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

USG COVID-19 Response Biostatistics Team

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Chapter 1

Disclaimers

- The data presented in the analysis originated from the Moderna Sponsored mRNA-1273-P301 clinical study and are provided to NIAID in accordance with Clinical Trial Agreement between the parties. The study was funded in part by BARDA under Government Contract No. 75A50120C00034
- The preliminary immunogenicity data presented here do not reflect the Sponsors statistical analysis plan and therefore should not be interpreted as a protocol defined read-out of the clinical study.
- These data are not to be disclosed without written permission of Moderna.



Chapter 2

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

Table 2.1: Variables considered for risk score analysis.

Variable.Name	Definition	Total.missing.values	Comments
MinorityInd	Baseline covariate underrepresented minority status (1=minority,	0/30000 (0.0%)	NA
EthnicityHispanic	0=non-minority) Indicator ethnicity = Hispanic (0 =	0/30000 (0.0%)	NA
EthnicityNotreported	Non-Hispanic) Indicator ethnicity = Not reported (0 =	0/30000 (0.0%)	NA
EthnicityUnknown	Non-Hispanic) Indicator ethnicity = Unknown (0 =	0/30000 (0.0%)	NA
Black	Non-Hispanic) Indicator race = Black $(0 = White)$	0/30000 (0.0%)	NA
Asian	Indicator race $=$ Asian (0 = White)	0/30000 (0.0%)	NA
NatAmer	Indicator race = American Indian or Alaska Native (0 = White)	0/30000 (0.0%)	NA
PacIsl	Indicator race = Native Hawaiian or Other Pacific Islander (0 = White)	0/30000 (0.0%)	NA
Multiracial	Indicator race = Multiracial (0 = White)	0/30000 (0.0%)	NA
Other	Indicator race = Other (0 = White)	0/30000 (0.0%)	NA
Notreported	Indicator race = Not reported (0 = White)	0/30000 (0.0%)	NA
Unknown	Indicator race = unknown (0 = White)	0/30000 (0.0%)	NA
HighRiskInd	Baseline covariate high risk pre-existing condition (1=yes, 0=no)	0/30000 (0.0%)	NA
Sex	Sex assigned at birth (1=female, 0=male)	0/30000 (0.0%)	NA
Age	Age at enrollment in years, between 18 and 85	0/30000 (0.0%)	NA
BMI	BMI at enrollment (kg/m^2)	0/30000 (0.0%)	NA

Table 2.2: All learner-screen combinations (28 in total) used as input to the Superlearner.

Learner	Screen*
SL.mean	all
SL.glm	all glmnet univar_logistic_pval highcor_random
SL.glm.interaction	glmnet univar_logistic_pval highcor_random
SL.glmnet	all
SL.gam	glmnet univar_logistic_pval highcor_random
SL.xgboost	all
SL.ranger.imp	all

Note:

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 ${\bf r}$

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

^{*}Screen details:



Figure 2.1: Cross-validated AUC (95% CI) of algorithms for predicting COVID-19 disease status starting 7 days after Day 57.



Figure 2.2: CV-estimated predicted probabilities of COVID-19 disease 7 days after Day 57 by case/control status for top 2 learners, SuperLearner and Discrete $^{\rm CI}$



Figure 2.3: ROC curves based off CV-estimated predicted probabilities for the top 2 learners, Superlearner and Discrete SL.

Table 2.3: Weights assigned by Superlearner.

Learner	Screen	Weight
SL.gam	screen_highcor_random	0.488
SL.glmnet	screen_all	0.253
SL.ranger.imp	screen_all	0.139
SL.glm.interaction	$screen_glmnet$	0.120
SL.mean	screen_all	0.000
SL.glm	screen_all	0.000
SL.xgboost	screen_all	0.000
SL.glm	screen_glmnet	0.000
SL.glm	screen_univariate_logistic_pval	0.000
SL.glm	screen_highcor_random	0.000
SL.glm.interaction	screen_univariate_logistic_pval	0.000
SL.glm.interaction	screen_highcor_random	0.000
SL.gam	screen_glmnet	0.000
SL.gam	screen_univariate_logistic_pval	0.000

Predictors Coefficient Odds.Ratio Importance Learner Screen Weight screen_highcor_random SL.gam 0.488(Intercept) -2.8730.057 NASL.gam screen_highcor_random 0.488s(Age,2) 0.8292.290 NAs(BMI,2) 0.955 NASL.gam screen_highcor_random 0.488-0.046 0.950 NΑ $_{\mathrm{SL.gam}}$ screen_highcor_random 0.488MinorityInd -0.0520.488EthnicityHispanic 0.0041.004 NASL.gam screen_highcor_random $_{\mathrm{SL.gam}}$ screen_highcor_random 0.488EthnicityNotreported 0.043 1.044 NASL.gamscreen_highcor_random 0.488EthnicityUnknown -0.073 0.930 NASL.gam0.488Black0.0041.004 NAscreen highcor random $_{\mathrm{SL.gam}}$ screen_highcor_random 0.488Asian 0.069 1.071 NASL.gamscreen_highcor_random 0.488NatAmer 0.0151.016 NAPacIsl SL.gam screen_highcor_random 0.488-0.013 0.987NΑ $_{\mathrm{SL.gam}}$ screen_highcor_random 0.488Multiracial -0.046 0.955 NASL.gam screen highcor random 0.488Other 0.0641.066 NA $_{\mathrm{SL.gam}}$ screen_highcor_random 0.488Notreported 0.003 1.003 NASL.gam screen_highcor_random 0.488Unknown 0.003 1.003 NAHighRiskInd-0.026 0.974 NASL.gam screen_highcor_random 0.488 SL.gam screen_highcor_random 0.488Sex-0.001 0.999 NA0.253 -2.665 0.070 NΑ SL.glmnet screen all (Intercept) SL.glmnetscreen_all 0.253MinorityInd 0.000 1.000 NASL.glmnet 0.253EthnicityHispanic 0.000 1.000 NΑ screen all SL.glmnet screen_all 0.253EthnicityNotreported 0.000 1.000 NA0.253 EthnicityUnknown NASL.glmnet screen_all 0.000 1.000 0.253 1.000 SL.glmnet Black 0.000 NAscreen all SL.glmnetscreen_all 0.253Asian 0.000 1.000 NA0.253NatAmer 1.000 NASL.glmnet screen all 0.000 SL.glmnetscreen_all 0.253PacIsl 0.000 1.000 NASL.glmnet 0.253Multiracial 0.000 1.000 NAscreen all SL.glmnetscreen_all 0.253Other 0.000 1.000 NASL.glmnet 0.253 Notreported 0.000 1.000 NAscreen_all 0.253 1.000 SL.glmnetscreen all Unknown 0.000 NASL.glmnetscreen_all 0.253HighRiskInd 0.000 1.000 NA0.253 0.000 1.000 NASL.glmnet screen all Sex SL.glmnetscreen all 0.253Age 0.4441.559 NASL.glmnetscreen_all 0.253 $_{\mathrm{BMI}}$ 0.000 1.000 NASL.ranger.imp screen all 0.139 MinorityInd NΑ NA9.100 SL.ranger.impscreen_all 0.139 EthnicityHispanic NANA9.685 0.139 EthnicityNotreported NΑ 9.247 SL.ranger.imp screen all NΑ SL.ranger.impscreen_all 0.139Ethnicity UnknownNANA6.908screen_all 0.139 Black NANA9.156 SL.ranger.imp 0.139NA SL.ranger.imp screen all Asian NA 7 256 SL.ranger.impscreen_all 0.139NatAmerNANA5.3810.139 PacIsl NA NA 2.553 SL.ranger.imp screen_all SL.ranger.imp0.139 Multiracial NANA5.054SL.ranger.imp0.139 Other NA NΑ 6.375 screen all

SL.ranger.imp

screen_all

0.139

Notreported

NA

NA

6.779

Table 2.4: Predictors in learners assigned weight > 0.0 by Superlearner.

Table 2.4: Predictors in learners assigned weight > 0.0 by Superlearner. (continued)

