THE LANCET Infectious Diseases

Supplementary appendix 2

This appendix formed part of the original submission and has been peer reviewed. We post it as supplied by the authors.

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Appendix to: Treatment outcomes 24 months after initiating short, all-oral bedaquiline-containing or injectable-containing rifampicin-resistant tuberculosis treatment regimens in South Africa: a retrospective cohort study

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Appendix Table 1: Distribution of included patients and select characteristics by province of treatment

	Regimen			Loss to Follow-		PLHIV Receiving
Province	Group	N	Success, %	up, %	PLHIV, %	ART, %
Eastern Cape -	Bedaquiline	201	75	5	64	98
Eastern Cape	Injectable	230	54	13	66	97
Free State -	Bedaquiline	31	71	3	74	96
Free State -	Injectable	20	50	5	65	85
Cautona	Bedaquiline	63	62	14	79	98
Gauteng -	Injectable	24	54	21	67	94
Kwazulu-Natal —	Bedaquiline	163	75	6	74	100
	Injectable	232	59	14	72	95
Limnono	Bedaquiline	16	62	6	62	100
Limpopo -	Injectable	11	55	0	64	71
Maumalanga	Bedaquiline	99	66	8	85	92
Mpumalanga -	Injectable	70	57	10	77	93
North West -	Bedaquiline	75	61	9	72	94
North West —	Injectable	41	63	15	66	63
Northern Cape —	Bedaquiline	29	59	0	55	100
	Injectable	30	37	20	67	95
Masters Cons	Bedaquiline	11	55	0	64	100
Western Cape -	Injectable	41	66	17	44	94

Appendix Table 2: Characteristics of patients eligible for and receiving a short regimen who were ultimately included vs excluded (N=2769)

	Included	Excluded	p-value
N	1387	1382	
Patient Characteristics			
Median (IQR) Age	37 (30 to 47)	36 (30 to 44)	0.0417
Male	812 (58.5%)	798 (57.7%)	0.6974
HIV Infection	967 (69.7%)	1013 (73.3%)	0.0409
If HIV Infected, on ART	918 (94.9%)	950 (93.8%)	0.3116
Microbiologic Findings at Baseline*			
If Result Available, AFB Smear Positive at Baseline	545 (46.4%)	563 (49.6%)	0.1423
If Result Available, Culture Positive at Baseline	676 (72.7%)	722 (77.9%)	0.0110
If Result Available, Culture or AFB or Xpert Positive at Baseline	1207 (91.2%)	1174 (90.8%)	0.7956
Previous Treatment			
Previously Treated for TB with First Line Drugs	560 (40.4%)	577 (41.8%)	0.4856
Treatment Received			
Received Moxifloxacin or Levofloxacin	1387 (100%)	1349 (97.6%)	
Received Bedaquiline	688 (49.6%)	946 (68.5%)	
Received Linezolid	0 (0%)	160 (11.6%)	
Received Cycloserine/Terizidone	0 (0%)	211 (15.3%)	
Received Clofazimine	1387 (100%)	1271 (92%)	
Received Injectable	699 (50.4%)	968 (70%)	
Received Amikacin	106 (7.6%)	115 (8.3%)	
Received Kanamycin	595 (42.9%)	854 (61.8%)	
Received Capreomycin	1 (0.1%)	3 (0.2%)	
Received Streptomycin	0 (0%)	0 (0%)	
Mean Number of Other Drugs Received	2.9	2.9	
End of Treatment Outcomes			
Number with Treatment Success	928	922	0.9466
Number with Failure	19	25	0.4402
Number Dying During Treatment	276	250	0.2440
Number Lost to follow-up During Treatment	164	185	0.2375

Abbreviations: AFB, acid fast bacilli; ART, antiretroviral treatment; IQR, interquartile range; TB, tuberculosis

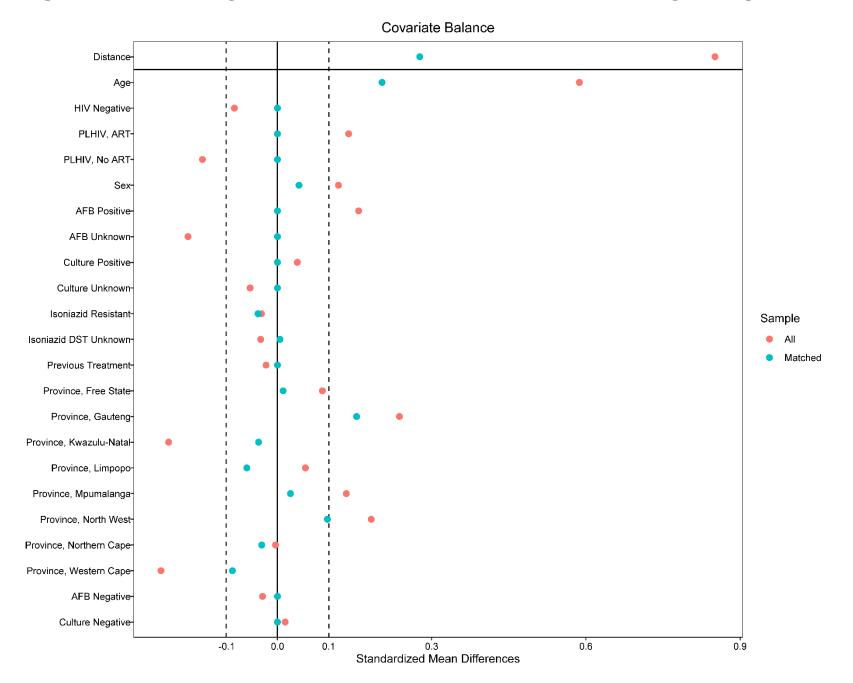
Notes: p-value calculated using Kruskal-Wallis test for medians and chi-square test for continuous variables

