Appendix:

Biparatopic Nanobodies protect mice from lethal challenge with SARS-CoV-2 variants of concern

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Appendix Table S1. Nb combinations for bipNbs

| Biparatopic (Bip)/ bivalent Nb | Single Nb combination       |
|--------------------------------|----------------------------|
| NM1267                         | NM1230-(G₄S)₄-NM1226       |
| NM1268                         | NM1228-(G₄S)₄-NM1226       |
| NM1251 (Control)               | Pep-Nb-(G₄S)₄-Pep-Nb       |
## Appendix Table S2. Primer Sequences

| Primer    | Sequence 5’-3’                                                                 |
|-----------|-------------------------------------------------------------------------------|
| NM1228Nfor| GGACGTCTCAACTCTGACGTGCAGCTGGTGGAAT                                             |
| NM1228Nrev| CACCACCAGCCAGATCCACCAGCCACCTGTACCTCCGCTCTTGAGGCACGTTGACCCAGGACC               |
| NM1226Cfor| GGTGGATCTGGCGGTGGTGGAAGTGGTGCGGAGGTAGTCAAGGTGCAGCTGGTGGAAT                   |
| NM1226Crev| GGGGAATTCAGTGAAGTGAGTGAGTGAGAGGACACGGTGACCGGGCC                              |
| RBDfor    | ATATCTAGAGCCACCATGTGCTGTTCGTTTCTGG                                            |
| RBDrev    | AAGATCTGCTAGCTGAGCTGCC                                                       |
| N501Yfor  | GGTGGTACATATGCGGTGGGCTGCTATAGCN                                              |
| N501Yrev  | CCACGCCATATGTGCGGCTGAAAGCCGTAG                                               |
| L452Rfor  | GGCACTACATATCCCGGTAATATAGGAGGCTCCGGCCAGG                                    |
| L452Rrev  | CGGTTACATATCCCGGTAATATAGGAGGCTCCGGCCAGG                                    |
| V367Ffor  | CGACTACTCTCTCTGAATCCCGGTACCGGCTCCGGCCAGG                                     |
| V367Frev  | CGGAGTTGTCAGCGAAGAGGTAGTGCAGCGCAGCCAGG                                      |
| T478Kfor  | CCGGCAGCAAGCCTTGTAACCGCGTGGAGGAGG                                          |
| T478Krev  | CGTTACAGGCTTCAGCGCAGAGGAGGAGG                                               |
| E484Kfor  | GTAACGGCGGCGAAAAGGCTTCAACTGCTACTTCC                                          |
| E484Krev  | GCACTGGAACCCCTTTCCACGCGTTACAGGAGG                                          |
| K417Nfor  | GACAGACAGCCACATCAGCGACTACCAACTACAAGG                                         |
| K417Nrev  | GTCGATAGCGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGT |
| K417Tfor  | GACAGACAGCCACATCAGCGACTACCAACTACAAGG                                         |
| K417Trev  | GTCGATAGCGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGTGTGCGT |