

Table S1. Characteristics of the Moderna and Pfizer vaccine groups

| Characteristics | Moderna group | Pfizer group | p-value |
|--|-----------------|-----------------|---------------------|
| Fist time point (~3 weeks) | n=16 | n=17 | |
| Sampling days post-dose 2 (mean [range]) | 19.3 [16–26] | 16.3 [12–22] | ^a p=0.02 |
| Age (years, mean [range]) | 35.7 [23–70] | 39.2 [21–69] | ^a p=0.94 |
| Gender (male/female) | 7/9 | 9/8 | ^b p=0.73 |
| Race/ethnicity (A/W/mixed/Hispanic) ^c | 11/5/0/0 | 10/5/0/2 | ^c p=1 |
| Second time point (~3 months) | n=13 | n=14 | |
| Sampling days post-dose 2 (mean [range]) | 94.4 [74–136] | 77.0 [42–124] | ^a p=0.16 |
| Age (years, mean [range]) | 40.9 [20–76] | 37.0 [20–69] | ^a p=0.34 |
| Gender (male/female) | 6/7 | 8/6 | ^b p=0.71 |
| Race/ethnicity (A/W/mixed/Hispanic) ^c | 10/3/0/0 | 9/5/0/4 | ^c p=0.68 |
| Third time point (~7 months) | n=15 | n=12 | |
| Sampling days post-dose 2 (mean [range]) | 215.7 [173–250] | 217.6 [184–247] | ^a p=0.74 |
| Age (years, mean [range]) | 40.4 [23–68] | 39.2 [20–69] | ^a p=0.42 |
| Gender (male/female) | 9/6 | 8/4 | ^b p=1 |
| Race/ethnicity (A/W/mixed/Hispanic) ^c | 9/5/1/0 | 7/5/0/2 | ^c p=1 |
| Sequential samples (from ~3 weeks to 7 months) | n=9 | n=9 | |
| Age (years, mean [range]) | 37.0 [23–68] | 36.9 [21–74] | ^a p=0.65 |
| Gender (male/female) | 4/5 | 6/3 | ^b p=0.64 |
| Race/ethnicity (A/W/mixed/Hispanic) ^c | 7/2/0/0 | 5/4/2/0 | ^c p=0.62 |

^a Mann-Whitney U test, two-tailed.

^b Fisher's exact test, two-tailed.

^c A: Asian, W: White, Fisher's exact test, two-tailed, comparing Asian and White.

Table S2. Days after the second dose of mRNA vaccine when NT₅₀ titers decline to 24

| Moderna group | | | | | | Pfizer group | | | | | |
|--|--------------|---|--------|--------|---------|--|--------------|---|--------|--------|---------|
| ID | Sampling day | Days NT ₅₀ titer=24 ^a | | | | ID | Sampling day | Days NT ₅₀ titer=24 ^a | | | |
| | | D614G | alpha | beta | delta | | | D614G | alpha | beta | delta |
| First sequential samples | | | | | | First sequential samples | | | | | |
| KMC08 | 16 | 390 | 470 | 256 | 368 | VX05 | 13 | 357 | 291 | 186 | 293 |
| VX21 | 16 | 251 | 287 | 186 | 230 | VX19 | 13 | 240 | 286 | 170 | 241 |
| VX22 | 24 | 243 | 303 | 156 | 233 | VX56 | 14 | 239 | 337 | 248 | 245 |
| VX29 | 16 | 316 | 332 | 342 | 266 | VX59 | 13 | 220 | 315 | 253 | 178 |
| VX53 | 26 | 225 | 276 | 87 | 255 | VX73 | 14 | 219 | 176 | 56 | 318 |
| VX30 | 19 | 246 | 313 | 98 | 242 | VX72 | 14 | 216 | 265 | 276 | 165 |
| VX69 | 15 | 286 | 320 | 174 | 371 | VX04 | 12 | 376 | 373 | 209 | 321 |
| VX134 | 19 | 358 | 353 | 189 | 255 | VX09 | 95 | 194 | 356 | 185 | 207 |
| VX136 | 19 | 483 | 446 | 207 | 253 | VX74 | 14 | 232 | 245 | 26 | 100 |
| Single time-point samples | | | | | | Single time-point samples | | | | | |
| KMC22 | 19 | 373 | 320 | 331 | 253 | VX38 | 22 | 361 | 318 | 211 | 273 |
| VX23 | 19 | 428 | 436 | 369 | 314 | VX40 | 22 | 227 | 191 | 103 | 237 |
| VX28 | 20 | 432 | 429 | 347 | 329 | VX42 | 17 | 263 | 251 | 177 | 234 |
| VX51 | 20 | 392 | 348 | 314 | 253 | VX44 | 19 | 354 | 277 | 208 | 314 |
| VX61 | 21 | 394 | 355 | 296 | 273 | VX50 | 21 | 268 | 191 | 151 | 210 |
| VX62 | 21 | 335 | 303 | 280 | 240 | VX85 | 21 | 239 | 182 | 93 | 203 |
| VX16 | 18 | 255 | 327 | 125 | 259 | VX35 | 14 | 214 | 223 | 110 | 251 |
| VX31 | 82 | 415 | 507 | 231 | 400 | VX82 | 19 | 176 | 160 | 64 | 128 |
| VX104 | 87 | 140 | 93 | – | 148 | VX75 | 15 | 222 | 62 | 235 | 290 |
| VX70 | 74 | 281 | 251 | 150 | 160 | VX07 | 88 | 134 | 269 | – | 139 |
| VX97 | 124 | 180 | 143 | – | – | VX88 | 82 | – | 147 | – | 120 |
| VX24 | 206 | 311 | 287 | – | 271 | VX108 | 76 | 268 | 268 | 246 | 246 |
| VX71 | 221 | – | 317 | – | – | VX112 | 118 | 164 | 175 | 126 | 122 |
| VX81 | 207 | 448 | – | 314 | 410 | VX126 | 124 | 273 | 266 | 179 | 212 |
| VX128 | 213 | 356 | 294 | – | 249 | VX79 | 202 | 227 | – | – | – |
| KMC33 | 222 | 298 | 263 | 279 | 258 | VX95 | 184 | 382 | 414 | 326 | 368 |
| VX105 | 173 | 285 | 324 | 205 | 274 | | | | | | |
| Average | | 325 | 324 | 235 | 274 | Average | | 253 | 252 | 174 | 226 |
| Range | | 140-483 | 93-507 | 87-369 | 148-410 | Range | | 134-382 | 62-414 | 26-326 | 100-368 |
| Second and third time-point sequential samples | | | | | | Second and third time-point sequential samples | | | | | |
| KMC08 | 136 | 273 | 397 | 285 | 181 | VX05 | 51 | 262 | 248 | 184 | 231 |
| | 208 | 331 | 345 | 257 | 230 | | 203 | 294 | 304 | – | 225 |
| VX21 | 83 | 239 | 298 | 83 | 195 | VX19 | 97 | 154 | 241 | – | 126 |
| | 234 | 332 | 334 | 351 | 266 | | 245 | – | – | – | – |
| VX22 | 108 | 320 | 393 | – | 345 | VX56 | 46 | 231 | 279 | 199 | 233 |
| | 227 | 339 | 447 | – | 357 | | 219 | 357 | 357 | – | 317 |
| VX29 | 86 | 331 | 361 | 253 | 300 | VX59 | 107 | 149 | 235 | – | 153 |
| | 227 | 337 | 367 | 313 | 299 | | 198 | 244 | 224 | – | 213 |
| VX53 | 58 | 209 | 243 | 85 | 192 | VX73 | 42 | 169 | 102 | 61 | 191 |
| | 211 | 221 | 217 | – | 211 | | 247 | – | 318 | 281 | – |
| VX30 | 86 | 207 | 247 | 92 | 190 | VX72 | 42 | 216 | 226 | 134 | 102 |
| | 212 | 324 | 355 | – | 257 | | 247 | 299 | 337 | – | – |
| VX69 | 131 | 164 | 191 | – | 272 | VX04 | 68 | 371 | 345 | 190 | 349 |
| | 250 | – | – | – | – | | 212 | 414 | 402 | 252 | 348 |
| VX134 | 86 | 359 | 325 | 175 | 155 | VX09 | 195 | – | 433 | 229 | 293 |
| | 212 | 367 | 438 | 281 | 281 | | 244 | 261 | 249 | 266 | 321 |
| VX136 | 86 | 323 | 399 | 165 | 147 | VX74 | 42 | 221 | 277 | – | 119 |
| | 212 | 362 | 373 | – | 261 | | 247 | – | – | – | – |

^a Based on the half-life and NT₅₀ titer against VOC on sampling days, the time after the second dose when NT₅₀ titer declined to 24 was calculated. –, not determined due to NT₅₀ titer <24.

Table S3. Characteristics of COVID-19-recovered plus vaccine and nature infection groups

| Characteristics^a | NI +Moderna or Pfizer group | NI group | p-value |
|--|------------------------------------|-----------------|---------------------|
| Sampling time point (~3 weeks) | n=12 | n=11 | |
| Sampling days post-dose 2 or NI (mean [range]) | 20.2 [16–26] | 25.3 [16–41] | ^b p=0.84 |
| Age (years, mean [range]) | 49.8 [27–70] | 48.2 [24–67] | ^b p=0.16 |
| Gender (male/female) | 5/7 | 7/4 | ^c p=0.41 |
| Race/ethnicity (A/W/mixed/Hispanic) ^d | 7/3/2/0 | 6/4/1/0 | ^c p=1 |

^d NI, natural infection.

^b Mann-Whitney U test, two-tailed.

^c Fisher's exact test, two-tailed.

^d A: Asian, W: White, Fisher's exact test, two-tailed, comparing Asian and White.

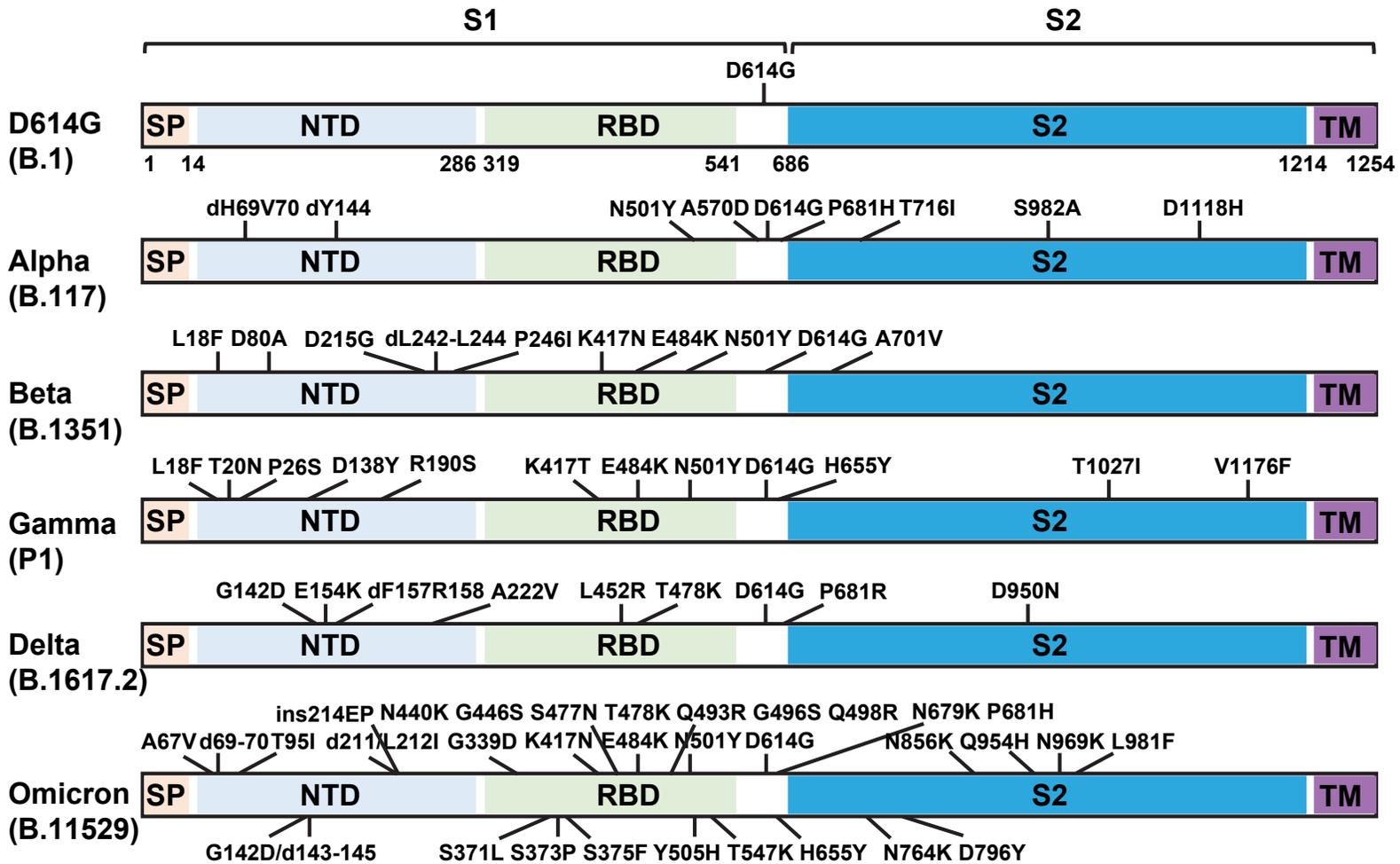


FIG S1 Plasmids expressing the S proteins of VOC and their mutations generated in this study. SP, signal peptide; S1 and S2, S1 and S2 subunits of the S protein; NTD, N-terminal domain; RBD, receptor binding domain; TM, transmembrane domain (43).

