



INTERNATIONAL JOURNAL OF CREATIVE RESEARCH THOUGHTS (IJCRT)

An International Open Access, Peer-reviewed, Refereed Journal

TO STUDY ON ETIOLOGY, PATHOPHYSIOLOGY, RISK FACTORS AND MANAGEMENT OF LEPROSY

MELAM.YESURATNAM¹, VALLAGLAJAY BABU², SHAIK.KHAJA RAHAMTHULLA³,
MEDANDRAO. MOUNIKA⁴, NAGANDLA.LAKSHMI CHAITANYA⁵.

CORRESPONDING ADDRESS:

Department of Pharmacy Practice, A.M Reddy Memorial College of
Pharmacy, Petlurivararipalem, Narasaraopet, Palnadu-522601, Andhra Pradesh.

ABSTRACT

Leprosy, also known as Hansen's disease (HD), is a long-term infection by the bacteria *Mycobacterium leprae* or *Mycobacterium lepromatosis*. Infection can lead to damage of the nerves, respiratory tract, skin, and eyes. This nerve damage may result in a lack of ability to feel pain, which can lead to the loss of parts of a person's extremities from repeated injuries or infection through unnoticed wounds. An infected person may also experience muscle weakness and poor eyesight. Leprosy symptoms may begin within one year, but, for some people, symptoms may take 20 years or more to occur.

Leprosy is spread between people, although extensive contact is necessary. Leprosy has a low pathogenicity, and 95% of people who contract *M. leprae* do not develop the disease. Spread is thought to occur through a cough or contact with fluid from the nose of a person infected by leprosy. Genetic factors and immune function play a role in how easily a person catches the disease. Leprosy does not spread during pregnancy to the unborn child or through sexual contact. Leprosy occurs more commonly among people living in poverty. There are two main types of the disease – paucibacillary and multibacillary, which differ in the number of bacteria present. A person with paucibacillary disease has five or fewer poorly pigmented, numb skin patches, while a person with multibacillary disease has more than five skin patches. The diagnosis is confirmed by finding acid-fast bacilli in a biopsy of the skin.

KEYWORDS:-LEPROSY,MYCOBACTERIUM, DISEASE,BACTERIA.

LEPROSY

Leprosy, also known as Hansen's disease (HD), is a long-term infection by the bacteria *Mycobacterium leprae* or *Mycobacterium lepromatosis*. Infection can lead to damage of the nerves, respiratory tract, skin, and eyes. This nerve damage may result in a lack of ability to feel pain, which can lead to the loss of parts of a person's extremities from repeated injuries or infection through unnoticed wounds. An infected person may also experience muscle weakness and poor eyesight.

ETIOLOGY:-

It is caused by a bacterium, *Mycobacterium leprae*.

Leprosy was once feared as a highly contagious and devastating disease, but now we know it doesn't spread easily and treatment is very effective. However, if left untreated, the nerve damage can result in crippling of hands and feet, paralysis, and blindness. *Mycobacterium leprae* and *Mycobacterium lepromatosis*

M. leprae, one of the causative agents of leprosy: As an acid-fast bacterium, *M. leprae* appears red when a Ziehl–Neelsen stain is used.

M. leprae and *M. lepromatosis* are the mycobacteria that cause leprosy. *M. lepromatosis* is a relatively newly identified mycobacterium isolated from a fatal case of diffuse lepromatous leprosy in 2008. *M. lepromatosis* is indistinguishable clinically from *M. leprae*.

M. leprae is an intracellular, acid-fast bacterium that is aerobic and rod-shaped. *M. leprae* is surrounded by the waxy cell envelope coating characteristic of the genus *Mycobacterium*.

Genetically, *M. leprae* and *M. lepromatosis* lack the genes that are necessary for independent growth. *M. leprae* and *M. lepromatosis* are obligate intracellular pathogens, and cannot be grown (cultured) in the laboratory. The inability to culture *M. leprae* and *M. lepromatosis* has resulted in a difficulty definitively identifying the bacterial organism under a strict interpretation of Koch's postulates.



