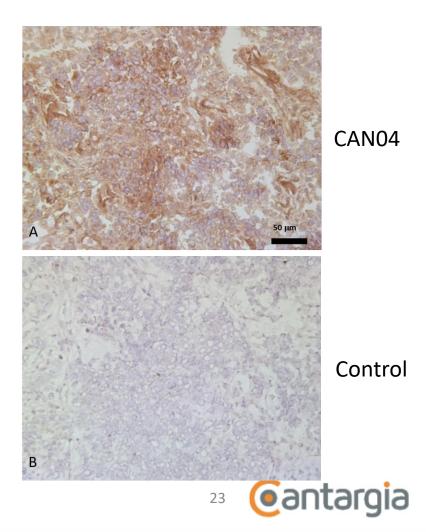


## Pancreatic ductal adenocarcinoma (PDAC)

#### PDAC hallmarks are connected

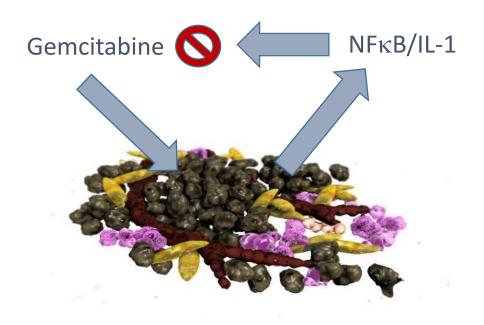


Ling et.al, KRAS<sup>G12D</sup>-Induced IKK2/ $\beta$ /NF- $\kappa$ B Activation by IL-1 $\alpha$  and p62 Feedforward Loops is Required for Development of Pancreatic Ductal Adenocarcinoma, Cancer Cell 2012



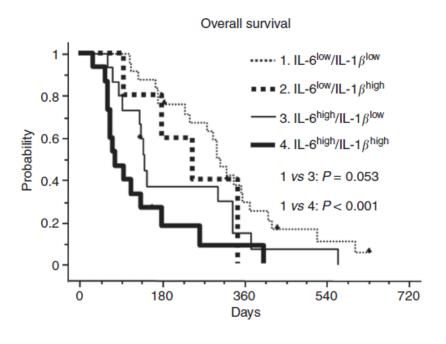
# Pancreatic ductal adenocarcinoma (PDAC)

#### PDAC hallmarks relate to chemoresistance



Zhuang et.al; IL1 Receptor Antagonist Inhibits Pancreatic Cancer Growth by Abrogating NF-kB Activation, Clinical Cancer Res 2016

Zhang et.al; Constitutive IRAK4 Activation Underlies Poor Prognosis and Chemoresistance in Pancreatic Ductal Adenocarcinoma, Clinical Cancer Res 2017



Mitsanuga et.al; Serum levels of IL-6 and IL-1 $\beta$  can predict the efficacy of gemcitabine in patients with advanced pancreatic cancer, Br J Cancer 2013



### Inflammation and metastasis

- Cancer cells (seeds) needs a good soil to form a metastasis
- The IL-1 system (inflammation) can provide such an enabling environment





### Summary

- CAN04 blocks IL-1 signaling and targets IL1RAP-expressing cells for immunemediated killing
- Both non-small cell lung cancer (NSCLC) and pancreatic cancer (PDAC) are tumors strongly coupled to IL-1 mediated tumor inflammation
- IL1RAP is highly expressed in NSCLC and PDAC
- IL-1 is involved in cancer development and mediates resistance to chemotherapy



# Clinical development



### CANFOUR - Press release FIM 2017-10-13

# Cantargia announces first patient treated and completed a three weeks safety evaluation period with immuno-oncology antibody CAN04

Cantargia AB ("Cantargia") today announces that the first patient in the CANFOUR clinical trial has received three cycles of treatment with the antibody CAN04. Thereby, the first patient has formally completed the safety evaluation period according to the clinical protocol. Two additional patients have received therapy with CAN04. No serious adverse events have been recorded. The ongoing clinical trial is a combined dose escalation/dose expansion phase I/IIa trial carried out in patients with non-small cell lung cancer, pancreatic cancer, colorectal cancer or triple negative breast cancer. The CAN04 antibody is targeted against IL1RAP, found in a number of cancer forms.

