



CORRELATION OF LYME DISEASE WITH IMMUNE DYSFUNCTION & REACTIVATION OF OLD VIRUSES

Dr. H.S. Mangat
Dr. H Kaur
H Sawhney
Dr. P Billa
S Matharu

BACKGROUND

- ❑ Lyme disease was first recognized in 1975 after researchers investigated why unusually large numbers of children were being diagnosed with Juvenile Rheumatoid Arthritis in Lyme, Connecticut, and two neighboring towns.
- ❑ Most of children lived and played near wooded areas infested with ticks. Most of the symptoms started in summer months. Many reported being bitten by a tick with a rash site.
- ❑ Further investigations discovered that tiny deer ticks infected with a spiral-shaped bacterium or spirochete (which was later named *Borrelia Burgodeferi*) were responsible for the outbreak of arthritis in Lyme.

BACKGROUND

- ❑ in the US, a CD4/CD8 ratio ranging from 0.9 to 1.9 is considered to be normal in non-immunocompromised individuals.
- ❑ Lyme disease is diagnosed based on symptoms, physical findings (e.g., rash), and the possibility of exposure to infected ticks.
- ❑ Currently, FDA has approved for two tier testing for Lyme.
- ❑ Screening with ELISA
- ❑ All positive and inconclusive results should be further tested with Western blot to confirm.
- ❑ ELISA is very sensitive. It does not give any information about current or past exposure.

2) RESEARCH FOCUS/GOAL

- ❑ to determine if there is a correlation between **Lyme Disease** (Western Blot IgG 41 positive) and **Immune Dysfunction** (CD4/CD8 ratio)
- ❑ Null Hypothesis: There is no correlation between Lyme Disease and Immune Dysfunction.
- ❑ Alternative Hypothesis: There is a correlation between Lyme Disease and Immune Dysfunction.

3) METHODOLOGY

- ❑ 183 patients at two medical centers were evaluated in two Lyme endemic communities of Maryland, US.
- ❑ Patients presenting with clinical features of Lyme in the past three years, with a positive titre were included in the study.
- ❑ Retrospective correlational study: for all the patients identified with confirmed Lyme Disease, their medical records were used to collect data on their Immunological tests, including a IgG and IgM panel, CD4/CD8 ratio, and the presence or absence of Ehrlichia, Bartonella, and Babesia microorganisms.
- ❑ Patients with Lyme Disease, but without a full Immunological panel done for each episode of illness, were excluded from study.
- ❑ Further investigation of 148 of these patients correlated CD4/CD8 ratio and their IgG 41 band, using both a one and two tailed Statistical tests.

4) STATISTICAL RESULTS

- ❑ The mean CD4/CD8 ratio in the 148 patients was 2.41; with a variance of 1.05 and a standard deviation of 1.025.
- ❑ Assuming a normal CD4/CD8 ratio of less than 2, with a 5% confidence interval, the p value on both a one tailed and two tailed test was shown to be $p=0.00001$.
- ❑ **The p-value was very statistically significant** (less than 0.05), thus there was enough evidence to reject the null hypothesis, that there is no correlation between Lyme Disease and Immune Dysfunction.
- ❑ Two patients with an initial CD4/CD8 ratio of 2.7 and 2.8 who were IgG 41 positive were subsequently tested with the Nanotrap Urine PCR and found to be positive for Lyme.

ADDITIONAL FINDINGS

- ❑ Many patients complained of extreme exhaustion as a symptom (staying in bed)
- ❑ In the cohort that was both IgG 41 positive and with altered CD4/8 ratio we ran viral titers for EBV and HHV6
- ❑ 55 of the 148 patients had viral titers run
- ❑ 41 had markedly elevated Ig G to EBV
- ❑ 38 had markedly elevated Ig G to HHV6
- ❑ Those with highly elevated EBV titers were treated with Valganciclovir/ Valacylovir.
- ❑ The majority of those treated with Valganciclovir felt better.

