SARS-CoV-2 Spike Ferritin Nanoparticle Vaccine

Kayvon Modjarrad, MD, PhD
Director, Emerging Infectious Diseases Program
Walter Reed Army Institute of Research

World Health Organization Pan-Sarbecovirus Vaccine Consultation

25 March 2022



Preclinical Summary

Cell Reports

SARS-CoV-2 ferritin nanoparticle vaccines elicit broad SARS coronavirus immunogenicity

npj | vaccines

A SARS-CoV-2 spike ferritin nanoparticle vaccine protects hamsters against Alpha and Beta virus variant challenge

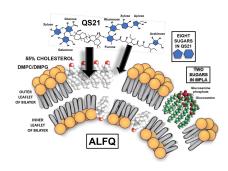


Science Translational Medicine

RESEARCH ARTICLES

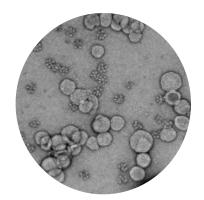
CORONAVIRUS

A SARS-CoV-2 ferritin nanoparticle vaccine elicits protective immune responses in nonhuman primates



npj | vaccines

SARS-CoV-2 ferritin nanoparticle vaccine induces robust innate immune activity driving polyfunctional spike-specific T cell responses



PNAS

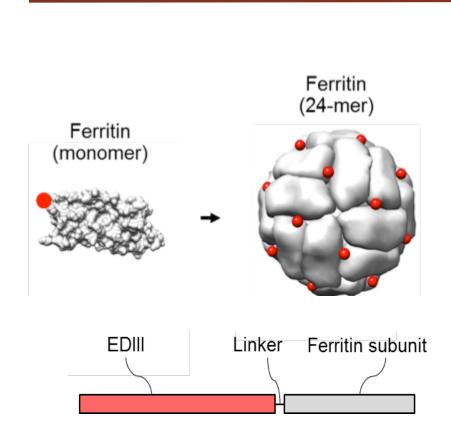
Proceedings of the National Academy of Sciences of the United States of America

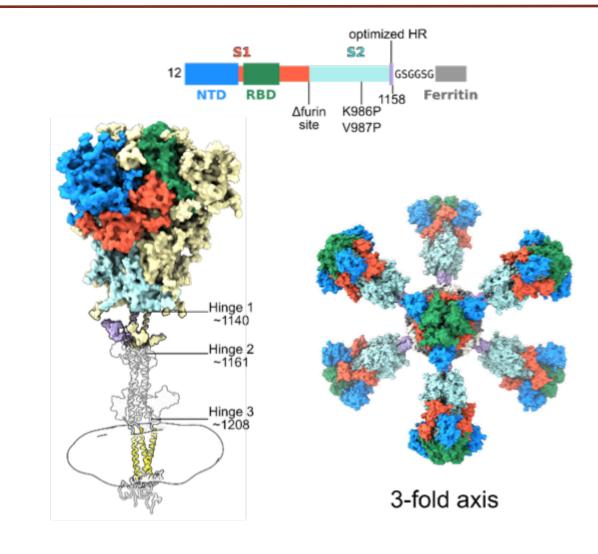
Efficacy and breadth of adjuvanted SARS-CoV-2 receptor-binding domain nanoparticle vaccine in macaques

Updated 25 February 2022



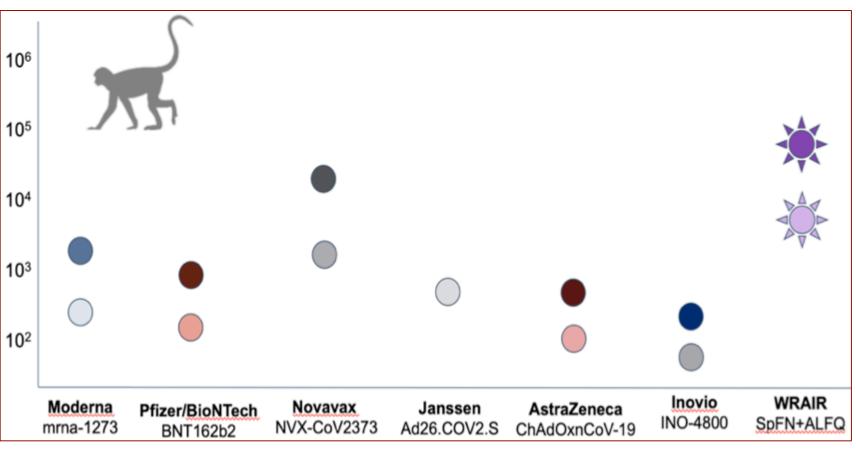
Design of SARS-CoV-2 Spike Ferritin Nanoparticle (SpFN)





