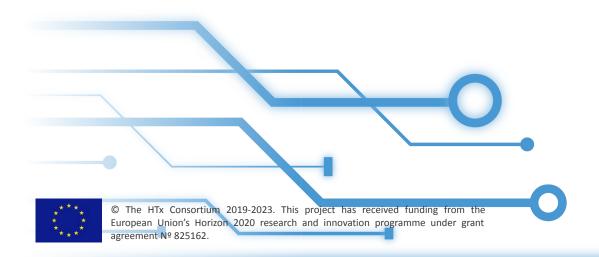


# Collinearity in Prognostic Models for Dysphagia

Artuur Leeuwenberg, PhD

Julius Center - University Medical Center Utrecht With Ewoud Schuit, Hans Reitsma, and Carl Moons





## Compared methods



- 1. Standard logistic regression (LR)
- 2. Lasso penalization
- 3. Ridge penalization
- 4. ElasticNet penalization
- 5. Principal component analysis logistic regression (PCLR)
- 6. Linear auto-encoder logistic regression (LAELR)
- 7. **Dropout** regularization
- 8. Non-negative constraints (**LRnn**)

# Simulation study



#### Case study (simulation study):

For **head and neck cancer patients**, we aim to predict the risk of experiencing **complications** 6 months **after radiotherapy**.

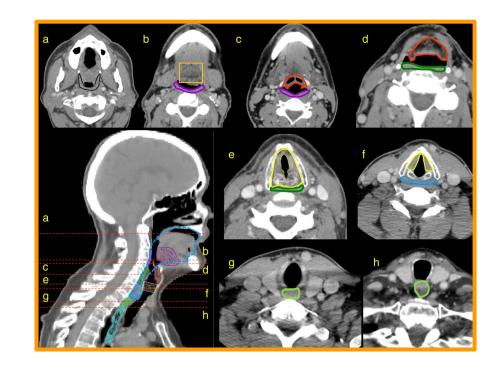
#### **Predictors** X:

Dosage per relevant organ

•

#### Outcome y:

• Dysphagia M<sub>6</sub>

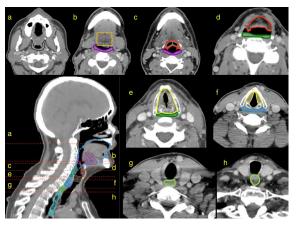




# High collinearity of dosage for OAR



Subm.L.Dm	-0.25											
Subm.R.Dm	-0.27	0.88										
Parotid.L.Dm	-0.26	0.85	0.72									
Parotid.R.Dm	-0.28	0.69	0.83	0.74								
PCM.Sup.Dm	-0.28	0.85	0.86	0.82	0.81							
PCM.Med.Dm	-0.25	0.87	0.89	0.74	0.72	0.79						
PCM.Inf.Dm	0.11	-0.11	-0.13	-0.11	-0.12	-0.35	0.11					
Supraglottic.Dm	-0.08	0.36	0.38	0.27	0.27	0.12	0.60	0.69				ı
OralCavity.Ext.Dm	-0.28	0.83	0.85	0.80	0.80	0.95	0.73	-0.40	0.07			
GlotticArea.Dm	0.15	-0.33	-0.34	-0.31	-0.31	-0.56	-0.16	0.89	0.54	-0.60	1	
DYSFAGIE.BSL.1	-0.04	0.24	0.25	0.25	0.21	0.24	0.23	-0.06	0.03	0.28	-0.12	
DYSFAGIE.BSL.2	-0.09	0.18	0.17	0.23	0.23	0.17	0.20	0.06	0.13	0.21	-0.01	-0.10
	AGE	Subm.L.Dm	Subm.R.Dm	Parotid.L.Dm	Parotid.R.Dm	PCM.Sup.Dm	PCM.Med.Dm	PCM.Inf.Dm	Supraglottic.Dm	OralCavity.Ext.Dm	GlotticArea.Dm	DYSFAGIE.BSL.1



# Simulation setup



Two levels of collinearity: 2 settings.

