

An Overview of Alpha-Gal Syndrome

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Fall 2023 CLPC series

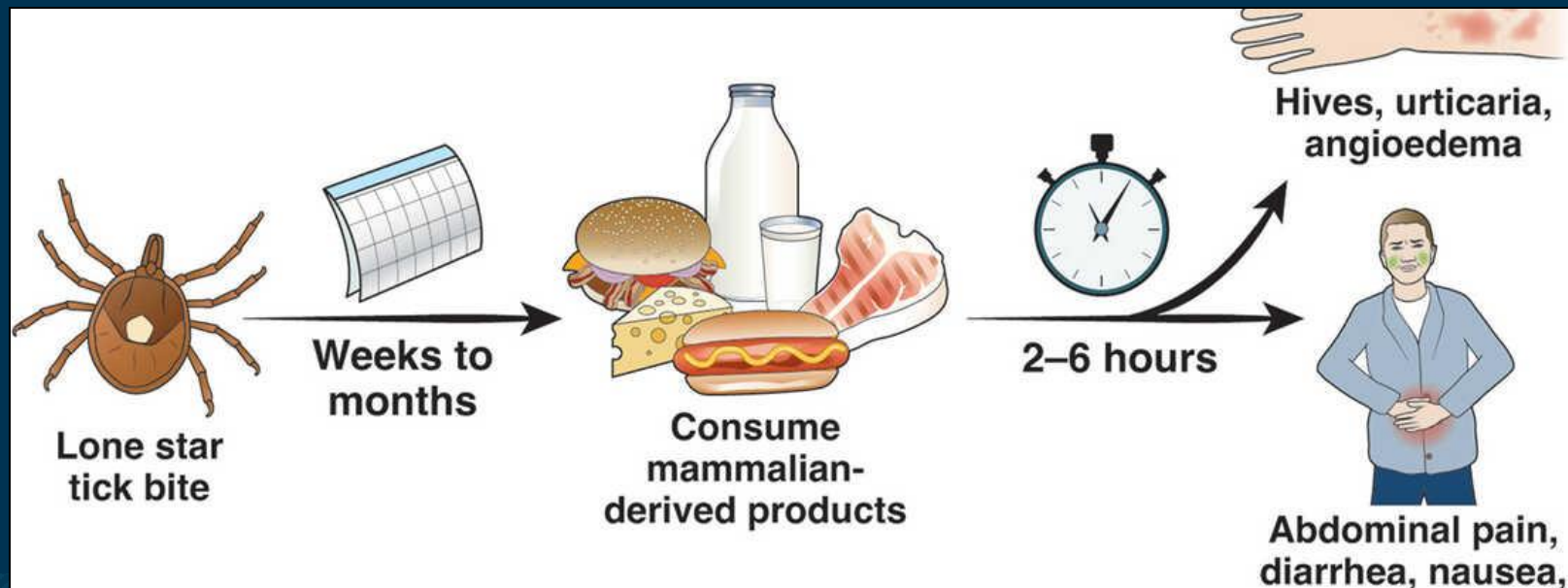
November 9, 2023

Learning Objectives

- Define/Describe Alpha-Gal Syndrome (α -Gal or AGS)/Mammalian Meat Allergy (MMA)/Red Meat Allergy (RMA)
- Describe pathophysiology of α -Gal Syndrome syndrome/MMA
- Describe laboratory testing, diagnostic approach, and treatment of α -Gal syndrome/MMA

<https://www.npr.org/sections/health-shots/2023/04/02/1166431106/ticks-ick-the-latest-science-on-the-red-meat-allergy-caused-by-some-tick-bites>

- CDC warns about red meat allergy caused by some tick bites
- Updated July 27, 2023 1:02 PM ET



CDC Press Release

For Immediate Release: Thursday, July 27, 2023

- Emerging Tick Bite-Associated Meat Allergy Potentially Affects Thousands
- Many healthcare providers not familiar with allergic condition
- 2 studies in Morbidity & Mortality Weekly Report (MMWR)

MMWR - the 2 studies

- Surveyed 1500 health care practitioners
 - Nearly half had not heard of AGS
 - 1/3 were not too confident in their ability to diagnose or manage patients with AGS
- Deidentified lab test results sIgE were examined from the commercial lab (Eurofins Viacor) which did all U.S. testing 2017-2021 for AGS
 - 300,000 samples
 - > 30% positive

Amblyomma Americanum -Lone Star Tick Family Ixodidae ('Hard Ticks')



“Questing”



Mouth parts of the lone star tick



<https://nigms.nih.gov/education/activities-and-multimedia/life-magnified>



Female Adult-stage Lone Star tick (*Amblyomma americanum*)

