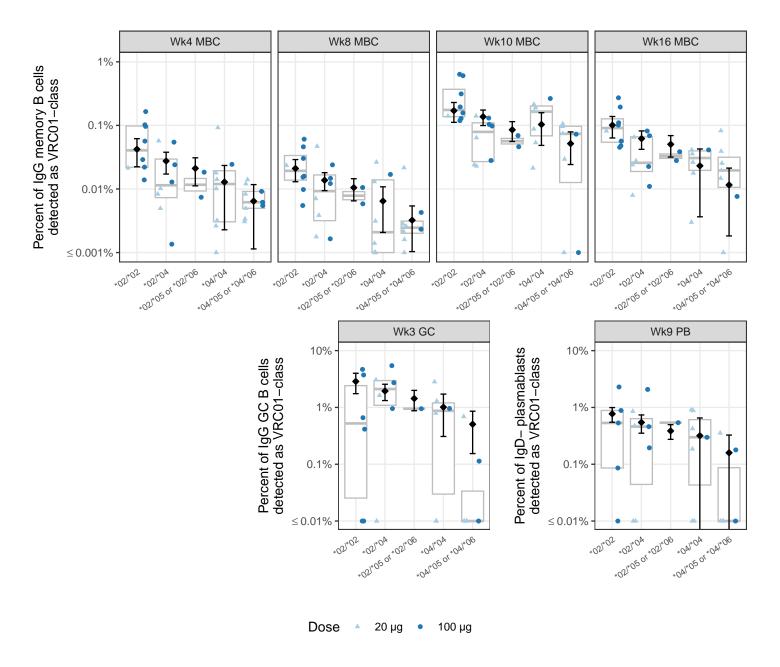
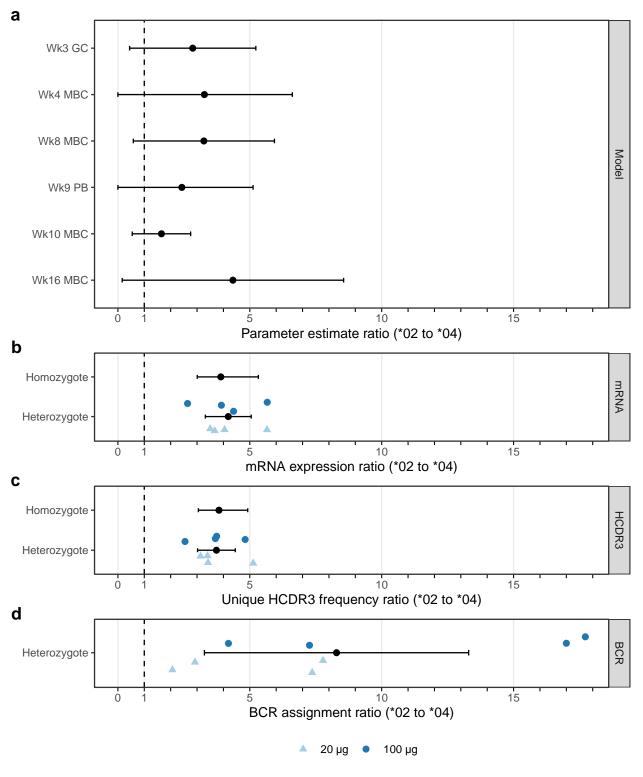


