Tickborne Disease!

- Be sure to consider in flu-like illness, especially in the absence of upper respiratory symptoms and when symptoms last longer than 4-5 days
- <u>Ixodes scapularis</u> and <u>Dermacentor variabilis</u> are the most common ticks in our area
 - How small can a tick be? Check out this picture of a larval *lxodes* tick I took from urgent care:



- Lyme
 - Presentation
 - Tick must be attached 36-48 hours to transmit disease
 - EM present in 70-80% of patients. Occurs at site of the bite, 3-30 days after bite.
 - Disseminated disease occurs days to weeks later: neurologic, MSK, CV symptoms and possibly multiple EM lesions
 - Prophylaxis
 - 200mg single dose doxycycline within 72 hours of tick removal, if tick attached for 36+ hours
 - Depending on the study, 0.2-0.4% of patients given prophylactic doxy subsequently developed Lyme, whereas 2.2-3.2% of non-prophylaxed patients developed Lyme This is an 87-90% Relative Risk Reduction with NNT of approximately 35-50 to prevent one case of Lyme.
 - Diagnosis
 - Presence of EM along with possible tick exposure is diagnostic for Lyme on its own
 - Use the 'Tick Borne Illness Work Up" order set will include the appropriate diagnostics
 - Two-tiered testing with ELISA followed by Western blot

REFERENCE FOR TICK BOURNE DISEASE – JUNE 2023

- Testing is less than 50% sensitive in early disease and greater than 90% in subsequent stages
- Treatment
 - Doxycycline (first line), Amoxicillin
 - Avoid doxy in patients less than 8 years old or pregnant, however AAP now supports doxy in young children if used 21 days or less due to the low-risk for dental staining
 - Ceftriaxone for late neurologic symptoms
 - 10-28 days of treatment depending on abx and indication (check UTD)
- Persistent Symptoms
 - 10-20% of patients are symptomatic even after treatment
 - "Post-Lyme Disease Syndrome" Encompasses symptoms that persist for greater than 6 months after completion of antibiotics. Antibiotics should not be extended in these patient since it will not improve symptoms; likely from inflammatory dysregulation.
 - Interestingly, patients with Lyme Arthritis are commonly found to have peptidoglycan (very inflammatory) from Borrelia bacteria remaining in the synovial fluid leading to the persistent inflammation, as opposed to actively growing bacteria.
- Anaplasma
 - Presentation
 - General viral-like syndrome, 5-14 days after exposure
 - Rash is rare
 - Diagnosis
 - Use the 'Tick Borne Illness Work Up" order set, PCR and serology are generally used
 - PCR is likely to positive in the first week of illness, whereas serology will likely be negative in the first week
 - In my experience, I have never had serology come back positive but I have had many PCR come back positive
 - o Treatment
 - Doxycycline for a minimum of 10 days, at least 3 days after resolution of the fever
 - If can't tolerate tetracycline, use 7-10 day course of rifampin
 - Prior infection may not confer long-lasting immunity
- Babesia

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- Presentation
 - Fever, Myalgias, lysis of erythrocytes
 - Symptoms develop 1-9 WEEKS after exposure
- o Diagnosis
 - PCR
 - Could diagnose by microscopy with Wright or Giemsa stain if available
 - Hemolysis, thrombocytopenia, elevated transaminases

- One of the local ID physicians recently recommended checking haptoglobin and LDH - if both normal, very unlikely to be Babesia. If low haptoglobin and elevated LDH, could be Babesia.
- o Treatment
 - Combination of atovaquone and azithromycin for 7-10 days
- Immunocompromised patients, those treated with rituximab, patients with malignancy (specifically B cell lymphoma), HIV, organ transplant may develop a relapse of Babesia despite a prior course of treatment.

TABLE 1

Incidence, Causative Agents, Vectors, and Geographic Distribution of Tickborne Diseases in the United States

Disease	Reported annual incidence	Causative agents	Vectors	Geographic distribution
Lyme disease	30,000	Borrelia burgdorferi	Black-legged/deer tick (Ixodes scapularis or Ixo- des pacificus)	New England, mid-Atlantic states, upper Midwest, north- ern California
Rocky Mountain spot- ted fever and other spotted fever Rick- ettsioses: <i>Rickettsia</i> <i>parkeri</i> rickettsiosis, Pacific Coast tick fever, Rickettsial pox	6,000	Rickettsia rickettsia R. parkeri Rickettsia species 364D Rickettsia akari	Wood tick (Dermacentor andersoni) American dog tick (Der- macentor variabilis) Brown dog tick (Rhipicephalus sanguineus)	Southeastern and south- central states (Oklahoma, Arkansas, Missouri, Ten- nessee, and North Carolina account for more than 60% of cases), Arizona and New Mexico
Anaplasmosis	5,000	Anaplasma phagocytophilum	Black-legged/deer tick (I. scapularis or I. pacificus)	Northeast, Midwest, and West Coast
Ehrlichiosis	1,500	Ehrlichia chaffeensis Ehrlichia ewingii Ehrlichia muris eauclairensis	Lone Star tick (<i>Ambly- omma americanum</i>) Black-legged/deer tick (<i>I. scapularis</i>) American dog tick (<i>D. variabilis</i>)	Southeastern and south- central states
Babesiosis	1,000- 2,000	Babesia microti Babesia duncani	Black-legged/deer tick (I. scapularis or I. pacificus)	Northeast, upper Midwest, few cases in Washington and California
Tularemia	Few hundred	Francisella tularensis	Lone star tick (A. americanum) Wood tick (D. andersoni) Dog tick (D. variabilis)	South-central states (Okla- homa, Arkansas, Missouri, and Kansas)
Colorado tick fever	200-300	Colorado tick fever virus (double-stranded RNA arbovirus)	Wood tick (<i>D. andersoni</i>)	Rocky Mountain region
Tickborne relapsing fever	20-30	Borrelia species: Borrelia hermsii, Borrelia parkerii, Borrelia turicatae	Ticks of <i>Ornithodoros</i> genus	Rocky Mountain region

Information from references 1, 2, and 4-18.

TABLE 2

Clinical Characteristics of Tickborne Diseases

Disease	Signs and symptoms	Diagnosis	Treatment
Lyme disease	Early localized: EM rash at site of inocula- tion, flulike symptoms Early disseminated: secondary EM lesions, neurologic (meningitis, facial palsy), muscu- loskeletal (arthralgias and myalgias), and cardiovascular symptoms (temporary atrioventricular block) Late disseminated: encephalomyelitis, polyarticular arthritis, Lyme carditis	Clinical diagnosis for early local- ized disease Enzyme-linked immunosorbent assay followed by Western blot assay for unclear or later stage diagnosis	Doxycycline 100 mg twice per day or 4 mg per kg in two divided doses for children > 8 years Amoxicillin 500 mg three times per day or 50 mg per kg in three divided doses for children Cefuroxime axetil (Ceftin) 500 mg twice per day or 30 mg per kg in two divided doses for children Azithromycin (Zithromax) 500 mg once per day or 10 mg per kg per day for children IV ceftriaxone (Rocephin) 2 g per day or 50 to 75 mg per kg per day for children used for neu- rologic manifestations of late disease Duration of therapy: Early localized: 14 days Early disseminated: 14 to 21 days Late disseminated: 14 to 28 days
Rocky Mountain spotted fever	Flulike symptoms with macular rash starting on wrists, forearms, and ankles, becomes petechial	Clinical signs and symptoms including thrombocytopenia and hyponatremia, elevated transami- nases, and hyperbilirubinemia IFA is confirmatory but should not delay treatment	Doxycycline 100 mg twice per day or 4 mg per kg for children in two divided doses Chloramphenicol if contraindication to doxycycline Duration of therapy: seven to 10 days
Anaplas- mosis and ehrlichiosis	Flulike symptoms with gastrointestinal predominance Rash in up to one-third of patients with ehrlichiosis, particularly children	Clinical signs and symptoms including thrombocytope- nia, leukopenia, and elevated transaminases IFA is confirmatory but should not delay treatment	Doxycycline 100 mg twice per day or 4 mg per kg for children in two divided doses Rifampin or chloramphenicol if contraindication to doxycycline Duration of therapy: minimum of 10 days, con- tinue for at least three days after fever subsides
Babesiosis	Nonspecific flulike symptoms; jaundice may be present	Laboratory findings of hemo- lytic anemia, thrombocytopenia, elevated transaminases Thin blood smear with charac- teristic "Maltese cross" pattern or PCR	Atovaquone (Mepron) 750 mg twice per day or 40 mg per kg in two divided doses for children plus azithromycin 500 mg on day one followed by 250 mg per day or 10 mg per kg on day one followed by 5 mg per kg per day for children Duration of therapy: seven to 10 days IV clindamycin plus oral quinine and/or exchange transfusion for severe disease
Tularemia	Flulike symptoms, cutaneous eschar at site of inoculation, and painful regional lymphadenopathy	History of exposure to rab- bits and other rodents or ticks, leukocytosis Culture is the dianostic standard for diagnosis of tularemia, but has biosafety concerns; PCR or paired serologies may also be used	Intramuscular streptomycin 2 g in two divided doses or 15 mg per kg in two divided doses for children Intramuscular gentamicin or IV 5 mg per kg in two divided doses Duration of therapy: seven to 10 days
Colorado tick fever	Triad of high fever (up to 104°F [40°C]), severe myalgias, and headache; fever is often biphasic; "saddle-back" pattern	Common laboratory find- ings of leukopenia and thrombocytopenia Reverse-transcriptase PCR or paired sample serologic testing	Supportive care only
Tickborne relapsing fever	Flulike symptoms with high fever in relapsing or remitting pattern	Detection of spirochetes in blood using dark field microscopy or specific staining	Doxycycline 100 mg twice per day or tetracy- cline 500 mg every 6 hours for adults Erythromycin 500 mg four times per day for pregnant women or 30 to 50 mg per kg in four divided doses for children IV ceftriaxone or penicillin G for patients with central nervous system involvement Duration of therapy: seven to 10 days Jarisch-Herxheimer reactions common with treatment

Information from references 1, 2, 4, 5, 7-9, 13, and 15-24.

References:

- <u>One Dose of Doxycycline for the Prevention of Lyme Disease: A Review of Clinical</u> <u>Effectiveness and Guidelines [Internet] - PubMed (nih.gov)</u>
- <u>Prophylaxis with Single-Dose Doxycycline for the Prevention of Lyme Disease after an</u> <u>Ixodes scapularis Tick Bite | NEJM</u>
- AAFP Tickborne Diseases Article
- <u>Persistence of Borrelia burgdorferi peptidoglycan in Lyme Arthritis</u>
- Meta-analysis for Lyme Prophylaxis