Unfed

Day 1.5

Day 2

Day 3

Day 4

Day 7

Day 9



**associated with STARI*
associated with red meat allergy

Blood engorged Female

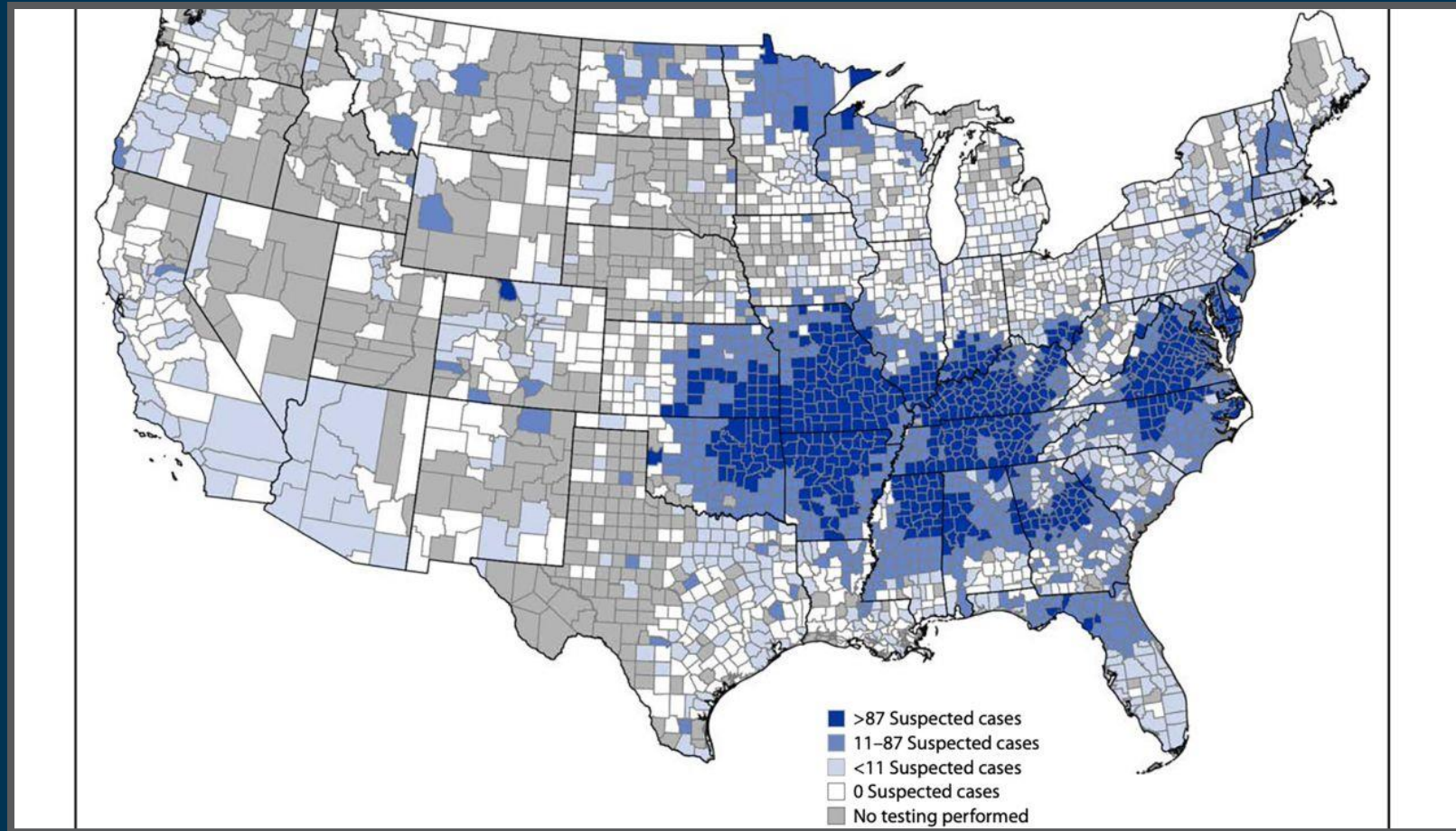
Female laying eggs



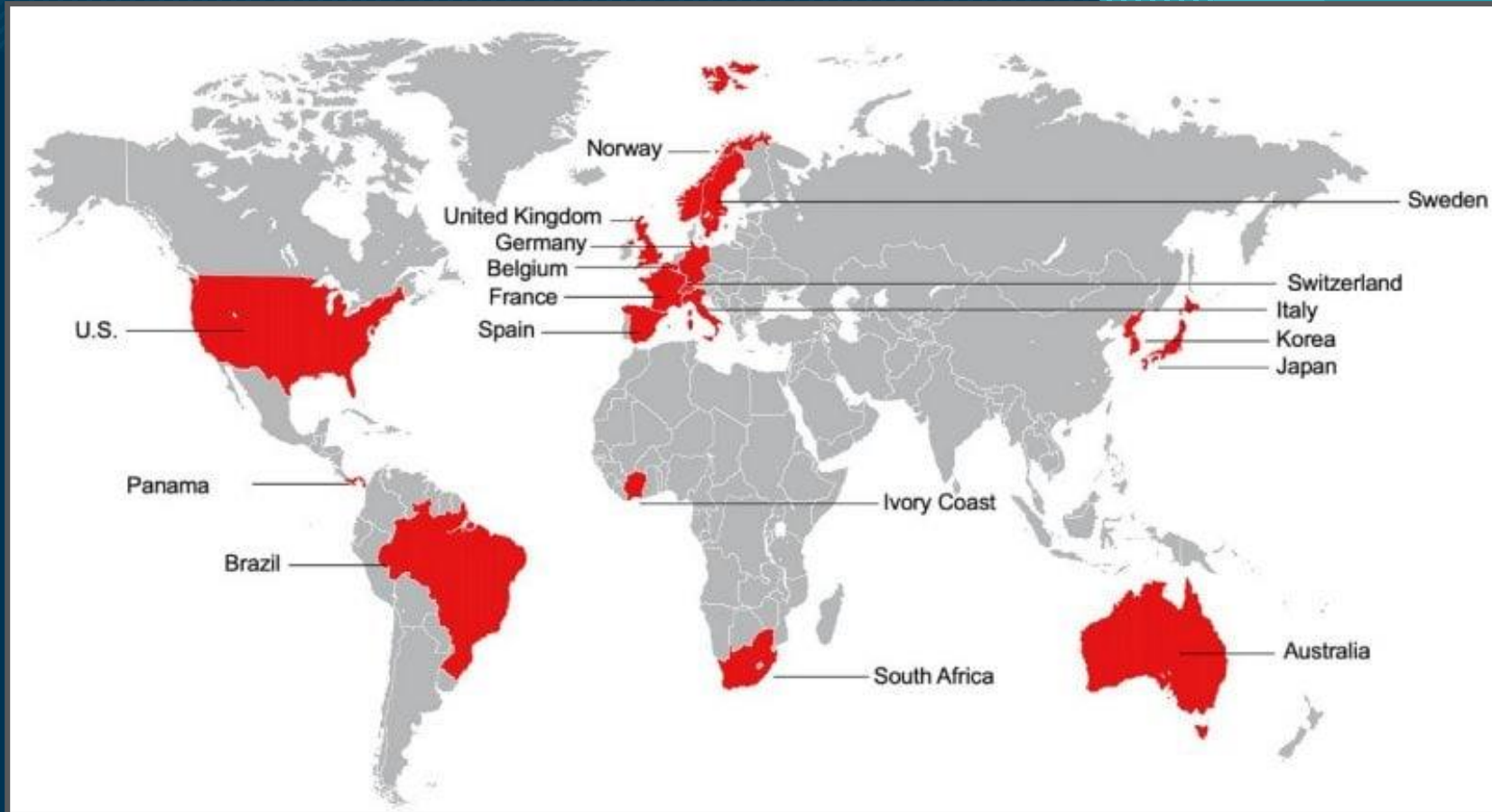
Distribution of Lone Star Tick



CDC map of cases/million 2017- 2022



World wide α -Gal Syndrome



- All continents except Antarctica

α -Gal syndrome tick associations

Scientific Name	Common Name(s) ^a	Geographic Range ^a
<i>Amblyomma americanum</i>	Lone Star Tick	North America (Southeastern US, Canada, Mexico)
<i>Amblyomma cajennense</i>	Cayenne Tick	North and Central America
<i>Amblyomma hebraeum</i> ^b	South African Bont Tick	South Africa
<i>Amblyomma sculptum</i>	N/A	South America (Brazil)
<i>Amblyomma testudinarium</i>	N/A	South Asia (India, Sri Lanka) and East Asia (including Japan)
<i>Amblyomma variegatum</i> ^b	Tropical Bont Tick	Southeast Asia, Africa
<i>Haemaphysalis longicornis</i>	Asian Longhorned Tick, Bush Tick	Japan
<i>Ixodes australiensis</i>	N/A	Australia
<i>Ixodes holocyclus</i>	Paralysis Tick	Australia, Southern Asia
<i>Ixodes nipponensis</i> ^b	Cattle Tick	Asia (including Korea, Japan)
<i>Ixodes ricinus</i>	Sheep Tick, Wood Tick, Deer Tick, Castor Bean Tick	North America, Europe and Northern Asia, Africa
<i>Ixodes scapularis</i>	Blacklegged Tick	Middle America, North America
<i>Rhipicephalus</i> spp (including <i>R. microplus</i> ; <i>R. evertsi</i>) ^b	Asian Blue Tick, Australian Cattle Tick, Southern Cattle Tick, Cuban Tick, Madagascar Blue Tick, Puerto Rican Tick	Southern Asia, South America, North America, Caribbean, Australia, Africa

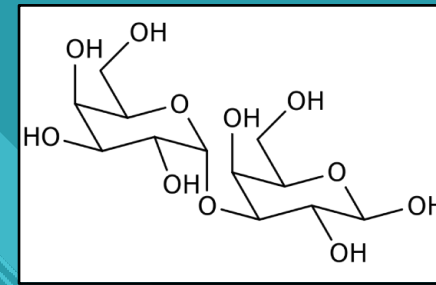
Notes: ^aCommon name and Geographic Range reported in Integrated Taxonomic Information System (ITIS, <https://www.itis.gov/>)¹³⁸ and/or reviewed in Carson AS, Gardner A, Iweala OI.¹⁹ ^bSuspected link, but not definitive association of tick with the development of alpha-gal syndrome.

Alpha Gal Syndrome - Not like usual tick diseases

- Most ticks contain bacteria, protozoans, or viruses that cause disease
- Other diseases caused by ticks:
 - Ehrlichiosis: (*Ehrlichia* sp.) Lone Star tick
 - Lyme Disease: (*Borrelia burgdorfi*) Black leg tick/ deer tick; US NE
 - Babesiosis: (*B. microti*) Black leg tick; NE US & upper Midwest
 - Rocky Mtn Spotted Fever: (*Rickettsia*) Dog tick; Lone Star tick, US, now in desert SW and Mexico
 - Powassan Virus Disease (*Flavivirus*) 3 Ixodes sp. NE and Great Lakes, Canada, Russia
 - Tick Borne Encephalitis (*Flavivirus*) Ixodes; Europe & UK, Far East, Siberia



What is Alpha Gal ?

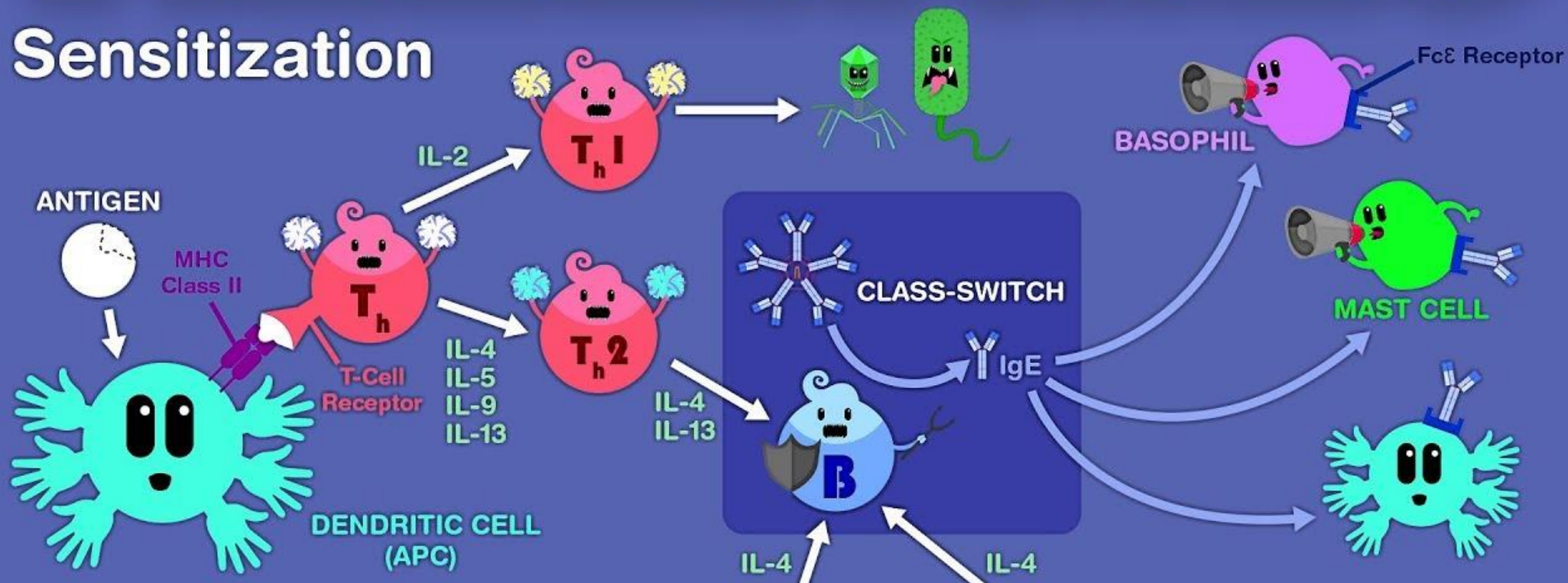


- A terminal sugar moiety linked to proteins or ceramide
- Found in most non-catarrhine mammals including cows, pigs, sheep, venison, rabbit, goat, bear, & in animal products
- NOT found in catarrhine - primate - mammals (humans, old world monkeys, apes) as well as birds, fish, reptiles
 - Mutation in the α -1,3-galactosyltransferase (α 1,3GT) gene
- Found in the saliva of ticks; more found when tick has fed
- **Alpha Gal Syndrome -Immunoglobulin E (IgE) mediated allergy to the disaccharide Alpha-gal (galactose- α -1,3-galactose)**
 - Allergic reactions to meats described in Australia 30 years before “discovery” in US

Th1 vs. Th2 responses.

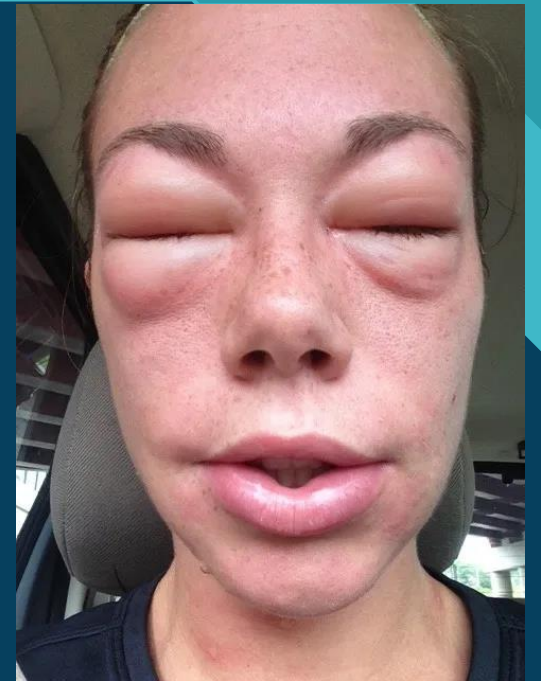
ALLERGIES

Sensitization



Definitions

- Allergic reaction – localized reaction, generally involves only one system such as the skin -Urticaria (hives) –swelling of the dermis
- Allergic angioedema –swelling underneath the skin (below the dermis) caused by fluid leakage into the surrounding skin and tissue. Most often occurs around the eyes, lips, tongue
- Anaphylaxis – a severe systemic allergic reaction involving two or more systems which may cause the collapse of circulatory &/or respiratory systems. (Due to massive release of inflammatory mediators; mediated by IgE)
- Anaphylactic shock –anaphylaxis plus hypotension and other signs of shock



Anaphylaxis



- Sense of 'impending doom'
- Acute onset of feeling unwell
- Change in demeanor (infants and toddlers)
- Cutaneous:
 - Flushing
 - Urticaria
 - Angioedema
 - Pruritus



- Nasal congestion
- Rhinorrhea
- Sneezing



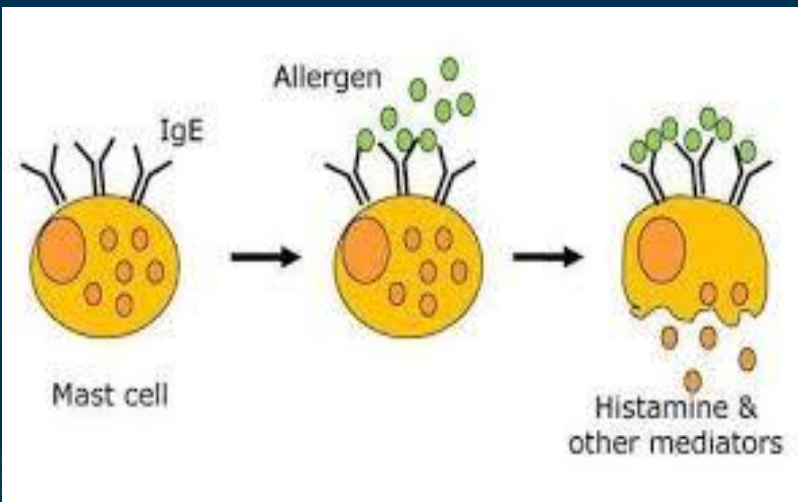
- Wheezing
- Persistent cough
- Chest tightness
- Respiratory distress