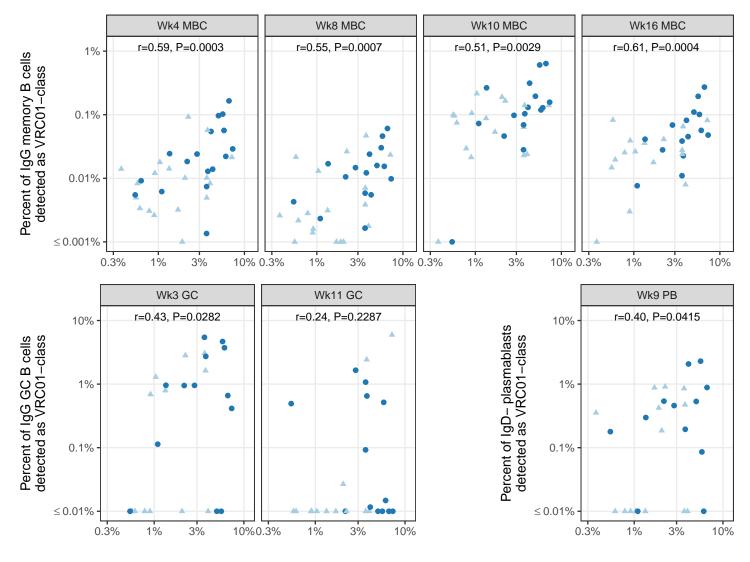
 $Composite\ Genotype-mRNA-HCDR3-Usage-Plot$



Allele model estimates by genotype.



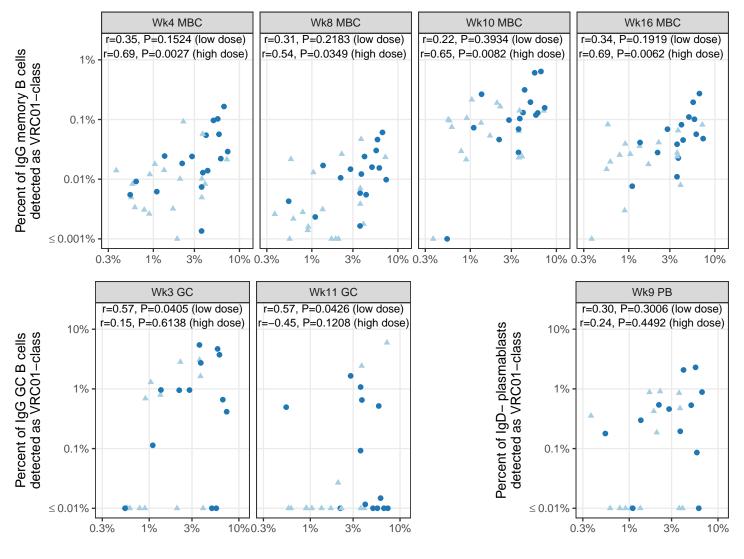
Ratio estimates



Unique HCDR3 frequency (IGHV1-2*02 or *04)

Dose Δ 20 μg • 100 μg

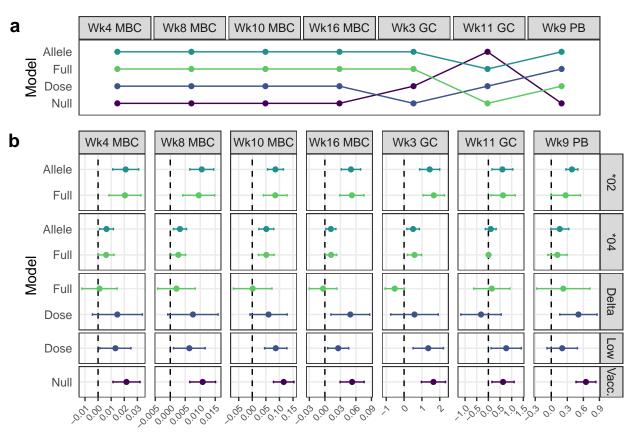
Naive vs. VRC01 response correlations



Unique HCDR3 frequency (IGHV1-2*02 or *04)

Dose Δ 20 μg • 100 μg

Naive vs. VRC01 response correlations



Parameter estimate and 95% confidence interval (percent scale)

Quasipoisson fit plot

Table S1

Treatment	*02/*02	*02/*04	*02/*05	*02/*06	*04/*04	*04/*05	*04/*06	*05/*06
100 μg	7	4	2	0	1	1	2	1
$20~\mu \mathrm{g}$	1	4	0	0	7	1	5	0
Placebo	3	5	0	2	2	0	0	0
Total	11	13	2	2	10	2	7	1

