

SPECIAL REPORT

Twenty Years of Global Surveillance of Antituberculosis-Drug Resistance

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Antimicrobial resistance represents a major threat to global health and security. In 2014, the World Health Assembly called on all nations and the international community to take every necessary measure to control it, including surveillance of its emergence and spread.¹

The development of drug resistance in *Mycobacterium tuberculosis* was first documented in the late 1940s, soon after antibiotic therapy was introduced for tuberculosis treatment.² It quickly became obvious that combination chemotherapy could prevent the emergence of drug resistance³ and that patients infected with drug-resistant strains were less likely to be cured.⁴ Nevertheless, it was only in the early 1990s that drug-resistant tuberculosis began to receive global attention as a public health threat. This coincided with the detection of outbreaks of multidrug-resistant (MDR) tuberculosis (defined as resistance to at least rifampin and isoniazid) that were associated with high mortality among patients coinfecting with the human immunodeficiency virus (HIV).^{5–8} The urgent need for a global mechanism to monitor the emergence and spread of resistance to antituberculosis drugs became clear.

In 1994, the Global Tuberculosis Program of the World Health Organization (WHO), with the support of the International Union against Tuberculosis and Lung Disease (the Union), established the Global Project on Anti-Tuberculosis Drug Resistance Surveillance (hereafter referred to as “the project”) to measure the magnitude of drug resistance and to monitor trends. This project remains the oldest and largest initiative on the surveillance of antimicrobial resistance in the world.⁹ In this article, we describe the history

of global surveillance of drug resistance in tuberculosis and discuss methods for surveillance, the quality of available data, the key achievements and findings to date, the main challenges that remain, and future directions.

HISTORY OF GLOBAL SURVEILLANCE OF RESISTANCE TO ANTITUBERCULOSIS DRUGS

Throughout its existence, the project has been hosted by the WHO and supported by the TB Supranational Reference Laboratory (SRL) Network and several technical agencies. Funding has been continuously provided by the U.S. Agency for International Development.

The foundations of the project were laid in 1993 and 1994. A first meeting of interested partners was organized at the annual international conference of the Union in October 1993. Investigators from the Dominican Republic, Kenya, and Zimbabwe committed to implement the first drug-resistance surveys using a standardized approach. In June 1994, at a meeting organized by the WHO and the Union in Mainz, Germany, a group of 14 laboratories established the SRL Network to conduct and ensure quality-assured testing of drug susceptibility, under the leadership of the Laboratory Centre for Disease Control in Ottawa. Meanwhile, the WHO issued guidelines on standardized methods for conducting surveys of antituberculosis-drug resistance, which have been regularly updated.^{10–14} Surveillance data are collected either through continuous surveillance systems based on routine testing of all patients with tuberculosis or periodic surveys, which are discrete studies measuring

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