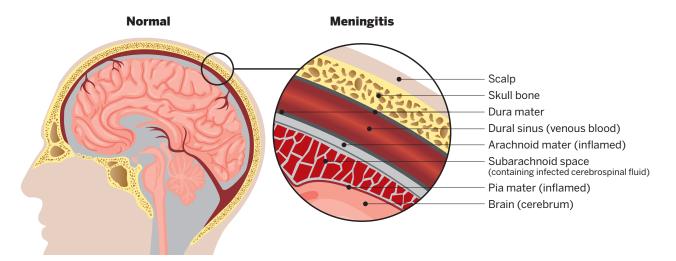




KEY FACTS ABOUT MENINGITIS^{1,2,3,4}

- Meningitis is a **severe infection of the central nervous system**. It occurs when bacteria, viruses, or fungi infect the meninges, or tissue membranes that cover the brain and spinal cord. This causes inflammation of the membranes and accumulation of purulence within the cerebrospinal fluid (see illustration below).
- Meningitis most commonly affects young children, the elderly, and immunocompromised people.
- Viral meningitis is the most common form and is usually a self-limited illness, although some viruses do respond to specific antiviral treatment.
- Bacterial or fungal meningitis is a medical emergency which, if not treated rapidly, can lead to irreversible brain damage and death.
- Meningococcal meningitis, a rapidly progressing form of community-acquired meningitis caused by *Neisseria meningitidis*, also has an elevated risk in adolescents and young adults in group settings, such as college dormitories, military barracks, or refugee camps and also in certain geographic regions such as sub-Saharan Africa.



THE BURDEN OF MENINGITIS 5,6,7,8,9,10

- Meningitis represents a heavy health burden for both patients and society.
- It is especially prevalent in an area of sub-Saharan Africa known as the "**meningitis belt**" that stretches across 26 countries from Senegal to Ethiopia.

GLOBAL BURDEN

- 2.5 million cases per year
 0.7-0.9 per 100,000 in US/Europe
- 250,000 deaths per year
- ~50% of cases and deaths in children <5 years old
- 1 in 5 survivors have permanent disability such as cognitive impairment, hearing loss, motor weakness or paralysis, incoordination, and epilepsy

BURDEN IN AFRICA

- 30,000 cases per year
 10 40 per 100,000
 - Epidemics of meningococcal meningitis up to 250,000 cases per year
- 2,000 deaths per year (up to 25,000 during epidemics)
- Many cases go undiagnosed





CAUSES AND CLINICAL PRESENTATION OF MENINGITIS 1,2,3,10

- Meningitis may be due to many different viruses, bacteria, mycobacteria, fungi and, less commonly, parasites.
- The most likely pathogen depends on several factors including a patient's age, immune and vaccination status, and geographic location.
- However, four bacteria are responsible for more than half of the deaths from meningitis globally.
- They are:
 - Neisseria meningitidis (meningococcus)
 - Streptococcus pneumoniae (pneumococcus)
 - Haemophilus influenzae type B (Hib)
 - Streptococcus agalactiae (group B streptococcus).

MOST COMMON SIGNS AND SYMPTOMS

NEWBORNS AND INFANTS

- High fever
- Constant crying
- Excessive irritability
- Lethargy
- A bulging fontanel
- Hyper/hypotonia
- · Poor appetite (refusing food or drink)
- Mottled skin

PATIENTS OLDER THAN 2 YEARS

- Sudden high fever
- Severe headache
- Stiff neck
- Nausea and/or vomiting
- Confusion or difficulty concentrating
- Decreased level of consciousness
- Seizures
- Photophobia
- · Petechial rash for older patients

A physical exam may produce a positive Kernig's and/or Brudzinski's signs





DIAGNOSTIC APPROACH 11,12,13,14

- Many guidelines for the diagnosis and management of meningitis are available. They may have some differences based on regional specifics.
- **Blood tests** provide information about the infection and the patient. A **blood culture** may be positive in up to 50% of meningitis cases.
- **A head CT**, when available, is recommended to rule out increased intracranial pressure for some patients. A CT or MRI may also be used to assess the damage to the brain or spinal cord.
- Whenever safely possible, **cerebrospinal fluid (CSF)** should be obtained via a **lumbar puncture** for microscopic and biochemical analysis as well as **pathogen identification** and **potential antimicrobial sensitivity testing**.
- Molecular diagnostics such as multiplex PCR are emerging as new and powerful diagnostic tools. The molecular syndromic testing approach enables rapid accurate identification of multiple target pathogens. This may be possible even when the patient has been pre-treated with antibiotics.



* Empiric therapy should be started prior to a head CT and lumbar puncture if there are delays in performing these investigations. ** ID/AST: identification and antimicrobial susceptibility testing.