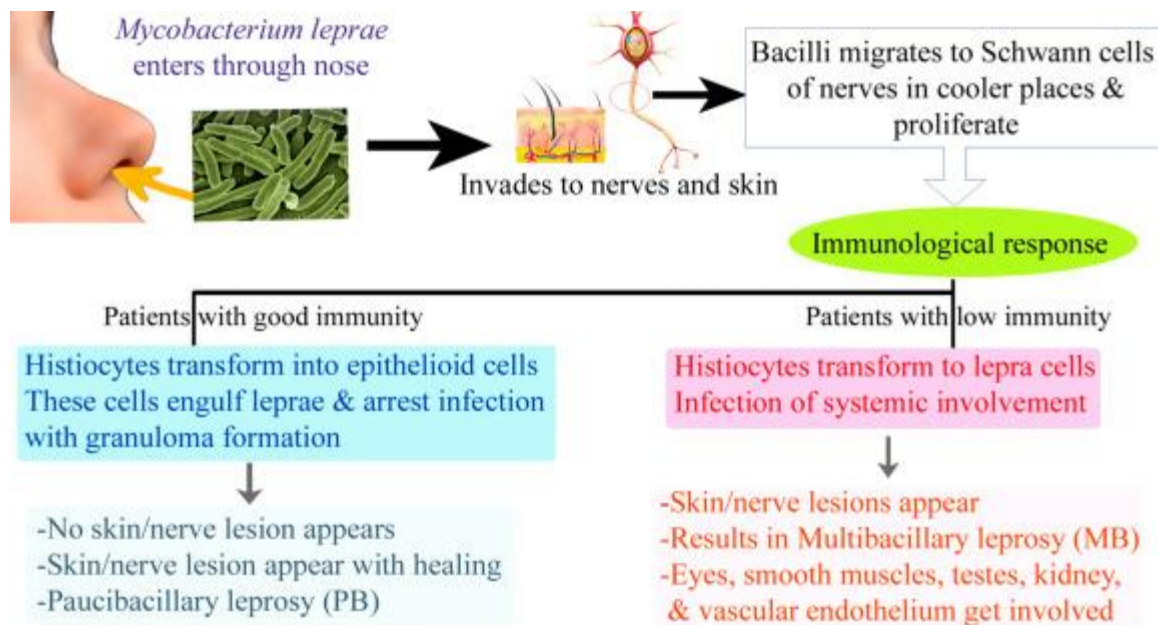
While the causative organisms have to date been impossible to culture in vitro, it has been possible to grow them in animals such as mice and armadillos.

Naturally occurring infection has been reported in nonhuman primates (including the African chimpanzee, the sooty mangabey, and the cynomolgus macaque), armadillos, and red squirrels. Multilocus sequence typing of the armadillo *M. leprae* strains suggests that they were of human origin for at most a few hundred years. Thus, it is suspected that armadillos first acquired the organism incidentally from early European explorers of the Americas. This incidental transmission was sustained in the armadillo population, and it may be transmitted back to humans, making leprosy a zoonotic disease (spread between humans and animals).

Red squirrels (*Sciurus vulgaris*), a threatened species in Great Britain, were found to carry leprosy in November 2016. It has been suggested that the trade in red squirrel fur, highly prized in the medieval period and intensively traded, may have been responsible for the leprosy epidemic in medieval Europe. A pre-Norman era skull excavated in Hoxne, Suffolk, in 2017 was found to carry DNA from a strain of *Mycobacterium leprae*, which closely matched the strain carried by modern red squirrels on Brownsea Island,

PATHOPHYSIOLOGY:-

In leprosy, the nerve is often thickened and involves areas proximal to the entrapment site. Motor weakness and wasting are often more severe in leprosy than in a carpal tunnel syndrome. Idiopathic entrapment neuropathy must not be misdiagnosed as neuritic leprosy (ie, leprosy without skin lesions).



Risk Factors

Those living in endemic areas with poor conditions such as inadequate bedding, contaminated water, and insufficient diet, or other diseases that compromise immune function are at highest risk for acquiring *M. leprae* infection. There has been concern that coinfection with HIV might exacerbate the pathogenesis of leprosy lesions and/or lead to increased susceptibility to leprosy as it is seen with tuberculosis. However, HIV infection has not been reported to increase susceptibility to leprosy, impact on immune response to *M. leprae*, or to have a significant effect on the pathogenesis of neural or skin lesions to date. On the contrary, initiation of antiretroviral treatment has been reported to be associated with activation of subclinical *M. leprae* infection and exacerbation of existing leprosy lesions (type I reaction) likely as part of immune reconstitution inflammatory syndrome.

