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(Research/Review) Article

The Role of Epidemiologic Surveillance in the Control of Drug-Resistant Tuberculosis (TB): A Literature Review

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Abstract: Drug-resistant tuberculosis (DR-TB) poses a serious threat to global public health and is a major barrier to achieving TB elimination targets. Epidemiological surveillance plays a central role in its control, yet its implementation faces various challenges, leading to gaps in detection and response. This literature review aims to synthesize the latest scientific evidence on the role, methods, and challenges of DR-TB surveillance to formulate a comprehensive overview as a basis for program improvement recommendations. A systematic literature search was conducted in PubMed and Google Scholar for articles published between 2015 and 2025. From an initial 347 articles, 6 relevant studies from various countries were selected based on inclusion and exclusion criteria for narrative analysis. The analysis reveals a significant gap between the estimated disease burden and detected cases, with the sensitivity of surveillance systems in some countries (e.g., Brazil) reported as low (~46.4%) and massive diagnostic failures (59% in Madagascar). Strong evidence indicates that active community transmission is a key driver of the epidemic, demonstrated by high primary resistance rates (9% in Bhutan) and the presence of genetic transmission clusters (37% of cases in Pará, Brazil). The studies also highlight the potential of innovative methods such as statistical correction to improve estimation accuracy and genomic surveillance for detecting new drug resistance and mapping transmission. Conventional DR-TB surveillance is no longer adequate to address the current complexity of the epidemic. Future effective control demands a dual approach: strengthening fundamental health systems (diagnostics and logistics) integrated with the adoption of innovative, data-driven surveillance methods such as statistical modeling and genomics for a more accurate, timely, and precise response.

Keywords: Epidemiological Surveillance; Drug-Resistant Tuberculosis; MDR-TB; Disease Control; Literature Review.

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1. Introduction

Tuberculosis (TB) continues to be one of the most persistent global health threats in the modern era. As an infectious disease that ranks among the top causes of death worldwide, TB claimed an estimated 1.3 million lives in 2022 alone, out of 10.6 million people who fell ill. The global commitment to end the epidemic is enshrined in the World Health Organization's "End TB Strategy", which targets a 90% reduction in incidence and 95% reduction in mortality by 2035 [1]. However, achieving these ambitious targets is significantly hampered by an increasingly complex challenge: the emergence and spread of Drug Resistant Tuberculosis (DR-TB).

The threat of DR-TB, particularly in the form of multidrug-resistant (MDR-TB) and extensively drug-resistant (XDR-TB), has fundamentally altered the TB control landscape [2]. MDR-TB, defined as resistance to the two most effective first-line drugs, isoniazid and rifampicin, is not only a clinical problem but also a public health crisis [3]. Treatment requires longer (up to 24 months), more toxic and much more expensive regimens, with global success rates still below 65% (Baum et al., 2024). With an estimated 410,000 new cases of MDR/RR-TB by 2022, DR-TB is a major barrier that could reverse the progress of TB programs and threaten global health security [4].

Epidemiological surveillance serves as a key navigation system for national TB control programs, especially in the face of such a crisis. Its role cannot be underestimated, as an effective surveillance system is the foundation of any measured and targeted public health

response [5]. Fundamentally, the function of epidemiologic surveillance in DR-TB control covers four critical domains: (1) Detection and Monitoring, to measure the magnitude of the problem, monitor resistance trends over time, and detect the emergence of new strains; (2) Resource Allocation and Policy, to direct investments to high-risk geographic areas and populations; (3) Program Evaluation, to assess the impact of interventions such as the use of new drug regimens or changes in diagnostic strategies; and (4) Understanding Transmission, to distinguish between cases of acquired resistance (due to inadequate treatment) and cases of primary resistance (due to direct transmission in the community) [6]. However, despite this ideal framework, the implementation of DR-TB surveillance in various countries shows a significant disconnect between objectives and capabilities in the field. An evaluative study in Brazil, for example, revealed that the national system was only able to detect less than half (46.4%) of estimated MDR-TB cases, indicating a massive hidden burden [7]. Meanwhile, a cascade of care analysis in Madagascar clearly shows that the biggest failure point is at the diagnosis stage, where 59% of MDR-TB patients who have accessed health services failed to be diagnosed correctly, exacerbated by extreme logistical delays [8]. On the other hand, studies in Bhutan and Pará, Brazil, highlighted that high primary transmission in the community is the main driver of the epidemic, a dynamic that is often missed by conventional surveillance systems [9]. This gap, coupled with methodological variations from passive systems to stateof-the-art genomic surveillance creates uncertainty in disease burden estimates and hinders the formulation of effective public health responses.