Table S2

	Allele	n	Mean	95% CI	Min	Max
*02	Homozygote Heterozygote	11 17	$3.15\% \ 3.34\%$	2.72% to 3.57% 2.87% to 3.82%	2.28% $1.98%$	4.52% $6.29%$
*04	Homozygote Heterozygote	10 22	0.86% $0.72%$	0.67% to 1.06% 0.63% to 0.81%	$0.41\% \\ 0.25\%$	1.22% 1.12%
*05	Heterozygote	5	0.09%	0.03% to $0.14%$	0.03%	0.14%
*06	Heterozygote	10	2.43%	1.87% to 2.99%	1.78%	4.18%

Table S3

Allele	Difference	95% CI	P-value
*02	-0.19%	-0.80% to 0.41%	0.51
*04	0.14%	-0.07% to 0.35%	0.17

Table S4

				P Value		
	Comparison	Sample Sizes	Median (Range)	Unadjusted	FDR Adjusted	
) μ g Wk4 MBC	*02/*04 vs. *04/*04	4 vs. 6	0.0092 [0.0049, 0.0572] vs. 0.0121 [0.0026, 0.0924]	1.0000	1.0000	
	*02/*04 vs. *04/*05 or *04/*06	4 vs. 6	0.0092 [0.0049, 0.0572] vs. 0.0066 [0.0031, 0.0140]	0.4762	0.8081	
	*04/*04 vs. *04/*05 or *04/*06	6 vs. 6	0.0121 [0.0026, 0.0924] vs. 0.0066 [0.0031, 0.0140]	0.4848	0.8081	
Wk8 MBC	*02/*04 vs. *04/*04	4 vs. 4	0.0054 [0.0018, 0.0469] vs. 0.0081 [0.0014, 0.0264]	0.8857	0.9797	
,, no 1120	*02/*04 vs. *04/*05 or *04/*06	4 vs. 6	0.0054 [0.0018, 0.0469] vs. 0.0024 [0.0007, 0.0216]	0.2571	0.8000	
	*04/*04 vs. *04/*05 or *04/*06	4 vs. 6	0.0081 [0.0014, 0.0264] vs. 0.0024 [0.0007, 0.0216]	0.3524	0.8000	
Wk10 MBC	*02/*04 vs. *04/*04	4 vs. 6	0.0440 [0.0230, 0.1404] vs. 0.1263 [0.0214, 0.2148]	0.3524	0.8000	
WRIO MIDO	*02/*04 vs. *04/*05 or *04/*06	4 vs. 6	0.0440 [0.0230, 0.1404] vs. 0.0854 [0.0005, 0.1057]	0.7619	0.9797	
	*04/*04 vs. *04/*05 or *04/*06	6 vs. 6	0.1263 [0.0214, 0.2148] vs. 0.0854 [0.0005, 0.1057]	0.3939	0.8000	
Wk16 MBC	*02/*04 vs. *04/*04	4 vs. 5	0.0259 [0.0080, 0.0653] vs. 0.0263 [0.0030, 0.0412]	0.9048	0.9797	
WKIO MIDO	*02/*04 vs. *04/*05 or *04/*06	4 vs. 6	0.0259 [0.0080, 0.0653] vs. 0.0225 [0.0001, 0.0826]	0.9143	0.9797	
	*04/*04 vs. *04/*05 or *04/*06	5 vs. 6	0.0263 [0.0030, 0.0412] vs. 0.0225 [0.0001, 0.0826]	0.9307	0.9797	
Wk3 GC	*02/*04 vs. *04/*04	2 vs. 3	2.3429 [1.6281, 3.0577] vs. 1.2946 [0.7917, 2.8275]	0.9307	0.9191	
Wk9 PB	*02/*04 vs. *04/*04	2 vs. 3	0.6629 [0.4731, 0.8527] vs. 0.6510 [0.1856, 0.9081]			
	02/ 04 VS. 04/ 04	2 VS. 4	0.0029 [0.4731, 0.8327] VS. 0.0310 [0.1630, 0.3081]			