Appendix Table 3: Univariable analysis results to support decisions on exact and propensity score matching in multivariable analysis

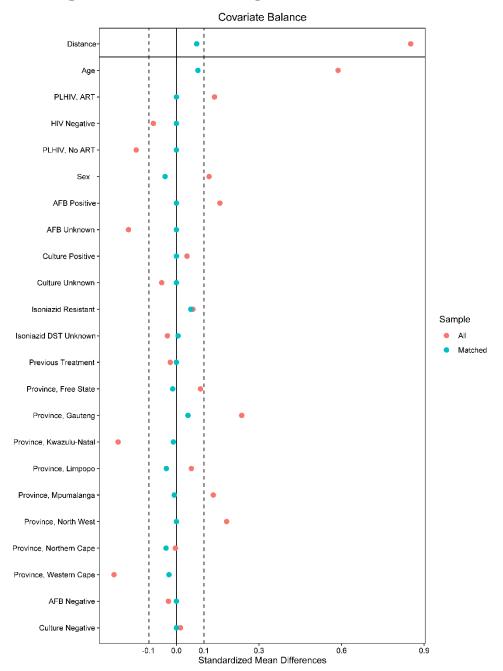
		Risk Difference per 100
Characteristic	Success/Total	(95% CI)
Age, median (IQR), per 10y increase	37 (30 to 47)	-1 (-3 to 1)
Sex		
Female	373/575	0 (reference)
Male	501/812	-3 (-8 to 2)
AFB Smear at Baseline		
Negative	413/629	0 (reference)
Positive	347/545	-2 (-7 to 3)
Unknown	114/213	-12 (-20 to -4)
Culture at Baseline		
Negative	182/254	0 (reference)
Positive	427/676	-8 (-15 to -2)
Unknown	265/457	-14 (-21 to -7)
HIV-Status		
Negative	275/420	0 (reference)
Positive on ART	582/918	-2 (-8 to 3)
Positive not on ART	17/49	-31 (-45 to -17)
Previous Treatment		
Never Treated	540/827	0 (reference)
Treated with First Line Drugs	334/560	-6 (-11 to -1)
Isoniazid Resistance		
Susceptible	325/518	0 (reference)
Resistant	362/553	3 (-3 to 8)
Unknown	187/316	-4 (-10 to 3)

Abbreviations: IQR, interquartile range; AFB, acid fast bacilli; ART, antiretroviral therapy; 95% CI, 95% confidence interval.

Appendix Figure 1: Love Plot of Changes in Standardized Mean Difference Before and After Matching with Caliper of 0.5



Appendix Figure 2: Love Plot of Changes in Standardized Mean Difference Before and After Matching with Caliper of 0.2



Appendix Table 4: Primary Analysis Outcomes from Multivariable Analysis using caliper distance of 0.2