The greatest risk factor for developing leprosy is contact with another person infected by leprosy. People who are exposed to a person who has leprosy are 5–8 times more likely to develop leprosy than members of the general population. Leprosy also occurs more commonly among those living in poverty. Not all people who are infected with *M. leprae* develop symptoms.

Conditions that reduce immune function, such as malnutrition, other illnesses, or genetic mutations, may increase the risk of developing leprosy. Infection with HIV does not appear to increase the risk of developing leprosy. Certain genetic factors in the person exposed have been associated with developing lepromatous or tuberculoid leprosy.

MANAGEMENT:-

A number of leprostatic agents are available for treatment. A three-drug regimen of rifampicin, dapsone and clofazimine is recommended for all people with leprosy, for six months for paucibacillary leprosy and 12 months for multibacillary leprosy.

Multidrug therapy (MDT) remains highly effective, and people are no longer infectious after the first monthly dose. It is safe and easy to use under field conditions because of its presentation in calendar blister packs. Post-treatment relapse rates remain low. Resistance has been reported in several countries, although the number of cases is small. People with rifampicin-resistant leprosy may be treated with second line drugs such as fluoroquinolones, minocycline, or clarithromycin, but the treatment duration is 24 months because of their lower bactericidal activity. Evidence on the potential benefits and harms of alternative regimens for drug-resistant leprosy is not available.

For people with nerve damage, protective footwear may help prevent ulcers and secondary infection. Canvas shoes may be better than PVC boots. There may be no difference between double rocker shoes and below-knee plaster. Topical ketanserin seems to have a better effect on ulcer healing than clioquinol cream or zinc paste, but the evidence for this is weak. Phenytoin applied to the skin improves skin changes to a greater degree when compared to saline dressings.

REFERENCES:-

1. Montoya D, Modlin RL (2010). Learning from leprosy: insight into the human innate immune response. *Advances in Immunology*. Vol. 105. pp. 1–24. doi:10.1016/S0065-2776(10)05001-7. ISBN 978-0-12-381302-2. PMID 20510728.
2. Byrne JP (2008). *Encyclopedia of pestilence, pandemics, and plagues*. Westport, Conn.[u.a.]: Greenwood Press. p. 351. ISBN 978-0-313-34102-1.
3. CDC (26 January 2018). "World Leprosy Day". Centers for Disease Control and Prevention. Archived from the original on 15 June 2019. Retrieved 4 July 2019.
4. "Global leprosy situation, 2012". *Weekly Epidemiological Record*. 87 (34): 317–328. August 2012. PMID 22919737.
5. Rodrigues LC, Lockwood DN (June 2011). "Leprosy now: epidemiology, progress, challenges, and research gaps". *The Lancet. Infectious Diseases*. 11 (6): 464–470. doi:10.1016/S1473-3099(11)70006-8. PMID 21616456.