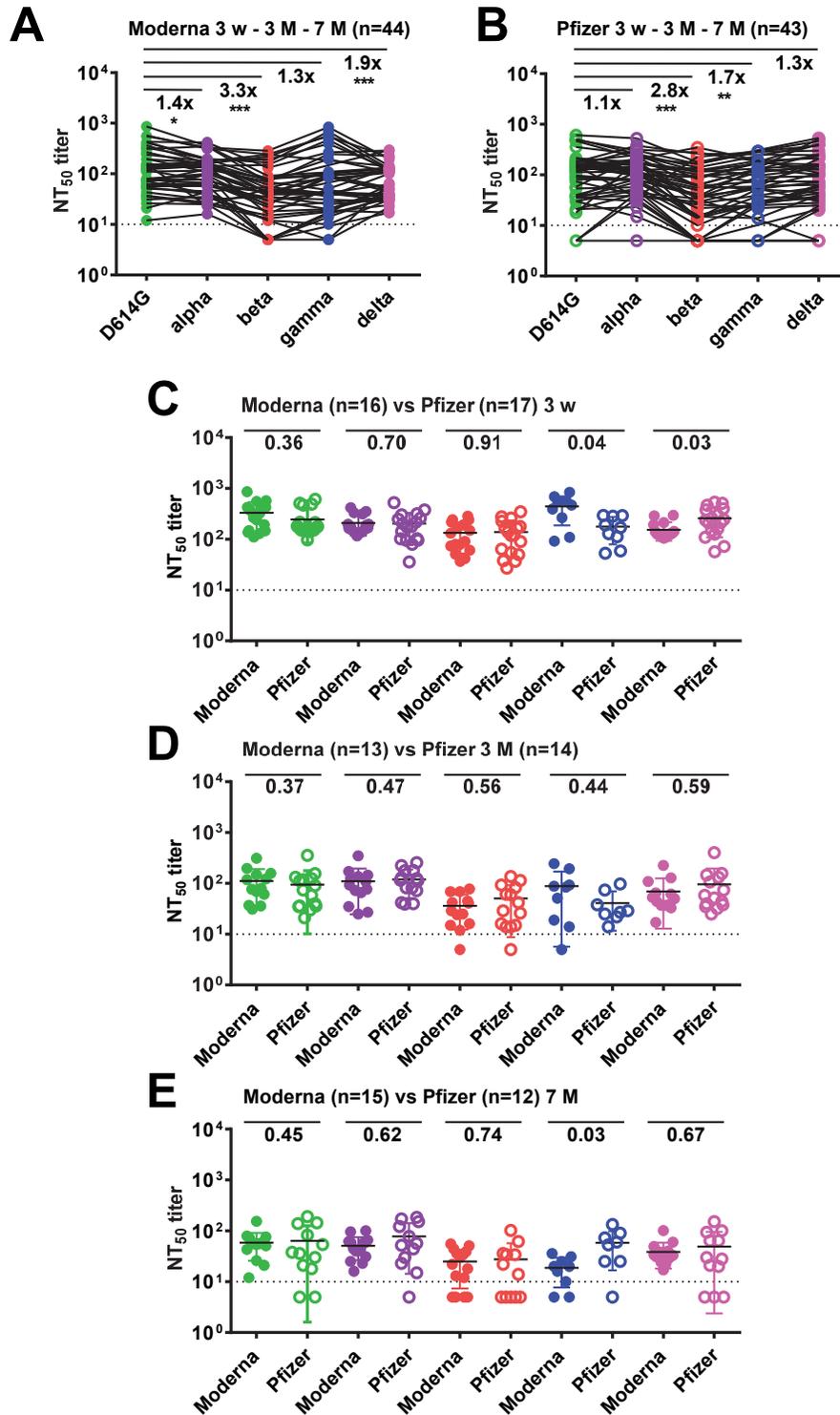


FIG S2 Comparison of NT₅₀ titers against SARS-CoV-2 D614G strain and VOC (alpha, beta, gamma and delta) between COVID-19-naïve participants receiving Moderna and Pfizer vaccine at three time points. NT₅₀ titers were based on pseudovirus neutralization test. (A,B) All samples together (n=44 and n=43 for the Moderna and Pfizer groups, respectively). (C) 3 weeks, (D) 3 months, and (E) 7 months. Number=p-value, the two-tailed Mann-Whitney test. Data are the means of duplicates from one experiment. Dot lines, NT₅₀ titer=10. Color code for each VOC: green, D614G; purple, alpha; red, beta; blue, gamma; pink, delta.