- Nausea
- Emesis
- Cramping
- Diarrhea



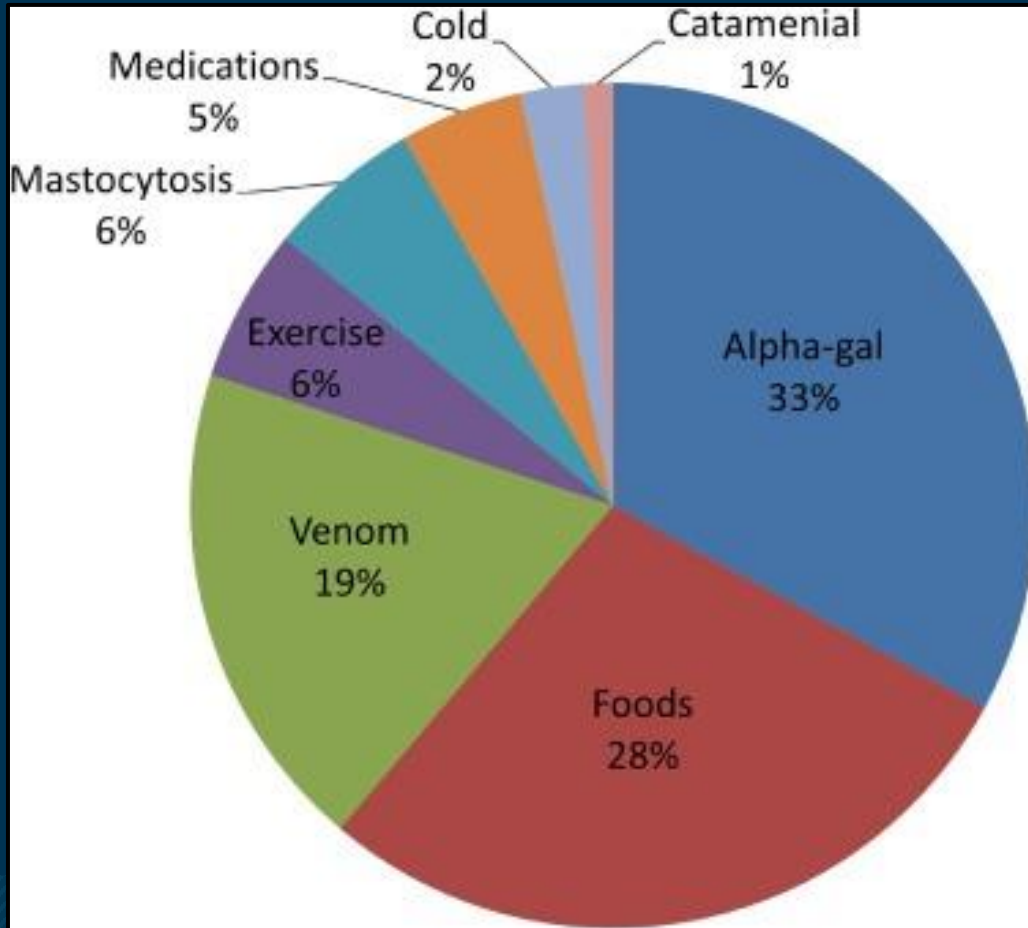
- Tachycardia
- Hypotension
- Syncope



Increasing cases of Anaphylaxis in US

Causes of Anaphylaxis, 2018

Definitive Cases



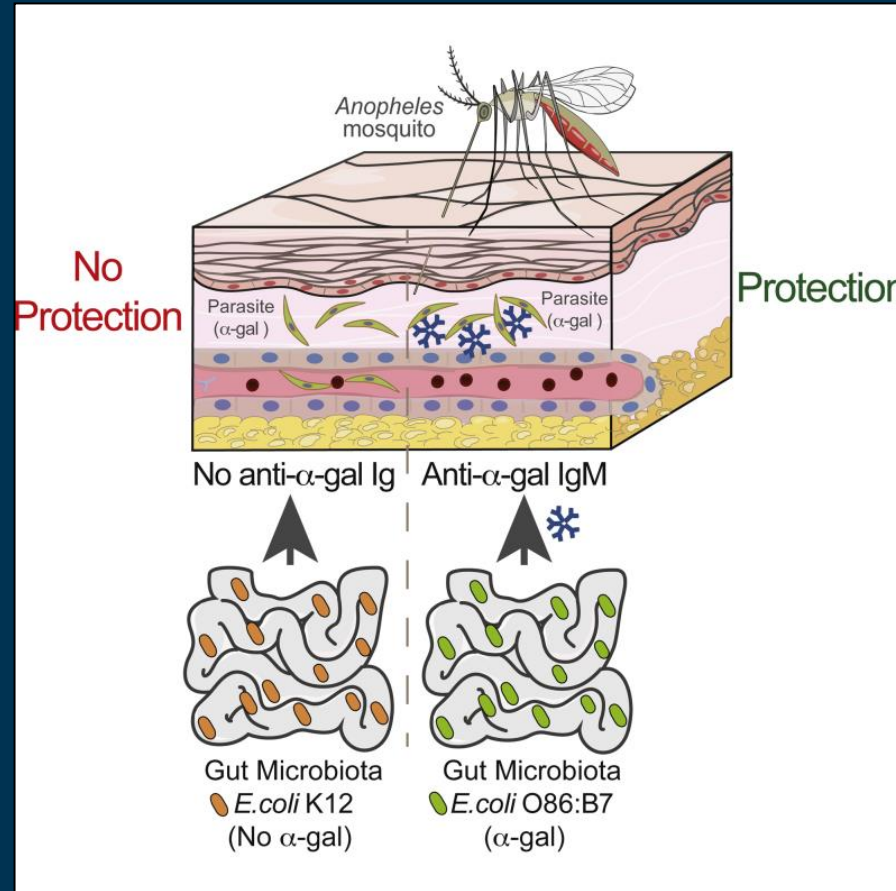
- U Tenn retrospective study 2006-2016
- 218 met NIAID & FAAN criteria for anaphylaxis
- **85 (39%) had “definitive cause”**; 57 (39%) were probable cause, and 76 (35%) were idiopathic
- AGS was found to be the number one trigger, accounting for 33% of cases with a definitive cause. The number two cause was all other food allergies combined at 28%
- Median age 42; 64% female
- Results differ from earlier retrospective studies
 - Idiopathic decreased to 35% (previously was 59%)

The changing face of anaphylaxis in adults and adolescents Annals of Allergy, Asthma & Immunology, 2018-11-01, Volume 121, Issue 5, Pages 594-597

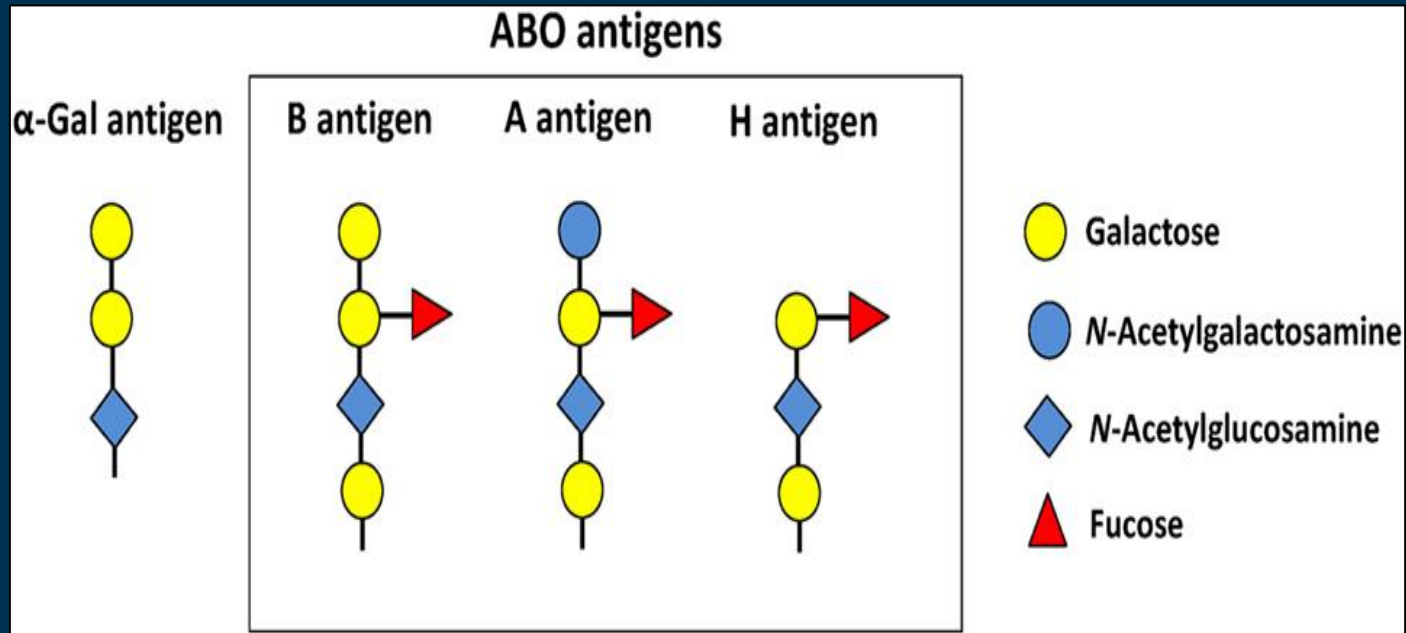
α -Gal

- All humans have IgG & IgM antibodies directed against α -Gal
 - Evolutionarily has thought to be protective against gut microbes and pathogens
 - Human pathogens (malaria, leishmaniasis, & Chagas disease)
 - Bacteria
 - Protozoa
 - Some fungi: *Aspergillus* sp. & *Schizosaccharomyces pombe* (but not in edible mushrooms)
 - Viruses incorporate alpha-gal into their envelope glycoproteins
 - Cobra venom
 - Amphibians

Hypothesis: Protection from malaria



α -Gal closely resembles the B group antigen structure



- Anti- α -Gal IgG antibodies can also recognize the blood group B antigen
- Blood group B antigen-expressing individuals (blood groups B and AB) have a lower level of α -Gal IgG antibody reactivity

The B antigen protects against the development of meat allergy

J Allergy Clin Immunol Pract. Sep-Oct 2018;6(5):1790-1791.e3.

- Patients expressing the B antigen of the ABO blood group system are protected from allergic sensitization to galactose- α -1,3-galactose and the development of red meat allergy.

TABLE I. Expected and observed frequencies of red meat allergy (RMA) according to ABO blood type or expression of the B antigen

Blood type	Control (n = 188)		RMA (n = 92)		<i>P</i> value
	Expected (%)	Observed (%)	Expected (%)	Observed (%)	
O	46.1	44.1	39.4	52.2	.212
A	37.1	38.3	40.4	43.5	.860
B	12.9	10.6	14.6	2.28	.026
AB	4.15	6.91	5.74	2.17	.126
O or A	83.2	82.4	79.8	95.7	.321
B or AB	17.0	17.6	20.3	4.35	.005

Expected frequencies are ethnicity-weighted according to population-based ABO distributions and representation of those populations in the control and RMA groups. *P* values are generated by χ^2 tests.

Transfusion risk?

An outbreak of anaphylactic transfusion reactions to group B plasma and platelets and its possible relationship to Alpha-Gal syndrome

- Case report : Three O+ patients receiving B type plasma or apheresis platelets emergently in two DC area hospitals From Nov 2022 – Feb 2023. All pts from S Maryland
- Chart review, pt interview (if poss), family interview; IgE specific to α -gal measured
 - Pt 1. Liver transplant, received B plasma, became worse, got other blood products, and **expired**. Serum tryptase elevated. IgE to α -Gal not done because were unaware of AGS
 - Pt. 2 Hx of cirrhosis, transferred with duodenal ulcer requiring transfusion, hx of 42 prior transfusions with allergic reactions to three. Received multiple transfusions. Another admission after collapsed after eating hamburgers for dinner. Got B plasma had severe reaction –responded to IV epi, solumedrol. **Hx tick bites**. Serum IgE to α -Gal elevated. **Survived**.
 - Pt. 3 - Found unconscious, Intracranial hemorrhage. Hx Crohn's Disease, and kidney transplant 6 months ago . 4 months prior had severe GI symptom & Increased ostomy output after eating beef/pork. Allergic to cats, dogs, horses, & grass. **Hx tick bites**. B apheresis platelets- Severe rxn requiring intubation. Elevated Serum IgE to α -Gal; got Klebsiella pneumonia; post transplant lymphoproliferative disorder; **expired** 25 days later

Australian Findings

- MMA after tick bite was first reported in 2007 in Australia by Dr Sheryl van Nunen et al.
 - Cases – 24/25 adults in Australia developed red meat allergy after reporting tick bites
 - She had a series of meat allergy cases dating back to the 80's
- Worldwide, Australia has the highest prevalence of MMA



Meanwhile... in Australia...

