

Supplementary Table 1: Characteristics of the population

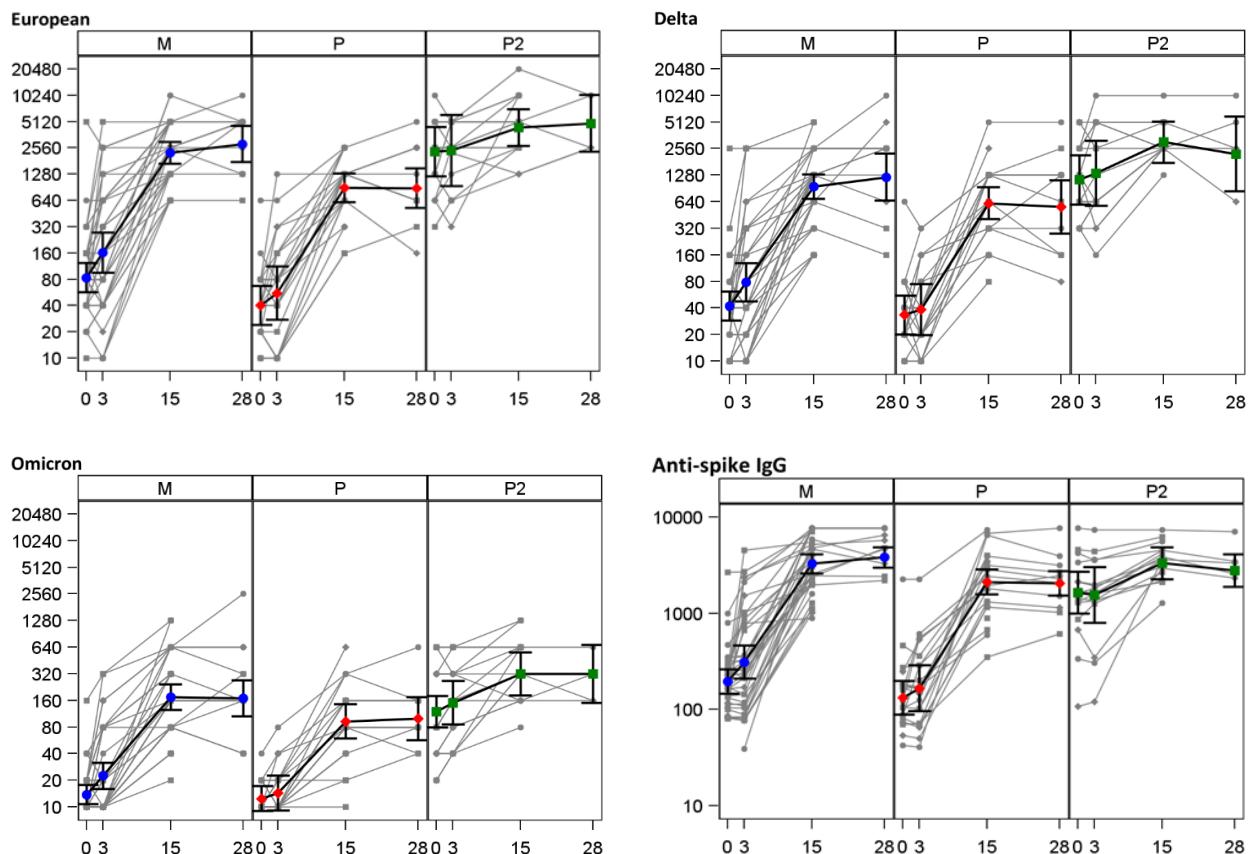
	SARS-CoV-2 naïve		SARS-CoV-2 infected before the primary vaccination	
	CoviCompareM (M) mRNA-1273 N = 34	CoviCompareP (P) BNT162b2 N = 19	CoviCompareP (P2) BNT162b2 N = 12	
Age (years)	71.5 [68.2, 76.0] (66 to 84)	72.0 [66.5, 75.5] (65 to 84)	0.56 <sup>1</sup>	70.0 [67.0, 75.0] (65 to 81)
Male sex	20 (59%)	8 (42%)	0.24 <sup>2</sup>	8 (67%)
Body Mass Index (BMI)	25.3 [22.6, 27.6]	24.7 [22.3, 26.6]	0.48 <sup>1</sup>	24.0 [20.7, 25.0]
BMI > 30	5 (15%)	0 (0%)	0.15 <sup>3</sup>	0 (0%)
Months between primary vaccination and boost	7.4 [7.0, 7.5]	6.8 [6.4, 7.0]	<0.001 <sup>1</sup>	7.8 [7.5, 7.9]
3 days after boost, days	4 (2 to 6)	3 (2 to 5)	0.12 <sup>1</sup>	3 (2 to 4)
15 days after boost, days	15 (12 to 17)	14 (13 to 20)	0.25 <sup>1</sup>	14 (14 to 17)
28 days after boost, n days	12 28 (26 to 29)	11 28 (25 to 28)	0.26 <sup>1</sup>	5 28 (27 to 28)

n (%); Median [IQR] (min to max)

<sup>1</sup>Wilcoxon rank sum test <sup>2</sup>Pearson's Chi-squared test <sup>3</sup>Fisher's exact test

Supplementary Table 2: **Geometric means (MMRM model estimates)**

Individual profiles are connected with grey lines and model-based geometric means estimates with 95% confidence intervals are connected with black lines at Day 0=before boost, 3, 15, 28 days after boost



#### European strain (neutralization titer)

Timepoint	CoviCompare-M mRNA-1273 SARS-CoV-2 naïve (M) N=34	CoviCompare-P BNT162b2 SARS-CoV-2 naïve (P) N=19	CoviCompare-P BNT162b2 previously SARS CoV-2 infected (P2) N=12
Before boost (Day 0), n	33	18	12
<b>Geometric mean</b>	<b>83.63</b>	<b>40.08</b>	<b>2317.65</b>
95%CI	56.79, 123.17	23.99, 66.96	1212.41, 4430.42
Day 3, n	34	18	12
<b>Geometric mean</b>	<b>162.30</b>	<b>55.66</b>	<b>2392.06</b>
95%CI	95.68, 275.32	27.46, 112.81	937.77, 6101.66
Day 15, n	34	19	12
<b>Geometric mean</b>	<b>2251.43</b>	<b>890.57</b>	<b>4375.14</b>
95%CI	1692.60, 2994.74	608.61, 1303.14	2701.65, 7085.25
Day 28, n	12	11	5
<b>Geometric mean</b>	<b>2828.78</b>	<b>883.94</b>	<b>4878.70</b>
95%CI	1753.91, 4562.38	523.47, 1492.62	2302.26, 10338.41

### Delta variant (neutralization titer)

Timepoint	CoviCompare-M mRNA-1273 SARS-CoV-2 naïve (M) N=34	CoviCompare-P BNT162b2 SARS-CoV-2 naïve (P) N=19	CoviCompare-P BNT162b2 previously SARS CoV-2 infected (P2) N=12
Before boost (Day 0), n	33	19	12
<b>Geometric mean</b>	<b>42.06</b>	<b>33.31</b>	<b>1135.41</b>
95%CI	28.68, 61.70	20.06, 55.32	598.24, 2154.94
Day 3, n	34	19	12
<b>Geometric mean</b>	<b>78.52</b>	<b>38.55</b>	<b>1350.24</b>
95%CI	47.59, 129.56	19.74, 75.27	580.60, 3140.14
Day 15, n	34	19	12
<b>Geometric mean</b>	<b>944.41</b>	<b>616.76</b>	<b>3031.19</b>
95%CI	688.45, 1295.52	404.55, 940.31	1777.21, 5169.96
Day 28, n	12	12	5
<b>Geometric mean</b>	<b>1217.55</b>	<b>558.93</b>	<b>2228.93</b>
95%CI	659.30, 2248.47	277.66, 1125.13	838.54, 5924.74

### Omicron BA.1 variant (neutralization titer)

Timepoint	CoviCompare-M mRNA-1273 SARS-CoV-2 naïve (M) N=34	CoviCompare-P BNT162b2 SARS-CoV-2 naïve (P) N=19	CoviCompare-P BNT162b2 previously SARS CoV-2 infected (P2) N=12
Before boost (Day 0), n	33	19	12
<b>Geometric mean</b>	<b>13.82</b>	<b>12.45</b>	<b>119.95</b>
95%CI	10.80, 17.69	8.99, 17.24	79.40, 181.23
Day 3, n	34	19	12
<b>Geometric mean</b>	<b>22.60</b>	<b>14.40</b>	<b>151.13</b>
95%CI	16.10, 31.70	9.16, 22.64	85.38, 267.51
Day 15, n	34	19	12
<b>Geometric mean</b>	<b>173.54</b>	<b>92.58</b>	<b>320.23</b>
95%CI	124.35, 242.20	59.31, 144.51	182.53, 561.82
Day 28, n	12	12	5
<b>Geometric mean</b>	<b>169.49</b>	<b>99.78</b>	<b>319.78</b>
95%CI	105.79, 271.56	57.40, 173.44	149.93, 682.03

### Anti-spike IgG (BAU/mL)

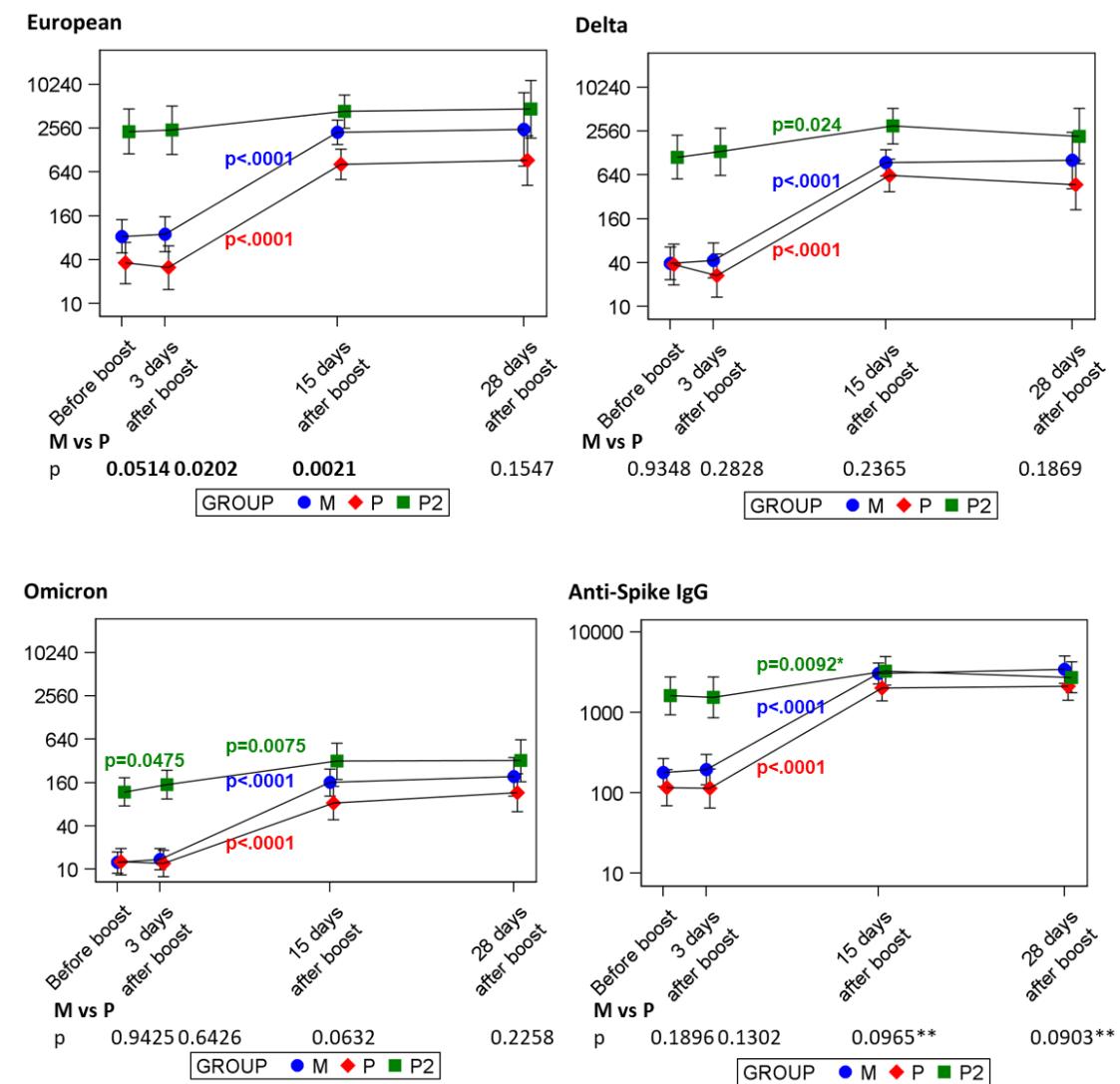
Timepoint	CoviCompare-M mRNA-1273 SARS-CoV-2 naïve (M) N=34	CoviCompare-P BNT162b2 SARS-CoV-2 naïve (P) N=19	CoviCompare-P BNT162b2 previously SARS CoV-2 infected (P2) N=12
Before boost (Day 0), n	33	18	12
<b>Geometric mean</b>	<b>195.77</b>	<b>132.24</b>	<b>1637.59</b>
95%CI	145.60, 263.21	88.46, 197.67	997.00, 2689.76
Day 3, n	34	18	12
<b>Geometric mean, n</b>	<b>311.72</b>	<b>166.85</b>	<b>1553.70</b>
95%CI	209.35, 464.13	96.75, 287.74	794.51, 3038.32
Day 15, n	34	19	12
<b>Geometric mean</b>	<b>3272.47</b>	<b>2121.77</b>	<b>3319.86</b>
95%CI	2604.99, 4110.96	1565.07, 2876.50	2258.85, 4879.25
Day 28, n	12	11	5
<b>Geometric mean</b>	<b>3813.58</b>	<b>2057.77</b>	<b>2793.87</b>
95%CI	3003.65, 4841.90	1532.71, 2762.69	1890.07, 4129.84

