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Role of type 1 fimbrial mutations in the pathogenesis of pandemic E. coli

Project Number	Contact PI/Project Leader	Awardee Organization
1R21AI147575-01	SOKURENKO, EVGENI VENIAMINOVIC	UNIVERSITY OF WASHINGTON
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Description

Abstract Text

ABSTRACT Adhesion is an essential trait for the pathogenesis of urinary tract infections and intestinal colonization by E. coli and mannose-specific type 1 fimbriae (T1F) is the most critical adhesive factor. Our goal is to understand how some of the unique features in T1F of the **pandemic** uropathogenic clonal group, ST131-H30, are adaptive for this sub-clone's urovirulence and intestinal fitness. We will concentrate on the structure-functional analysis of the mannose-binding adhesive subunit of the fimbriae, FimH, and the regulatory mechanism of fimbrial expression that are specific to ST131-H30. We will characterize in great detail the functional properties of the fimbrial adhesin, FimH, by comparing its ability to mediate adhesion under various flow and inhibitory conditions. We also will determine the functional significance of genetic insertion that inactivated the gene coding for positive regulator of T1F, FimB. We will attempt to understand in what way this mutation affected expression of T1F itself as well as other genes across the genome and the E. coli's virulence and colonization properties. Finally, we will use animal studies to determine the adaptive significance of the T1F and its variants in ST131-H30.

Public Health Relevance Statement

The bacterial attachment traits are called type 1 fimbriae and are critical for the pathogenesis of E. coli virulence in the urinary tract infections. This project will study how is one of the most successful uropathogenic E. coli differ from others in the fine attachment and regulation mode. This will be investigated by using genetically manipulated strains and animal models.

NIH Spending Category

- Digestive Diseases
- Genetics
- Infectious Diseases
- Urologic Diseases
- Women's Health

Project Terms










- Adhesions
- Adhesives
- Affect
- Animal Model
- Animals
- Antibiotic Resistance
- Bacterial Adhesins
- Bacterial Attachment Site
- Binding
- Blood Circulation
- Code
- Data
- Escherichia coli
- Fimbrial Adhesins
- Gene Expression
- Genes
- Genetic
- Genetic Enhancement
- Genetic Transcription
- Genome
- Goals
- Horizontal Gene Transfer
- Human
- IS Elements
- Individual
- Intestines
- Invaded
- Mannose
- Mediating
- Microbial Biofilms
- Modeling
- Morphology
- Mus
- Mutate
- Mutation
- Nature
- Pathogenesis
- Phenotype
- Point Mutation
- Population
- Probiotics
- Property
- Regulation
- Role
- Seminal
- Structure
- Urinary tract

Details

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Project Number

1R21AI147575-01

Contact PI/Project Leader

SOKURENKO, EVGENI
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Awardee Organization

UNIVERSITY OF
WASHINGTON

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Organization

Name

UNIVERSITY OF
WASHINGTON

Department Type

MICROBIOLOGY/IMMUN/VIROLOGICAL
SCHOOLS OF MEDICINE

State Code

WA

City

SEATTLE

Congressional District

07

Country

UNITED STATES (US)

Other Information

FOA
[PA-18-489](#)

Administering Institutes or
Centers
NATIONAL INSTITUTE OF
ALLERGY AND INFECTIOUS
DISEASES

Project Start
Date
15-July-
2019

Study Section
[Special Emphasis
Panel\[ZRG1 IDM-V \(02\)\]](#)

DUNS Number CFDA Code
605799469 855

Project End
Date
30-June-
2021

Award Notice
Date

Fiscal Year
2019

Budget Start
Date

Budget End
Date

10-July-2019

15-July-
2019

30-June-
2020

Project Funding Information for 2019

Total Funding	Direct Costs	Indirect Costs
\$233,250	\$150,000	\$83,250

Year	Funding IC	
2019	NATIONAL INSTITUTE OF ALLERGY AND INFECTIOUS DISEASES	\$233,250

NIH Categorical Spending [Click here for more information on NIH Categorical Spending](#)










Funding IC	FY Total Cost by IC	NIH Spending Category
NATIONAL INSTITUTE OF ALLERGY AND INFECTIOUS DISEASES	\$174,938	Women's Health
NATIONAL INSTITUTE OF ALLERGY AND INFECTIOUS DISEASES	\$233,250	Digestive Diseases; Genetics; Infectious Diseases; Urologic Diseases;

 Sub Projects

No Sub Projects information available for 1R21AI147575-01

Thank you for your feedback!

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1R21AI147575-01	SOKURENKO, EVGENI VENIAMINOVIC	UNIVERSITY OF WASHINGTON



Patents

No Patents information available for 1R21AI147575-01



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 1R21AI147575-01



Clinical Studies

No Clinical Studies information available for 1R21AI147575-01



News and More

Related News Releases

No news release information available for 1R21AI147575-01



History

No Historical information available for 1R21AI147575-01



Similar Projects

No Similar Projects information available for 1R21AI147575-01

Thank you for your feedback!