



# **Analysis of dog serologic data reveals critical insights into the epidemiology of human coccidioidomycosis, 2012-2023**

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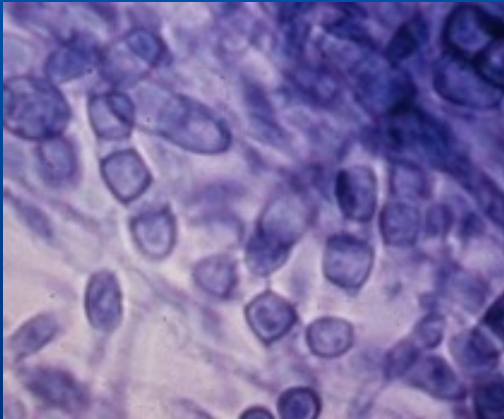
*This presentation contains preliminary data  
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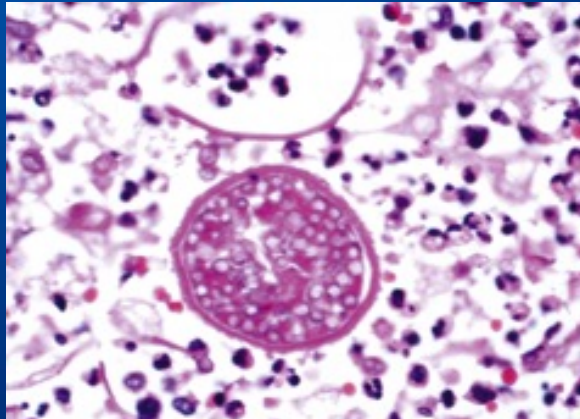
# Background Coccidioidomycosis

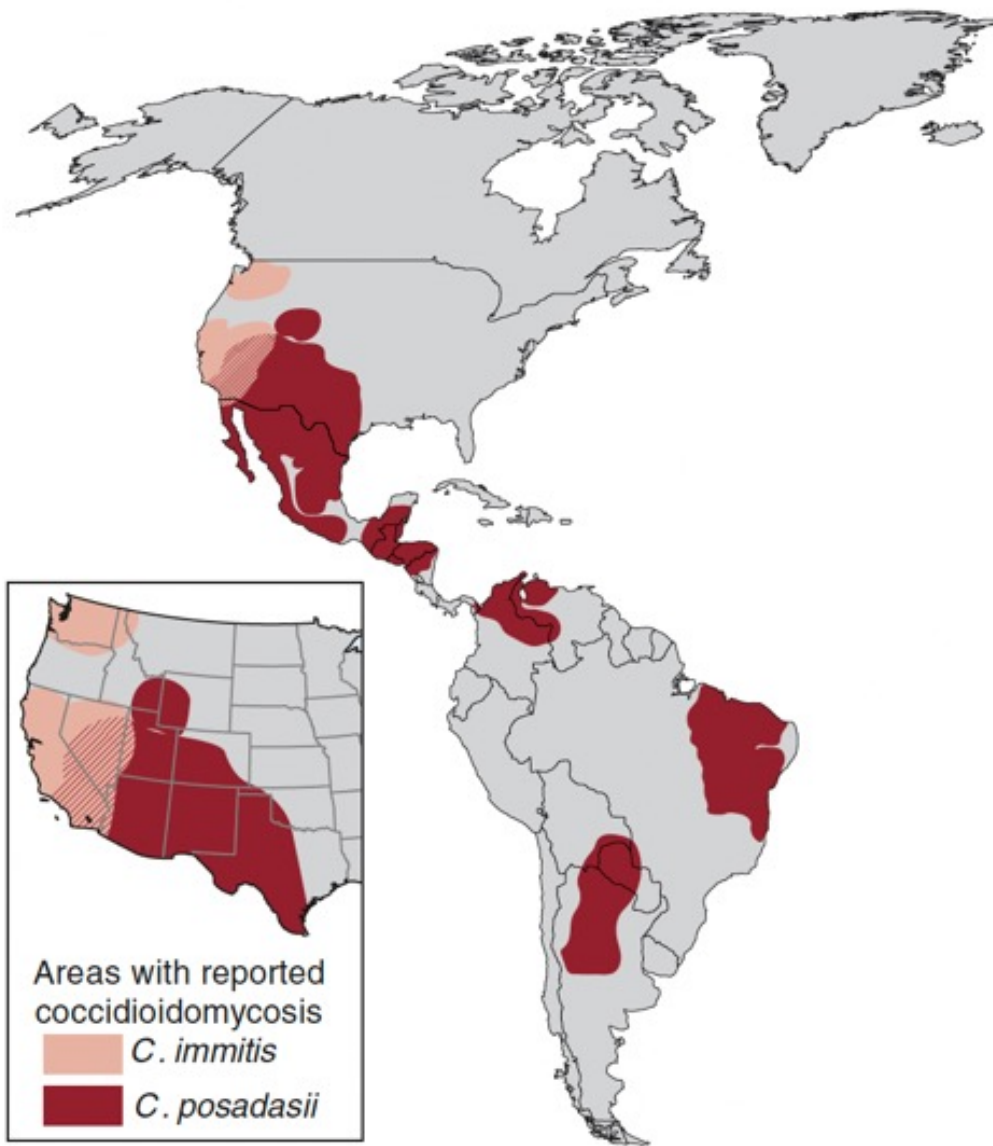
- Systemic fungal disease of humans and animals<sup>1,2</sup>
  - *Coccidioides immitis* and *Coccidioides posadasii*
  - Soil-borne, dimorphic fungi
  - Humans, dogs, horses, camelids, marine mammals
  - Other domestic animals, wildlife

Arthrospores – Infectious



Mature spherule – Non-infectious





## Background Epidemiology

- Mexico, parts Central and South America
- Southwestern United States
  - AZ, TX, CA; also NM, UT, NV
  - San Joaquin Valley (“Valley fever”)
  - Phoenix and Tucson hyperendemic
- Lower Sonoran life zone
  - Hot summers
  - Low elevation
  - Semi-arid to arid soil

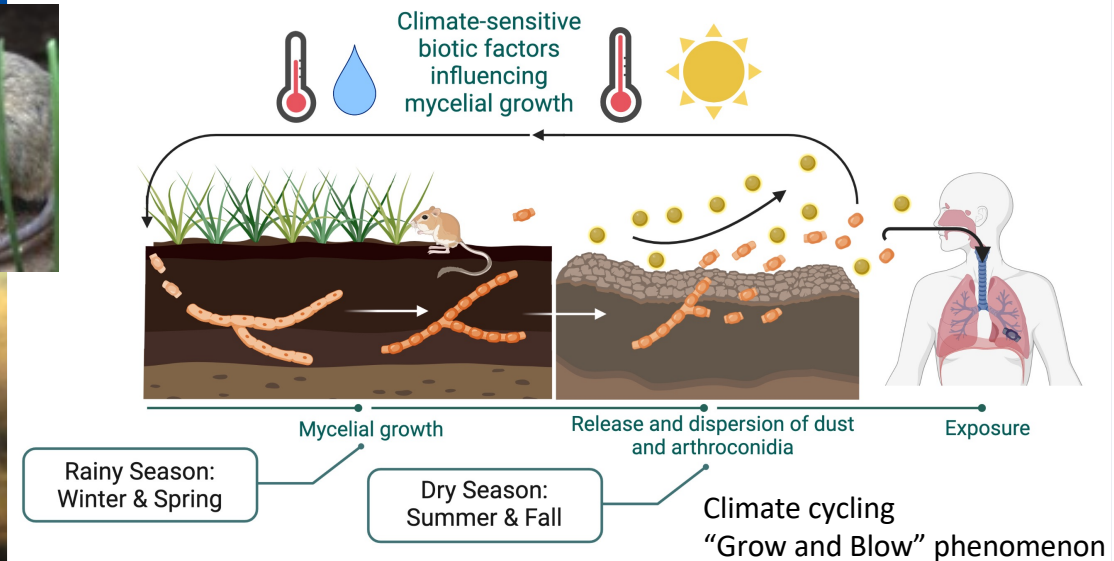


Ref 2

# Background Epidemiology

## The “Dust Devil”

- Dust storms, haboobs, earthquakes, heavy rains, construction, agriculture, other soil disturbance



Credit: A Weaver, UC Berkeley

# Background Epidemiology

- Severe disease/dissemination
  - Adults 60+
  - Males
  - Pregnancy
  - Diabetes
  - HIV/AIDS
  - Organ transplant recipients
  - Pharmacologic immunosuppression
  - African Americans and Filipinos
    - Genetic variations in *CLEC7A* and *PLCG2* genes<sup>4</sup>
    - Encode DECTIN-1 and PLC $\gamma$ 2
    - Fungal  $\beta$ -glucan pattern recognition



**Coccidioidomycosis outbreak among inmate wildland firefighters: California, 2017**

# Background Epidemiology

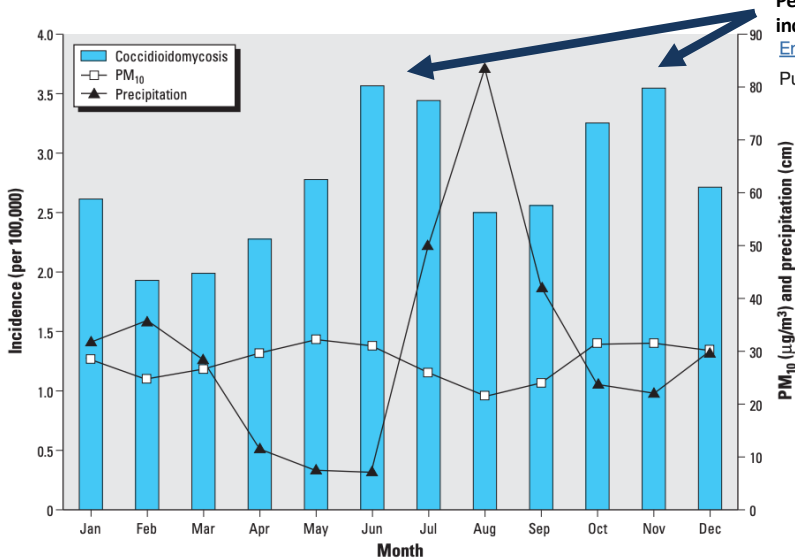
## Research | Article

### Climate Factors Influencing Coccidioidomycosis Seasonality and Outbreaks

Andrew C. Comrie

Department of Geography and Regional Development, University of Arizona, Tucson, Arizona, USA

**Peak Valley fever incidence**  
*Environ Health Perspect.* 2005 Jun; 113(6): 688–692.  
Published online 2005 Mar 3. doi: [10.1289/ehp.7786](https://doi.org/10.1289/ehp.7786)



**Figure 2.** Mean monthly coccidioidomycosis incidence in Pima County, Arizona, based on estimated exposure date, with mean monthly precipitation and mean monthly PM<sub>10</sub> concentrations, 1992–2003.

EcoHealth 15, 840–852, 2018  
<https://doi.org/10.1007/s10393-018-1375-9>



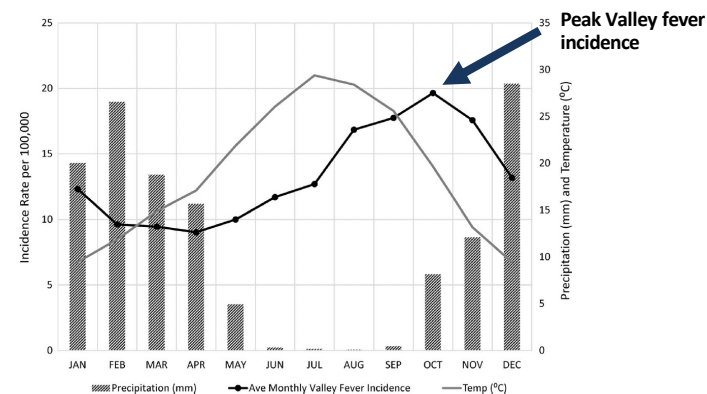
Original Contribution

### Investigating the Relationship Between Climate and Valley Fever (Coccidioidomycosis)

Elizabeth A. Weaver<sup>1</sup> and Korine N. Kolivras<sup>1</sup>

<sup>1</sup>Virginia Polytechnic Institute and State University, 115 Major Williams Hall, Blacksburg, VA 4061

Refs 5, 6



**Figure 3.** Disease and climate seasonality based on 2000–2015 monthly averages with the incidence having a single peak in the month of October, precipitation having a single peak in the month of February and temperature having a single peak in July.

Credit: A Weaver, UC Berkeley

# Background Impact of Climate Change

## Expansion of Coccidioidomycosis Endemic Regions in the United States in Response to Climate Change

Morgan E. Gorris<sup>1</sup> , Kathleen K. Treseder<sup>2</sup> , Charles S. Zender<sup>1</sup> , and James T. Randerson<sup>1</sup> 

## Economic Valuation of Coccidioidomycosis (Valley Fever) Projections in the United States in Response to Climate Change

Morgan E. Gorris<sup>a</sup>, James E. Neumann<sup>b</sup>, Patrick L. Kinney<sup>c</sup>, Megan Sheahan<sup>b</sup>, Marcus C. Sarofim<sup>d</sup>

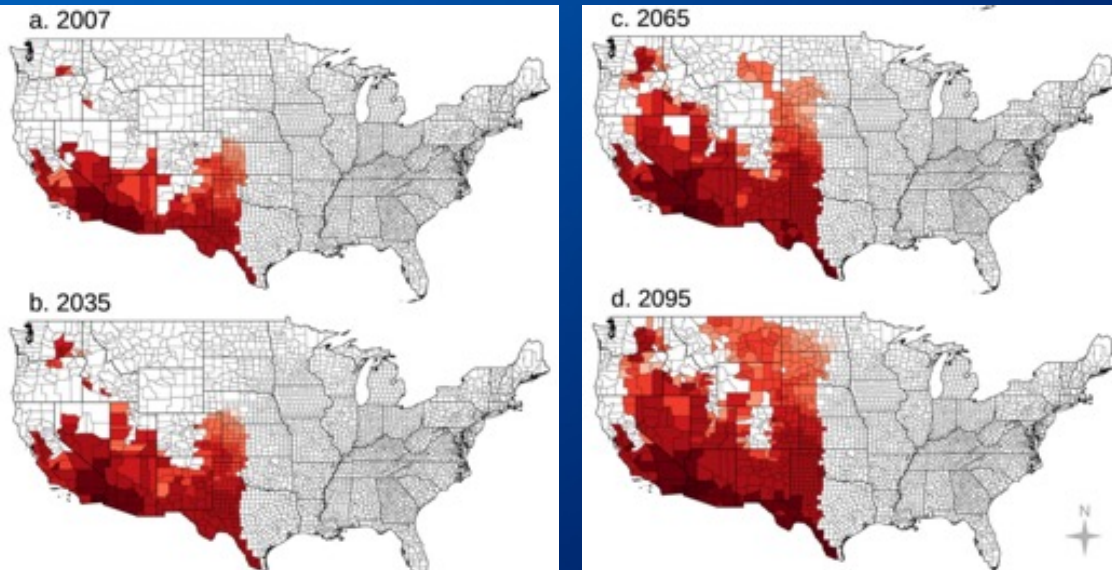
### GeoHealth

#### RESEARCH ARTICLE

10.1029/2019GH000209

##### Key Points:

- We created a niche model to estimate climate limits on the spatial extent of Valley fever endemicity in the United States
- For a high warming scenario, the area of climate-limited endemicity will more than double by 2100, expanding northward into dry western states
- Our predictive model of Valley fever endemic regions may help mitigate disease impacts as the disease spreads into new regions

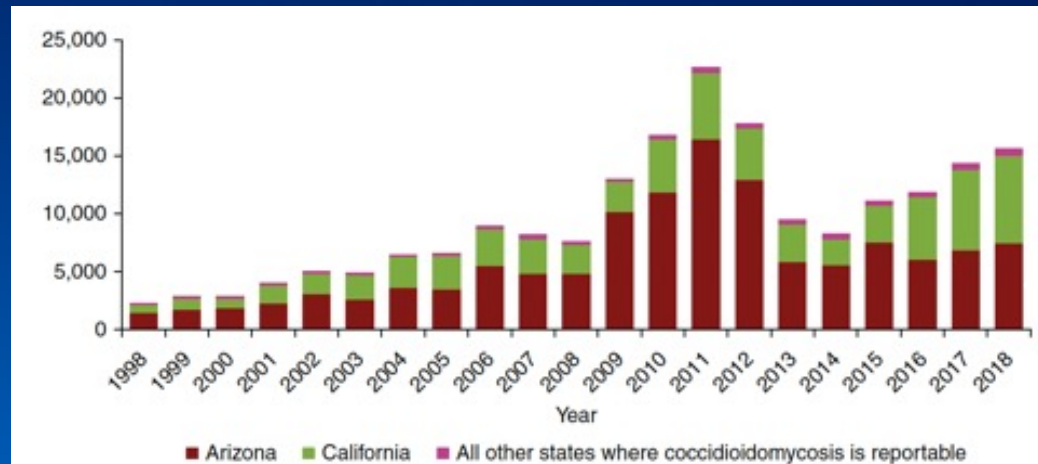


| Time Period                  | Cost       |
|------------------------------|------------|
| 2005-2019                    | \$3.9B/yr  |
| End 21 <sup>st</sup> century | \$18.5B/yr |