There are 1,275 animals that can kill you in this picture, and a further 800 that can't kill you but definitely want to.

imgflip.com

“Discovery” in U.S. Cetuximab

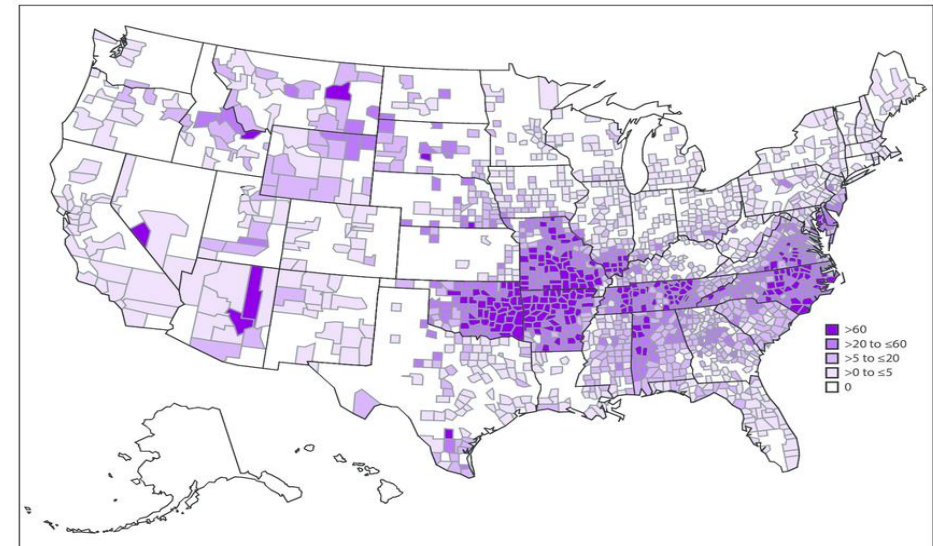
- Chimeric mouse-human IgG1 monoclonal antibody to EGFR
- Used for metastatic colorectal cancer and squamous cell carcinoma of head and neck
- **Caused death on first infusion in one patient in Arkansas**
- AR MD contacted her mentor @ UVa to help ID the cause. Formed a research team to investigate
- Cetuximab was causing severe urticarial or anaphylactic reactions in up to 15% of patients in the area: VA, NC, TN, AR, MO
- One of the team pointed out that these reactions to the drug overlapped the geographic area for incidence of Rocky Mtn Spotted Fever
- People who didn't get Cetuximab started coming to research clinic with symptoms and sometimes anaphylaxis several hours after eating meat. All had been bitten by ticks.
- 2008 NEJM identified alpha gal as the cause
- In 2009, the US team described patients with Alpha-Gal IgE who experienced delayed reactions after eating meat in a history of tick bites

States where anaphylactic reactions to Cituximab occurred compared to Rocky Mtn Spotted Fever



Incidence of Spotted Fever Rickettsiosis

Reported to CDC by County 2000 - 2013

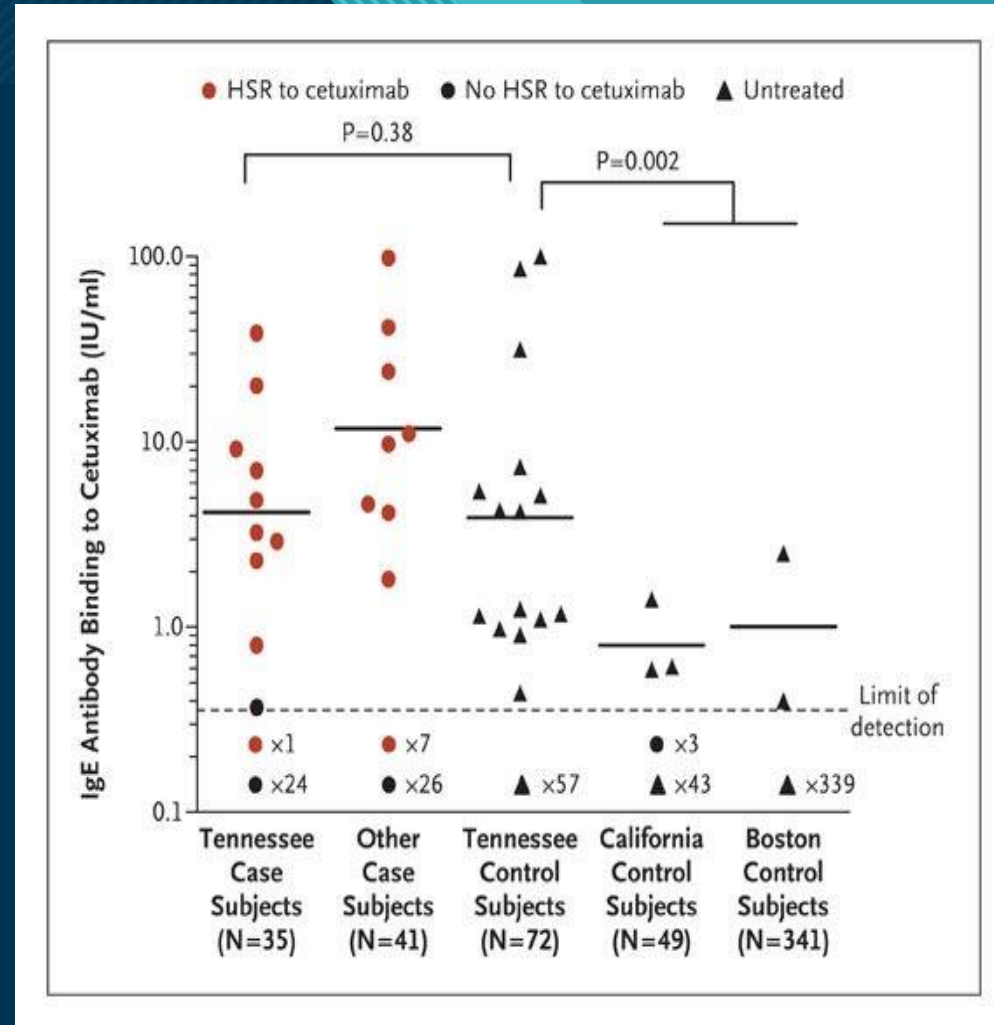


* As reported through national surveillance, per 1,000,000 persons per year. Cases are reported by county of residence, which is not always where the infection was acquired.

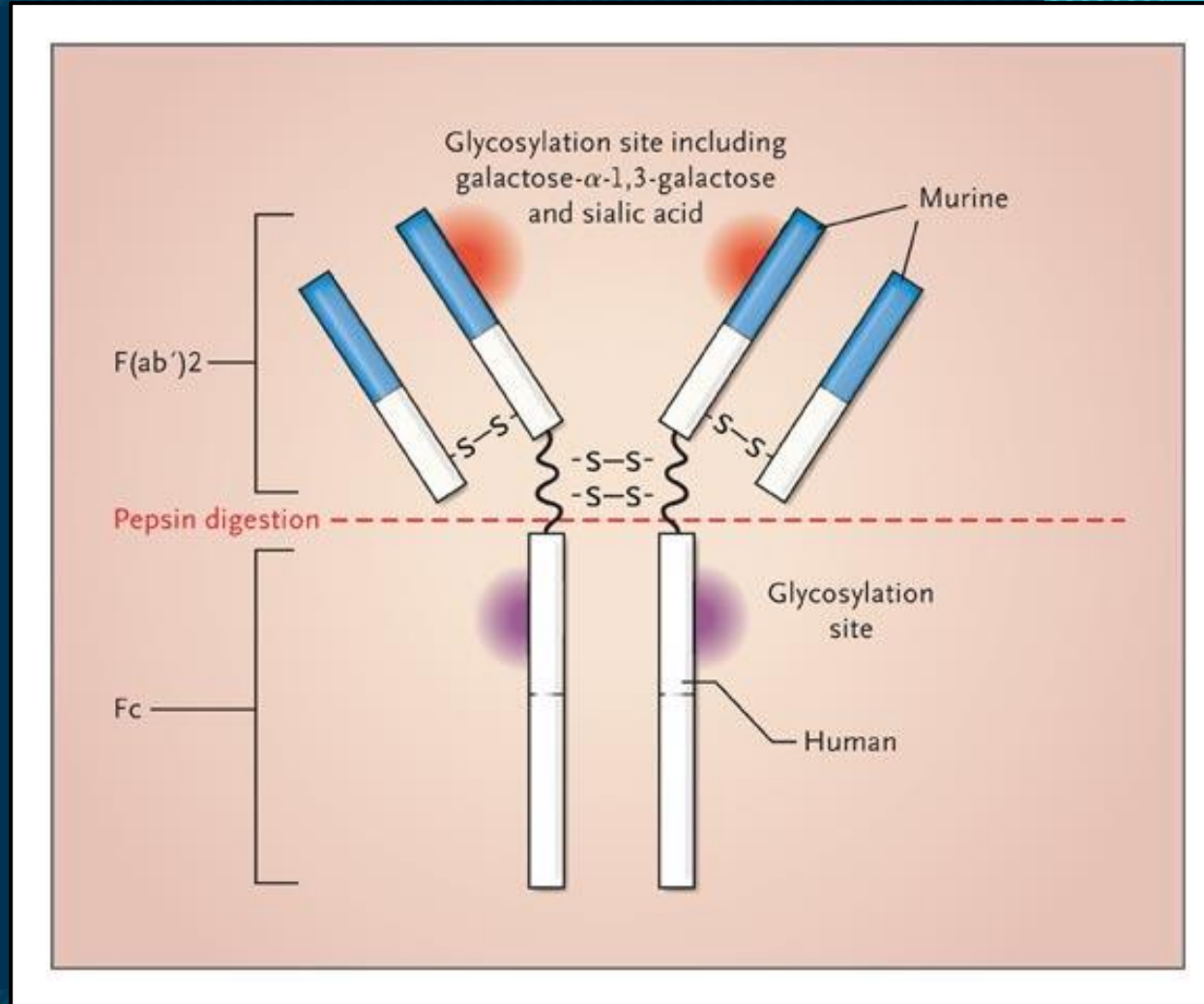
+ Includes Rocky Mountain spotted fever (RMSF) and other spotted fever group rickettsioses. In 2010, the name of the reporting category changed from RMSF to spotted fever rickettsiosis.

