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## **Future priorities in tackling infections due to brain-eating amoebae**

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21 **Abstract**

22 Brain-eating amoebae (*Acanthamoeba* spp., *Balamuthia mandrillaris* and *Naegleria*  
23 *fowleri*) can cause opportunistic infections involving the central nervous system. It is  
24 troubling that the mortality rate is more than 90% despite advances in antimicrobial  
25 chemotherapy over the last few decades. Here, we describe urgent key priorities for  
26 improving outcomes from infections due to brain-eating amoebae.

27 **Dear Editor**

28 Whilst brain infections due to pathogenic free-living amoebae are rare, the mortality  
29 remains very high leading almost always to death.<sup>1-4</sup> Defining the global burden of infections  
30 due to brain-eating amoebae presents a major challenge, as infections are rare but insidious in  
31 nature leading to inherent difficulty in their diagnosis due to a global lack of capacity for  
32 diagnostics especially in developing countries. Lack of effective drugs and/or their delivery  
33 to the site of infection results in mortality rate of more than 95%, highlighting global failure  
34 in tackling this infection over the past several decades. Despite exceptionally high mortality  
35 rate, brain-eating amoebae have not had the expected level of focus from the global  
36 community. There is a need for renewed efforts for:

- 37 (i) Better epidemiology data involving collaborative efforts between basic  
38 scientists and clinical researchers to accelerate translational medicine.
- 39 (ii) Improved laboratory and point-of-care testing. It is obvious that, without  
40 point-of-care testing, these infections will remain difficult to diagnose, and  
41 treat, and their true global burden will remain undetermined.
- 42 (iii) Better access to drugs. Access to established medicines, as well as  
43 development of new medicines. Access, in particular to Miltefosine is  
44 particularly, and liposomal amphotericin B (Ambisome) remains very

45 expensive in many countries. Acceleration of vaccination programmes should  
46 be a key priority, but will be challenging due to the rarity of the disease.

47 (iv) Capacity building for pathogenic free-living amoebae. Whilst there are several  
48 groups working in the area of brain-eating amoebae, better cohesion and  
49 extension within basic scientists and practicing physicians will enable more  
50 rapid progress in this area.

51 (v) Funding for development of diagnosis, treatment strategies, and  
52 implementation programmes, especially in resource-limited settings. In this  
53 regard, establishment of advocacy groups and public engagement will lead to  
54 infrastructure development programme for disease surveillance and to devise  
55 treatment strategies.

56 (vi) Fundamental research in genomics-based studies of amoebal evolution,  
57 parasite-host interactions, and resistance in the host including metabolic  
58 adaptation and understanding the innate and acquired immune responses  
59 remain priority areas.

60 Although there are some encouraging novel therapies on the horizon including intranasal  
61 delivery of anti-amoebic molecules to bypass blood-brain barrier selectivity,<sup>5</sup> there is an  
62 urgent need in delivering novel diagnostic and therapeutic strategies to limit mortality from  
63 these infections. However, engagement of major funding bodies and governmental and non-  
64 governmental agencies is needed to enable substantial reductions in the unacceptably high  
65 mortality from infections due to brain-eating amoebae.

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