00 μg Wk4 MBC	*02/*02 vs. *02/*04	7 vs. 4	0.0566 [0.0139, 0.1649] vs. 0.0185 [0.0014, 0.0545]	0.1091	0.7273	
	*02/*02 vs. *02/*05 or *02/*06	7 vs. 2	0.0566 [0.0139, 0.1649] vs. 0.0129 [0.0074, 0.0184]	_		
	*02/*02 vs. *04/*05 or *04/*06	7 vs. 3	0.0566 [0.0139, 0.1649] vs. 0.0062 [0.0055, 0.0092]	***0.0167**	* 0.2424	
	*02/*04 vs. *02/*05 or *02/*06	4 vs. 2	0.0185 [0.0014, 0.0545] vs. 0.0129 [0.0074, 0.0184]	_	_	
	*02/*04 vs. *04/*05 or *04/*06	4 vs. 3	0.0185 [0.0014, 0.0545] vs. 0.0062 [0.0055, 0.0092]	0.4000	0.8000	
	*02/*05 or *02/*06 vs. *04/*05 or *04/*06	2 vs. 3	0.0129 [0.0074, 0.0184] vs. 0.0062 [0.0055, 0.0092]		_	
Wk8 MBC	*02/*02 vs. *02/*04	7 vs. 4	0.0159 [0.0055, 0.0608] vs. 0.0134 [0.0016, 0.0240]	0.3152	0.8000	
	*02/*02 vs. *02/*05 or *02/*06	7 vs. 2	0.0159 [0.0055, 0.0608] vs. 0.0082 [0.0058, 0.0105]		_	
	*02/*02 vs. *04/*05 or *04/*06	7 vs. 2	0.0159 [0.0055, 0.0608] vs. 0.0033 [0.0023, 0.0043]	_	_	
	*02/*04 vs. *02/*05 or *02/*06	4 vs. 2	0.0134 [0.0016, 0.0240] vs. 0.0082 [0.0058, 0.0105]	_	_	
	*02/*04 vs. *04/*05 or *04/*06	4 vs. 2	0.0134 [0.0016, 0.0240] vs. 0.0033 [0.0023, 0.0043]		_	
	*02/*05 or *02/*06 vs. *04/*05 or *04/*06	2 vs. 2	0.0082 [0.0058, 0.0105] vs. 0.0033 [0.0023, 0.0043]	_	_	
Wk10 MBC	*02/*02 vs. *02/*04	7 vs. 4	0.1951 [0.1188, 0.6376] vs. 0.1007 [0.0281, 0.1305]	***0.0242**	* 0.2424	
	*02/*02 vs. *02/*05 or *02/*06	7 vs. 2	0.1951 [0.1188, 0.6376] vs. 0.0577 [0.0461, 0.0692]	_		
	*02/*04 vs. *02/*05 or *02/*06	4 vs. 2	0.1007 [0.0281, 0.1305] vs. 0.0577 [0.0461, 0.0692]		_	
Wk16 MBC	*02/*02 vs. *02/*04	7 vs. 4	0.1010 [0.0452, 0.2718] vs. 0.0457 [0.0110, 0.0817]	0.1636	0.8000	
	*02/*02 vs. *02/*05 or *02/*06	7 vs. 2	0.1010 [0.0452, 0.2718] vs. 0.0332 [0.0280, 0.0385]		_	
	*02/*04 vs. *02/*05 or *02/*06	4 vs. 2	0.0457 [0.0110, 0.0817] vs. 0.0332 [0.0280, 0.0385]		_	
Wk3 GC	*02/*02 vs. *02/*04	4 vs. 3	2.1944 [0.4144, 4.6746] vs. 2.7265 [0.9546, 5.4440]	0.6286	0.9670	
Wk11 GC	*02/*02 vs. *02/*04	2 vs. 4	0.2658 [0.0148, 0.5167] vs. 0.3717 [0.0116, 1.6519]	_		
Wk9 PB	*02/*02 vs. *02/*04	4 vs. 3	0.7086 [0.0858, 2.2943] vs. 0.4593 [0.1946, 2.0745]	0.8571	0.9797	

Table S5

					P Va	ılue
		Comparison	Sample Sizes	Median (Range)	Unadjusted	FDR Adjusted
02/*04						
Wk	k4 MBC	20 μg vs. 100 μg	4 vs. 4	$0.0092\ [0.0049,\ 0.0572]\ vs.\ 0.0185\ [0.0014,\ 0.0545]$	0.8857	0.9048
Wk	k8 MBC	20 μg vs. 100 μg	4 vs. 4	0.0054 [0.0018, 0.0469] vs. 0.0134 [0.0016, 0.0240]	0.8857	0.9048
Wk	k10 MBC	20 μg vs. 100 μg	4 vs. 4	0.0440 [0.0230, 0.1404] vs. 0.1007 [0.0281, 0.1305]	0.4857	0.9048
Wk	k16 MBC	20 μg vs. 100 μg	4 vs. 4	$0.0259 \; [0.0080, 0.0653] \; \mathrm{vs.} \; \; 0.0457 \; [0.0110, 0.0817]$	0.6857	0.9048
Wk	k3 GC	20 μg vs. 100 μg	2 vs. 3	2.3429 [1.6281, 3.0577] vs. 2.7265 [0.9546, 5.4440]	_	_
Wk	k9 PB	20 μg vs. 100 μg	2 vs. 3	$0.6629\ [0.4731,\ 0.8527]\ vs.\ 0.4593\ [0.1946,\ 2.0745]$	_	_
04/*05	or *04/*0	6				
,	k4 MBC	$20~\mu \mathrm{g}$ vs. $100~\mu \mathrm{g}$	6 vs. 3	$0.0066\ [0.0031,\ 0.0140]\ vs.\ 0.0062\ [0.0055,\ 0.0092]$	0.9048	0.9048
Wk	k8 MBC	20 μg vs. 100 μg	6 vs. 2	0.0024 [0.0007, 0.0216] vs. 0.0033 [0.0023, 0.0043]	_	_

Table S6

Model	n	K	LL	Dispersion	QAICc
Wk4 M	BC				·
Allele	35	3	-954.5112	67.96823	33.63438
Full	35	4	-953.5981	71.07256	36.16783
Dose	35	3	-1241.2651	102.81975	41.70371
Null	35	2	-1353.1930	110.50062	42.45420
Wk8 M	\mathbf{BC}				
Allele	34	3	-467.4507	27.70813	37.91914
Full	34	4	-461.2600	30.04265	40.08632
Dose	34	3	-615.3809	47.22338	47.76716
Null	34	2	-684.2942	44.60637	49.94195
Wk10 N	ıвс				
Allele	33	3	-3913.2854	275.82562	34.31452
Full	33	4	-3913.0473	284.73788	36.91383
Dose	33	3	-4507.3975	325.05221	38.48756
Null	33	2	-4997.3051	397.41236	39.50109
Wk16 N	IВС				
Allele	31	3	-1192.3772	95.31918	31.38919
Full	31	4	-1189.6335	97.33573	33.98238
Dose	31	3	-1638.2275	127.00529	40.55027
Null	31	2	-2049.7041	178.03677	46.54474
Wk3 G0	C				
Allele	26	3	-7480.1546	486.41216	37.62126
Full	26	4	-7182.3606	490.01431	39.21966
Null	26	2	-11266.3607	803.67639	50.50554
Dose	26	3	-10976.7429	779.39438	51.89263
Wk11 G	\mathbf{c}				
Null	26	2	-8550.4050	1001.94407	26.62847
Allele	26	3	-7690.3863	705.77005	26.97410
Dose	26	3	-8333.1437	881.98514	28.63592
Full	26	4	-7474.9931	773.55665	29.23106
Wk9 PE	3				
Allele	26	3	-628.4218	45.03017	34.45853
Dose	26	3	-655.0902	46.71709	35.61993
Full	26	4	-613.5097	45.92447	36.62296
Null	26	2	-759.2088	49.71729	37.58511