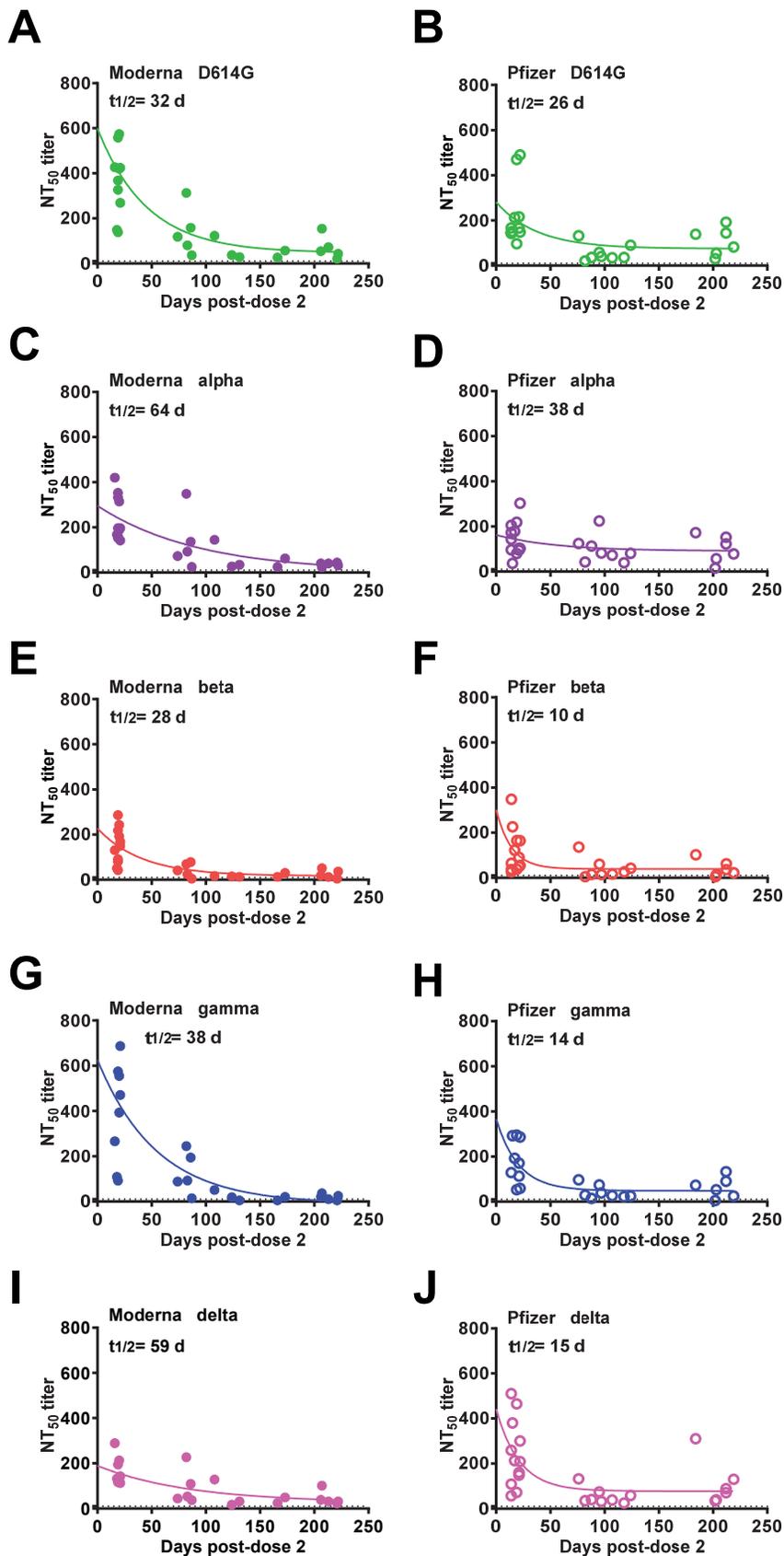


FIG S3 Decline of NT_{50} titers against VOC in the Moderna and Pfizer groups. For the initial samples of the Moderna (A,C,E,G, $n=26$) and Pfizer (B,D,F,H, $n=25$), NT_{50} titers against the D614G strain (A,B) and each VOC including the alpha (C,D), beta (E,F), gamma (G,H) and delta (I,J) variants were plotted against sampling days for each individual following the second dose of the Moderna and Pfizer vaccine. A non-linear regression method (one-phase decay, least squares) was used to determine the best fit curve and estimate the half-life (GraphPad 6.0). Data are the means of duplicates from one experiment. Dot lines, NT_{50} titer=10.

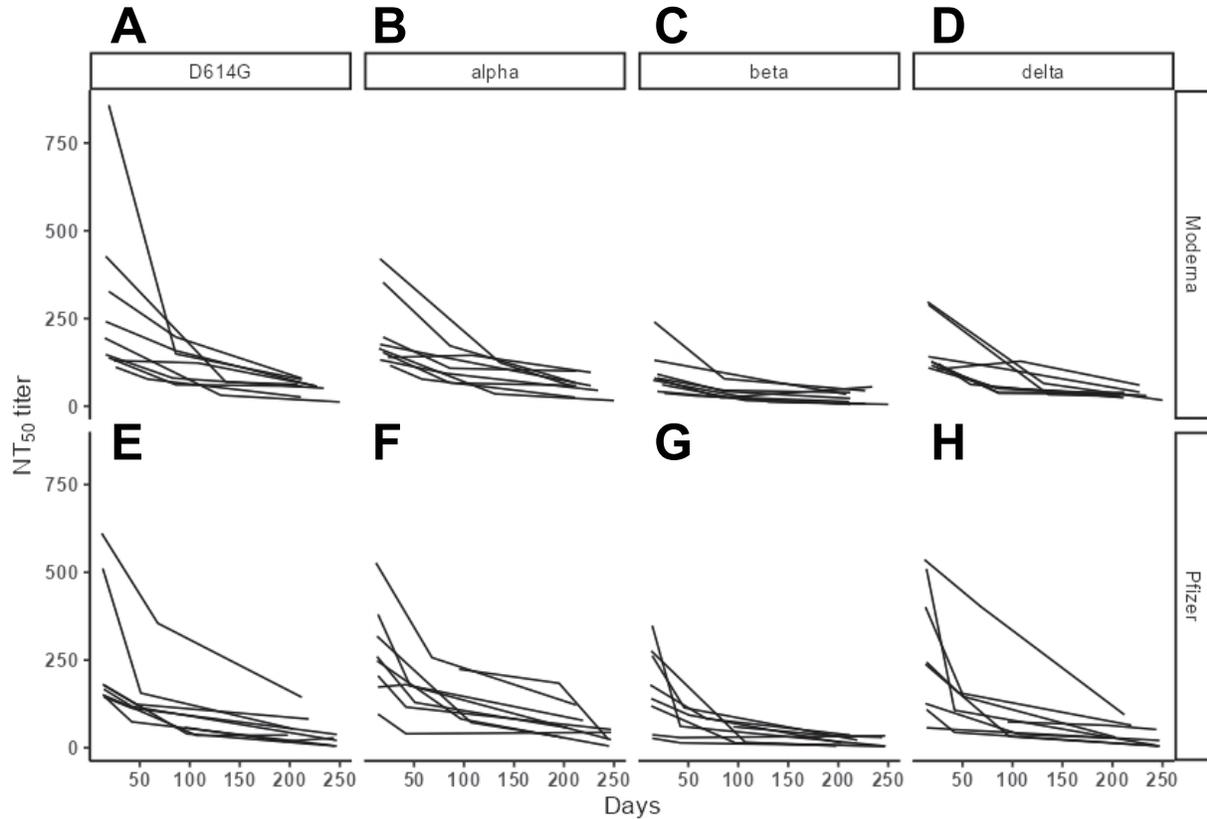


FIG S4 Decline of NT₅₀ titers against VOC in the Moderna and Pfizer groups. For the sequential samples of the Moderna (A–D) and Pfizer (E–H) groups (n=27 from 9 participants in each group), NT₅₀ titers (before log-transformation) against different VOC were plotted against sampling days for each individual following the second dose of the Moderna and Pfizer vaccine. Data are the means of duplicates from one experiment.

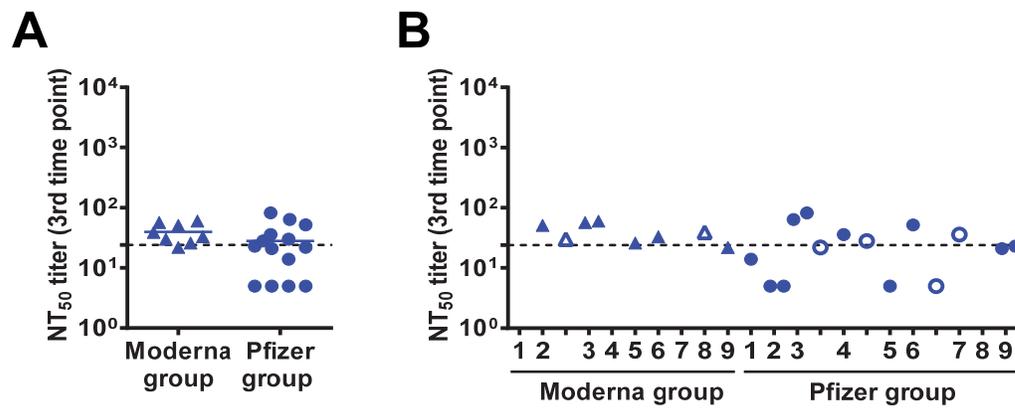
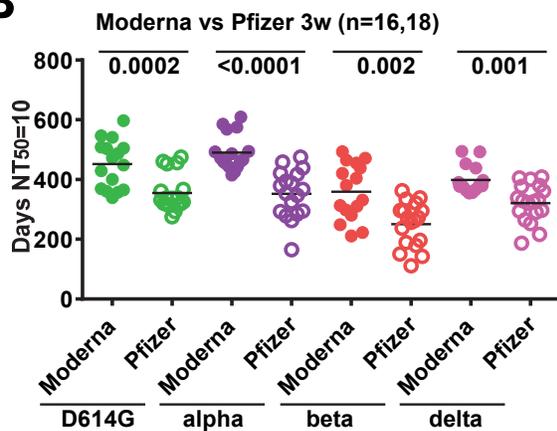


FIG S5 Comparison of NT₅₀ titers of the third time point, which were within 30 days of the projected times based on the first time point, to 24. (A,B) Available data from the Moderna and Pfizer vaccinees as a group (A) or from each individual (B). NT₅₀ titers against either D614G strain, alpha, beta or delta VOC were included in the analysis. Data are the means of duplicates from one experiment. Dash lines indicate NT₅₀ titer=24. For panel B, NT₅₀ titers higher or lower than 24 when the third time points were before or after the projected times, respectively, are presented as close symbol; otherwise open symbols. Numbers indicate each individual.

A

| Moderna group | | | | | | Pfizer group | | | | | |
|---------------------------|--------------|--------------------------------|---------|---------|---------|---------------------------|--------------|--------------------------------|---------|---------|---------|
| ID | Sampling day | Days NT ₅₀ titer=10 | | | | ID | Sampling day | Days NT ₅₀ titer=10 | | | |
| | | D614G | alpha | beta | delta | | | D614G | alpha | beta | delta |
| First sequential samples | | | | | | First sequential samples | | | | | |
| KMC08 | 16 | 503 | 609 | 380 | 492 | VX05 | 13 | 456 | 394 | 272 | 380 |
| VX21 | 16 | 365 | 425 | 310 | 354 | VX19 | 13 | 339 | 388 | 256 | 329 |
| VX22 | 24 | 357 | 442 | 280 | 356 | VX56 | 14 | 338 | 439 | 334 | 332 |
| VX29 | 16 | 429 | 471 | 465 | 390 | VX59 | 13 | 318 | 417 | 338 | 265 |
| VX53 | 26 | 339 | 415 | 211 | 378 | VX73 | 14 | 317 | 278 | 142 | 405 |
| VX30 | 19 | 360 | 452 | 222 | 366 | VX72 | 14 | 315 | 367 | 362 | 252 |
| VX69 | 15 | 400 | 459 | 298 | 494 | VX04 | 12 | 475 | 475 | 295 | 408 |
| VX134 | 19 | 472 | 492 | 313 | 378 | VX09 | 95 | 293 | 458 | 271 | 294 |
| VX136 | 19 | 597 | 585 | 331 | 377 | VX74 | 14 | 331 | 347 | 111 | 187 |
| Single-time-point samples | | | | | | Single-time-point samples | | | | | |
| KMC22 | 19 | 487 | 459 | 455 | 377 | VX38 | 22 | 460 | 421 | 296 | 361 |
| VX23 | 19 | 541 | 575 | 493 | 438 | VX40 | 22 | 325 | 293 | 189 | 325 |
| VX28 | 20 | 546 | 568 | 471 | 452 | VX42 | 17 | 361 | 353 | 262 | 321 |
| VX51 | 20 | 505 | 486 | 438 | 377 | VX44 | 19 | 452 | 379 | 294 | 401 |
| VX61 | 21 | 508 | 494 | 420 | 397 | VX50 | 21 | 367 | 294 | 237 | 297 |
| VX62 | 21 | 448 | 442 | 404 | 364 | VX85 | 21 | 337 | 284 | 179 | 290 |
| VX16 | 18 | 369 | 466 | 248 | 383 | VX35 | 14 | 313 | 325 | 196 | 338 |
| VX31 | 82 | 529 | 646 | 355 | 523 | VX82 | 19 | 275 | 262 | 150 | 216 |
| VX104 | 87 | 253 | 232 | – | 272 | VX75 | 15 | 320 | 165 | 321 | 377 |
| VX70 | 74 | 394 | 389 | 273 | 283 | VX07 | 88 | 232 | 371 | 128 | 226 |
| VX97 | 124 | 294 | 282 | 181 | 199 | VX88 | 82 | 165 | 250 | – | 207 |
| VX24 | 206 | 425 | 426 | 289 | 395 | VX108 | 76 | 366 | 370 | 332 | 333 |
| VX71 | 221 | 317 | 456 | 221 | 332 | VX112 | 118 | 262 | 277 | 212 | 209 |
| VX81 | 207 | 562 | 339 | 437 | 534 | VX126 | 124 | 371 | 368 | 265 | 299 |
| VX128 | 213 | 469 | 433 | 239 | 373 | VX79 | 202 | 326 | 249 | – | 271 |
| KMC33 | 222 | 411 | 402 | 403 | 382 | VX95 | 184 | 480 | 516 | 412 | 455 |
| VX105 | 173 | 399 | 463 | 328 | 398 | | | | | | |
| Average | | 434 | 458 | 339 | 387 | Average | | 344 | 350 | 255 | 311 |
| Range | | 253-597 | 232-646 | 211-493 | 199-534 | Range | | 165-480 | 165-516 | 111-412 | 187-455 |