Refs 7, 8



# Background Human Surveillance



One of only two nationally notifiable fungal diseases

Reportable to local health jurisdictions in 26 states (including KS) and District of Columbia

Ref 9

## Just One Breath

**Valley fever cases reach epidemic levels, but harm remains hidden**

Reporting on Health Collaborative

Saturday, September 8, 2012

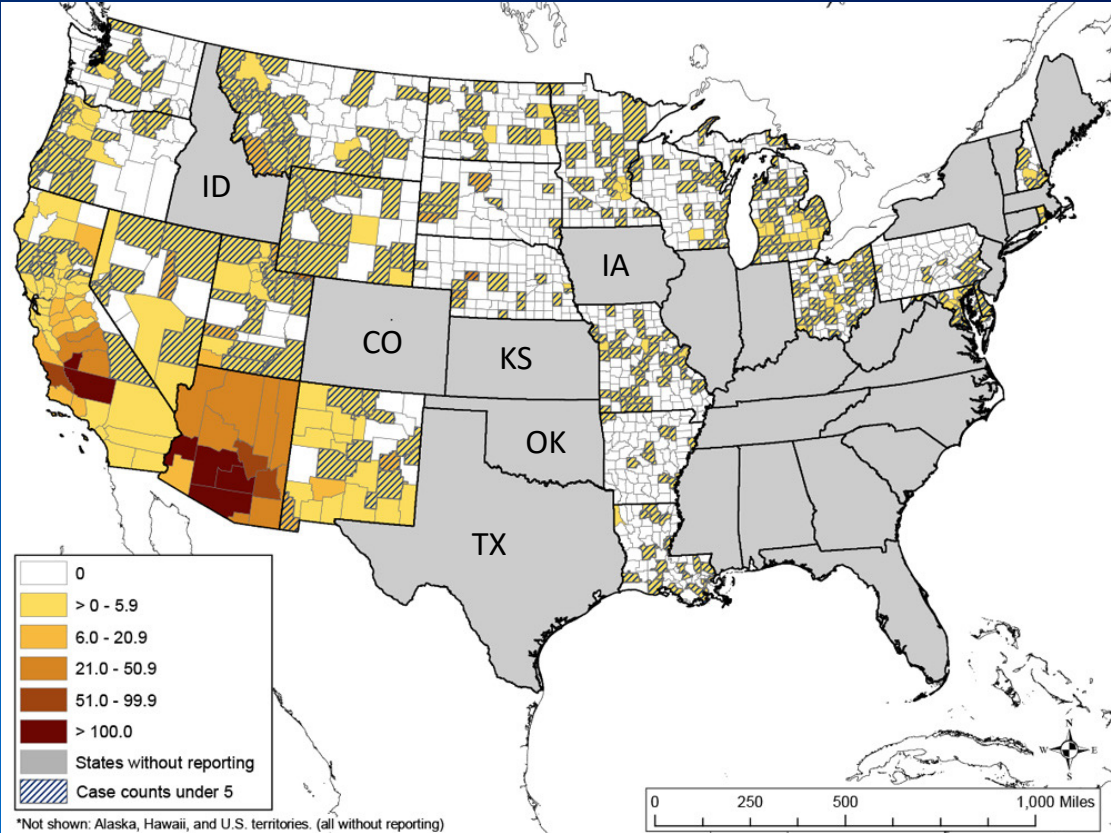
By **Kellie Schmitt, Rebecca Plevin and Tracy Wood**

Valley fever starts with the simple act of breathing.

Ref 10

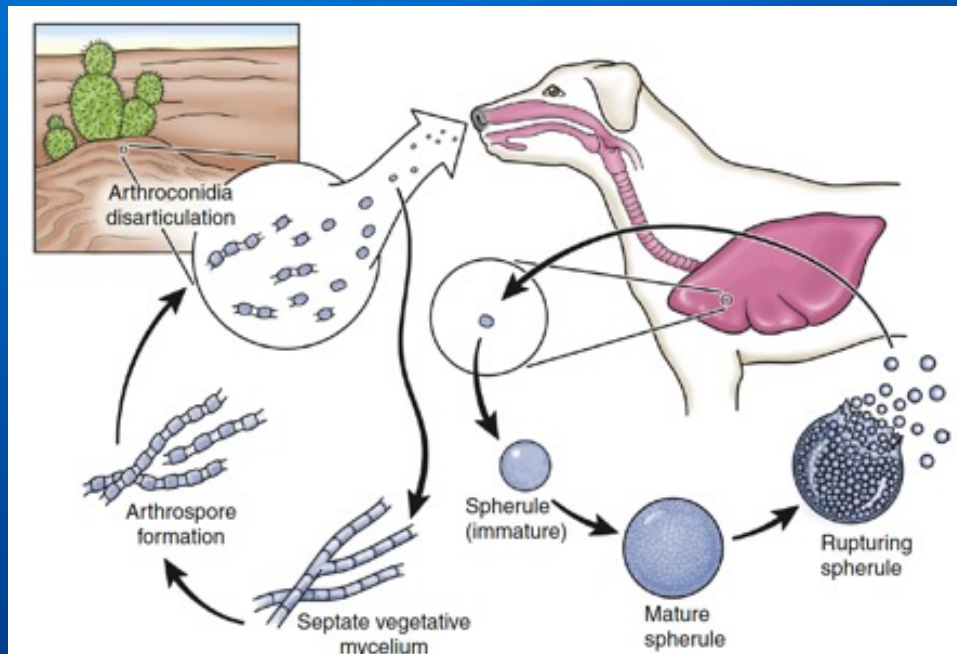


# Background Human Surveillance

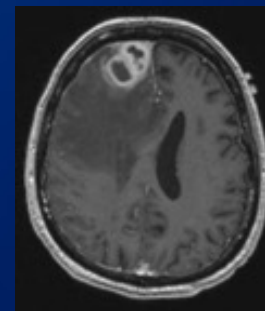
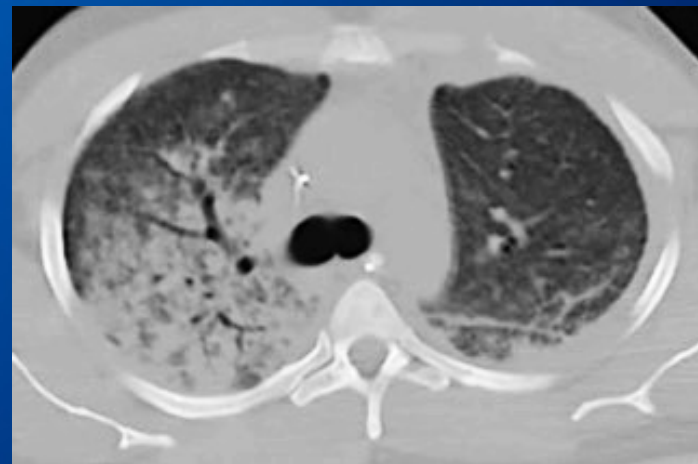


Average annual incidence (per 100,000), 2011-2017 (CDC)

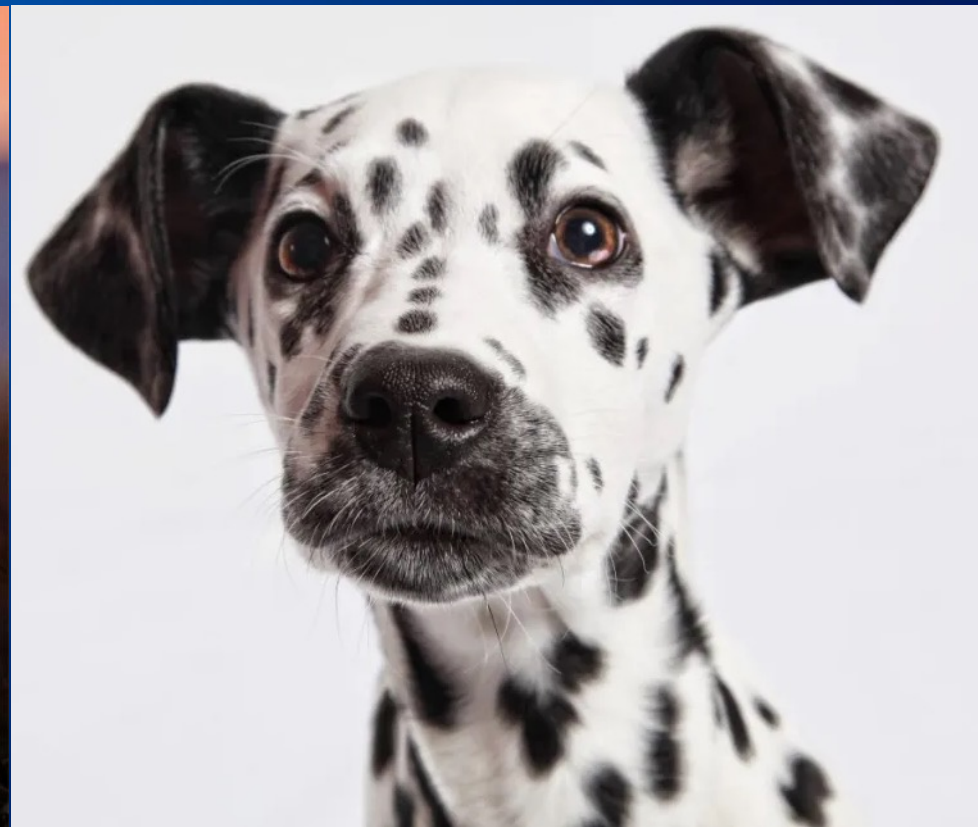
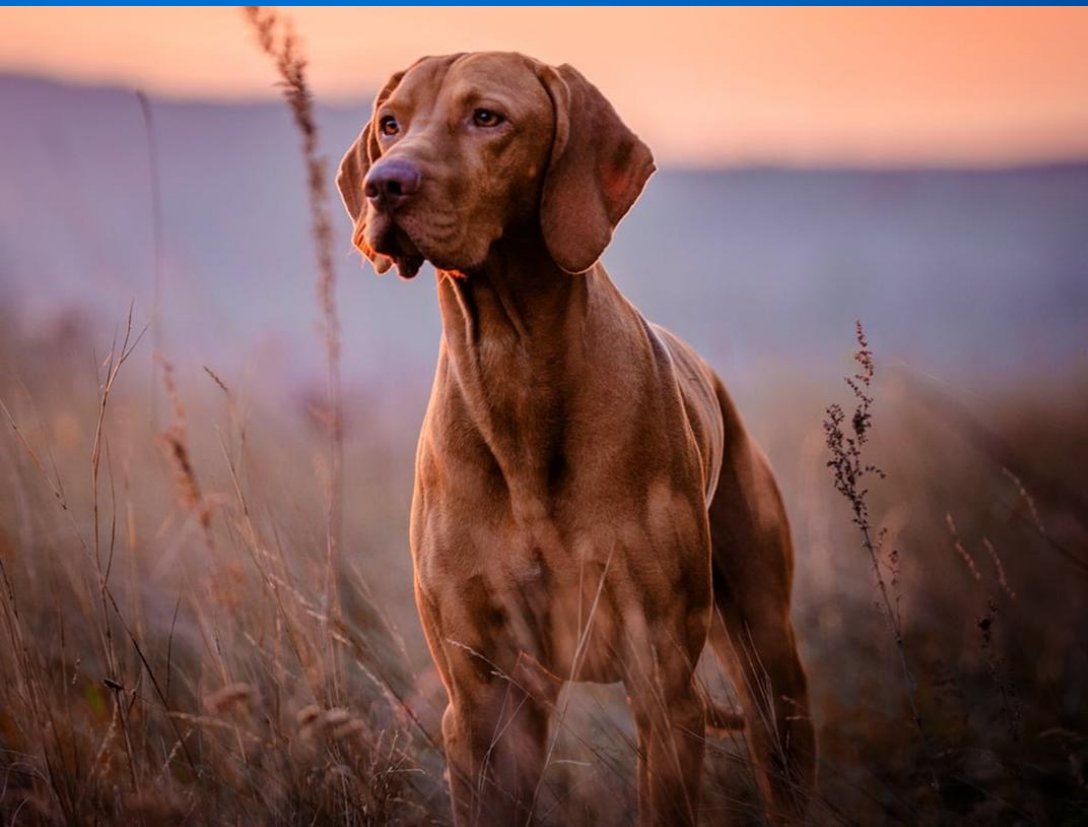
# Background Dogs as a Model



Ref 3



# Background Dogs as a Model



# Background Dogs as a Model

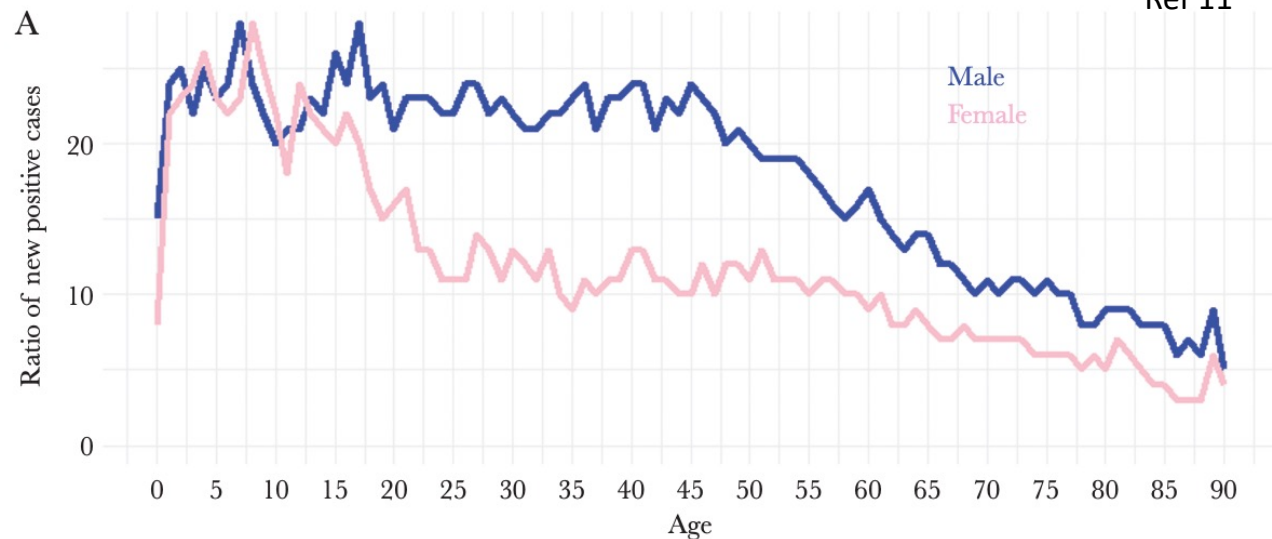
Open Forum Infectious Diseases

## BRIEF REPORT

### Sex Differences in Susceptibility to Coccidioidomycosis

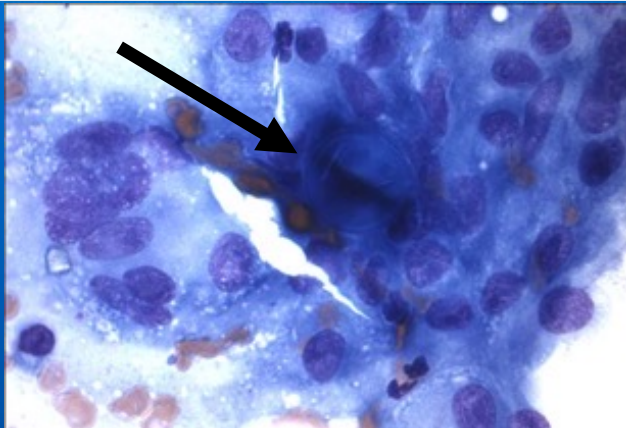
Ian McHardy,<sup>1,2,3</sup> Krystle L. Reagan,<sup>4</sup> Jamie F. Sebastian,<sup>4</sup> Bridget Barker,<sup>5</sup> Derek J. Bays,<sup>2,6</sup> Satya Dandekar,<sup>2,3</sup> Stuart H. Cohen,<sup>2,3,6</sup> Kathleen E. Jennings,<sup>7</sup> Jane Sykes,<sup>2,5</sup> and George R. Thompson III<sup>2,3,6</sup>

<sup>1</sup>Scripps Medical Laboratory, Scripps Health, San Diego, California, USA, <sup>2</sup>University of California Davis Center for Valley Fever, Sacramento, California, USA, <sup>3</sup>Department of Medical Microbiology and Immunology, University of California – Davis, Davis, California, USA, <sup>4</sup>School of Veterinary Medicine, University of California – Davis, Davis, California, USA, <sup>5</sup>Northern Arizona University, Flagstaff, Arizona, USA, <sup>6</sup>Division of Infectious Diseases, Department of Internal Medicine, University of California Davis Medical Center, Sacramento, California, USA, and <sup>7</sup>Born Free USA Primate Sanctuary, Cotulla, Texas, USA