Learner	Screen	Weight	Predictors	Coefficient	Odds.Ratio	Importance
SL.ranger.imp	screen_all	0.139	Unknown	NA	NA	4.201
SL.ranger.imp	screen_all	0.139	HighRiskInd	NA	NA	17.116
SL.ranger.imp	screen_all	0.139	Sex	NA	NA	17.890
SL.ranger.imp	screen_all	0.139	Age	NA	NA	182.028
SL.ranger.imp	screen_all	0.139	BMI	NA	NA	231.544
SL.glm.interaction	screen_glmnet	0.12	(Intercept)	-2.963	0.052	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported	-0.075	0.928	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown	-0.283	0.753	NA
SL.glm.interaction	screen_glmnet	0.12	Black	-0.043	0.958	NA
SL.glm.interaction	screen_glmnet	0.12	Asian	0.023	1.023	NA
SL.glm.interaction	screen_glmnet	0.12	Multiracial	-0.422	0.656	NA
SL.glm.interaction	screen_glmnet	0.12	Other	0.074	1.077	NA
SL.glm.interaction	screen_glmnet	0.12	HighRiskInd	-0.057	0.945	NA
SL.glm.interaction	screen_glmnet	0.12	Age	0.849	2.337	NA
SL.glm.interaction	screen_glmnet	0.12	BMI	-0.065	0.937	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:EthnicityUnknown	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:Black	0.039	1.039	NA
SL.glm.interaction	screen glmnet	0.12	EthnicityNotreported:Asian	0.002	1.002	NA
SL.glm.interaction	screen glmnet	0.12	EthnicityNotreported:Multiracial	-0.525	0.592	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:Other	0.034	1.035	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:HighRiskInd	-0.069	0.934	NA
SL.glm.interaction	screen glmnet	0.12	EthnicityNotreported:Age	0.001	1.001	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:BMI	0.043	1.044	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown:Black	0.008	1.008	NA
SL.glm.interaction	screen glmnet	0.12	EthnicityUnknown:Asian	-0.067	0.936	NA
SL.glm.interaction	screen glmnet	0.12	EthnicityUnknown:Multiracial	-0.529	0.589	NA
SL.glm.interaction	screen glmnet	0.12	EthnicityUnknown:Other	0.066	1.068	NA
SL.glm.interaction	screen glmnet	0.12	EthnicityUnknown:HighRiskInd	-0.009	0.991	NA
SL.glm.interaction	screen glmnet	0.12	EthnicityUnknown:Age	0.114	1.121	NA
SL.glm.interaction	screen glmnet	0.12	EthnicityUnknown:BMI	-0.061	0.941	NA
SL.glm.interaction	screen_glmnet	0.12	Black:Asian	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	Black:Multiracial	NA	NA	NA
SL.glm.interaction	screen glmnet	0.12	Black:Other	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	Black:HighRiskInd	-0.018	0.982	NA
SL.glm.interaction	screen glmnet	0.12	Black:Age	0.012	1.012	NA
SL.glm.interaction	screen_glmnet	0.12	Black:BMI	-0.028	0.972	NA
SL.glm.interaction	screen glmnet	0.12	Asian:Multiracial	NA	NA	NA
SL.glm.interaction	screen glmnet	0.12	Asian:Other	NA	NA	NA
SL.glm.interaction	screen_gimnet screen glmnet	0.12	Asian:Other Asian:HighRiskInd	0.031	1.032	NA NA
SL.glm.interaction	screen_gimnet	0.12	Asian:Age	0.018	1.032	NA NA
SL.glm.interaction	screen_glmnet screen_glmnet	0.12	Asian: Age Asian: BMI	-0.025	0.976	NA NA
SL.glm.interaction	screen_gimnet	0.12	Multiracial:Other	NA	NA	NA
SL.glm.interaction	screen_gimnet screen_glmnet	0.12	Multiracial:HighRiskInd	-0.082	0.921	NA NA
SL.glm.interaction SL.glm.interaction	screen_gimnet screen glmnet	0.12	Multiracial: Age	0.103	1.108	NA NA

Learner	Screen	Weight	Predictors	Coefficient	Odds.Ratio	Importance
SL.glm.interaction	screen_glmnet	0.12	Other:HighRiskInd	0.036	1.037	NA
SL.glm.interaction	screen_glmnet	0.12	Other:Age	-0.019	0.981	NA
SL.glm.interaction	screen_glmnet	0.12	Other:BMI	0.036	1.037	NA
SL.glm.interaction	screen_glmnet	0.12	HighRiskInd:Age	0.025	1.026	NA
SL.glm.interaction	screen_glmnet	0.12	HighRiskInd:BMI	-0.090	0.914	NA
SL.glm.interaction	screen_glmnet	0.12	Age:BMI	0.012	1.012	NA
SL.glm.interaction	screen_glmnet	0.12	(Intercept)	-2.963	0.052	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported	-0.075	0.928	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown	-0.283	0.753	NA
SL.glm.interaction	screen_glmnet	0.12	Black	-0.043	0.958	NA
SL.glm.interaction	screen_glmnet	0.12	Asian	0.023	1.023	NA
SL.glm.interaction	screen_glmnet	0.12	Multiracial	-0.422	0.656	NA
SL.glm.interaction	screen_glmnet	0.12	Other	0.074	1.077	NA
SL.glm.interaction	screen_glmnet	0.12	HighRiskInd	-0.057	0.945	NA
SL.glm.interaction	screen_glmnet	0.12	Age	0.849	2.337	NA
SL.glm.interaction	screen_glmnet	0.12	BMI	-0.065	0.937	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:EthnicityUnknown	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:Black	0.039	1.039	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:Asian	0.002	1.002	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:Multiracial	-0.525	0.592	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:Other	0.034	1.035	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:HighRiskInd	-0.069	0.934	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:Age	0.001	1.001	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityNotreported:BMI	0.043	1.044	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown:Black	0.008	1.008	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown:Asian	-0.067	0.936	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown:Multiracial	-0.529	0.589	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown:Other	0.066	1.068	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown:HighRiskInd	-0.009	0.991	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown:Age	0.114	1.121	NA
SL.glm.interaction	screen_glmnet	0.12	EthnicityUnknown:BMI	-0.061	0.941	NA
SL.glm.interaction	screen_glmnet	0.12	Black:Asian	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	Black:Multiracial	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	Black:Other	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	Black:HighRiskInd	-0.018	0.982	NA
SL.glm.interaction	screen_glmnet	0.12	Black:Age	0.012	1.012	NA
SL.glm.interaction	screen_glmnet	0.12	Black:BMI	-0.028	0.972	NA
SL.glm.interaction	screen_glmnet	0.12	Asian:Multiracial	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	Asian:Other	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	Asian:HighRiskInd	0.031	1.032	NA
SL.glm.interaction	screen_glmnet	0.12	Asian:Age	0.018	1.018	NA
SL.glm.interaction	screen_glmnet	0.12	Asian:BMI	-0.025	0.976	NA
SL.glm.interaction	screen_glmnet	0.12	Multiracial:Other	NA	NA	NA
SL.glm.interaction	screen_glmnet	0.12	Multiracial:HighRiskInd	-0.082	0.921	NA
SL.glm.interaction	screen_glmnet	0.12	Multiracial:Age	0.103	1.108	NA

Table 2.4: Predictors in learners assigned weight > 0.0 by Superlearner. (continued)

Table 2.4: Predictors in learners assigned weight > 0.0 by Superlearner. (continued)

_						
Learner	Screen	Weight	Predictors	Coefficient	Odds.Ratio	Importance
SL.glm.interaction	screen_glmnet	0.12	Multiracial:BMI	0.000	1.000	NA
SL.glm.interaction	screen_glmnet	0.12	Other:HighRiskInd	0.036	1.037	NA
SL.glm.interaction	screen_glmnet	0.12	Other:Age	-0.019	0.981	NA
SL.glm.interaction	screen_glmnet	0.12	Other:BMI	0.036	1.037	NA
SL.glm.interaction	screen_glmnet	0.12	HighRiskInd:Age	0.025	1.026	NA
SL.glm.interaction	screen_glmnet	0.12	HighRiskInd:BMI	-0.090	0.914	NA
SL.glm.interaction	screen_glmnet	0.12	Age:BMI	0.012	1.012	NA



Figure 2.4: Superlearner predicted probabilities of COVID-19 disease in vaccinees 7 days after Day 57 by case/control status.



Figure 2.5: ROC curve based off Superlearner predicted probabilities in vaccinees.



Chapter 3

Appendix

- This report was built from the CoVPN/correlates_reporting repository with commit hash 77cdcc434759b9a204d42459f69123a23bbfb215. A diff of the changes introduced by that commit may be viewed at https://github.com/CoVPN/correlates_reporting/commit/77cdcc434759b9a204d42459f69123a23bbfb215
- The sha256 hash sum of the raw input file, "COVID_VEtrial_practicedata_primarystage1.csv": 83d0f55d1745ffd42be124d8f9ec9a9903abcc13cd22f95e537542a08b41300a
- The sha256 hash sum of the processed file, "moderna_mock_data_processed.csv": 28964ce20cfcd70a621aff9df412c42b1178d5e749e0f49ddd79fbff2dc7c90a