

## CHALLENGES FACED IN THE PLANNING, DEPLOYMENT, AND ADOPTION OF A LAYERED MEDICAL DEFENSE STRATEGY

## Characterization Of A Cynomolgus Macaque Model Of Q Fever

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A robust animal model that recapitulates human clinical disease is essential for screening novel medical countermeasures. Although Q fever, caused by the bacterium Coxiella burnetii, is rarely lethal, it can lead to a debilitating acute and chronic infection that is naturally recalcitrant to antibiotics. The only vaccine available is currently approved for use in Australia and requires a skin test prior to administration to avoid a potential hypersensitive response in previously sensitized individuals. This requirement precludes it from being utilized on a wider scale or for rapid deployment. In order to test next-generation vaccines, this portfolio of work has generated characterized challenge stock, determined the median aerosol lethal dose, and defined Q fever disease progression in an aerosol nonhuman primate (NHP) model. To date, the NHP clinical model demonstrates respiratory illness as measured through increased respiratory rate, clinical signs, and radiographs, coupled with sustained fever over approximately seven days. Although overt clinical signs typically resolve by day 28 post-exposure, histopathology on major organs show continual damage is present in the lungs and heart. The developed aerosol NHP model shares many key parameters that mimic C. burnetii infection in humans, demonstrating its future utility in next-generation medical countermeasure efficacy studies.

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