



### **DNB's 8th annual Nordic Healthcare Conference**

14 December 2017



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This report contains certain forward-looking statements based on uncertainty, since they relate to events and depend on circumstances that will occur in future and which, by their nature, will have an impact on the results of operations and the financial condition of Targovax. Such forward-looking statements reflect the current views of Targovax and are based on the information currently available to the company. Targovax cannot give any assurance as to the correctness of such statements.

There are a number of factors that could cause actual results and developments to differ materially from those expressed or implied in these forward-looking statements. These factors include, among other things, risks or uncertainties associated with the success of future clinical trials; risks relating to personal injury or death in connection with clinical trials or following commercialization of the company's products, and liability in connection therewith; risks relating to the company's freedom to operate (competitors patents) in respect of the products it develops; risks of non-approval of patents not yet granted and the company's ability to adequately protect its intellectual property and know-how; risks relating to obtaining regulatory approval and other regulatory risks relating to the development and future commercialization of the company's products; risks that research and development will not yield new products that achieve commercial success; risks relating to the company's ability to successfully commercialize and gain market acceptance for Targovax's products; risks relating to the future development of the pricing environment and/or regulations for pharmaceutical products; risks relating to the company's ability to secure additional financing in the future, which may not be available on favorable terms or at all; risks relating to currency fluctuations; risks relating to the company's ability to retain key personnel; and risks relating to the impact of competition.



## Immunotherapy has the potential to cure cancer

Patient example – Yervoy® checkpoint inhibitor trial



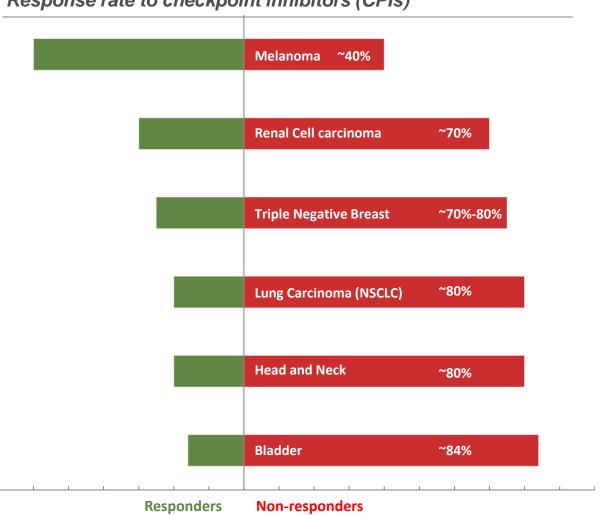
Prior to Yervoy®

1 year after



## Most patients do not respond to currently available immunotherapies

Response rate to checkpoint inhibitors (CPIs)



**Boosting T-cells in** tumors may make checkpoint inhibitors effective in more patients



## Targovax has two immuno-oncology programs in clinical development

#### **ONCOS** Oncolytic virus

- Genetically designed adenovirus
- Makes cancer antigens visible to immune system
- **Induces T-cells** specific to patients' tumor



#### TG RAS neoantigen vaccine

- Cocktail of synthetic peptides
- Mimics cancer causing RAS neoantigens
- **Induces T-cells** specific to **RAS mutations**





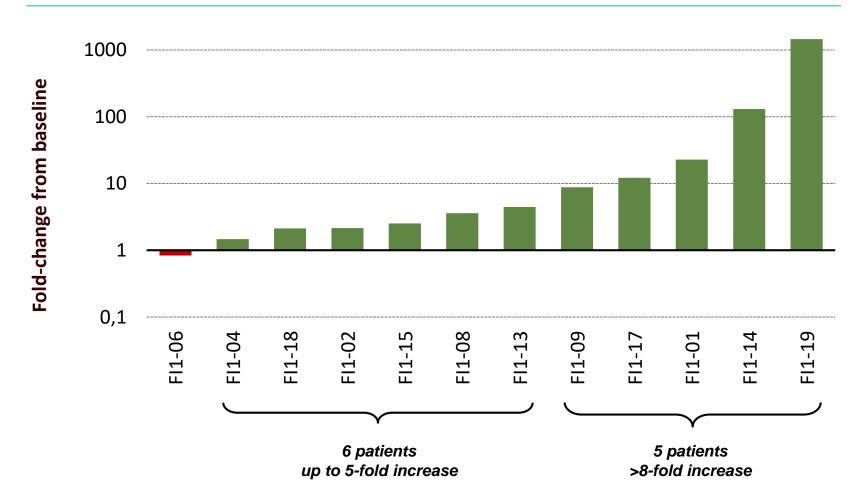
## **Agenda**

- ONCOS oncolytic virus platform
- TG mutRAS neoantigen vaccine platform
- Targovax clinical program overview



