



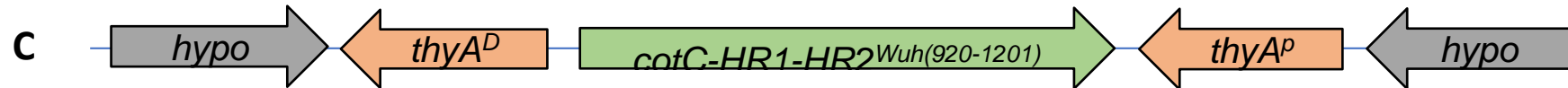
CotB-RBD^{Wuh(319-530)} MW = 61.8 kDa.

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SNSFISSKSYSSSKSSKRSKSSDDQSSKSGRSSRSKSSSKSSKRSKSSDYQSSKSGRSSRSKSSSKSSKRSKSSDYQSSKSSKRSRPSRVQPTESIVRFPNITNLCPFGEVFNATRFA
SVYAWNRKRISNCVADYSVLVNSASFSTFKCYGVSPTKLNDLCFTNVYADSFVIRGDEVQRQIAPGQTGKIADYNYKLPDDFTGCVIAWNSNNLDSKVGGNYNYLYRLFRKSNLKPFERDIST
EIYQAGSTPCNGVEGFNCYFPLQSYGFQPTNGVGYQPYRVVLSFELLHAPATVCGPKKS



CotB-RBD^{Omi(316-538)} MW = 63.36 kDa.

MSKRRMKYHSNNEISYYNFLHSMKDKIVTVYRGGPESKKGKLTAVKSDYIALQAEKKIIYYQLEHVKSITEDTNNSTTTIETEEMLDADDFHSLIGHLINQSVQFNQGGPESKKGRLVWLGGD
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SNSFISSKSYSSSKSSKRSKSSDDQSSKSGRSSRSKSSSKSSKRSKSSDYQSSKSGRSSRSKSSSKSSKRSKSSDYQSSKSSKRSRPSRVQPTESIVRFPNITNLCPFD^DEVFNATRFA
SVYAWNRKRISNCVADYSVLVNL^LAP^FFTFKCYGVSPTKLNDLCFTNVYADSFVIRGDEVQRQIAPGQTG^NIADYNYKLPDDFTGCVIAWNSN^KLDSKV^SGNVNYLYRLFRKSNLKPFERDIST
EIYQAG^{NK}PCNGV^AGFNCYFPL^RSY^SFR^PTYGVGH^HQPYRVVLSFELLHAPATVCGPKKSTNLVKNKCVNF

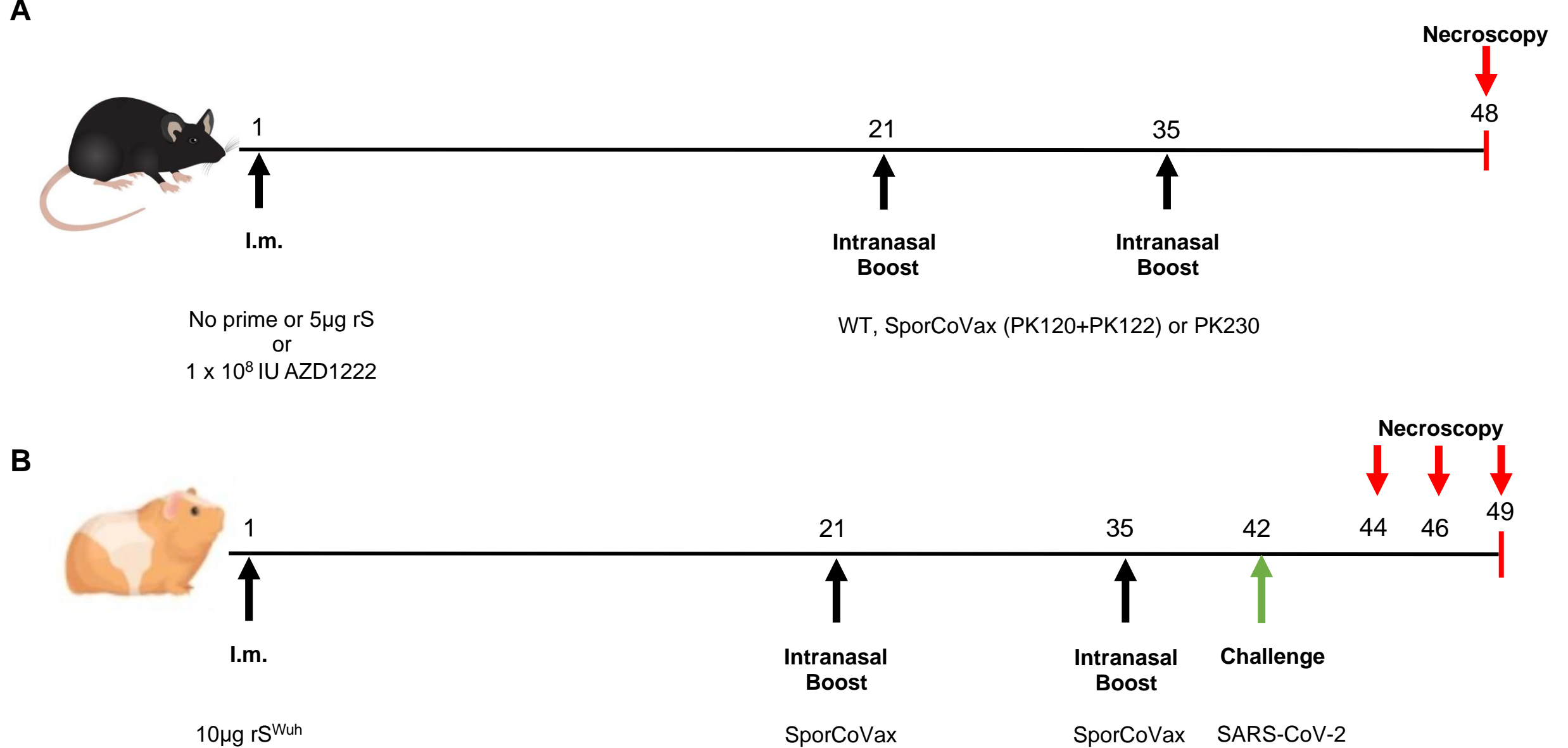


CotC-HR1-HR2^{Wuh(920-1201)} MW = 39.87 kDa.

MGYYKKYKEEYYTVKKTYKKYYEYDKKDYDCDYDKKYDDYDKKYDHDKKDYDYVVEYKKHKKHYQKLIANQFN^SAI^GKI^QDSLS^STASALGKLQDVVNQNAQALNTLVKQLSSNFGAIS
SVLNDILSRDKVEAEVQIDRLITGRLQSLQTYVTQQLIRAAEIRASANLAATKMSECVLGQSKRVDFCGKGYHLMSPQSAPHGVVFLHVTYVPAQEKNFTTAPAICHGKAHFPREGVVF
SNGTHWFVTQRNFYEPQIITDNTFVSGNCDVVIGIVNNTVYDPLQPELDSFKEELDKYFKNHTSPD^VDLGDISGINASV^VNIQKEIDRLNEVAKNLN^ESLIDLQ

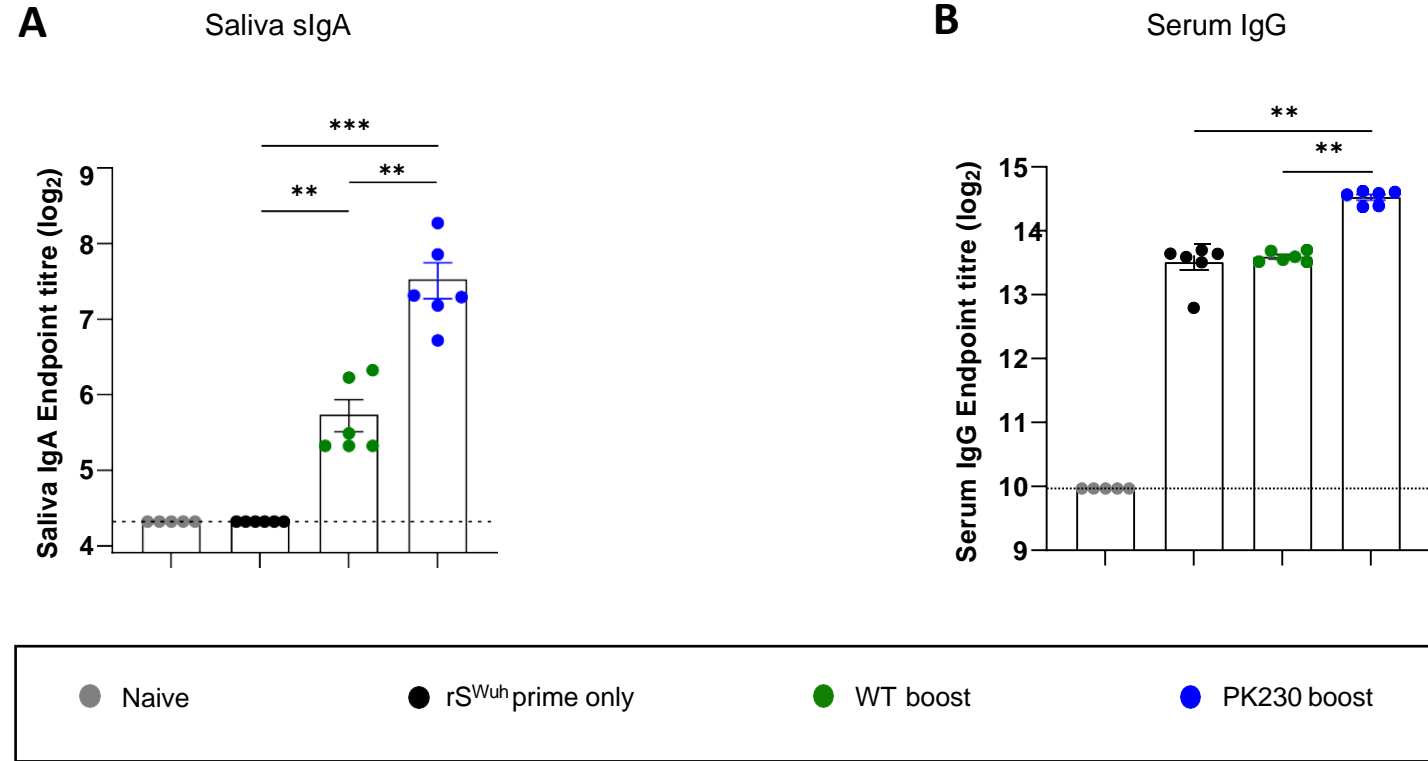
Supplementary Figure S1: Recombinant Spore Vaccines

Chimeric genes inserted at the *thyA* loci of *B. subtilis* are shown. The Wuhan and Omicron-derived Spike RBD domains were fused in frame to the C-termini of the CotB spore coat protein and the HR1-HR2 domain (Wuhan) of Spike fused in frame to the C-terminus of the CotC spore coat protein. The 15 amino acid changes present in the Omicron RBD variant are highlighted. The Mw of the chimera are shown.



Supplementary Figure S2: Animal Study Schedules

For murine immunisations intra-nasal boosts were given three and five weeks after the prime. Each boost consisted of three daily administrations of 10 μ L/mouse, 5 μ L/nare). Boost 1, days 21, 22, 23 and Boost 2, days 35, 36, 37). For hamster boosts animals were given a single intranasal dose of 0.1 mL (50 μ L/nare) three and five weeks post-prime.



Supplementary Figure S3: Intranasal Boost of the AZD1222 Vaccine with Spores Expressing Omicron-specific Antigens

Three groups (n=6) of mice (female Balb/c) were primed (i.m.) with the AZD1222 vaccine. Two groups were then boosted (intranasal) with spores (5×10^9 CFU/dose) spores of either WT (naked spores, no antigen expression) or PK230 (CotB-RBD^{Omi}) at 21 and 35-days post-prime. Panels show rS^{Omi}-specific responses determined by ELISA (OD_{450nm}) 48-days post-immunization. **Panel A)** rS^{Omi}-specific sIgA in longitudinal saliva samples, **B)** rS^{Omi}-specific IgG in serum. Significance was tested using a two-tailed Mann-Whitney U test, ** $p < 0.01$, *** $p < 0.001$