Given this complex landscape, this literature review was developed to systematically review and synthesize the current scientific evidence on the role, methods, challenges, and innovations in epidemiological surveillance for drug-resistant TB control. By deeply analyzing various surveillance approaches and their findings across different countries, this study aims to formulate a conceptual framework that can guide future improvements in DR-TB control program policies and practices.

2. Proposed Method

This study used a literature review method to review and synthesize information from various scientific sources on the role of epidemiological surveillance in drug-resistant tuberculosis (TB) control efforts. This approach allowed for an in-depth analysis of existing studies to gain a comprehensive overview of the topic.

Article Search and Selection Strategy

A systematic literature search was conducted in June 2025 through two major electronic databases, PubMed and Google Scholar. The search process used a combination of specific keywords with the formula: ("epidemiological surveillance" OR "disease surveillance") AND ("drug-resistant tuberculosis" OR "MDR-TB" OR "XDR-TB") AND ("control" OR "prevention"). The article selection process followed the steps adapted from the PRISMA (Preferred Reporting Items for Systematic Reviews and Meta-Analyses) flow chart. In the identification stage, an initial search of the PubMed and Google Scholar databases yielded 19 and 328 articles, respectively, for a total of 347 articles. Next, at the screening stage, the articles were evaluated based on title, abstract, and duplication. Articles that were not relevant to the topic or were duplicates were excluded. Next, at the eligibility stage, the remaining articles were examined in their entirety (full text) based on predefined inclusion and exclusion criteria. Inclusion criteria included original research or review articles that addressed the role of epidemiologic surveillance in controlling drug-resistant TB, published between 2015-2025, and available in full-text format. After going through all these selection stages, at the final stage (included), 6 articles were obtained that were considered to meet the criteria for further analysis.

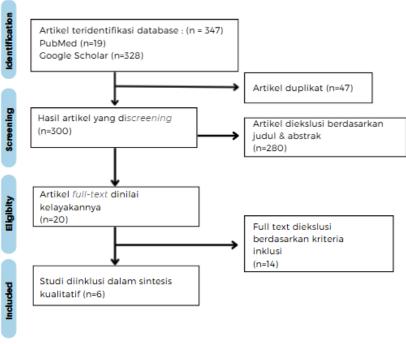


Figure 1. Flowchart of Search

Data Extraction and Analysis

From the six selected articles, a systematic data extraction process was carried out, which included the author's name, year of publication, study design, research location, and main findings. The collected data were then synthesized and analyzed descriptively to answer the research objectives.

3. Results and Discussion

Table 1. Summary of Epidemiologic Surveillance Studies on Drug-Resistant TB (2015-2025)

