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

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 = complement) | $0/19501 \ (0.0\%)$ | NA |
| EthnicityNotreported | Indicator ethnicity = Not reported (1 = Not reported, 0 = complement) | $0/19501 \ (0.0\%)$ | NA |
| EthnicityUnknown | Indicator ethnicity = Unknown (1 = Unknown, 0 = complement) | 0/19501 (0.0%) | NA |
| Black | Indicator race = Black (1=Black, 0=complement) | $0/19501 \ (0.0\%)$ | NA |
| Asian | Indicator race = Asian (1=Asian, 0=complement) | $0/19501 \ (0.0\%)$ | NA |
| NatAmer | Indicator race = American Indian or Alaska Native (1=NatAmer, 0=complement) | 0/19501 (0.0%) | NA |
| Multiracial | Indicator race = Multiracial (1=Multiracial, 0=complement) | 0/19501 (0.0%) | NA |
| Notreported | Indicator race = Not reported (1=Notreported, 0=complement) | 0/19501 (0.0%) | NA |
| Unknown | Indicator race = unknown (1=Unknown, 0=complement) | 0/19501 (0.0%) | NA |
| URMforsubcohortsampling | Indicator of under-represented minority (1=Yes, 0=No) | $0/19501 \ (0.0\%)$ | NA |
| HighRiskInd | Baseline covariate indicating >= 1 Co-existing conditions (1=yes, 0=no, NA=missing) | 0/19501 (0.0%) | NA |
| Sex | Sex assigned at birth (1=female, 0=male/undifferentiated/unknown | 0/19501 (0.0%) | NA |
| Age | Age at enrollment in years (integer $>= 18$, NA=missing). Note that the randomization strata included Age 18-59 vs. Age $>= 60$. | 0/19501 (0.0%) | NA |

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

| Variable.Name | Definition | Total.missing.values |
|----------------------------------|---|----------------------|
| BMI | BMI at enrollment (Ordered categorical 1,2, 3, 4, | 0/19501 (0.0%) |
| | NA=missing); $1 = \text{Underweight BMI} < 18.5$; $2 =$ | |
| | Normal BMI $18.5 \text{ to} < 25; 3 = \text{Overweight BMI } 25 \text{ to}$ | |
| | < 30; 4 = Obese BMI >= 30 | |
| Country | Country of the study site of enrollment (0=United | $0/19501 \ (0.0\%)$ |
| | States, 1=Argentina,2=Brazil, 3=Chile,4=Columbia, | |
| | 5=Mexico, 6=Peru, 7=South Africa) | |
| HIVinfection | Indicator HIV infected at enrollment (1=infected, | $0/19501 \ (0.0\%)$ |
| | 0=not infected) | |
| ${\bf Calendar Date Enrollment}$ | Date variable (used to control for calendar time trends | $0/19501 \ (0.0\%)$ |
| | in COVID incidence). Coded as number of days since | , , , |
| | first person enrolled until the ppt is enrolled. | |

Note:

Variables with more than 5% missing values were dropped from analysis; missing values for other variables values for other variables values are 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_all | 0.833 |
| SL.glm | screen_glmnet | 0.167 |
| SL.mean | screen_all | 0.000 |
| SL.glm | $screen_univariate_logistic_pval$ | 0.000 |
| SL.glm | screen_highcor_random | 0.000 |