- Multivariate Gaussian to simulate predictor (cov from real data of 750 pts.)
- True predictor-outcome relation: Ridge fit on real data of 750 pts.
- ~600 training samples, 10.000 test samples, 100 repeats

# Results: spot the differences





VIF=5 (EPV:6)



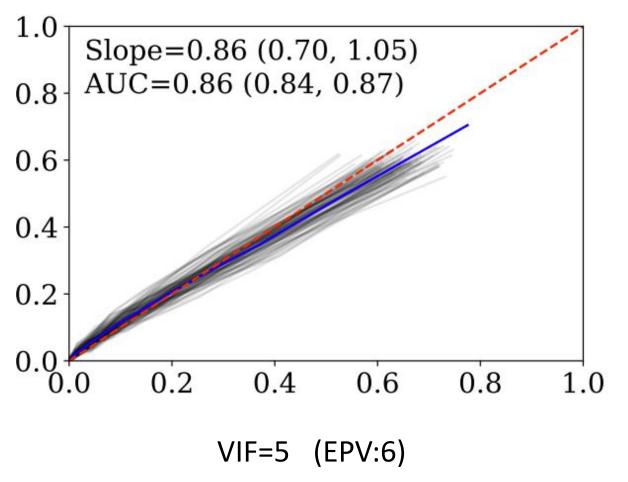
© The HTx Consortium 2019-2023. This project has received funding from the European Union's Horizon 2020 research and innovation programme under grant agreement Nº 825162.

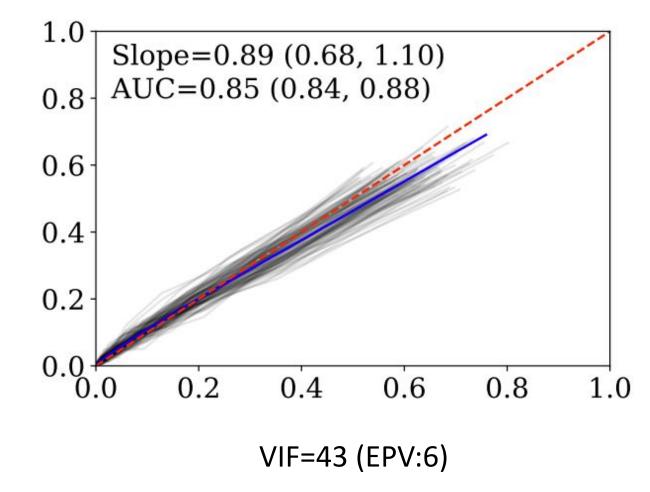


VIF=43 (EPV:6)