Table S7

Parameter	Allele	Full	Dose	Null
Wk4 MB0	C			
*02	$0.021\ (0.011,\ 0.031)$	$0.021\ (0.008,\ 0.033)$		
*04	$0.006\ (0.001,\ 0.012)$	$0.006\ (0.000,\ 0.012)$		
Low			$0.013\ (0.001,\ 0.025)$	
Delta		$0.001 \ (-0.012, \ 0.015)$	0.015 (-0.004, 0.034)	
Vacc.				$0.022 \ (0.011, \ 0.032)$
Wk8 MB0	\mathbb{C}			
*02	$0.011\ (0.006,\ 0.015)$	$0.010\ (0.004,\ 0.015)$		
*04	$0.003\ (0.001,\ 0.005)$	$0.003\ (0.000,\ 0.005)$		
Low			$0.006 \ (0.001, \ 0.012)$	
Delta		$0.002 \ (-0.004, \ 0.008)$	$0.008 \ (-0.001, \ 0.016)$	
Vacc.				$0.011\ (0.007,\ 0.015)$
Wk10 ME	3C			
*02	$0.085 \ (0.056, \ 0.115)$	$0.085 \ (0.041, \ 0.128)$		
*04	$0.052\ (0.024,\ 0.079)$	$0.052\ (0.023,\ 0.080)$		
Low			$0.086 \ (0.045, \ 0.127)$	
Delta		$0.002 \ (-0.069, \ 0.073)$	$0.060 \ (-0.008, \ 0.128)$	
Vacc.				$0.116 \ (0.078, \ 0.153)$
Wk16 ME	BC			
*02	$0.050\ (0.032,\ 0.069)$	$0.052\ (0.029,\ 0.076)$		
*04	$0.012\ (0.002,\ 0.021)$	$0.012\ (0.001,\ 0.023)$		
Low			$0.026 \ (0.005, \ 0.046)$	
Delta		-0.004 (-0.030, 0.023)	$0.049 \ (0.012, \ 0.087)$	
Vacc.				$0.053 \ (0.030, \ 0.076)$
Wk3 GC				
*02	$1.435 \ (0.874, 1.996)$	1.665 (1.070, 2.261)		
*04	$0.506 \ (0.154, \ 0.858)$	$0.584 \ (0.185, \ 0.983)$		
Low			$1.353 \ (0.500, \ 2.206)$	
Delta		-0.523 (-1.067, 0.020)	$0.584 \ (-0.747, \ 1.914)$	
Vacc.				$1.645 \ (0.970, \ 2.320)$
Wk11 GC				
*02	$0.602 \ (0.157, \ 1.047)$	$0.630 \ (0.113, \ 1.147)$		
*04	$0.103 \ (-0.127, \ 0.333)$	$0.008 \ (-0.071, \ 0.087)$		
Low			0.764 (0.116, 1.411)	
Delta		$0.147 \ (-0.631, \ 0.925)$	-0.317 (-1.183, 0.550)	0.000 (0.471 + 151)
Vacc.				$0.629 \ (0.154, \ 1.104)$
Wk9 PB				
*02	$0.386 \ (0.274, \ 0.497)$	0.268 (-0.009, 0.544)		
*04	$0.159 \ (-0.008, \ 0.326)$	$0.115 \ (-0.064, \ 0.295)$		
Low		0.00* (0.0=; 0.=0=;	0.205 (-0.080, 0.489)	
Delta		$0.225 \ (-0.274, \ 0.723)$	$0.508 \ (0.160, \ 0.856)$	0.040 (0.400 0.001)
Vacc.				$0.648 \ (0.463, \ 0.834)$

