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Project Number

1R01AI158177-01

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Novel delivery platform and antigen design for an effective COVID-19 vaccine

Contact PI/Project Leader MITTAL, SURESH K

Awardee Organization PURDUE UNIVERSITY

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Abstract Text

PROJECT SUMMARY For effective management of the COVID-19 pandemic and its second wave, the design and implementation of multiple intervention approaches are crucial. They include the development of effective antivirals, high-affinity SARS-CoV-2-neuralizing human or humanized monoclonal antibodies, rapid diagnostic assays, immunogenic and protective vaccines, strategies to mitigate virus transmissibility, and enhancing capacity related to trained medical personnel, facilities, and supplies. Due to the possibility of antibody-dependent enhancement (ADE) of COVID-19, vaccine efforts should consider the use of a novel **vaccine** platform and design of a relevant antigen strategy. It is essential to note that the elderly are the most vulnerable segment of the population that is at a higher risk of COVID-19 severity; the vaccine development efforts should, therefore, consider the decline in the immune competence in the elderly. We have developed a novel replication-defective (E1 & E3 deleted) bovine adenovirus (Ad) type 3 (BAd3)- based vaccine platform, which is better than the currently available Ad vector systems for providing heterologous influenza protection with dose sparing and is not impacted by the pre-existing human Ad vector immunity. Recently, we have revealed that the BAd vaccine platform provides the expression of significantly higher levels of the immunogen and innate and adaptive immunity-related factors compared to that of human Ad vectors in mice. This work suggests that the BAd vector system could serve as an excellent delivery vehicle for the development of recombinant vaccines against emerging pathogens for the elderly and other segments of the population. We have also identified a 22 amino acid residues Autophagy-Inducing Peptide (AIP) C5 (AIP-C5) from the CFP10 protein of M. tuberculosis that enhances robust T cell immune responses in mice to NP of H7N9 influenza virus when delivered through an Ad vector. It conferred complete protection against H1N1, H3N2, H5N2, H7N9, and H9N2 influenza viruses. The proposal is based on the hypothesis that immunization with the autophagy-inducing replicationdeficient BAd vector expressing relevant antigen/s of SARS-CoV-2 will strengthen an effective mucosal (lung) and systemic anti-COVID-19 immunity. Under Aim 1, we will evaluate the immunogenicity and protective efficacy of a novel vaccine platform and antigen design in animal models for developing an effective COVID-19 vaccine. Whereas under Aim 2, we will investigate the vaccine-induced antibody-dependent enhancement (ADE) of SARS-CoV-2 infection, the quality of memory innate, B and T cell responses, and the durability of protective immunity in the best animal model. We believe that the use of a unique nonhuman Ad vaccine platform and novel antigen design containing AIP-C5 will yield an effective COVID-19 vaccine for all segments of the population. This effort will be of significant value to effectively flatten the COVID-19 pandemic's trajectory and its second wave.

Public Health Relevance Statement

PROJECT NARRATIVE For effective management of the COVID-19 pandemic and its second wave, the design and implementation of multiple intervention approaches are crucial. We believe that the use of a unique nonhuman adenoviral vaccine platform and novel antigen design will yield an effective COVID-19 vaccine for all segments of the population. This effort will be of significant value to effectively flatten the COVID-19 pandemic's trajectory and its second wave.

NIH Spending Category

Aging Biotechnology Coronaviruses Emerging Infectious Diseases

Immunization Infectious Diseases Influenza Lung Pneumonia

Pneumonia & Influenza Prevention Vaccine Related

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COVID-19 pandemic COVID-19 **COVID-19 vaccine** Cattle **Cessation of life** China Chiroptera **Coronavirus Infections** Country **Development Dose Feline Coronavirus Fever Elderly Equilibrium Failure Ferrets**

Glycoproteins Health Personnel Human Immune response Immune system

Infection Influenza **Immunity Immunization Immunocompetence**

Read More

Details

Contact PI/ Project Other Pls Program Official

Leader Not Applicable Name

Name

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Organization Type Congressional District City

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Country **UNITED STATES (US)**

Other Information

FOA Administering Institutes or **Project Start** PAR-20-178 Centers Date

NATIONAL INSTITUTE OF Study Section **ALLERGY AND INFECTIOUS** Special Emphasis Panel ZAI1 **DISEASES**

<u>JHM-X (S3)</u>] **DUNS Number CFDA Code**

Award Notice Budget Start 072051394 855 Date Date Fiscal Year 11-August-**Budget End**

2020 Date 2021

Project Funding Information for 2020

Funding IC

Total Funding Direct Costs Indirect Costs \$792,476 \$660,247 \$132,229

2020 NATIONAL INSTITUTE OF ALLERGY AND INFECTIOUS DISEASES \$792,476

Year

2020

11-August-

2020

2025

2020

31-July-

31-July-

11-August-

Project End

Date

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Diseases; Immunization; Infectious Diseases; Influenza; Lung; Pneumonia; Pneumonia & Influenza; Prevention; Vaccine Related;

品 Sub Projects

No Sub Projects information available for 1R01Al158177-01

Publications

No Publications available for 1R01Al158177-01

∀ Patents

No Patents information available for 1R01AI158177-01

Outcomes

The Project Outcomes shown here are displayed verbatim as submitted by the Principal Investigator (PI) for this award. Any opinions, findings, and conclusions or recommendations expressed are those of the PI and do not necessarily reflect the views of the National Institutes of Health. NIH has not endorsed the content below.

No Outcomes available for 1R01AI158177-01

Clinical Studies

No Clinical Studies information available for 1R01Al158177-01

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Related News Releases

No news release information available for 1R01AI158177-01

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Contact PI/Project Leader Awardee Organization
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