COVID-19 Baseline Risk Score Analysis Report $$_{\rm Mock ENSEMBLE\ Study}$$

USG COVID-19 Response Biostatistics Team

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Chapter 1

Baseline Risk Score (Proxy for SARS-CoV-2 Exposure)

Table 1.1: Variables considered for risk score analysis.

| Variable.Name | Definition | Total.missing.values | Comment |
|-------------------------|---|----------------------|---------|
| EthnicityHispanic | Indicator ethnicity = Hispanic (1 = Hispanic, $0 = \text{complement}$) | 0/19462 (0.0%) | NA |
| EthnicityNotreported | Indicator ethnicity = Not reported (1 = Not reported, 0 = complement) | $0/19462 \ (0.0\%)$ | NA |
| EthnicityUnknown | Indicator ethnicity = Unknown (1 = Unknown, 0 = complement) | 0/19462 (0.0%) | NA |
| Black | Indicator race = Black (1=Black, 0=complement) | $0/19462 \ (0.0\%)$ | NA |
| Asian | Indicator race = Asian (1=Asian, 0=complement) | $0/19462 \ (0.0\%)$ | NA |
| NatAmer | Indicator race = American Indian or Alaska Native (1=NatAmer, 0=complement) | 0/19462 (0.0%) | NA |
| Multiracial | Indicator race = Multiracial (1=Multiracial, 0=complement) | 0/19462 (0.0%) | NA |
| Notreported | Indicator race = Not reported (1=Notreported, 0=complement) | 0/19462 (0.0%) | NA |
| Unknown | Indicator race = unknown (1=Unknown, 0=complement) | 0/19462 (0.0%) | NA |
| URMforsubcohortsampling | Indicator of under-represented minority (1=Yes, 0=No) | $0/19462 \ (0.0\%)$ | NA |
| HighRiskInd | Baseline covariate indicating >= 1 Co-existing conditions (1=yes, 0=no, NA=missing) | 0/19462 (0.0%) | NA |
| HIVinfection | Indicator HIV infected at enrollment (1=infected, 0=not infected) | 0/19462 (0.0%) | NA |
| Sex | Sex assigned at birth (1=female, 0=male/undifferentiated/unknown | $0/19462 \ (0.0\%)$ | NA |
| Country.X1 | Dummy indicator country = Argentina (1 = Argentina, $0 = \text{complement}$) | 0/19462 (0.0%) | NA |
| | | | |

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Table 1.1: Variables considered for risk score analysis. (continued)

| Variable.Name | Definition | Total.missing.values |
|-------------------|--|----------------------|
| Country.X2 | Dummy indicator country = Brazil (1 = Brazil, 0 = complement) | 0/19462 (0.0%) |
| Country.X3 | Dummy indicator country = Chile $(1 = \text{Chile}, 0 = \text{complement})$ | $0/19462 \ (0.0\%)$ |
| Country.X4 | Dummy indicator country = Columbia (1 = Columbia, 0 = complement) | $0/19462 \ (0.0\%)$ |
| Country.X5 | Dummy indicator country = Mexico (1 = Mexico, 0 = complement) | $0/19462 \ (0.0\%)$ |
| Country.X6 | Dummy indicator country = Peru (1 = Peru, 0 = complement) | $0/19462 \ (0.0\%)$ |
| Country.X7 | Dummy indicator country = South Africa $(1 = South Africa, 0 = complement)$ | $0/19462 \ (0.0\%)$ |
| Region.X1 | Dummy indicator region = Latin America (1 = Latin America, $0 = \text{complement}$) | $0/19462 \ (0.0\%)$ |
| Region.X2 | Dummy indicator country = Southern Africa (1 = Southern Africa, 0 = complement) | $0/19462 \ (0.0\%)$ |
| CalDtEnrollIND.X1 | Dummy indicator variable representing enrollment occurring between 4-8 weeks periods of first subject enrolled (1 = Enrollment between 4-8 weeks, 0 = complement). | 0/19462 (0.0%) |
| Age | Age at enrollment in years (integer >= 18, NA=missing). Note that the randomization strata included Age 18-59 vs. Age >= 60. | 0/19462 (0.0%) |
| ВМІ | BMI at enrollment (Ordered categorical 1,2, 3, 4, NA=missing); 1 = Underweight BMI < 18.5; 2 = Normal BMI 18.5 to < 25; 3 = Overweight BMI 25 to < 30; 4 = Obese BMI >= 30 | 0/19462 (0.0%) |

Note:

Variables with more than 5% missing values were dropped from analysis; missing values for other variables values for other variables and Indicator variables not meeting the threshold, such that under the null of not a risk factor there were less the subgroup with value 1 (or 0), were dropped from analysis.

Table 1.2: All learner-screen combinations (14 in total) used as input to the Superlearner.

| Learner | Screen* |
|---------|---|
| SL.mean | all |
| SL.glm | all glmnet univar_logistic_pval highcor_random |

Note:

*Screen details:

all: includes all variables

glmnet: includes variables with non-zero coefficients in the standard implementation of SL.glmnet that optimizes the lasso tuning parameter via cross-validation

univar_logistic_pval: Wald test 2-sided p-value in a logistic regression model $<0.10\,$

high cor_random: if pairs of quantitative variables with Spearman rank correlation > 0.90, select one of the variables at random

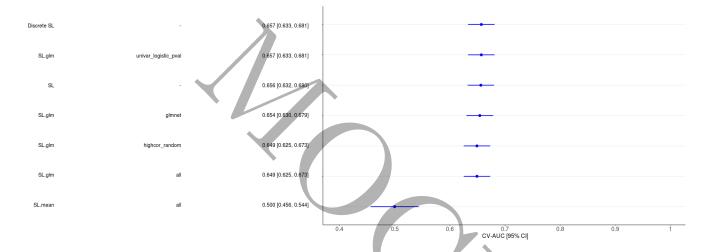


Figure 1.1: Cross-validated AUC (95% CI) of algorithms for predicting COVID-19 disease status starting 7 days after Day 29.

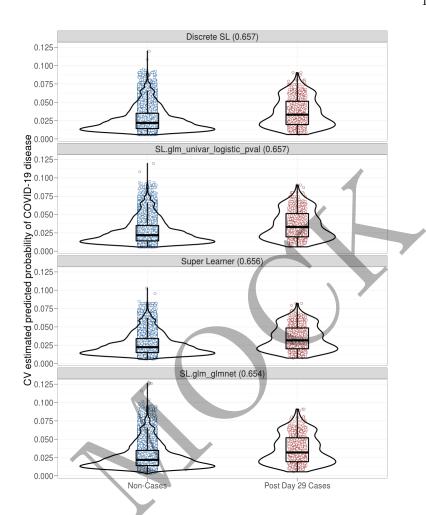


Figure 1.2: CV-estimated predicted probabilities of COVID-19 disease 7 days after Day 29 by case/control status for top 2 learners, SuperLearner and Discrete SL.

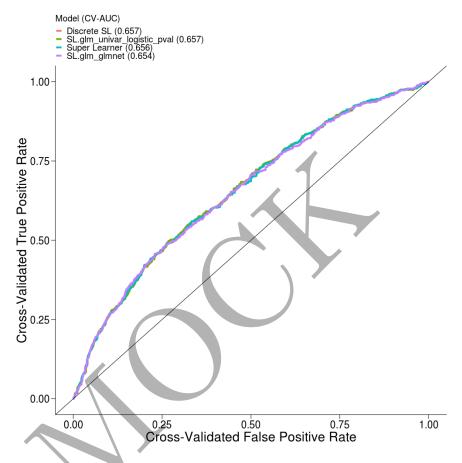


Figure 1.3: ROC curves based off CV-estimated predicted probabilities for the top 2 learners, Superlearner and Discrete SL.