Table S8

Genotype	Percent (95% CI)
Wk4 MBC	
*02/*02	$0.042\ (0.022,\ 0.062)$
*02/*04	0.028 (0.017, 0.038)
*02/*05 or *02/*06	0.021 (0.011, 0.031)
*04/*04	0.013 (0.002, 0.023)
*04/*05 or *04/*06	$0.006\ (0.001,\ 0.012)$
Wk8 MBC	
*02/*02	$0.021\ (0.013,\ 0.029)$
*02/*04	0.014 (0.009, 0.018)
*02/*05 or *02/*06	$0.011 \ (0.006, \ 0.015)$
*04/*04	$0.006 \ (0.002, \ 0.011)$
*04/*05 or *04/*06	$0.003 \ (0.001, \ 0.005)$
Wk10~MBC	
*02/*02	$0.171\ (0.112,\ 0.229)$
*02/*04	$0.137 \ (0.100, \ 0.174)$
*02/*05 or *02/*06	$0.085 \ (0.056, \ 0.115)$
*04/*04	$0.103\ (0.048,\ 0.158)$
*04/*05 or $*04/*06$	$0.052\ (0.024,\ 0.079)$
Wk16~MBC	
*02/*02	$0.101\ (0.063,\ 0.138)$
*02/*04	$0.062\ (0.042,\ 0.082)$
*02/*05 or *02/*06	$0.050\ (0.032,\ 0.069)$
*04/*04	$0.023\ (0.004,\ 0.043)$
*04/*05 or $*04/*06$	$0.012\ (0.002,\ 0.021)$
Wk3 GC	
*02/*02	$2.870\ (1.747,\ 3.993)$
*02/*04	$1.941\ (1.321,\ 2.561)$
*02/*05 or *02/*06	$1.435\ (0.874,\ 1.996)$
*04/*04	$1.012\ (0.308,\ 1.716)$
*04/*05 or $*04/*06$	$0.506 \ (0.154, \ 0.858)$
Wk9 PB	
*02/*02	$0.771\ (0.548,\ 0.994)$
*02/*04	$0.545 \ (0.352, \ 0.738)$
*02/*05 or *02/*06	$0.386\ (0.274,\ 0.497)$
*04/*04	$0.318 \ (-0.016, \ 0.652)$
*04/*05 or *04/*06	$0.159 \ (-0.008, \ 0.326)$

Table S9

Visit	Ratio (95% CI)
Wk3 GC	2.836 (0.443, 5.229)
Wk4~MBC	3.278 (0.000, 6.613)
Wk8~MBC	$3.259 \ (0.584, 5.934)$
Wk9 PB	$2.425 \ (0.000, \ 5.124)$
Wk10~MBC	$1.650 \ (0.544, \ 2.757)$
$Wk16\ MBC$	$4.360 \ (0.163, \ 8.557)$

Table S10

Visit	Difference (95% CI)	P Value
Wk3 GC	0.929 (0.226, 1.632)	***0.0095***
Wk4 MBC	0.015 (0.003, 0.027)	***0.0169***
Wk8 MBC	0.007 (0.002, 0.012)	***0.0030***
Wk9 PB	0.227 (0.018, 0.435)	***0.0333***
Wk10 MBC	0.034 (-0.009, 0.076)	0.1239
Wk16 MBC	0.039 (0.017, 0.061)	***0.0006***

Table S11

Frequency	Group	Ratio	95% CI	P-value
mRNA	Homozygote	3.9	3.0 to 5.3	< 0.0001
mRNA	Heterozygote	4.2	3.3 to 5.1	0.0001
Unique HCDR3	Homozygote	3.8	3.1 to 4.9	< 0.0001
Unique HCDR3	Heterozygote	3.7	3.0 to 4.5	< 0.0001

