Reasons for False Negative (Seronegative) Test Results in Lyme Disease

Nine Reasons for False Negative Lyme Disease Blood Test Results

Reprinted from the Lyme Disease Foundation's brochure, *Frequently Asked Questions About Lyme Disease*, available online at www.lyme.org

- 1. Antibodies against *Borrelia burgdorferi* (Bb) are present, but the laboratory is unable to detect them.
- 2. Antibodies against Bb may not be present in detectable levels in a patient with Lyme disease because the patient is currently on, or has recently taken, antibiotics. The antibacterial effect of antibiotics can reduce the body's production of antibodies.
- 3. Antibodies against Bb may not be present in detectable levels in a patient with Lyme disease because the patient is currently on or has previously taken anti-inflammatory steroidal drugs These can suppress a person's immune system, thus reducing or preventing an antibody response.
- 4. Antibodies against Bb may not be present in detectable levels in a patient with Lyme disease because the patient's antibodies may be bound with the bacteria with not enough free antibodies available for testing.
 - For this reason, some of the worst cases of Lyme disease test negative -- there is too much bacteria for the immune system to handle.
- 5. Antibodies against Bb may not be present in detectable levels in a patient with Lyme disease because the patient could be immunosuppressed for a number of other reasons, and the immune system is not reacting to the bacteria.
- 6. Antibodies against Bb may not be present in detectable levels in a patient with Lyme disease because the bacteria has changed its makeup (antigenic shift) limiting recognition by the patient's immune system.
- 7. Antibodies against Bb may not be present in detectable levels in a patient with Lyme disease because the patient's immune response has not been stimulated to produce antibodies, i.e., the blood test is taken too soon after the tick-bite (8-6 weeks).
 - Please do not interpret this statement as implying that you should wait for a positive test to begin treatment.
- 8. Antibodies against Bb may not be present in detectable levels in a patient with Lyme disease because the laboratory has raised its cutoff too high.
- 9. Antibodies against Bb may not be present in detectable levels in a patient with Lyme disease because the patient is reacting to the Lyme bacteria, but is not producing the "right" bands to be considered positive.

Reasons Why A Seronegative Test Result Might Occur

Reprinted from Tom Grier's Lyme Alliance site at www.lymealliance.org

- 1. Recent infection before immune response
- 2. Antibodies are in immune complexes
- 3. Spirochete encapsulated by host tissue (i.e. lymphocytic cell walls)
- 4. Spirochetes are deep in host tissue
- 5. Blebs in body fluid, no whole organisms needed for PCR
- 6. No spirochetes in body fluid on day of test
- 7. Genetic heterogeneity (300 strains in U.S.)
- 8. Antigenic variability
- 9. Surface antigens change with temperature
- 10. Utilization of host protease instead of microbial protease
- 11. Spirochete in dormancy phase
- 12. Recent antibiotic treatment
- 13. Recent anti-inflammatory treatment
- 14. Concomitant infection with *Babesia* may cause immunosuppression
- 15. Other causes of immunosuppression
- 16. Lab with poor technical capability for Lyme disease
- 17. Lab tests not standardized for late stage disease
- 18. Lab tests labeled "for investigational use only"
- 19. CDC criteria is epidemiological, not a diagnostic criteria

The majority of laboratories, including state and county public health laboratories, use the CDC epidemiological criteria for reporting Bb Western Blot IgM and IgG test results. Some of these labs may automatically insert a statement to physicians similar to the following one included by the Sonoma County Department of Health Services Public Health Laboratory:

The diagnosis of Lyme disease must include careful clinical evaluation and should not be based only on the detection of antibodies to *Borrelia burgdorferi*.

Unfortunately, however, most physicians *do* use these tests as definitively diagnostic, rather than making a clinical diagnosis based on patient symptomology and response to the various antimicrobial protocols used to treat Lyme and common tickborne co-infections.