A Suite of Animal Studies: Consistent Potency & Protection





Next Set of Animal Studies

- Multivalent formulations mouse/hamster studies
- Heterologous prime-boost NHP Omicron challenge study
- mRNA formulation mouse studies
- Passive transfer human IgG hamster SARS1/2 challenge
- Adjuvant and Immunogen combination comparison NHP

Ferritin Influenza HA Clinical Trial



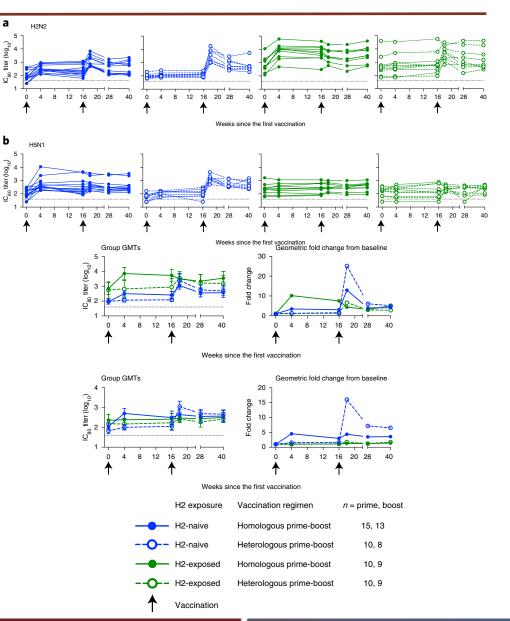
ARTICLES

https://doi.org/10.1038/s41591-021-01660-8

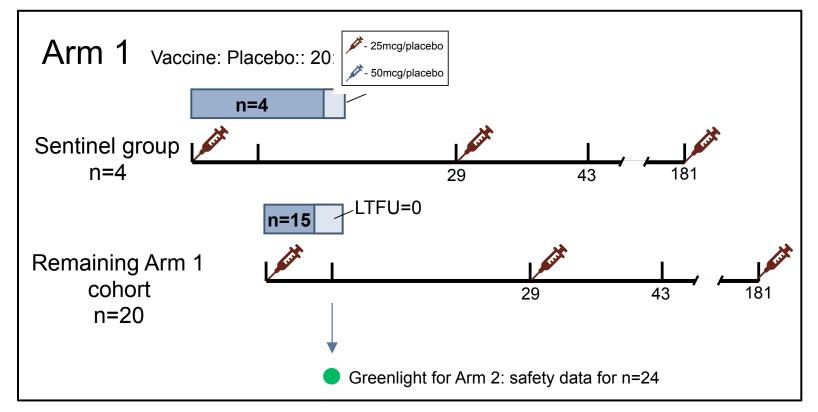


Safety and immunogenicity of a ferritin nanoparticle H2 influenza vaccine in healthy adults: a phase 1 trial

Katherine V. Houser ^{1,7} [∞], Grace L. Chen^{1,7}, Cristina Carter¹, Michelle C. Crank¹, Thuy A. Nguyen¹, Maria Claudia Burgos Florez¹, Nina M. Berkowitz¹, Floreliz Mendoza¹, Cynthia Starr Hendel¹, Ingelise J. Gordon¹, Emily E. Coates¹, Sandra Vazquez¹, Judy Stein¹, Christopher L. Case², Heather Lawlor¹, Kevin Carlton¹, Martin R. Gaudinski¹,³, Larisa Strom¹, Amelia R. Hofstetter ¹, C. Jason Liang⁴, Sandeep Narpala¹, Christian Hatcher¹, Rebecca A. Gillespie ¹, Adrian Creanga ¹, Masaru Kanekiyo ¹, Julie E. Raab¹, Sarah F. Andrews ¹, Yi Zhang¹, Eun Sung Yang ¹, Lingshu Wang¹, Kwanyee Leung¹, Wing-Pui Kong¹, Alec W. Freyn⁵, Raffael Nachbagauer ¹, Peter Palese ^{5,6}, Robert T. Bailer¹, Adrian B. McDermott ¹, Richard A. Koup¹, Jason G. Gall¹, Frank Arnold¹, John R. Mascola¹, Barney S. Graham¹, Julie E. Ledgerwood¹ and the VRC 316 Study Team*

