



Arming the patient's immune system to fight cancer

4Q & FY 2016 presentation

16 February 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.



# Fourth quarter highlights

**Patents** 

- Targovax granted an European patent for ONCOS-102, extending patent coverage
- Follows a similar US patent in June
- O Both expire in 2029

**Finances** 

- Cash NOK 172m
- Operating expenses NOK 31m
- Operating cash flow NOK -23m

People

Øystein Soug appointed as CEO on 1 November

Post-period

- Erik Digman Wiklund appointed CFO, starting April 2017
- Encouraging top line two-year survival data from TG01 clinical trial in resected pancreatic cancer patients



\* PoC = Proof of Concept

# TG01 Phase I/II resected pancreatic trial

 Encouraging top line two-year survival data -



# TG01 in resected pancreatic cancer: Encouraging survival rate and "signal" of efficacy

	First Cohort	Modified Cohort
1 Immunization schedule	<ul> <li>26 vaccinations over 2 years</li> </ul>	<ul> <li>15 vaccinations over 2 years</li> </ul>
Patient population	<ul> <li>15 eligible patients</li> <li>19 ITT<sup>1</sup>, 4 lost to follow up due to lack of consent</li> </ul>	<ul><li>Recruitment completed</li><li>13 patients</li></ul>
Immune activation	<ul><li>DTH response: 15 of 18</li><li>T-cell response: 6 of 8</li></ul>	<ul> <li>DTH response at 8 weeks: 4 of first 5</li> <li>T-cell response: not yet available</li> </ul>
Interim 1-year survival	<ul> <li>14 of 15 patients alive after 1 year</li> <li>No patients died from pancreatic cancer during the first year</li> </ul>	Not planned
2-year survival	<ul> <li>13 of 19 patients (68%) alive after 2 year</li> <li>Published* historical rate 30-53% suggests a signal of clinical efficacy for TG01</li> <li>Abstract submitted to ASCO 2017: efficacy, safety, immune activation data</li> </ul>	) 1H18
6 Safety	<ul><li>Generally well tolerated</li><li>4 allergic reactions triggering the "modified cohort"</li></ul>	Not yet available

<sup>&</sup>lt;sup>1</sup> ITT – Intention to treat



<sup>&</sup>lt;sup>2</sup> J Neoptolemos 2010, J van Loethem 2010, H Oettle 2013, M Sinn 2015, K Uesaka 2016 (In these reported studies overall survival is measured either from surgery or treatment randomization).

# TG – background – "reasons to believe"

**RAS** 

- RAS mutations are neoantigens
- Regulate cell proliferation. Mutations cause abnormal cell growth
  - definition of cancer
- Exclusively found in cancer cells

**TG-peptides** 

- Activate both RAS specific CD4+ and CD8+ T cells
  - recognize and destroy mutated RAS cells

History

- 120 patients treated with TG peptides in 1990's
- Encouraging long-term survival for resected patients treated with TG01 or single TG peptides<sup>1</sup>



# Encouraging survival rate and "signal" of efficacy in TG01 trial

CT TG01-01; A Phase I/II Trial of TG01 and Gemcitabine as Adjuvant Therapy for Treating Patients with Resected Adenocarcinoma of the Pancreas

- 68% (13 of 19) of the patients in cohort 1 were alive two years after the resection
  - Published historical rate 30-53% suggests a signal of clinical efficacy for TG01<sup>1</sup>
- Abstract submitted to ASCO 2017 (June) from this 1<sup>st</sup> cohort
  - Efficacy, safety, immune activation
- In summary: encouraging survival rate and "signal" of efficacy