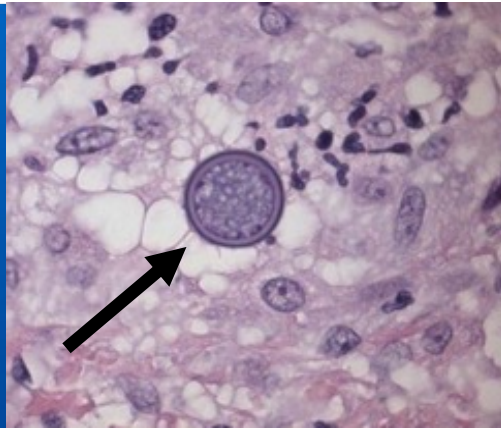


- 220,240 human cases; significant differences in risk between males and females only apparent between 19 and 80 (OR 1.8,  $P < 0.00001$ )
- 82961 male dogs, 80390 female dogs, 2000-2020
- 99 male dog cases, 60 female dog cases (OR 1.6,  $P = 0.0041$ ); risk abolished for male neutered dogs

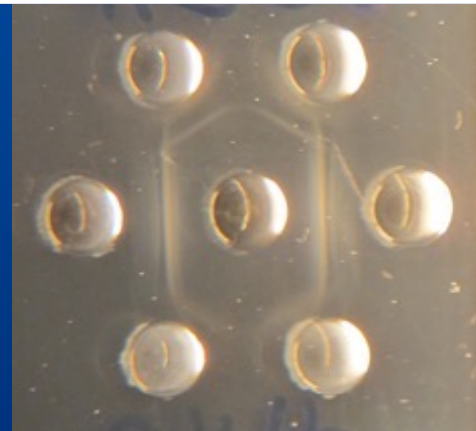
# Background Diagnosis



Cytology



Histopathology



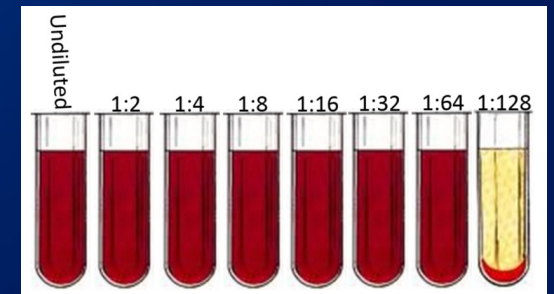
Serology (immunodiffusion)



Serology (lateral flow immunoassay)



Serology (EIA)



Serology (complement fixation)

# Background Dogs as Sentinels

Transboundary and Emerging Diseases

Transboundary

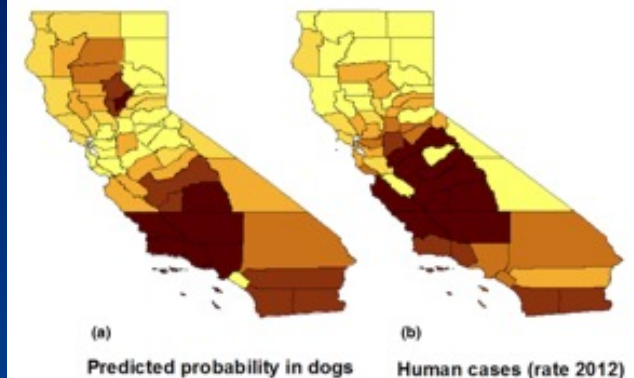
ORIGINAL ARTICLE

## Risk Factors and Spatial Distribution of Canine Coccidioidomycosis in California, 2005–2013

S. E. Grayzel<sup>1</sup>, B. Martinez-López<sup>2</sup> and J. E. Sykes<sup>2</sup>

**Table 1.** Association between dog demographic and environmental variables and coccidioidomycosis occurrence in dogs in California as a result of a multivariable logistic regression model

| Variable                                       | Descriptive statistics |          | Odds ratio (95% confidence interval) | P value <sup>a</sup> |
|------------------------------------------------|------------------------|----------|--------------------------------------|----------------------|
|                                                | Cases                  | Controls |                                      |                      |
| Age in months (mean ± SD) <sup>b</sup>         | 64 ± 36                | 99 ± 44  | 0.3 (0.16–0.56)                      | <0.001               |
| Digging behavior                               |                        |          |                                      |                      |
| No <sup>c</sup>                                | 16                     | 59       | 6.68 (2.47–18.06)                    | <0.001               |
| Yes                                            | 25                     | 20       |                                      |                      |
| Travel to Arizona or California central valley |                        |          |                                      |                      |
| No <sup>c</sup>                                | 14                     | 39       | 4.42 (1.46–13.38)                    | 0.005                |
| Yes                                            | 27                     | 40       |                                      |                      |



Ref 12



# Background

Zoonoses and Public Health

## ORIGINAL ARTICLE

### Identifying Areas of High Risk of Human Exposure to Coccidioidomycosis in Texas Using Serology Data from Dogs

R. Gautam<sup>1</sup>, I. Srinath<sup>1</sup>, A. Clavijo<sup>2</sup>, B. Szonyi<sup>1</sup>, M. Bani-Yaghoub<sup>1</sup>, S. Park<sup>1</sup> and R. Ivanek<sup>1</sup>

<sup>1</sup> Department of Veterinary Integrative Biosciences, College of Veterinary Medicine and Biomedical Sciences, Texas A&M University, College Station, TX, USA

<sup>2</sup> Texas Veterinary Medical Diagnostic Laboratories, College Station, TX, USA

#### Impacts

- Described the spatial distribution of Valley Fever among dogs in Texas.
- Identified an area with a high rate of seropositivity for Valley Fever in dogs in the western and southwestern parts of Texas.
- Confirmed that dogs can be used to determine the spatial distribution of human exposure risk to Valley Fever.

Ref 13. Gautam R, et al. Zoonoses Public Health 2013;60:174-181

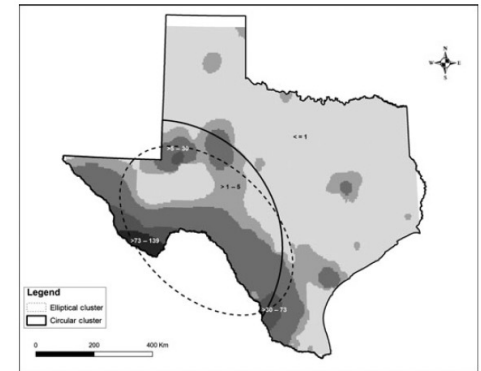


Fig. 3. Isopleth map obtained through kriging of the empirical Bayesian-smoothed seropositive rates of VF among dogs in Texas over the study period (July 1999–December 2009). The seropositive rate distribution is represented by graduated scale of shading in which rate decreases as the shade becomes lighter. The map is overlaid with the identified spatio-temporal cluster that spans the period from August 2004 to October 2009 and from September 2003 to November 2008, based on the cylindrical and ellipsoidal scanning windows, respectively.

July 1999 – December 2009  
Seroprevalence 2,547/5,871 (43%)

# Background Dogs as Sentinels

Ref 14

## Coccidioidomycosis Acquired in Washington State

**Nicola Marsden-Haug,<sup>1</sup> Marcia Goldoft,<sup>1</sup> Cindy Ralston,<sup>2</sup> Ajit P. Limaye,<sup>4</sup> Jimmy Chua,<sup>3</sup> Heather Hill,<sup>2</sup> Larry Jecha,<sup>2</sup> George R. Thompson III,<sup>5</sup> and Tom Chiller<sup>6</sup>**

<sup>1</sup>Office of Communicable Disease Epidemiology, Washington State Department of Health, Shoreline, <sup>2</sup>Communicable Disease Program, Benton-Franklin Health District, and <sup>3</sup>Infectious Diseases Department, Kennewick General Hospital, Kennewick, and <sup>4</sup>Division of Allergy and Infectious Diseases, University of Washington, Seattle, Washington; <sup>5</sup>Coccidioidomycosis Serology Laboratory, University of California, Davis; and <sup>6</sup>Mycotic Diseases Branch, Centers for Disease Control and Prevention, Atlanta, Georgia



- Acute disease, three unrelated people from SE WA, no travel
  - 2010: 12-year-old boy with pneumonia, biked, dug in canyon
  - 2010: 15-year-old boy with knee cellulitis after ATV crash
  - 2011: 58-year-old man with pneumonia, construction excavator

# Background Dogs as Sentinels



Review

## Coccidioidomycosis in Brazil: Historical Challenges of a Neglected Disease

Rossana Cordeiro <sup>1,\*</sup>, Santiago Moura <sup>1,†</sup>, Débora Castelo-Branco <sup>1,\*</sup>, Marcos Fábio Rocha <sup>1,2</sup>, Reginaldo Lima-Neto <sup>3</sup> and José Júlio Sidrim <sup>1</sup>

“The international scientific community turned its attention to coccidioidomycosis in Brazil after an outbreak...

In August 1991, three individuals and eight of their dogs who went on an armadillo hunt developed severe pneumonia; three dogs died a few days later”

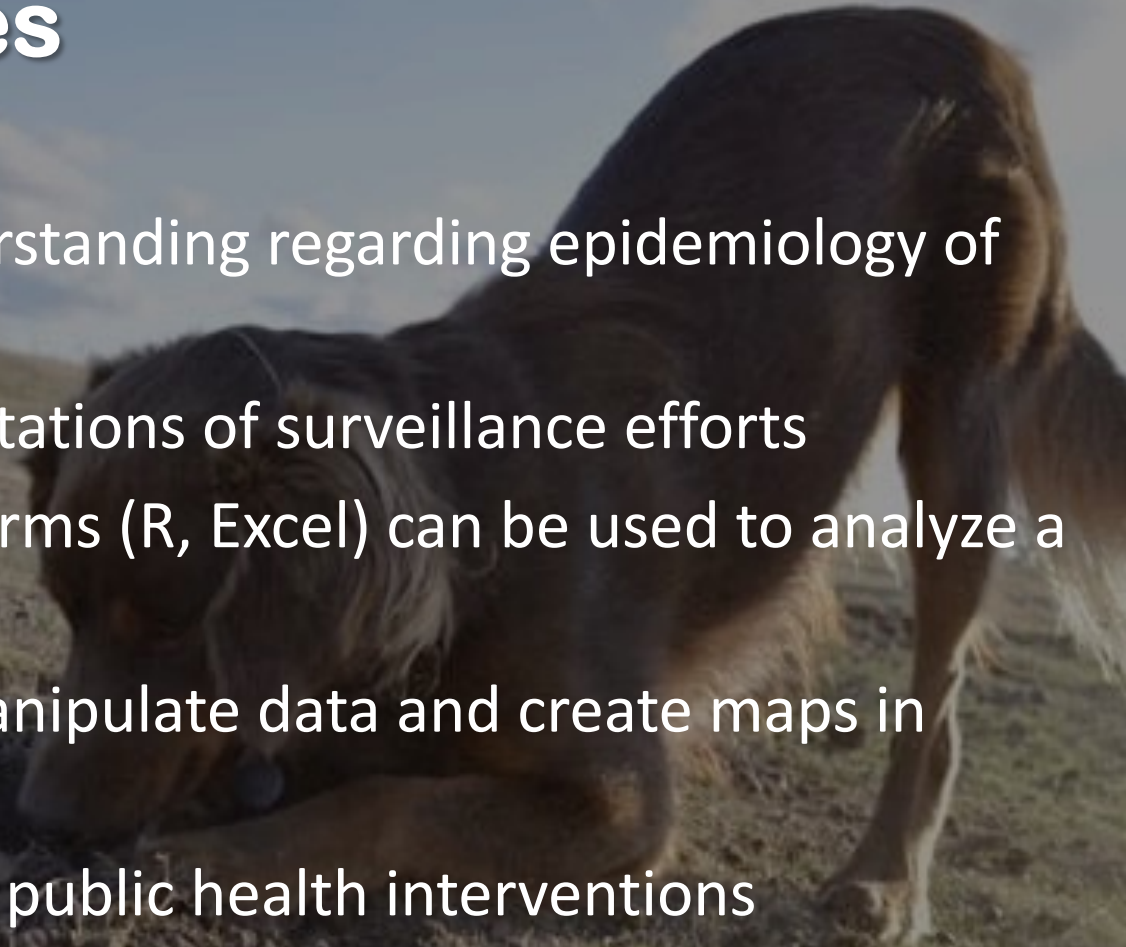
Ref 15

# Hypothesis and Study Goals

- Serosurveillance efforts in dogs will provide insight into the epidemiology of coccidioidomycosis in humans
- Spatial analysis of serologic test data will correlate with that found for humans and reveal emerging regions of endemicity
- The inter-annual and seasonal pattern of seroreactivity in dogs will match those known for people
- Inter- and intra-annual variation in seroreactivity will correlate with climatic variables
- Overarching goal: demonstrate value of dogs as a sentinel

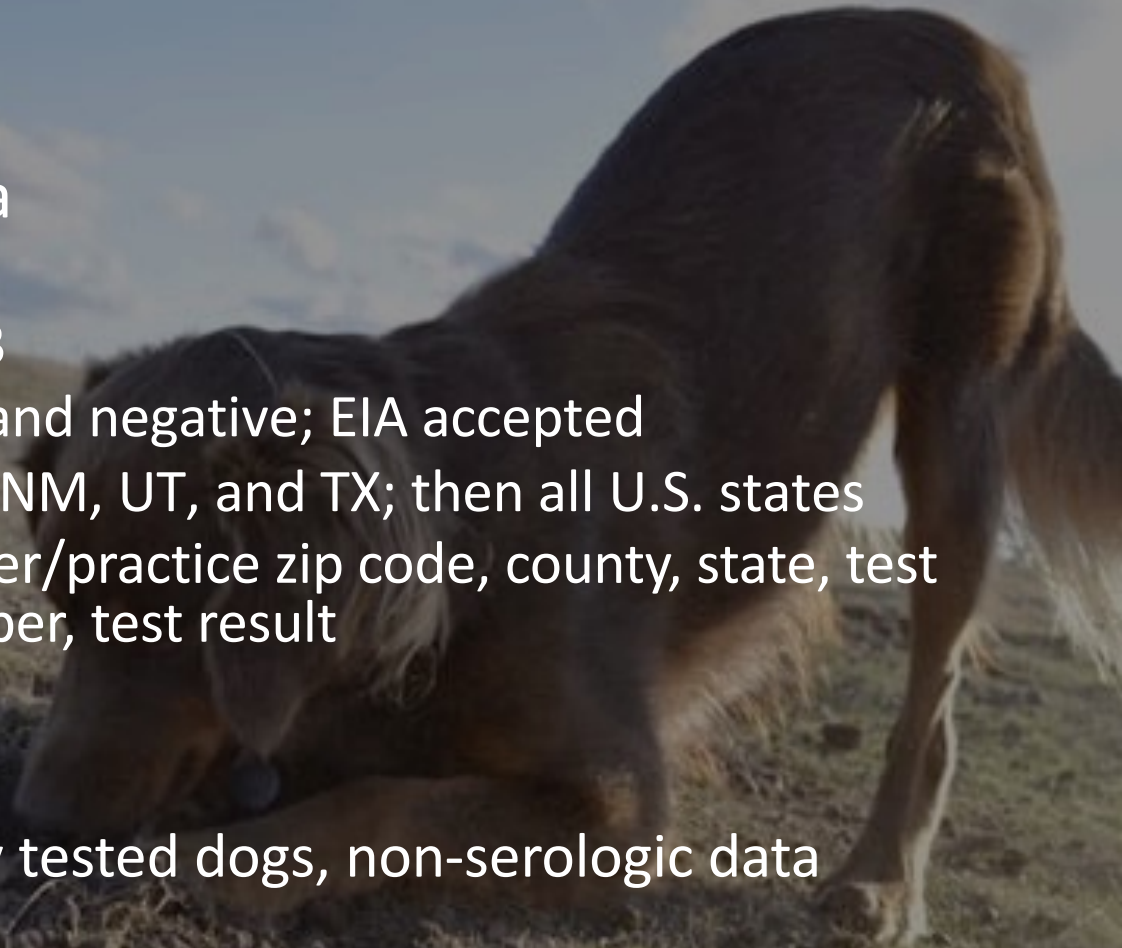
# Learning Objectives

- Develop an in-depth understanding regarding epidemiology of coccidioidomycosis
- Understand value and limitations of surveillance efforts
- Learn how software platforms (R, Excel) can be used to analyze a large set of health data
- Know steps required to manipulate data and create maps in ArcGIS Pro
- Interpret results to inform public health interventions
- Refine writing and oral presentation skills



# Methods

- Gathered 'big' serologic data
  - 7 different laboratories
  - January 2012 to August 2023
  - AGID, IgM and IgG, positive and negative; EIA accepted
  - Initially CA, AZ, OR, WA, NV, NM, UT, and TX; then all U.S. states
  - Dog ID, age, breed, sex, owner/practice zip code, county, state, test date (month, year), lab number, test result
- Data clean-up (R)
  - Raw 2,141,055 visits
  - Removed non-dogs, serially tested dogs, non-serologic data
  - Final 887,764 dogs
  - Harmonized location data



