

# COVID-19 Baseline Risk Score Analysis Report

## 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.

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## 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=non-minority)	0/30000 (0.0%)	NA
EthnicityHispanic	Indicator ethnicity = Hispanic (0 = Non-Hispanic)	0/30000 (0.0%)	NA
EthnicityNotreported	Indicator ethnicity = Not reported (0 = Non-Hispanic)	0/30000 (0.0%)	NA
EthnicityUnknown	Indicator ethnicity = Unknown (0 = Non-Hispanic)	0/30000 (0.0%)	NA
Black	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 ( $\text{kg}/\text{m}^2$ )	0/30000 (0.0%)	NA

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

<b>Learner</b>	<b>Screen*</b>
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:*

\*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  $< 0.10$

highcor\_random: if pairs of quantitative variables with Spearman rank correlation  $> 0.90$ , select one of the variables at random



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 SL.



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

Table 2.4: Predictors in learners assigned weight &gt; 0.0 by Superlearner.

Learner	Screen	Weight	Predictors	Coefficient	Odds.Ratio	Importance
SL.gam	screen_highcor_random	0.488	(Intercept)	-2.873	0.057	NA
SL.gam	screen_highcor_random	0.488	s(Age,2)	0.829	2.290	NA
SL.gam	screen_highcor_random	0.488	s(BMI,2)	-0.046	0.955	NA
SL.gam	screen_highcor_random	0.488	MinorityInd	-0.052	0.950	NA
SL.gam	screen_highcor_random	0.488	EthnicityHispanic	0.004	1.004	NA
SL.gam	screen_highcor_random	0.488	EthnicityNotreported	0.043	1.044	NA
SL.gam	screen_highcor_random	0.488	EthnicityUnknown	-0.073	0.930	NA
SL.gam	screen_highcor_random	0.488	Black	0.004	1.004	NA
SL.gam	screen_highcor_random	0.488	Asian	0.069	1.071	NA
SL.gam	screen_highcor_random	0.488	NatAmer	0.015	1.016	NA
SL.gam	screen_highcor_random	0.488	PacIsl	-0.013	0.987	NA
SL.gam	screen_highcor_random	0.488	Multiracial	-0.046	0.955	NA
SL.gam	screen_highcor_random	0.488	Other	0.064	1.066	NA
SL.gam	screen_highcor_random	0.488	Notreported	0.003	1.003	NA
SL.gam	screen_highcor_random	0.488	Unknown	0.003	1.003	NA
SL.gam	screen_highcor_random	0.488	HighRiskInd	-0.026	0.974	NA
SL.gam	screen_highcor_random	0.488	Sex	-0.001	0.999	NA
SL.glmnet	screen_all	0.253	(Intercept)	-2.665	0.070	NA
SL.glmnet	screen_all	0.253	MinorityInd	0.000	1.000	NA
SL.glmnet	screen_all	0.253	EthnicityHispanic	0.000	1.000	NA
SL.glmnet	screen_all	0.253	EthnicityNotreported	0.000	1.000	NA
SL.glmnet	screen_all	0.253	EthnicityUnknown	0.000	1.000	NA
SL.glmnet	screen_all	0.253	Black	0.000	1.000	NA
SL.glmnet	screen_all	0.253	Asian	0.000	1.000	NA
SL.glmnet	screen_all	0.253	NatAmer	0.000	1.000	NA
SL.glmnet	screen_all	0.253	PacIsl	0.000	1.000	NA
SL.glmnet	screen_all	0.253	Multiracial	0.000	1.000	NA
SL.glmnet	screen_all	0.253	Other	0.000	1.000	NA
SL.glmnet	screen_all	0.253	Notreported	0.000	1.000	NA
SL.glmnet	screen_all	0.253	Unknown	0.000	1.000	NA
SL.glmnet	screen_all	0.253	HighRiskInd	0.000	1.000	NA
SL.glmnet	screen_all	0.253	Sex	0.000	1.000	NA
SL.glmnet	screen_all	0.253	Age	0.444	1.559	NA
SL.glmnet	screen_all	0.253	BMI	0.000	1.000	NA
SL.ranger.imp	screen_all	0.139	MinorityInd	NA	NA	9.100
SL.ranger.imp	screen_all	0.139	EthnicityHispanic	NA	NA	9.685
SL.ranger.imp	screen_all	0.139	EthnicityNotreported	NA	NA	9.247
SL.ranger.imp	screen_all	0.139	EthnicityUnknown	NA	NA	6.908
SL.ranger.imp	screen_all	0.139	Black	NA	NA	9.156
SL.ranger.imp	screen_all	0.139	Asian	NA	NA	7.256
SL.ranger.imp	screen_all	0.139	NatAmer	NA	NA	5.381
SL.ranger.imp	screen_all	0.139	PacIsl	NA	NA	2.553
SL.ranger.imp	screen_all	0.139	Multiracial	NA	NA	5.054
SL.ranger.imp	screen_all	0.139	Other	NA	NA	6.375
SL.ranger.imp	screen_all	0.139	Notreported	NA	NA	6.779

