The Impact of Age and Comorbidity on Effect of Treatment, Adverse Effects and QoL in Danish Lung Cancer Patients Receiving Immunotherapy.

#### PhD group:

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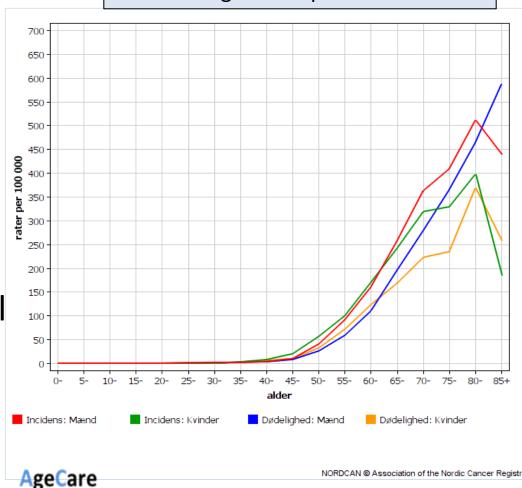
Region Syddanmark

# Lung Cancer (NSCLC)

• Medium age at diagnosis = 70 years.

### <u>Up until 2015:</u>

- Stage IV NSCLC
- No long-term survival



Danish Lung Cancer patients 2015

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## Immunotherapy for NSCLC

The NEW ENGLAND JOURNAL of MEDICINE

#### ORIGINAL ARTICLE

### Nivolumab versus Docetaxel in Advanced Squamous-Cell Non–Small-Cell Lung Cancer

Julie Brahmer, M.D., Karen L. Reckamp, M.D., Paul Baas, M.D., Lucio Crinò, M.D., Wilfried E.E. Eberhardt, M.D., Elena Poddubskaya, M.D., Scott Antonia M.D. Bh.D. Adam Bluzanski, M.D. Bh.D. Everett E. Vokes, M.D. n engl j med 373;2 nejm.org july 9, 2015

### The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

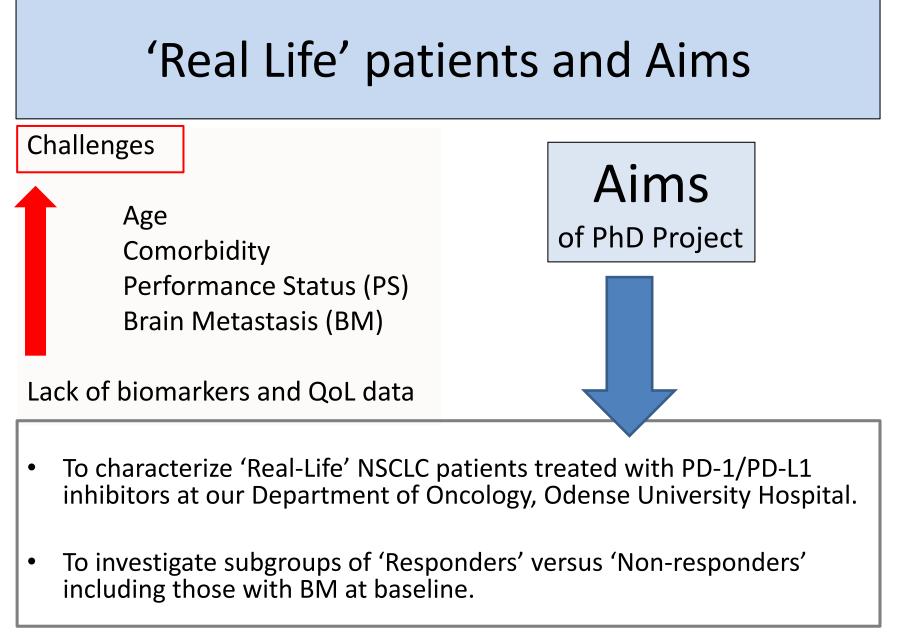
NOVEMBER 10, 2016

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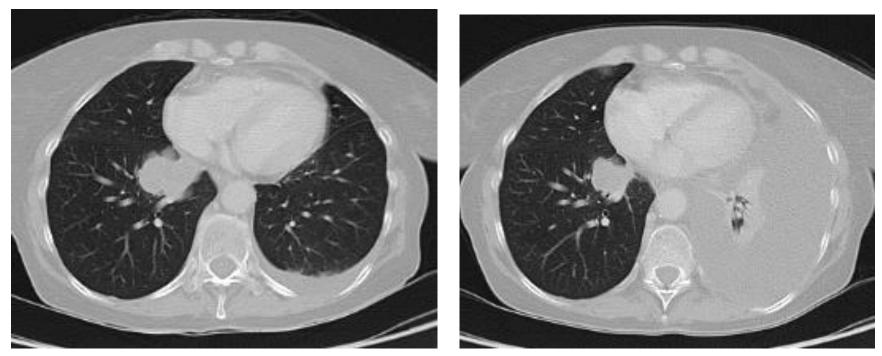
#### Pembrolizumab versus Chemotherapy for PD-L1–Positive Non–Small-Cell Lung Cancer

Martin Reck, M.D., Ph.D., Delvys Rodríguez-Abreu, M.D., Andrew G. Robinson, M.D., Rina Hui, M.B., B.S., Ph.D., Tibor Csőszi, M.D., Andrea Fülöp, M.D., Maya Gottfried, M.D., Nir Peled, M.D., Ph.D., Ali Tafreshi, M.D., Sinead Cuffe, M.D., Mary O'Brien, M.D., Suman Rao, M.D., Katsuyuki Hotta, M.D., Ph.D., Melanie A. Leiby, Ph.D., Gregory M. Lubiniecki, M.D., Yue Shentu, Ph.D., Reshma Rangwala, M.D., Ph.D., and Julie R. Brahmer, M.D., for the KEYNOTE-024 Investigators\*

- Checkpoint- or PD-1/PD-L1 inhibition.
- 2015: For selected patients based on Phase III Studies.



### PD-1 inhibition. Response evaluation?



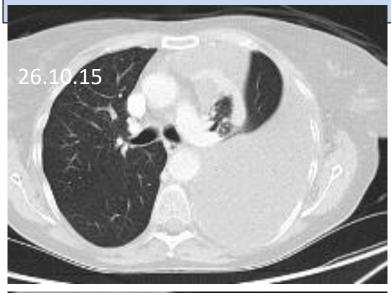
- Radiologic tumor regression but....
- PD of pleural effusion and...
- Clinical Benefit.

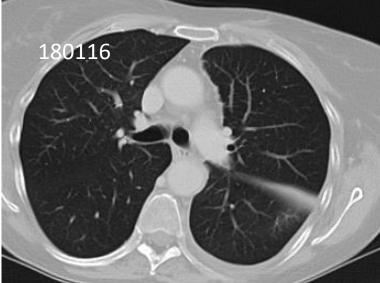
**F** 

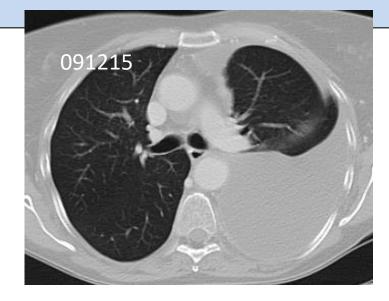
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### **Continued PD-1 inhibition**









## Patients and Methods

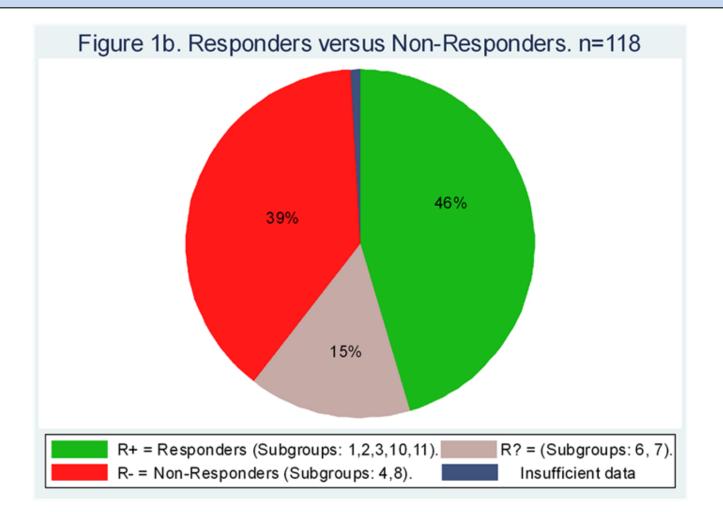
 All NSCLC (incurable IIIA-IVB) patients treated with PD-1/PD-L1 inhibitors from January 2015-April 2018 (n=118). Stratified into groups:

R+: Radiologic Responders with Clinical Benefit or No-Change Clinical (n=54).

R-: Radiologic Non-responders independent of clinical response (n=46).

R?: An intermediate 'discrepancy' group with Radiologic Response but Clinical Deterioration (n=18).

## **Preliminary Results**



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# Preliminary results/conclusion

### NSCLC in general (n=118):

- Patients with PS=2 (n=10) seem to respond to treatment as well.
- No patients with PS=3 (n=2) responded to treatment.
- There might be more discrepancy between clinical benefit and radiologic response with higher age and higher comorbidity.
- More prospective studies are needed and subgroup analysis.

### Preliminary results

Subgroup with Brain Metastasis at baseline n=22		R+ n= 10 (45%)	R- n= 11 (55%)	R? n= 1 (5%)
<b>Time on treatment</b> Days, mean, range	187 [0-700]	358 [51-700]	37 [0-105]	126 [126]
<b>Status,</b> n (%): - Ongoing - Follow-up	5 (23%) 3 (14%)	5 (50%) 3 (30%)	0 (0%) 0 (0%)	0 (0%) 0 (0%)

- They respond to checkpoint-inhibitors (comparative mean treatment time of 187 days/164 days).
- 'Responders' seem to benefit from treatment (mean ToT > 300 days/37 days , and 30%/0% are in FU).

## Perspectives



### Part 1: Retrospective data analysis

 More data analysis on OS, PFS and toxicity in all patients (n=118) and those with BM (n=22).

#### Part 2: Biomarker, toxicity, comorbidity.



- A prospective study was initiated in May 2018 (20/150 patients have been included so far).
- Blood samples consecutively for biobank, MRI and expanded CT scans are performed (for BM and comorbidity).



#### Part 3: Quality of Life

- EORTC-QoL 30 og Euro-5Q-5D-5L. Consecutively.
- Clinical benefit is relevant also for patients not fulfilling criteria for an objective treatment response.

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