## Results: LR



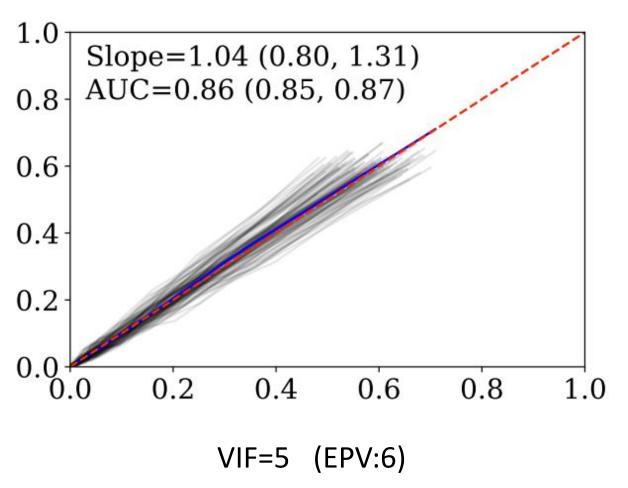


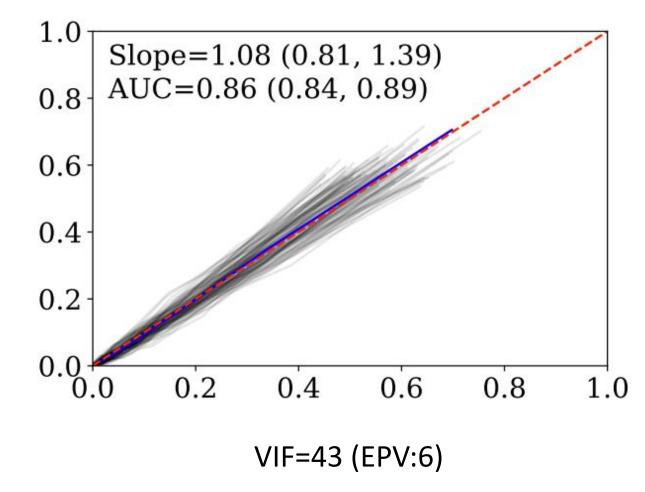




## Results: Lasso



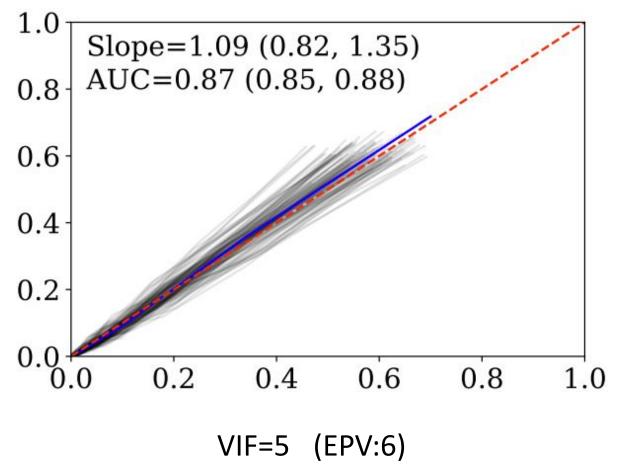


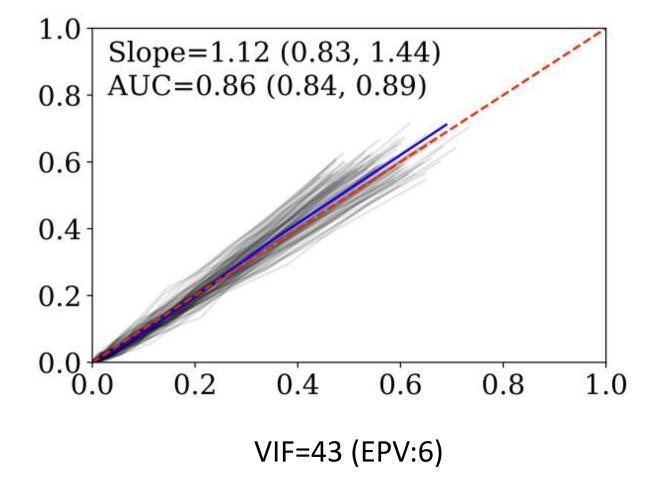




# Results: Ridge



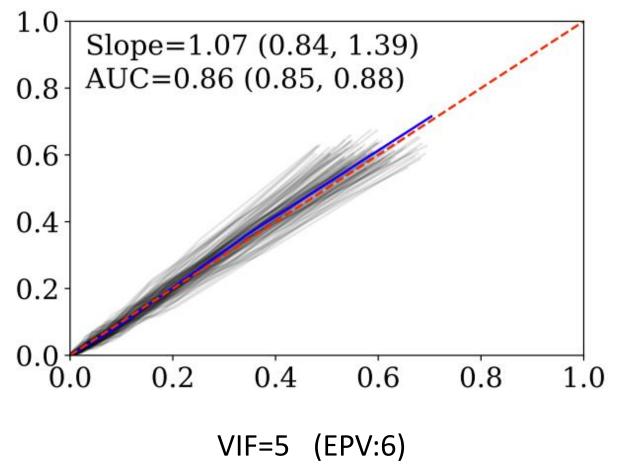