### **ONCOS-102** can increase T-cell count in tumors

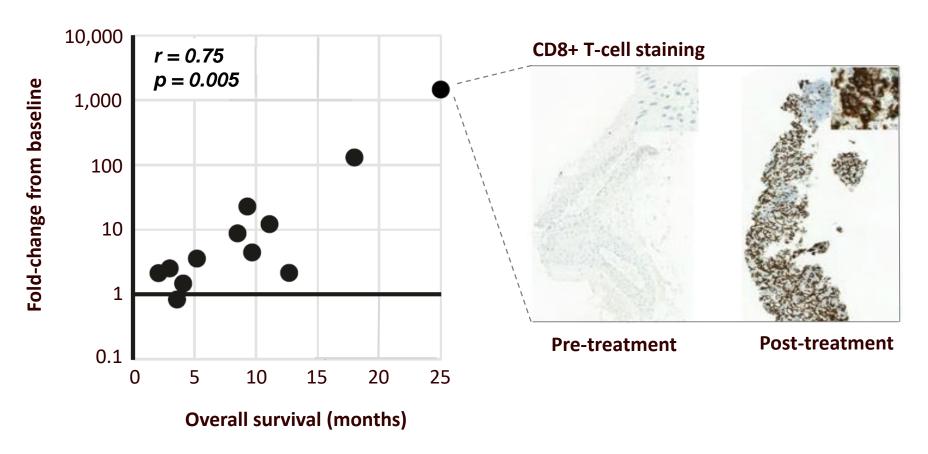
#### Phase I trial data: change in CD8+ T-cell count after treatment with ONCOS-102





### The T-cell increase correlates with survival

Phase I trial data: Fold-change CD8+ T-cell count vs. survival





## Clinical trial program overview

Completed trials
Ongoing trials
Starting trials

Compassionate
use program
Finland
115 patients

- Individual clinical responses
- Reassuring safety data

Phase I trial
7 Solid tumors
12 patients

 Correlation between immune activation and survival Ovarian / colorectal
Phase I/II

Collaboration with Ludwig, CRI
and MedImmune (AstraZeneca)

Intraperitoneal administration

Mesothelioma
Phase Ib/II
30 patients

up to 78 patients

- 1st line combination with chemo
- Randomized controlled trial

Prostate
Phase I
10 patients

- Partnered with Sotio
- Combination with DC therapy

Melanoma
Phase I
12 patients

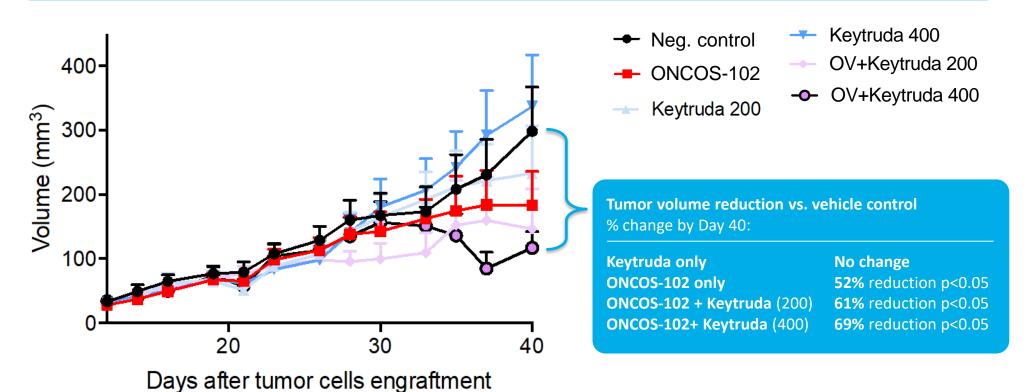
- Combination with PD-1
   CPI in refractory patients
- Memorial Sloan Kettering



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## 70% reduction in tumor volume with CPI combination in mouse melanoma model

Effect of ONCOS-102 and Keytruda in humanized mouse melanoma model, change in tumor volume