6. "Hansen's Disease Data & Statistics". Health Resources and Services Administration. Archived from the original on 4 January 2015. Retrieved 12 January 2015.
7. "Central Florida is a hot spot for leprosy, report says". CNN. August 2023.
8. Walsh F (31 March 2007). "The hidden suffering of India's lepers". BBC News. Archived from the original on 29 May 2007.
9. Lyn TE (13 September 2006). "Ignorance breeds leper colonies in China". Independat News & Media. Archived from the original on 8 April 2010. Retrieved 31 January 2010.
10. Pisuthipan A (6 July 2020). "Forgotten victims of the virus". Bangkok Post. Archived from the original on 28 August 2021. Retrieved 6 July 2020.
11. Nunzi E, Massone C, eds. (2012). *Leprosy a practical guide*. Milan: Springer. p. 326. ISBN 978-88-470-2376-5. Archived from the original on 8 September 2017.
12. "Neglected Tropical Diseases". cdc.gov. 6 June 2011. Archived from the original on 4 December 2014. Retrieved 28 November 2014.
13. McMenamin D (2011). *Leprosy and stigma in the South Pacific: a region-by-region history with first person accounts*. Jefferson, N.C.: McFarland. p. 17. ISBN 978-0-7864-6323-7. Archived from the original on 19 May 2016.
14. "Signs and Symptoms | Hansen's Disease (Leprosy) | CDC". www.cdc.gov. 22 October 2018. Archived from the original on 22 July 2019. Retrieved 22 July 2019.
15. "Pathogenesis and Pathology of Leprosy". *International Textbook of Leprosy*. 11 February 2016. Archived from the original on 22 July 2019. Retrieved 22 July 2019.
16. WHO Expert Committee on Leprosy – Eight report (PDF). World Health Organization (WHO). 2012. pp. 11–12. ISBN 978-9241209687. Archived from the original (PDF) on 5 August 2013. Retrieved 9 May 2018.
17. Talhari C, Talhari S, Penna GO (2015). "Clinical aspects of leprosy". *Clinics in Dermatology*. 33 (1): 26–37. doi:10.1016/j.clindermatol.2014.07.002. PMID 25432808.
18. Kulkarni GS (2008). *Textbook of Orthopedics and Trauma (2nd ed.)*. Jaypee Brothers Publishers. p. 779. ISBN 978-81-8448-242-3.
19. "Q and A about leprosy". American Leprosy Missions. Archived from the original on 4 October 2012. Retrieved 22 January 2011. Do fingers and toes fall off when someone gets leprosy? No. The bacillus attacks nerve endings and destroys the body's ability to feel pain and injury. Without feeling pain, people injure themselves on fire, thorns, rocks, even hot coffee cups. Injuries become infected and result in tissue loss. Fingers and toes become shortened and deformed as the cartilage is absorbed into the body.
20. de Sousa JR, Sotto MN, Simões Quaresma JA (28 November 2017). "Leprosy As a Complex Infection: Breakdown of the Th1 and Th2 Immune Paradigm in the Immunopathogenesis of the Disease". *Frontiers in Immunology*. 8: 1635. doi:10.3389/fimmu.2017.01635. PMC 5712391. PMID 29234318.
21. Reinar LM, Forsetlund L, Lehman LF, Brurberg KG (July 2019). "Interventions for ulceration and other skin changes caused by nerve damage in leprosy". *The Cochrane Database of Systematic Reviews*. 2019 (7): CD012235. doi:10.1002/14651858.CD012235.pub2. PMC 6699662. PMID 31425632.
22. Ryan KU, Ray CJ, eds. (2004). *Sherris Medical Microbiology (4th ed.)*. McGraw Hill. pp. 451–53. ISBN 978-0-8385-8529-0. OCLC 61405904.