No.	Author (Year)	Study Title	Study Design	Location &	Surveillance	Main Findings	Conclusion & Relevance
				Population	Method		
1	Sarah E. Baum, et	Surveillance For	Methodological	Brazil; more	Propose a new	The corrected	This statistical correction
	al. (2024) [4]	TB Drug	development	than 800,000	surveillance	prevalence estimates of	method can be generally
		Resistance Using	study applying a	reported TB	method that uses	rifampicin resistance	used to utilize routine RDT
		Routine Rapid	statistical	cases,	routine rapid	were statistically higher	data to produce more
		Diagnostic Testing	correction model	categorized into	diagnostic test	(28-44% for new cases,	accurate and timely
		Data:	(hierarchical	new and treated	(RDT) data that is	2-17% for old cases)	estimates of drug-resistant
		Methodological	generalized additive	cases.	not universal in	than the naive	TB prevalence. It is
		Development and	regression model) to		coverage. This	estimates (directly from	particularly relevant for
		Application in	individual-level		method corrects	RDT data), suggesting	countries where drug
		Brazil	TB case data in		for selection bias	that RDT testing is	resistance testing coverage
			Brazil from 2017-		by: (1) modeling	more common in	is not yet universal,
			2023.		the probability of	populations with lower	allowing them to get a
					resistance in	risk of resistance, so	better picture of the
					tested patients,	the naive estimates	epidemiology without
					and (2) using the	underestimate the true	having to wait for
					model results to	burden of resistance;	expensive and infrequent
					impute (estimate)	This method produces	national prevalence
					the resistance	estimates with	surveys.
						narrower uncertainty	

					status in untested	intervals than the WHO	
					patients.	estimates; There was a	
						downward trend in	
						rifampicin-resistant	
						cases between 2017-	
						2023 for both new and	
						treated cases.	
2	Mei-Hua Wu, et al.	Surveillance Of	Population-based	Taiwan; 1,511	Population-based	There was a significant	The MDR-TB management
	(2022) [10]	Multidrug-	retrospective	MDR-TB cases,	continuous	decline in total MDR-	program in Taiwan is
		Resistant	study analyzing	consisting of	surveillance	TB cases from 2008-	considered effective, as
		Tuberculosis In	reported MDR-	941 new cases	system. Includes	2019, with an	indicated by the overall
		Taiwan, 2008-	TB case data from	and 485	universal Drug	annualized rate of	downward trend in cases.
		2019	2008 to 2019.	previously	Sensitization	decline (APC) of -	However, the slow decline
				treated cases.	Testing (DST) for	4.17%; the steepest	in new cases remains a
					almost all culture-	decline was in	challenge. It shows the
					positive TB cases	previously treated cases	success of structured
					(98.5%), with	(APC -9.18%), while	program interventions
					confirmation by	the decline in new cases	(such as DOTS-Plus and
					the National	was slower (APC -	universal DST). These
					Reference Laboratory	1.41%); among MDR-	surveillance results are
					(NRL). Data were	TB patients, resistance	important for guiding drug
					collected from	to other drugs remains	procurement policies,
					various national	high: Ethambutol	updating treatment
					registry systems.	(47.2%), Streptomycin	regimens, and designing
						(42.4%), and	diagnostic algorithms to
						Pyrazinamide (28.9%);	achieve TB elimination.
						There is a downward	
						trend in resistance to	
						some second-line drugs,	
						especially to injectable	
						drugs (SLIDs) in new	
						cases and	
						fluoroquinolones in	
						previously treated cases;	
						The XDR-TB rate is	
						relatively low (1.9%).	
3	Knoblauch, A.M.,	Multidrug-	This study was a	Madagascar;	Passive & active	59% of MDR-TB	The study concludes that it
	Grandjean	Resistant	retrospective	The main	surveillance;	patients who reached	is urgent to expand
	Lapierre, S.,	Tuberculosis	review that	population in	national NTP &	health centers failed to	coverage and strengthen
	Randriamanana,	Surveillance And	analyzed data	this study were	lab data.	be accurately	diagnostic and management
	D., et al. (2020) [7]	Cascade Of Care In	from the	"presumptive		diagnosed; in 2017, only	capacity for MDR-TB
		*	<u> </u>	*	<u> </u>		= :