DISCUSSION AND IMPLICATIONS

- ❑ Overall, the alternative hypothesis was supported by this research: in that there **is** a correlation between Lyme Disease and Immunological Dysfunction (which can have future implications for including Immunological testing as part of routine Diagnostic protocol for detecting Lyme Disease in patients).
- ❑ Decreased CD4/CD8 ratio with a positive IgG 41 band appears to be a strong predictor of a subsequent diagnosis of Lyme disease despite current diagnostic guidelines.
- ❑ This is evidenced by the progression of patients who did not meet current CDC diagnostic guidelines to “neurolyme”, but presented with full blown involuntary movements, hallucinations and subsequent thirst episodes.

1

Lyme was initially termed as a "Rheumatoid arthritis type picture" in tick borne in 1975 there was a limited understanding of the role of CD4 and CD8 T lymphocytes then.

2

Rheumatoid Arthritis today is recognized as an auto-immune condition with a decreased CD8 count clinically causes the body to attack itself manifesting in synovial joint degradation and pain.

3

Syphilis a fellow spirochete & cousin to Lyme.

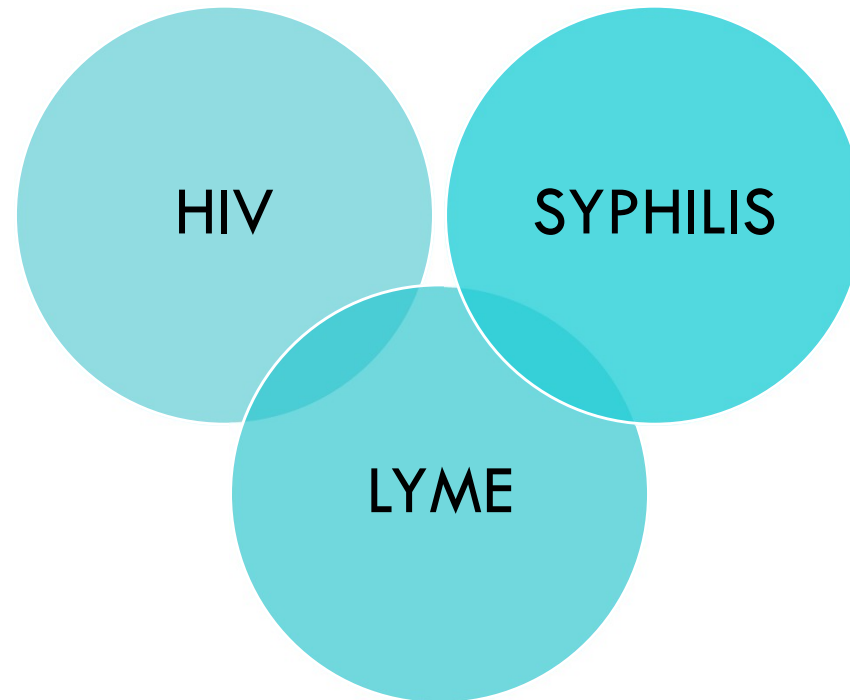
Lyme has many clinical correlates including Argyll Robertson Pupil, Keratitis.

4

HIV affects CD4 function and the correlation of EBV with Kaposi Sarcoma is well recognized.

HISTORICAL PERSPECTIVE

CLINICAL CORRELATES



6) TAKE HOME POINTS

- ❑ DO A 3 POINT LYME TEST
- ❑ CONVENTIONAL WESTERN BLOT
- ❑ CD4, CD8 AND CD4/8 RATIOS LIKE HIV MEDECINE
- ❑ VIRAL TITERS FOR EBV AND HHV6

❑ CAVEATS

YOUNG HAVE GOOD IMMUNE RESPONSE HENCE THIS CD4/8 RATIO MAY NOT BE VALID

OLDER INDIVIDUALS WHO WERE NEVER EXPOSED TO EBV IN ADOLESCENCE WILL HAVE NEGATIVE TITERS(EXCEPT THOSE WHO MIGHT HAVE SHARED DRINKS)