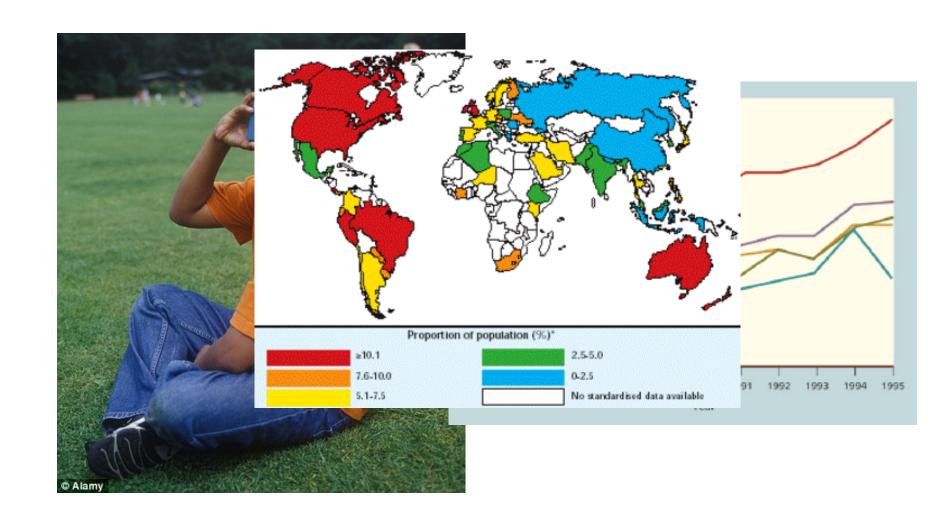
# Household Dust Exposure Modifies Gut Microbiome Composition and Airway Health

### Susan V. Lynch, PhD

Associate Professor of Medicine
Director Colitis and Crohn's Disease Research Core
Division of Gastroenterology,
Department of Medicine,
University of California San Francisco.

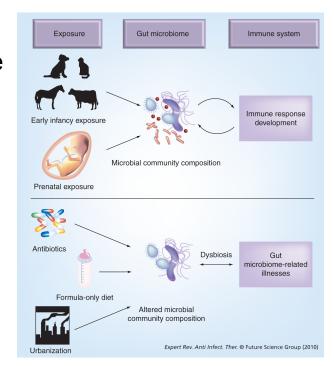


### **Pediatric Asthma**



### Childhood Allergic Disease and Asthma

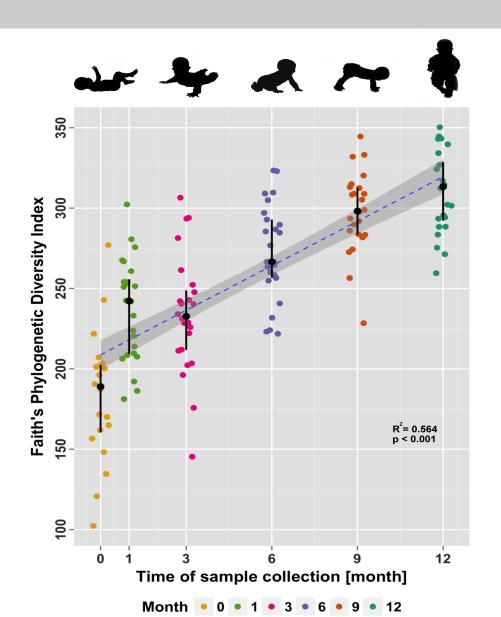
- Factors associated with childhood allergic disease development:
  - Early life antibiotic exposure
  - Formula feeding
  - Caesarian section delivery
  - Lack of maternal exposure to animals during pregnancy
  - Maternal antibiotic use
  - Lack of early life furred pet exposure



Fujimura et al, ERAI 2010

 Early-life gastrointestinal microbial outgrowth is associated with childhood allergic disease development

## **Early Life Microbiome Development**



# Is the Indoor Microbial Environment Related to Allergic Disease?



86.9% (68.7% in residence)

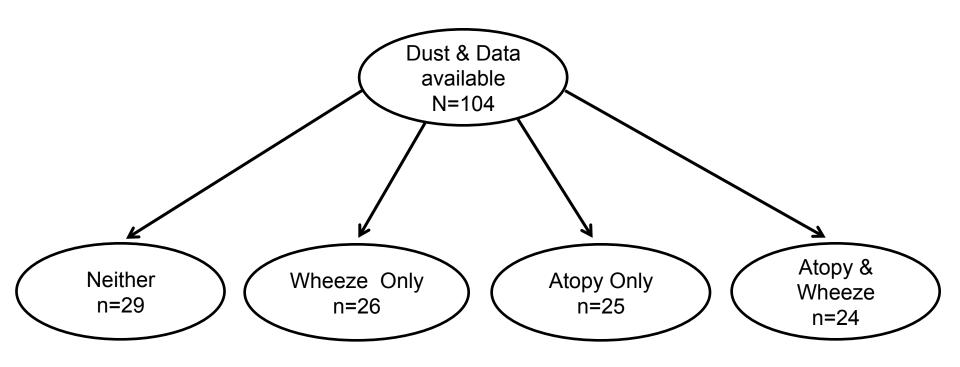






5.5%

# Is House Dust Microbiome Related to Childhood Allergic Outcomes?

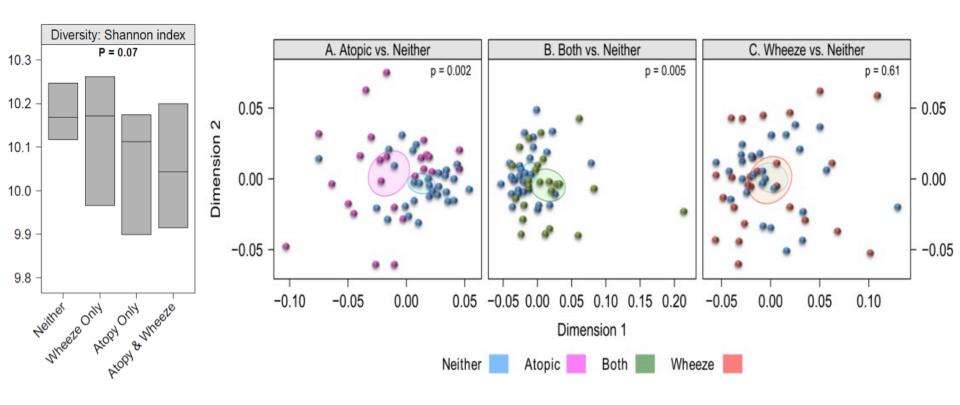


**Atopy defined as:**  $IgE \ge 0.35$  for any aeroallergen was considered positive

IgE < 0.10 for all aeroallergens was considered negative

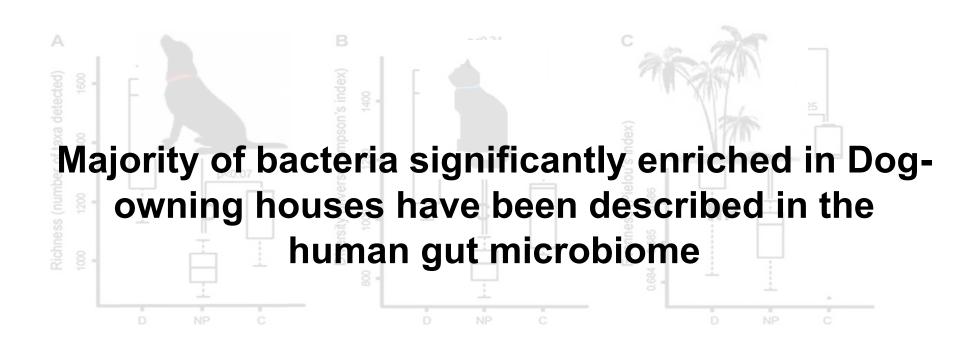
Wheeze defined as: ≥ 2 episodes of wheeze, with at least 1 occurring at age 3

## Dust Microbiome Composition is Related to Clinical Outcomes



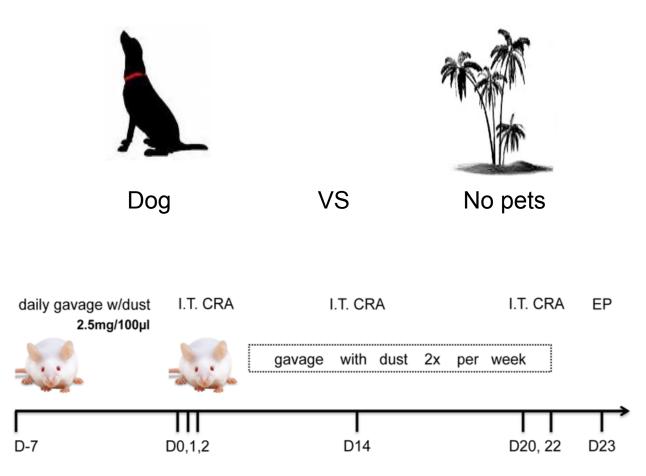
~ 70 taxa in household dust that are associated with protection against disease development – human gut inhabitants