NEJM study - examined IgE Abs against Cetuximab

- Group 1 Pre-treatment samples -TN, AR, NC case subjects, 76 – who got the drug
- Group 2 Tennessee controls, 72
- Group 3 California H & N cancer pts, 49
- Group 4 Boston female controls, 341
- In most subjects who had a hypersensitivity reaction to cetuximab, **IgE antibodies against cetuximab were present in serum before therapy. The antibodies were specific for galactose- α -1,3-galactose**



Characterization of the epitopes on Cituximab



IgE-mediated food allergies

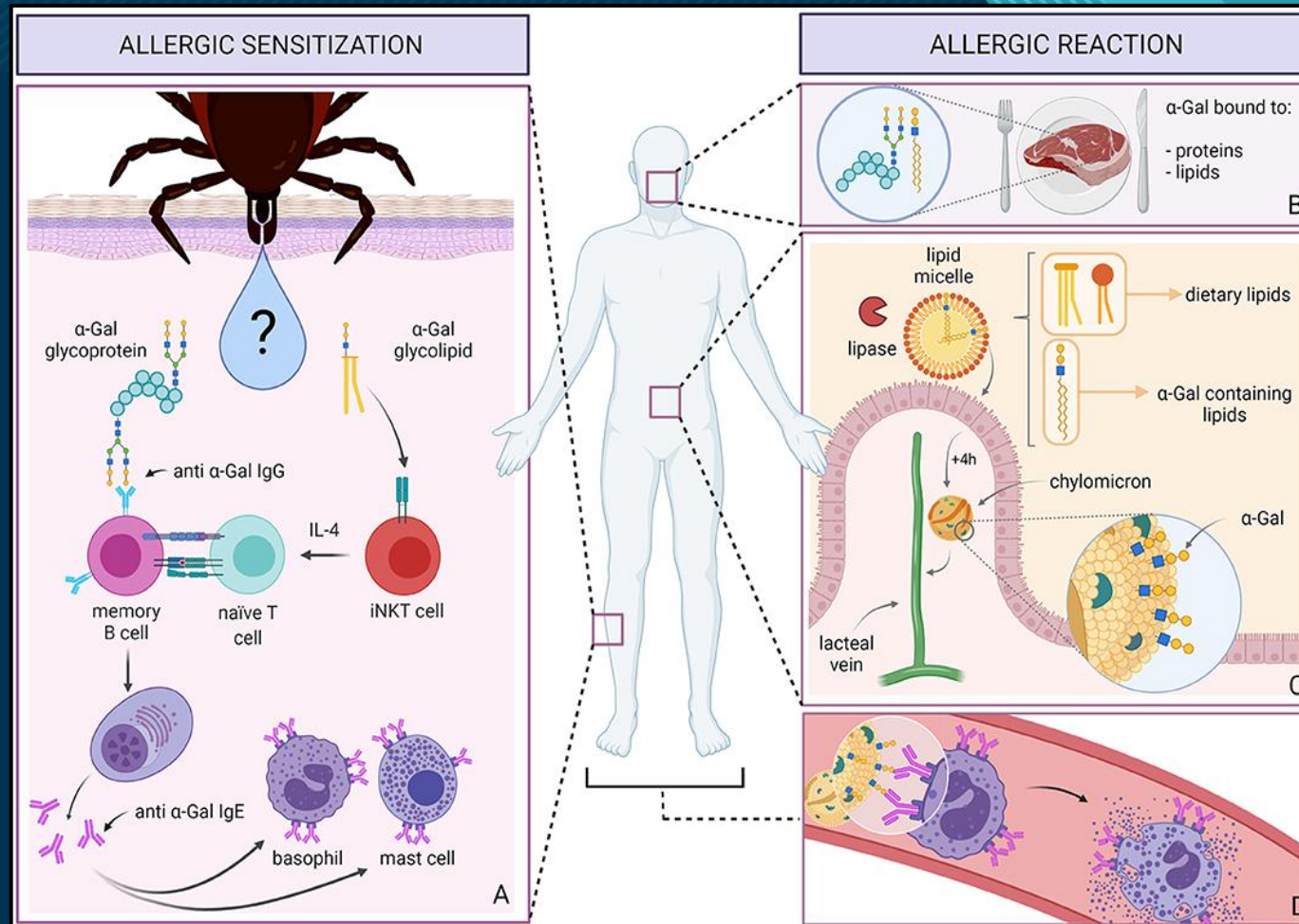
Conventional

- IgE Abs form against a protein antigen
- IgE Abs associated with allergen exposure
- **Immediate** hypersensitivity (anaphylaxis) is characteristic

AGS

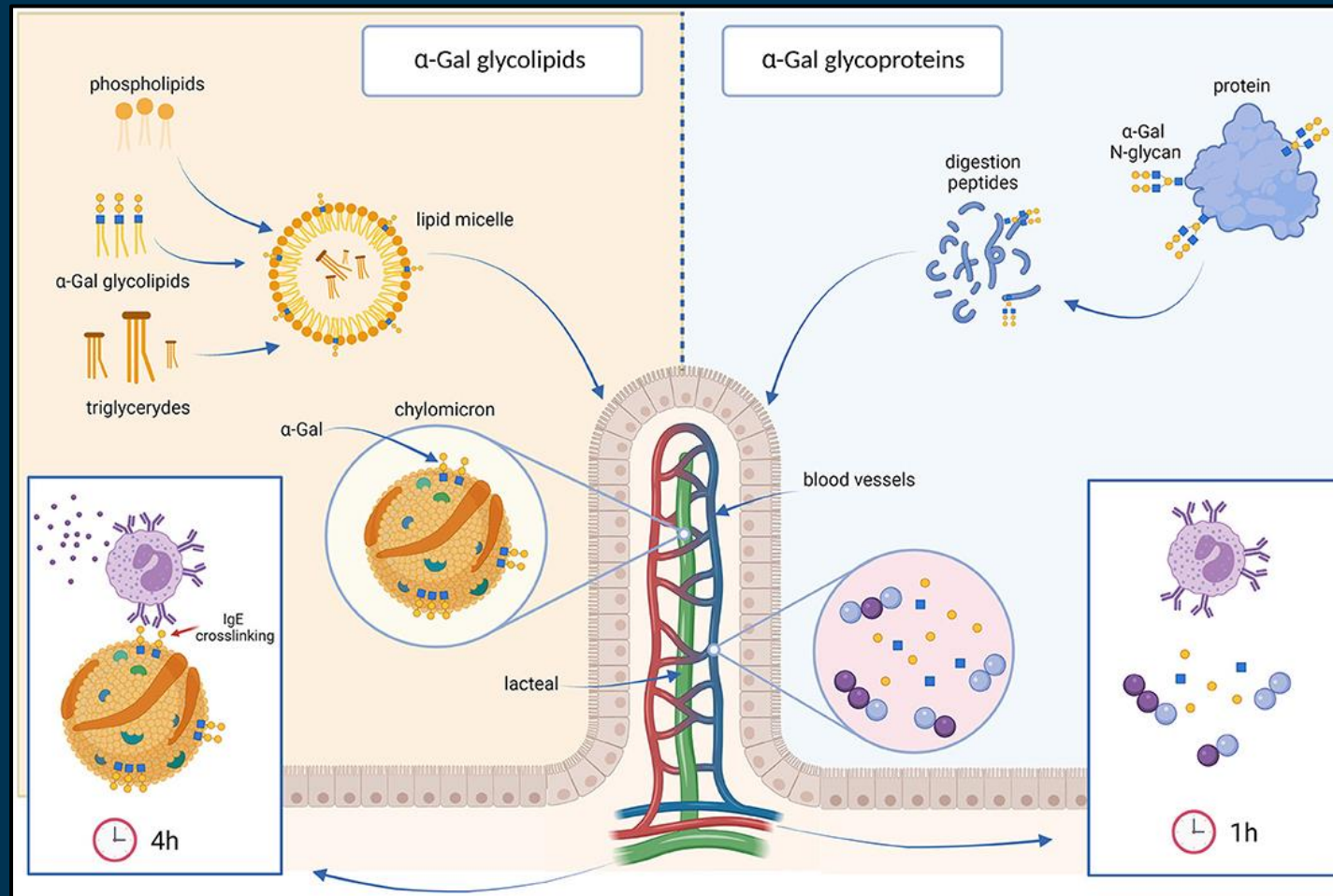
- IgE Abs form against the α -gal sugar
- IgE Abs associated with exposure to a **tick bite**
- **Delayed onset**; typically >2 hrs; generally 3-6 hrs after ingestion
- Immediate onset –after infusion with Cituximab

Allergic sensitization & allergic reaction to α -Gal

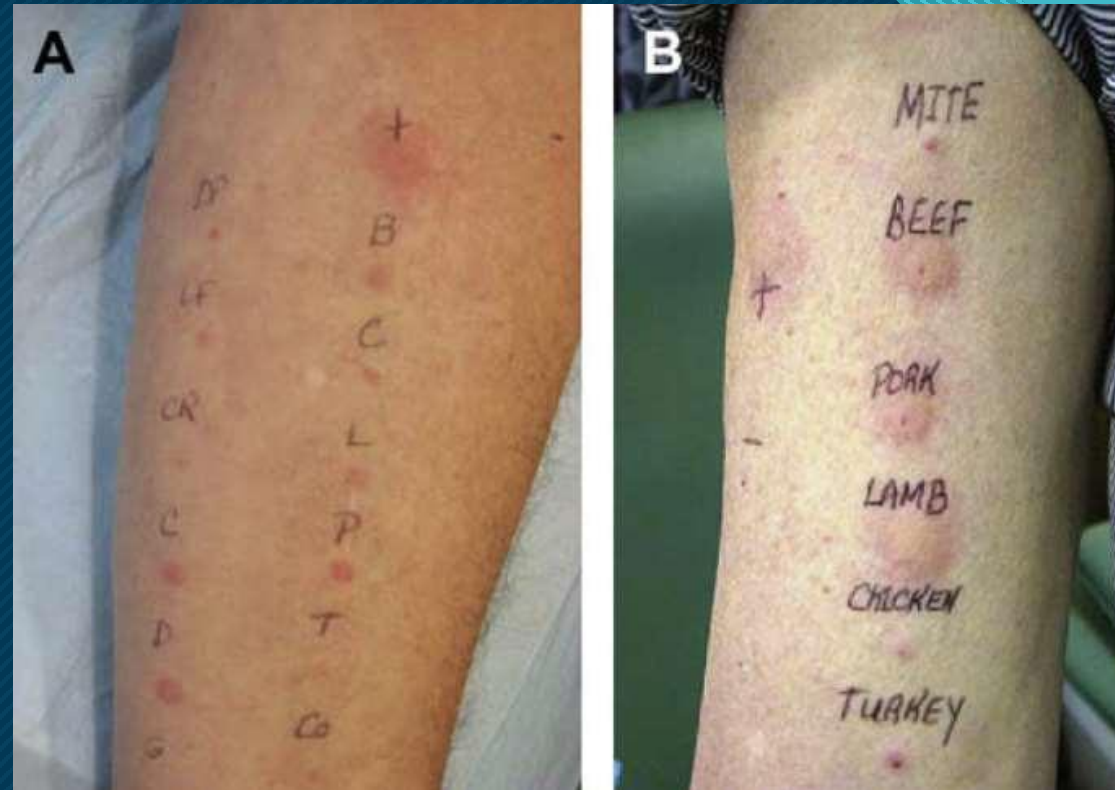


Romain-Carrasco P et al. 2021. The α -Gal Syndrome and Potential Mechanisms. Frontiers in Allergy. Vol 2 Article 783279

Why does the reaction take so long? α -Gal bound to Glycolipids “Glycolipid Hypothesis”



Skin Prick Test (SPT) and Intradermal (ID test) in patients with IgE antibodies to α -gal



Commins SP, et al. 2009 Delayed anaphylaxis, angioedema, or urticaria after consumption of red meat in patients with IgE antibodies specific for galactose- α -1,3-galactose J Allergy Clin Immunol 2009;123:426-33