The first sites in the phase I/lla clinical trial CANFOUR have been initiated and patient recruitment is ongoing. According to the protocol, patients are recruited in groups of three. Following the start of patient recruitment, all three patients in the first group has now each been given at least two infusions of CAN04. The first patient has completed three infusions and has been followed through a safety evaluation period of 21 days. No serious adverse events have been noted and once all three patients have received three infusions and have completed their 21-day safety evaluation period, the next dose group can be recruited

### **CANFOUR**

#### **Coordinating Investigator:**

Professor Ahmad Awada Jules Bordet Institute, Brussels, Belgium

#### **Belgium**

Institut Jules Bordet, Brussels Recruiting
Contact: Dr Christiane Jungels

#### Denmark

Rigshospitalet, Department of Oncology, Copenhagen Recruiting

Contact: Dr Morten Mau Sørensen

#### **The Netherlands**

Netherlands Cancer Institute, Amsterdam Recruiting

Contact: Prof Dr Jan H.M. Schellens

Erasmus MC, Rotterdam Recruiting

Contact: Dr Ferry ALM Eskens

#### Norway

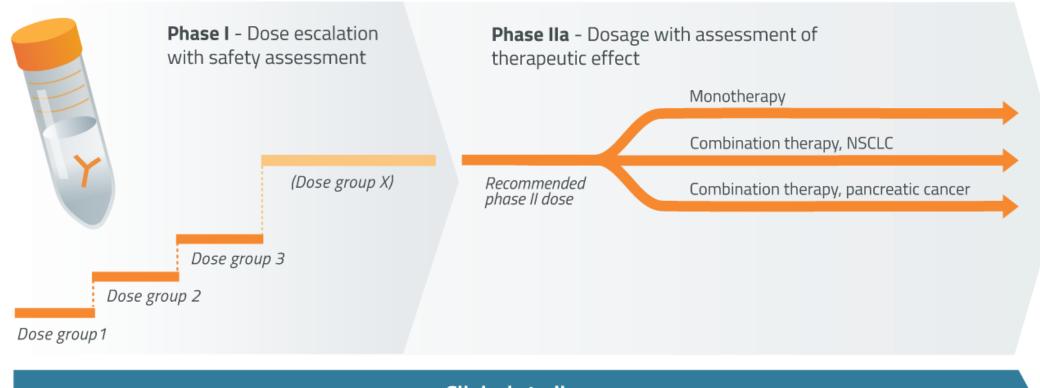
Oslo University Hospital, Radiumhospitalet, Oslo Recruiting

Contact: Dr Tormod Kyrre Guren





### CANO4 – CANFOUR clinical trial



**Clinical studies** 

Details on www.clinicaltrials.gov

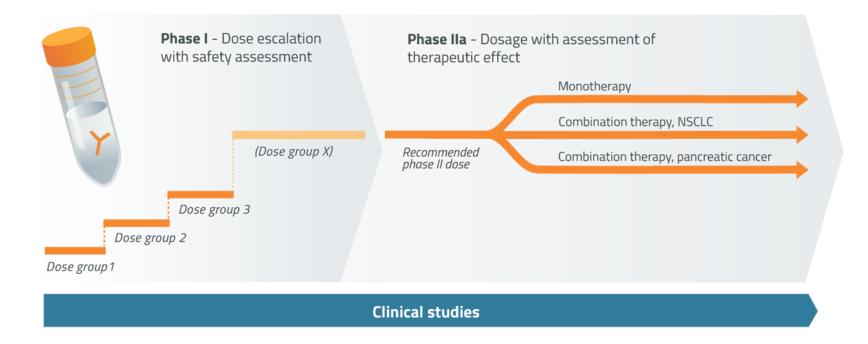
### **CANFOUR**

#### Part I

- NSCLC, PDAC, CRC, TNBC
- Safety and tolerability
- Pharmacokinetics
- Biomarkers
- Efficacy
- Identify RP2D

#### Part II

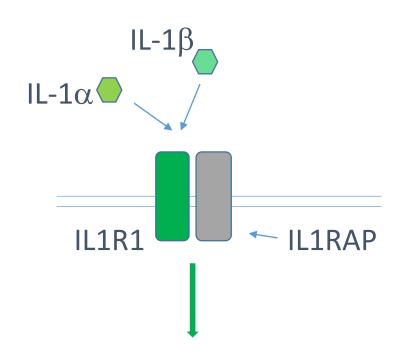
- NSCLC, PDAC
- Safety and tolerability
- Pharmacokinetics
- Biomarkers
- Efficacy
  - Monotherapy
  - Combination SoC



Patients will stay on treatment until disease progression, unacceptable toxicity, or discontinuation for any other reason



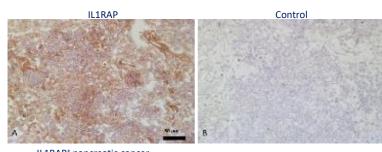
### Biomarkers and additional assessments



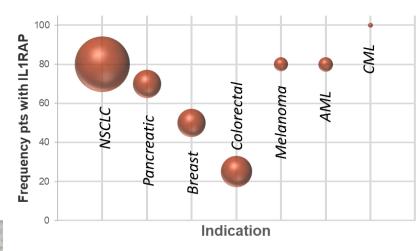
Cancer, Inflammation (IL-6 & IL-8)

- Pharmacokinetics
- Anti-drug Antibodies (ADA)
- Biomarkers:

- Expression of IL1RAP
- Expression of other IL1RAP associated biomarkers

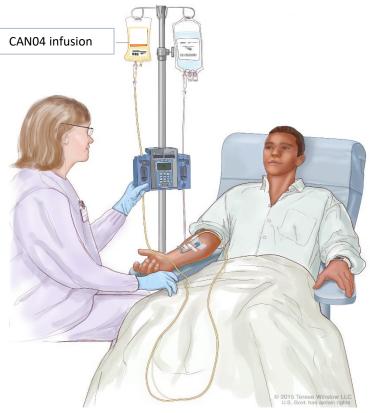


Biological matrix: Archived formalin fixed paraffin embedded tumor tissues



## CAN04 patient administration



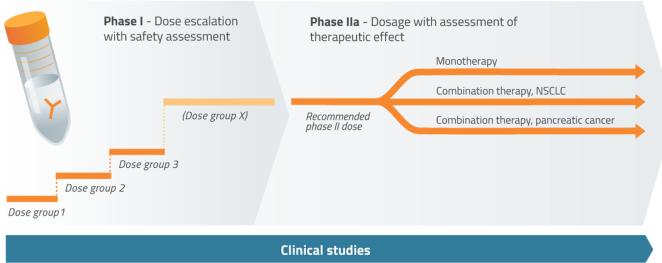


- CAN04 dose is calculated based on patient weight and the per protocol dose level
- CAN04 drug product is diluted in 0,9% NaCl infusion bag
- CAN04 study preparation is administered to the patient by intravenous infusion over 60 minutes

### Combinations with SoC

"Combination therapies may dramatically improve the outcome for cancer patients, and indeed it is expected that such therapies will eventually become the standard of care for cancer treatment, but the discovery of effective combinations is a challenging endeavor"

KM Morrissey et al. Clin Transl Sci (2016); 9: 89-104



### Combination with SoC - PAC

#### First line treatments available:

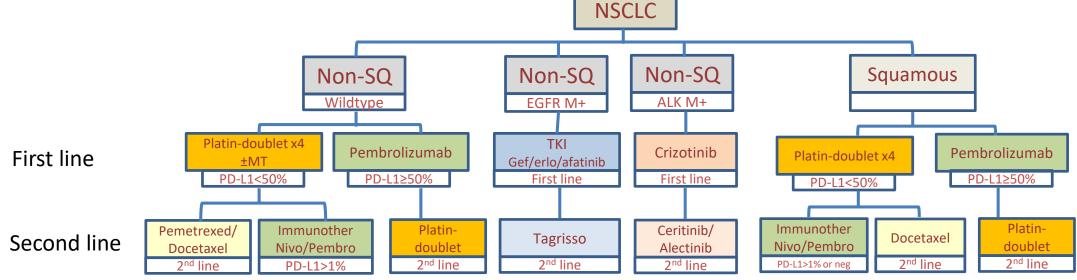
- FOLFIRINOX (5-FU, leucovorin, irinotecan and oxaliplatin)
- Gemcitabine and *nab*-paclitaxel (Abraxane)
- Gemcitabine monotherapy



### Combination with SoC - NSCLC

Three major options available for treatment:

- ImmunotherapyAnti-EGFR/ALK
- Chemotherapy 🔲 🔲



Current regimens in the treatment of NSCLC.

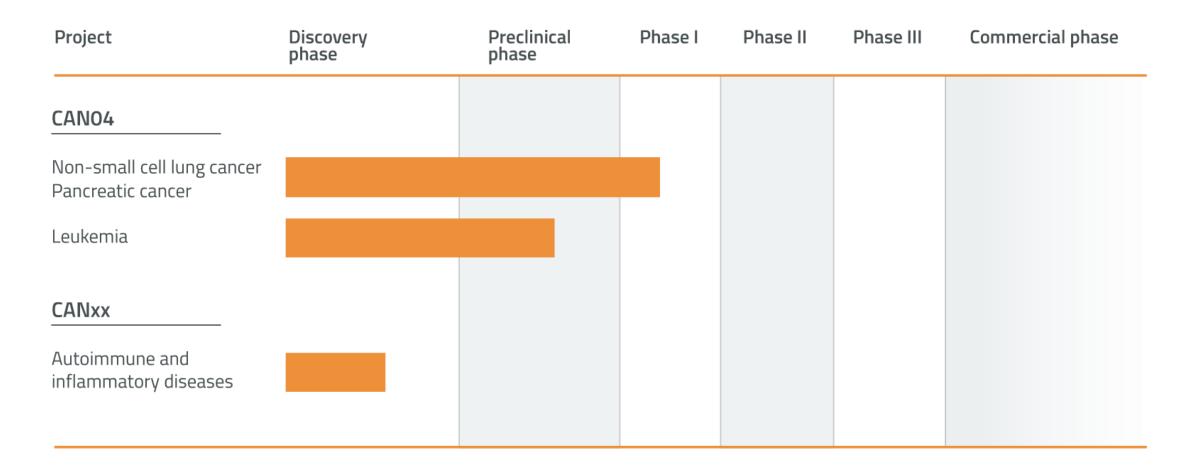
Adapted from Dr M Mau-Sørensen at Rigshospitalet, Copenhagen, Denmark

### Summary

- First patient treated with CAN04 September 2017
- Dose escalation phase (Part I) to be completed during summer 2018
- Dose expansion phase (Part II)
  - to follow directly after clinical completion of Part I
  - to evaluate CAN04 as monotherapy and in combination with SoC
  - to be completed by end of 2019

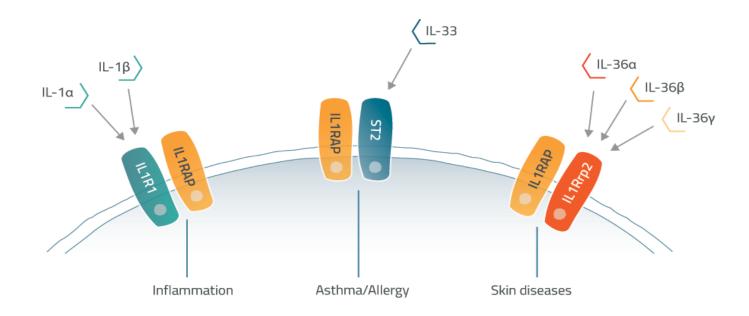


# Cantargia pipeline



# IL1RAP - additional potential indications to leverage the value of our asset

- Three different systems signal through IL1RAP
- These systems contribute to various inflammatory diseases
- Can be blocked by Cantargia's antibodies against IL1RAP



# Strategic partnership with Panorama Research Inc

#### Panorama Research Inc.

- Privately-owned biomedical R&D company in Silicon Valley California
- Leader in antibody technology, managed by Dr James W. Larrick

Deal structure: Panorama share risk in exchange for a fraction of future incomes



Panorama Research, Inc.

A Biotechnology Research & Development Company

- Development of new antibody binding IL1RAP with high affinity/potent inhibition of signaling
- Focus on autoimmune/inflammatory disease
- Selection of clinical candidate 2019
- Cantargia IL1RAP antibody, affinity matured and humanized using Panorama's proprietary technology
- Panorama also generates cell lines optimized for high level GMP production
- Cantargia responsible for subsequent development



## Significant value inflection points ahead

#### 2018

- Preclinical data on combination therapy
- Clinical progress
- Preclinical studies
- Phase I clinical data (summer 2018)
- Initiation of Phase IIa portion of the clinical trial (summer 2018)
- US regulatory and clinical strategy

# Cantargia IP

Use of IL1RAP as target hematological cancers

Filed 2009

Granted (EPO, USA, Japan, China)

Use of IL1RAP as target solid tumors

**Filed 2011** 

Granted (EPO, Japan, USA, China)

The product candidate CAN04

Filed 2014

Granted (EPO, USA)

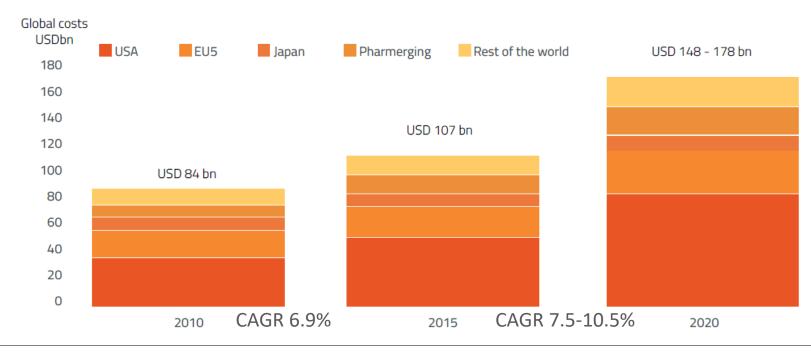
New IL1RAP antibodies (e.g. CAN03)

Filed 2014





# Lead project CAN04 in the highest growth segment— Oncology antibodies



World's most sold cancer drugs are antibodies (2016)

Immuno-oncology driving market growth 2016 (2015)

Rituxan/MabThera \$8.58bn
Avastin \$6.75bn
Herceptin \$6.75bn
Source IMS Health

Opdivo \$3.77bn (\$0.94bn)
Keytruda \$1.40bn (\$0.57bn)



## Why invest in Cantargia?

- Focus on immuno-oncology the strongest growing pharmaceutical segment
  - Taking advantage of the well established antibody technology to design novel targeted pharmaceuticals
- Lead candidate antibody CAN04 with double mechanism of action in clinical trial with multiple value inflection points 2018-2019
  - Initial development in NSCLC and pancreatic cancer (cancer forms with poor prognosis)
  - Important clinical data expected
  - Recent external validation of pathway
- Second generation antibodies for autoimmune disease
- Unique and strong IP
- Strong lead investors with high competence and well known track record
- Recent new share issue of 232 MSEK funding until mid 2020.