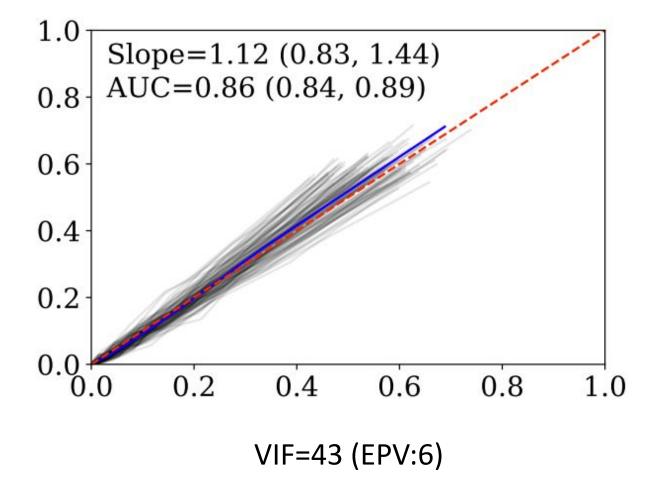




#### Results: ElasticNet



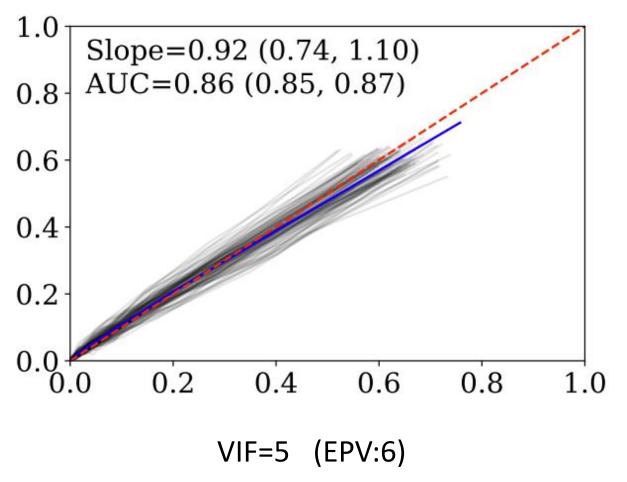


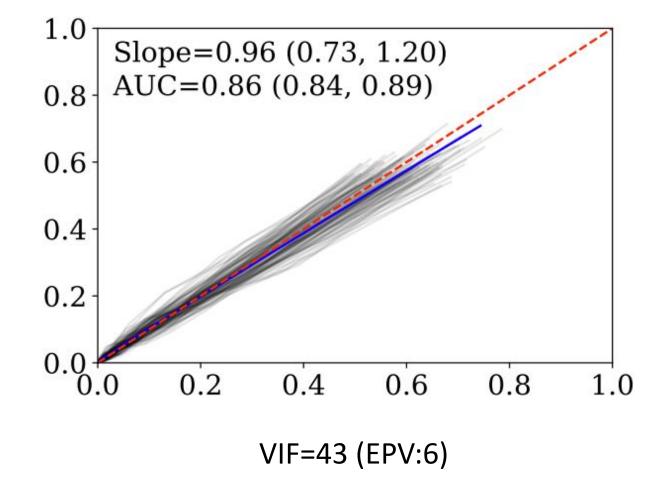




### Results: PCLR



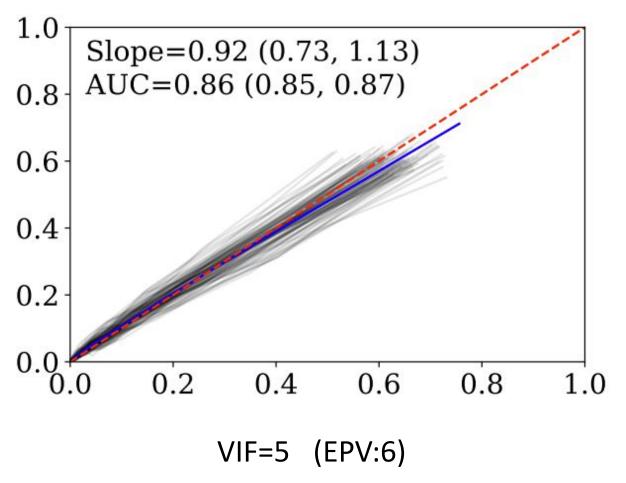


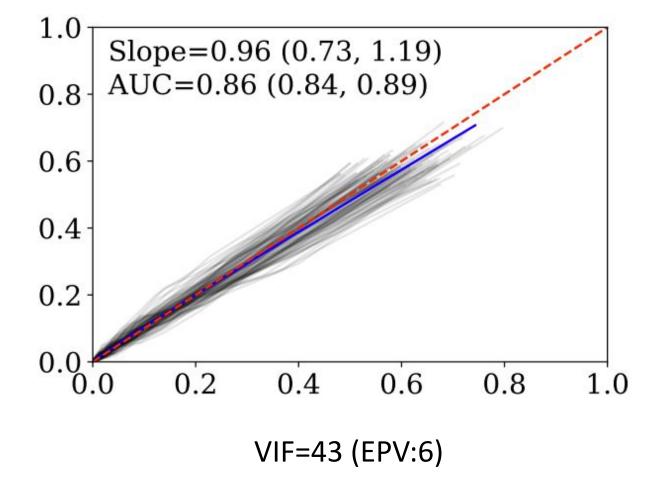




## Results: LAELR



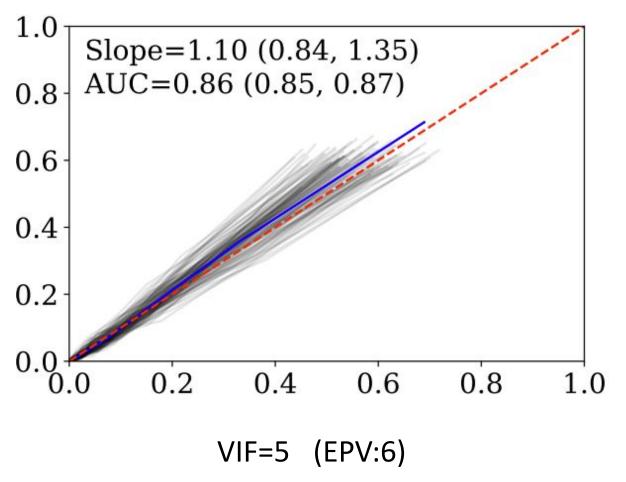


