





Machine learning based identification of high risk patients (WP8)

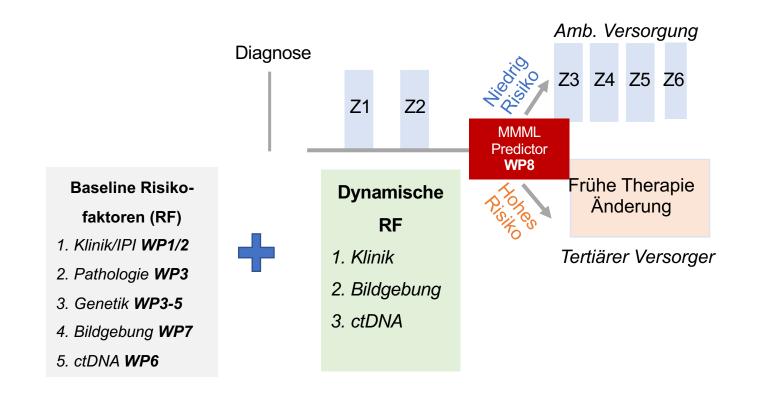
Für die Antragsteller des MMML Verbundes / GLA Rainer Spang

German Lymphoma Alliance e.V.

Arbeitsgruppen: Aggressive Lymphome, Pathologie, Biologie, Biometrie, Bildgebung

The MMML Predictor





Very High Risk



2y PFS: 67%

Prevalence: 24/100

Costs: 900 Euro

2y PFS: 10% Prevalence: 1/500

Costs: 999 Euro

2y PFS: 47%

Prevalence: 17/100

Costs: 31.100 Euro

2y survival: 49%

Prevalence: 15/100%

Costs: 1.419 Euro









Machine Learning based Risk Predictor



Age

Initial PET

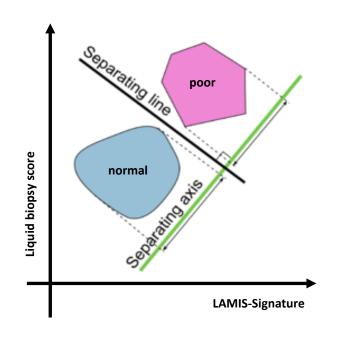
Standard Pathology

Expression Signatures

Genomics Host/Somatic

Liquid biopsies

Second PET/CT



Data Integration

2y survival < 50%

< 1500 Euro

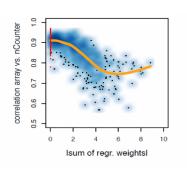
Prediction Accuracy

Validation

Umsetzungsstrategien



$$y_i = \beta_0 + \sum_j \beta_j (x_{ij} + \gamma_i)$$
$$y_i = \beta_0 + \sum_j \beta_j x_{ij} + \gamma_i \sum_j \beta_j$$



Our work horse is Zero-Sum Regression a more robust derivative of the **LASSO**

Altenbuchinger et al. Bioinformatics 2019a Altenbuchinger et al. Bioinformatics 2019b Staiger et al. Leukemia 2019 (LAMIS)

Work Package

Plan A

Plan B/Improvements

Data Integration

Late Integration

Intermediate Integration (train on residuals)

2y survival < 50%

Binary Zero-Sum Regression

Constrained Coordinate Descent (Cox Zero-Sum)

< 1500 Euro

Weighted L1-Penalization

Nested Monotone Regression

Constrained Coordinate Descent (Zero-Sum)

Prediction Accuracy

Validation/Leipzig **Independent Cohort**

Risikoeinschätzung



Fall: 04-YB13-71641b

Hoch Risiko: 91,3%

Genexpression: 75,9%

Liquid Biopsy: 99,9%

Pathologie: 98,1%

Fall: X14-9950-AZG9c

Hoch Risiko: 1,2%

Genexpression: 0,9%

Liquid Biopsy: 3,0%

Pathologie: 5,8%

Fall: 66-9UJ50-HPARe

Hoch Risiko: 69,9%

Genexpression: 18,1%

Liquid Biopsy: 99,6%

Pathologie: 2,0%