23. "Genomics Insights into the Biology and Evolution of Leprosy Bacilli". International Textbook of Leprosy. 11 February 2016. Archived from the original on 12 February 2019. Retrieved 11 February 2019.
24. McMurray DN (1996). "Mycobacteria and Nocardia". In Baron S; et al. (eds.). *Baron's Medical Microbiology* (4th ed.). Univ of Texas Medical Branch. ISBN 978-0-9631172-1-2. OCLC 33838234. Archived from the original on 12 February 2009.
25. Bhattacharya S, Vijayalakshmi N, Parija SC (October 2002). "Uncultivable bacteria: implications and recent trends towards identification". *Indian Journal of Medical Microbiology*. 20 (4): 174–177. doi:10.1016/S0255-0857(21)03184-4. PMID 17657065.
26. "WHO | Microbiology: culture in vitro". World Health Organization (WHO). Archived from the original on 9 August 2020. Retrieved 22 July 2019.
27. "The Armadillo Model for Leprosy". International Textbook of Leprosy. 11 February 2016. Archived from the original on 22 July 2019. Retrieved 22 July 2019.
28. Loughry WJ, Truman RW, McDonough CM, Tilak MK, Garnier S, et al. (2009) "Is leprosy spreading among nine-banded armadillos in the southeastern United States?" *J Wildl Dis* 45: 144–52.
29. Meredith A, Del Pozo J, Smith S, Milne E, Stevenson K, McLuckie J (September 2014). "Leprosy in red squirrels in Scotland". *The Veterinary Record*. 175 (11): 285–286. doi:10.1136/vr.g5680. PMID 25234460. S2CID 207046489.
30. Monot M, Honoré N, Garnier T, Araoz R, Coppee JY, et al. (2005). "On the origin of leprosy". *Science* 308: 1040–42.
31. Han XY, Silva FJ (February 2014). "On the age of leprosy". *PLOS Neglected Tropical Diseases*. 8 (2): e2544. doi:10.1371/journal.pntd.0002544. PMC 3923669. PMID 24551248.
32. "Red squirrels in the British Isles are infected with leprosy bacilli" Archived 12 June 2022 at the Wayback Machine, Dr. Andrej Benjak, Prof Anna Meredith and others. *Science*, 11 November 2016. [1] Archived 12 June 2022 at the Wayback Machine. Retrieved 11 November 2016.
33. "Could squirrel fur trade have contributed to England's medieval leprosy outbreak?". *ScienceDaily*. Archived from the original on 22 November 2018. Retrieved 21 November 2018.
34. Inskip S, Taylor GM, Anderson S, Stewart G (November 2017). "Leprosy in pre-Norman Suffolk, UK: biomolecular and geochemical analysis of the woman from Hoxne". *Journal of Medical Microbiology*. 66 (11): 1640–1649. doi:10.1099/jmm.0.000606. PMID 28984227. S2CID 33997231.
35. Penna ML, Penna GO, Iglesias PC, Natal S, Rodrigues LC (May 2016). "Anti-PGL-1 Positivity as a Risk Marker for the Development of Leprosy among Contacts of Leprosy Cases: Systematic Review and Meta-analysis". *PLOS Neglected Tropical Diseases*. 10 (5): e0004703. doi:10.1371/journal.pntd.0004703. PMC 4871561. PMID 27192199.
36. Alcaïs A, Mira M, Casanova JL, Schurr E, Abel L (February 2005). "Genetic dissection of immunity in leprosy". *Current Opinion in Immunology*. 17 (1): 44–48. doi:10.1016/j.coi.2004.11.006. PMID 15653309.
37. Lockwood DN, Lambert SM (January 2011). "Human immunodeficiency virus and leprosy: an update". *Dermatologic Clinics*. 29 (1): 125–128. doi:10.1016/j.det.2010.08.016. PMID 21095536.
38. "Epidemiology of Leprosy". International Textbook of Leprosy. 11 February 2016. Archived from the original on 23 July 2019. Retrieved 30 July 2019.

39. Chavarro-Portillo B, Soto CY, Guerrero MI (September 2019). "Mycobacterium leprae's evolution and environmental adaptation". *Acta Tropica*. 197: 105041. doi:10.1016/j.actatropica.2019.105041. PMID 31152726. S2CID 173188912.
40. Eichelmann K, González González SE, Salas-Alanis JC, Ocampo-Candiani J (September 2013). "Leprosy. An update: definition, pathogenesis, classification, diagnosis, and treatment". *Actas Dermo-Sifiliograficas*. 104 (7): 554–563. doi:10.1016/j.adengl.2012.03.028. PMID 23870850. S2CID 3442319.
41. Joel Carlos Lastória JC, Milanez Morgado de Abreu MA (March–April 2014). "Leprosy: review of the epidemiological, clinical, and etiopathogenic aspects - Part 1". *An Bras Dermatol*. 89 (2): 205–218. doi:10.1590/abd1806-4841.20142450. PMC 4008049. PMID 24770495.
42. "Hansen's Disease (Leprosy) Transmission". cdc.gov. 29 April 2013. Archived from the original on 13 March 2015. Retrieved 28 February 2015.
43. Lockwood DN, Kumar B (June 2004). "Treatment of leprosy". *BMJ*. 328 (7454): 1447–1448. doi:10.1136/bmj.328.7454.1447. PMC 428501. PMID 15205269.
44. "What Is Leprosy?" | from News-Medical.Net – Latest Medical News and Research from Around the World. Web. 20 Nov. 2010. "What is Leprosy?". News-Medical.net. 18 November 2009. Archived from the original on 6 June 2013. Retrieved 14 May 2013..
45. Truman RW, Singh P, Sharma R, Busso P, Rougemont J, Paniz-Mondolfi A, et al. (April 2011). "Probable zoonotic leprosy in the southern United States". *The New England Journal of Medicine*. 364 (17): 1626–1633. doi:10.1056/NEJMoa1010536. PMC 3138484. PMID 21524213.
46. "Hansen's Disease (Leprosy) Transmission". cdc.gov. 29 April 2013. Archived from the original on 13 March 2015. Retrieved 28 February 2015.
47. Cambri G, Mira MT (20 July 2018). "Genetic Susceptibility to Leprosy-From Classic Immune-Related Candidate Genes to Hypothesis-Free, Whole Genome Approaches". *Frontiers in Immunology*. 9: 1674. doi:10.3389/fimmu.2018.01674. PMC 6062607. PMID 30079069.
48. Cook GC (2009). *Manson's tropical diseases* (22nd ed.). [Edinburgh]: Saunders. p. 1056. ISBN 978-1-4160-4470-3. Archived from the original on 4 September 2017.
49. Buschman E, Skamene E (June 2004). "Linkage of leprosy susceptibility to Parkinson's disease genes". *International Journal of Leprosy and Other Mycobacterial Diseases*. 72 (2): 169–170. doi:10.1489/1544-581X(2004)072<0169:LOLSTP>2.0.CO;2 (inactive 31 January 2024). PMID 15301585. S2CID 43103579.
50. Bhat RM, Prakash C (2012). "Leprosy: an overview of pathophysiology". *Interdisciplinary Perspectives on Infectious Diseases*. 2012: 181089. doi:10.1155/2012/181089. PMC 3440852. PMID 22988457.
51. Moschella SL, Garcia-Albea V (September 2016). "International Textbook of Leprosy" (PDF). *Differential Diagnosis of Leprosy*. p. 3, Section 2.3. Archived (PDF) from the original on 16 July 2020. Retrieved 4 July 2019.
52. U.S. Department of Health and Human Services, Health Resources and Services Administration. (n.d.). National Hansen's disease (leprosy) program Retrieved from "National Hansen's Disease (Leprosy) Program". Archived from the original on 10 February 2011. Retrieved 12 May 2013.
53. Martinez AN, Talhari C, Moraes MO, Talhari S (April 2014). "PCR-based techniques for leprosy diagnosis: from the laboratory to the clinic". *PLOS Neglected Tropical Diseases*. 8 (4): e2655. doi:10.1371/journal.pntd.0002655. PMC 3983108. PMID 24722358.

54. Tatipally S, Srikantam A, Kasetty S (October 2018). "Polymerase Chain Reaction (PCR) as a Potential Point of Care Laboratory Test for Leprosy Diagnosis-A Systematic Review". *Tropical Medicine and Infectious Disease*. 3 (4): 107. doi:10.3390/tropicalmed3040107. PMC 6306935. PMID 30275432.
55. Smith DS (19 August 2008). "Leprosy: Overview". *eMedicine Infectious Diseases*. Archived from the original on 18 February 2010. Retrieved 1 February 2010.
56. Singh N, Manucha V, Bhattacharya SN, Arora VK, Bhatia A (June 2004). "Pitfalls in the cytological classification of borderline leprosy in the Ridley-Jopling scale". *Diagnostic Cytopathology*. 30 (6): 386–388. doi:10.1002/dc.20012. PMID 15176024. S2CID 29757876.
57. Ridley DS, Jopling WH (1966). "Classification of leprosy according to immunity. A five-group system". *International Journal of Leprosy and Other Mycobacterial Diseases*. 34 (3): 255–273. PMID 5950347.
58. James WD, Berger TG, Elston DM, Odom RB (2006). *Andrews' Diseases of the Skin: clinical Dermatology*. Saunders Elsevier. pp. 344–46. ISBN 978-0-7216-2921-6.
59. Lastória JC, Abreu MA (2014). "Leprosy: a review of laboratory and therapeutic aspects--part 2". *Anais Brasileiros de Dermatologia*. 89 (3): 389–401. doi:10.1590/abd1806-4841.20142460. PMC 4056695. PMID 24937811.
60. Kumar, Bhushan; Uprety, Shraddha; Dogra, Sunil (11 February 2016). "Clinical Diagnosis of Leprosy". *International Textbook of Leprosy*. Archived from the original on 13 February 2019. Retrieved 12 February 2019.