

6. Incubation period—From 1–7 days (mean 5 days) in documented cases of *Naegleria* infection; usually longer in infections of *Acanthamoeba* and *Balamuthia*.

7. Transmission—*Naegleria* occurs through exposure of the nose to contaminated water, most commonly by diving or swimming in warm freshwater, especially ponds or lakes in warm climate areas or during summer, thermal springs or bodies of water warmed by the effluvia of industrial plants, or inadequately maintained swimming pools. *Naegleria* has also occurred following direct nasal irrigation with contaminated water with solutions made from contaminated water. After entering the nose, *Naegleria* trophontes invade brain and meninges by extension along olfactory nerves. *Acanthamoeba* and *Balamuthia* trophontes reach the CNS through hematogenous spread, probably from a skin lesion or other site of primary colonization, such as the respiratory tract. *Acanthamoeba* frequently occurs in chronically ill or immunosuppressed patients with a history of swimming or known source of infection. *Balamuthia* also occurs in such patients but develops in immunocompetent patients as well. Additionally, transmission of *Balamuthia* through solid organ transplant has been documented in the USA on 3 separate occasions since 2009. Contact lens use, homemade saline used as a cleaning or wetting solution, a broad-spectrum contact lens solution no longer on the market, and contact lens exposure to water have been implicated as risk factors for non-*Acanthamoeba* infection. Other than solid organ transplant transmission, no person-to-person transmission has been observed with any of the living amebae.

8. Risk groups—*Naegleria* occurs mainly in active immunocompetent children and young men and women who have had recent contact with warm, fresh water.

Transmission of *Acanthamoeba* and *Balamuthia*, although balamuthia occurs in immunocompetent persons as well and appears to occur more frequently in persons of Hispanic ethnicity. Eye infections associated with *Acanthamoeba* have occurred primarily in contact lens wearers.

9. Prevention—

1) PAM and swimming: the only sure way to prevent warm-water associated PAM is to refrain from swimming and other water-related activities in warm freshwater. Infections may be reduced by educating the public about the risks of swimming in lakes or ponds during periods of warm water temperatures (particularly where infection is known or presumed to have been acquired) or of allowing such water to go up the nose through diving or underwater swimming. Protect the nasopharynx from exposure

water likely to contain *N. fowleri* (e.g., through the use of nose-clips). In practice, this is difficult, since the amebae may occur in a wide variety of aquatic bodies, including inadequately treated swimming pools. Swimming pools containing residual free chlorine of 1–2 ppm are considered safe. No infection is known to have been acquired in a well-maintained chlorinated swimming pool.

- 2) PAM and nasal irrigation: solutions used for nasal irrigation should be made with boiled, distilled, or sterile water.
- 3) *Acanthamoeba* keratitis: contact lens wearers should not wear lenses while bathing, showering, or swimming or while using hot tubs and should follow strictly the wear and care procedures recommended by lens manufacturers and health care professionals.

10. Management of patient—Despite the sensitivity of the organism to antibiotics in laboratory studies, effective treatment for PAM and GAE has not been established and recoveries have been rare. Recovery from PAM occurred in two case patients (one each from the USA and Mexico) following intravenous (with and without intrathecal) administration of amphotericin B at various dosages, oral rifampicin, and an intravenous azole (miconazole or flucytosine) along with dexamethasone to treat cerebral edema. *N. fowleri* is also sensitive to aztreonam, which could be added to the other medications. GAE recoveries have occurred with varying combinations of pentamidine, sulfadiazine, flucytosine, flucosazole, and miltefosine. Like *N. fowleri*, *Balamuthia* is sensitive to aztreonam, which has also been successfully added to the drug cocktail. For eye infections due to *Acanthamoeba*, topical chlorhexidine and/or polyhexamethylene biguanide have been used successfully, often with the addition of the diamidine griseofulvin, or hexamidine.

11. Management of contacts and the immediate environment—A history of swimming or introducing water into the nose (e.g., nasal irrigation) within 1 week prior to the onset of PAM symptoms may suggest the source of infection. Sources of *Acanthamoeba* and *Balamuthia* infection are less well understood but a history of soil exposure (either recreationally or occupationally) is sometimes elicited from GAE patients; freshwater exposure might also be a source of infection.

12. Special considerations—Any grouping of cases warrants prompt epidemiological investigation. If GAE occurs in a recent transplant patient, investigation of the donor and other organ recipients from the same donor should be initiated.

[S. Roy, J. Voder, M. Beach, G. Vlachouli]



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