Total Population Success vs. All Other Outcomes 478/868 396/699 394 (67%) 16 (10.	Outcome	Bedaquiline Group (events/total)	Injectable Group (events/total)	Matched Pairs (% Matched)	Adjusted RD per 100 (95% CI)
Success vs. Ail Other Outcomes 478/688 396/699 394 (57%) 16 (10.2	Total Population	(events/total)	(events/total)	(% Matched)	(95% CI)
Survival vs. Death Speak Fire Survival with Treatment Failure/Recurrence 528688 500/699 394 (57%) 10 (4.)		478/688	396/699	394 (57%)	16 (10, 23)
Disease-Free Survival vs. Survival with Treatment Failure/Recurrence \$22/526 483/500 281 (55%) 2 (-0.1, Lost to Follow-up vs. All Other Outcomes 478/688 396/699 334 (57%) -6. (-10. PLHIV Receiving ART Success vs. All Other Outcomes 337/478 245/440 289 (63%) 18 (10. Survival vs. Death 245/440 289 (63%) 13 (6. Survival vs. Death 245/440 289 (63%) 13 (6. Survival vs. Death 245/440 289 (63%) 13 (6. Survival vs. Death 245/440 289 (63%) 14 (-10. Survival vs. Death 245/440 289 (63%) 14 (-10. Survival vs. Death 245/440 289 (63%) 4 (-9. Survival vs. Death 245/440 289 (63%) 4 (-9. Survival vs. Death 245/440 289 (63%) 4 (-9. Survival vs. Death 245/440				, ,	10 (4, 16)
Lost to Follow-up vs. All Other Outcomes	Disease-Free Survival vs. Survival with Treatment Failure/Recurrence			` ,	2 (-0.1, 4)
PLHIV Receiving ART Success vs. All Other Outcomes 337/478 302/440 289 (63%) 18 (10, 2		478/688		. ,	-6 (-10, -2)
Survival vs. Death 372/478 302/440 289 (63%) 13 (6.2	PLHIV Receiving ART			, ,	, , ,
Survival vs. Death 372/478 302/440 289 (63%) 13 (6.2)	Success vs. All Other Outcomes	337/478	245/440	289 (63%)	18 (10, 26)
Lost to Follow-up vs. All Other Outcomes 31/478 51/440 289 (63%) 4-(-9)	Survival vs. Death	372/478	302/440	289 (63%)	13 (6, 21)
People HIV-Negative Success vs. All Other Outcomes 138/195 137/225 105 (50%) 19 (6, 5	Disease-Free Survival vs. Survival with Treatment Failure/Recurrence	368/372	296/302	197 (58%)	1 (-1, 3)
Success vs. All Other Outcomes 138/195 137/225 105 (50%) 19 (6, 3	Lost to Follow-up vs. All Other Outcomes	31/478	51/440	289 (63%)	-4 (-9, 1)
Survival vs. Death					
Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 147/147 169/179 71 (44%) 6 (0.3.)	Success vs. All Other Outcomes	138/195	137/225	105 (50%)	19 (6, 32)
Lost to Follow-up vs. All Other Outcomes 91/95 32/225 105 (50%) -9 (-16, AFB Smear Positive at Baseline Success vs. All Other Outcomes 205/297 142/248 170 (62%) 15 (5, 2 (5	Survival vs. Death	147/195	179/225	105 (50%)	6 (-6, 17)
AFB Smear Positive at Baseline Success vs. All Other Outcomes 205/297 142/248 170 (62%) 15 (5, 2 Survival vs. Death 224/297 186/248 170 (62%) 17	Disease-Free Survival vs. Survival with Treatment Failure/Recurrence	147/147	169/179	71 (44%)	6 (0.3, 11)
Success vs. All Other Outcomes 205/297 142/248 170 (62%) 15 (5, 2)	Lost to Follow-up vs. All Other Outcomes	9/195	32/225	105 (50%)	-9 (-16, -1)
Survival vs. Death 224/297 186/248 170 (62%) 6 (-3, 170 (62%) 5 (-4, 170 (62%) 5 (-4, 170 (62%	AFB Smear Positive at Baseline				
Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 223/224 177/186 108 (53%) 5 (1, Lost to Follow-up vs. All Other Outcomes 18/297 35/248 170 (62%) -5 (-11, AFB Smear Negative at Baseline Success vs. All Other Outcomes 224/307 189/322 173 (55%) 16 (6, 2 Survival vs. Death 244/307 230/322 173 (55%) 16 (7, 2 Survival vs. Death 244/307 230/322 173 (55%) 16 (7, 2 Survival vs. Death 244/307 230/322 173 (55%) 16 (7, 2 Survival vs. Death 242/244 223/230 118 (50%) 0 (-2, Lost to Follow-up vs. All Other Outcomes 18/307 34/322 173 (55%) 0 (-6, Previously Treated with First-Line Drugs Success vs. All Other Outcomes 181/274 153/286 158 (56%) 11 (1, 2 Survival vs. Death 198/274 188/286 158 (56%) 9 (-1, 3 Survival vs. Death 198/274 188/286 158 (56%) 9 (-1, 3 Survival vs. Death 198/274 188/286 158 (56%) 9 (-1, 3 Survival vs. Disease-Free Survival vs. All Other Outcomes 15/274 28/286 158 (56%) -1 (-7, Never Previously Treated for Tuberculosis Success vs. All Other Outcomes 297/414 243/413 245 (59%) 14 (6, 2 Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 4 Survival vs. Death 10 Survival vs. Survival with Treatment Failure/Recurrence 297/414 243/413 245 (59%) 14 (6, 2 Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 4 Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 4 Survival vs. Death 328/414 312/413 245 (59%) 2 (-1, 5 Survival vs. Death 10 Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 5 Survival vs. Death 10 Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 5 Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 5 Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%)	Success vs. All Other Outcomes	205/297	142/248	170 (62%)	15 (5, 25)
Lost to Follow-up vs. All Other Outcomes 18/297 35/248 170 (62%) -5 (-11, AFB Smear Negative at Baseline	Survival vs. Death	224/297	186/248	170 (62%)	6 (-3, 16)
AFB Smear Negative at Baseline Success vs. All Other Outcomes 224/307 189/322 173 (55%) 16 (6, 2 Survival vs. Death 244/307 230/322 173 (55%) 16 (7, 2 Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 242/244 223/230 118 (50%) 0 (-2, 2 Lost to Follow-up vs. All Other Outcomes 18/307 34/322 173 (55%) 0 (-6, 7 Previously Treated with First-Line Drugs Success vs. All Other Outcomes 181/274 153/286 158 (56%) 11 (1, 2 Survival vs. Death 198/274 188/286 158 (56%) 9 (-1, 1 Lost to Follow-up vs. All Other Outcomes 15/274 28/286 158 (56%) 4 (-1, 2 Never Previously Treated for Tuberculosis Success vs. All Other Outcomes 297/414 243/413 245 (59%) 7 (-1, 2 Disease-Free Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 2 Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 2)	Disease-Free Survival vs. Survival with Treatment Failure/Recurrence	223/224	177/186	108 (53%)	5 (1, 9)
