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Lyme Disease and Other Tickborne Diseases in New Hampshire

Key Points and Recommendations:

1. NH continues to have one of the highest rates of Lyme disease in the nation and ~60% of blacklegged ticks sampled in NH are infected with *Borrelia burgdorferi*, the bacterium that causes Lyme disease.
2. Prevent tickborne disease through the use of DEET insect repellent, wearing long pants and sleeves outdoors, and daily tick checks followed by prompt removal of any ticks.
3. Healthcare providers should diagnose early Lyme disease clinically when an erythema migrans rash because diagnostic serologies (including IgM) may not yet be positive.
4. Report all tickborne diseases, confirmed or suspected, to the NH DPHS Bureau of Infectious Disease Control at 603-271-4496 (after hours 1-800-852-3345, x5300).

Background:

Lyme disease (*Borrelia burgdorferi*), babesiosis (*Babesia microti* and other species), anaplasmosis (*Anaplasma phagocytophilum*), and Powassan virus are transmitted by the bite of the blacklegged tick (*Ixodes scapularis*), also known as the deer tick. Although the lifespan of this tick is two years, people are most likely to be infected between May and August when the aggressive nymph stage is active. Nymphs are very small (< 2mm) and difficult to see unless they become engorged with blood.

Epidemiology:

The Centers for Disease Control and Prevention (CDC) reported 33,461 confirmed and probable cases of Lyme disease in the United States in 2014 (the most recent available data), which is an increase from the 2013 count of 27,203 cases. Over the last decade, reported Lyme disease cases have increased significantly in NH. In 2015, there were an estimated 1,373 new cases reported statewide, with the highest rates in Rockingham, Strafford and Hillsborough counties. New Hampshire Lyme disease data and maps by county and town from 2006-2014 are available at: <http://www.dhhs.nh.gov/dphs/cdcs/lyme/publications.htm>. In 2015, 110 cases of anaplasmosis, and 53 cases of babesiosis were reported in NH. The only case of locally-acquired Powassan virus infection was reported in 2013.

An individual's risk of tickborne disease depends on their outdoor activities and the abundance of infected ticks. Tick surveillance performed during 2013-2014 in NH counties showed that >50% of ticks tested in most counties were infected with the bacteria causing Lyme disease with the exception of Carroll, Cheshire, Coos and Sullivan counties where very low numbers of ticks were collected, precluding prevalence assessment. Tick surveillance maps by county from 2013-2014 are available at:

<http://www.dhhs.nh.gov/dphs/cdcs/lyme/documents/blacklegged13-14.pdf>.

Babesia and *Anaplasma* have been detected in ticks in NH, though reliable prevalence data for these pathogens in ticks is not available due to small sample size.

Lyme Disease Diagnosis and Treatment

Clinical Presentation: In approximately 70-80% of patients, illness begins within the first 30 days after a tick bite with a red “bull’s-eye” rash that expands slowly, often with central clearing (erythema migrans [EM]). Early systemic manifestations may also include malaise, fever, headache, stiff neck, muscle and joint pains, and lymphadenopathy. At this early stage, serologic testing is often negative and treatment should be initiated based on clinical features. Early treatment generally leads to complete and rapid recovery, and may prevent seroconversion (so that later testing is negative). Patients who are not treated in the early stage of infection may develop a variety of syndromes weeks to months after onset of symptoms, including aseptic meningitis, encephalitis, cranial neuritis, and cardiac abnormalities such as heart block or myopericarditis. Without treatment, a patient may develop chronic or intermittent episodes of arthritis or neurological symptoms months to years after onset.

In 2013, CDC released a report of three Lyme disease carditis cases in the northeastern United States that resulted in sudden cardiac death. While rare, these cases highlight the importance of prompt diagnosis and treatment for Lyme disease. Healthcare providers should ask patients with suspected Lyme disease about cardiac symptoms and obtain an electrocardiogram (EKG) if indicated. Healthcare providers should also consider Lyme disease for patients with unexplained heart block. The full report on this rare clinical presentation is available at: <http://www.cdc.gov/mmwr/preview/mmwrhtml/mm6249a1.htm>

