585 SUPPLEMENTARY TABLES

Table S1. Extended COVID Symptoms and Underlying Conditions by Vaccination Status

Variable	Overall , N = 1.957^{1}	Unvaccinated, N = 1.024^{1}	Vaccinated, $N = 933^{1}$	p ²	
COVID Symptoms		-,			
Fever	720 (47%)	430 (52%)	290 (42%)	< 0.001	
(Missing)	435	197	238	0.001	
Chills	537 (35%)	314 (38%)	223 (32%)	0.02	
(Missing)	435	198	237		
Rigors	29 (2.0%)	17 (2.1%)	12 (1.8%)	0.67	
(Missing)	491	221	270		
Myalgia	501 (34%)	279 (35%)	222 (33%)	0.59	
(Missing)	484	218	266		
Headache	521 (35%)	299 (37%)	222 (33%)	0.12	
(Missing)	476	215	261		
Sore Throat	286 (20%)	138 (17%)	148 (22%)	0.02	
(Missing)	496	228	268		
Nausea / Vomiting	451 (30%)	282 (35%)	169 (25%)	< 0.001	
(Missing)	455	209	246		
Diarrhea	369 (25%)	212 (26%)	157 (23%)	0.13	
(Missing)	470	220	250		
Fatigue	630 (41%)	337 (41%)	293 (41%)	0.81	
(Missing)	418	195	223		
Runny Nose or Nasal Congestion	519 (35%)	217 (27%)	302 (45%)	< 0.001	
(Missing)	473	215	258		
Cough	1,087 (70%)	576 (69%)	511 (72%)	0.37	
(Missing)	414	195	219		
Shortness of Breath or Difficulty Breathing	690 (44%)	408 (49%)	282 (39%)	< 0.001	
(Missing)	401	188	213		
Loss of Taste	264 (18%)	151 (19%)	113 (17%)	0.33	
(Missing)	505	233	272		
Underlying Conditions					
Pregnant	69 (3.6%)	59 (6.0%)	10 (1.1%)	< 0.001	
(Missing)	66	42	24		
Chronic Lung Disease	387 (21%)	193 (20%)	194 (22%)	0.33	
(Missing)	134	75	59		
Hypertension	874 (47%)	377 (39%)	497 (56%)	< 0.001	
(Missing)	95	50	45		
Overweight	584 (32%)	315 (33%)	269 (32%)	0.50	
(Missing)	150	70	80		
Cardiovascular Disease	534 (29%)	217 (23%)	317 (36%)	< 0.001	
(Missing)	126	65	61		
Diabetes	426 (23%)	180 (19%)	246 (28%)	< 0.001	
(Missing)	101	53	48		

Renal Disease	316 (17%)	106 (11%)	210 (24%)	< 0.001
(Missing)	132	69	63	
Liver Disease	83 (4.5%)	37 (3.8%)	46 (5.2%)	0.16
(Missing)	107	59	48	
Autoimmune Disease	101 (5.5%)	42 (4.4%)	59 (6.8%)	0.03
(Missing)	135	69	66	
Immunocompromised ³	269 (15%)	89 (9.3%)	180 (21%)	< 0.001
(Missing)	137	69	68	
Systemic Immunosuppressive Therapy or Medications	293 (16%)	117 (12%)	176 (20%)	< 0.001
(Missing)	115	59	56	

¹ n (%) or median (Interquartile Range [IQR])

² Pearson's Chi-squared test or Fisher's exact test for categorical variables and Wilcoxon rank-sum test for continuous variables

³ Immunocompromised (e.g., HIV infection, active cancer, solid organ transplant, hematopoietic stem cell transplant)

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Table S2	2. Circulation of VOC/VOI	in Georgia, USA as measur	ed by available sec	juences on GISAID
VOC/VOI	Period variant circulated in GA ¹	Date range of sequences obtained from GISAID	Sequences available on GISAID	Total sequences available from May 2021 to May 2022 (%)
Alpha	12/10/2020-08/26/2021	05/01/2021-08/26/2021	1,361	2.66
Beta	02/01/2021-05/25/2021	05/01/2021-05/25/2021	16	0.03
Delta	4/29/2021-03/10/2022	05/01/2021-03/10/2022	28,320	55.5
Gamma	03/05/2021-09/24/2021	05/01/2021-09/24/2021	185	0.36
Lamba	02/07/2021-08/05/2021	05/01/2021-08/05/2021	18	0.04
Mu	04/23/2021-09/14/2021	05/01/2021-09/14/2021	142	0.28
Omicron	11/30/2021-4/18/2023	11/30/2021-04/18/2023	20,759	40.6
		Total VOC/VOI	50,801	99.46
Fotal sequenc	es available on GISAID fro	m May 2021 to May 2022 from the state of Georgia	51,075	

¹Period of circulation based on sequences available on GISAID including Emory Healthcare samples.

