

Figure S1 (related to main Figure 2A, D). Comparison of the immunogenicity of S0 and S0*. NAb titres against prototypic (grey circles), VOC Beta (blue circles), VOC Gamma (orange circles) and VOC Delta (purple circles) spike pseudotyped virus on day 21 after vaccination with prototype YF-S0 ($n=32$) and YF-S0* ($n=24$). Bar graphs denote median \pm IQR. Dotted line represents the lower limit of quantification. Differences between groups were analyzed using non-parametric two-tailed Mann-Whitney.

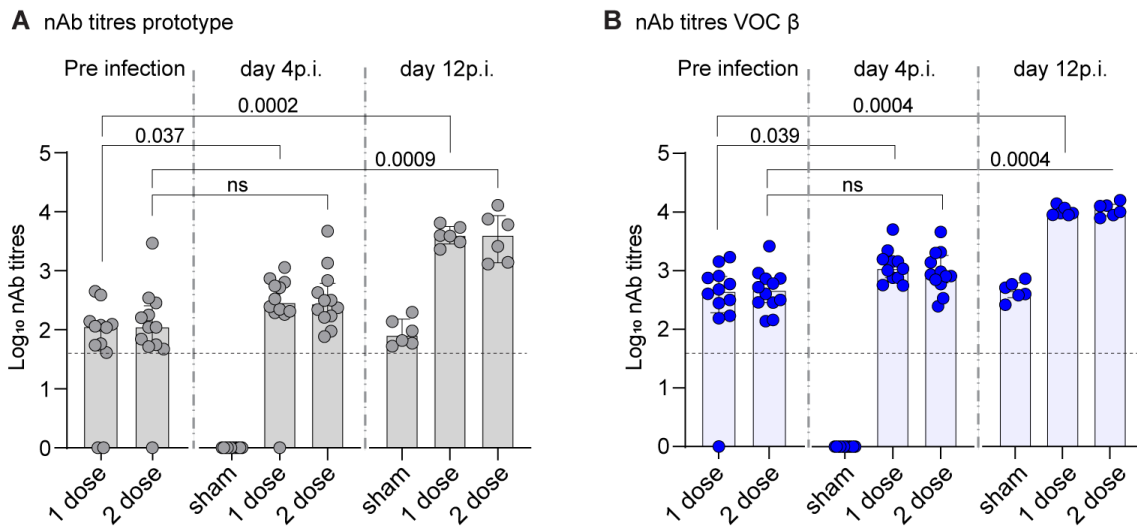


Figure S2 (related to main Figure 3). Comparison of nAb levels pre and post challenge. A, B, nAb titres against prototypic (grey circles) (A) and VOC Beta (blue circles) (B) spike pseudotyped virus on day 21 after vaccination with 1 ($n=12$) or 2 doses ($n=12$) of YF-S0* (pre infection) and day 4 ($n=24$ vaccinated and $n=12$ sham vaccinated) or day 12 ($n=12$ vaccinated and $n=6$ sham vaccinated) after infection. Bar graphs denote median \pm IQR. Dashed line (horizontal) represents the lower limit of quantification. Pre infection nAb data have been reused from main Figure 3B. Differences between groups were analyzed using non-parametric Kruskal Wallis test uncorrected for ties.

A nAb titres YF-S0 **B** nAb titres YF-S0*

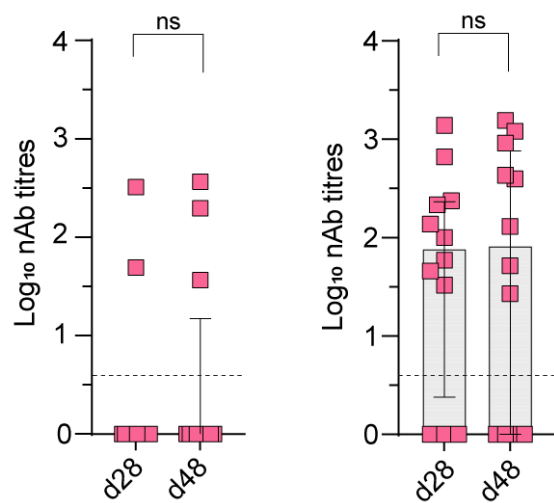


Figure S3 (related to main Figure 4). Increased potency of vaccine candidate S0* against VOC Omicron. **A, B**, nAb titres against omicron spike pseudotyped virus (red squares) on day 28 and day 48 after vaccination with two doses (day 0 and day 7) of prototype YF-S0 ($n=12$) (**A**) or YF-S0* ($n=12$) (**B**). Bar graphs denote median \pm IQR. Dashed line represents the lower limit of quantification. Differences between groups were analyzed using non-parametric two-tailed Mann-Whitney.