Testing: Although treatment of early disease should be given based on clinical suspicion, clinicians should obtain laboratory testing. Within 4 weeks of infection, *Borrelia burgdorferi* antibodies can be detected using FDA-approved serologic testing. The CDC currently recommends a two-stage testing approach with the first step involving an ELISA as the screening test, and if positive or equivocal confirmed by Western Blot. A Western Blot should not be run if the initial ELISA test is negative. A patient is considered to have a positive Lyme Western Blot test if 2 of 3 IgM bands are reactive (24, 39, 41 kDa) OR if 5 of 10 IgG bands are reactive (18, 21, 28, 30, 39, 41, 45, 58, 66, 93 kDa). An isolated positive IgM (without a positive IgG) in a patient with tick exposure more than 8 weeks prior is likely to be a false positive test, since IgG antibodies are produced within 4-8 weeks after exposure. Only laboratories with validated and FDA-approved testing methods for Lyme ELISA and confirmatory Western blot should be used for diagnosis of disease.

Treatment: In 2010, an independent panel reviewed and approved the Infectious Disease Society of America (IDSA) guidelines for tickborne diseases. A summary of these recommendations is attached and the full guidelines are available at: <http://cid.oxfordjournals.org/content/43/9/1089.full.pdf+html>

Antibiotic Prophylaxis: Based on the high prevalence of Lyme disease in NH, providers can consider prescribing single dose doxycycline prophylaxis (200 mg) for patients who meet all four criteria outlined in the attached Lyme disease prophylaxis guidelines. Note that single-dose doxycycline is not 100% effective for prevention of Lyme disease; consequently, patients who receive this therapy should monitor themselves for the development of Lyme disease as well as other tickborne diseases, including anaplasmosis and babesiosis. Testing the tick for tickborne infectious agents is available in certain labs but is not recommended for guiding individual prophylaxis or treatment decisions.

Anaplasmosis

Clinical Presentation: Anaplasmosis (human granulocytic anaplasmosis [HGA], previously known as human granulocytic ehrlichiosis) is an infection of neutrophils caused by the rickettsial bacteria *Anaplasma phagocytophilum*, and is transmitted by the blacklegged tick. Symptoms typically occur 5-21 days following the bite of an infected tick, and may include fever, chills, headache, and myalgia. Some people, particularly elderly persons or those with weakened immune systems, may have severe illness. Laboratory abnormalities may include leukopenia, lymphopenia, thrombocytopenia, and mild elevation in liver enzyme levels.

Testing: Identification of the characteristic intragranulocytic inclusions on blood smear is the most rapid diagnostic method, but requires lab expertise. Acute and convalescent antibody assays are the most sensitive diagnostic method.

Treatment: Doxycycline is the first line therapy for anaplasmosis (see attached treatment guideline table). If co-infected with Lyme disease, doxycycline will treat both infections. Antibiotic therapy should not be delayed in a patient with a suggestive clinical presentation pending the results of diagnostic testing.

Babesiosis

Clinical Presentation: Babesiosis is caused by the intraerythrocytic protozoan *Babesia microti* (or other *Babesia* species) and is transmitted by the blacklegged tick. Clinical presentation can range in severity from asymptomatic to a rapidly fatal illness. Most people with *Babesia* present within 1-6 weeks after bite of an infected tick with fever, chills, sweats, myalgia, arthralgias, anorexia, nausea, vomiting, and/or fatigue. Severe and fatal cases most often occur in patients who are older or have a weakened immune system, particularly those without a spleen. Complications of *Babesia* can include acute respiratory failure, congestive heart failure, renal failure, and disseminated intravascular coagulation. Laboratory abnormalities may include findings of a hemolytic anemia, thrombocytopenia, elevated liver enzymes, and renal dysfunction. Rare cases of relapsing disease have been reported.

Testing: Diagnosis is based on identification of *Babesia* parasites in a blood smear or by PCR amplification of babesial DNA.

Treatment: Babesiosis can be successfully treated with antimicrobial therapy (see attached treatment guideline table). RBC exchange transfusion may also be needed in cases with a high parasitemia ($\geq 10\%$), or in persons with severe clinical disease.

Powassan Virus Infection

Clinical Presentation: Powassan (POW) virus is an RNA virus of the genus *Flavivirus* with an incubation period of 7-30 days following bite of an infected tick. Although most infections are subclinical, symptoms may include fever, headache, vomiting, and generalized weakness that can progress to meningoencephalitis. About half of those that survive clinical disease have permanent neurological sequelae.

Testing: Cerebrospinal fluid (CSF) findings include normal or mildly elevated protein, normal glucose concentration, and lymphocytic pleocytosis typically with <500 white blood cells/mm³. Brain magnetic resonance imaging (MRI) reveals changes consistent with microvascular ischemia or demyelinating disease in the parietal or temporal lobes. Electroencephalography (EEG) shows generalized slow wave activity.

Diagnosis can be made by the detection of POW virus-specific IgM antibody in serum or CSF, combined with a consistent clinical presentation. Currently, POW virus testing is not commercially available but can be arranged through the NH Public Health Laboratories.

Treatment: Treatment is supportive.

Reporting Tickborne Diseases:

Clinicians should report suspected and confirmed cases of Lyme disease, anaplasmosis, babesiosis, and Powassan virus infection to the NH DPHS Bureau of Infectious Disease Control at 603-271-4496 (after hours 1-800-852-3345, x5300). When filling out the Lyme disease case report form, it is important to record the date of symptom onset because this information is used to determine whether a case meets the CDC case definition for surveillance. A copy of the most recent Lyme disease case report form is attached. The case report form is also available at: <http://www.dhhs.nh.gov/dphs/cdcs/documents/lymediseasereport.pdf>

Prevention Messages for Patients:

- Avoid tick-infested areas when possible and stay on the path when hiking to avoid brush.
- Wear light-colored clothing that covers arms and legs so ticks can be more easily seen.
- Tuck pants into socks before going into wooded or grassy areas.
- Apply insect repellent (20-30% DEET) to exposed skin. Other repellent options may be found here: Outdoor workers in NH are at particular risk of tickborne diseases and they should be reminded about methods of prevention.
- Do daily tick checks to look for ticks on the body, especially warm places like behind the knees, the groin, and the back and neck.
- Pets returning inside may also bring ticks with them. Performing tick checks and using tick preventatives on pets will minimize this occurrence.
- Shower soon after returning indoors to wash off any unattached ticks and check clothes for any ticks that might have been carried inside. Placing clothes in the dryer on high heat for an hour effectively kills ticks. A recent study suggests that if clothing is not wet, shorter drying times (minimum of 6 minutes) may effectively kill ticks.
- Remove ticks promptly using tweezers. Tick removal within 36 hours of attachment can prevent disease.
- Monitor for signs and symptoms of tickborne diseases for 30 days after a tick bite. Patients should contact their healthcare provider if symptoms develop.

Additional background information about tickborne diseases and prevention can be found in the State of New Hampshire Tickborne Disease Prevention Plan at:

<http://www.dhhs.nh.gov/dphs/cdcs/lyme/documents/tbdpreventionplan.pdf>

For any questions regarding the contents of this message, please contact NH DHHS, DPHS, Bureau of Infectious Disease Control at 603-271-4496 (after hours 1-800-852-3345 ext.5300).

To change your contact information in the NH Health Alert Network, contact Thom Flynn at 603-271-7499 or tdflynn@dhhs.nh.gov.

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From: Benjamin Chan, MD, MPH- State Epidemiologist
Originating Agency: NH Department of Health and Human Services, Division of Public Health Services

Attachments: 1. Tickborne diseases treatment table, 2. Lyme disease prophylaxis guidelines, 3. Lyme disease case report form

ATTACHMENT 1

Tick bites and single-dose doxycycline as prophylactic treatment for Lyme disease in NH (Based on the 2006 Infectious Disease Society of America guidelines)

A **single** dose of doxycycline (200 mg) may be offered to adult patients and to children ≥ 8 years of age (4 mg/kg up to a maximum dose of 200 mg) when ALL of the following conditions exist:

1. The attached tick is a blacklegged tick (deer tick, *Ixodes scapularis*). Tick identification is most accurately performed by an individual trained in this discipline. However, blacklegged ticks are very common in southeastern and central New Hampshire and there are many images available online to help identification. AND
2. The tick has been attached for at least 36 hours. This determination can be made by asking the patient about outdoor activity in the time before the tick bite was noticed to estimate attachment time, or by asking about degree of engorgement. Unengorged (unfed) blacklegged ticks are typically flat. Any deviation from this “flatness,” which is often accompanied by a change in color from brick red to a gray or brown, is an indication that the tick has been feeding. AND
3. Prophylaxis can be started within 72 hours of the time that the tick was removed. This time limit is suggested because of an absence of data on the efficacy of prophylaxis for tick bites following longer time intervals after tick removal. AND
4. Doxycycline prophylaxis is not contraindicated. Doxycycline is contraindicated in pregnant women and children less than 8 years old. The other common antibiotic treatment for Lyme disease, amoxicillin, is not recommended for prophylaxis because of an absence of data on an effective short-course prophylaxis regimen, the likely need for a multi-day regimen along with its possible adverse effects, and the excellent efficacy of treatment if signs or symptoms do develop.

Note that single-dose doxycycline is not 100% effective for prevention of Lyme disease; consequently, patients who receive this therapy should monitor themselves for the development of Lyme disease as well as other tickborne diseases including anaplasmosis and babesiosis.

Adapted from: Wormser GP, et al. The Clinical Assessment, Treatment, and Prevention of Lyme Disease, Human Granulocytic Anaplasmosis, and Babesiosis: Clinical Practice Guidelines by the Infectious Diseases Society of America. *Clinical Infectious Diseases*;2006;43:1089 –1134. Available online at: <http://cid.oxfordjournals.org/content/43/9/1089.full>

ATTACHMENT 2

**NH DPHS Treatment Recommendations for Tickborne Diseases
 Summary of 2006 Infectious Disease Society of America Guidelines**

Disease	Treatment Regimens for Adults	Treatment Regimens for Children
Lyme disease	Oral Options	
	Doxycycline 100 mg PO twice daily*	For children 8 years and older: Doxycycline 4 mg/kg/day in 2 divided doses (max 100 mg/dose)
	Amoxicillin 500 mg PO three times daily	Amoxicillin 50 mg/kg/day in 3 divided doses (max 500 mg/dose)
	Cefuroxime axetil 500 mg PO twice daily	Cefuroxime axetil 30 mg/kg/day in 2 divided doses (max 500 mg/dose)
	Parenteral options	
	Preferred: Ceftriaxone 2g IV Once daily	Preferred: Ceftriaxone 50-75 mg/kg IV Once daily (max 2g)
	Alternative: Cefotaxime 2g IV every 8 hrs	Alternative: Cefotaxime 150-200 mg/kg/day IV in 3-4 divided doses (max 6g/day)
	Alternative: Penicillin G 3-4 million units IV every 4 hrs	Alternative: Penicillin G 200,000-400,000 U/kg/day divided every 4 hr (max 18-24 million units per day)
Note: Choice of regimen, route and length of treatment for Lyme disease depends on symptoms and stage of disease.		
Anaplasmosis	Doxycycline 100 mg PO twice daily for 10 days* Alternatives: <u>Mild disease:</u> Rifampin 300 mg PO twice a day for 7-10 days (note: rifampin is not effective for Lyme disease. If coinfection exists, Amoxicillin or Cefuroxime would need to be <i>added</i> for treatment of Lyme disease as well)	8 years and older: Doxycycline 4 mg/kg/day PO in 2 divided doses for 10 days (max 100 mg/dose) Under 8 years old (without concomitant Lyme disease): <u>Mild disease:</u> Rifampin 10 mg/kg PO twice daily (max 300 mg/dose) for 7-10 days, with close monitoring after for resolution of symptoms. <u>Severe disease:</u> Doxycycline (dose as above) for 4-5 days, with close monitoring after for resolution of symptoms. Under 8 years old (with concomitant Lyme disease): --If Doxycycline is used for 4-5 days (dose as above) then complete a 14-day course with Amoxicillin OR Cefuroxime axetil (doses as above) after treating with doxycycline. --If Rifampin is used, add Amoxicillin or Cefuroxime for treatment of Lyme disease as above.
Babesiosis	Atovaquone 750 mg PO every 12 hrs plus Azithromycin 500-1000 mg on day 1, then 250 mg PO daily thereafter <u>Severe disease:</u> Clindamycin 300-600 mg IV every 6 hrs (or 600 mg PO every 8 hrs)	Atovaquone 20 mg/kg PO every 12 hrs (max 750 mg/ dose) plus azithromycin 10 mg/kg/day once daily on day 1 (max 500 mg/dose) then 5 mg/kg once daily (max 250 mg/dose) thereafter.