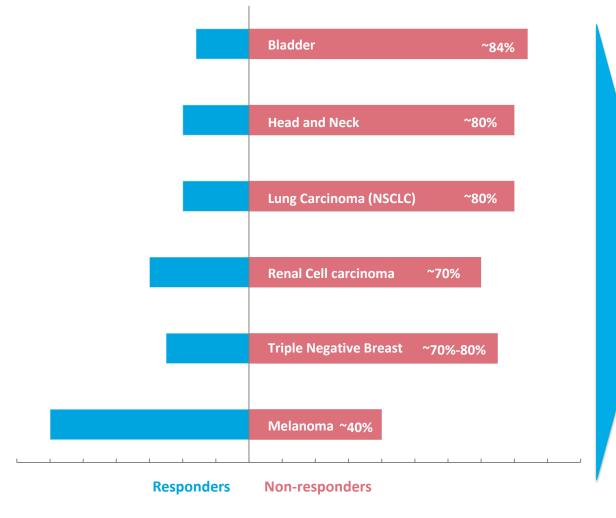
# **ONCOS-102** Phase I Melanoma trial

- Clinical proof of platform -



# Large unmet need for checkpoint inhibitor refractory patients

#### Response rate to checkpoint inhibitors (CPIs)



ONCOS-102 can potentially activate non-responders to become susceptible to CPI's



# **ONCOS-102: CPI refractory melanoma trial details**

# Background

No standard of care for patients not responding to CPI

### Setting

Advanced malignant melanoma patients not responsing to CPIs

 Immune activate CPI non-responders with ONCOS-102, then rechallenge with a CPI (Keytruda)

### **Cohorts**

Six patients with prior PD1 monotherapy

Six patients with prior PD1 plus Yervoy combination therapy

## Key endpoints

Safety

Immune activation and clinical response data

Correlation of immune activation and clinical response data

### Sequence

ONCOS-102 - 3 weeks

Keytruda – 5 months



## **How does ONCOS-102 work?**

#### At the tumor:

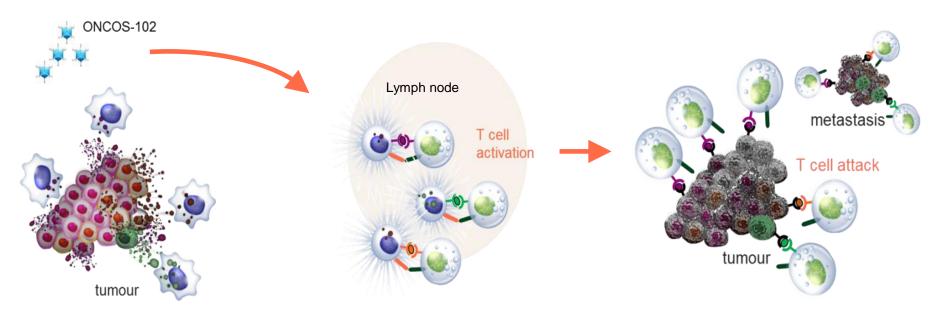
Virus injected directly into tumor, replicates, lyses cells and releases antigens. Immune system picks up antigens

#### At the lymph node:

Immune system starts production of tumor specific T-cells

#### At the tumor lesions:

T-cells find tumor lesions with corresponding tumor antigens and kill the cancer cells





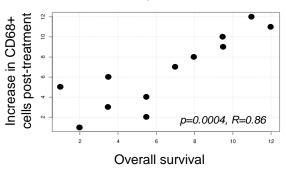
# Initial ONCOS-102 trial showed strong T-cell response

Evidence that immune system recognizes tumor threat

Innate Immune System (biopsy)

- Induction of proinflammatory cytokines + fever (all patients)
- Infiltration of innate immune cells into tumors in 11 out of 12 patients

Scatterplot of ranks



Correlation between post-treatment increase in innate immune cells and OS

Evidence that T-cells find the tumor and are cell killing

Adaptive immune system (biopsy)

- Increase in T-cell infiltration into tumors (including CD8+ killer T-cells) in 11 out of 12 patients
- Observation in one non-injected distant metastasis

OvCa. patient (FI1-19)





Correlation between post-treatment increase in CD8+ T-cells and OS (p=0.008, R=0.74)

Evidence that newly produced T-cells are tumor specific

Anti-tumor immune response (blood)

