

Table S1 CRISPR-based diagnostics for TB and drug-resistant TB

Effector	Strategy	Target	DNA extraction	Sample	Time	LOD	Sensitivity	Specificity	Ref
Cas1 2a	A recombinase polymerase amplification (RPA) step with CRISPR-based detection	<i>Mtb</i> IS6110	A combination strategy of beads beating, chemical lysis, and heating	Sputum	1.5h	5 copies/ μ L	79%	98%	²³⁷
Cas1 2b	TB-QUICK: loop-mediated isothermal amplification (LAMP) and CRISPR/Cas12b detection	<i>Mtb</i> IS6110	QIAamp Blood Mini Kit and QIAamp Circulating Nucleic Acid Kit	Pulmonary (sputum, bronchoalveolar lavage fluid) or plasma samples	2h	1.3 copy/ μ L	Pulmonary: 86.8% Plasma: 41.2% for AFB-positive and 31.7% for AFB-negative patients	95.2%	²³⁸
Lba Cas1 2a	CRISPR detection of circulating cell-free Mycobacterium tuberculosis DNA in adults and children, including children with HIV	<i>Mtb</i> IS6110, <i>esxB</i> , <i>gyrB</i>	The Quick-cfDNA Serum & Plasma	Serum	2h	0.06 copy/ μ L	68%-100%	-	¹⁶⁰
LwCas13a	A sensitive Mycobacterium tuberculosis (MTB) complex polymerase chain reaction (PCR)-CRISPR/Cas13a detection method (CRISPR-MTB)	<i>Mtb</i> IS1081	Conventional DNA extraction	Sputum, BALF, and pus samples		1 target sequence copy/ μ L	97.2%-100%	95.5%	²³⁹
Cas9	A CRISPR/Cas-9-mediated fluorescent strategy utilizing fluorescence resonance energy transfer (FRET)	High-variable region of <i>M. tuberculosis</i> 16S rDNA fragment	-	Simulated sputum samples	2h	20 CFU mL ⁻¹	-	-	²⁴⁰
Cas1 2b	CRISPR/CRISPR-associated 12b nuclease CRISPR/Cas12b-based multiple cross displacement amplification technique (CRISPR-MCDA)	<i>Mtb</i> IS6110	Genomic DNA extraction kits	Sputum	70min	5 fg/ μ L of genomic DNA extracted from the MTB reference strain H37Rv.	-	100%	²⁴¹
Cas9	Finding Low Abundance Sequences by Hybridization (FLASH)	FLASH to amplify 52 candidate genes probably associated with resistance to first- and second-line drugs in the <i>Mtb</i> reference strain (H37Rv)	Mechanical disruption method	The laboratory H37Rv <i>Mtb</i> reference strain, cultured isolates, and sputum	-	-	-	Drug resistance predictions for 15/16 (93.7%) clinical samples	¹⁷¹
Cas9	A Cas9/gRNA-assisted quantitative real-time PCR (qRT-PCR) (CARP) assay	S531 and H526 positions in the rifampicin (RIF)-resistance-determining region (RRDR) of the <i>Mtb</i> <i>rpoB</i> gene	-	<i>M. tuberculosis</i> genomic DNA template	~4h	less than 0.0001 femtogram of mycobacterial genomic DNA.	-	100%	¹⁷⁰