# Does pet ownership influence the microbial exposures in households?



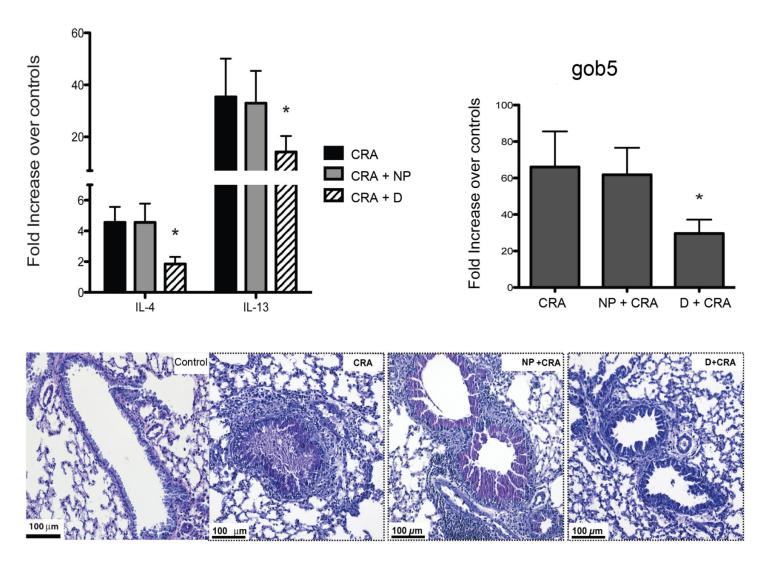
Fujimura KE et al. J Allergy Clin Immunol. 2010 126(2) 410-2

## Can Environmental Microbial Exposure Influence Airway Responses?

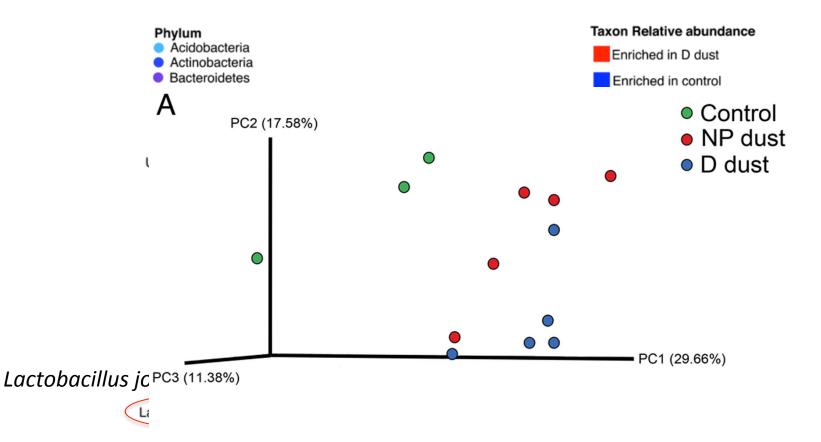


Berlin, A.A., et al, Lab Invest. 2006 Jun;86(6):557-65.

# House Dust Exposure Changes Airway Response to Allergen



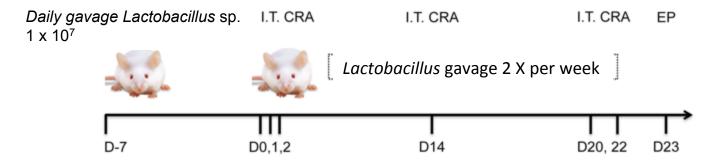
### **Gut Microbiome is Distinct in Protected Mice**



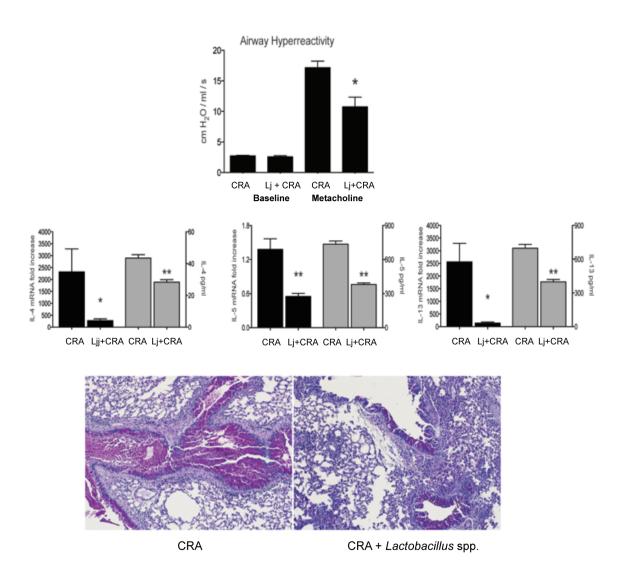
## Lactobacillus Isolation & Supplementation



