Cryptococcus: Screening for Opportunistic Infection among People Living with HIV/ AIDS





National Center for Emerging and Zoonotic Infectious Diseases Division of Foodborne, Waterborne, and Environmental Diseases

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Cryptococcal Infection

What is Cryptococcus?

Cryptococcus is a fungus that is found in the soil throughout the world. Because *Cryptococcus* is common in the environment, most people probably breathe in small amounts of microscopic, airborne spores every day. Sometimes these spores cause symptoms of a respiratory infection, but other times there are no symptoms at all. In healthy people, the fungus usually does not cause serious illness because the immune system can fight off the infection. However, in people with weakened immune systems, the fungus can stay hidden in the body and later reactivate, spreading to other parts of the body and causing serious disease.

Cryptococcal meningitis

An infection with the fungus *Cryptococcus* is known as cryptococcosis, and it is a serious opportunistic infection among people with advanced HIV/AIDS. Cryptococcosis is not contagious, meaning it cannot spread from person-to-person. Cryptococcal meningitis specifically occurs after *Cryptococcus* has spread from the lungs to the brain. Meningitis can also be caused by a variety of other organisms, including bacteria, viruses, and other fungi.

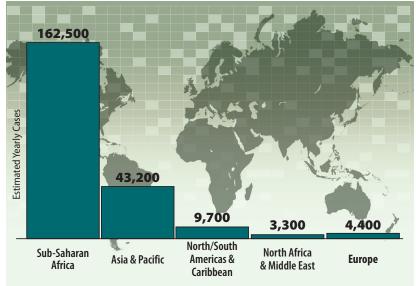
Symptoms of meningitis include:

- Headache
- Fever
- Neck pain
- Nausea and vomiting
- Sensitivity to light
- Altered mental status (ranging from confusion to coma)



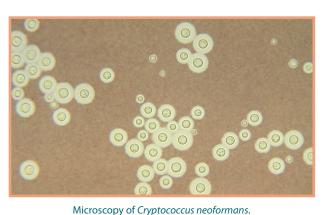
Worldwide, an estimated 220,000 new cases of cryptococcal meningitis occur each year, resulting in 181,000 deaths. Most cases are opportunistic infections that occur among people with HIV/AIDS. Although the widespread availability of antiretroviral therapy (ART) in developed countries has helped reduce cryptococcal infections in these areas, it is still a major problem in developing countries where access to healthcare is limited. Throughout much of sub-Saharan Africa, for example, Cryptococcus is now the most common cause of meningitis. Cryptococcal meningitis is one of the leading causes of death in HIV/AIDS patients; in sub-Saharan Africa, it may kill more people each year than tuberculosis.

The global burden of cryptococcal meningitis









Page 2 of 4



Preventing Deaths Due to Cryptococcus



Targeted screening: a cost-effective strategy

One approach to prevent deaths is known as "targeted screening." Research suggests that *Cryptococcus* is able to live in the body undetected, especially when a person's immune system is weaker than normal. In a targeted screening program, HIV-infected patients are tested for cryptococcal antigen, an indicator of cryptococcal infection, before starting ART. A patient who tests positive for cryptococcal antigen can take oral fluconazole to help the body fight the early stage of the infection. This could prevent the infection from developing into meningitis.

A simple new test for Cryptococcus

A new "dipstick" test for detecting cryptococcal antigen is simple to use on a small sample of serum (a component of blood). The test accurately detects both early and advanced cryptococcal infections more than 95% of the time. In addition, the test is inexpensive, and the results are ready in just 10 minutes. Using this test, targeted screening is a cost-effective approach to prevent deaths from cryptococcal meningitis.





Access to essential care

In order to screen patients for cryptococcal disease, as well as diagnose other patients with cryptococcal meningitis, healthcare facilities must have access to diagnostic tests. Currently, these tests are unavailable in most district and provincial laboratories in sub-Saharan Africa. Equipping these facilities with the ability to perform the new dipstick test is an important step in combating cryptococcal meningitis.



Essential medications for the treatment of cryptococcal disease are also lacking in areas of the world that are most in need. Amphotericin B and flucytosine, antifungal medications that have been shown to improve survival in patients with advanced disease and are the standard-of-care in developed countries, are widely unavailable in sub-Saharan Africa.

Integration with HIV/AIDS care and treatment

Like co-trimoxazole prophylaxis and isoniazid preventive therapy (IPT), cryptococcal screening and treatment of antigenemia with oral fluconazole can be part of an integrated care and treatment strategy for people living with HIV/AIDS. In areas where cryptococcal disease prevalence is high, an integrated approach can help reduce early mortality among HIV/AIDS patients starting ART.

CDC's response

CDC is focusing its efforts on countries in sub-Saharan Africa and Southeast Asia, the areas with the highest number of deaths due to cryptococcal disease. In these regions, CDC is helping build laboratory capacity to reliably diagnose cryptococcal infections and is assisting with the implementation of targeted cryptococcal screening programs. In the future, it is hopeful that diagnostic tests will be available even in remote, rural areas.

CDC's call to action is to equip half of all HIV clinics in Africa and Asia to perform cryptococcal screening and treatment, which could save 50,000 to 100,000 lives every year. Early identification of cryptococcalinfected patients in resource-limited settings may lead to more timely treatment, reduced mortality due to cryptococcal meningitis, and overall improved quality of life.

What You Can Do

Physicians and other healthcare providers can:

- Familiarize themselves with the World Health Organization's most upto-date guidelines on the diagnosis, prevention, and management of cryptococcal disease in HIV-infected individuals, available on the web at: <u>http://whqlibdoc.who.int/publications/2011/9789241502979_eng.pdf</u>
- Be aware of the benefits of targeted screening for cryptococcal antigenemia among patients at risk
- Advocate to implement cryptococcal screening programs at clinics where the prevalence of HIV/AIDS is high
- Teach at-risk patients about *Cryptococcus* and the importance of screening for cryptococcal antigenemia
- Advocate for increased availability of antifungal treatment and cryptococcal diagnostic capabilities in resource-limited settings
- Visit <u>http://preventcrypto.org/</u> to learn more about how to prevent deaths due to *Cryptococcus*

Non-governmental organizations (NGOs) can:

- Partner with healthcare facilities to consider implementing targeted cryptococcal screening programs in areas of the world where the prevalence of HIV/AIDS is high
- Get involved with monitoring and evaluation of existing cryptococcal screening programs
- Advocate for increased availability of antifungal treatment and cryptococcal diagnostic capabilities in resource-limited settings

Government organizations can:

- Create policies that support the development of sustainable cryptococcal screening programs
- Ensure adequate access to antifungal medications and cryptococcal diagnostic capabilities through partnerships and contracts with pharmaceutical companies







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Page 4 of 4