# Methods

## Statistical Analysis

- Risk factors: breed, body size, sex, age
  - Breed categories
    - Closely-related breeds (n = 155)
    - Behavior-associated groups (n = 53)
    - American Kennel Club (AKC) breed groups (n = 9) (e.g., working, sporting, herding, mixed breed, hound, toy)
    - AKC breed size groupings (n = 5)
  - Sex categories
    - MI, FI, MN, FS, MU, FU
  - Age by year (continuous), groups
  - Chi square test (GraphPad Prism), unpaired t-test (Excel)

iScience

CellPress  
OPEN ACCESS

### Article

## Breed, age, and social environment are associated with personality traits in dogs

Milla Salonen,<sup>1,2,3,4,\*</sup> Salla Mikkola,<sup>1,2,3</sup> Julia E. Niskanen,<sup>1,2,3</sup> Emma Hakanen,<sup>1,2,3</sup> Sini Sulkama,<sup>1,2,3</sup> Jenni Puurunen,<sup>1,2,3</sup> and Hannes Lohi<sup>1,2,3,5,\*</sup>

**Table S1.** Breeds and breed groups, related to STAR Methods.

|                                                                                                                                                                                              |                                                                                                                                                                            |                                                                                                                                                                                                                                  |                                                                                                                                                                                                                                                                                                                                         |
|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <b>Australian Shepherd</b><br>Australian Shepherd<br>Miniature American Shepherd                                                                                                             | <b>Belgian shepherd dogs</b><br>Groenendael<br>Laekenois<br>Malinois<br>Tervueren                                                                                          | <b>Bernese Mountain Dogs</b><br>Appenzel Cattle Dog<br>Bernese Mountain Dog<br>Entlebucher Mountain Dog<br>Greater Swiss Mountain Dog                                                                                            | <b>Bichon type dogs</b><br>Bichon Frise<br>Bolognese<br>Coton de Tulear<br>Havanese<br>Lowchen<br>Maltese<br>Russian Tsvetnaya Bolonka                                                                                                                                                                                                  |
| <b>Brachycephalic dogs</b><br>Boston Terrier<br>Bulldog<br>French Bulldog<br>Olde English Bulldogge<br>Pug                                                                                   | <b>Bull type terriers</b><br>Bull Terrier<br>Miniature Bull Terrier<br>Staffordshire Bull Terrier                                                                          | <b>Dachshunds</b><br>Miniature Longhaired<br>Miniature Shorthaired<br>Miniature Wirehaired<br>Rabbit Longhaired<br>Rabbit Shorthaired<br>Rabbit Wirehaired<br>Standard Longhaired<br>Standard Shorthaired<br>Standard Wirehaired | <b>English herders</b><br>Australian Cattle Dog<br>Australian Kelpie<br>Australian Koolie<br>Bearded Collie<br>Lancashire Heeler<br>Old English Sheepdog<br>Welsh Sheepdog                                                                                                                                                              |
| <b>European sighthounds</b><br>Borzoi<br>Deerhound<br>Greyhound<br>Hungarian Greyhound<br>Irish Wolfhound<br>Italian Sighthound<br>Polish Greyhound<br>Silken Windhound<br>Spanish Greyhound | <b>Fighting dogs</b><br>American Bulldog<br>American Bully<br>American Pit Bull Terrier<br>American Staffordshire Terrier<br>Cane Corso<br>Dogo Argentino<br>Presa Canario | <b>German spitz related breeds</b><br>Eurasier<br>Giant Spitz<br>Medium size Spitz<br>Miniature Spitz<br>Japanese Spitz<br>Keeshond<br>Pomeranian<br>Volpino Italiano                                                            | <b>Hunting terriers</b><br>Bedlington Terrier<br>Border Terrier<br>Cairn Terrier<br>Cesky Terrier<br>Dandie Dinmont Terrier<br>Fox Terrier Smooth<br>Fox Terrier Wire<br>German Hunting Terrier<br>Glen of Imaal Terrier<br>Lakeland Terrier<br>Norfolk Terrier<br>Norwich Terrier<br>Scottish Terrier<br>Skye Terrier<br>Welsh Terrier |



Ref 16



# Methods

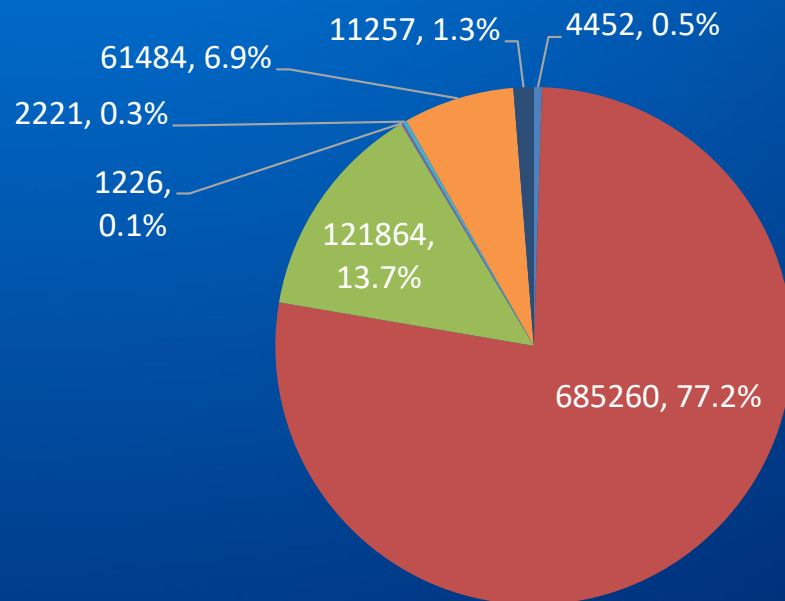
## Statistical Analysis

- Examined total tests, test positivity, incidence/10,000 households – annual and seasonal
- Inter-annual and monthly/seasonal incidence compared with human data for CA and AZ
  - CA Department of Public Health (monthly county-level) and CA Department of Finance
  - AZ Department of Health Services (annual county-level; dog data aggregated)
  - Pearson's correlation coefficient (R)
- Spatial analysis (ArcGIS Pro)
  - Data joined with county shapefiles (US Census Bureau)
  - Choropleth mapping (tests, prevalence, incidence)
    - USA Albers Equal Area Conic projection
  - Spatial autocorrelation
    - Moran's I
  - Hot spot analysis
    - Getis Ord-Gi\*, fixed distance band method



# Results Laboratories

Proportion of Samples by Laboratory, 2012-2023 (n = 887,764)

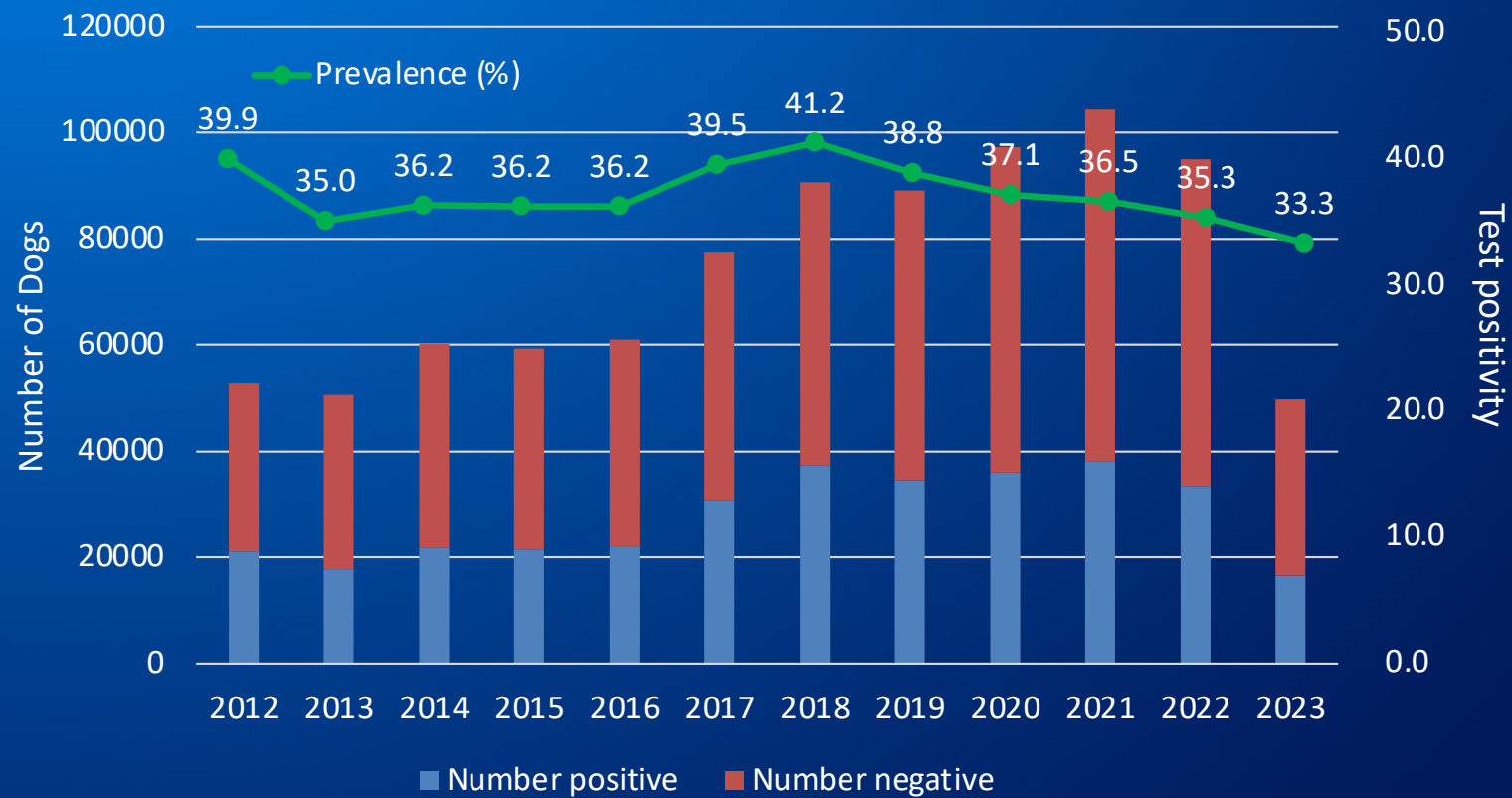


Test positivity 330,959/887,764 (37.3%)

■ Lab1 ■ Lab2 ■ Lab3 ■ Lab4 ■ Lab5 ■ Lab6 ■ Lab7

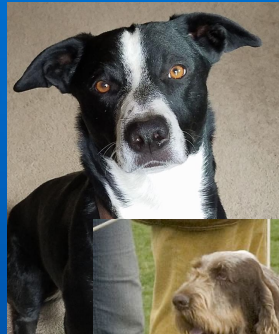
# Results

## Total Tests and Test Positivity

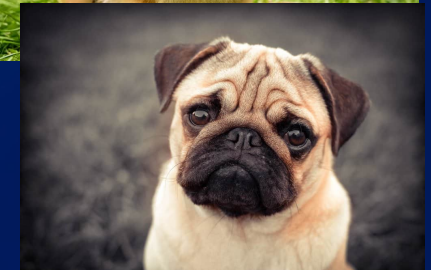


# Results

## Individual Breed Analysis: 682,078 dogs (77%)



| Breed                          | Number positive | Number negative | Test Positivity | OR (95% CI)         | P value  |
|--------------------------------|-----------------|-----------------|-----------------|---------------------|----------|
| McNab                          | 29              | 6               | 82.9%           | 7.62 (3.25 – 17.85) | < 0.0001 |
| Spinone Italiano               | 71              | 41              | 63.4%           | 2.89 (1.96 – 4.26)  | < 0.0001 |
| Viszla                         | 1852            | 1082            | 63.1%           | 2.87 (2.60 – 3.18)  | < 0.0001 |
| Polish Lowland sheepdog        | 22              | 15              | 59.5%           | 2.44 (1.27 – 4.67)  | 0.0094   |
| Golden retriever mix           | 2834            | 1970            | 59.0%           | 2.41 (2.21 – 2.64)  | < 0.0001 |
| Scottish terrier               | 964             | 684             | 58.5%           | 2.37 (2.10 – 2.66)  | < 0.0001 |
| Weimaraner                     | 2245            | 1877            | 54.5%           | 2.01 (1.83 – 2.20)  | < 0.0001 |
| Bull terrier                   | 1014            | 856             | 54.2%           | 1.99 (1.78 – 2.23)  | < 0.0001 |
| Afghan hound                   | 144             | 125             | 53.5%           | 1.93 (1.51 – 2.48)  | < 0.0001 |
| Saluki                         | 73              | 68              | 51.8%           | 1.80 (1.29 – 2.52)  | 0.0005   |
| Shetland sheepdog              | 432             | 1999            | 17.8%           | 0.36 (0.32 – 0.41)  | < 0.0001 |
| Pomeranian                     | 1367            | 5913            | 18.8%           | 0.39 (0.35 – 0.42)  | < 0.0001 |
| Pug                            | 1700            | 7258            | 19.0%           | 0.39 (0.36 – 0.42)  | < 0.0001 |
| Japanese Chin                  | 84              | 340             | 19.8%           | 0.42 (0.33 – 0.53)  | < 0.0001 |
| Brussels Griffon               | 139             | 547             | 20.3%           | 0.43 (0.35 – 0.52)  | < 0.0001 |
| Maltese                        | 1853            | 7282            | 20.3%           | 0.43 (0.39 – 0.46)  | < 0.0001 |
| Italian greyhound              | 240             | 919             | 20.7%           | 0.44 (0.37 – 0.51)  | < 0.0001 |
| French bulldog                 | 1016            | 3558            | 22.2%           | 0.48 (0.44 – 0.53)  | < 0.0001 |
| Chihuahua                      | 10419           | 36358           | 22.3%           | 0.48 (0.45 – 0.51)  | < 0.0001 |
| Cavalier King Charles Spaniels | 884             | 2793            | 24.0%           | 0.53 (0.48 – 0.59)  | < 0.0001 |



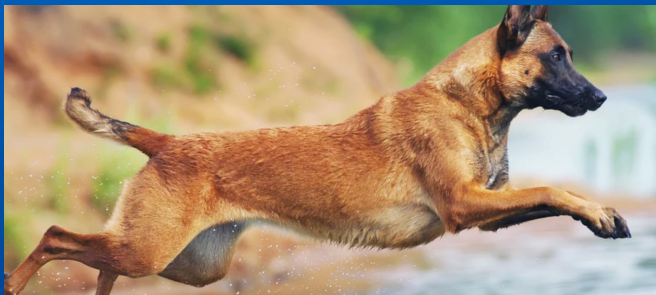
Reference group greyhounds, 1340/3589, prevalence 37.3%

# Results

## American Kennel Club Size Group Analysis: 568,997 dogs (64%)

| Body Size | Number positive | Number negative | Test Positivity | OR (95% CI)        | P value  |
|-----------|-----------------|-----------------|-----------------|--------------------|----------|
| Large     | 78674           | 102693          | 43.4%           | 1.24 (1.21 – 1.27) | < 0.0001 |
| Medium    | 58284           | 76478           | 43.2%           | 1.23 (1.20 – 1.26) | < 0.0001 |
| Small     | 38315           | 86826           | 30.6%           | 0.71 (0.70 – 0.73) | < 0.0001 |
| Toy       | 22004           | 68891           | 24.2%           | 0.51 (0.50 – 0.53) | < 0.0001 |

Reference group Giant dogs  
(14087/36832, prevalence  
38.2%)





# Results

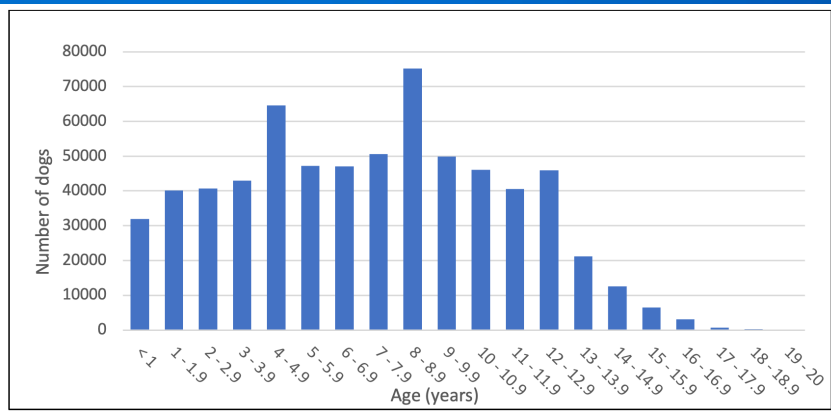
## Sex Analysis: 804,426 (91%)

| Sex               | Number positive | Number negative | Test Positivity | OR (95% CI)        | P value  |
|-------------------|-----------------|-----------------|-----------------|--------------------|----------|
| Male not stated   | 30318           | 43068           | 41.3%           | 1.16 (1.14 – 1.19) | < 0.0001 |
| Male neutered     | 113705          | 178328          | 38.9%           | 1.1 (1.03 – 1.08)  | < 0.0001 |
| Female not stated | 24730           | 40885           | 37.7%           | 1.00 (0.99 – 1.02) | 0.98     |
| Female neutered   | 108482          | 188595          | 36.5%           | 0.95 (0.93 – 0.97) | < 0.0001 |
| Female intact     | 11122           | 20977           | 34.6%           | 0.88 (0.85 – 0.90) | < 0.0001 |