TREATMENT STRATEGIES^{11,12,13,14}

- Parenteral antibiotics should be given within one hour when bacterial meningitis is suspected.
- Appropriate antimicrobial therapy should not be postponed for diagnostic delays, such as when a head CT is needed but cannot be obtained quickly.
- Antimycobacterial regimens for tuberculosis, antifungal medication for cryptococcus or antivirals such as acyclovir for herpes simplex virus and varicella zoster virus infection are given when there is an increased suspicion for these pathogens.
- Ancillary treatments such as steroids and strict fluid management to reduce inflammation and cerebral edema are sometimes used, but their usefulness is limited.

PREVENTION 14,15,16

- Vaccines have led to substantial declines in many of the most common causes of bacterial meningitis including *Streptococcus pneumoniae*, *Haemophilus influenzae* type b, and *Neisseria meningitidis*.
- Historically, these vaccines have only been largely available in more developed countries and regions.
- However, recently a mass immunization program using a **conjugate vaccine against group A meningococcus**, which has caused >80% of epidemics in the African meningitis belt, is having a significant and dramatic impact in breaking the cycle of these epidemics.
- Where feasible, **close contacts of cases of meningococcus meningitis** may be given **chemoprophylaxis** with an appropriate antibiotic in order to prevent epidemic spread.

References:

- 1. WHO. Meningitis Fact Sheet. https://www.who.int/news-room/fact-sheets/detail/meningitis Accessed on September 29, 2023.
- FCCMG Meningitis Blog. https://www.fccmg.com/blog/meningitis-symptoms-treatments/ Accessed on September 29, 2023.
- NINDS. Meningitis and Encephalitis Fact Sheet. https://www.ninds.nih.gov/health-information/ disorders/meningitis Accessed on September 29, 2023.
- CDC. Meningococcal Disease. https://www.cdc.gov/meningococcal/about/risk-factors.html Accessed on September 29, 2023.
- GBD 2019 Meningitis and Antimicrobial Resistance Collaborators. Global, regional, and national burden of meningitis and its aetiologies, 1990–2019: a systematic analysis for the Global Burden of Disease Study 2019. Lancet Neurol. 2023;22(8):685–711. https://doi.org/10.1016/S1474-4422(23)00195-3
- 6. Barichello T, Rocha Catalão CH, Rohlwink UK, *et al.* Bacterial meningitis in Africa. *Front Neurol.* 2023;14:822575 https://doi.org/10.3389/fneur.2023.822575
- GBD 2016 Causes of Death Collaborators. Global, regional, and national age-sex specific mortality for 264 causes of death, 1980–2016: a systematic analysis for the Global Burden of Disease Study 2016. Lancet Neurol. 2017;390 1151-1210. https://doi.org/10.1016/S0140-6736(17)32152-9
- Brouwer MC, van de Beek D. Epidemiology of community-acquired bacterial meningitis. Curr Opin Infect Dis. 2018;31(1):78-84. https://doi.org/10.1097/qco.000000000000417
- Mazamay S, Guégan JF, Diallo N, et al. An overview of bacterial meningitis epidemics in Africa from 1928 to 2018 with a focus on epidemics "outside- the-belt". BMC Infect Dis. 2021;21:1027 https://doi.org/10.1186/s12879-021-06724-1

- Wright C, Blake N, Glennie L, et al. The Global Burden of Meningitis in Children: Challenges with Interpreting Global Health Estimates. *Microorganisms*. 2021; 9(2):377 https://doi.org/10.3390/microorganisms9020377
- Tunkel RA, Hasbun R, Bhimraj A, et al. 2017 Infectious Diseases Society of America's Clinical Practice Guidelines for Healthcare-Associated Ventriculitis and Meningitis, *Clin Infect Dis.* 2017;64(6):e34–e65 https://doi.org/10.1093/cid/ciw861
- 12. van de Beek D, Cabellos C, Dzupova O, *et al.* ESCMID Study Group for Infections of the Brain (ESGIB). ESCMID guideline: diagnosis and treatment of acute bacterial meningitis. *Clin Microbiol Infect.* 2016;22 Suppl 3:S37-62 https://doi.org/10.1016/j.cmi.2016.01.007
- Bremer M, Kadernani YE, Wasserman S, et al. Strategies for the diagnosis and management of meningitis in HIV-infected adults in resource limited settings. Expert Opin. Pharmacother. 2021;22(15):2053-2070 https://doi.org/10.1080/14656566.2021.1940954
- Govender I, Steyn C, Maricowitz G, et al. (2018) A primary care physician's approach to a child with meningitis. S Afr J Infect Dis. 2018;33(2):31-37 https://doi.org/10.1080/23120053.2017.1394610
- World Health Organization. Defeating meningitis by 2030: baseline situation analysis. Feb, 2019. https://www.who.int/publications/m/item/defeating-meningitis-2030-baseline-situation-analysis Accessed on September 29, 2023.
- Stuart J. Impact of serogroup A meningococcal conjugate vaccine for Africa. Hum Vaccin Immunother. 2018;14(5):1116–1117 https://doi.org/10.1080/21645515.2017.1412022

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