



METABOLIC PHENOTYPING BY ^1H - NMR SPECTROSCOPY DETECTS LUNG CANCER VIA A SIMPLE BLOOD SAMPLE

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CONTENT

- Lung cancer
- Research topic
- Results
- Future perspectives

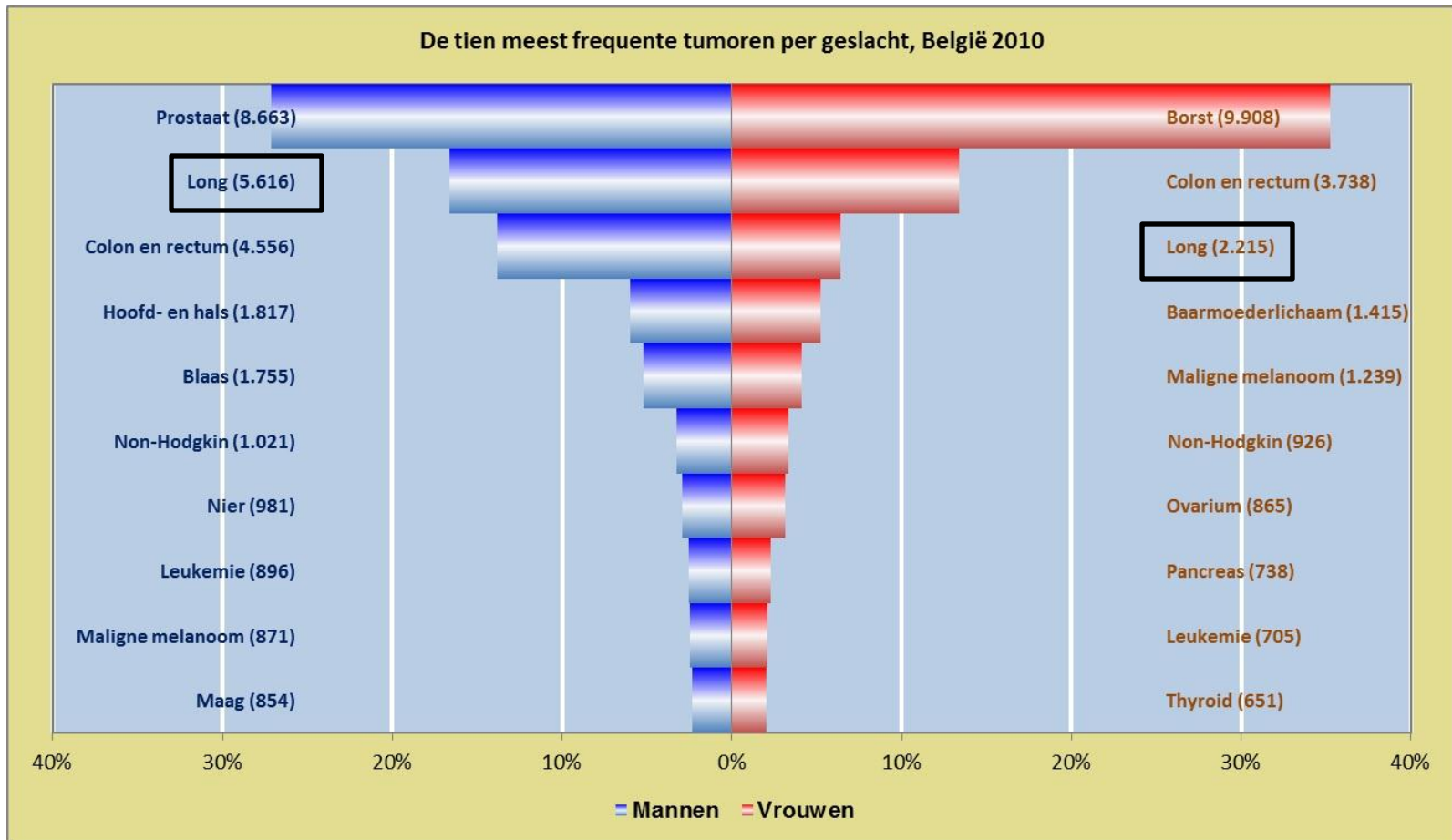


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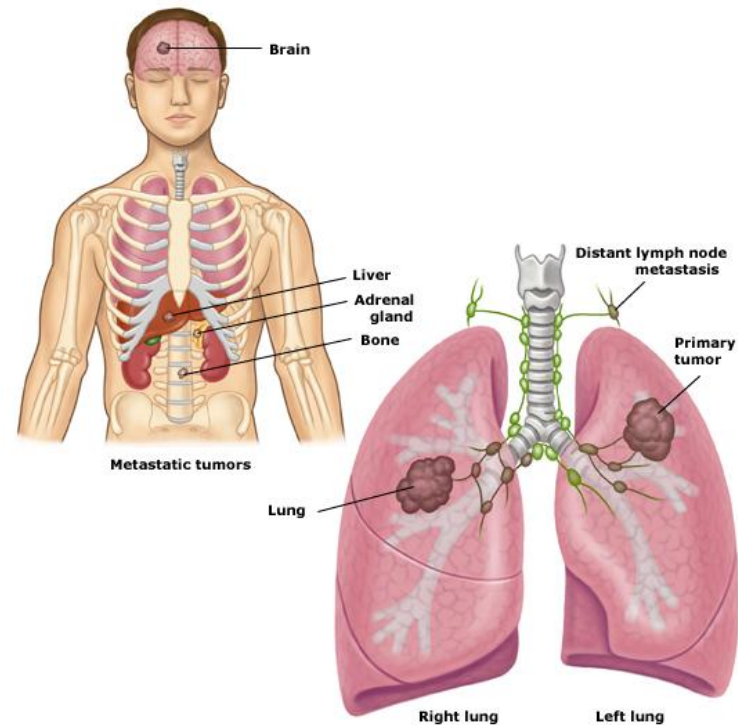


NUMBERS IN BELGIUM



LUNG CANCER

- Average 5-year survival of 15%
- Diagnosed in a metastatic stage
- Harmful and expensive chemotherapies



LINK BETWEEN TUMOR STAGE AND PROGNOSIS

FIGURE 70 - LUNG CANCER: RELATIVE SURVIVAL BY STAGE IN MALES (BELGIUM, 2004-2008)

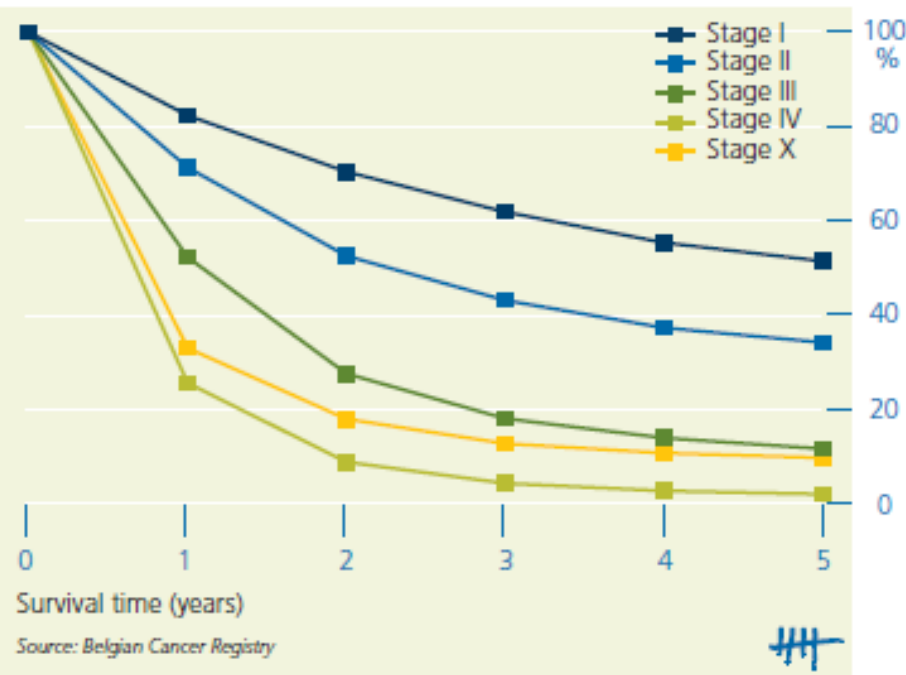
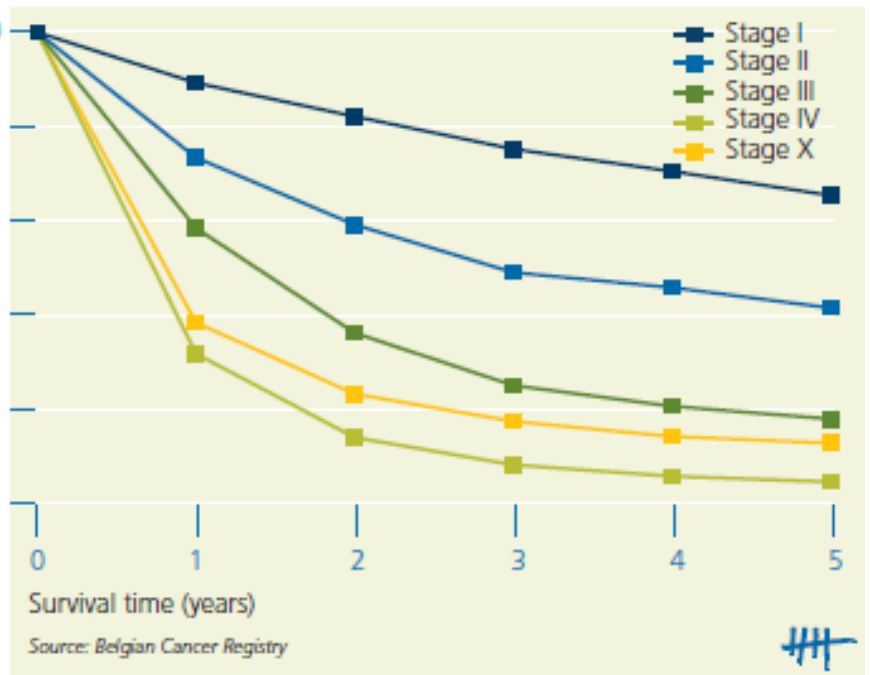


FIGURE 71 - LUNG CANCER: RELATIVE SURVIVAL BY STAGE IN FEMALES (BELGIUM, 2004-2008)



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RESEARCH TOPIC

- Problem:

Urgent need of effective methods to detect lung cancer in an early stage

- Hypothesis:

Analyses of the metabolic composition of blood plasma by $^1\text{H-NMR}$ spectroscopy allows to detect lung cancer



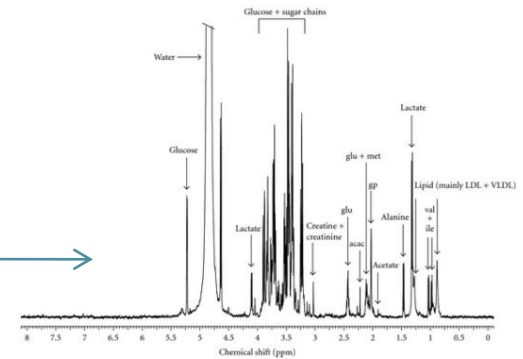
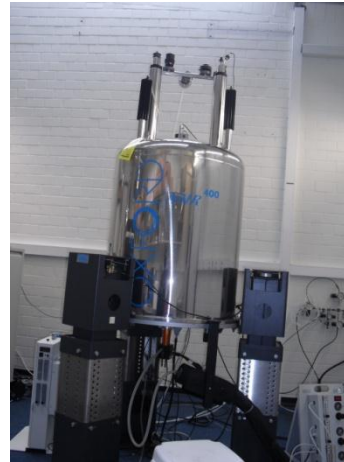
RESEARCH TOPIC



Control subjects



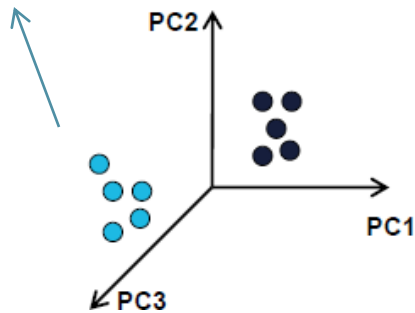
Lung cancer patients



$^1\text{H-NMR}$ spectrum

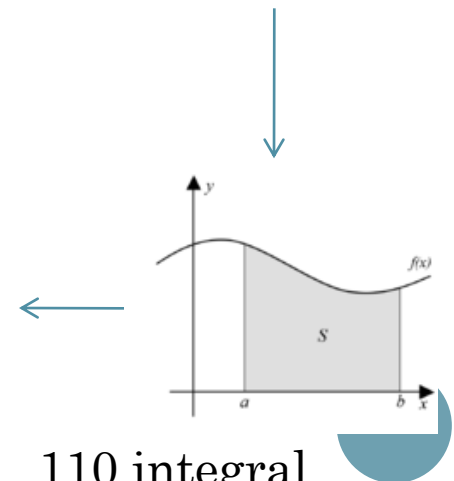
Analysis by $^1\text{H-NMR}$ spectroscopy

Metabolic interpretation



Statistical analyses

| | Patient 1 | Patient 2 | Patient 3 | Patient 4 | Patient 5 | Patient 6 | Patient 7 | Patient 8 | Patient 9 | Patient 10 |
|---------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|------------|
| Metab 1 | | | | | | | | | | |
| Metab 2 | | | | | | | | | | |
| Metab 3 | | | | | | | | | | |
| Metab 4 | | | | | | | | | | |
| Metab 5 | | | | | | | | | | |
| Metab 6 | | | | | | | | | | |
| ... | | | | | | | | | | |



110 integral segments

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SUBJECT CHARACTERISTICS

| | Lung cancer patients | Control subjects |
|-----------------------|--|---|
| Number | 78 | 78 |
| Gender | Male: 53 (67%) Female: 26 (33%) | Male: 45 (58%) Female: 33 (42%) |
| Average age | 68 ± 9 | 64 ± 13 |
| Smoking habits | Active: 50 Stopped > 6m: 25 Never: 3 | Active: 19 Stopped > 6m: 27 Never: 32 |



HISTOLOGY AND STAGE - 83 LUNG TUMORS

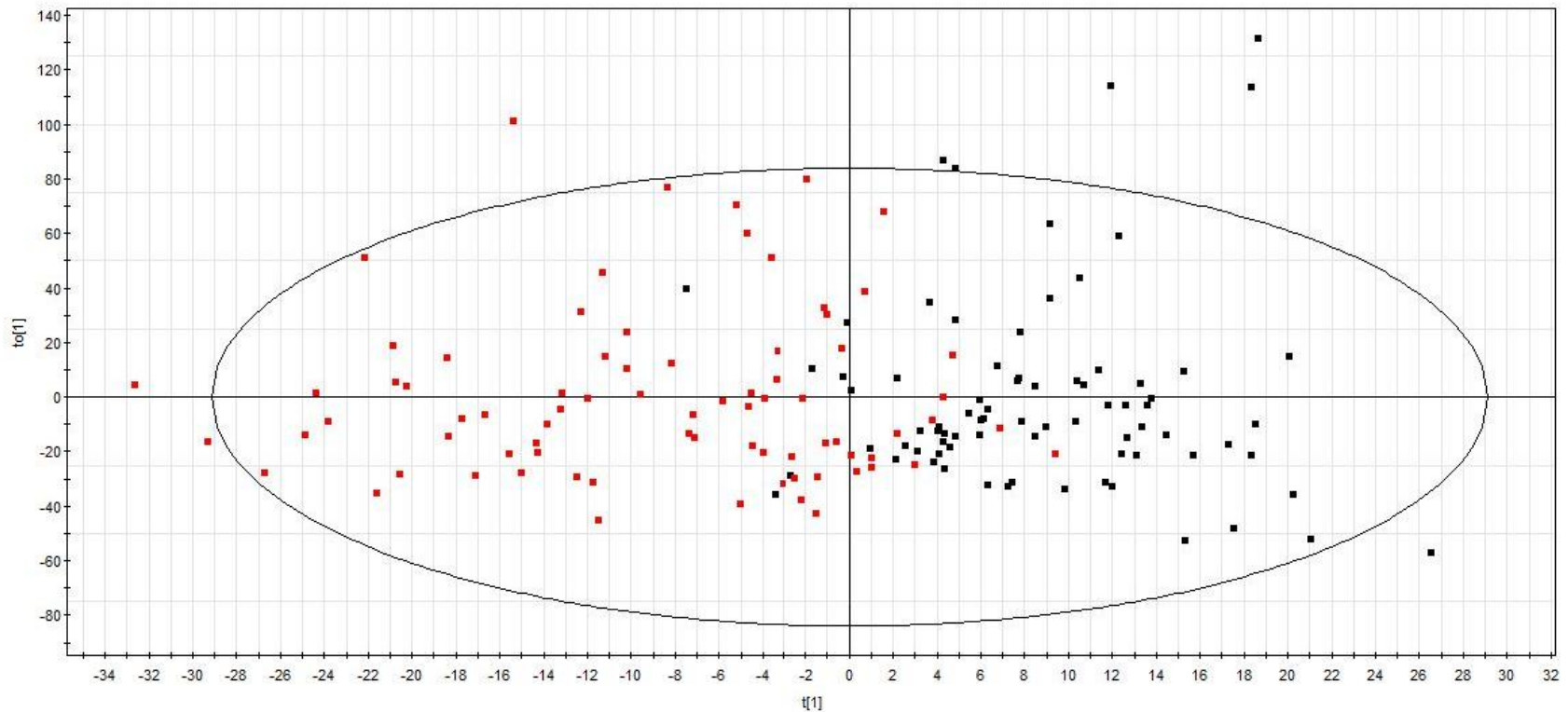
| | |
|---|------------------------------|
| Non-small cell lung cancer or NSCLC (59) | Adenocarcinoma (29) |
| | Spinocellular carcinoma (22) |
| | Large cell carcinoma (4) |
| | Adenosquamous carcinoma (3) |
| | Carcinoid (1) |
| Small cell lung cancer or SCLC (14) | |
| Unknown histological diagnosis (10) | |

| | | | | |
|-------------------------|-----------------------|-----------|----------------------|---------|
| Lung tumors (83) | Stage I (25) | | Stage II (7) | |
| | IA (20) | IB (5) | IIA (5) | IIB (2) |
| | Stage III (30) | | Stage IV (21) | |
| | IIIA (17) | IIIB (13) | | |



MULTIVARIATE DATA ANALYSIS - ALL 110 INTEGRAL SEGMENTS

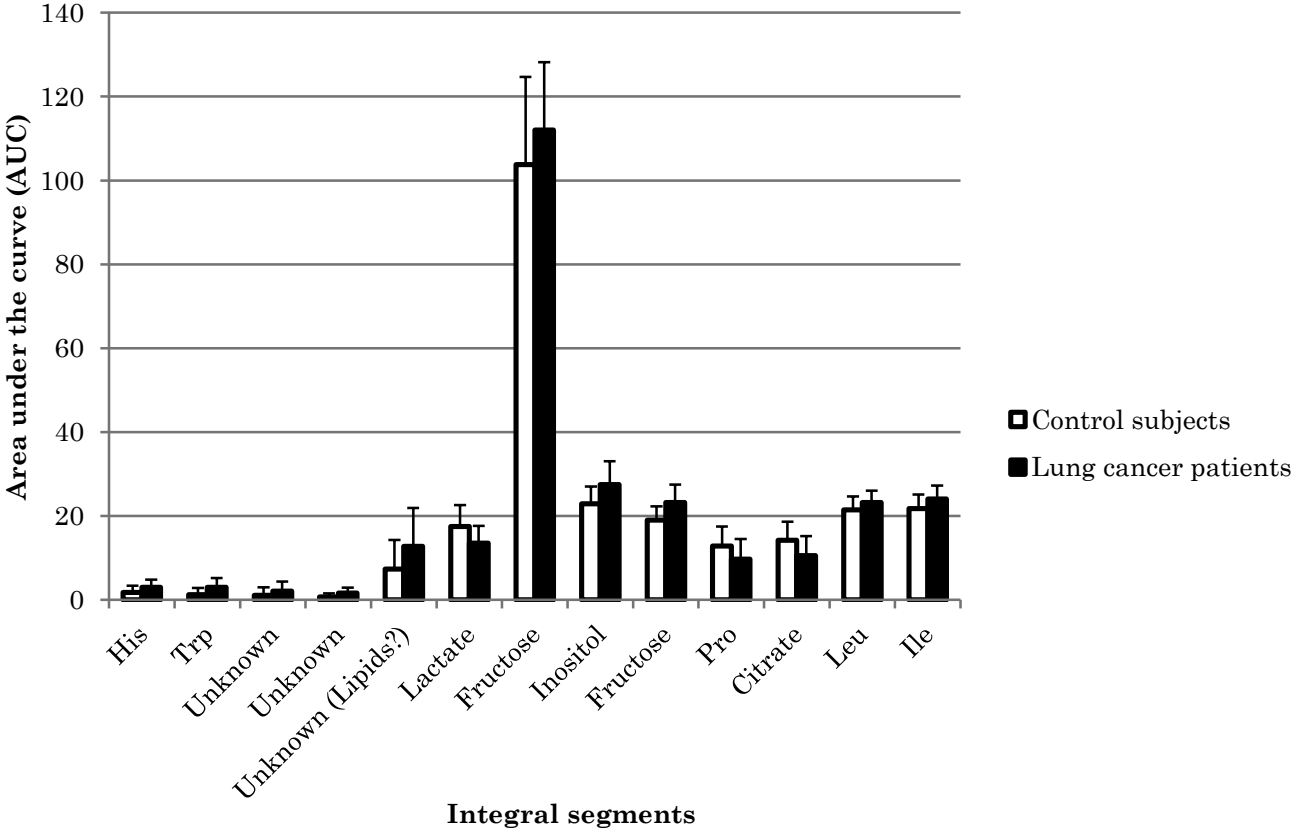
■ = Control subjects
■ = Lung cancer patients



Specificity: 92% (72/78)

Sensitivity: 83% (65/78)

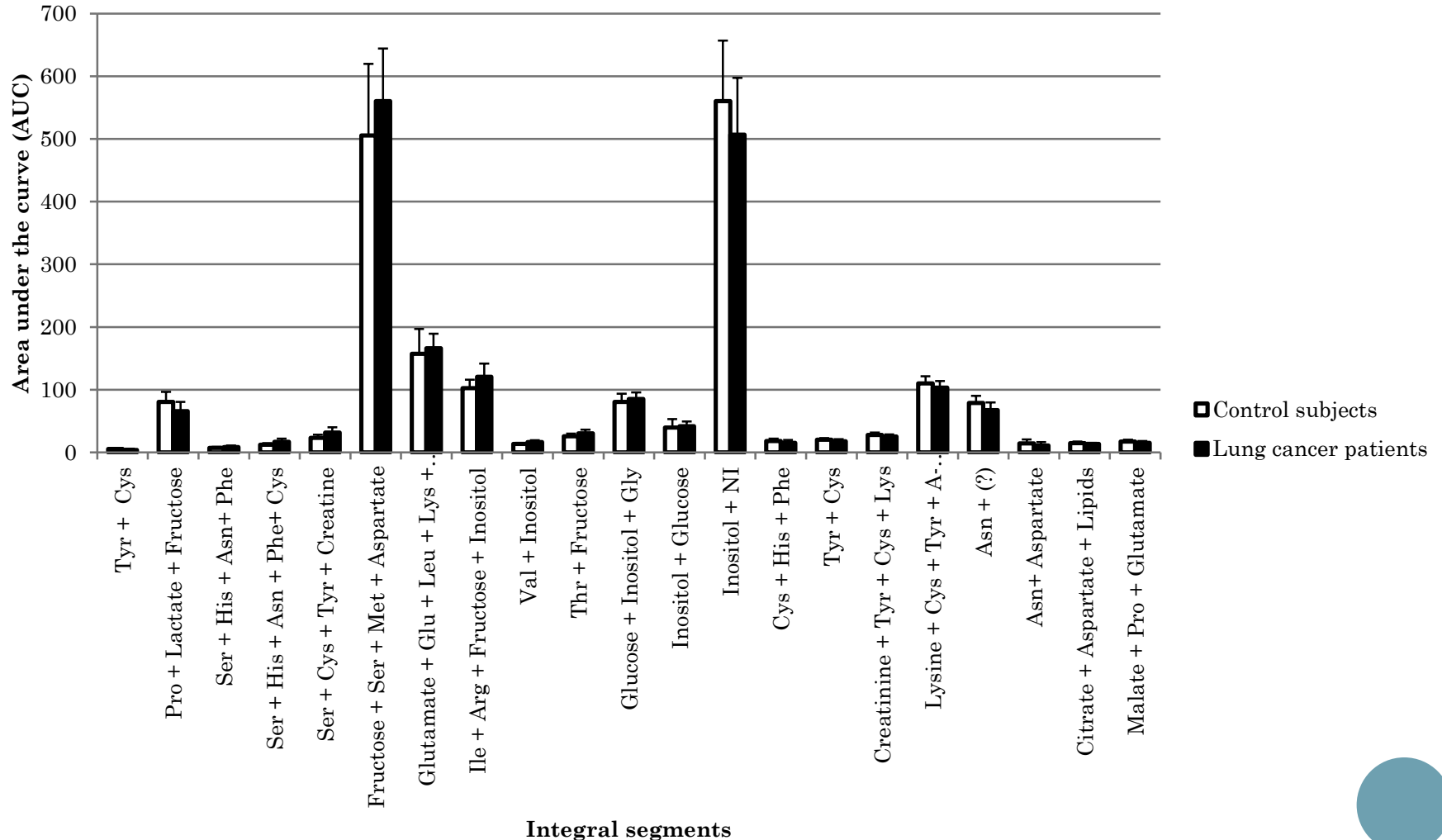
SIGNIFICANTLY DIFFERENT INTEGRAL SEGMENTS



13 integral segments with peaks of one metabolite



SIGNIFICANTLY DIFFERENT INTEGRAL SEGMENTS

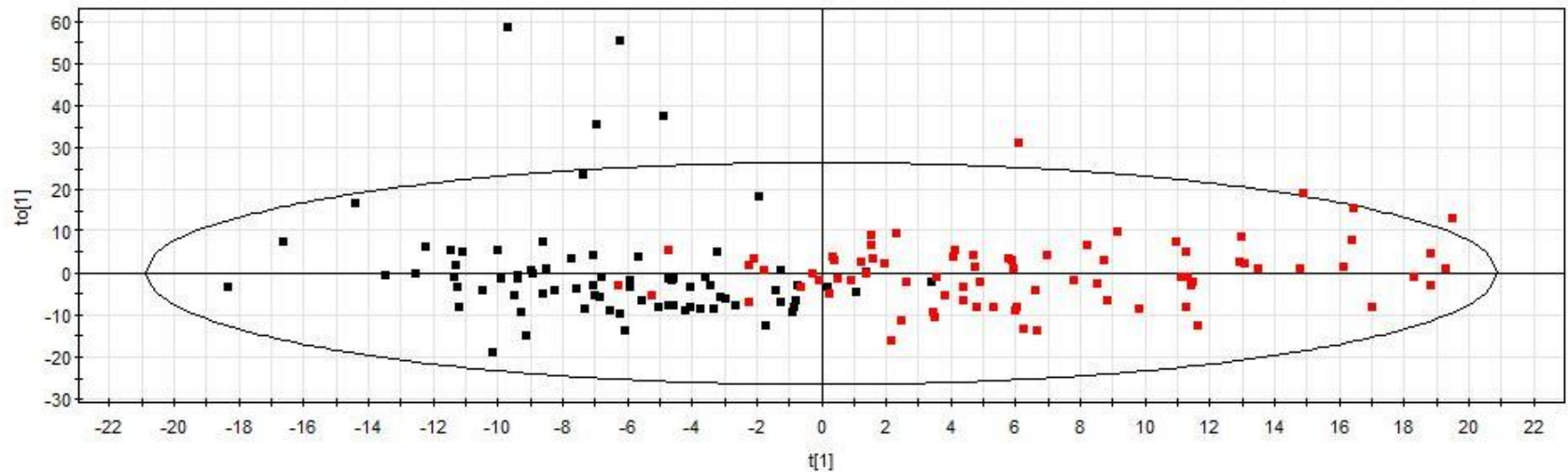


21 integral segments with peaks of 2 or more metabolites



MULTIVARIATE DATA ANALYSIS - SIGNIFICANTLY DIFFERENT INTEGRAL SEGMENTS

■ = Control subjects
■ = Lung cancer patients



Specificity: 95% (74/78)

Sensitivity: 87% (68/78)



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FUTURE PERSPECTIVES

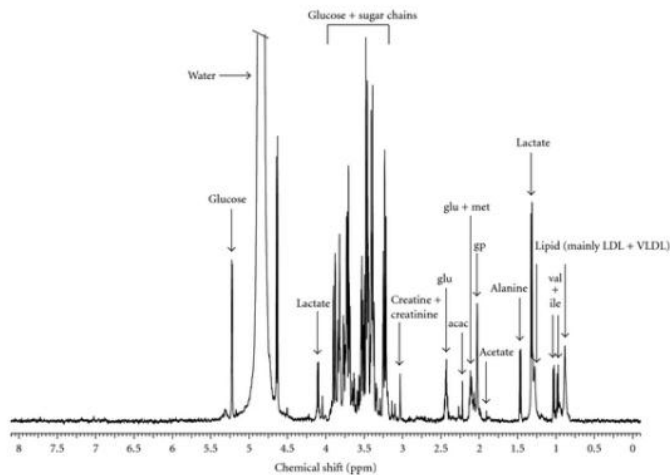
- Investigate the impact of gender, age, weight and smoking habits on the build model (classifier)
- **Validation** of the classifier in an independent study population
 - Aim: 250 lung cancer patients - 250 control subjects
- Current situation

| | |
|---|---------|
| Number of lung cancer patients (Limburg) | 92 |
| Number of lung cancer patients (Leuven) | 68 |
| Number of lung cancer patients | 160/250 |
| Number of control subjects | 119/250 |



FUTURE PERSPECTIVES

- Investigate whether the metabolic changes in the blood correlate with metabolic changes in the tumor, visible on a **PET/CT-scan**



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