Reference group  
intact male dogs  
(16668/27548,  
prevalence 37.7%)

# Results

## Age Analysis: 667,279 (75%)

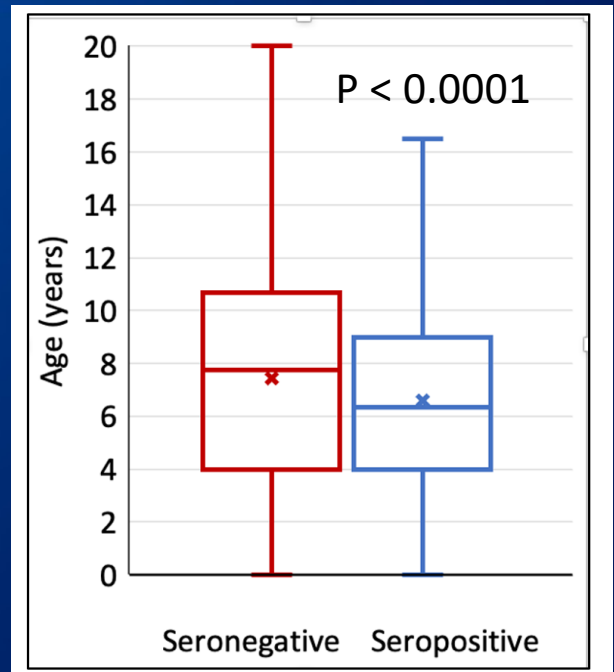


Distribution of age categories for dogs tested for *Coccidioides* spp. 2012-2023



| Age Group (years) | Number positive | Number negative | Test Positivity | OR (95% CI)        | P value  |
|-------------------|-----------------|-----------------|-----------------|--------------------|----------|
| Less than 1       | 8836            | 23102           | 27.7%           | 0.63 (0.61 – 0.65) | < 0.0001 |
| 1 to 3            | 33857           | 46944           | 41.9%           | 1.19 (1.17 – 1.21) | < 0.0001 |
| 3 to 6            | 70356           | 84296           | 45.5%           | 1.36 (1.36 – 1.40) | < 0.0001 |
| 6 to 8            | 42075           | 55605           | 43.1%           | 1.25 (1.23 – 1.27) | < 0.0001 |
| 10 to 15          | 47928           | 118419          | 28.8%           | 0.67 (0.66 – 0.68) | < 0.0001 |
| 15 to 20          | 1940            | 8688            | 18.3%           | 0.37 (0.35 – 0.39) | < 0.0001 |

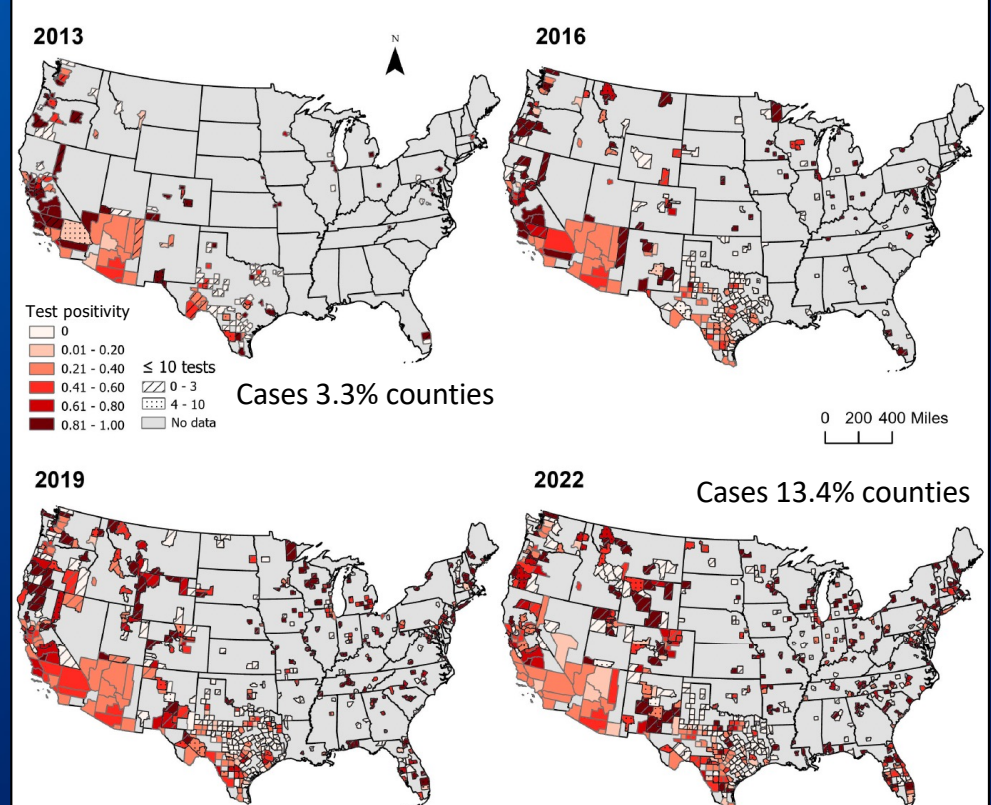
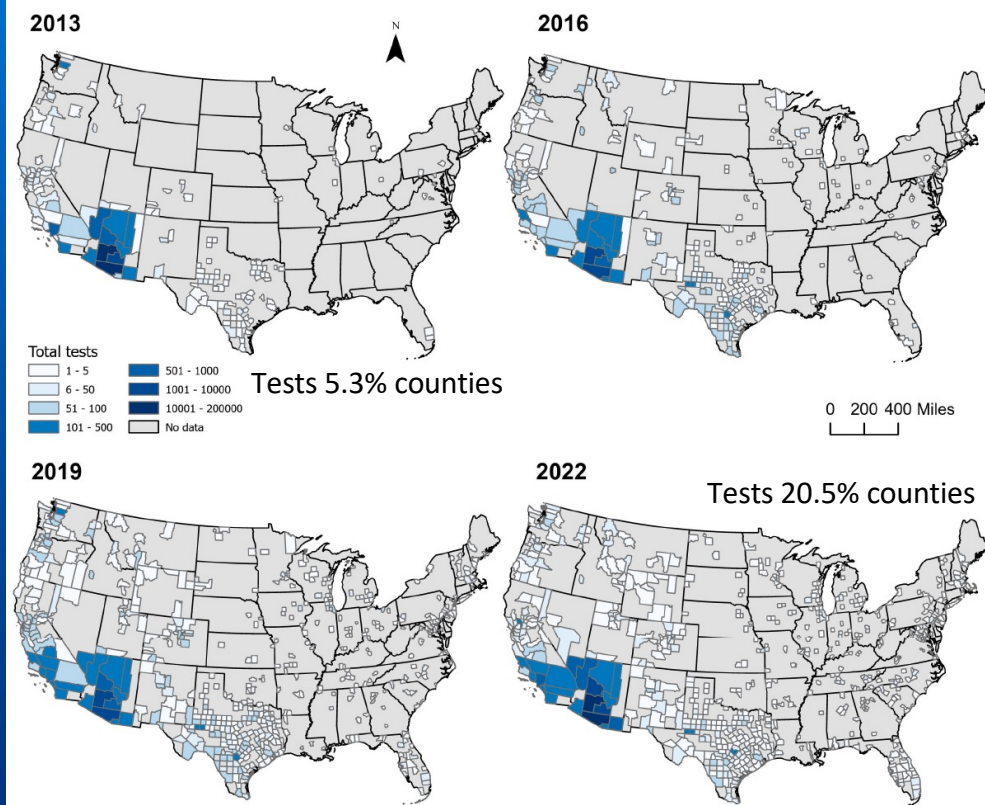
Reference group dogs 8 to 10 years of age (47239/125123, prevalence 37.8%)



Comparison of age distributions in dogs seronegative and seropositive for *Coccidioides* spp. antibodies

# Results

## Total Tests and %Test Positivity Maps





# Results

## Overall Test Positivity, 2012-2023

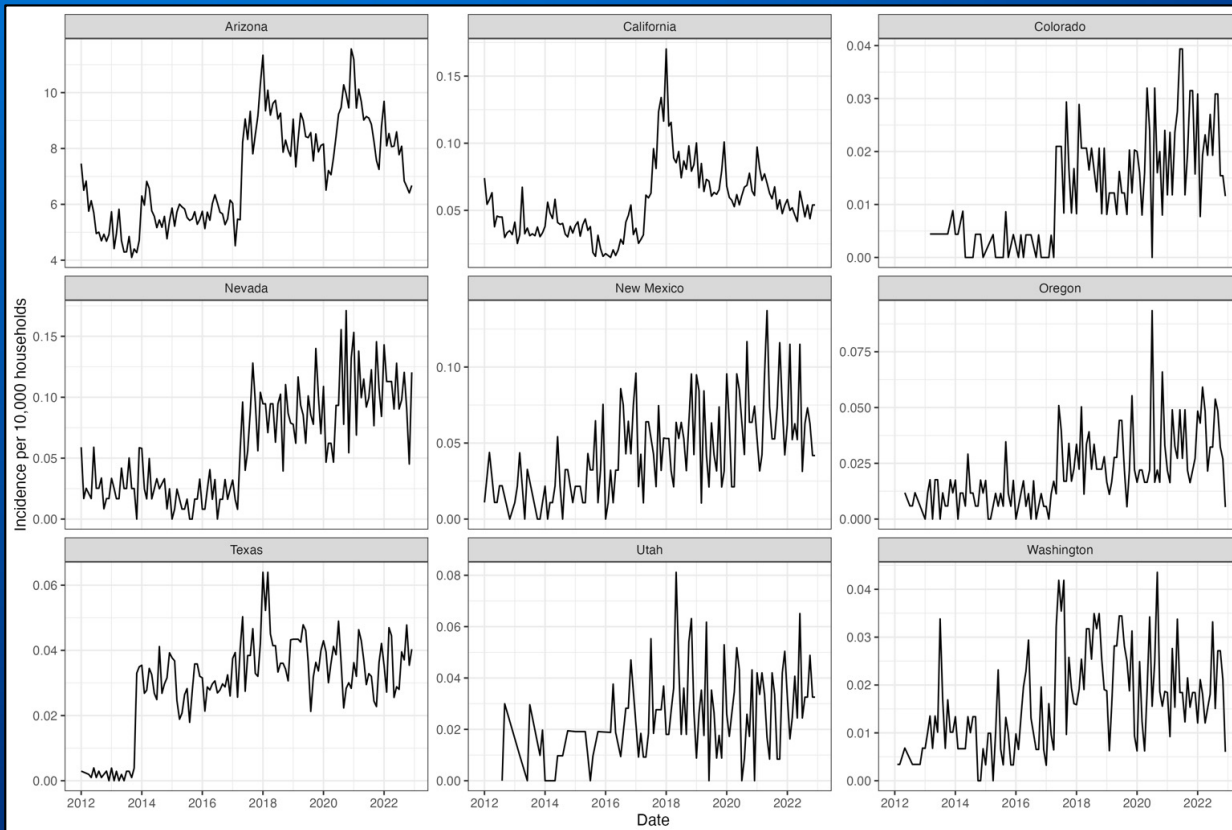
| State      | Number positive | Number negative | Test Positivity | OR (95% CI)        |
|------------|-----------------|-----------------|-----------------|--------------------|
| Texas      | 4998            | 10776           | 31.7%           | 0.87 (0.84 – 0.90) |
| Washington | 695             | 1145            | 37.8%           | 1.14 (1.04 – 1.26) |
| Nevada     | 526             | 747             | 41.3%           | 1.33 (1.19 – 1.48) |
| New Mexico | 780             | 807             | 49.1%           | 1.82 (1.65 – 2.01) |
| California | 8528            | 8464            | 50.2%           | 1.90 (1.84 – 1.96) |
| Oregon     | 612             | 424             | 59.1%           | 2.72 (2.40 – 3.08) |
| Utah       | 300             | 149             | 66.8%           | 3.79 (3.12 – 4.62) |
| Colorado   | 869             | 256             | 77.2%           | 6.39 (5.56 – 7.35) |

with 95% CI

Reference group AZ, 248987/717930, test positivity 34.7%; all P values < 0.0001

# Results

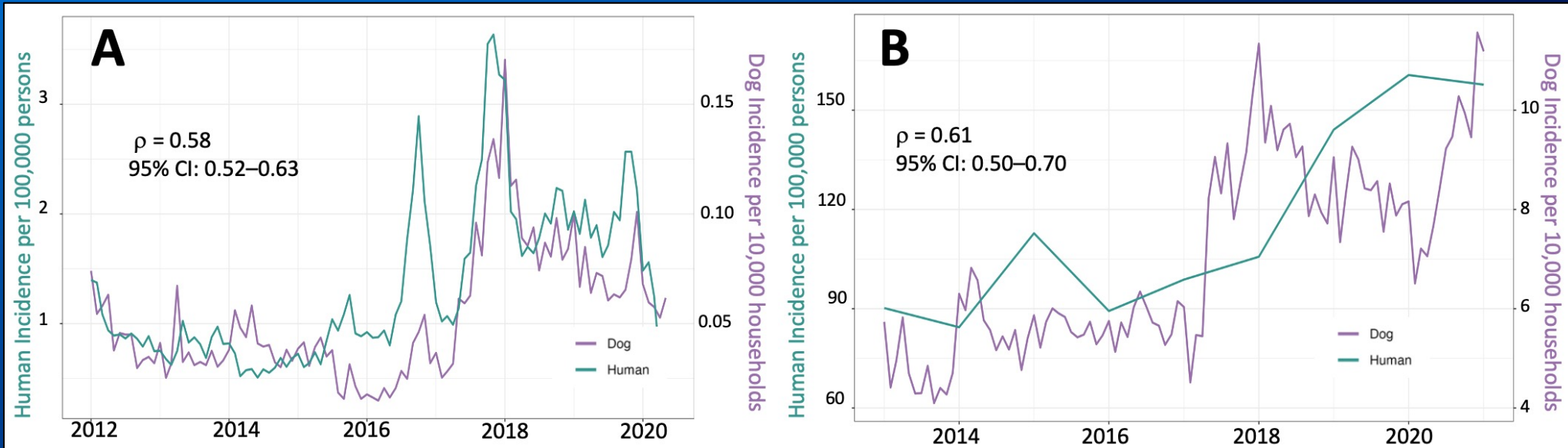
## Incidence Curves, 2012-2023



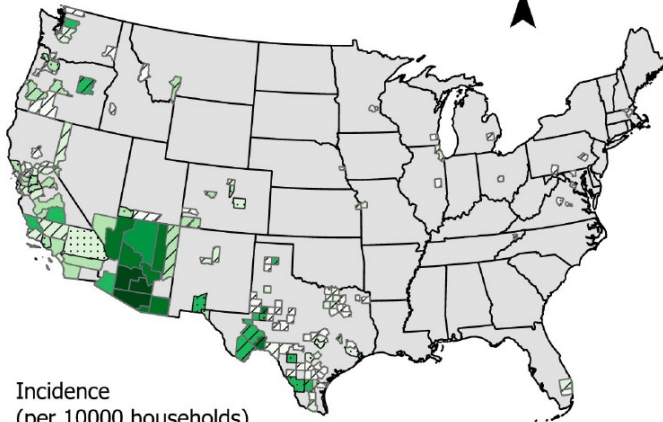
- AZ incidence rates 80 to 100-fold higher than CA, 100- to 1000-fold other SW states
- 2018 peak in AZ, CA, TX
- Incidence rates in NV and NM exceeded CA 2019-2023
- Rates in WA, OR, UT, TX low

# Results

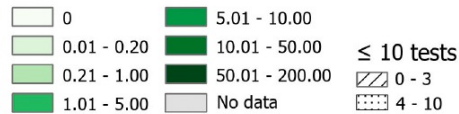
## Dog vs. Human Incidence Curves, 2012-2023



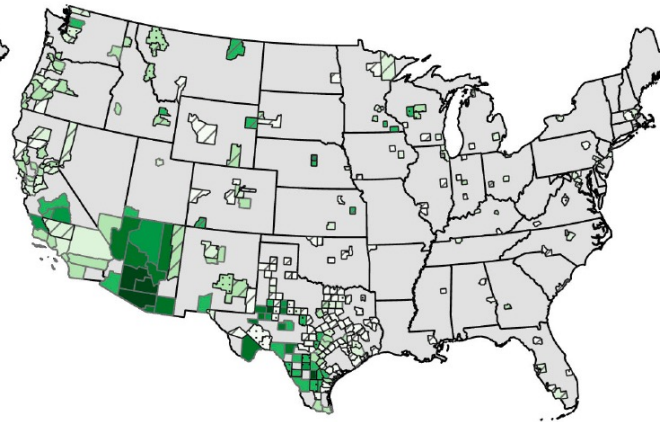
2013



Incidence  
(per 10000 households)

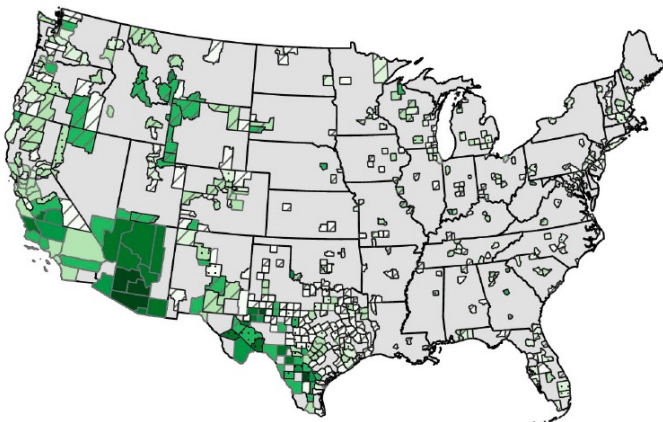


2016

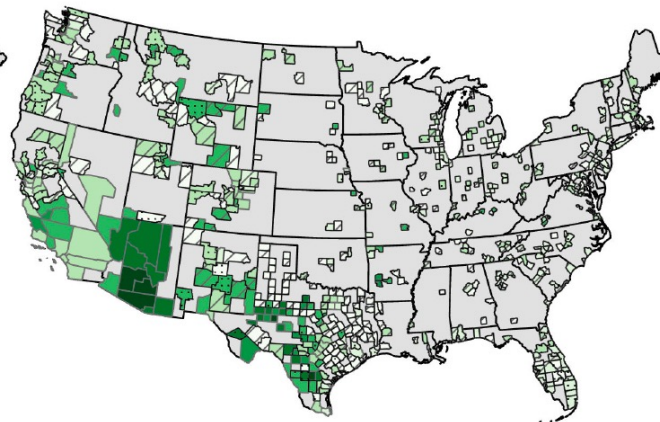


0 200 400 Miles

2019



2022



# Results Incidence Maps (Annual)

## Overall Incidence:

Arizona (87.1)

Nevada (0.78)

California (0.67)

New Mexico (0.53)

Montana (0.35)

Texas (0.34)

Oregon (0.29)

Colorado (0.26)

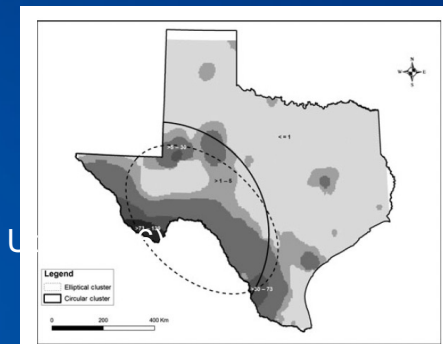
Washington (0.20)

Wyoming (0.19)

Idaho (0.17)

Remaining:

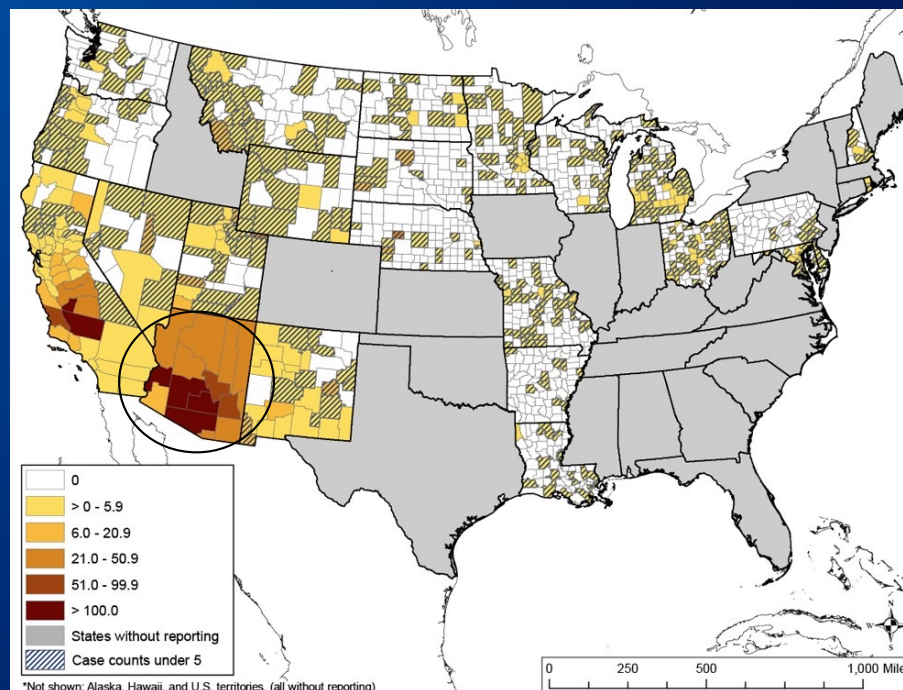
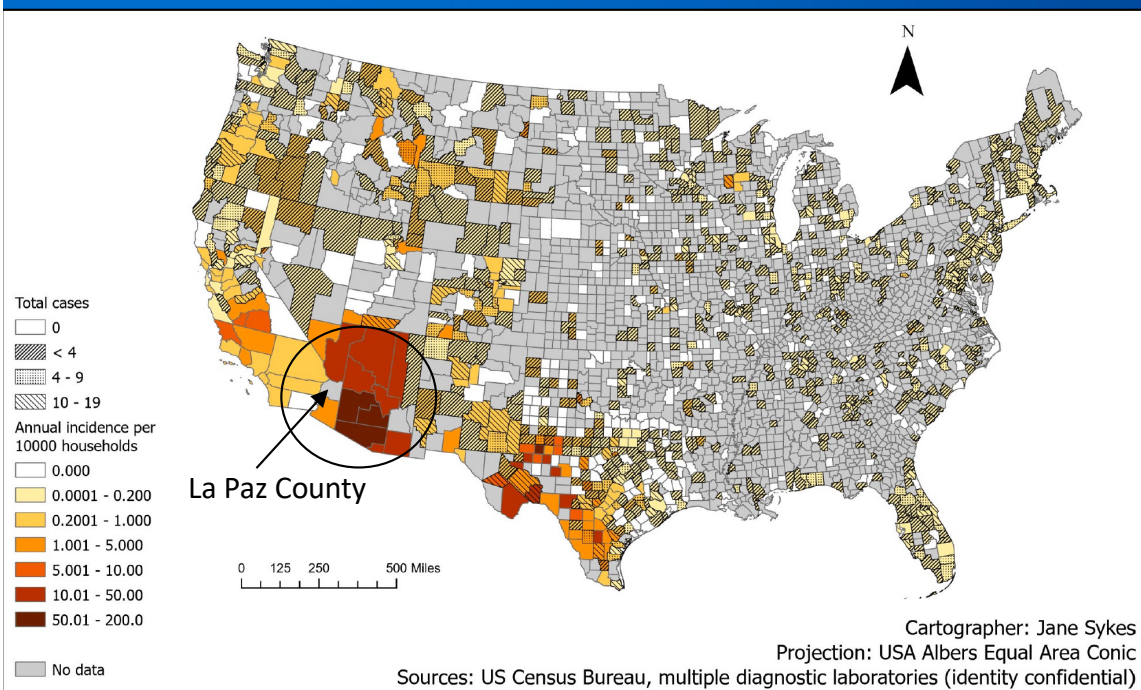
0.007 (AL) to 0.09 (KS)



Gautam et al. 2013

# Results

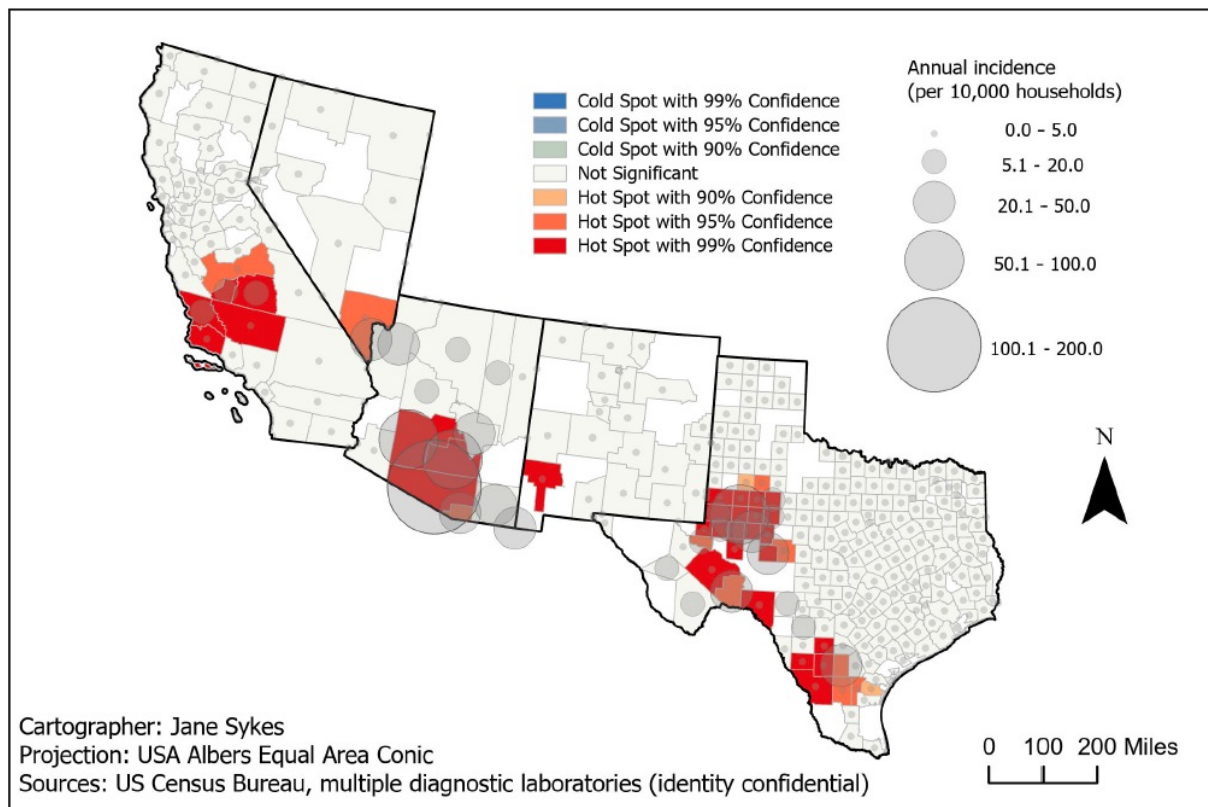
## Dog vs. Human Incidence Maps, 2012-2023



La Paz County population 16,557 (Maricopa 4.6 million)  
Largest city, Parker, 3,417 people  
Cases from Parker listed as Mohave county

# Results

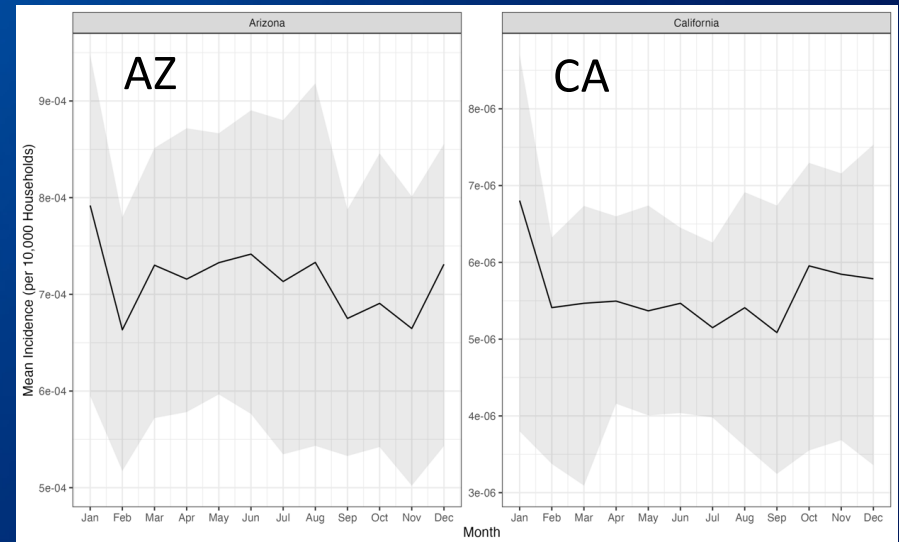
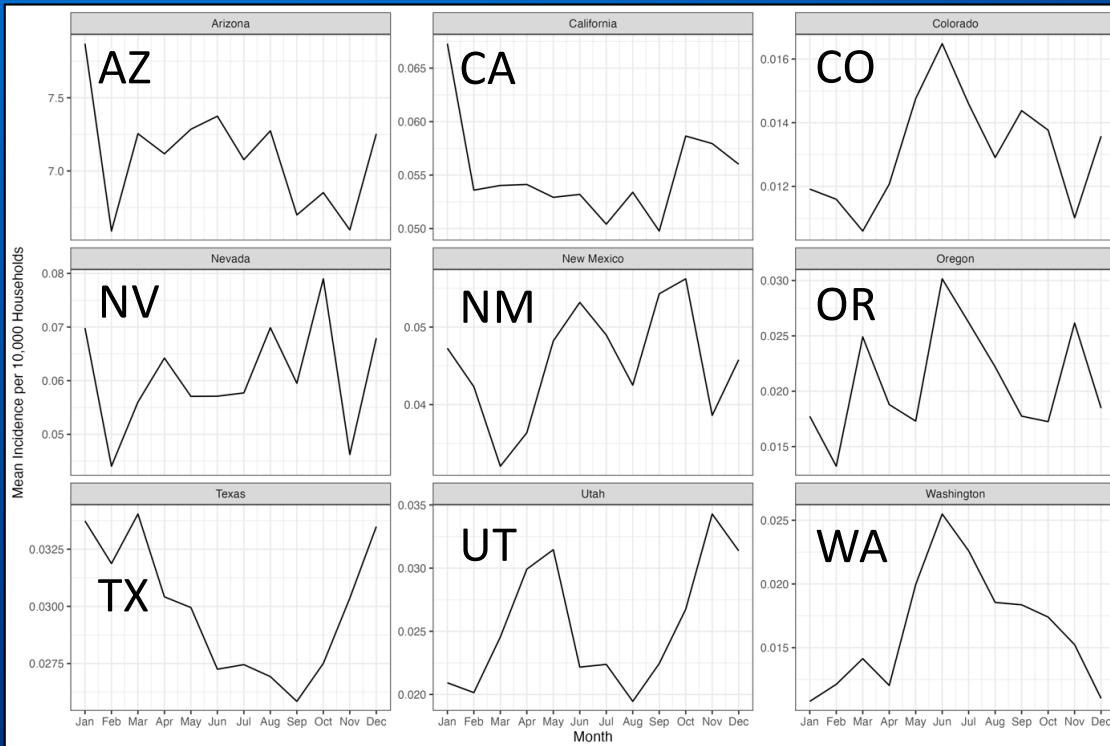
## Hot Spot Analysis, 2012-2023