 Systemic induction of tumor-specific CD8+ T-cells

#### Ovarian patient:

NY-ESO-1, MAGE-A1, MAGE-A3, and Mesothelin specific CD8+ cells

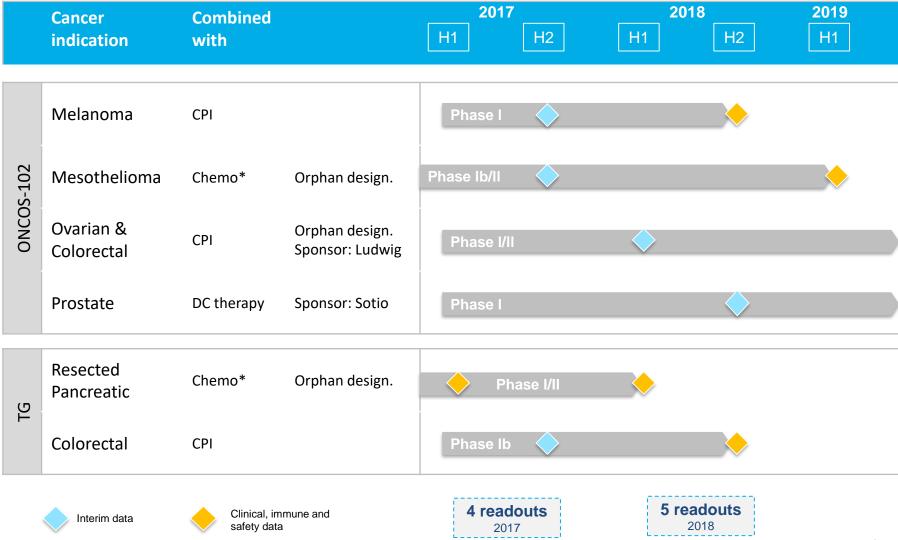
#### Mesothelioma patient:

MAGE-A3 specific CD8+ cells

Associated with clinical benefit

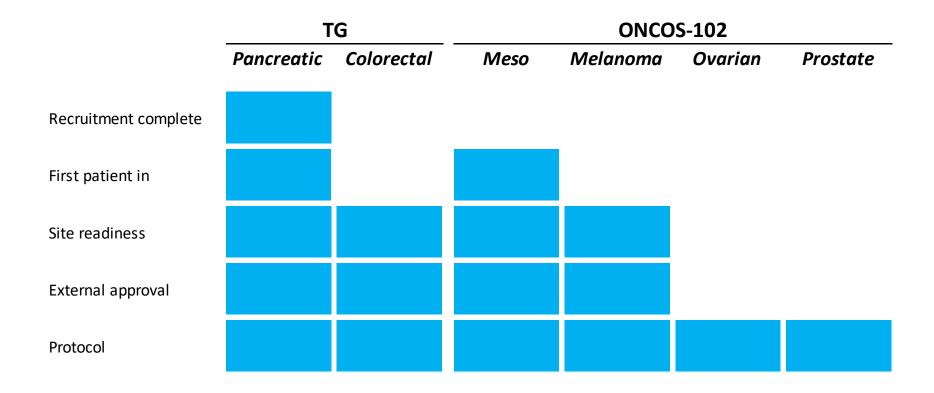


# Six shots on goal





# Where are we with the clinical trials?





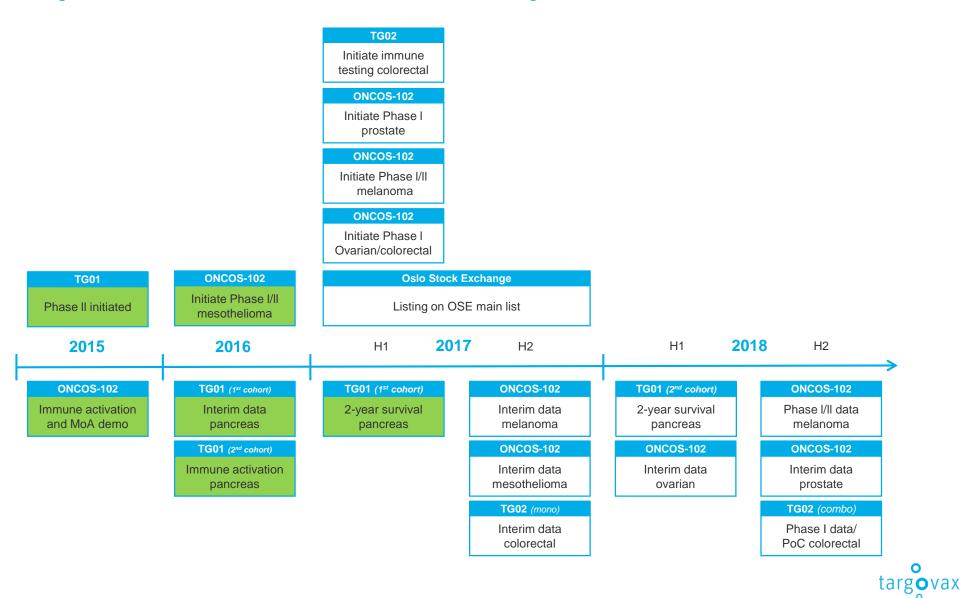
# **Financial summary**

Operations			
Cash	NOK 172m	USD 20m	
Yearly run rate	NOK 110m	USD 13m	Last four quarters
Annual opex	NOK 120m	USD 14m	Last four quarters

