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Novel Influenza nano vaccines for broad cross protection

Project Number Contact PI/Project Leader 5R01AI101047-08 WANG, BAOZHONG

Awardee Organization
GEORGIA STATE
UNIVERSITY

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Abstract Text

SUMMARY Influenza is a major public health risk. The current seasonal influenza vaccine is effective against closely matched viruses in healthy adults, but it cannot prevent the outbreaks of epidemics or pandemics because influenza viruses mutate frequently and zoonotic strains can jump the species barrier into humans. Other disadvantages of the seasonal influenza vaccine include the need to produce new vaccines every season, the uncertainty in selecting vaccine strains, and the compromised efficacy for mismatched viruses. A novel generation of influenza vaccines, termed universal influenza vaccines, will overcome these challenges. In the previous grant period, we have produced layered protein nanoparticles (nanoclusters) from conserved HA stalk antigens and the M2 protein ectodomain of influenza A. Nanocluster immunizations induced cross protection against viruses from both phylogenic groups of influenza A, including pandemic-potential avian strains. Both influenza A and influenza B cause influenza epidemics in humans. In this proposal, we propose to construct a multivalent layered nanocluster formulation composed of newly designed antigenic proteins from both influenza A and influenza B as a universal influenza vaccine. The new vaccine will induce broad cross- protection against both influenza types. We have three specific aims: Aim 1. To design and construct conserved antigens from influenza A and B, fabricate nanoclusters from these and previously designed antigenic proteins, and characterize these new nanoclusters. We will optimize the orchestration, composition, and stability of these nanoclusters for the physiologicallyactivated release of free antigenic proteins, antigen-processing and presentation after the uptake by dendritic cells, distribution of these nanoclusters to draining lymph nodes, and induction of strong antigen-specific immune responses in mice. Aim 2. To test whether these layered nanoclusters or an optimal combination will induce broadly reactive immune responses and whether the immunity will grant cross-protection against viruses spanning both influenza A and influenza B in mice. Aim 3. To test whether the leading multivalent nanocluster combinations will induce robust immune responses which confer broad cross-protection in ferrets. Overall, our research will develop a broadly crossprotective universal influenza vaccine.

Public Health Relevance Statement

Narrative: Influenza is the leading cause of death by infection. We will develop a protein nanoparticle universal influenza vaccine composed of newly designed influenza antigens from both influenza A and influenza B to induce broadly reactive immunity. The implementation and success of the project will improve public health by granting broad cross-protection against both influenza epidemics and pandemics.

Project Terms

Adaptive Immune System **Antibody titer measurement** Adult **Animal Model Antigen Presentation Antigen Presentation Pathway Antigen-Presenting Cells Antigens Birds Dendritic Cells** Cause of Death **Chimeric Proteins** Disadvantaged **Ethanol Disease Outbreaks Drug Delivery Systems Epidemic** Generations **Ferrets Filtration Formulation** Head Genes Grant Immune response Hemagglutinin Human **Immune Immunity Immunization** Inflammation Influenza **Immunize** Infection Influenza B Virus Influenza A virus **Innate Immune Response** Intramuscular

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2021

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Awardee Organization **GEORGIA STATE UNIVERSITY**

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Organization

Name Department Type State Code **MISCELLANEOUS GEORGIA STATE** GA UNIVERSITY

Organization Type ORGANIZED RESEARCH

UNITS

Budget End

Date

Congressional District

15-May-

30-June-

01-July-

30-June-

2012

2024

2021

2022

05

Other Information

UNITED STATES (US)

FOA Administering Institutes or **Project Start** PA-18-859 Centers Date NATIONAL INSTITUTE OF Study Section **ALLERGY AND INFECTIOUS Project End Vaccines Against Microbial DISEASES** <u>Diseases Study</u> Date Section[VMD] **DUNS Number CFDA Code Budget Start** 837322494 855 **Award Notice** Date Fiscal Year Date

Project Funding Information for 2021

21-June-2021

Total Funding Indirect Costs Direct Costs \$181,660 \$630,929 \$449,269

Year **Funding IC**

\$630,929 2021 NATIONAL INSTITUTE OF ALLERGY AND INFECTIOUS DISEASES

品 Sub Projects

No Sub Projects information available for 5R01Al101047-08

Publications

No Publications available for 5R01Al101047-08



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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 5R01Al101047-08

Clinical Studies

No Clinical Studies information available for 5R01Al101047-08

News and More

Related News Releases

No news release information available for 5R01Al101047-08

History

No Historical information available for 5R01Al101047-08

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