Spatial autocorrelation  
(Moran's I),  $p < 0.0001$

# Results

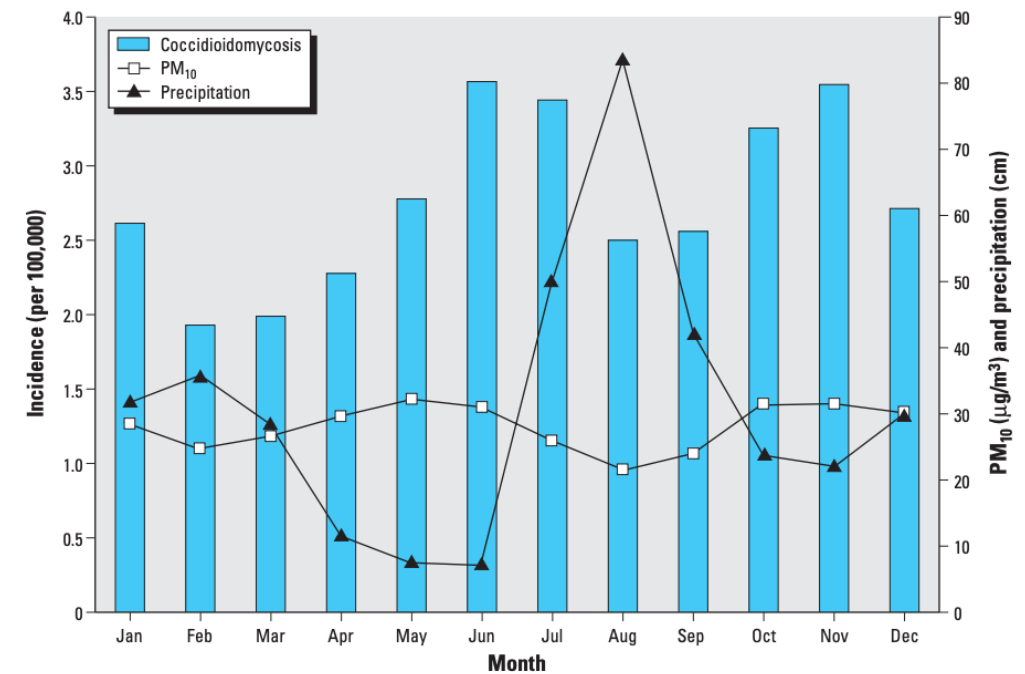
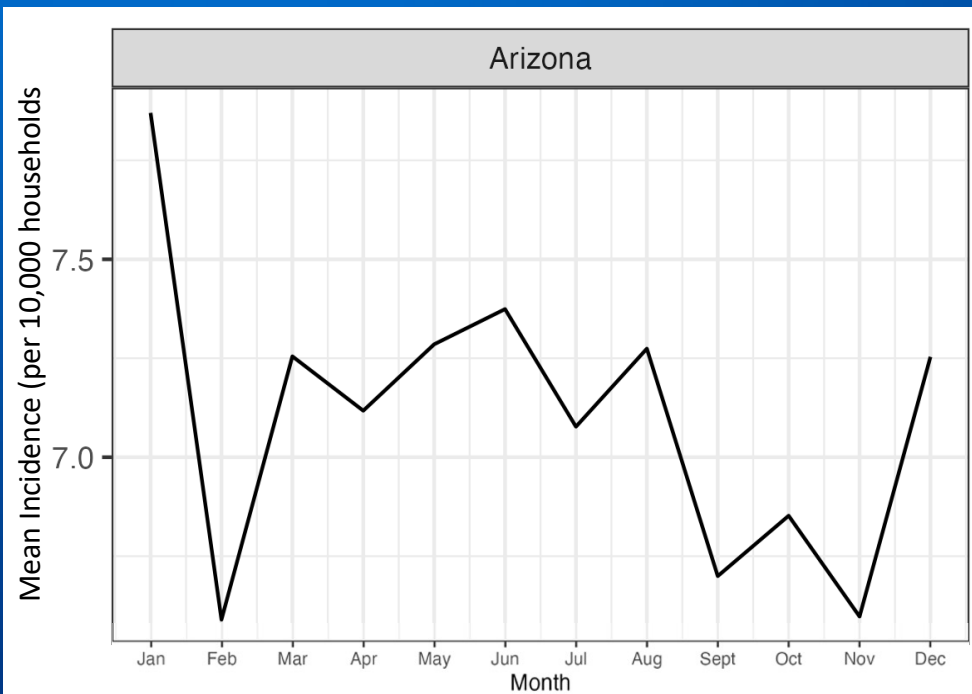
## Seasonal Incidence Curves, 2012-2023



AZ and CA shown with interquartile ranges  
Wide variation in annual incidence

# Results

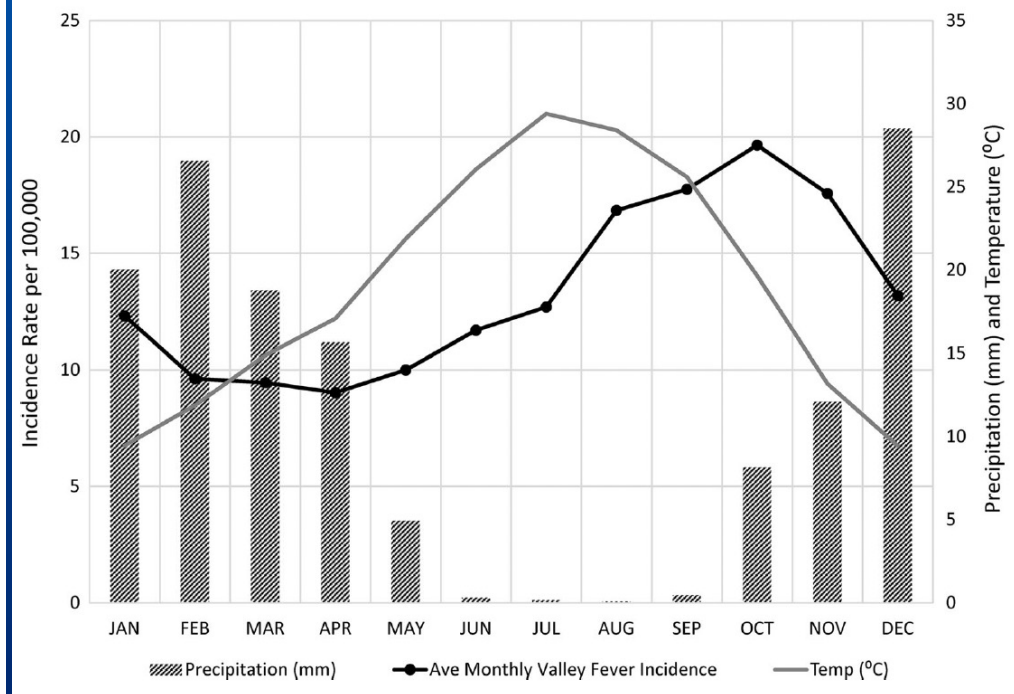
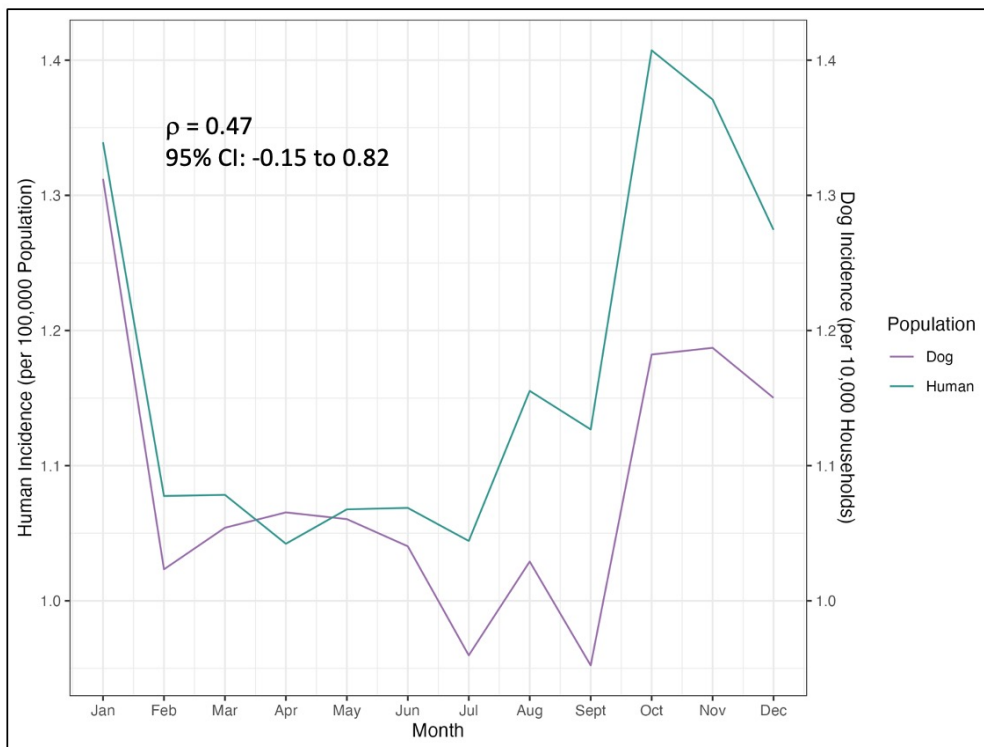
## Seasonal Incidence Curves Dog vs Human, AZ



Comrie et al. Environ Health Perspect 2005;113:688-692 (ref 5)

# Results

## Seasonal Incidence Curves Dog vs Human, AZ



Weaver and Kolivras. Ecohealth 2018;15:840-852 (ref 6)



# Conclusions

- Capture and analysis of almost all serologic test results
- High overall prevalence of seroreactivity
- Test numbers and % counties testing increased between 2012 and 2023
- Signalment risk factors
  - Medium and large dogs
  - Small breed protective, except terriers
  - Specific breed predispositions
  - Age 1 to 8 years
  - Intact female status protective
  - Highlight value of dogs as model





# Conclusions

- Dog incidence and spatial data correlated with human data
  - Including hot spots in CA, TX, AZ, NM, NV
- Incidence in AZ at least 80 to 100-fold greater than other states
- Possible fungal presence in new regions in OR, WA, ID, MT, CO, TX
- Increased incidence in existing regions
  - Climate niche model may be an underestimate
- Monthly incidence curves followed precipitation patterns

# Limitations

- Dogs are not humans
- Dog travel
- All cases not captured
  - Especially 2012 and 2023
- Spatial and temporal variations in dog ownership
  - Imperfect denominator
- Incubation period in dogs vs. humans?
  - ‘lag’ effect
- Variability in location granularity
- Inter-laboratory variation in test results
- Effect of increased testing
- Predictive value of positive test results





## Next Steps

- Further spatiotemporal analysis (R and ArcGIS Pro), lag model
  - Temperature, precipitation, drought cycling, soil pH, soil moisture, dust exposure
  - Land cover/land use
  - Zip code level studies
- Incorporation of AVMA pet ownership data
- Spatial interpolation studies
  - Predictive model incorporating climate and dog data
- Active surveillance efforts in new emerging regions
  - Non-traveled dogs



# MPH Applied Practice Experience Foundational Competency Attainment

| MPH Foundational Competency |                                                                                                                     | Description                                                                                                                                                                                                                                                                                                                                           |
|-----------------------------|---------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| 2                           | Select quantitative and qualitative data collection methods appropriate for a given public health context           | Gathered signalment information and serologic test data for 10-year period with sufficient spatial and seasonal granularity to identify how dog data might inform public health decision <a href="#">making</a>                                                                                                                                       |
| 3                           | Analyze quantitative and qualitative data using biostatistics, informatics, computer-based programming and software | Used Excel, <del>Graphpad</del> Prism, and ArcGIS Pro to perform statistics and map distribution of <i>Coccidioides</i> <del>seroreactivity</del> in dogs. Analyzed spatial and temporal distribution of <i>Coccidioides</i> exposure in dogs and made comparisons to findings in <a href="#">humans</a>                                              |
| 4                           | Interpret results of data analysis for public health research, policy, or practice                                  | Interpreted results considering previous studies on risk factors and predicted spread of coccidioidomycosis. Made conclusions regarding how results could influence future modelling strategies, public health policy (reporting procedures), and public health practice (increased awareness/education, environmental controls and PPE, vaccination) |
| 19                          | Communicate audience-appropriate public health content, in writing and through oral presentation                    | Wrote abstract, draft manuscript, ILE report; delivered oral presentations to communicate study findings and implications to public health professionals, academics, and industry members                                                                                                                                                             |
| 21                          | Perform effectively on interprofessional teams                                                                      | Engaged a multidisciplinary collaborative team (epidemiologists, veterinarians, physicians, public health professionals, statisticians, geographers, laboratory diagnosticians) to gather and analyze data using state-of-the-art technology                                                                                                          |



# MPH Applied Practice Experience Emphasis Area Competency Attainment

| MPH Emphasis Area Competency |                                       | Description                                                                                                                                                                                                                                                                                       |
|------------------------------|---------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| 1                            | Pathogens/pathogenic mechanisms       | In-depth understanding of causative agents of coccidioidomycosis and pathogenesis of infection                                                                                                                                                                                                    |
| 2                            | Host response to pathogens/immunology | Understanding of host factors that influence susceptibility to coccidioidomycosis, especially hormonal, genetic, and immunologic (e.g., HIV, transplantation, age); influences of host immunity on serologic test results; <u>current status</u> of vaccine development                           |
| 3                            | Environmental/ecological influences   | Understanding of “grow and blow” hypothesis, effects of climate variables and dust exposure on incidence, rodent activity, animal movement                                                                                                                                                        |
| 4                            | Disease surveillance                  | Understanding of disease risk factors (occupational/racial, sex, age, immune status), current surveillance methods at state and national levels and limitations of surveillance, made recommendations for reporting of dog serologic data                                                         |
| 5                            | Disease vectors                       | Understanding of the role of rodents as disseminators of <i>Coccidioides</i> spp., association between environmental pollutants such as fine particulate matter and disease outbreaks, and influence of immunosuppressive and antineoplastic drugs on immune response to <i>Coccidioides</i> spp. |

# Acknowledgements

## Laboratories that provided anonymized data, including

Carlos Rodriguez at Texas A&M VMDL  
Arizona Veterinary Diagnostic Laboratory  
Wade Burton and David Denton, IDEXX Laboratories  
Antech Laboratories  
Kelly Crucillo and Allen Bryan at UC Davis Cocci Serology Laboratory  
Protatek Laboratory  
MiraVista Diagnostic Laboratory

**CDPH:** Gail Sondermeyer-Cooksey

**UC Berkeley:** Simon Campo, Amanda Weaver, Jennifer Head, Justin Remais

**Kansas State University:** Ellyn Mulcahy, Hélène Avocat, Zach Mills, Christopher Carter, Amelia Brady, Rebecca Burks

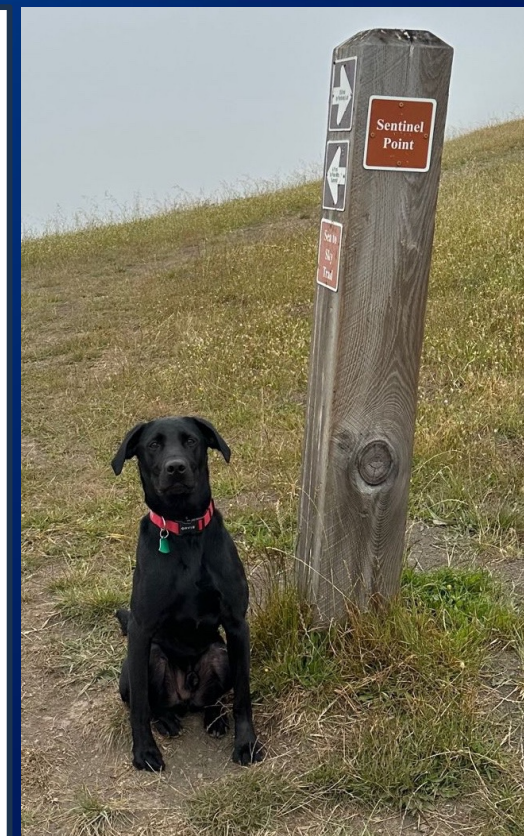
**UC Davis:** GR Thompson, Beatriz Martinez-Lopez, Kristen Aiemjoy



Berkeley Public Health

KANSAS STATE  
UNIVERSITY

UC DAVIS  
UNIVERSITY OF CALIFORNIA



# Questions





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15. Cordeiro R, Moura S, Castelo-Branco D, et al. Coccidioidomycosis in Brazil: historical challenges of a neglected disease. *J Fungi* 2021;7:85.
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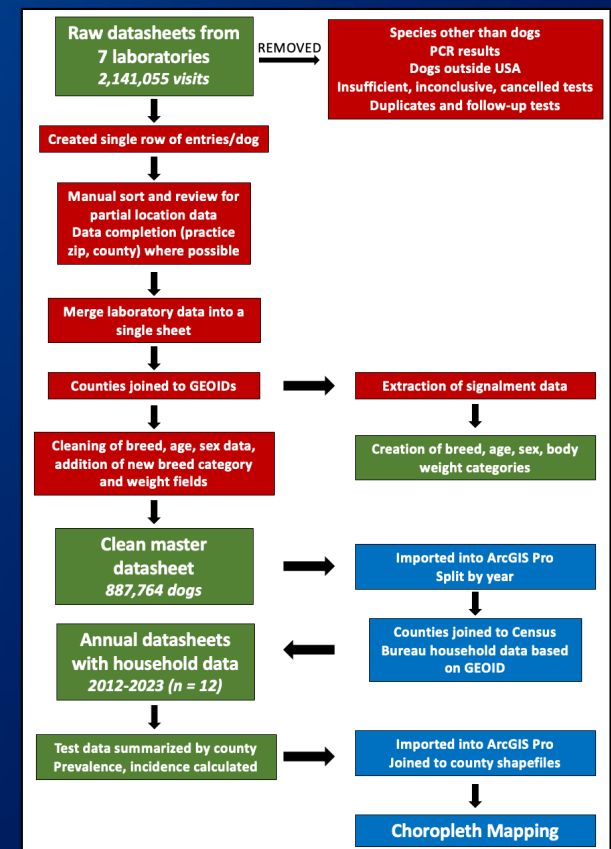
# MPH Applied Practice Experience Products

