

The Use of Artificial Intelligence in Tuberculosis Care: A Scoping Review

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Abstract

Tuberculosis (TB) is a subacute to chronic respiratory infection with insidious onset and protean symptoms. Treatment is complex and requires a multi-drug, multi-month regimen in which adherence is critical. Artificial intelligence (AI) offers promising solutions to challenges across the TB care cascade including screening, diagnosis and treatment. We conduct a scoping review of the literature published from 2017 to 2025 on the use of AI in TB screening, diagnosis, drug resistance diagnosis, treatment monitoring, and regimen design. We then extract data on study characteristics, AI methodology, input data, and sample size and describe AI tool performance and technology readiness level (TRL). Ninety studies are included, representing 803,383 study participants across 24 countries. Most studies (n=46) focus on radiological imaging for TB screening or diagnosis, but a burgeoning number of studies address drug resistance diagnosis (n=11), regimen design (n=4), treatment monitoring (n=12), and treatment adherence (n=8). Reported accuracy of AI interpretation of chest imaging for TB diagnosis was high at a median Area Under the receiver operating Curve (AUC) of 0.94 [IQR 0.12, range 0.81-0.99] for internal validation and 0.89 [IQR 0.14, range 0.66-0.98] for external validation, and a median TRL of 5 (IQR 1, range 4-7). AI demonstrates promise for advancing TB care towards World Health Organization (WHO) End TB targets, but several gaps remain, including AI-ready fit-for-task data availability, limited external validation and challenges in clinical integration. Closing these gaps will be critical for realizing the full potential of AI in TB care towards WHO End TB targets.

Keywords: tuberculosis; artificial intelligence; machine learning; statistical learning; deep learning; AI platform; neural network; automatic; diagnosis; screening; regimen design; treatment monitoring; drug resistance

1. Introduction

Tuberculosis (TB) is the top infectious disease killer globally, causing more than 10.5 million new cases of disease in 2023 (1). TB demonstrates stark regional variation in incidence ranging from 2 to >800 per 100,000 population by country (2). The World Health Organization (WHO) has proposed the End TB Strategy which aims to reduce global TB incidence to <10 per 100,000 by 2035, however, modelling has projected that even with maximal deployment of currently available treatment and preventative strategies for TB, this degree of reduction will not be possible (3). Further, the sustainability of current TB interventions is at risk given the dependence of TB control programs on non-governmental funding and international aid (4). More than ever before, innovations that

improve delivery of TB screening, diagnosis, and treatment are needed. AI is at the forefront of such innovations as it offers cross-cutting data processing and interpretation capability promising to increase access to healthcare, increase healthcare reliability, and reduce costs. Here, we review AI innovations in TB care with the aim of charting an AI research agenda that can support achievement of WHO End TB targets (5). Given the rapidly evolving and heterogeneous nature of AI applications in TB care, a scoping review is an appropriate approach to map existing evidence, identify knowledge gaps, and classify AI tools across the care cascade.

The TB care cascade comprises four main steps: screening, diagnosis, treatment regimen design, and treatment monitoring, with the goal of TB cure (Figure 1). Many gaps currently exist along this care cascade (6). Gaps in screening include the lack of low cost, high sensitivity tools for early detection of TB prior to symptom onset (*i.e.*, subclinical or asymptomatic TB) which is increasingly recognized as a critical step in interrupting transmission to reduce disease incidence (7,8). Prompt and accurate TB diagnosis in symptomatic patients presenting to healthcare facilities is crucial for the timely initiation of TB therapy needed not only for cure but also for the reduction of long-term disease complications (9). Diagnosis classically relies on microbiological detection of TB bacilli. Yet, the low burden of bacilli in relevant human samples like sputum, and gastric aspirate (especially for children and those with HIV) requires improvements in sensitivity (10). Tests that leverage new sample types such as tongue swabs are expected to be recommended by WHO soon, and exhaled breath condensates also hold promise. Other adjunctive non-microbiological diagnostics have the potential to expedite diagnosis or boost accuracy. These include a variety of novel assays, such as assays of the host immune response, ultrasound manifestations of disease, and audio recordings of breath sounds and/or cough. Several of these approaches are in early-stage development and/or have faced challenges in implementation due to cost or complexity.

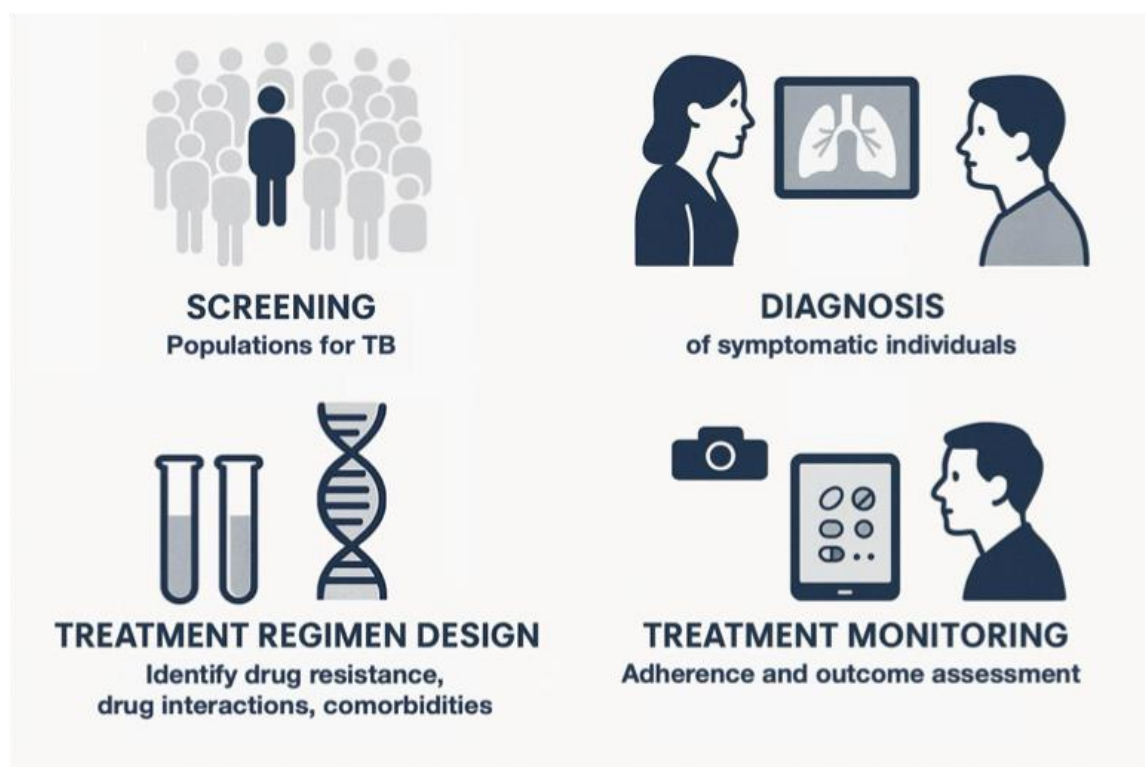


Figure 1. The TB care cascade comprises four main steps: screening, diagnosis, treatment regimen design, and treatment monitoring with the goal of TB cure. Source: Authors' schematic of the tuberculosis care cascade.

Challenges in regimen design include effectiveness of the regimen, drug resistance, drug interactions, and comorbidities. Globally, 10% of TB is resistant to the drug isoniazid, with 3% resistant to both isoniazid and rifampicin, i.e., multidrug resistant (MDR) (1). Recent reports on MDR-TB describe over 10% resistance to bedaquiline in some countries (11,12). Rapid and comprehensive drug resistance detection is critical for the design of an effective treatment regimen, but this has been difficult to achieve using traditional microbiological tools. In the last decade, pathogen sequencing has emerged as a viable rapid and low biohazard approach to comprehensive resistance diagnosis, but cost, output complexity, and interpretability are still major barriers.

After regimen design, treatment monitoring for adherence, side effects, and adequate microbiological and symptom response is needed. Monitoring is highly resource-intensive and limited by availability and accuracy of sputum testing including culture as well as human resources needed for outreach (14). Lastly, innovation around patient support, counseling, and education can facilitate adherence, management of side effects, and patient-centered care across the entire cascade.

In this study, we define AI broadly as encompassing statistical learning, machine learning, deep learning, and Large Language Models (15). Through automation and advanced analytical capabilities, AI can support interpretation of complex clinical data streams such as radiological images and genome sequencing data (16,17). AI tools can also integrate with compact medical or personal devices such as handheld ultrasound machines to support clinical decision-making, and smartphones to provide direct patient counselling or record medication adherence, bridging gaps in healthcare delivery in resource-limited settings.

This scoping review aims to map and characterize AI applications across the TB care cascade, describing their data inputs, reported performance, and technological readiness level for clinical use on a standardized scale. We also highlight key research and implementation gaps to inform future development and deployment of AI in global TB control.

2. Methods

The review protocol was registered on the Open Science Framework (OSF) and is available at <https://osf.io/wah6d/overview> (OSF registration number: wah6d).

2.1. Search Strategy and Study Identification

A literature search was conducted in PubMed for English-language studies published between January 1, 2017 and August 1, 2025. We limited inclusion to studies published from 2017 onward to reflect contemporary AI methodologies and clinical applications relevant to current TB care. Only English-language, peer-reviewed original research articles were included to ensure methodological transparency and feasibility of review. Reviews and non-original publications were excluded to avoid duplication of evidence. PubMed was chosen as it is known to be the largest indexed database of biomedical, peer-reviewed publications.

The search strategy targeted titles containing the following terms: (“tuberculosis,” or “TB”) and (“artificial intelligence,” or “machine learning,” or “deep learning,” or “neural network,” or “AI platform,” or “automatic,” or “smart pillbox,” or “statistical learning,” or “machine-learning”). This search yielded 380 unique records. The final search was conducted on August 1, 2025. The full PubMed search strategy, including field tags and limits, is provided in Table S1.

Articles were included if they met all the following criteria:

- 1) Original research article
- 2) Describing the development and testing of AI or a related method (as detailed by the search terms above)
- 3) The AI is intended to improve screening, active case finding, diagnosis, drug resistance diagnosis, treatment monitoring, drug regimen design, or resistance prediction of TB
- 4) The AI is applied to data derived from human subjects with active tuberculosis (clinical or subclinical, pulmonary with or without extrapulmonary) and an appropriate control population (e.g.

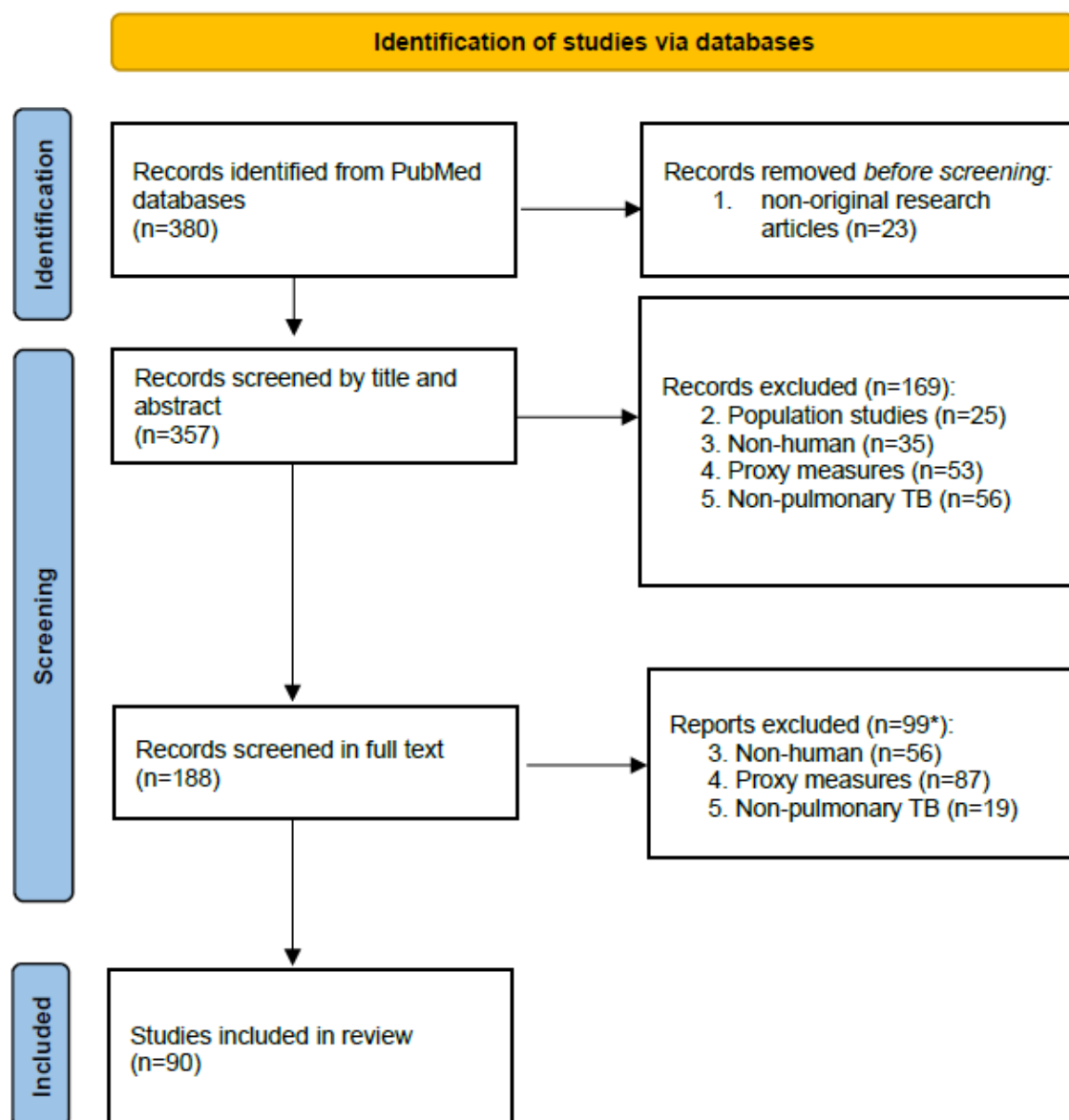
for TB screening: healthy individuals, for TB diagnosis: patients meeting criteria for TB testing but do not have TB, etc.).

Articles were excluded if they met any of the following criteria:

- 1) Any of the following type of articles: systematic review, scoping review, meta-analysis, editorial, commentary, letter, opinion piece, policy-focused paper
- 2) Focused primarily on population level incidence or forecasting and did not address individual and direct patient care
- 3) Analyzed data exclusively from animal or *in vitro* experiments that are not directly derived from human subjects
- 4) Studied proxy measures without explicit or direct clinical application in one of the four care categories, for example, predicted specific radiographic abnormalities (e.g., pleural effusion) rather than TB/ no-TB diagnosis
- 5) Focused on a disease other than pulmonary tuberculosis (including studies that focused solely on extrapulmonary TB) or if they compared tuberculosis to a specific other disease rather than the desired general non-TB control group e.g., COPD, pneumonia, HIV-only cohorts, or other non-TB comparator diseases.

2.2. Screening and Selection

Study selection was performed in stages in accordance with PRISMA guidelines (Figure 2). First, duplicates, retracted papers, author corrections, letters to the editor, resource articles, and comments were removed (n=23), resulting in 357 records for title screening. Then two reviewers (ASR and KRVH) independently reviewed titles and abstracts. ASR reviewed abstracts with the support of ChatGPT-4 and a standardized prompt detailing the relevant inclusion and exclusion criteria (Figure S1). The inclusion and exclusion rationale for each paper was manually reviewed and categorized by ASR.



*Some papers meet more than one of the exclusion criteria, hence the discrepancies in totals.

Figure 2. PRISMA flow diagram outlining the number of records identified, screened, excluded (with reasons), and included at each stage. Source: Authors' PRISMA diagram for this scoping review.

2.3. Study Data Extraction

Two reviewers (ASR, KRVH) independently extracted data from each included study. Data extraction forms were piloted on a subset of studies then refined prior to full extraction to ensure consistency and completeness. We captured the following variables: clinical application (screening, diagnosis, treatment monitoring or adherence, regimen design), AI methodology, type and source of input data, patient population and validation setting, sample size (and key subgroups, where relevant), training and/or external validation performance (such as sensitivity, specificity, accuracy, and Area Under the receiver operating characteristic Curve [AUC]), reference standard, stage of AI development, and study identifiers (year, country, setting). Any discrepancies were resolved by consensus. Extracted data were compiled into structured tables to enable comparison of clinical application, performance, and readiness for implementation across studies.

In addition to the metrics extracted above, we assessed each AI based on a standardized Technology Readiness Level (TRL). The TRL enables a semi-quantitative assessment of technological maturity and enables comparison of AI across studies and care categories. We adapted the US Department of Defense biomedical and digital health 8-level TRL scale (18) that ranges from initial scientific discovery (TRL 1) to clinical trial or advanced real-world evidence with full clinical implementation (TRL 8) (Table S2). For summarization and descriptive purposes only, we aggregated internal model performance, external model performance, or TRL across data input types and care categories (Table S3). These aggregates are not meant for clinical use or as endorsement of the accuracy or readiness of the AI tools, but rather as a description of the reported literature. We used Wilcoxon rank-sum tests to compare TRL distributions between groups. Consistent with scoping review methodology, no formal critical appraisal of study quality or risk of bias was conducted; instead, we extracted several quality-related metrics such as sample size, the gold standard metric the AI was compared to, study setting including multi-site setting, whether external validation was performed, and the reported difference in performance between training and external validation (Table S3).

3. Results

3.1. Overview of AI Applications in TB Care

The literature search identified 380 total studies of which 90 met inclusion criteria and from which we extracted study characteristics (Table 1). By care category, studies most commonly focused on the use of AI for TB diagnosis (n=33) and TB screening (n=23) (Table 2A, Table S3). We identified 20 studies focused on treatment monitoring, 11 on drug resistance diagnosis and four on regimen design; one study focused on a combination of care categories (diagnosis and regimen design) (Table 2B, Table S3). By data type, most AI tools interpreted imaging data (n=46; 38 (82%) chest X-ray [CXR]). Fewer studies analyzed other clinical or demographic data, including blood-based laboratory data, or symptom data collected from electronic health records or registries (n=17). Seven studies evaluated microbiological images or markers, six evaluated host or pathogen omic/molecular data, three clinical audio recordings, and in three there was direct patient interaction. Eight other studies used multimodal data sources, with some combining imaging with clinical variables. The cumulative sample size across all studies was 803,383, with a median sample size per study of 898.5 (IQR: 2,045, range 50-243,726) (Table S3). Technology readiness levels (TRLs) were used as a descriptive proxy for the maturity of AI tools. Across TB care categories, TRLs ranged from 4 to 8, with the most mature tools being concentrated in screening and diagnosis (Figure 3), whereas drug resistance diagnosis, regimen design, and monitoring applications showed greater variability and lower TRLs. Imaging-based tools for screening and diagnosis have higher median and upper-range TRLs than models relying on other input data types (Wilcoxon rank sum test, $p=0.02$, Figure 3).

Table 1. Summary of included studies evaluating AI tools for TB care. For each study, the table details the clinical category (screening, diagnosis, drug resistance diagnosis, treatment monitoring or adherence, and regimen design), and the type of input data. Source: Authors' compilation based on the 90 studies included in the scoping review.

Study category	Study Count	Input Data Type							References
		Imaging	Clinical Variables*	Micro	Pathogen or host Omics	Audio	Patient Facing	Multi-modal*	
Screening	23	17	2	-	-	3	-	1	(20,27-29,48-66)
Diagnosis	33	24	2	6	1	-	-		(19,26,45,67-96)
Drug Resistance Diagnosis	11	3	2	1	2	-	-	3	(16,46,97-107)

Monitoring (Outcome)	12	1	6	-	2	-	-	3	(17,47,106–115)
Monitoring (Adherence)	8	-	5	-	-	-	3	-	(42–44,116–120)
Regimen Design	4	2	-	-	1	-	-	1	(95,121–123)
Total Studies	90***	46	17	7	6	3	3	8	

* Symptom data, electronic health record data, blood-based laboratory testing data. **Multimodal indicates studies that combined two or more data types (e.g. imaging and clinical variables). ***Sums for categories and data types exceed total studies due to overlapping classifications of study categories (one study classified as diagnosis and regimen design).

Table 2. AI Tools summarized by care category, data input, AI model, country, performance (AUC, external AUC), and technology readiness level (TRL). (A) AI Tools for TB screening and diagnosis. (B) AI Tools for TB drug resistance diagnosis, monitoring, and regimen design.

(A)									
Cate gory	Data Input	Specific Input	AI Tool Name	Country	AUC	Ext AUC	TR L	Ref	
Diag nosis	Clinica l variabl es	Demogra phic + blood data	RF + ANN	Multi- country	1	0.946	6	(67)	
			RF (conditional)	China	0.978	0.963	6	(68)	
			JF CXR-1 v2	China	-	0.98	7	(26)	
			qXR v3.2	India	0.944	-	5	(82)	
			U-Net (DL)	-	-	0.91	6	(96)	
			DL (custom, Google)	Multi- country	-	0.89	7	(72)	
	Imagin g CXR		CNN	China, USA	0.87	-	5	(70)	
			EfficientNetV2	Taiwan, China, USA	0.878	0.806	6	(75)	
			qXR v2	India	0.81	-	5	(78)	
			CheXaid	South Africa	0.83	0.71	6	(93)	
			HaMLET	Multi- country	0.97	0.94	6	(81)	
			DLAD	Multi- country	0.988	0.977	6	(77)	
			ResNet50 (DL)	South Korea	0.98	0.815	6	(92)	
			LUNIT v3.1	INSIGHT Pakistan	0.85	-	7	(69)	

	CT		Lunit INSIGHT CXR v3	South Korea	0.924	-	5	(83)
			TB-CAD	South Korea	0.85	-	5	(80)
			EfficientNetB4	India	-	-	5	(76)
			EfficientNetB3	Multi-country	-	-	5	(73)
			EfficientNet-DNN	South Korea	-	0.82	6	(90)
			EfficientNetB4-CNN	India	0.975	0.665	6	(89)
			SVM (GA-optimized)	Multi-country	-	-	5	(71)
			Attention-ResNet	China	0.97	-	5	(74)
			3D ResNet-50	China	0.992	0.946	6	(91)
			U-Net (DL)	China	0.98	-	5	(79)
			DL (custom pipeline)	China	0.971	-	6.5	(94)
			DeepTB	China	0.943	-	5	(95)
	Microbiological	Sputum	MiniLab TB assay	China	-	-	5	(88)
		Microscopy images	ANN	Mexico	1	-	4	(86)
			Depth-wise CNN	Brazil	-	-	5	(84)
		Digital pathology	TB-AI	China	-	-	4	(45)
			Modified EfficientNet	Taiwan	-	-	6	(87)
		Histopathology	HALO AI	Japan	-	-	5	(85)
	Omics data	Metabolo						
	mics + clinical data	RF, SVM, MLP	China	-	-	5	(19)	
Screening	Clinical (audio)	Cough sound	LR (MFCC-based)	South Africa	0.94	-	6.5	(50)
			CNN	South Africa	0.95	-	4	(48)
			Swaasa AI	India	0.94	0.9	7	(49)
	Clinical variables	Symptoms + demographics	CDSS	Brazil	0.79	-	5	(51)

Imaging	CXR	EHR	RF (risk model)	Switzerland, Austria	0.72	0.67	5.5	(52)
			DL (Google Teachable Machine)	Taiwan	-	0.951	5	(61)
			qXR v3/v4	Peru	-	0.78	7	(59)
			ResNet (normalization-free)	Multi-country	0.993	-	5.5	(55)
			F-CAD	Cambodia	-	0.86	7	(56)
			Genki (DeepTek)	India	-	-	7	(65)
			ResNet- DCNN	China	0.994	-	6.5	(54)
			CAD4TB v5	Zambia	-	0.87	7	(64)
			qXR v2; CAD4TB v6	Pakistan	-	0.92	8	(27)
			XmarTB	Public/China	-	-	5	(62)
			qXR v3	Nigeria	-	-	7	(58)
			DLAD	South Korea	-	0.999	6.5	(60)
			qXR-based AI	Nepal	-	-	6.5	(57)
			JF CXR-1 v2	China	-	0.978	7	(28)
			CAD4TB v6/ v7	South Africa, Lesotho	-	0.865	7	(63)
			Multiple CAD tools	Bangladesh	-	0.908	7	(29)
			Multiple CAD tools	Nepal, Cameroon	-	0.94	6.5	(66)
			AI4TB	Nepal	0.72	-	5.5	(53)
		Imaging + clinical data	CNN (demographic)	South Korea	-	0.971	5	(20)

(B)

Cat	eg Data Input	Specific Input	AI Tool Name	Country	AUC	Ext AUC	TRL	Ref	
Drug resistance diagnosis algorithms	Clinical variables	EHR	XGB, RF, SVM, Adaboost, LR, kM	Brazil	0.888	-	5.5	(98)	
		Epidemiological data	ANN	Moldova	0.87	-	4	(97)	
	Imaging	CXR	Ens- DL	Multi-country	0.951	-	5	(99)	
			Ens-DL (AMIS)	Multi-country	0.889	-	5.5	(100)	
		CT	Patch-CNN + SVM	Multi-country	-	-	4	(46)	
	Imaging + clinical	CXR + clinical data		SVM	Multi-country	0.784	-	5	(102)
		CT scan + demographics	Radiomics		China	0.881	0.834	6	(101)
			Multimodal fusion		China	0.899	-	5.5	(103)

Monitoring (Outcome)	Micro-biological	Xpert Ultra (Tm profiles)	Tm-Dist	Multi-country	-	-	5	(104)
	Omics	WGS-derived resistance	GenTB	Multi-country	-	0.94	6	(16)
	Omics + clinical	WGS SNPs + clinical data	LR, GBC, XGB	Uganda	0.95	0.63	5	(105)
	Clinical variables	Symptoms + demographics	ANN, SVM, kNN, RF, DT	Multi-country	-	-	5	(111)
		Demographics + blood data	RF, SGB, CART, MARS	Multi-country	0.808	-	5.5	(112)
		Registry data	Landmark+ RF	Moldova	-	-	5	(109)
		Clinical trial data	DT, RF, SVM, NB	India	0.909	-	5	(110)
		EHR	DT	South Africa	-	-	5	(113)
			XGB, RF, MLP, LGBM, LR, SVM	Taiwan	0.834	-	5	(108)
	Imaging	CT	PM1-3	China	0.815	0.735	6	(106)
	Imaging + clinical	Chest X-ray + clinical data	DL (DeepCatch-X-TB v1)	South Korea	-	-	5	(114)
			Ens- DL (DenseNet121)	Multi-country	0.86	0.714	5.5	(17)
		CT + demographics	RF	China	0.913	0.815	6	(107)
	Omics data	Metabolomics + clinical data	kNN	China	0.87	-	4	(115)
		Proteomics	SVM- MMC	China	-	-	4	(47)
	Clinical variables	Registry data	SVM, AdaBoost, RF, ANN, LR	Uganda	0.87	-	5	(117)
		Epidemiological data	LGBM	Brazil	0.72	-	5	(116)
		Demographic + blood data	RF, SVM, ANN	Pakistan	-	-	5	(118)
		EHR	RF, ANN, NB, kNN, LR	Indonesia	0.808	-	5.5	(120)
			LR (risk model)	Ethiopia	0.78	-	5	(119)
Patient-facing	Video- based ingestion	3D-ResNet	Uganda	0.87	-	4	(42)	
	Dispensation events	STAMP	India	-	-	6.5	(43)	
	Device datalog	EvriMED	Ethiopia	-	-	7	(44)	
Imaging	CT	DeepTB	China	0.943	-	5	(95)	
	CXR	TB- DR CDSS	Multi-country	0.947	-	6	(121)	
Imaging + clinical	CXR + clinical data	DL-Risk	China	0.7	0.77	5	(95)	
Omics data	WGS-derived resistance	TR- CDSS	South Africa	-	-	6	(123)	

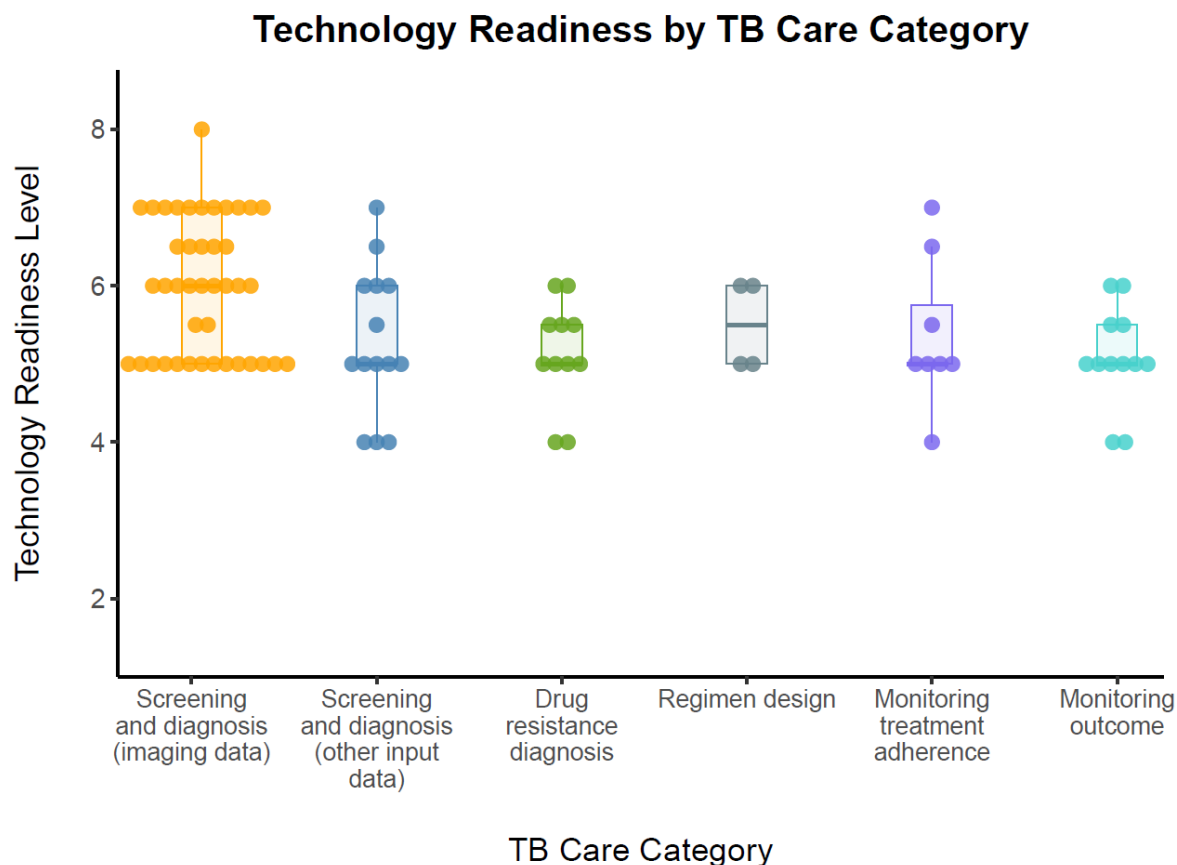


Figure 3. Technology readiness level by TB care category. Swarm plot of technology readiness levels for individual AI tools, grouped by TB care category. Each dot represents a single tool; vertical position indicates its technology readiness level, and horizontal jitter is added to reduce overlap of points within each category. Source: Authors' analysis of technology readiness levels for 90 AI tools identified in the scoping review.

3.2. Screening and Diagnosis

Overview and input data: Studies of AI-based screening and diagnosis (n=56) primarily analyzed CXR images (n=35) and, less commonly, computed tomography (CT) scan images (n=6). Other data input types included microbiological data (n=6), clinical variables (n=4), and cough/audio recordings (n=3); one study utilized patient omics data (19) and one study integrated clinical variables with imaging to boost accuracy (20). AI TB diagnosis and screening are benchmarked against microbiology (culture and/or GeneXpert, n=19), expert clinician assessment (including radiologists or pathologists, n=18) or a predefined composite microbiological and clinical criterion (n=19). Imaging-based studies typically report large sample sizes, some exceeding 150,000 participants, with a median study size of 1,780 (IQR 8,828, range 100-168,012) for internal training and validation, and 1,173 (IQR 5,528, range 36-25,598) for external validation (Table S3).

Performance and external validation: Reported accuracy, sensitivity, and specificity were high for diagnostic imaging studies on internal training and validation (AUC 0.94 [IQR 0.12, range 0.81-0.99], sensitivity 92% [IQR 10.2, range 67-100], and specificity 93% [IQR 17.1, range 64.3-98.7]). 58.5% of diagnostic or screening imaging studies reported model external validation (Figure 4A). Performance for diagnostic imaging studies on external validation was lower but still high (AUC 0.89 [IQR 0.14, range 0.66-0.98], sensitivity 90% [IQR 7, range 19-100], and specificity 85% [IQR 29, range 26-100]). A similar trend is noted for all screening studies (all data modalities), where the median AUC for training data was 0.94 (IQR 0.21, range 0.72-0.99), compared with 0.91 (IQR 0.10, range 0.67-1.00) for external validation.

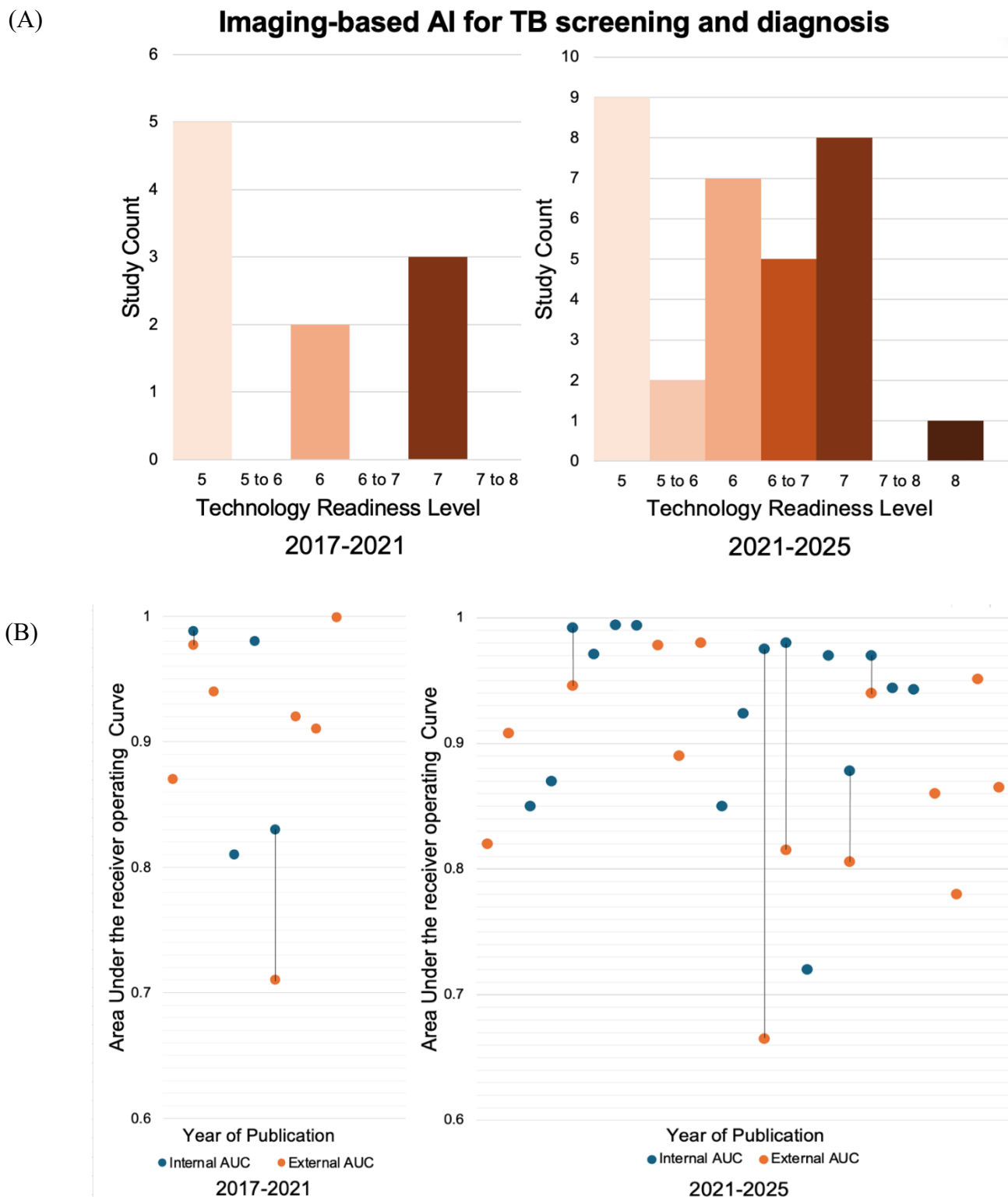


Figure 4. Imaging-based AI Tools for TB screening and diagnosis. Panel A shows technology readiness level (TRL) by publication year for AI tools interpreting imaging data. Panel B shows reported area under the curve (AUC) by year of publication; internal validation AUCs are in blue and external validation AUCs in orange. Vertical lines connect internal-external pairs from the same study, illustrating changes in performance between internal and external evaluation over time.

Technology readiness: The median TRL for screening and diagnostic imaging studies is 6 (IQR 1.5, range 5-8), with several commercial CXR AI platforms (e.g., CAD4TB, qXR, Lunit INSIGHT CXR) reaching TRL 7-8 and being prospectively evaluated and implemented in high-burden, real-world

settings. In contrast, clinical, omics, and microbiological AI screening and diagnostic studies are relatively immature (median TRL = 5, IQR 1.25, range 4-7). Median TRLs for imaging studies increased between the early publication period (January 2017-June 2021) and the later period (July 2021-August 2025) (Wilcoxon rank-sum test, $p=0.003$, Figure 4B).

Key gaps: The studies identified in the review report training of AI for TB diagnosis and screening in a limited number of countries (20 distinct countries), raising concerns about the generalizability to other contexts. Deployment of imaging AI relies on digital capture of chest radiography, which is currently limited in very low resource settings.

3.3. Regimen Design: Drug Resistance Diagnosis

Overview and input data: The 11 studies in this domain aim to diagnose resistance either to individual drugs ($n=4$) or classify the patient's presentation into one of several drug resistance classes e.g., drug-susceptible (DS), MDR or extensively drug-resistant (XDR) ($n=7$). The tools use pathogen whole-genome-sequencing (WGS) ($n=2$) or PCR probe melting temperatures ($n=1$) that detect pathogen DNA mutations as input for predicting individual drug resistance profiles. Other studies used radiography: CXR ($n=3$) and CT scan ($n=3$), of which three studies combined imaging with clinical data to classify presentation into resistance classes; two studies used only clinical variables at presentation ($n=2$). The reference standard is phenotypic drug susceptibility testing (DST) ($n=11$), with others also using Sanger sequencing ($n=1$) or comparison with other tools (TB-Profiler and Mykrobe) ($n=1$). The median study sample size for training was 540 (IQR 2,050, range 33-5,039).

Performance and external validation: Performance metrics indicate moderate to good accuracy in internal validation, with median AUC of 0.89 (IQR 0.04, range 0.78-0.95). For the three of the 11 models with external validation, the median AUC was 0.83 (IQR 0.31, range 0.63-0.94).

Technology readiness: Most AI models for drug resistance diagnosis are in the clinical validation phase, with a median TRL of 5 (IQR 0.5, range 4-6). No tools have achieved regulatory approval or broad adoption, but some demonstrate high accuracy in large-scale external validation efforts (e.g., GenTB externally validated on 20,403 patient isolates with an AUC of 0.94 [Table S3]), indicating readiness for clinical deployment in select contexts.

Key gaps: Gaps include limited causal formulation of the AI tasks. This can limit the identification of fit-for-task data for building generalizable models (e.g., CXR alone is unlikely to be sufficiently discriminatory for drug resistance diagnosis given it can only capture lung disease severity). Drug-resistant TB bacteria differ from susceptible bacteria by a few mutations, and these mutations have very limited or subtle effects on pathogenesis (21). Another related gap is limited external validation, including across geographic settings where resistance frequencies and TB genotypes vary.

3.4. Regimen Design: Regimen Composition

Overview and input data: Only a handful of studies met criteria for inclusion into this care category ($n=4$) and all have been published since 2020. These studies aimed to use AI to synthesize clinical, microbiological, and genomic data (e.g., WGS-based resistance profiles) to propose individualized drug regimens, with two specifically for MDR-TB. Three of these tools benchmark against laboratory gold standards (DST, microbiological, or molecular confirmation), whilst clinical decision support systems (CDSS) benchmark AI recommendations against clinician-designed regimens to support expert decision-making, especially in resource-limited settings. Studies reported a wide range of sample sizes, with a median study size of 1,228 (IQR 3,546, range 20-7,008).

Performance and external validation: Median internal AUC was 0.94 (IQR 0.25, range 0.70-0.95), with external validation performed in two studies. Among these, only one study reported performance in external validation (AUC 0.77).

Technology readiness: Similar to drug resistance diagnosis, the median TRL is 5.5 (IQR 1, range 5-6) indicating that technology is still in the prototype phase and requires clinical validation.

Key gaps: The identified regimen design models are limited by predominantly retrospective or prototype evaluations with minimal multi-country prospective external validation, and reliance on specific imaging/genomic datasets that may not generalize to diverse or low resource clinical settings.

3.5. Treatment Monitoring (Outcome & Adherence)

Overview and input data: The identified AI tools (n=12) used clinical variables (n=6), omics data (n=2), imaging data (n=1), or multimodal inputs combining imaging and clinical variables (n=3) to estimate cure, failure, relapse, mortality, or adverse reactions. Adherence-focused tools (n=8) used baseline clinical characteristics (typically from registry data) or digital adherence technologies (e.g., smart pillboxes, video observed therapy) to predict or detect non-adherence, respectively. The goal is to improve adherence in a personalized way and improve treatment outcomes. Median sample size was 790 (IQR 1,650, range 114-17,958) and 678 (IQR 2,388, range 100-243,726) for outcome and adherence prediction studies, respectively. The largest study by Rodrigues et al. (2024) predicted low adherence using a large national TB registry cohort (243,726 cases) relying on 8 baseline predictors, primarily comprising basic sociodemographic and self-reported clinical variables.

Performance and external validation: For internal validation, outcome prediction AI tools had a median AUC of 0.86 (IQR 0.09, range 0.81-0.91), and a median sensitivity and specificity of 89% (IQR 20, range 69-96) and 83% (IQR 21, range 70-93), respectively. Three of these 12 models underwent external validation, and the median AUC for external validation was lower at 0.74 (IQR 0.05, range 0.71-0.82). Adherence prediction tools had a median AUC of 0.81 (IQR 0.12, range 0.72-0.87), with none of the tools externally validated.

Technology readiness: Outcome models had a median TRL of 5 (IQR 0.5, range 4-6), typically reflecting analytical or retrospective validation. Adherence-monitoring tools similarly had a median TRL of 5 (IQR 1, range 4-7), with some tools (e.g., EvriMED smart pillbox) piloted in operational settings (TRL 7).

Key gaps: In contrast to outcome models, no adherence models were externally validated. Adherence models are largely developed and evaluated retrospectively, with limited external generalizability; additional research is needed to assess real-world utility and integration into existing care pathways.

4. Discussion

This scoping review demonstrates that AI applications in TB care are advancing rapidly but unevenly across the care cascade. AI tools for TB screening and diagnosis, particularly CXR interpretation, are comparatively mature, while applications addressing drug resistance diagnosis, regimen design, treatment monitoring, and adherence remain at earlier stages of development. This pattern mirrors broader trends in clinical AI, where image-based diagnostic tasks have progressed more quickly than tools designed to support longitudinal care and complex clinical decision-making (22). The dominance of diagnostic imaging applications also reflects structural advantages common to radiology-focused AI: Imaging data are standardized, reference standards are well defined, and tasks such as disease classification or triage are narrowly scoped (23). This has led to faster performance gains, more external validation, and earlier clinical adoption than AI applications that aim to predict treatment outcomes, guide therapy selection, or support adherence. The latter require integration of heterogeneous and longitudinal data, which are capabilities that remain challenging for current AI approaches and implementation in practice (24,25).

Within TB care, and more generally within AI decision support in healthcare, AI-based CXR interpretation represents a notable success story. Several tools have achieved high performance, undergone external validation, and been deployed in real-world, high-burden settings (26–30). Importantly, this progress has been driven not only by algorithmic innovation but also by coordinated investments in data infrastructure and evaluation. The FIND-led AI4Health initiative, which created large, curated, and geographically diverse AI-ready datasets for CXR-based TB diagnosis, exemplifies how standardized, high-quality data can accelerate both development and

clinical validation (31). By enabling benchmarking, external validation, and regulatory engagement, this effort has helped move CXR AI from proof-of-concept to implementation.

Our findings also underscore the limitations of concentrating AI innovation primarily on diagnosis and screening. Improvements in case detection alone are unlikely to achieve substantial population-level impact if downstream challenges (such as inappropriate regimen selection, undetected drug resistance, poor adherence, or delayed recognition of treatment failure) persist (32). Many of the largest gaps in the TB care cascade occur after diagnosis, particularly in settings with constrained laboratory capacity and limited clinical follow-up resources (33).

This review has highlighted that the relative immaturity of AI tools outside diagnosis and screening is linked to the lack of benchmarks and AI-ready datasets for these tasks. We found the few models for regimen design or treatment monitoring were trained on small or single-center, context-specific datasets, and had limited external validation, modest technology readiness levels, and uncertain generalizability (34,35). This stands in contrast to diagnostic imaging, where purpose-built datasets and benchmarks have enabled rapid iteration and comparison across tools. The success of CXR AI suggests that similar investments in AI-ready datasets and benchmarks could substantially accelerate progress in other domains of TB care. For example, globally representative datasets, e.g., from clinical trials, linking pathogen whole-genome sequencing, phenotypic DST, treatment regimens, and outcomes would support more robust resistance prediction and regimen optimization models (36,37). Likewise, longitudinal datasets integrating clinical variables, microbiological results, digital adherence technologies, and patient-reported outcomes could enable more effective AI tools for treatment monitoring and adherence support (38). Without such coordinated data efforts, advances in model architecture alone are unlikely to overcome current translational barriers.

Several additional cross-cutting gaps emerged across the TB AI literature. External validation remains inconsistent outside imaging-based diagnosis, raising concerns about robustness across diverse epidemiological and health system contexts (34). Also, we note a measurable drop in performance between internal and external validation across all care domains. TB is a heterogeneous disease, with clinical presentation and risk profiles shaped by local epidemiologic and health system factors (39,40). This heterogeneity may underlie the performance drop in external validation (39,40). As such, future development and evaluation of tools should prioritize robust multi-country validation prior to broader implementation, to ensure that performance estimates are reliable and that tools are appropriately tailored to diverse contexts. Many models rely on correlational features with limited causal relevance, particularly for resistance and outcome prediction (41). Our search did not identify studies that yet leverage LLM- or chatbot-based tools for patient monitoring or support. Recently, such tools have appeared promising in other chronic illnesses, including mental health; we identify this as an area of potential growth for AI in TB care (42–44). In addition, few studies assess integration into clinical workflows or evaluate downstream effects on patient outcomes or health system performance (45–48). These limitations are well recognized in clinical AI more broadly but are particularly consequential for TB, where failures at any stage of care directly contribute to transmission, resistance amplification, and mortality.

This scoping review has several limitations. We focused our search on PubMed to efficiently capture a comprehensive set of high-quality, peer-reviewed biomedical studies; while this approach focused the search, it is possible that additional studies indexed in other databases were not identified. We included only English-language publications, which may have limited representation of non-English literature. Abstract screening was assisted by ChatGPT-4, which helped streamline the review process, although all final selections were carefully reviewed manually. Finally, consistent with scoping review methodology, no formal exclusion was made based on critical appraisal or risk-of-bias assessment. However, we do describe several key quality metrics including sample size, gold standard metric, setting of the study and the conduct of external validation, which helped assess gaps and future directions in this area.

5. Conclusions and Implications

In conclusion, this scoping review suggests that the next phase of AI innovation in TB should move towards tools that address the full care cascade. The experience of AI-based CXR interpretation demonstrates what is possible when high-quality data, rigorous validation, and implementation pathways are aligned. Extending this model, especially through the development of AI-ready datasets for non-diagnostic tasks, represents a promising opportunity to accelerate clinically meaningful AI applications and support progress towards WHO End TB goals.

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Abbreviations

Model types and methods: AI indicates artificial intelligence; ML, machine learning; DL, deep learning; ANN, artificial neural network; CNN, convolutional neural network; DNN, deep neural network; MLP, multilayer perceptron; RF, random forest; DT, decision tree; CART, classification and regression tree; LR, logistic regression; SVM, support vector machine; NB, naïve Bayes; kNN, k-nearest neighbor; GBM, gradient boosting machine; XGB, extreme gradient boosting (XGBoost); LGBM, light gradient boosting machine; AdaBoost, adaptive boosting; Ens-DL, ensemble deep learning.

Imaging: CXR, chest radiograph; CT, computed tomography; DLAD, deep-learning-based automatic detection; U-Net, U-shaped convolutional neural network; ResNet, residual neural network; DenseNet, densely connected convolutional neural network; EfficientNet, efficient convolutional neural network family.

Data types: EHR, electronic health record; WGS, whole-genome sequencing; SNP, single-nucleotide polymorphism.

Performance and development: AUC, area under the receiver-operating-characteristic curve; Ext AUC, external-validation AUC; TRL, technology readiness level.

Clinical systems: CDSS, clinical decision support system; TB-DR, tuberculosis drug resistance.

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