Table 1.3: Weights assigned by Superlearner.

| Learner | Screen | Weight |
|---------|---------------------------------|--------|
| SL.glm | screen_univariate_logistic_pval | 0.942 |
| SL.mean | screen_all | 0.058 |
| SL.glm | screen_all | 0.000 |
| SL.glm | screen_glmnet | 0.000 |
| SL.glm | screen_highcor_random | 0.000 |

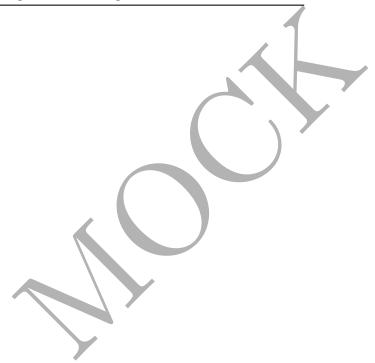


Table 1.4: Predictors in learners assigned weight > 0.0 by Superlearner.

| Learner | Screen | Weight | Predictors | Coefficient | Odds.Ratio |
|---------|--------------------------------------|--------|-------------|-------------|------------|
| SL.glm | screen_univariate_logistic_pval | 0.942 | (Intercept) | -3.760 | 0.023 |
| SL.glm | screen_univariate_logistic_pval | 0.942 | HighRiskInd | 0.476 | 1.609 |
| SL.glm | screen_univariate_logistic_pval | 0.942 | Country.X2 | 0.097 | 1.101 |
| SL.glm | $screen_univariate_logistic_pval$ | 0.942 | Age | 0.343 | 1.409 |

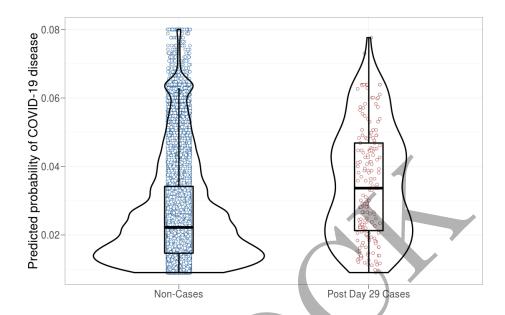


Figure 1.4: Superlearner predicted probabilities of COVID-19 disease in vaccinees 7 days after Day 29 by case/control status.

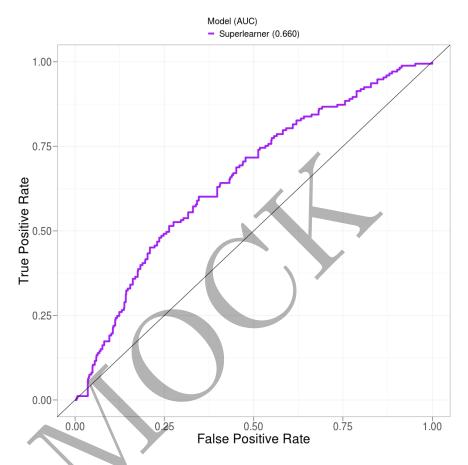


Figure 1.5: ROC curve based off Superlearner predicted probabilities in vaccinees.

Table 1.5: Cases per treatment arm prior to risk score analysis.

| Study-Arm | Non-Cases | Post-Day 29-Cases |
|-----------|-----------|-------------------|
| Placebo | 18743 | 516 |
| Vaccine | 19183 | 173 |

Table 1.6: Cases per treatment arm post risk score analysis.

| | | | - 4 |
|-----------|-----------|-------------------|-----|
| Study-Arm | Non-Cases | Post-Day 29-Cases | |
| Placebo | 18743 | 516 | - |
| Vaccine | 19183 | 173 | |
| | | | |



Chapter 2

Appendix

- This report was built from the CoVPN/correlates_reporting repository with commit hash 220ac78ab74ec2ca9ec2747c09a4f84e68765fda. A diff of the changes introduced by that commit may be viewed at https://github.com/CoVPN/correlates_reporting/commit/220ac78ab74ec2ca9ec2747c09a4f84e68765fda
- \bullet The sha256 hash sum of the raw input file, "COVID_ENSEMBLE_practicedata.csv": 847161e464e2488f2d36717254de9e0d885d56cbe7205a3d174d747b6cb828d8
- The sha256 hash sum of the processed file, "janssen_pooled_mock_data_processed.csv": c05d0a8b66052e68358fc5fb76888a80b0437b151ae016a1b42f0072ac94d1eb