Table S12

dose	ID	*02	*04	Ratio
<u>20μg</u>	046	40.5	5.5	7.364
$20\mu g$	077	38.0	13.0	2.923
$20\mu g$	100	140.0	18.0	7.778
$20\mu g$	187	31.0	15.0	2.067
$100 \mu g$	062	88.0	21.0	4.190
$100 \mu g$	064	138.0	19.0	7.263
$100 \mu g$	112	68.0	4.0	17.000
$100 \mu g$	193	221.5	12.5	17.720
	Median			7.313

Table S13

dose	ID	*02	*04	Ratio			
Wk4 MBC							
$20\mu g$	046	6.0	1.0	6.000			
$20\mu g$	077	5.5	1.5	3.667			
20μg	100	30.0	4.0	7.500			
$20\mu \mathrm{g}$	187	2.0	5.0	0.400			
$100 \mu g$	062	1.0	2.0	0.500			
$100 \mu g$	064	27.0	5.0	5.400			
$100 \mu g$	112	8.0	2.0	4.000			
$100 \mu g$	193	37.0	3.0	12.333			
	Median			4.700			
Wk8 MBC							
$20\mu g$	046	1.0	0.0	Inf			
$20\mu g$	077	7.5	0.5	15.000			
$20\mu g$	100	24.0	2.0	12.000			
$20\mu g$	187	2.0	2.0	1.000			
$100 \mu g$	062	3.0	0.0	Inf			
$100 \mu g$	064	22.0	1.0	22.000			
$100 \mu g$	112	17.0	1.0	17.000			
100µg	193	15.0	2.0	7.500			
	Median			16.000			
Wk10 MBC							
$20\mu g$	046	27.0	4.0	6.750			
$20\mu g$	077	1.0	2.0	0.500			
$20\mu g$	100	16.0	7.0	2.286			
$20\mu g$	187	18.0	3.0	6.000			
$100 \mu g$	062	11.0	7.0	1.571			
$100 \mu g$	064	18.0	4.0	4.500			
$100 \mu g$	112	11.0	0.0	Inf			
100µg	193	34.5	2.5	13.800			
	Median			5.250			
Wk16 M							
$20\mu g$	046	6.5	0.5	13.000			
$20\mu g$	077	4.0	1.0	4.000			
$20\mu g$	100	28.0	5.0	5.600			
$20\mu g$	187	9.0	5.0	1.800			
$100 \mu g$	062	8.0	5.0	1.600			
100µg	064	34.0	2.0	17.000			
$100 \mu g$	112	8.0	0.0	Inf			
100µg	193	23.0	3.0	7.667			
	Median			6.633			

Table S13 (continued)

dose	ID	*02	*04	Ratio	
Wk3 GC					
$20\mu g$	046				
$20\mu g$	077	19.0	8.0	2.375	
$20\mu g$	100	32.0	0.0	Inf	
$20\mu g$	187				
$100 \mu g$	062	49.0	7.0	7.000	
$100 \mu g$	064				
$100 \mu g$	112	2.0	0.0	Inf	
100µg	193	31.0	0.0	Inf	
	Median			Inf	
Wk11 GC					
$20 \mu g$	046				
$20\mu g$	077				
$20\mu g$	100	8.0	0.0	Inf	
$20\mu g$	187				
$100 \mu g$	062	16.0	0.0	Inf	
$100 \mu g$	064	1.0	0.0	Inf	
$100 \mu g$	112	22.0	0.0	Inf	
100µg	193	51.0	0.0	Inf	
	Median			Inf	
Wk9 PB					
$20\mu g$	046				
$20\mu g$	077	1.0	0.0	Inf	
$20\mu g$	100	2.0	0.0	Inf	
$20\mu g$	187				
$100 \mu g$	062				
$100 \mu g$	064	36.0	7.0	5.143	
$100 \mu g$	112	0.0	1.0	0.000	
100µg	193	30.0	2.0	15.000	
	Median			15.000	