- Master and cleaned datasheets
- Over 40 ArcGIS Pro-generated maps
- Test positivity and incidence curves
- Conference abstract
- Applied Practice Experience and Integrated Learning Experience reports
- Paper for submission to Lancet Planetary Health (in preparation)
- Oral presentations
  - Remais-Barker collaborative research group (July 11 2023, one hour)
  - Anivive Life Sciences and University of Arizona Center for Valley Fever (August 21 2023, 30 minutes)
  - American Veterinary Medical Association leadership and data scientists (August 28 2023, 30 minutes)
  - Centers for Disease Control and Prevention endemic mycoses group quarterly meeting for state partners (October 24 2023, 48 nationwide public health participants)
  - Final defense presentation

# Methods

## Statistical Analysis

- Inter-annual and monthly/seasonal incidence
  - Compared with human data for AZ and CA
    - CA Department of Public Health (monthly county-level) and CA Department of Finance
    - AZ Department of Health Services (annual county-level; dog data aggregated)
    - Pearson's correlation coefficient (R)
- Spatial analysis (ArcGIS Pro, USA Albers Equal Area Conic)
  - Data joined with county shapefiles (US Census Bureau)
  - Choropleth mapping (tests, test positivity, incidence)
  - Spatial autocorrelation (Moran's I)
  - Hot spot analysis (Getis Ord-Gi\*, fixed distance band method)



# Results

## Behavior Group Breed Analysis

| Breed                       | Number positive | Number negative | Test Positivity | OR (95% CI)        | P value  |
|-----------------------------|-----------------|-----------------|-----------------|--------------------|----------|
| Pointers                    | 9432            | 8318            | 53.1%           | 1.91 (1.80 – 2.02) | < 0.0001 |
| Bull Type Terriers          | 3458            | 3272            | 51.4%           | 1.78 (1.66 – 1.90) | < 0.0001 |
| Parson Type Terriers        | 44              | 43              | 50.6%           | 1.72 (1.12 – 2.64) | 0.01     |
| Golden Retrievers           | 12103           | 12115           | 50.0%           | 1.68 (1.59 – 1.77) | < 0.0001 |
| Primitive Sighthounds       | 2446            | 2698            | 47.6%           | 1.52 (1.42 – 1.64) | < 0.0001 |
| Shetland sheepdog           | 432             | 1999            | 17.8%           | 0.36 (0.32 – 0.41) | < 0.0001 |
| Japanese Chin               | 84              | 340             | 19.8%           | 0.41 (0.33 – 0.53) | < 0.0001 |
| German Spitz Related Breeds | 1861            | 6772            | 21.6%           | 0.46 (0.43 – 0.50) | < 0.0001 |
| Miniature Pinscher          | 1060            | 3354            | 24.0%           | 0.53 (0.49 – 0.58) | < 0.0001 |
| Teacup                      | 15453           | 46692           | 24.9%           | 0.56 (0.53 – 0.59) | < 0.0001 |

Reference group Jack Russell Terriers (2820/7562, prevalence 37.3%)



# Results

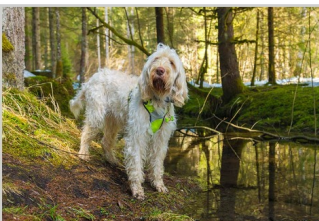
## AKC Breed Group Analysis

| Breed Group              | Number positive | Number negative | Test Positivity | OR (95% CI)        | P value  |
|--------------------------|-----------------|-----------------|-----------------|--------------------|----------|
| Foundation Stock Service | 1261            | 1536            | 45.1%           | 1.31 (1.22 – 1.41) | < 0.0001 |
| Hound                    | 16046           | 29931           | 34.9%           | 0.86 (0.84 – 0.88) | < 0.0001 |
| Miscellaneous            | 1852            | 228             | 34.7%           | 0.85 (0.68 – 1.06) | 0.14     |
| Mix Breed                | 40384           | 53543           | 43.0%           | 1.20 (1.18 – 1.23) | < 0.0001 |
| Non-Sporting             | 20204           | 40665           | 33.2%           | 0.79 (0.78 – 0.81) | < 0.0001 |
| Sporting                 | 53850           | 66193           | 44.9%           | 1.30 (1.28 – 1.32) | < 0.0001 |
| Terrier                  | 27265           | 38964           | 41.2%           | 1.12 (1.09 – 1.14) | < 0.0001 |
| Toy                      | 30461           | 93440           | 24.6%           | 0.52 (0.51 – 0.53) | < 0.0001 |
| Working                  | 36220           | 48352           | 42.8%           | 1.20 (1.17 – 1.22) | < 0.0001 |

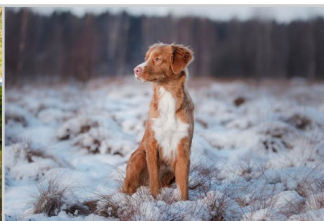
Reference group Herding group (32080/83291, prevalence 38.5%)



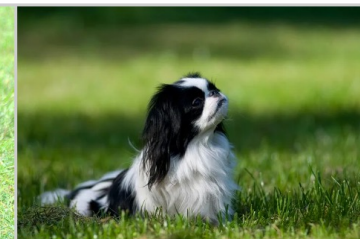
Spinone Italiano



Nova Scotia Duck Tolling Retriever



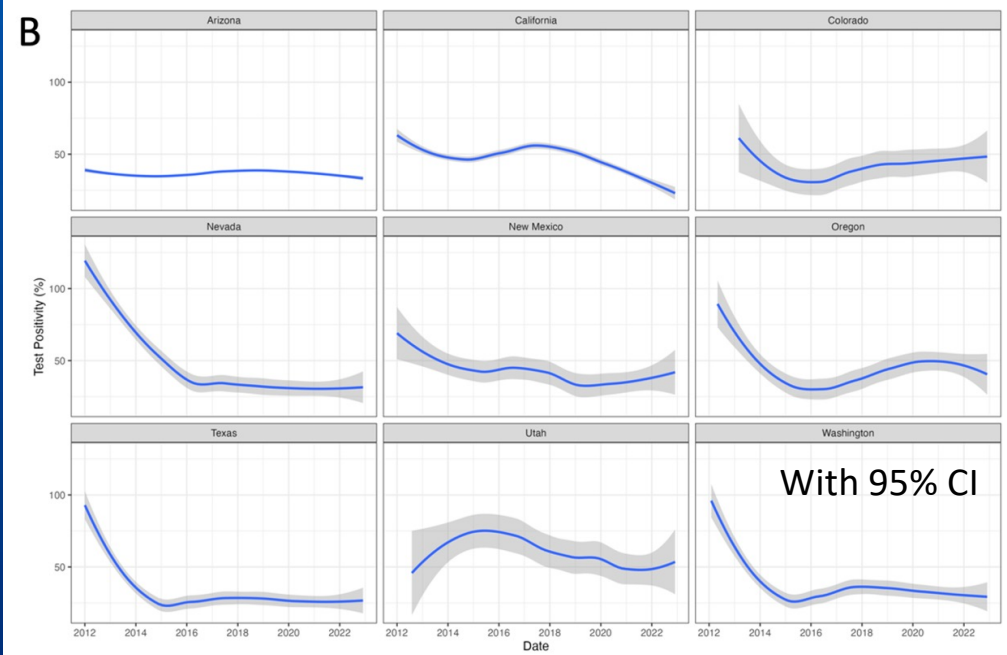
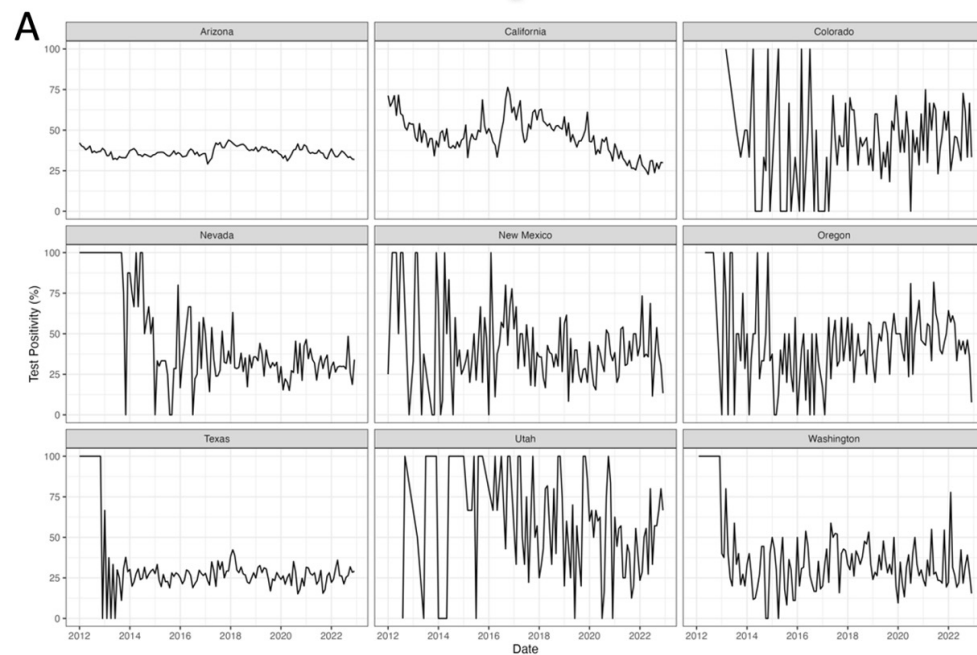
Affenpinscher



Japanese Chin

# Results

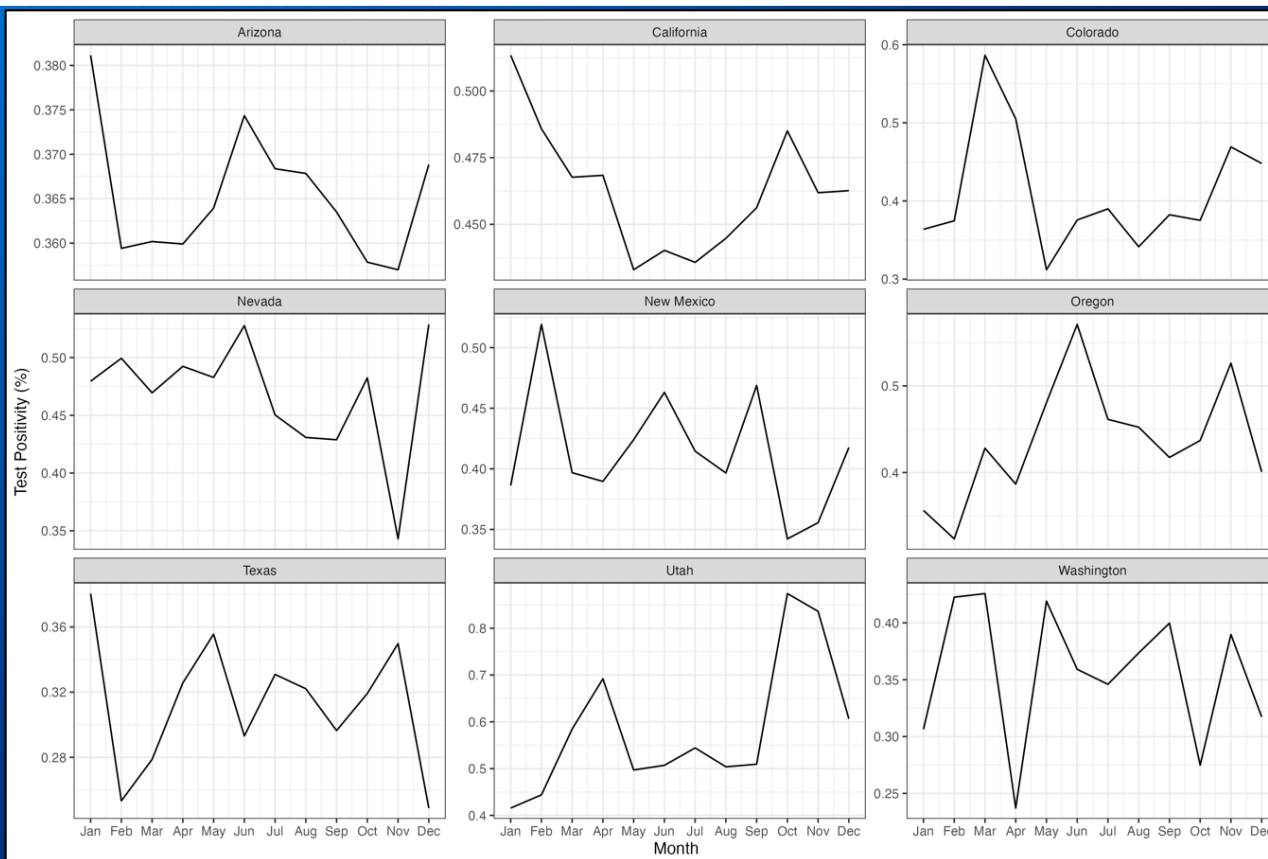
## Raw and Smoothed Test Positivity Curves, 2012-2023



Test positivity stable in AZ (mean 35%), declining in CA since 2018 (75% to 25%), mean 50%

# Results

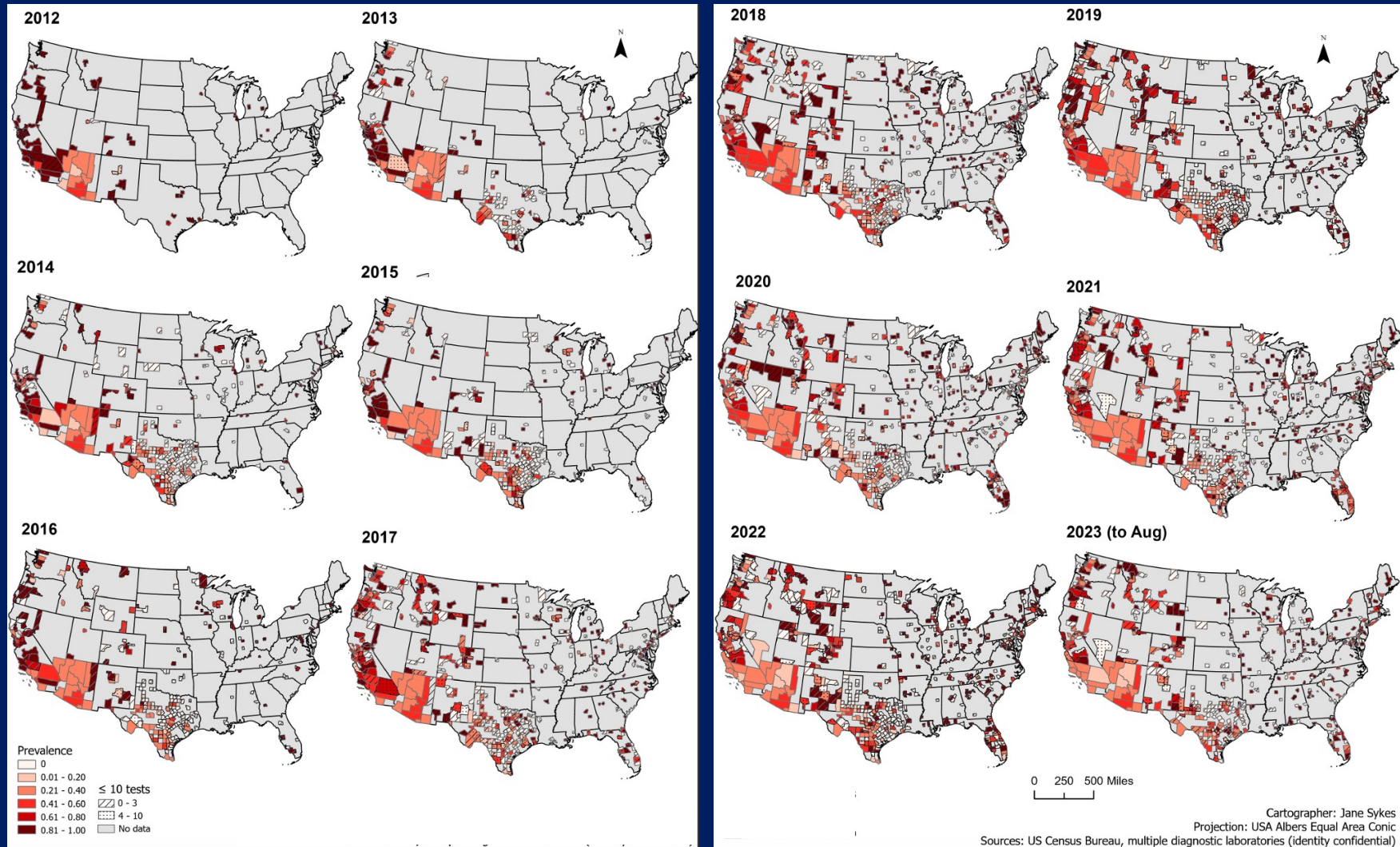
## Seasonal Test Positivity Curves, 2012-2023



# County Total Tests for Antibodies to *Coccidioides* spp., 2012-2023



# County Test Positivity of Seroreactivity to *Coccidioides* spp., 2012-2023



# County Incidence of Seroreactivity to *Coccidioides* spp., 2012-2023