Success vs. All Other Outcomes 224/307 189/322 173 (55%) 16 (6, 2 Survival vs. Death 244/307 230/322 173 (55%) 16 (7, 2 Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 242/244 223/230 118 (50%) 0 (-2, 2 Lost to Follow-up vs. All Other Outcomes 18/307 34/322 173 (55%) 0 (-6, 2 Previously Treated with First-Line Drugs 8 181/274 153/286 158 (56%) 11 (1, 2 Survival vs. Death 198/274 188/286 158 (56%) 9 (-1, 3) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 196/198 181/188 97 (50%) 4 (-1, 4) Lost to Follow-up vs. All Other Outcomes 15/274 28/286 158 (56%) -1 (-7, 8) Never Previously Treated for Tuberculosis 297/414 243/413 245 (59%) 14 (6, 2, 2, 2, 2, 2, 2, 2, 2, 2, 2, 2, 2, 2,	Lost to Follow-up vs. All Other Outcomes	18/297	35/248	170 (62%)	-5 (-11, 1)
Survival vs. Death 244/307 230/322 173 (55%) 16 (7, 2) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 242/244 223/230 118 (50%) 0 (-2, 2) Lost to Follow-up vs. All Other Outcomes 18/307 34/322 173 (55%) 0 (-6, 2) Previously Treated with First-Line Drugs 5 5 5 5 11 (1, 2) 5 5 11 (1, 2) 5 5 11 (1, 2) 5 6 158 (56%) 11 (1, 2) 11 (1, 2) 5 11 (1, 2) 5 11 (1, 2) 11	AFB Smear Negative at Baseline				
Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 242/244 223/230 118 (50%) 0 (-2, 18) Lost to Follow-up vs. All Other Outcomes 18/307 34/322 173 (55%) 0 (-6, 18) Previously Treated with First-Line Drugs Success vs. All Other Outcomes 181/274 153/286 158 (56%) 11 (1, 2 (1, 2 (1) (1) (1) (1) (1) (1) (1) (1) (1) (1)	Success vs. All Other Outcomes	224/307	189/322	173 (55%)	16 (6, 25)
Lost to Follow-up vs. All Other Outcomes 18/307 34/322 173 (55%) 0 (-6, Previously Treated with First-Line Drugs Success vs. All Other Outcomes 181/274 153/286 158 (56%) 11 (1, 2 survival vs. Death Survival vs. Death 198/274 188/286 158 (56%) 9 (-1, 3 survival vs. Survival with Treatment Failure/Recurrence Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 196/198 181/188 97 (50%) 4 (-1, 2 survival vs. All Other Outcomes Lost to Follow-up vs. All Other Outcomes 15/274 28/286 158 (56%) -1 (-7, 3 survival vs. All Other Outcomes Success vs. All Other Outcomes 297/414 243/413 245 (59%) 14 (6, 2 survival vs. Death Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 3 survival vs. Survival with Treatment Failure/Recurrence Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 3 survival vs. Survival with Treatment Failure/Recurrence	Survival vs. Death	244/307	230/322	173 (55%)	16 (7, 25)
Previously Treated with First-Line Drugs Success vs. All Other Outcomes 181/274 153/286 158 (56%) 11 (1, 2) Survival vs. Death 198/274 188/286 158 (56%) 9 (-1, 2) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 196/198 181/188 97 (50%) 4 (-1, 2) Lost to Follow-up vs. All Other Outcomes 15/274 28/286 158 (56%) -1 (-7, 2) Never Previously Treated for Tuberculosis 297/414 243/413 245 (59%) 14 (6, 2) Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 2) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 2)	Disease-Free Survival vs. Survival with Treatment Failure/Recurrence	242/244	223/230	118 (50%)	0 (-2, 2)
Success vs. All Other Outcomes 181/274 153/286 158 (56%) 11 (1, 2) Survival vs. Death 198/274 188/286 158 (56%) 9 (-1, 2) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 196/198 181/188 97 (50%) 4 (-1, 2) Lost to Follow-up vs. All Other Outcomes 15/274 28/286 158 (56%) -1 (-7, 2) Never Previously Treated for Tuberculosis 297/414 243/413 245 (59%) 14 (6, 2) Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 2) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 2)	Lost to Follow-up vs. All Other Outcomes	18/307	34/322	173 (55%)	0 (-6, 6)
Survival vs. Death 198/274 188/286 158 (56%) 9 (-1, -7) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 196/198 181/188 97 (50%) 4 (-1, -7) Lost to Follow-up vs. All Other Outcomes 15/274 28/286 158 (56%) -1 (-7, -7) Never Previously Treated for Tuberculosis Success vs. All Other Outcomes 297/414 243/413 245 (59%) 14 (6, 2, 2, 2, 2, 2, 2, 2, 2, 2, 2, 2, 2, 2,					
Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 196/198 181/188 97 (50%) 4 (-1, 20%) Lost to Follow-up vs. All Other Outcomes 15/274 28/286 158 (56%) -1 (-7, 20%) Never Previously Treated for Tuberculosis Success vs. All Other Outcomes 297/414 243/413 245 (59%) 14 (6, 20%) Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 20%) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 20%)	Success vs. All Other Outcomes	181/274	153/286	158 (56%)	11 (1, 22)
Lost to Follow-up vs. All Other Outcomes 15/274 28/286 158 (56%) -1 (-7, Never Previously Treated for Tuberculosis Success vs. All Other Outcomes 297/414 243/413 245 (59%) 14 (6, 2 32) Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 2 32) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 2 32)	Survival vs. Death	198/274	188/286	158 (56%)	9 (-1, 19)
Never Previously Treated for Tuberculosis Success vs. All Other Outcomes 297/414 243/413 245 (59%) 14 (6, 2) Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 2) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 2)	Disease-Free Survival vs. Survival with Treatment Failure/Recurrence	196/198	181/188	97 (50%)	4 (-1, 9)
Success vs. All Other Outcomes 297/414 243/413 245 (59%) 14 (6, 2 (59%)) Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 2 (59%)) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 2 (59%))	Lost to Follow-up vs. All Other Outcomes	15/274	28/286	158 (56%)	-1 (-7, 4)
Survival vs. Death 328/414 312/413 245 (59%) 7 (-1, 7) Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1, 7)	•				
Disease-Free Survival vs. Survival with Treatment Failure/Recurrence 326/328 302/312 178 (56%) 2 (-1,	Success vs. All Other Outcomes			` '	14 (6, 22)
					7 (-1, 14)
Lost to Follow-up vs. All Other Outcomes 29/414 59/413 245 (59%) -7 (-12,					2 (-1, 5)
	Lost to Follow-up vs. All Other Outcomes	29/414	59/413	245 (59%)	-7 (-12, -2)

Abbreviations: RD, risk difference; AFB, acid fast bacilli; ART, antiretroviral therapy; CI, confidence interval. Note: Estimates comes from a model where patients were matched exactly on HIV, ART, previous treatment, and baseline smear and culture results, while propensity score matched on age, sex, province of treatment, and isoniazid resistance.

Appendix Table 5: Risk of mortality during or post-treatment estimated via multivariable analysis among patients in the injectable group

	Survived	Death During Treatment	Death Post-Treatment	aRD per 100 (95%CI)	aRD per 100 (95%CI) Death During Treatment	aRD per 100 (95%CI) Death Post-Treatment vs.
Characteristic	(n=500)	(n=159)	(n=40)	Death Ever vs. Survival	vs. Survival	Survival
Age, median (IQR), per 10y	33 (27 to 40)	38 (31 to 47)	33 (27 to 40)	6 (3 to 9)	6 (4 to 8)	0.4 (0.01 to 1)
Sex						
Female	228 (73.5%)	63 (20.3%)	19 (6.1%)	0 (reference)	0 (reference)	0 (reference)
Male	272 (69.9%)	96 (24.7%)	21 (5.4%)	1 (-4 to 7)	2 (-1 to 6)	-1 (-2 to 1)
AFB Smear at Baseline						
Negative	230 (71.4%)	72 (22.4%)	20 (6.2%)	0 (reference)	0 (reference)	0 (reference)
Positive	186 (75%)	52 (21%)	10 (4%)	-1 (-7 to 5)	2 (-2 to 5)	-2 (-4 to 0.4)
Unknown	84 (65.1%)	35 (27.1%)	10 (7.8%)	3 (-8 to 14)	1 (-7 to 10)	1 (-3 to 6)
Culture at Baseline						
Negative	96 (76.2%)	26 (20.6%)	4 (3.2%)	0 (reference)	0 (reference)	0 (reference)
Positive	244 (73.1%)	69 (20.7%)	21 (6.3%)	8 (1 to 14)	4 (-0.2 to 9)	2 (-1 to 4)
Unknown	160 (66.9%)	64 (26.8%)	15 (6.3%)	9 (1 to 18)	6 (-1 to 13)	3 (0.2 to 5)
HIV-Status						
Negative	179 (79.6%)	39 (17.3%)	7 (3.1%)	0 (reference)	0 (reference)	0 (reference)
Positive on ART	302 (68.6%)	107 (24.3%)	31 (7%)	8 (2 to 15)	6 (2 to 10)	2 (0.1 to 4)
Positive not on ART	19 (55.9%)	13 (38.2%)	2 (5.9%)	22 (4 to 41)	18 (1 to 35)	0 (-3 to 2)
Previous Treatment						
Never Treated	312 (75.5%)	84 (20.3%)	17 (4.1%)	0 (reference)	0 (reference)	0 (reference)
Treated with First Line Drugs	188 (65.7%)	75 (26.2%)	23 (8%)	8 (1 to 15)	4 (-1 to 9)	2 (-0.3 to 5)
Isoniazid Resistance						
Susceptible	177 (70.5%)	56 (22.3%)	18 (7.2%)	0 (reference)	0 (reference)	0 (reference)
Resistant	217 (76.4%)	51 (18%)	16 (5.6%)	-2 (-8 to 5)	-3 (-8 to 2)	0 (-3 to 3)
Unknown	106 (64.6%)	52 (31.7%)	6 (3.7%)	7 (-1 to 15)	8 (1 to 15)	-2 (-4 to 1)

Abbreviations: AFB, acid fast bacilli; aRD, adjusted risk difference; 95% CI, 95% confidence interval; ART, antiretroviral treatment; IQR, interquartile range

^{*}Notes: Comparisons were done only on patients who were alive during the analytical period (e.g., analyses of post-treatment death vs. survival were only done on patients who were alive at their treatment end date). Percentages represent row percentages. Model is adjusted for all factors listed here, including province of treatment.

Appendix Table 6: Risk of mortality during or post-treatment estimated via multivariable analysis among patients in the bedaquiline group

					aRD per 100 (95%CI)	aRD per 100 (95%CI)
	Survived	Death During Treatment	Death Post Treatment	aRD per 100 (95%CI)	Death During Treatment vs.	Death Post-Treatment vs.
Characteristic	(n=526)	(n=117)	(n=45)	Death Ever vs. Survival	Survival	Survival
Age, median (IQR), per 10y	41 (33 to 51)	44 (35 to 57)	38 (31 to 56)	4 (2 to 6)	4 (2 to 5)	0 (-1 to 1)
Sex						
Female	203 (76.6%)	45 (17%)	17 (6.4%)	0 (reference)	0 (reference)	0 (reference)
Male	323 (76.4%)	72 (17%)	28 (6.6%)	0 (-5 to 4)	-2 (-5 to 1)	-1 (-3 to 0.1)
AFB Smear at Baseline						
Negative	244 (79.5%)	46 (15%)	17 (5.5%)	0 (reference)	0 (reference)	0 (reference)
Positive	224 (75.4%)	52 (17.5%)	21 (7.1%)	3 (-3 to 8)	2 (-1 to 6)	-1 (-2 to 1)
Unknown	58 (69%)	19 (22.6%)	7 (8.3%)	7 (-6 to 19)	3 (-2 to 9)	2 (-3 to 7)
Culture at Baseline						
Negative	108 (84.4%)	16 (12.5%)	4 (3.1%)	0 (reference)	0 (reference)	0 (reference)
Positive	263 (76.9%)	50 (14.6%)	29 (8.5%)	8 (2 to 14)	0 (-3 to 2)	7 (4 to 10)
Unknown	155 (71.1%)	51 (23.4%)	12 (5.5%)	10 (1 to 18)	6 (-0.3 to 11)	2 (-0.1 to 3)
HIV-Status						
Negative	147 (75.4%)	35 (17.9%)	13 (6.7%)	0 (reference)	0 (reference)	0 (reference)
Positive on ART	372 (77.8%)	75 (15.7%)	31 (6.5%)	1 (-4 to 6)	-1 (-4 to 1)	-3 (-5 to -0.4)
Positive not on ART	7 (46.7%)	7 (46.7%)	1 (6.7%)	25 (-7 to 57)	16 (-16 to 48)	6 (-21 to 32)
Previous Treatment						
Never Treated	328 (79.2%)	62 (15%)	24 (5.8%)	0 (reference)	0 (reference)	0 (reference)
Treated with First Line Drugs	198 (72.3%)	55 (20.1%)	21 (7.7%)	5 (0.4 to 10)	2 (-1 to 5)	2 (-1 to 4)
Isoniazid Resistance						
Susceptible	200 (74.9%)	52 (19.5%)	15 (5.6%)	0 (reference)	0 (reference)	0 (reference)
Resistant	213 (79.2%)	35 (13%)	21 (7.8%)	-4 (-9 to 2)	-5 (-9 to -1)	1 (-1 to 2)
Unknown	113 (74.3%)	30 (19.7%)	9 (5.9%)	2 (-7 to 11)	0 (-7 to 7)	1 (-0.5 to 3)

Abbreviations: AFB, acid fast bacilli; aRD, adjusted risk difference; 95% CI, 95% confidence interval; ART, antiretroviral treatment; IQR, interquartile range

^{*}Notes: Comparisons were done only on patients who were alive during the analytical period (e.g., analyses of post-treatment death vs. survival were only done on patients who were alive at their treatment end date). Percentages represent row percentages. Model is adjusted for all factors listed here, including province of treatment.

Appendix Table 7: Factors for mortality by early vs. late death as estimated by multivariable analysis