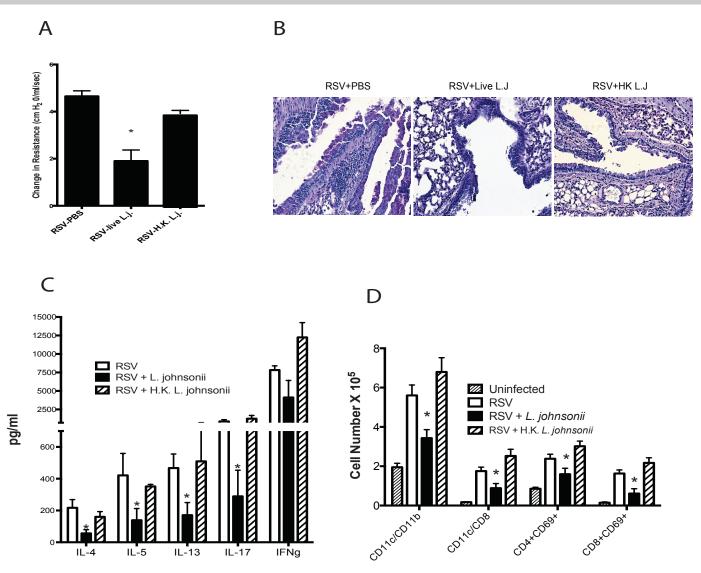




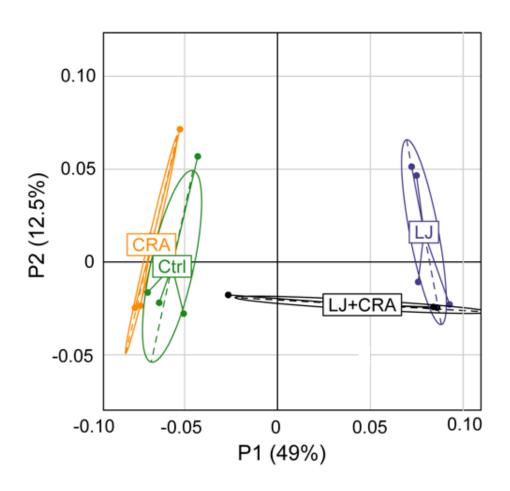
# Lactobacillus Supplementation Protects against Airway Allergen Challenge



# L. johnsonii Supplementation Protects Against Viral Infection



# L. johnsonii Reshapes the Murine Cecal Microbiome



- No enrichment of Lachnospiraceae, Peptococcaceae or Bacillaceae
- Environmentally-sourced species required for full protection

### **Conclusions**

- Local environmental exposures are related to clinical outcomes of childhood allergic disease
- Gut microbiome can be influenced by environmental exposures
- Gut microbiome manipulation leads to altered host immune response at remote mucosal surfaces

## What's next?









### **Acknowledgements**

#### **UCSF**

### **University of Michigan**

Nick Lukacs

Tine Demoor

Kei Fujimura Marcus Rauch Michelle McKean Homer Boushey Michael Cabana

**URECA** study

Jim Gern
Bob Wood
Homer Boushey
Augustin Calatroni
Henry Lynn
Bill Busse

### **Henry Ford Hospital**

Chris Johnson
Suzanne Havstad
Ed Zoratti
Kim Woodcroft
Kevin Bobbitt

Ganessa Wegienka

### Georgia Health Sciences Health System

**Dennis Ownby** 

#### **Lawrence Berkeley National Lab**

**Eoin Brodie** 

#### **Second Genome**

Jannet Warrington Todd DeSantis

#### **Funding**

NIH/NIAID
NIH/NIAID
NIH/NCCAM
UCSF Program for Breakthrough
Biomedical Research
Broad Foundation
Sloan Foundation
Cystic Fibrosis Foundation
Janssen Pharmaceuticals Inc.
Pfizer Inc.