**Supplementary Figure 1: Geometric means and 95% Confidence Intervals (MMRM model estimates) for participants with 2-4 days post boost only (n=48)**

Neutralizing antibody titers for European strain, Delta, Omicron (BA.1) variants and anti-spike IgG (BAU/mL) in SARS-CoV-2 naïve mRNA-1273 recipients (M in blue), BNT162b2 recipients (P in red) and previously SARS-CoV-2 infected BNT162b2 recipients (P2 in green) before booster dose (day 0) and at 3, 15 and 28 days after boost.

Timepoints comparisons: significant p-values within each group are shown (<.05)

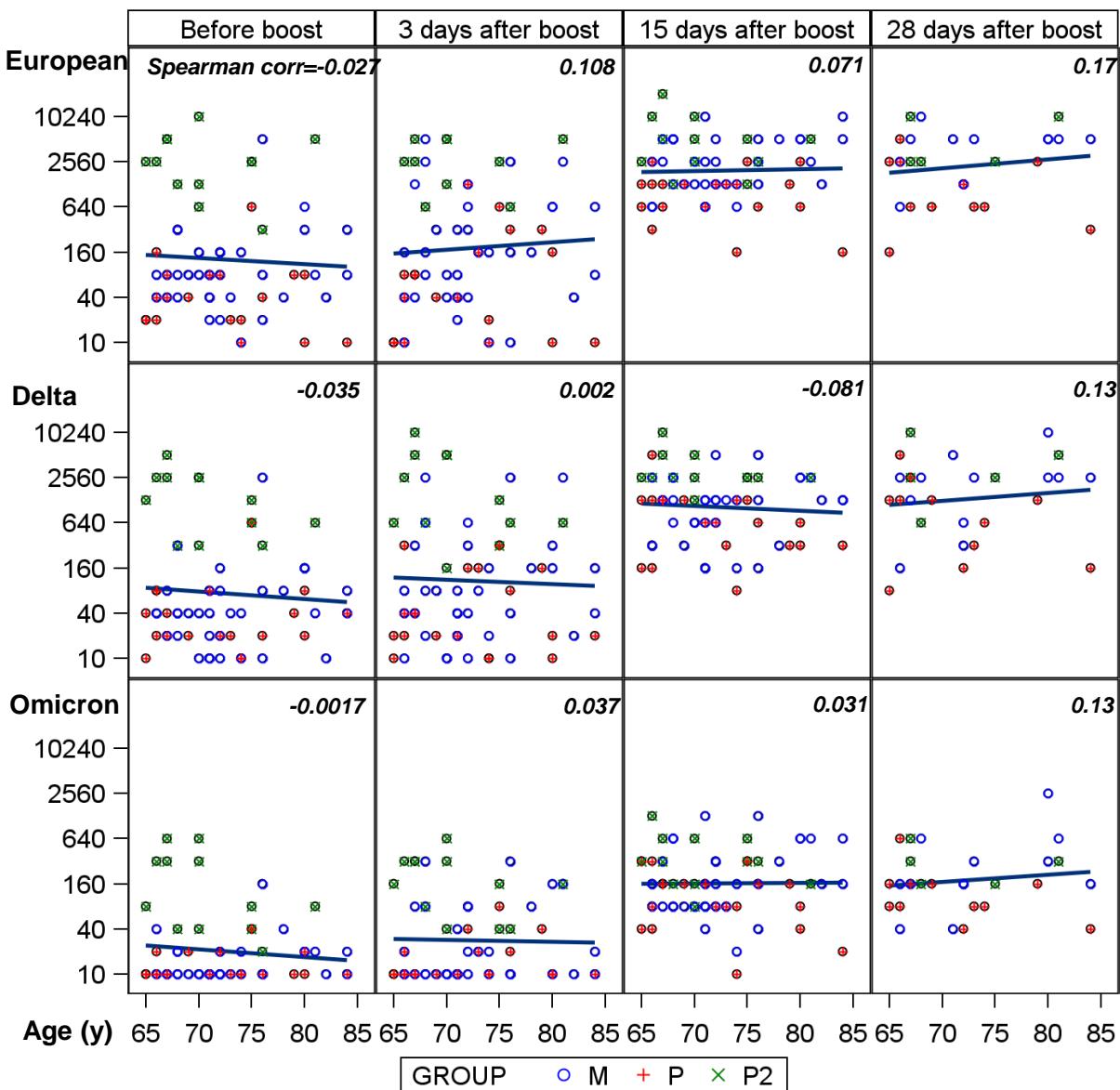
M vs P comparisons: p-values at each timepoint below the figures



\* no time effect slice (in P2)

\*\* no group effect slice (at D15 and D28)

**Supplementary Figure 2: Neutralizing antibody titers for European strain, Delta, Omicron (BA.1) variants against the age at the first dose of vaccine (years) in SARS-CoV-2 naïve mRNA-1273 recipients (M in blue), BNT162b2 recipients (P in red) and previously SARS-CoV-2 infected BNT162b2 recipients (P2 in green) before booster dose and days 3, 15 and 28 after boost. Regression slopes and Spearman correlations are not statistically significant.**



Supplementary Figure 3: Neutralizing antibody titers for European strain, Delta, Omicron (BA.1) variants at D1 (1<sup>st</sup> dose); D29 (28 days after 1<sup>st</sup> dose and 2<sup>nd</sup> dose in SARS-CoV-2 naïve); D57 (28 days after 2<sup>nd</sup> dose). SARS-CoV-2 naïve mRNA-1273 recipients (M in blue), BNT162b2 recipients (P in red) and previously SARS-CoV-2 infected BNT162b2 recipients (P2 in green). Sample sizes (N) and medians are indicated at the bottom and p-values are for paired signed rank tests between D15 post-boost and D28 post primary-vaccination when n>5. Black triangles show administration of vaccine.