B



C

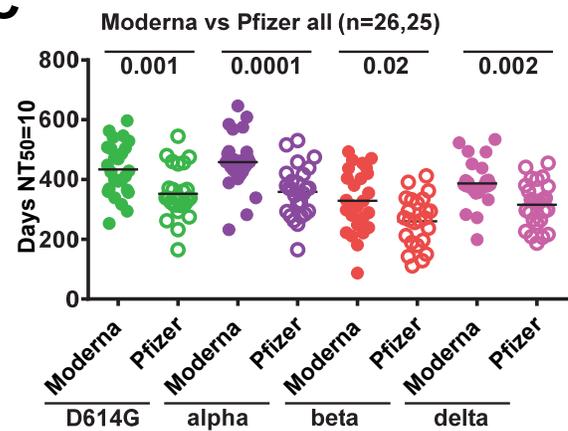


FIG S6 The time following the second dose of the Moderna or Pfizer vaccine when NT₅₀ titers against each VOC declined to 10. (A) Based on the half-life and NT₅₀ titer against each VOC on different sampling days (including the first sequential samples and single-time-point samples), the time after the second dose when NT₅₀ titer declined to 10 was calculated for each participant in the Moderna and Pfizer groups. –, not determined due to NT₅₀ titer <10. (B,C) Comparison of the time when NT₅₀ titers declined to 10 for each VOC between the Moderna and Pfizer groups based on the 3 week samples (B) and all initial samples (C). Number=p-value, the two-tailed Mann-Whitney U test.

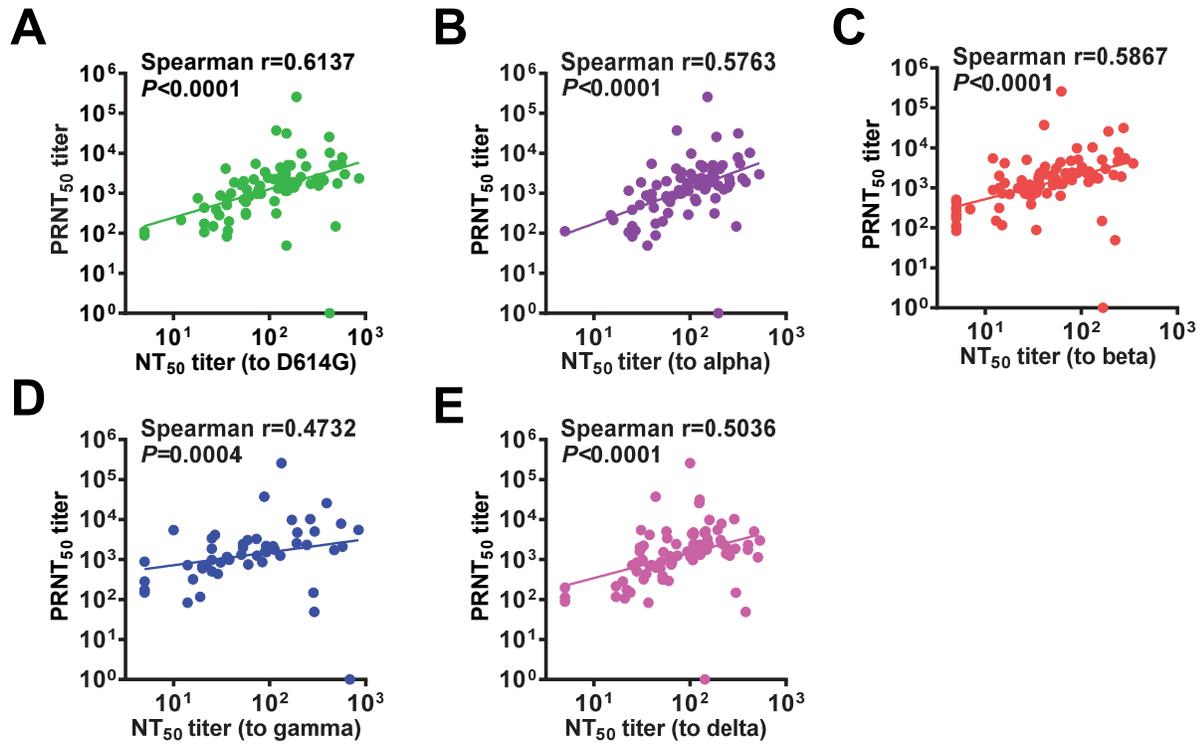


FIG S7 Correlation between PRNT₅₀ titers and NT₅₀ titers against different VOC. (A–E) Relationship between PRNT₅₀ titers to USA-WA1 strain (containing D614) and pseudovirus NT₅₀ titers against D614G strain (A), alpha (B), beta (C), gamma (D) or delta (E) variant. Data are the means of duplicates from one experiment. The two-tailed Spearman correlation test (GraphPad 6.0).

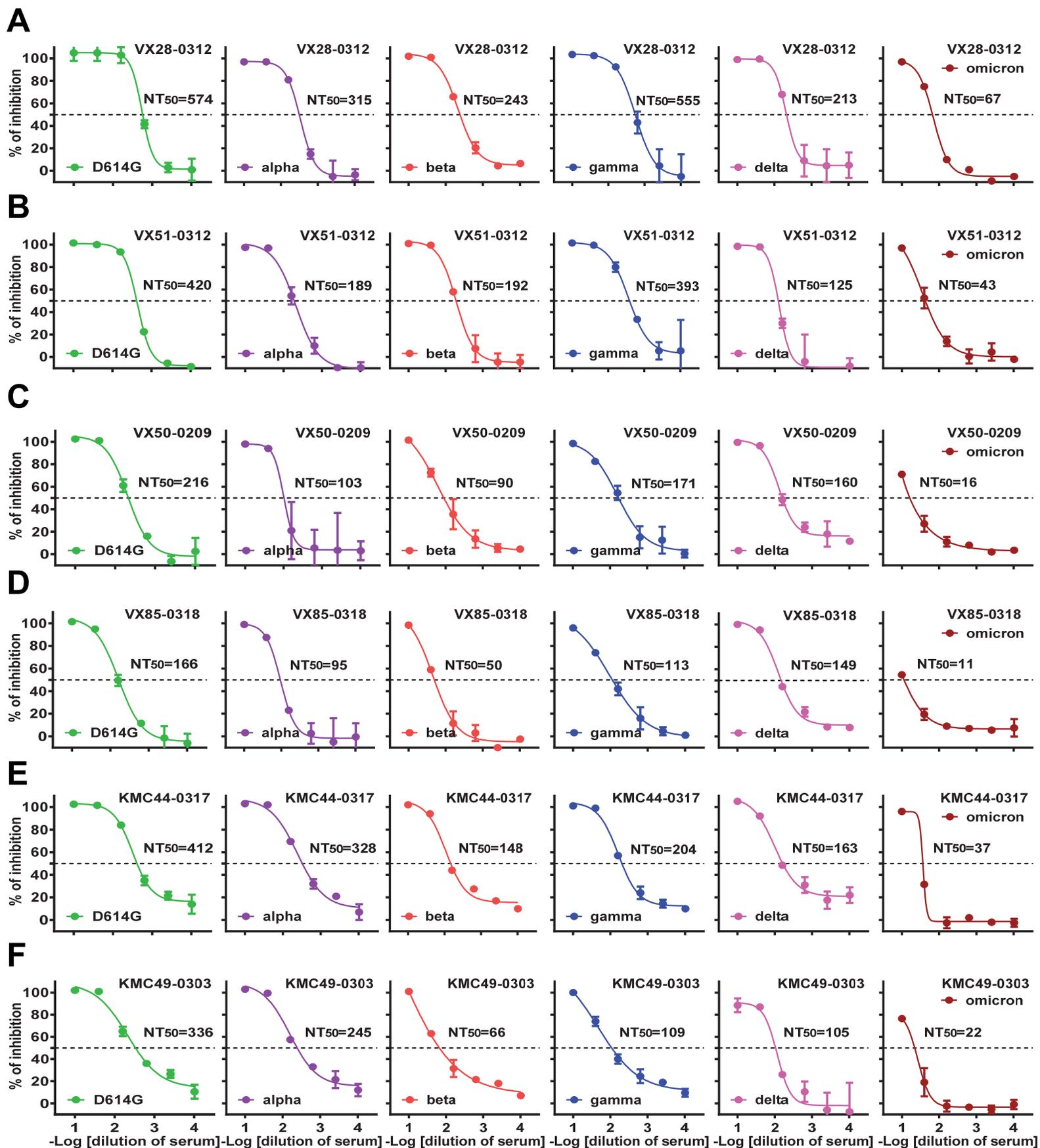


FIG S8 Neutralization curves and NT_{50} titers against different VOC in vaccinees by SARS-CoV-2 pseudovirus neutralization test. (A–F) Vaccinees including COVID-19-naïve participants after two doses of Moderna (A,B) or Pfizer (C,D) vaccine and COVID-19 recovered participants after two doses of Moderna vaccine (E,F) vaccine. Data are the means standard deviations of duplicates from one experiment.