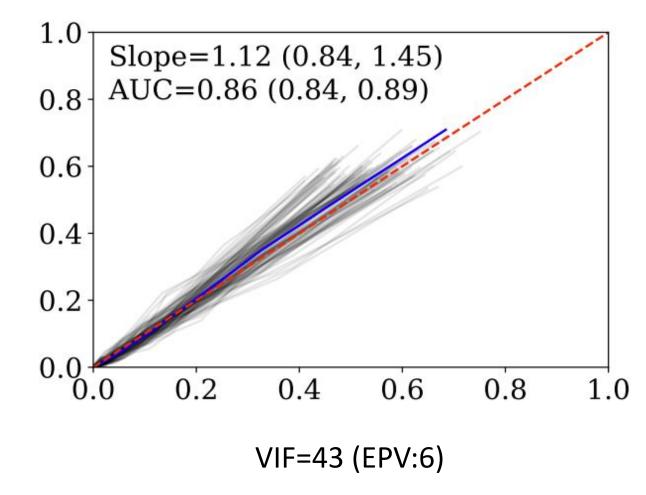




# Results: Dropout



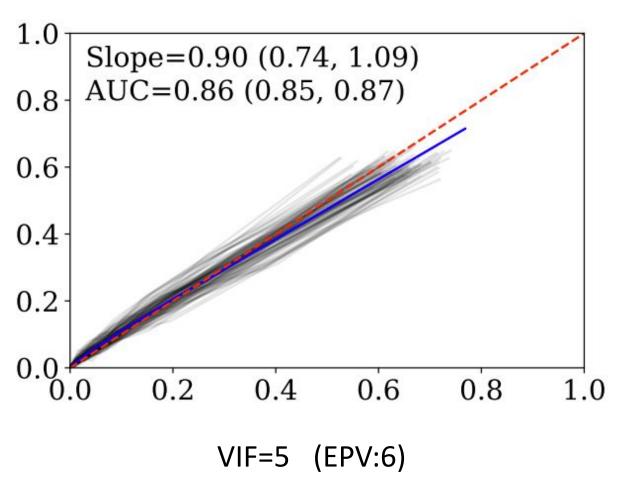


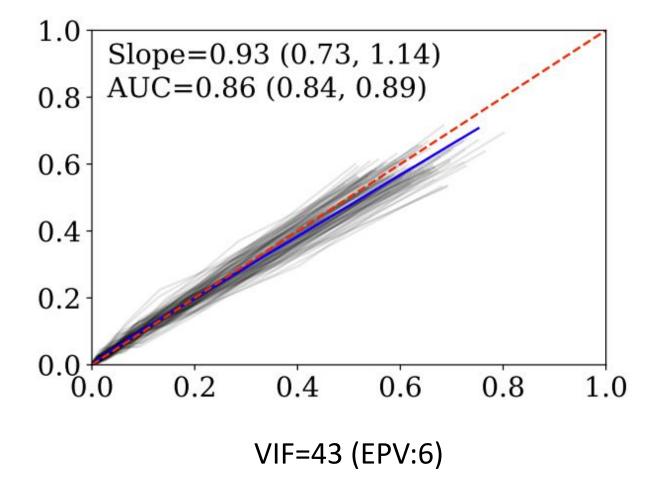




# Results: LR<sub>NN</sub>







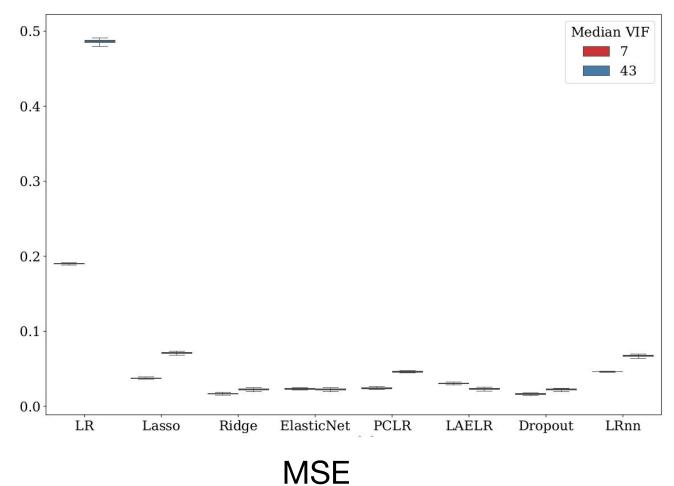


### Confirmation



Collinearity did not impact predictive performance in the simulations.

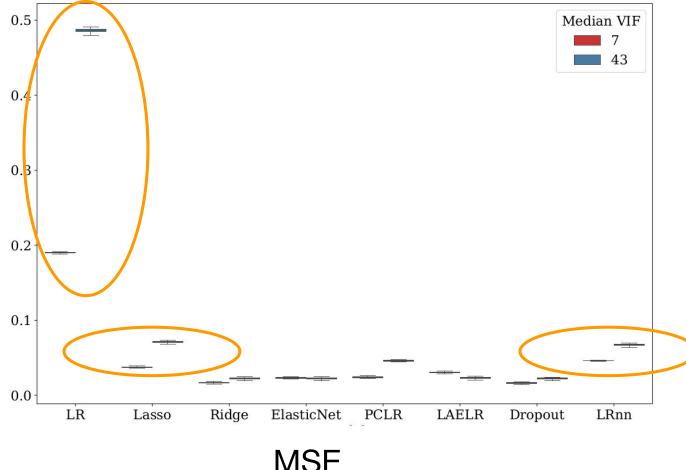




How far are the estimated coefficients from the true coefficients?





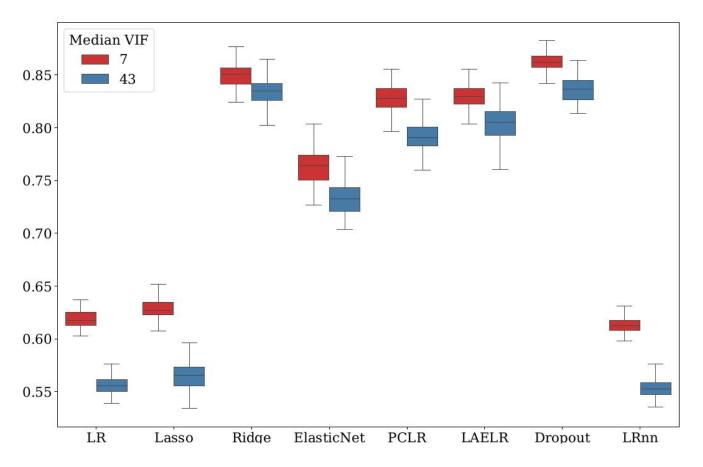


How far are the estimated coefficients from the true coefficients?