	plus Quinine 650 mg PO every 6-8 hrs. Consider exchange transfusion.	<u>Severe disease:</u> Clindamycin 7-10 mg/kg every 6-8 hrs PO or IV (max 600 mg/dose) plus Quinine 8 mg/kg PO every 8 hrs (max 650 mg/dose). Consider exchange transfusion.
Note: patients coinfecting with <i>B. burgdorferi</i> or <i>A. phagocytophilum</i> should be treated with additional antibiotics as described above.		

***NOTE: For pregnant women, doxycycline should not be used.**

Adapted from: Wormser GP, et al. The Clinical Assessment, Treatment, and Prevention of Lyme Disease, Human Granulocytic Anaplasmosis, and Babesiosis: Clinical Practice Guidelines by the Infectious Diseases Society of America. *Clinical Infectious Diseases*;2006;43:1089 –1134. Available online at: <http://cid.oxfordjournals.org/content/43/9/1089.full>

**NEW HAMPSHIRE LYME DISEASE CASE REPORT FORM
HEALTH CARE PROVIDER**

Patient's Name _____ Report Date _____
 (Last Name) (First Name)
 Date of Birth _____ Age _____ Male Female Unknown
 Address _____
 City / Town _____ County _____ State _____ Zip _____
 Home Phone _____ Work Phone _____
 Occupation: _____

Race
 White
 African American
 Asian
 Hawaiian or Pacific Islander
 Native Am./Alaskan Native
 Other
 Unknown
Ethnicity
 Hispanic
 Not Hispanic
 Unknown

SYMPTOMS AND SIGNS OF CURRENT EPISODE (Please answer each question)

Is this person being diagnosed with Lyme Disease?..... Yes No
 Date of symptom onset _____ Onset date unknown Date of Lyme Disease diagnosis _____

DERMATOLOGIC:

Erythema migrans (physician diagnosed EM at least 5 cm in diameter)?..... Yes No Unknown

RHEUMATOLOGIC:

Arthritis characterized by recurrent brief attacks of joint swelling? Yes No Unknown

NEUROLOGIC:

Bell's palsy or other cranial neuritis? Yes No Unknown
 Radiculoneuropathy? Yes No Unknown
 Lymphocytic meningitis? Yes No Unknown
 Encephalitis/Encephalomyelitis? Yes No Unknown
 CSF tested for antibodies to *B. burgdorferi*? Yes No Unknown
 Antibody to *B. burgdorferi* higher in CSF than serum Yes No Unknown

CARDIOLOGIC:

Acute onset 2nd or 3rd degree atrioventricular block? Yes No Unknown

Pregnant: Yes No Unknown
Hospitalized: Yes No Unknown If yes, where _____
Treatment: Doxycycline Amoxicillin Other: _____

Duration of treatment in days: _____

Has this patient been diagnosed with Lyme Disease prior to this diagnosis? Yes, date (mm/yyyy) _____ No Unknown

EXPOSURE HISTORY

Tick Bite reported within 30 days of onset: Yes No Unknown
 In the 30 days prior to symptom onset, did this individual travel outside of NH: Yes, out of state Yes, out of country
 No Unknown
County and state most likely exposed? _____

LABORATORY RESULTS (Check all that apply)

EIA/IFA: Positive Equivocal Negative Not done/Unknown Date if positive: _____
Western Blot: IgM Positive IgM Negative Not done/Unknown Date if positive: _____
 IgG Positive IgG Negative Not done/Unknown Date if positive: _____
Culture Results/Other: _____

HEALTH CARE PROVIDER REPORTING INFORMATION:

Reported by _____
 Ordering Provider _____ Phone _____
 Provider Facility _____
 City/Town _____ State _____ Zip _____

**For NH DHHS Staff Only
Imported**

Acquired in NH
 Acquired Outside US
 Acquired in Another State
 Unknown

Case Status

Confirmed (meets CDC definitions)
 Probable (meets CDC definitions)
 Suspected (meets CDC definitions)
 Not A Case
 Out of state

Notes: