



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

DnB Healthcare conference

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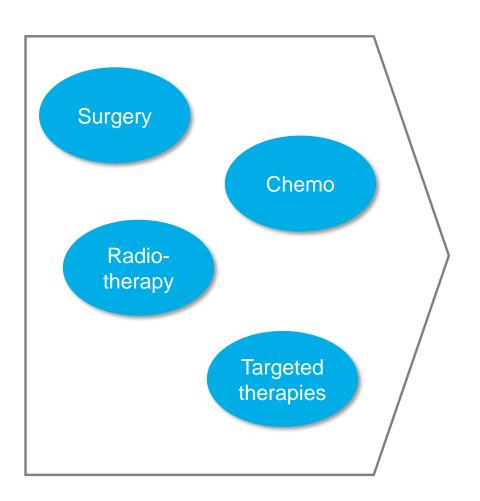
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## Immunotherapy – enables the immune system to kill cancer cells

#### **Traditional cancer treatment**

## **New approach - Immunotherapy**



#### **Enables the immune system to kill cancer cells:**

#### Oncolytic viruses

- Release cancer antigens
- Imlygic, ONCOS-102

#### Peptide vaccines

- Mimic cancer antigens
- TG01, TG02

#### Cell therapies

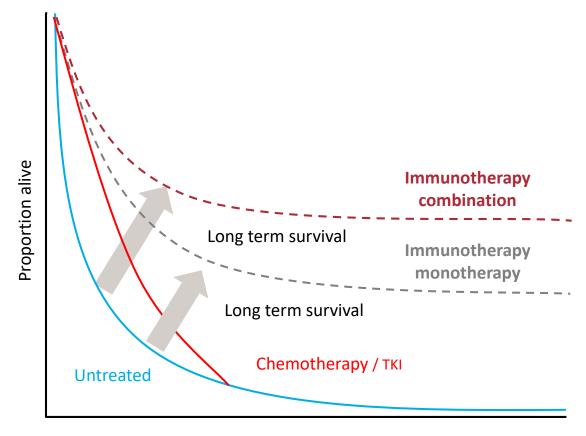
- Load T-cells with antigen receptors
- Chimeric antigen receptors, CARs

### Checkpoint inhibitors

- General upgrade of immune system
- Yervoy, Keytruda, Opdivo, Tecentriq



# The goal is to make cancer a chronic disease by combining immuno-oncology therapies



Time from treatment

- Yervoy started the revolution in cancer treatment in 2011
- Due to immuno-oncology combination the number of addressable cancers is expected to increase to at least 60%



# Checkpoint inhibitors show signs of "curing" some cancers - example of Yervoy treated melanoma



Prior to Yervoy



4 weeks



8 weeks



20 weeks



8 months

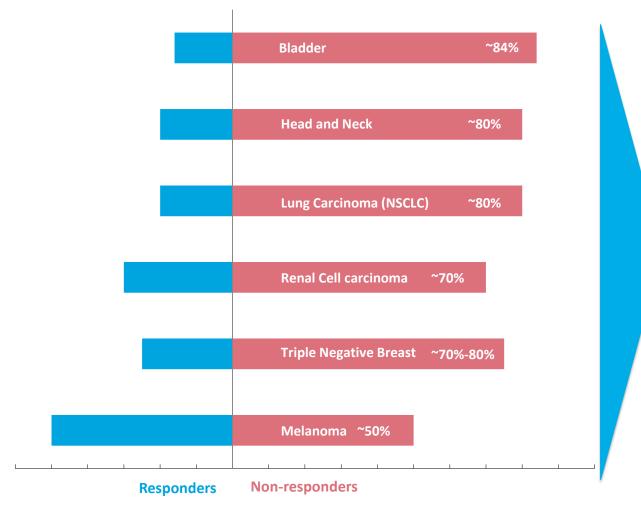


1 year



## 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

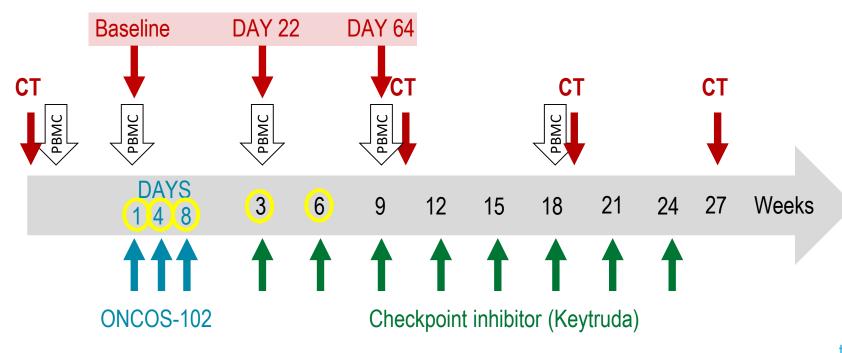


## **ONCOS-102: CPI refractory melanoma trial details**

#### Open-label Phase I trial

- ONCOS-102: 3 injections at day 1, 4 & 8
- O CPI (Keytruda) at day 22, then every 3 weeks for 5 months

## 3 biopsies per patient



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

#### At the tumor:

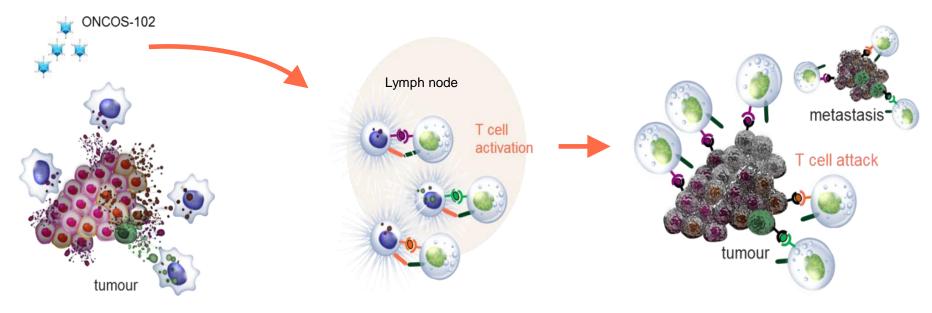
Virus injected directly into tumor, replicates, lyses cells and releases antigens. Immune system picks up antigens

#### 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





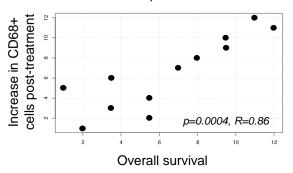
## Initial ONCOS-102 trial showed strong T-cell response

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

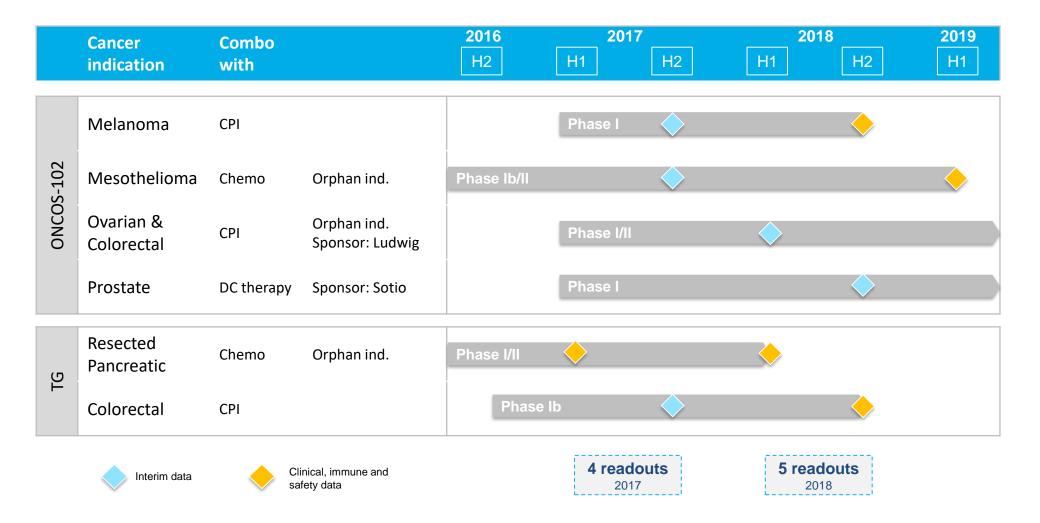
#### Mesothelioma patient:

MAGE-A3 specific CD8+ cells

Associated with clinical benefit



## Six shots on goal





## TG01 upcoming data: Two-year survival in resected pancreatic cancer

#### 2015-16 data demonstrated

- 14 of 15 patients alive after 1 year (19 ITT, 15 evaluable patients)
- DTH response 15 of 18 patients
- RAS specific T-cell response: 6 of first 8 patients

First half 2017

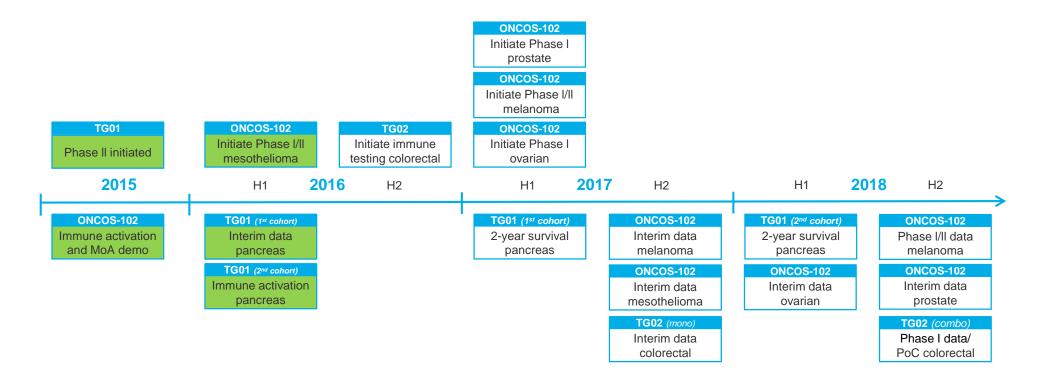
Two-year survival

First cohort: 19 patients

TG has potential to enable the immune system to identify and destroy tumors bearing any RAS mutations.



## Multiple near term value inflection points





## **Financial summary**

Operations	
Cash	<b>NOK 193m</b> at 30 September 2016
Cash run rate	NOK 121m (last four quarters)
Annual opex	NOK 129m (last four quarters)

The Targovax share	OSE: TRVX
Daily liquidity	NOK 4m (last month's avg.)
Market Cap	NOK 600m
Number of shares	42,2m (44,4m fully diluted)
Analyst coverage	DNB, ABGSC, Arctic, Redeye, Norske Aksjeanalyser



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

Two year survival data of TG01 in resected pancreatic cancer TG Data in 1H17 Important proof of concept trial in CPI refractory melanoma **ONCOS** Data in 2H17 ✓ Six shots on goal **Clinical trials** 