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	Moderate ¹		Sever	Severe ¹			Death ¹		
Variable	OR ²	95% CI ²	р	OR ²	95% CI ²	р	OR ²	95% CI ²	р
Unadjusted Models									
Age	1.04	1.03, 1.05	<0.001	1.05	1.04, 1.05	<0.001	1.07	1.05, 1.09	<0.001
Male ³	1.40	1.13, 1.74	<0.01	1.30	0.95, 1.78	0.10	1.26	0.72, 2.22	0.40
White ⁴	0.86	0.67, 1.10	0.20	1.27	0.91, 1.77	0.20	1.04	0.57, 1.91	0.90
Chronic Lung Disease	2.07	1.59, 2.68	<0.001	1.95	1.35, 2.81	<0.001	2.65	1.44, 4.86	<0.01
Hypertension	2.94	2.35, 3.68	<0.001	4.19	2.98, 5.90	<0.001	11.4	5.09, 25.6	<0.001
Overweight	1.45	1.15, 1.83	<0.01	1.58	1.14, 2.19	0.01	2.02	1.13, 3.61	0.02
Cardiovascular Disease	2.89	2.28, 3.67	<0.001	3.46	2.48, 4.81	<0.001	4.80	2.70, 8.54	<0.001
Diabetes	2.04	1.58, 2.63	<0.001	3.21	2.29, 4.50	<0.001	4.35	2.43, 7.80	<0.001
Renal Disease	3.88	2.90, 5.19	<0.001	3.86	2.62, 5.67	<0.001	9.86	5.46, 17.8	<0.001
Liver Disease	1.47	0.87, 2.50	0.20	2.36	1.25, 4.45	<0.01	3.08	1.16, 8.21	0.02
Autoimmune Disease	1.60	1.01, 2.56	0.046	1.58	0.82, 3.03	0.20	2.45	0.93, 6.47	0.07
Immunocompromised	2.65	1.97, 3.58	<0.001	2.44	1.61, 3.69	<0.001	3.69	1.93, 7.09	<0.001
Systemic Immunosuppressive Therapy/Meds.	3.49	2.61, 4.67	<0.001	3.24	2.18, 4.81	<0.001	3.48	1.79, 6.78	<0.001
Vaccination Status									
Unvaccinated	Ref			Ref	_		Ref		
Vaccinated ⁵	0.80	0.64, 1.01	0.07	0.86	0.61, 1.21	0.40	1.00	0.54, 1.84	>0.90
Vaccinated & boosted ⁵	2.28	1.52, 3.43	<0.001	3.24	1.95, 5.40	<0.001	3.69	1.58, 8.59	<0.01
<u>Days Since Most Recent</u> <u>Vaccination/Booster</u>									
Unvaccinated	_	_			_		_	_	
Within past 90 Days ⁵	1.34	0.86, 2.08	0.20	1.68	0.93, 3.03	0.09	2.81	1.17, 6.79	0.02
91 - 180 Days Ago ⁵	0.98	0.74, 1.29	0.90	0.92	0.60, 1.40	0.70	1.14	0.55, 2.37	0.70
181 - 270 Days Ago ⁵	0.72	0.51, 1.01	0.06	0.74	0.45, 1.24	0.30	0.71	0.27, 1.89	0.50
More than 270 Days Ago ⁵	1.18	0.72, 1.94	0.50	2.49	1.43, 4.36	< 0.01	1.91	0.64, 5.72	0.20
<u>Lineage</u>									
Delta	Ref	_		Ref	_		Ref		
Omicron ⁶	1.51	1.09, 2.07	0.01	2.25	1.46, 3.48	<0.001	4.55	2.25, 9.20	<0.001

Table S3. Unadjusted Multinomial Logistic Regression Models Testing the Association between Demographics, Underlying
Health Conditions, and COVID-Related Characteristics with Disease Severity

¹" Mild" is the Reference Category

² OR = Odds Ratio, CI = Confidence Interval, Ref = Reference Level ³ Compared to "Female" ⁴ Compared to "Black"

⁵Compared to "Unvaccinated individuals"

⁶Compared to "Delta"

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unvaccinated ("unvax") individuals are labeled. Each square in the heatmap represents the

number of single nucleotide polymorphisms (SNPs) in each gene, labeled on the x-axis. Each

row represents one sample.





599 Figure S2. Phylogenetic analysis does not reveal differences in SARS-CoV-2 sequences

600 **from vaccinated and unvaccinated individuals.** Maximum likelihood tree containing

sequences from vaccinated (green) and unvaccinated (yellow) individuals in the context of 2000

602 global sequences from GISAID (orange) selected by a custom Nextstrain subsampling scheme

603 and rooted to NC_045512.