Screen Weight Predictors Coefficient Odds.Ratio Learner screen_all SL.glm 0.833 (Intercept) -19.397 0 SL.glmscreen_all 0.833 EthnicityHispanic -0.037 0.964 SL.glm 0.833 EthnicityNotreported 0.013 1.013 screen_all $_{\mathrm{SL.glm}}$ 0.833 ${\bf Ethnicity Unknown}$ $screen_all$ 0.05 1.051 SL.glm 0.833 -0.16 0.852screen_all SL.glmscreen_all 0.833 Asian -0.066 0.936 SL.glm screen_all 0.833NatAmer-0.056 0.945SL.glm 0.833 Multiracial-0.039 0.961screen_all $_{\mathrm{SL.glm}}$ $screen_all$ 0.833 Notreported -1.403 0.2460.833 -0.056 SL.glm $screen_all$ Unknown 0.946URMforsubcohortsampling SL.glm $screen_all$ 0.833 0.063 1.065 SL.glmscreen_all 0.833 HighRiskInd 0.3791.461 SL.glm screen all 0.833 Sex 12.071 174754.678 $_{\mathrm{SL.glm}}$ $screen_all$ 0.833Age 0.3541.425SL.glm 0.833 BMI -0.046 0.955 screen all -0.017 0.983 SL.glm screen_all 0.833 Country SL.glm screen_all 0.833HIVinfection 0.0111.011 ${\bf Calendar Date Enrollment}$ 1.027SL.glm $screen_all$ 0.8330.027 $_{\mathrm{SL.glm}}$ screen_glmnet 0.167(Intercept) -19.395 SL.glm $screen_glmnet$ 0.167 EthnicityNotreported 0.015 1.015 SL.glm $screen_glmnet$ 0.167EthnicityUnknown 0.052 1.053 SL.glm screen_glmnet 0.167Black -0.136 0.873 $_{\mathrm{SL.glm}}$ 0.167 Asian -0.0670.936 screen_glmnet SL.glm screen_glmnet 0.167NatAmer-0.055 0.947-0.04 0.961 SL.glm $screen_glmnet$ 0.167Multiracial $_{\mathrm{SL.glm}}$ screen_glmnet 0.167Notreported -1.4030.246SL.glm Unknown -0.056 0.946 $screen_glmnet$ 0.167 $_{\mathrm{SL.glm}}$ $screen_glmnet$ 0.167URMforsubcohortsampling 0.031.031 SL.glm 0.167 HighRiskInd 0.379 1.461 screen_glmnet 174555.444 SL.glm0.167 Sex 12.07screen_glmnet SL.glm screen_glmnet 0.1670.3541.425Age $_{\mathrm{BMI}}$ -0.046 0.955 SL.glm screen_glmnet 0.167 $_{\mathrm{SL.glm}}$ $screen_glmnet$ 0.167 Country -0.013 0.987 SL.glm screen_glmnet 0.167HIVinfection0.011 1.011 ${\bf Calendar Date Enrollment}$ $_{\mathrm{SL.glm}}$ $screen_glmnet$ 0.167 0.0261.027 SL.glm $screen_glmnet$ 0.167(Intercept) -19.395 0 0.015 1.015 SL.glm $screen_glmnet$ 0.167 EthnicityNotreported $_{\mathrm{SL.glm}}$ $screen_glmnet$ 0.167Ethnicity Unknown0.0521.053 SL.glm 0.167 Black -0.136 0.873 screen glmnet -0.067 0.936 SL.glm screen_glmnet 0.167 Asian SL.glm screen_glmnet 0.167NatAmer-0.055 0.947SL.glm0.167 Multiracial -0.04 0.961 screen_glmnet $_{\mathrm{SL.glm}}$ screen_glmnet 0.167Notreported -1.403 0.246

0.167

0.167

screen glmnet

 $screen_glmnet$

Unknown

URMforsubcohortsampling

-0.056

0.03

0.946

1.031

SL.glm

SL.glm

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

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

| Learner | Screen | Weight | Predictors | Coefficient | Odds.Ratio |
|---------|---------------|--------|------------------------|-------------|------------|
| SL.glm | screen_glmnet | 0.167 | HighRiskInd | 0.379 | 1.461 |
| SL.glm | screen_glmnet | 0.167 | Sex | 12.07 | 174555.444 |
| SL.glm | screen_glmnet | 0.167 | Age | 0.354 | 1.425 |
| SL.glm | screen_glmnet | 0.167 | BMI | -0.046 | 0.955 |
| SL.glm | screen_glmnet | 0.167 | Country | -0.013 | 0.987 |
| SL.glm | screen_glmnet | 0.167 | HIVinfection | 0.011 | 1.011 |
| SL.glm | screen_glmnet | 0.167 | CalendarDateEnrollment | 0.026 | 1.027 |



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.

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

| Study-Arm | Non-Cases | Post-Day 29-Cases |
|-----------|-----------|-------------------|
| Placebo | 18756 | 521 |
| Vaccine | 19198 | 170 |

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

| Study-Arm | Non-Cases | Post-Day | 29-Cases |
|-----------|-----------|----------|----------|
| Placebo | 18756 | | 521 |
| Vaccine | 19198 | | 170 |

Chapter 2

Appendix

- This report was built from the CoVPN/correlates_reporting repository with commit hash 51bf6dfc91119f05596750ff15382361cf2fb5e6. A diff of the changes introduced by that commit may be viewed at https://github.com/CoVPN/correlates_reporting/commit/51bf6dfc91119f05596750ff15382361cf2fb5e6
- The sha256 hash sum of the raw input file, "COVID_ENSEMBLE_practicedata.csv": 186c6fefe0d7ee781c3b0bf5cedee3686398eb67ddc3da19dfa0c9bf3ad88fcc
- The sha256 hash sum of the processed file, "janssen_pooled_mock_data_processed.csv": 168c3beacf95500be32a6d6866656b93759de1680e60d61f8a34341b2a4458a6