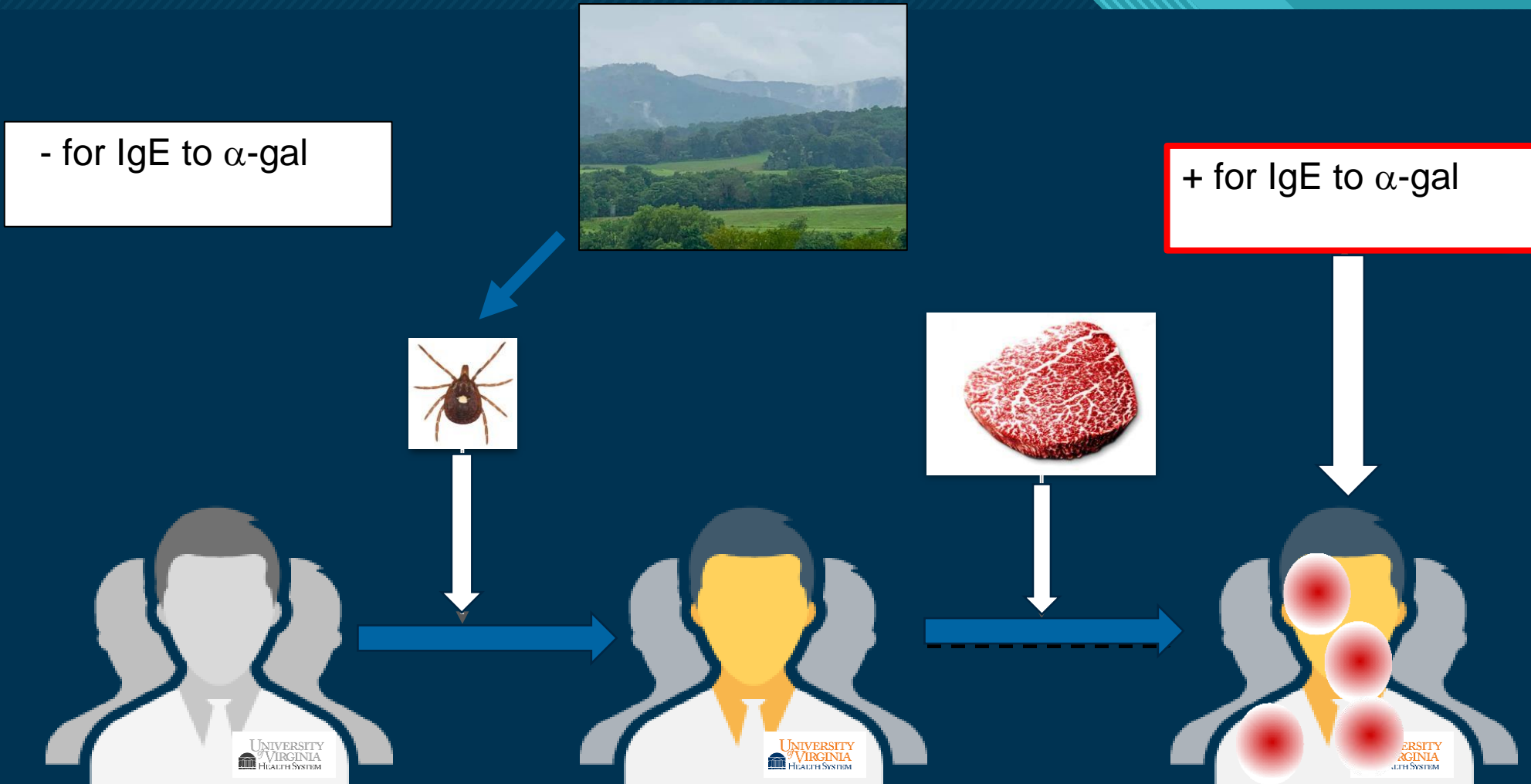
Results from expanded study

People who live in rural areas where tick bites are more common have a higher prevalence of IgE Abs to α -Gal than people who live in cities

TABLE II. Prevalence of IgE antibodies to alpha-gal in different populations related to local prevalence of tick bites

	Prevalence*	Percentage
Virginia clinic populations		
Patients presenting with delayed anaphylaxis to red meat ^{†9}	117/121 (99)	97
Asthma in the clinic [‡]	6/56 (3)	11
Random populations in the Southeast [§]		
Tennessee ⁵	24/107 (11)	22
North Carolina	15/75 (9)	20
Virginia (ER) ^{16,17}	25/136 (8)	18
Southeastern populations, total	64/318 (28)	20
Areas where tick bites are rare		
Boston Women's Study (adults) ¹⁸	2/341 (0)	<1
Norrbottn, Sweden (age 18 y) ¹⁹	1/150 (0)	<1
Northern California		
Stanford University cancer patients ⁵	3/49 (0)	6
Travis Air Force Base, asthmatic patients and control subjects ²⁰	1/117 (0)	1
University of California, San Francisco, asthmatic patients and control subjects	3/136 (1)	2
Northern California, total	7/302 (1)	2
Tropical areas where tick bites are common		
Kabati, Kenya (rural) ²¹	100/131 (50)	76
Thika, Kenya (moderately sized industrial town) ²¹	36/123 (10)	29
Esmeraldas Province, Ecuador ²²	110/295 (36)	37

“3 researchers developed red-meat allergy and recalled being bitten by a tick weeks-months before”



Alpha-gal syndrome (AGS) reactions can include:

- Rash
- Hives
- Nausea or vomiting
- Heartburn
- Severe stomach pain
- Diarrhea
- Difficulty breathing
- Drop in blood pressure
- Dizziness or faintness
- Recurrent angioedema
- Anaphylaxis
- **Symptoms commonly appear two to six hours after eating meat or exposure to products containing alpha-gal**
- Reactions can be different from person to person and can range from mild to severe or even life-threatening.
- People may not have an allergic reaction after every alpha-gal exposure.
- A person who thinks they may have AGS (MMA) should talk to a doctor (CDC recommendations, 2020)

Sometimes ONLY GI symptoms

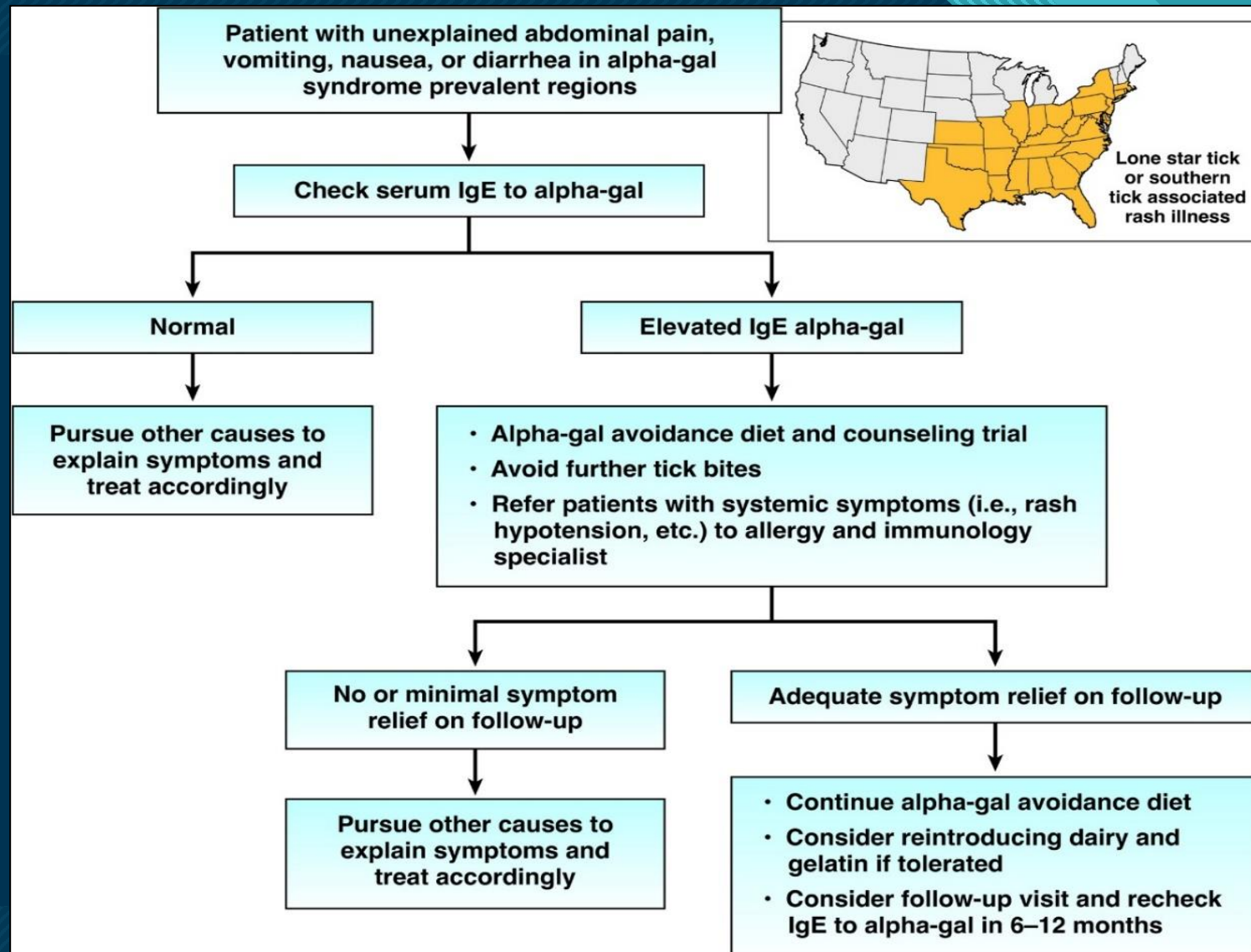
Modulating/amplifying co-factors

- Ingesting a greater amount of allergen (mammalian meat or meat product)
- The effect of cooking (slow cooking or reheating of the meat dish breaks down the connective tissue in meat, increasing alpha-gal availability)
- Offal consumption (e.g. sausage encasings, liver, kidney), because offal contains higher concentrations of the alpha-gal allergen
- Co-ingestion of alcohol
- Sleep deprivation
- Exercise, particularly within two hours
- Inclusion of spices, usually chilli and capsicum (but not nutmeg, black pepper or bay leaf)
- Prior administration of non-steroidal anti-inflammatory agents (within 24 hours)
- Being otherwise unwell/recent illness (e.g. upper respiratory tract infection)
- Being in the premenstrual/perimenstrual phase.

Alpha Gal Sensitization - other

- Tick bites; also Chiggers (?) -larvae in a species of mites, family Trombiculidae
- Allergy can occur in kids or adults
- Time course may be confounding
- # of tick bites may play role
- Unclear role for underlying atopy, conflicting studies

Diagnostic Approach



Pertinent History for Alpha-Gal Syndrome

- Timing
 - Night time symptoms are most common
 - Itching often the earliest symptom
- Ticks
 - Known tick bites or a history of outdoor activities in areas with ticks
 - Persistent irritation at site of tick bite
- Diet
 - Red meat, fatty meats, high fat dairy in diet
- If patient affirms ≥ 2 of these, consider testing for alpha-gal IgE

α -Gal IgE

- **Positive test result** Using the **cut-off of >0.1 IU/mL** (specificity of 92.3% and sensitivity of 100%)
- α -Gal IgE levels **higher than 2% of the total IgE levels**
 - α -Gal allergic individuals are frequently non-atopic individuals with low total IgE levels >> suggested to compare with total IgE levels
- α -Gal specific IgE declines over time (Possible reason for seronegative testing despite a history of symptoms)
- Additional tick bites lead to increases in α -Gal specific IgE

Other helpful tests

- Basophil Activation Test
 - Differentiate between α -Gal allergic patients and asymptomatic α -Gal sensitized patients
 - Higher basophil reactivity and sensitivity in patients with the α -Gal syndrome
- Tryptase – (IgE dependent MC activation) Serial measurement of total mast cell tryptase may help differentiate Anaphylaxis from other mimics

Oral food challenge

- Gold standard
- Risk of life-threatening anaphylactic reactions requiring multiple doses of epinephrine or emergency medical transport
- Typical challenge is 2 pork sausage patties prior to arrival in clinic



Problem - Health Care Team not aware of Alpha-Gal Syndrome

- Alpha Gal – not the same as alpha galactosidase
- Problem – coding test requests properly

2022 ICD-10-CM Diagnosis Code

Z91.014

Allergy to mammalian meats

The Wrong Test

- ~~Alpha-Galactosidase~~ — this is the wrong test!
- ~~α -Galactosidase A Deficiency~~ — this is the wrong test!
- The main reasons why this test is sometimes mistakenly ordered are:
 - Even though a-galactosidase *has nothing to do* with alpha-gal syndrome, the names sound similar (and abbreviations are the same!) Remember **Alpha-gal is (galactose- α -1,3-galactose)**
 - Doctors and laboratory staff have trouble finding the correct test codes

Updated 5/6/23

Quest and Labcorp alpha-gal IgE test codes

Quest Alpha-gal IgE Test Code: 10554

Labcorp Alpha-gal IgE Test Code: 650001

Updated 5/6/23

Quest and Labcorp Alpha-gal Panel test codes

Quest Alpha-gal Panel Test Code: 10555

Labcorp Alpha-gal Panel Test Code: 650003

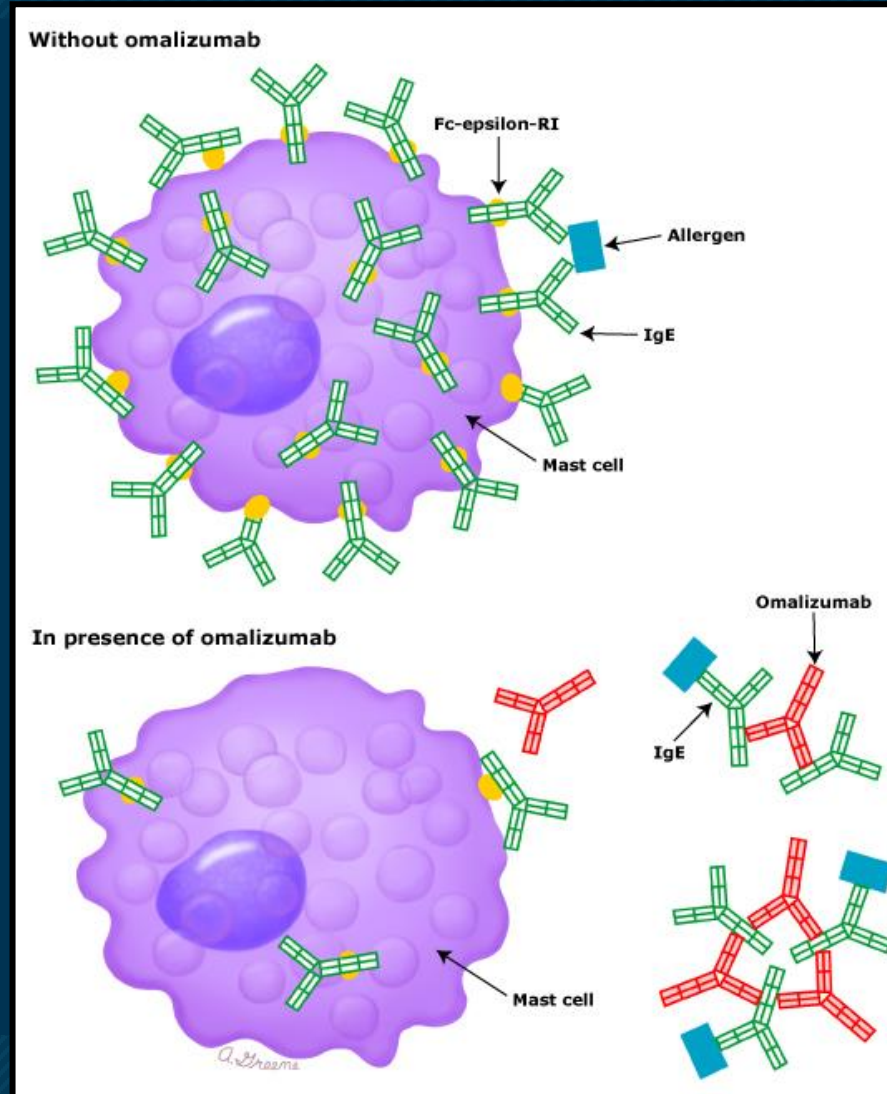
Note: experts rarely order the full panel, which is more expensive and not usually needed for diagnosis.

Approach to treatment

- Avoidance of meats, and other α -Gal containing products
- Avoidance of ticks
- Oral antihistamines
- Oral corticosteroids
- Oral Cromolyn (mast cell stabilizer)
- EpiPen
- Newer anti-IgE drug, Omalizumab



Omalizumab – binds free IgE



Tick bite prevention tips:

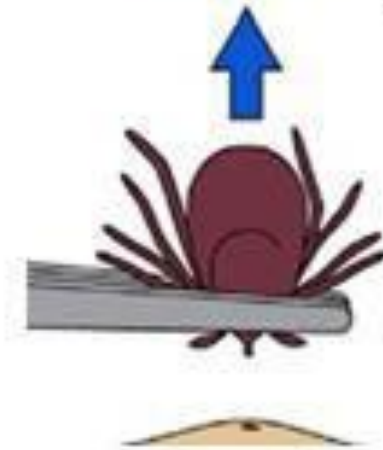
- When possible, avoid areas where ticks can be found, including tall grasses, bushes, shrubs and leaf litter
- Use a bug spray or repellent with 30% or more of DEET (can be used on skin)
- CDC recommends permethrin treatment of clothes, boots, gear. (Do not use on skin). Can buy pre-treated clothing (\$)
- Wear clothing that covers your arms and legs to give ticks fewer places to bite.
- Tuck pants into socks while hiking.
- Examine your children, pets and gear after an outdoor excursion.
- Check your scalp, behind your knees, belly button and around your body after being outdoors.

How to Remove a Tick

Step 1: Use fine-tipped tweezers to grasp the tick as close to the skin's surface as possible. The goal is to remove the entire tick including its head and mouth.



Step 2: Pull upward with steady, even pressure. Do not twist or jerk the tick!



Step 3: Clean the bite area and your hands with rubbing alcohol, an iodine scrub, or soap and water.

Other meat products

- Gelatins made from beef or pork collagen
 - Jello, gummy bears
 - Gelatin casings for medicines
 - Gummy vitamins/supplements
 - Tooth paste
 - Topical numbing agents for dental procedures
- Products containing mammalian fats, like lard, tallow or suet
- Broths, stocks, gravies and bouillons made with meat
- Snake anti-venoms produced in horse or sheep
- Drugs produced in/from animals – Heparin, Cituximab
- Red algae (carrageenan) -used as a thickener in many food products including milk substitutes



Hidden Gelatin/a-Gal

- Nuts –may use gelatin to make salt stick
- Marshmallows
- Altoids brand mints
- Kellogg's Frosted Pop Tarts and Frosted Shredded Mini Wheats
- Candy corn
- Starburst
- Ice cream
- Puddings
- Jiffy Corn Muffin Mix
- Airborne fumes from cooking (bacon; barbecue)



<https://spoonfulapp.com/>

Other products may contain a-Gal

- Lanolin in lipstick & lotions
- Clothing –wool, cashmere, alpaca, suede
- Animal feed
- Milk and butter
- Personal care items: lotions, soaps, detergents, deodorants, cosmetics, & cosmetic brushes
- Vaccines– Excipients for stabilization or during manufacture (CDC) - MMR and Zoster Zostavix
- Drugs manufactured in animals – Abatacept, Infliximab
- Pancreatic enzymes
- Heparin
- Some thyroid hormone preps
- Bioprosthetic heart valves

Biggest challenges

- Greater training for health care professionals
 - Is the medical team aware of alpha-gal syndrome?
 - Awareness of medicines that may contain meat products
- Drug labeling –mammalian products used in the manufacture of drugs
- Food labeling – hidden alpha gal
- Testing – send outs
- Awareness campaigns for the public
- Some people may become allergic to their pets
- **Psychological impact of alpha gal allergy: Long journey to diagnosis; considerable distress at continued allergic and anaphylactic reactions –patient information/advocacy websites have filled the gap**

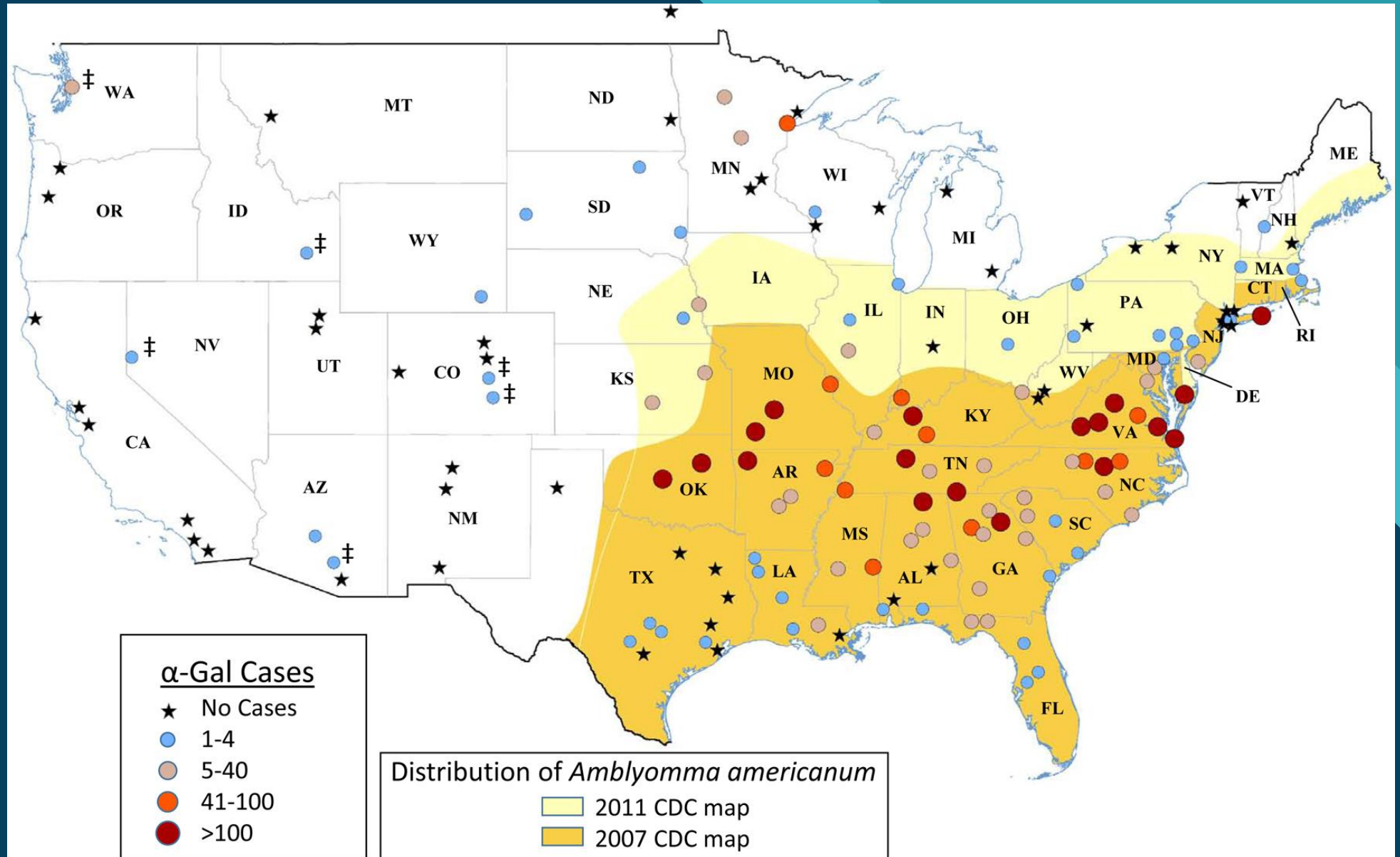
I live in La. and we are outdoors all the time. Should I be worried?



Pick Your Poison:
Fire Ants vs Lone Star Ticks

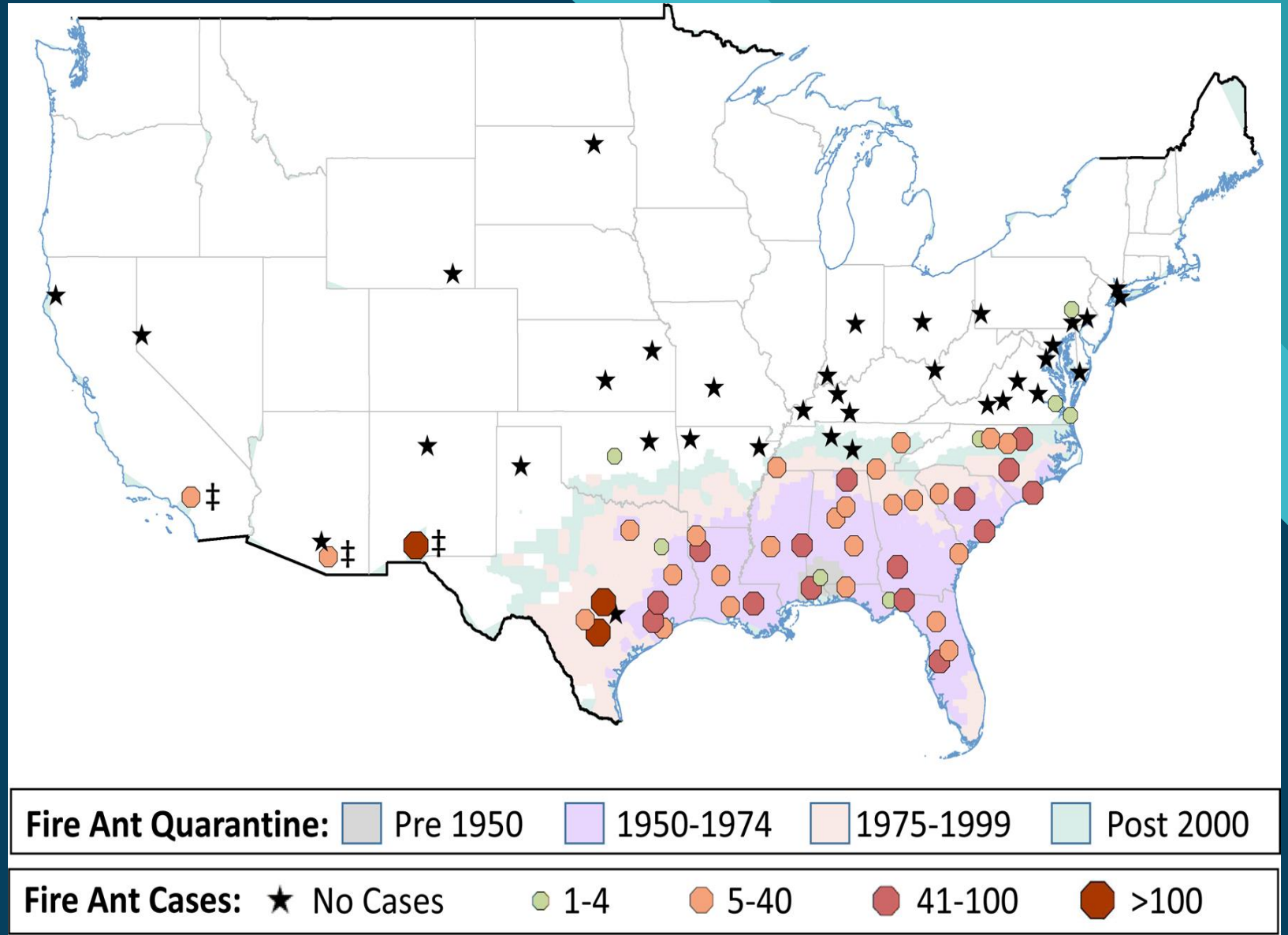


Cases of α -Gal syndrome identified by a survey of U.S. allergists



IFA(Imported Fire Ants) Quarantine Areas and Fire Ant Anaphylaxis cases from second survey

- Two types of IFA came to America in 1918 & 1930 through the port of Mobile, AL.
- Cargo boats coming from S. America used soil as ballast.
- Damage crops, agricultural equipment, impact wildlife, painful bites that can cause anaphylaxis.
- APHIS – Animal and Plant Health Inspection Service (USDA) – works to prevent spread of IFA by enforcing Federal Quarantine & works with states to regulate high risk commodities such as nursery stock, sod, hay, & soil moving equipment.



Delayed Anaphylaxis to Red Meat Questionnaire

Are you aware of the basic details of delayed anaphylaxis to red meat? Yes _ No ___ How many have you tested on the basis of history? ___ ; or

0 1-4 5-40 40-100 >100

Have you diagnosed cases in your practice? Yes _ No ___ If so, how many in the last 3 years? # ___ or

0 1-4 5-40 40-100 >100

Are you aware of any other practices in your region or state that are seeing cases? _____; contact information for those practices

If you have seen cases of delayed anaphylaxis to red meat:

a) How did you confirm cases: Skin tests _ Laboratory tests _ Approximately how many of the cases you tested because of a suggestive history have been negative for IgE to alpha-gal?

b) What is the range of times before start of reactions after eating mammalian products?

c) Do you question patients about: Tick bites _ Insect bites _ Chiggers

d) How common is a positive answer about tick or chigger bites? < 50% 50-70% >70%

In confirmed cases:

What % become free of attacks on a diet avoiding red meat? <50% 50- 70% 70-90% >90%

If symptoms persist, do you recommend: Dairy Free diet ___ or other treatment

Experience IFA allergy:

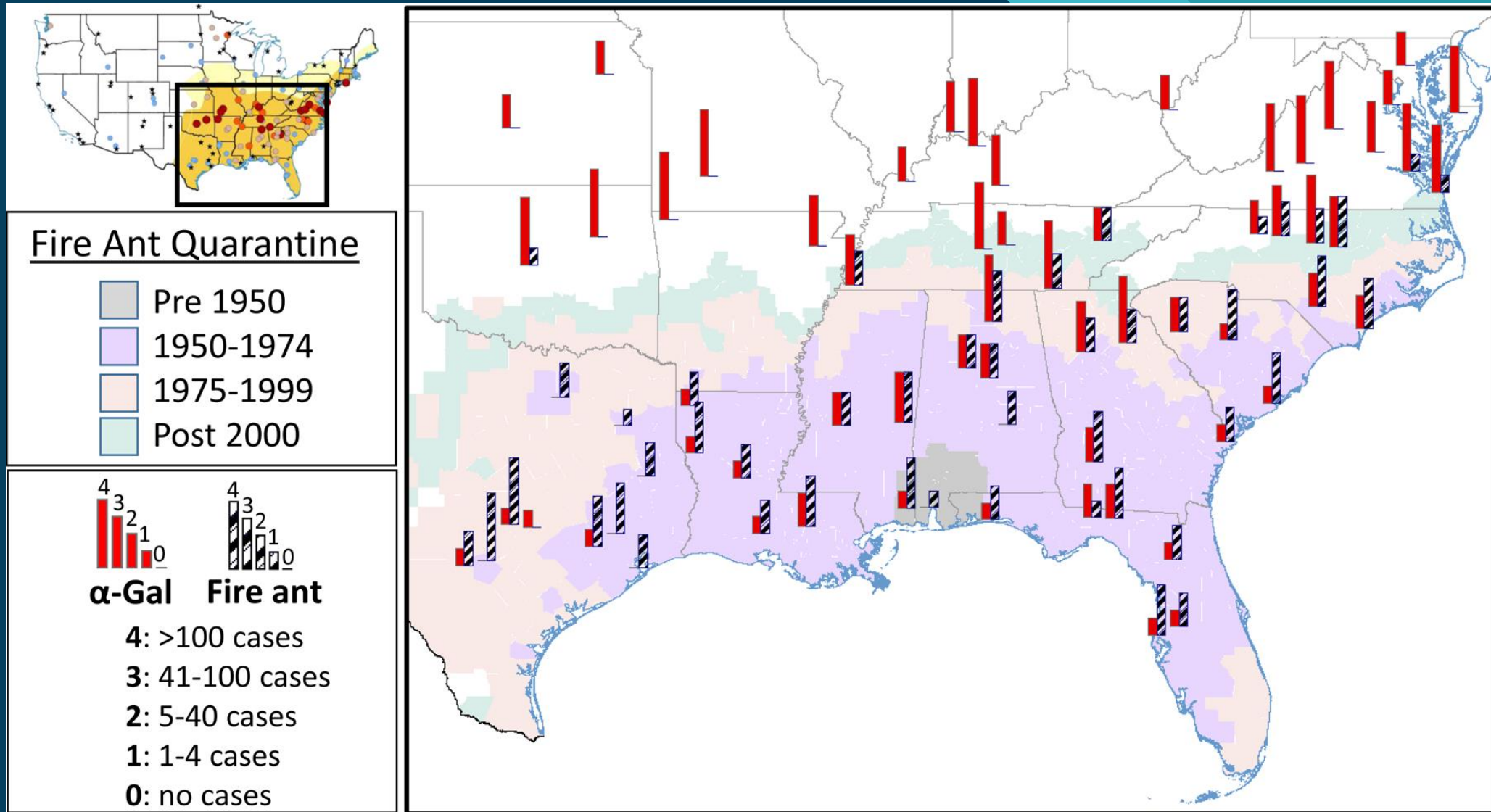
How many cases have you seen that relate to allergic reactions to fire ant bites in the last 3 years?

0 1-4 5-40 40-100 >100

How many on ImmunoTherapy (IT) for Fire Ant? _____ Are you aware of any other practices in your region or state that are seeing cases? ___;
contact information for those practices

FIG E2. Questionnaire of allergists in practice for second phase of the survey.

Epidemiological study -overlapping maps of Specific IgE for α -Gal & IFA overlaid on the quarantine data



Why? Fire Ant predation on ticks!

- Tests in BR and Pine Grove LA - Lone star ticks released into two areas with IFA 1.A. Untreated; B. Untreated but with barrier 2. Mirex treated to suppress fire ants
 - More larvae & ticks survived in the Mirex treated areas; None survived in non-treated areas. Barrier allowed survival
- *Environmental Entomology*, Volume 1, Issue 3, 1 June 1972, Pages 362–365,
- Mirex is no longer used due to its damage to the natural environment

Imported Fire Ants (IFA)

https://www.aphis.usda.gov/aphis/ourfocus/planthealth/plant-pest-and-disease-programs/pests-and-diseases/imported-fire-ants/CT_Imported_Fire_Ants#:~:text=APHIS%20works%20to%20prevent%20artificial,hay%20and%20soil%2Dmoving%20equipment.

Dec 2020 - FDA approves “GAL SAFE” pigs made by Revivacor using
“Intentional Genomic Alteration”



<https://www.medpagetoday.com/allergyimmunology/allergy/90211>

FYI: Informative University websites

- <https://web.uri.edu/tickencounter/fieldguide/tick-growth-comparison-charts/>
- <https://bugoftheweek.com/blog>
- https://entnemdept.ufl.edu/creatures/urban/medical/lone_star_tick.htm

Questions?

- <https://amaroohills.com>
- <https://alphagalinformation.org>
- <https://www.twoalphagals.com>



Thank You !



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