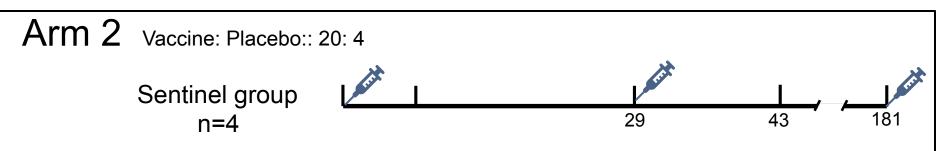


SpFN Phase I Trial Design

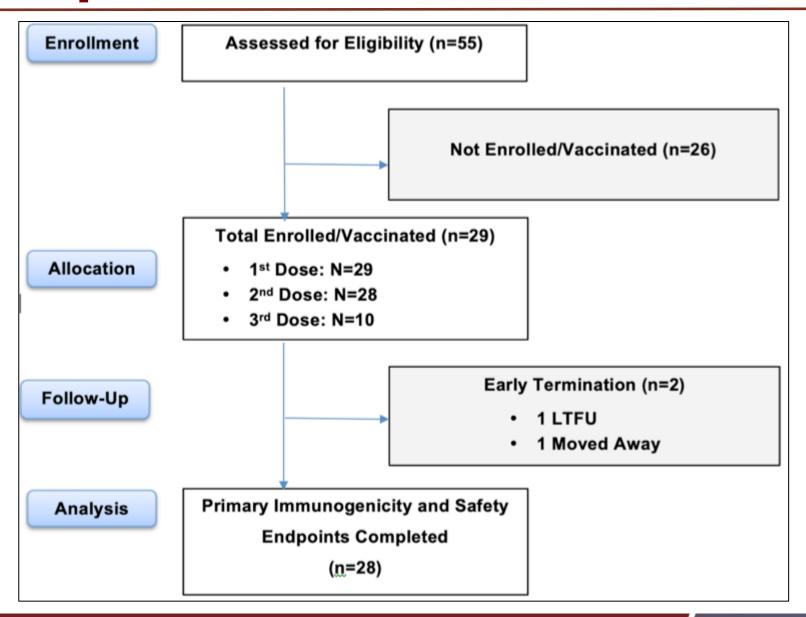








SpFN Phase I Trial Flow



SpFN Phase I Trial Endpoints

- Local and Systemic reactogenicity
- Binding antibody (MSD)
- SARS-CoV-2 VOC Pseudovirus Neutralization
- Cell mediated immunity (ICS)
- Sarbecovirus RBD binding antibody (BLI (Octet)) panel
- Broader sarbecovirus neutralization panel
- Purified IgG passive transfer/challenge (Delta, Omicron, SARS1)

Acknowledgements

Emerging Infectious Diseases Branch (EIDB)

Gordon Joyce Paul Scott **Brittany Shepherd** Mihret Amare Sandhya Vasan Erifile Zografos Heather Liu Jarrett Headley Greg Bujak Jenny Lay Kara Lombardi Leigh Anne Eller

MHRP

Gary Matyas Mangala Rao Vicky Polonis Sebastian Molnar **PBF**

LTCJeffrey Froude Rafael de la Barrera Stasya Zarling-Bejma MAJ Nathan Jansen

Amy Castellano Daniel Choi Elizabeth Bianchi Erica Sondergaard Mark Milazzo Mekdi Taddese Melanie McCauley Paul Zaremba Farooq Nasir Shilpa Hakre Tsedal Mebrahtu Theresa Lins

VDB

Greg Gromowski Aslaa Ahmed

CTC

Melissa Greenleaf Evie Guzman MAJ Aaron Sanborn MAJ Jack Hutter MAJ Mike Koren

Misook Choe Rajeshwer Sankhala Wei-Hung Chen Paul V. Thomas Agi Hajduczki Caroline Peterson William Chang Paul Thomas Elizabeth Martinez Phyllis Rees Petra Prins Leilani Francisco

DHVI

David Montefiori Hongmei Gao Amanda Eaton Kelli Greene

WRAIR / CIDR

Nelson Michael COL Robert O'Connell COL Chad Koenig **BG Clinton Murray** COL(P) Deydre Teyhen

NIAID/VRC

John Mascola Barney Graham Rick Koup Julie Ledgerwood Adrian McDermott Kizzmekia Corbett Masaru Kanekiyo Sarah O'Connell Sandeep Narpala Sandra Vazquez Somia Hickman

FHCC

Julie McElrath Stephen De Rosa Jason Gall **Kevin Carlton** Paula Lei Lisa Kueltzo Dan Gowetski Janel Holland Linn Jacob Demirji Kristin Leach Michael Pratt Florence Kaltovich Jessica Bahorich William Shadrick Yanhong Yang

















10