		Madagascar: A	tuberculosis (TB)	MDR-TB cases".		about one-third	throughout Madagascar.
		Five-Year (2012-	control program	A total of 2,391		(32.7%) of cases that	Current surveillance data
		2017) Retrospective	in Madagascar	samples from		should have been	likely underestimates the
		Study	over a five-year	presumptive		screened were actually	true burden of MDR-TB in
			period, from	MDR-TB		referred; Only 75% of	the country, so the
			September 2012	patients referred		diagnosed patients	implementation of a new
			to December	to the national		successfully started	national drug resistance
			2017. We	reference		treatment, and in the	survey is highly
			collected and	laboratory		end, only 33% achieved	recommended to obtain
			analyzed	between 2012-		a long-term relapse-free	accurate prevalence data.
			notification data	2017 were		cure; Among the high-	Relevantly, this study
			from the national	analyzed in this		risk groups tested, the	provides clear evidence of
			TB control	study.		MDR-TB rate was	weaknesses in the health
			program, clinical			relatively stable, ranging	system that directly impact
			management			from 3.9% to 4.4%	disease control. Using a
			data, and data			during the study period.	cascade of care approach
			from the national				and GIS, the study
			reference				effectively identified critical
			laboratory.				points that require
							intervention, such as
							increased access to rapid
							diagnosis (e.g. GeneXpert),
							improved logistical referral
							systems, and strengthened
							treatment monitoring.
							These findings are not only
							important for Madagascar
							but can also serve as lessons
							for other resource-limited
							countries facing similar
							challenges in MDR-TB
							control.
4	Tourinho et al.	Evaluation of the	Descriptive	Brazil; 6,078	A passive and	The data quality of the	While the system is useful,
	(2020) [9]	Drug-Resistant	evaluative study	Drug-Resistant	universal	systems was rated as	its low sensitivity suggests
		Tuberculosis	based on data for	Tuberculosis	surveillance	very good (average	significant gaps in case
		Surveillance System,	the period 2013-	(DRTB) cases	system that uses	completeness of 95%);	detection. Improved access
		Brazil, 2013-2017	2017, using	reported to the	two information	The sensitivity of the	to DRTB diagnosis is
			guidelines from	SITETB	systems: SINAN	system was low,	urgently needed. The
			the	information	(for initial	detecting only about	results of this evaluation
			Centers for Disease	system.	notification of all	46.4% of the WHO	provide strong evidence for
			Control and		TB cases) and	estimated MDR-TB	health program managers in
			Prevention (CDC).		SITETB (an online	cases; The flexibility,	Brazil to improve case
			Trouming (CDC).		CIIII (an omine	cases, The healphilty,	Diam to improve twe

				system specialized	acceptability, timeliness,	detection, strengthen
				for cases	and stability of the	laboratory networks, and
				diagnosed as	system were rated as	simplify surveillance system
				drug-resistant).	"moderate";	workflows.
					Compliance for follow-	
					up sputum culture	
					testing declined	
					dramatically with time	
					on treatment (from	
					94.8% on the first	
					culture to 43.6% on the	
					fourth culture).	
5 Thinley Dorji, e	al. High Incidence Of	Retrospective	Location :	Based on data	The incidence of MDR-	Bhutan has a high and
(2024) [5]	Multidrug-	cohort study	Bhutan,	from the national	TB was high (10.6%),	uneven incidence of MDR-
	Resistant	using national	Population:	surveillance	while Isoniazid	TB. There is a need to
	Tuberculosis In	surveillance data	2,290 samples	system (TBISS).	resistance (HR-TB) was	decentralize diagnostic
	Bhutan: A Cohort	from 2018-2021	from TB cases	Initial diagnosis	lower (3.5%), in	facilities to areas with high
	Study Based On	to analyze	tested for drug	using microscopy	contrast to patterns in	case load to expedite
	National TB	resistance	resistance	and Xpert	neighboring countries;	diagnosis and close
	Surveillance Data	patterns and	between 2018-	MTB/RIF.	Significant risk factors	monitoring of at-risk
		identify risk	2021.	Samples were	for MDR/pre-XDR-	groups. This study has the
		factors using		then sent centrally	TB were age 18-39	relevance of providing the
		logistic		to the National	years, female gender,	first comprehensive
		regression.		Tuberculosis	and previous TB	national data on the burden
				Reference Laboratory	treatment history;	and risk factors of DR-TB
				(NTRL) for	Distribution of cases	in Bhutan, which is crucial
				genotypic	was uneven, with the	for policy making,
				(MTBDRplus)	highest incidence in	intervention planning, and
				and phenotypic	Thimphu, Samtse, and	strengthening the national
				(pDST) testing.	Sarpang districts; The	TB control program.
					proportion of primary	
					MDR-TB (9%) was	
					higher than the global	
					average, suggesting the	
					possibility of active	
					transmission in the	
					community; There was	
					a significant diagnostic	
					gap, with only about	
					half of the samples	
					successfully tested for	
					resistance due to	
					resistance due to	

6	Davi	Josué	Comprehensive	Retrospective	Location: Pará	Using genomic	First report on	Genomic surveillance
	Marcon, et al.	[8]	Genomic	cross-sectional	state, Brazil.	surveillance by	detection and	(WGS) is essential to detect
			Surveillance Reveals	observational	Population: 103	performing Whole-	transmission of XDR-	and monitor the emergence
			Transmission	study using Whole-	patients treated	Genome Sequencing	TB in Pará, Brazil,	and transmission of DR-TB
			Profiles of	Genome Sequencing	for Drug	(WGS) on	based on 2021 WHO	strains, including XDR-TB.
			Extensively Drug-	(WGS) for	Resistant TB	bacterial isolates	definition; WGS	There are critical gaps in
			resistant	genomic analysis	(DR-TB)	from DR-TB	identified a complex	existing standardized
			Tuberculosis Cases		between	patients. WGS	resistance profile: MDR	diagnostic protocols. This
			in Para, Brazil		October 2021 -	results were	(52%), pre-XDR (20%),	has prompted the
					December	compared with	and XDR (7%);	implementation of WGS as
					2022, with 40	existing Standard of	Evidence of active	a routine tool in TB
					isolates	Care (SOC)	transmission was found	surveillance in Brazil to
					randomly	diagnostic results	(37% of cases were in	speed up diagnosis,
					selected for	(such as Xpert	genetic clusters),	understand transmission
					WGS analysis.	and pDST) for	including clusters of	patterns, detect resistance
						resistance and	pre-XDR and XDR	to new drugs, and guide
						transmission	strains; WGS	more effective public health
						analysis.	successfully detected	interventions.
							mutations related to	
							resistance to a new drug	
							(Bedaquiline), which	
							were not detected by	
							standard methods;	
							There was a significant	
							discrepancy (41%)	
							between WGS results	
							and standard diagnostic	
							methods, largely due to	
							not testing for second-	
							line drugs.	

Discussion

A comparative analysis of six drug-resistant tuberculosis (DR-TB) surveillance studies from different countries highlights a complex and challenging picture of global TB control efforts. The review systematically revealed significant gaps between estimated disease burden and detected cases, the importance of understanding local transmission patterns, and the urgency to adopt more sophisticated and integrated surveillance methods. Despite programmatic successes in some regions such as Taiwan, studies from Brazil, Madagascar, and Bhutan collectively demonstrate that surveillance systems in many high TB burden countries still struggle to achieve adequate sensitivity and timeliness.

One of the most consistent and alarming findings of this review is the low sensitivity of conventional surveillance systems in detecting DR-TB cases. An evaluative study in Brazil by Tourinho et al. (2020) quantitatively showed that the national surveillance system was only able to detect about 46.4% of the estimated MDR-TB cases estimated by WHO. This low sensitivity indicates that a large proportion of drug-resistant cases in the population are likely to go undiagnosed, untreated, and potentially continue to transmit the disease [9]. This problem was further detailed by a case study from Madagascar, Knoblauch et al. (2020), which used a cascade of care approach. The study identified that the biggest "gap" in the flow of care occurred at the diagnosis stage, where 59% of MDR-TB patients who had reached a

health facility failed to be accurately diagnosed. Logistical delays, with an average of 28 days from sample collection to testing, further exacerbated this situation and contributed to poor treatment outcomes, with only 33% of patients achieving a long-term cure [7].

Facing the challenge of bias due to non-universal test coverage, a methodological study by Baum et al. (2024) in Brazil offers an innovative solution. By applying a statistical correction model, they showed that the true estimated prevalence of rifampicin resistance was 28-44% higher in new cases and 2-17% higher in previously treated cases compared to naive estimates from existing rapid diagnostic test (RDT) data. These findings unequivocally confirm that the current system underestimates the burden of DR-TB due to test selection bias more often in populations with lower risk of resistance. This underscores the importance of not only expanding test coverage but also applying sophisticated analytical methods to interpret incomplete surveillance data, an approach that is particularly relevant for many low- and middle-income countries [4].

Understanding the source of DR-TB cases whether due to treatment failure (secondary resistance) or direct transmission in the community (primary resistance) is key to designing effective interventions. A cohort study from Bhutan by Dorji et al. (2024) provides worrying evidence, where the proportion of primary MDR-TB (in new cases) reached 9%, which is much higher than the global average. This indicates that active transmission of MDR-TB strains in the community is a major driver of the epidemic in Bhutan, not just individual treatment failure. This high proportion of primary MDR-TB also makes the epidemic pattern in Bhutan (10.6% MDR-TB vs 3.5% HR-TB) unique when compared to neighboring countries such as India and Thailand, where Isoniazid resistance (HR-TB) is predominant [5]. This suspicion of community transmission was further confirmed by a genomic surveillance study in Pará, Brazil Marcon et al., (2024) using Whole-Genome Sequencing (WGS), this study definitively identified that 37% of the analyzed cases were part of a genetic transmission cluster, including the spread of pre-XDR and XDR-TB strains. WGS not only proved the existence of active transmission but also revealed the evolution of a locally endemic strain (SIT 2517) that had evolved into pre-XDR. These findings provide strong molecular evidence that DR-TB control must shift from a reactive approach (treating the resistant) to a proactive one (breaking the chain of transmission) [8].

The evolution of surveillance methods is another important theme in this review. Wu et al. (2023), Taiwan, represents the gold standard with a continuous surveillance system that includes universal Drug Sensitization Testing (DST) for 98.5% of culture-positive cases, which has proven effective in significantly reducing MDR-TB cases. However, in countries where resources are limited, innovative approaches are crucial. The study by Marcon et al. (2024) demonstrated the power of WGS as a superior surveillance tool, which was able to detect resistance to new drugs such as Bedaquiline missed by standard methods, as well as identify diagnostic discrepancies of up to 41%. Meanwhile, the statistical correction method by Baum et al. (2024) shows how existing RDT data can be utilized to produce more accurate estimates, even when test coverage is not ideal. This suggests that there is a spectrum of solutions that can be applied, ranging from basic system strengthening to the adoption of cutting-edge genomic technologies.

The implications of these findings for public health policy are clear. First, closing diagnostic gaps is a top priority. Recommendations from the study in Madagascar to strengthen capacity and the study in Bhutan to decentralize diagnostic facilities to high-burden areas should be addressed. Second, the adoption of more advanced technologies should no longer be considered a luxury, but rather a necessity. WGS, as demonstrated in Pará, provides insights that cannot be obtained from other methods and is crucial for dealing with XDR strains and resistance to new drugs. Third, the focus of control programs must expand to include breaking the chain of transmission in communities, given the strong evidence of primary resistance transmission in Bhutan and Brazil. This means more aggressive contact investigations and possibly active screening in high-risk populations. Fourth, integration of information systems, as successfully implemented in Taiwan and a challenge in Brazil, is the foundation for effective surveillance and accurate program evaluation.

This review certainly has limitations. The studies analyzed come from different geographical contexts and timescales, with varying health systems and disease burdens, so direct comparisons should be made with caution. The diverse methodologies ranging from retrospective cohort studies, system evaluations, to the development of statistical methods and genomic surveillance present different puzzle pieces of the same problem, rather than a single complete picture. Reliance on routine surveillance data also carries inherent risks regarding data quality and completeness, a limitation recognized in some of the studies themselves.

Taken together, these six studies collectively confirm that DR-TB surveillance is at a crossroads. On the one hand, there are mature systems that are successfully curbing the epidemic. On the other hand, many countries still face a hidden burden of disease that continues to spread in communities. Future progress will not come from a single solution, but rather from a dual approach: strengthening the foundations of conventional surveillance systems as outlined in the Madagascar study and the Brazil evaluation, while strategically adopting innovations such as genomic surveillance and advanced statistical methods to make TB control programs more sensitive, accurate, and responsive to the evolving DR-TB threat.

4. Conclusions

A comprehensive review of various drug-resistant tuberculosis (DR-TB) surveillance systems around the world confirms a crucial paradigm shift. DR-TB control can no longer rely solely on passive approaches and treatment of detected cases. Evidence from Brazil, Madagascar, and Bhutan consistently suggests a much larger disease burden than reported with massive diagnostic gaps and low system sensitivity being major barriers. Furthermore, genomic and epidemiologic analyses in Pará (Brazil) and Bhutan revealed that active transmission in the community rather than treatment failure is the main driver of the DR-TB epidemic, including for pre-XDR and XDR strains. Therefore, the future of effective DR-TB surveillance lies in the integration of two main pillars. First, fundamental strengthening of public health systems, including decentralization of diagnostics, accelerated sample logistics, and enhanced contact investigation capacity, as is urgently needed in Madagascar and Bhutan. Second, the adoption of smarter and more accurate innovative surveillance methods. The use of statistical correction models for incomplete data, as applied in Brazil, and the implementation of genomic surveillance as a routine tool not just for research have proven to be able to provide a truer picture of epidemiology, detect new drug resistance, and map the chain of transmission with precision. Only by marrying conventional system strengthening with datadriven innovation can TB control programs move from merely managing the epidemic to actually having a chance to eliminate it according to global targets.

Based on the synthesis of findings from various TB control program contexts, several strategic suggestions and recommendations can be formulated to strengthen DR-TB surveillance systems and accelerate global elimination efforts. These recommendations are designed to address identified gaps, ranging from the fundamental level to the application of cutting-edge technologies.

- 1. Strengthening Diagnostic Capacity at the Primary Care Level. Expanding access to rapid diagnostic tests (RDTs) such as GeneXpert and decentralizing culture and Drug Sensitivity Testing (DST) facilities to regional or district levels should be a priority. This is crucial to address massive diagnostic gaps such as those identified in Madagascar (59% of MDR-TB cases missed) and Bhutan, where logistical challenges impede timely testing.
- 2. Adoption of Statistical Correction Methods for Disease Burden Estimation. For countries with non-universal coverage of drug resistance testing, National TB Programs (NTPs) are recommended to adopt statistical correction models on existing routine RDT data to generate more accurate DR-TB prevalence estimates. As demonstrated in Brazil, this method corrects for selection bias and reveals a higher disease burden than initially estimated, allowing for more targeted resource allocation without having to wait for a national survey.
- 3. Programmatic Implementation of Genomic Surveillance.
 Genomic surveillance (Whole-Genome Sequencing/WGS) should begin to be integrated as a routine public health tool, especially in DR-TB hotspot areas or in high-risk cases. Utilization of WGS has proven to be very effective for: (a) map transmission clusters in real-time, (b) detect early resistance to new drugs (such as Bedaquiline) not covered by standard tests, and (c) identify diagnostic discrepancies for continuous improvement of laboratory workflows, as clearly demonstrated in the study in Pará, Brazil.
- 4. Increased Focus on Contact Investigation and Breaking the Chain of Transmission. Given the strong evidence of active transmission of DR-TB in communities, as demonstrated by the high incidence of primary resistance in Bhutan and the presence of genetic clusters in Brazil, programs should improve the capacity and quality of contact investigations. Interventions should not only focus on treatment, but also proactively find source cases and break the chain of transmission, a crucial step to reduce the rate of new MDR-TB cases.

5. Development of Integrated Information Systems and Program Evaluation. Investing in the development of an integrated and digitized health information system is necessary to ensure that patient, laboratory, and treatment data are seamlessly linked. A solid system, as implemented in Taiwan, provides the backbone for accurate data analysis, standardized program performance evaluation, and evidence-based decision-making essential for effective disease control.

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