## **Agenda**

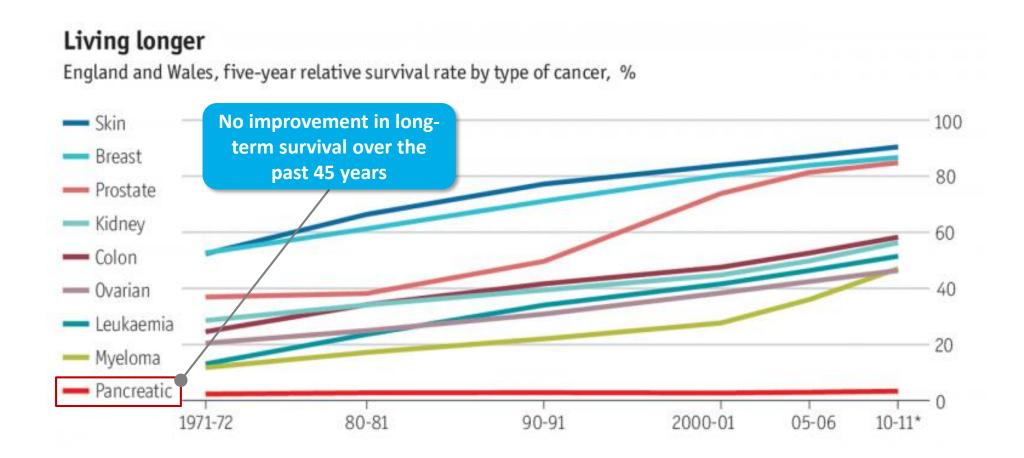
ONCOS oncolytic virus platform

**○** TG mutRAS neoantigen vaccine platform

Targovax clinical program overview



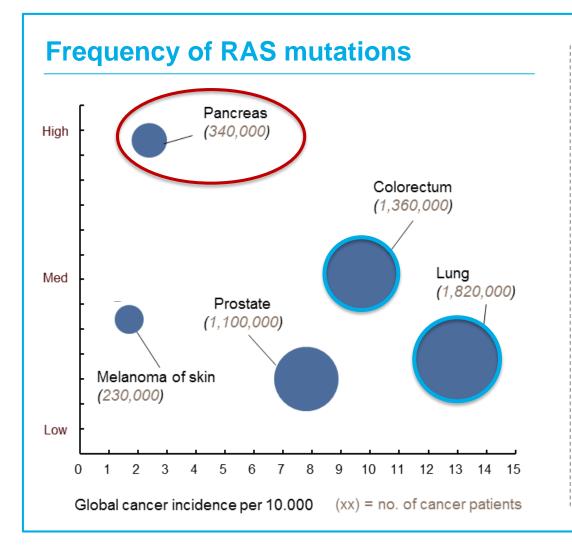
# The five year survival rate for pancreatic cancer patients has not improved since the 1970s





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# The RAS gene is mutated in 90% of pancreatic cancer patients, making it an ideal target

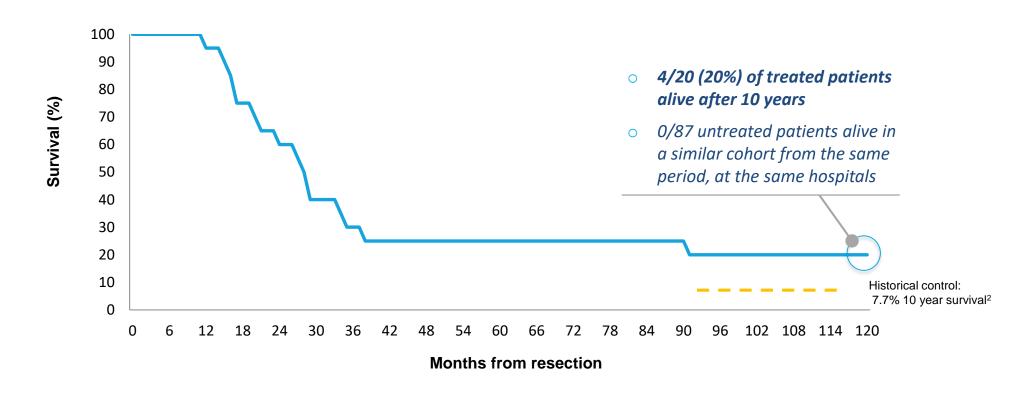


- RAS mutations result in uncontrolled cell division
- There are no existing therapies targeting RAS
- Targovax has developed a unique vaccine against mutant RAS

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# In previous trials in resected pancreatic cancer, TG vaccination has shown 20% 10 year survival

**10** year survival in historical TG trials in resected pancreatic cancer (n=20, TG monotherapy)





# These promising results are now being validated in an ongoing phase I/II trial with adjuvant chemotherapy

1<sup>st</sup> cohort (19 patients)

- Median survival 33.1 months vs. 27.6 for historical control
- 13 of 19 patients (68%) alive 2 years after surgery, vs. 30-53% in historical controls

**2**<sup>nd</sup> **cohort** (13 patients)

13 of 13 patients (100%) alive 1 year after surgery

mutRAS immune response (1 yr)

90% of patients (29/32) had RAS-specific immune activation

Safety

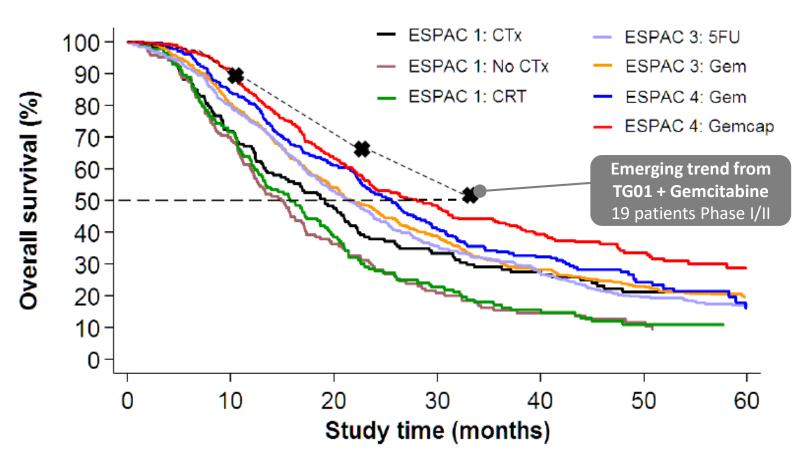
- TG01 and gemcitabine combination treatment is well-tolerated
- Four allergic reactions reported in 1<sup>st</sup> cohort, none in 2<sup>nd</sup> cohort (up to 1 year)



### **TG01** data in context

As presented by TG01 PI Prof. Daniel Palmer, London, June 2017

#### Comparative survival rates across trials in resected pancreatic cancer



NOTE: Relative survival curves across studies (ESPAC), meant for indicative comparisons only. No Kaplan Meier analysis has been done of the TG01 study data. Instead 1 and 2 year survival as well as median OS have been plotted.



### Why TG may succeed where others have failed

#### **Lessons Learned**

### The TG approach

**Target often poorly defined** and not cancer specific



Mutated **RAS** is a well-defined neoantigen, and a driving cause of cancer

Insufficient immune activation of CD4+ helper and CD8+ killer T-cells



TG peptides are **proven** to induce both CD4+ and CD8+ mutRAS T-cells

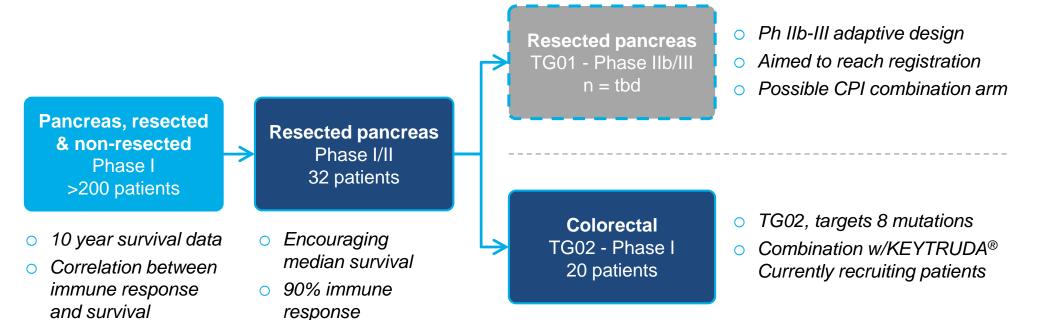
Most clinical trials have been done in advanced disease



Initial focus on resected patients, with stronger immune system



## Clinical trial program overview





# Resected pancreatic cancer is the lead indication, but all RAS mutated cancers are potential TG targets





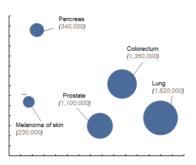












- TG01 lead indication
- Completing phase I/II
- Planning phase IIb/III
- 40.000 patients

- TG02 lead indication
- Phase I trial recruiting
- 50% RAS mutated
- O Up to 500.000 patients

- TG02 potential future indication
- 30% RAS mutated
- O Up to 500.000 patients

- TG02 + TG03 ultimate long-term potential
- 30% of all cancers
- Up to 30% of all cancer patients



Source: Global data, Riva et al. Plos One 2017

Estimated total addressable patient number with RAS mutations in US, EU and China

## **Agenda**

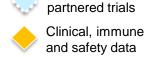
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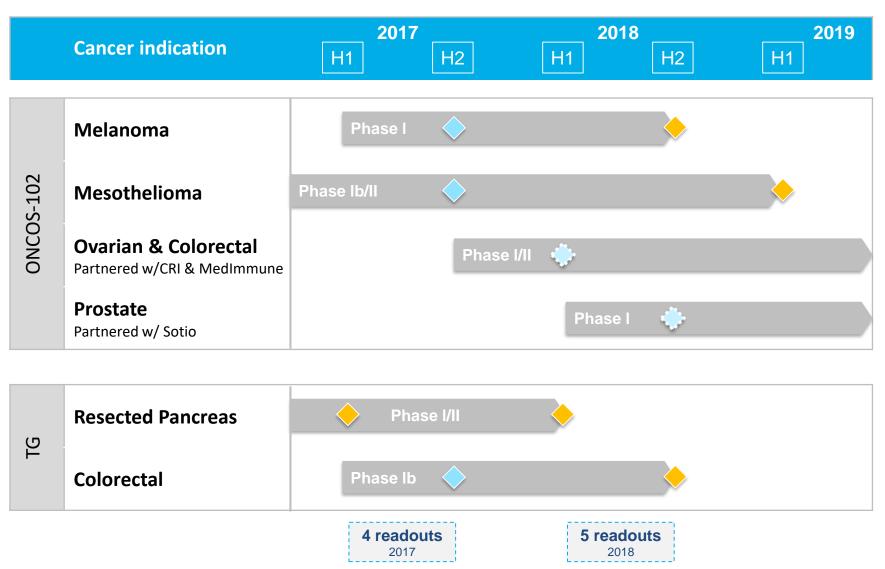


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## Interim data Interim data,

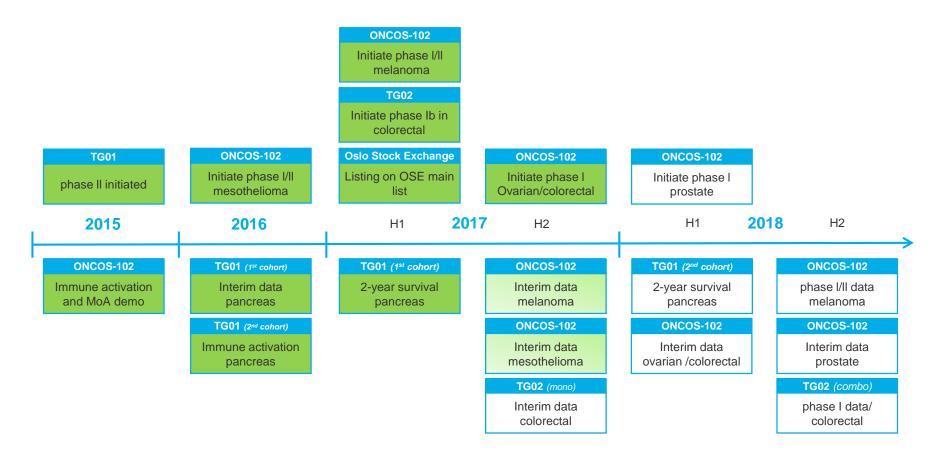
## Overview of Targovax' full clinical program





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# Strong upcoming news flow, with multiple near term value inflection points





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## Arming the patient's immune system to fight cancer

## Broad clinical program



- ✓ Six shots on goal
- ✓ Several upcoming data points

#### **ONCOS**



- ✓ Demonstrated ability to increase T-cell count
- ✓ Potential to make CPIs effective in more indications

**TG** 



- ✓ Unique approach for targeting RAS mutations
- ✓ Potential to benefit up to 1/3 of all cancer patients

