Hackaton Non-small cell lung cancer (NSCLC)

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Outline

- What is non-small cell lung cancer (NSCLC)?
- What causes NSCLC?
- How do we diagnose NSCLC?
- How do we treat NSCLC?

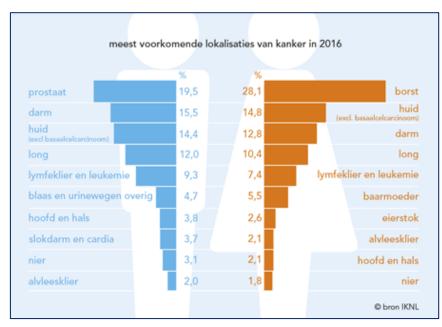


Introduction

Lung cancer incidence Netherlands:

- 12.000 new patients/ year
- 85% of all lung cancer: non-small cell lung cancer (NSCLC)
- 1-yr OS 44% (2006-2010; so after 1 year 44% is alive, 56% died)
- 5-yr OS 17% (2006-2010)

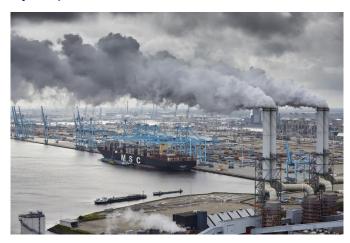






What causes non-small cell lung carcinoma (NSCLC)?¹

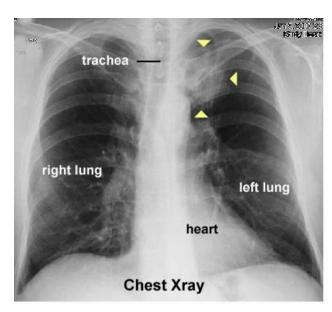
- Tobacco smoking
- Secondhand tobacco smoking
- Diet
- Alcohol
- Air pollution
 - Proportion of lung cancers attributable to urban air pollution in Europe is estimated to be 11%² (Netherlands: contributes to 1200 new cases per year)
- Occupational exposure (asbestosis)
- Genetic factors



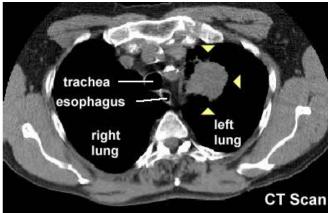


Clinical manifestations¹

- Cough
- Hemoptysis (coughing blood)
- Shortness of breath
- Chest pain

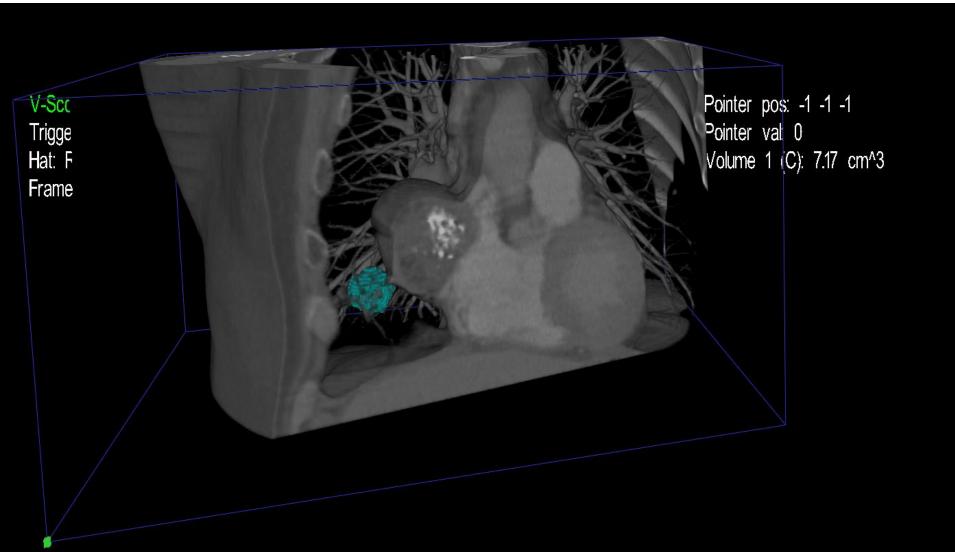






Left Upper Lobe Cancer note how much more obvious the tumor is on the CT scan compared to chest Xray

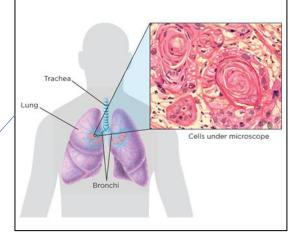




Treatment

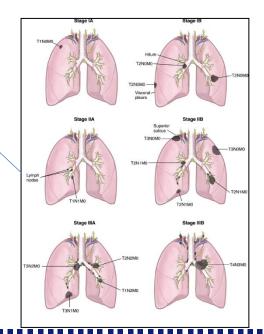
Treatment based on:

Tumor type by microscope(morphology)/ mutation analysis



Squamous/ non-squamous



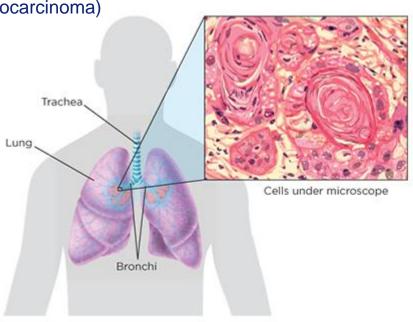




Non-small cell lung cancer

- Two types (obtained by biopsy, imaging by microscope):
 - Squamous cell carcinoma (Nederlands: plaveiselcelcarcinoom)

Non-squamous cell carcinoma (mainly adenocarcinoma)

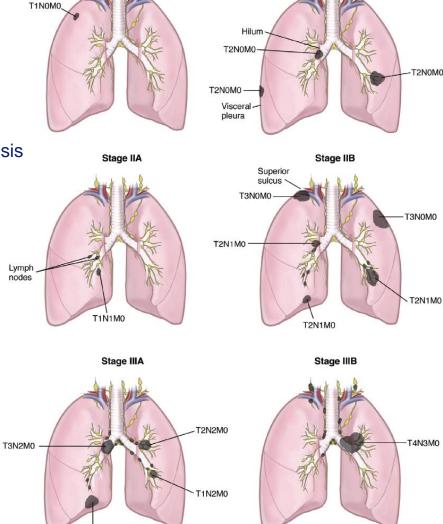




Staging NSCLC based on TNM classification

- T= tumor size, N=lymph node, M= distant metastasis
- The more extensive, the worse prognosis (chance to survive after 5 years at stage IV is 1%)

Stage	TMN	5-yr Survival Rates
IA	T1N0M0	~70%
IB	T2N0M0	~60%
IIA	T1N1M0	55%
IIB	T2N1M0 T3N0M0	~40%
IIIA	T1-3N2M0 T3N1M0	~25%
IIIB	Any T4,N3, M0	<5%
IV	Any M1	1%



Stage IA

T3N1M0



Stage IB

Treatment

Treatment NSCLC based on staging:

Treatment 1400E0 based on stagning



- Adjuvant chemotherapy
 - Squamous cell carcinoma: 4x chemotherapy (cisplatin/gemcitabin)
 - Non-squamous carcinoma: 4x chemotherapy (cisplatin/pemetrexed)

Stage III irresectable → radiotherapy + chemotherapy

- 3x chemotherapy (cisplatine/etoposide) and concurrent radiotherapie
- Stage III irresectable at recurrence: first/second line therapy stage IV







Treatment

IV

Stage IV → Systemic therapy (chemotherapy, targeted therapy, immunotherapy)



First line therapy

- Treatable driver mutations:
 - E.g. HER2, BRAF, RET, MET, EGFR, ELM-ALK, ROS1 → TKI
- Without treatable driver mutations:
 - Platinum-based chemotherapy
 - Pembrolizumab for pts with a high level (>50%) PD-L1 expression (1)

Second line therapy

- Treatable driver mutations: E.g. T790M → TKI
- Without treatable driver mutations:
 - Nivolumab (3,4)
 - Pembrolizumab for pts with a level >1% PD-L1 expression (5)
 - Pemetrexed, docetaxel

Erasmus MC

1. Reck et al., NEJM 2016, 2. Hellman et al., NEJM 2018, 3. Brahmer et al., NEJM 2015, 4. Borghaei et al., NEJM 2015

5. Herbst et al., Lancet oncol 2016.

To memorize !



- TNM classification
 - <u>Tumor size</u>, lymph <u>node involvement</u>, distant <u>metastases</u>
- Staging
 - Based on TNM
 - I, II, III and IV (bad to worse)
- Treatment
 - Based on stage
 - Stage I/II: chemo surgery
 - Stage III chemo radio, chemo
 - Stage IV chemotherapy/ targeted therapy/ immunotherapy