Characteristic	Survived (n=1026)	Early Death: died ≤3 months after treatment start (n=171)	Late Death: died >3 to 24 months after treatment start (n=190)	aRD per 100 (95%CI) Death Ever vs. Survival	aRD per 100 (95%CI) Early Death vs. Survival	aRD per 100 (95%CI) Late Death vs. Survival
Regimen	(1323)	()	()			
Injectable-Containing	500 (71.5%)	108 (15.5%)	91 (13%)	0 (reference)	0 (reference)	0 (reference)
Bedaquiline-Containing	526 (76.5%)	63 (9.2%)	99 (14.4%)	-8 (-11 to -5)	-6 (-9 to -4)	0 (-3 to 3)
Age, median (IQR), per 10y	36 (29 to 46)	40 (34 to 53)	38 (31 to 49)	5 (3 to 6)	3 (2 to 3)	2 (1 to 3)
Sex			· · · · · ·	<u> </u>	<u> </u>	<u> </u>
Female	431 (75%)	68 (11.8%)	76 (13.2%)	0 (reference)	0 (reference)	0 (reference)
Male	595 (73.3%)	103 (12.7%)	114 (14%)	0 (-3 to 4)	0 (-1 to 1)	0 (-1 to 1)
AFB Smear at Baseline						
Negative	474 (75.4%)	68 (10.8%)	87 (13.8%)	0 (reference)	0 (reference)	0 (reference)
Positive	410 (75.2%)	65 (11.9%)	70 (12.8%)	2 (-2 to 5)	1 (-1 to 4)	-2 (-4 to 1)
Unknown	142 (66.7%)	38 (17.8%)	33 (15.5%)	6 (1 to 11)	3 (0.1 to 6)	4 (-2 to 9)
Culture at Baseline						
Negative	204 (80.3%)	21 (8.3%)	29 (11.4%)	0 (reference)	0 (reference)	0 (reference)
Positive	507 (75%)	71 (10.5%)	98 (14.5%)	7 (3 to 10)	-1 (-3 to 1)	4 (1 to 6)
Unknown	315 (68.9%)	79 (17.3%)	63 (13.8%)	9 (3 to 15)	3 (-0.2 to 7)	2 (-1 to 5)
HIV-Status						
Negative	326 (77.6%)	51 (12.1%)	43 (10.2%)	0 (reference)	0 (reference)	0 (reference)
Positive on ART	674 (73.4%)	104 (11.3%)	140 (15.3%)	6 (3 to 10)	0 (-2 to 1)	5 (2 to 8)
Positive not on ART	26 (53.1%)	16 (32.7%)	7 (14.3%)	22 (6 to 38)	6 (-7 to 19)	8 (-5 to 21)
Previous Treatment						
Never Treated	640 (77.4%)	99 (12%)	88 (10.6%)	0 (reference)	0 (reference)	0 (reference)
Treated with First Line Drugs	386 (68.9%)	72 (12.9%)	102 (18.2%)	8 (4 to 12)	1 (-1 to 3)	7 (3 to 11)
Isoniazid Resistance						
Susceptible	377 (72.8%)	62 (12%)	79 (15.3%)	0 (reference)	0 (reference)	0 (reference)
Resistant	430 (77.8%)	55 (9.9%)	68 (12.3%)	-4 (-8 to 1)	-1 (-3 to 0.5)	-2 (-5 to 1)
Unknown	219 (69.3%)	54 (17.1%)	43 (13.6%)	2 (-4 to 9)	0 (-3 to 4)	0 (-4 to 5)

Abbreviations: AFB, acid fast bacilli; aRD, adjusted risk difference; 95% CI, 95% confidence interval; ART, antiretroviral treatment; IQR, interquartile range

^{*}Notes: Comparisons were done only on patients who were alive during the analytical period (e.g., analyses of death >3 months vs. survival were only done on patients who were alive after 3 months of treatment). Percentages represent row percentages. Model is adjusted for all factors listed here, including province of treatment.

Appendix Table 8: Risk of early or late mortality as estimated by multivariable analysis among patients in the injectable group

		Early Death: died ≤3 months	Late Death: died >3 to 24			
	Survived	after treatment start	months after treatment start	aRD per 100 (95%CI)	aRD per 100 (95%CI)	aRD per 100 (95%CI)
Characteristic	(n=500)	(n=108)	(n=91)	Death Ever vs. Survival	Early Death vs. Survival	Late Death vs. Survival
Age, median (IQR), per 10y	33 (27 to 40)	39 (31 to 46)	35 (29 to 44)	6 (3 to 9)	4 (3 to 6)	2 (0 to 3)
Sex						
Female	228 (73.5%)	42 (13.5%)	40 (12.9%)	0 (reference)	0 (reference)	0 (reference)
Male	272 (69.9%)	66 (17%)	51 (13.1%)	1 (-4 to 7)	0 (-3 to 4)	-1 (-3 to 1)
AFB Smear at Baseline						
Negative	230 (71.4%)	42 (13%)	50 (15.5%)	0 (reference)	0 (reference)	0 (reference)
Positive	186 (75%)	40 (16.1%)	22 (8.9%)	-1 (-7 to 5)	2 (-2 to 5)	-4 (-9 to -0.3)
Unknown	84 (65.1%)	26 (20.2%)	19 (14.7%)	3 (-8 to 14)	2 (-5 to 9)	-1 (-8 to 5)
Culture at Baseline						
Negative	96 (76.2%)	15 (11.9%)	15 (11.9%)	0 (reference)	0 (reference)	0 (reference)
Positive	244 (73.1%)	48 (14.4%)	42 (12.6%)	8 (1 to 14)	3 (-1 to 6)	5 (1 to 9)
Unknown	160 (66.9%)	45 (18.8%)	34 (14.2%)	9 (1 to 18)	5 (-2 to 11)	6 (1 to 10)
HIV-Status						
Negative	179 (79.6%)	31 (13.8%)	15 (6.7%)	0 (reference)	0 (reference)	0 (reference)
Positive on ART	302 (68.6%)	67 (15.2%)	71 (16.1%)	8 (2 to 15)	2 (-2 to 5)	7 (2 to 11)
Positive not on ART	19 (55.9%)	10 (29.4%)	5 (14.7%)	22 (4 to 41)	11 (-4 to 26)	4 (-5 to 14)
Previous Treatment						
Never Treated	312 (75.5%)	63 (15.3%)	38 (9.2%)	0 (reference)	0 (reference)	0 (reference)
Treated with First Line Drugs	188 (65.7%)	45 (15.7%)	53 (18.5%)	8 (1 to 15)	0 (-3 to 3)	5 (-0.3 to 9)
Isoniazid Resistance			· · · · · ·		· · · · ·	
Susceptible	177 (70.5%)	34 (13.5%)	40 (15.9%)	0 (reference)	0 (reference)	0 (reference)
Resistant	217 (76.4%)	37 (13%)	30 (10.6%)	-2 (-8 to 5)	-1 (-5 to 2)	-1 (-3 to 2)
Unknown	106 (64.6%)	37 (22.6%)	21 (12.8%)	7 (-1 to 15)	6 (-0.4 to 12)	0 (-2 to 3)