The share	OSE: TRVX				
Daily liquidity	NOK 9m	USD 1m Last two month's avg.			
Market Cap	NOK ~1 bn	USD 123m	At share price NOK ~24		
Debt	NOK 40m	USD 5m	EUR 6m conditional		
No. of shares	42.2m		44.9m fully diluted		
Analysts	DNB, ABG Sundal Collier, Arctic, Redeye, Norske Aksjeanalyser				



# Multiple near term value inflection points



# Arming the patient's immune system to fight cancer

- Core focus on immuno-oncology
- ✓ Lead product is an differentiated oncolytic adenovirus
- ✓ Targeting refractory solid, injectable tumors

Proprietary platforms and pipeline

- ✓ Promising Phase I data from two platform technologies
- ✓ Immunological findings linked to clinical benefit
- Multiple near term
  value inflection
  points
- ✓ Six combination trials (Phase I and II)
- ✓ All six trials read out in 2017-2018

4

**Corporate** 

- ✓ Oslo IPO in July 2016 (OSE:TRVX)
- ✓ Cash at approx. NOK 172m



# **Appendix**



18

# **Financial Snapshot**

### NOK m

	4Q15	1Q16	2Q16	3Q16	4Q16
Total revenue	0	-	-	0	0
		-	_	-	
External R&D expenses	-15	-11	-12	-11	-12
Payroll and related expenses	-15	-13	-12	-10	-13
Other operating expenses	-11	-7	-8	-4	-6
Total operating expenses	-41	-31	-32	-25	-31
Operating loss	-41	-31	-32	-25	-31
Net financial items	-1	-1	-1	-1	-1
Loss before income tax	-42	-32	-33	-26	-32
	-	-	-	-	
Net change in cash	-33	-33	-34	85	-21
Net cash EOP	174	141	107	193	172

# Strong shareholder base as per 6 February 2017

**Estimated ownership** 

Shareholder HealthCap RadForsk Nordea Rasmussengruppen KLP Nordnet Livsforsikring	No. of shares  11 155 584 4 077 255 2 594 239 1 820 000 1 703 333 1 207 802	26,4 % 9,7 % 6,1 % 4,3 % 4,0 % 2,9 %
RadForsk Nordea Rasmussengruppen KLP	4 077 255 2 594 239 1 820 000 1 703 333 1 207 802	9,7 % 6,1 % 4,3 % 4,0 %
Nordea Rasmussengruppen KLP	2 594 239 1 820 000 1 703 333 1 207 802	6,1 % 4,3 % 4,0 %
Rasmussengruppen KLP	1 820 000 1 703 333 1 207 802	4,3 % 4,0 %
KLP	1 703 333 1 207 802	4,0 %
	1 207 802	-
Nordnet Livsforsikring		2,9 %
	015 001	•
Statoil	915 981	2,2 %
Danske Bank (nom.)	770 916	1,8 %
Nordnet Bank AB (nom.)	739 998	1,8 %
Timmuno AS	724 650	1,7 %
Prieta AS	720 000	1,7 %
Sundt AS	400 000	0,9 %
Pohjola	320 966	0,8 %
ONB	291 993	0,7 %
Tobech Invest AS	286 449	0,7 %
Γhorendahl Invest AS	260 000	0,6 %
Netfonds Livsforsikring AS	253 639	0,6 %
Avanza Bank AB (nom.)	251 102	0,6 %
Danske Bank (nom.)	182 791	0,4 %
Molnar	181 800	0,4 %
Other shareholders (~2790)	13 332 302	31,6 %
Гotal	42 190 800	100,0 %

#### 42.2m ordinary shares

- Management ownership: 2.1%
- Approx. ~2810 shareholders

#### 44.4m<sup>1,2</sup> shares fully diluted

- Average strike price on options ~NOK 21
- Total dilutive effect of options is 5.6%<sup>1</sup>



<sup>&</sup>lt;sup>1</sup> As per 30 December 2016

<sup>&</sup>lt;sup>2</sup> Includes outstanding options (2,513,170) and Restricted Stock Units (129,991) to Board members