Table 2.4: Predictors in learners assigned weight &gt; 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_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
SL.glm.interaction	screen_glmnet	0.12	Multiracial:BMI	0.000	1.000	NA

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	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)*

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

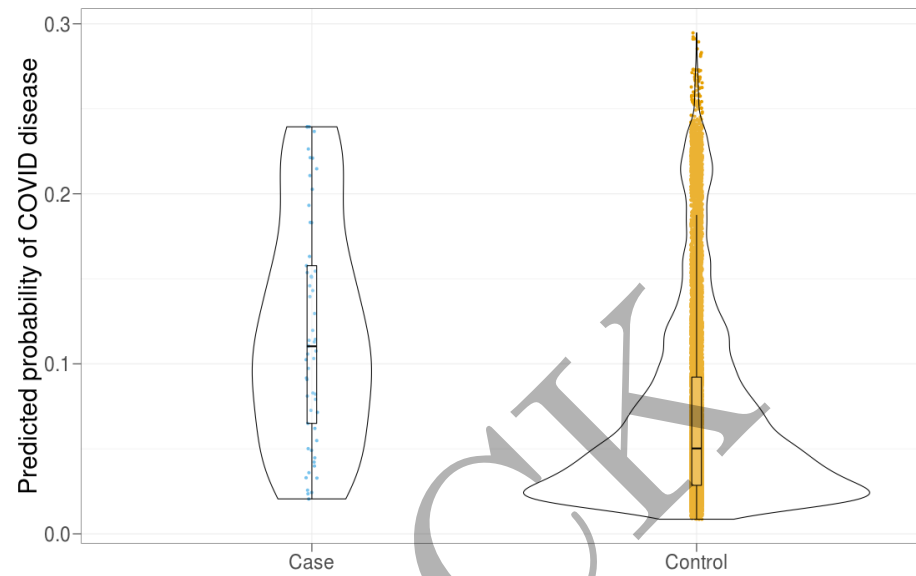


Figure 2.4: Superlearner predicted probabilities of COVID-19 disease in vaccinees 7 days after Day 57 by case/control status.



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

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## Chapter 3

# Appendix

- This report was built from the [CoVPN/correlates\\_reporting](https://github.com/CoVPN/correlates_reporting) repository with commit hash 24ed61884c57033177845492cccb7a2820196bd2. A diff of the changes introduced by that commit may be viewed at [https://github.com/CoVPN/correlates\\_reporting/commit/24ed61884c57033177845492cccb7a2820196bd2](https://github.com/CoVPN/correlates_reporting/commit/24ed61884c57033177845492cccb7a2820196bd2)
- 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”: 28964ce20cfd70a621aff9df412c42b1178d5e749e0f49ddd79fbff2dc7c90a