**MSE** 







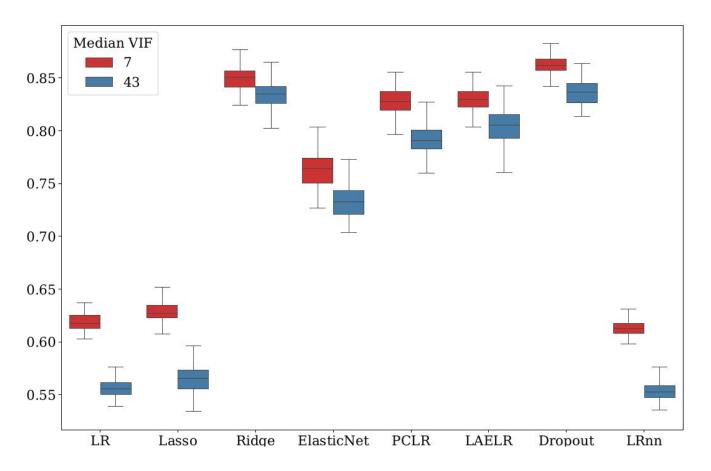
What percentage of coefficients is expected to get the same sign when repeating the experiment?

Model A	Model B
0	0.45
0.33	0.22
-0.13	0.34

#### **Expected Percentage of Similar Coefficients**







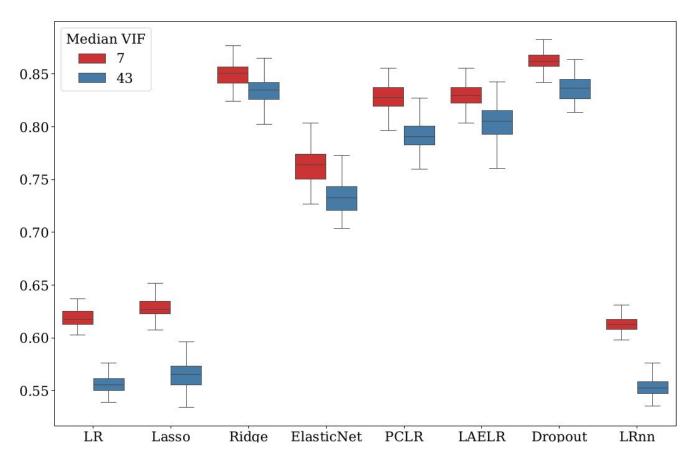
What percentage of coefficients is expected to get the same sign when repeating the experiment?

Model A	Model B
0	+
+	+
-	+

#### **Expected Percentage of Similar Coefficients**







What percentage of coefficients is expected to get the same sign when repeating the experiment?

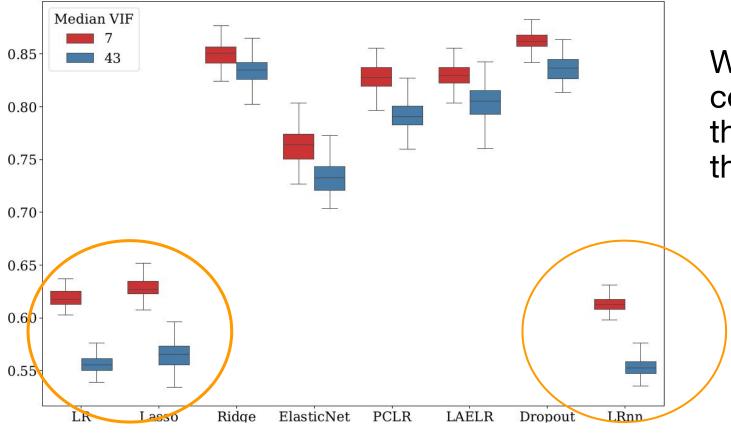
Model A	Model B
0	+
+	+
-	+

#### **Expected Percentage of Similar Coefficients**









What percentage of coefficients is expected to get the same sign when repeating the experiment?

#### Expected Percentage of Similar Coefficients



#### Conclusions



(in presence of high collinearity)

#### Predictive performance:

Similar across methods.

#### Coefficient estimation:

For mostly LR, but also Lasso and LRnn

- Increase in MSE
- Less consistent selection of coefficients

# Acknowledgements







Ewoud Schuit

Maarten van Smeden

Hans Reitsma

Carl Moons

Lisa van den Bosch Arjen van der Schaaf Hans Langendijk