Abbreviations: AFB, acid fast bacilli; aRD, adjusted risk difference; 95% CI, 95% confidence interval; ART, antiretroviral treatment; IQR, interquartile range

^{*}Notes: Comparisons were done only on patients who were alive during the analytical period (e.g., analyses of death >3 months vs. survival were only done on patients who were alive after 3 months of treatment). Percentages represent row percentages. Model is adjusted for all factors listed here, including province of treatment.

Appendix Table 9: Risk of early or late mortality as estimated by multivariable analysis among patients in the bedaquiline group

	المعدنات منا	Early Death: died ≤3 months after treatment start	Late Death: died >3 to 24 months after treatment start	aDD man 400 (050/ CI)	-DD 400 (050/ CI)	-DD 400 (05% CI)
Characteristic	Survived (n=526)	(n=63)	(n=99)	aRD per 100 (95%CI) Death Ever vs. Survival	aRD per 100 (95%CI) Early Death vs. Survival	aRD per 100 (95%CI) Late Death vs. Survival
Age, median (IQR), per 10y	41 (33 to 51)	47 (36 to 59)	41 (33 to 54)	4 (2 to 6)	2 (1 to 3)	1 (-1 to 3)
Sex	(======	(**************************************	(**************************************	()	(:: :/	(22 2)
Female	203 (76.6%)	26 (9.8%)	36 (13.6%)	0 (reference)	0 (reference)	0 (reference)
Male	323 (76.4%)	37 (8.7%)	63 (14.9%)	0 (-5 to 4)	-1 (-2 to -0.1)	1 (-4 to 6)
AFB Smear at Baseline	,	,		, ,	, ,	` ,
Negative	244 (79.5%)	26 (8.5%)	37 (12.1%)	0 (reference)	0 (reference)	0 (reference)
Positive	224 (75.4%)	25 (8.4%)	48 (16.2%)	3 (-3 to 8)	1 (-0.3 to 2)	3 (-3 to 9)
Unknown	58 (69%)	12 (14.3%)	14 (16.7%)	7 (-6 to 19)	2 (-0.1 to 4)	6 (-4 to 16)
Culture at Baseline						
Negative	108 (84.4%)	6 (4.7%)	14 (10.9%)	0 (reference)	0 (reference)	0 (reference)
Positive	263 (76.9%)	23 (6.7%)	56 (16.4%)	8 (2 to 14)	-2 (-3 to -1)	6 (-1 to 12)
Unknown	155 (71.1%)	34 (15.6%)	29 (13.3%)	10 (1 to 18)	4 (0.2 to 7)	1 (-6 to 8)
HIV-Status						
Negative	147 (75.4%)	20 (10.3%)	28 (14.4%)	0 (reference)	0 (reference)	0 (reference)
Positive on ART	372 (77.8%)	37 (7.7%)	69 (14.4%)	1 (-4 to 6)	0 (-1 to 1)	0 (-6 to 5)
Positive not on ART	7 (46.7%)	6 (40%)	2 (13.3%)	25 (-7 to 57)	6 (-23 to 35)	2 (-29 to 34)
Previous Treatment						
Never Treated	328 (79.2%)	36 (8.7%)	50 (12.1%)	0 (reference)	0 (reference)	0 (reference)
Treated with First Line Drugs	198 (72.3%)	27 (9.9%)	49 (17.9%)	5 (0.4 to 10)	-1 (-1 to 0.1)	6 (1 to 12)
Isoniazid Resistance						
Susceptible	200 (74.9%)	28 (10.5%)	39 (14.6%)	0 (reference)	0 (reference)	0 (reference)
Resistant	213 (79.2%)	18 (6.7%)	38 (14.1%)	-4 (-9 to 2)	0 (-1 to 1)	-2 (-8 to 3)
Unknown	113 (74.3%)	17 (11.2%)	22 (14.5%)	2 (-7 to 11)	-1 (-3 to 0.2)	3 (-4 to 11)

Abbreviations: AFB, acid fast bacilli; aRD, adjusted risk difference; 95% CI, 95% confidence interval; ART, antiretroviral treatment; IQR, interquartile range

^{*}Notes: Comparisons were done only on patients who were alive during the analytical period (e.g., analyses of death >3 months vs. survival were only done on patients who were alive after 3 months of treatment). Percentages represent row percentages. Model is adjusted for all factors listed here, including province of treatment.