Prevalence and Associated Risk Factors of Multidrug Resistant Tuberculosis in Malaysia

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Abstract

Objective: Multidrug resistant tuberculosis is currently a significant public health concern. This research aims to determine the prevalence of MDR-TB in patients with tuberculosis infection.

Materials and methods: Data for this retrospective study were collected from the national TB Information System (TBIS) between 2009 and 2019. A total of 989 cases of MDR-TB have been reported and associated risk characteristics have been determined, such as marital status, gender, ethnicity, employment status, consumption of alcohol, diabetic status and smoking status. A multivariate, descriptive and Pearson's chi-square statistical analysis was conducted using SPSS version 20 software.

Results: The incidence of MDR-TB among patients with TB infections in Malaysia was 0.34%. The findings showed major differences in MDR-TB incidence between male and female patients (0.44% vs. 0.20%, p<0.001), single and married patients (1.63% vs. 0.24%, p<0.001), race (p<0.001), working and nonworking patients (0.48% vs. 0.32%, p<0.001), alcoholic and non-alcoholic patients (0.44% vs. 0.32%, p<0.001), diabetic patients and non-diabetic patients (0.39% vs. 0.27%, p<0.001), followed by smoking and non-smoking patients (0.13% vs. 0.27%, p<0.001).

Conclusion: This study provides a significant assessment of the prevalence of MDR-TB and related risk factors that could be useful in Malaysia's national TB strategy.

Keywords: Multidrug resistant tuberculosis • Prevalence data • Quality of life • Risk factors • TB surveillance and urban health

Introduction

In 2019 about 600,000 MDR-TB cases were reported worldwide with an estimated 250 million annual deaths. Global TB report has stated that incidents of MDR-TB have continued to rise, potentially threatening to undermine the advances made in TB control. A variation of socio economic and health risk conditions account for the high incidences of MDR-TB worldwide. Multidrug resistant TB is basically an infectious disease caused by inadequate drug sensitive treatment of tuberculosis.

in 2018, almost 480,000 new MDR-TB incidents with 100,000 Rifampicin Resistant (RR) incidents were reported worldwide. An approximate 500,000 new Isoniazid Resistant (IR) cases have been reported in 2018, of which 78% were MDR-TB cases. Despite this increase, however, the majority of patients diagnosed in 2017 and 2018 were only one in three or 32% of about 500,000 patients who have developed MDR-TB [1].

MDR-TB is known for its higher medication costs, a longer period of time for therapy and poor effectiveness as compared to other medical conditions. The MDR-TB epidemic is exacerbated by inadequate health education given to TB patients, an increasing number of respiratory infections and other health factors related to this disease. The frequency of MDR-TB cases is projected to increase in 2021 due to the lack of knowledge among TB patients and poor awareness of its contributing factors. In order to develop a comprehensive strategy for health care interventions, a broader understanding of the potential risk factors linked to MDR-TB infection is extremely essential [2].

However, most of MDR-TB related studies investigated were mainly in United States, Europe and Asia region. Hence, it is interesting and important to study on the prevalence of MDR-TB in Malaysia since the numbers of studies done in Malaysia are limited. The purpose of this study is to determine the prevalence of MDR-TB and highlight the relevant health risk factors associated to the development of MDR-TB infection in Malaysia as a tropical country [3].

Materials and Methods

The occurrence of MDR-TB infection in patients with TB was identified between years 2009 to year 2019. All registered TB patients who have been infected with MDR-TB are retrieved from national TB Information System (TBIS), Malaysia. Socio demographic characteristics such age, gender, ethnicity, marital status and employment, followed by clinical characteristics were recorded into a Microsoft excel data sheet [4].

Malaysian TB information system is a TB patient tracking system in the country that was introduced in 2003. MDR-TB data were collected and reported regularly by these TB clinics or treatment centers throughout the country using a uniform method. Throughout this process, all MDR-TB cases were reported to health department office which has served as a data center for the collection, surveillance, monitoring and recording of all MDR-TB cases in all the states. The data collected will be used to monitor cases involving MDR-TB, not only in treatment facilities but also at a national level. The mechanism of collecting data has helped in evaluating data at a national and global level. Furthermore, the collected data is contributed to the annual reports of the World Health Organization (WHO) [5].

Initial information on TB treatment (TBIS 10A1) forms were used to collect data from individuals with MDR-TB in this study. TBIS 10A1 forms were also used with the intention of enrolling individuals with TB as well as a means to input information into the national TBIS database. Accurate and detailed information regarding the individual with MDR-TB, such as socio demographic, clinical data and initiation of TB therapy were included in the forms. Data obtained from TBIS is used to approximate the prevalence of MDR-TB as well as to identify the possible risk factors among TB patients. The data collected from TBIS were cleaned and converted into an excel spreadsheet. The Statistical Software (SPSS version 20) was used to analyze the data. A 95% confidence interval was used to describe the frequency of MDR-TB. Prevalence is a term used to describe the overall number of incidents in which the disease is transmitted by the population at risk. Cases that were not MDR-TB were selected randomly as

control measures and to make comparisons. In order to calculate the prevalence, the following formula or equation (1) was used [6].

$$Prevalance = \frac{\text{Count of MDR_TB patients}}{(\text{Count of MDR_TB patients} + \text{Count of Non_MDR_TB patients})} \qquad \qquad \text{Equation. (1)}$$

Seven risk factors for MDR-TB infected patients in Malaysia were evaluated: Gender, marital status, ethnicity, employment status, alcohol consumption, diabetic status and smoking status. For each risk factor, Pearson's *chi-square* analysis was performed. The analysis involved the following steps:

- Extraction of observed values.
- · Formulation of hypothesis.
- Calculation of expected values.
- · Calculation of chi-square test significance.

The observed values and expected values were calculated based on equation (2).

To calculate the significance level, p-value, of the Pearson *chi-square* test, CHISQ.TEST was used [7].

= CHISQ.TEST (Observed-values / Expected-values $\chi_c^2 = \sum \left(\frac{o_i - E_i}{E_i}\right)^2$ Equation. (2)

Results

A total of 989 MDR-TB cases were reported to Malaysian national TB surveillance from 2009 to 2019. The MDR-TB cases that were reported during the timeframe included the seven independent risk factors: Gender, marital status, ethnicity, employment status, alcohol consumption, diabetic status and smoking status as shown in Table 1. Meanwhile the total number of TB cases from 2009 to 2019 was 287,092. Similarly, data from TB cases from 2009 to 2019 also included the seven independent risk factors: Gender, marital status, ethnicity, employment status, alcohol consumption, diabetic status and smoking status as shown in Table 2 [8].

Table 1. MDR-TB cases from year 2009 to 2019.

Risk factors/ Years	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019
MDR-TB	55	64	141	74	124	104	101	74	116	64	72
Male	37 (67%)	45	92	57	92	84	82	61	77	51	54
		(70%)	(65%)	(77%)	(74%)	(81%)	(81%)	(82%)	(66%)	(80%)	(75%)
Female	18	19	49	17	32	20	19	13	33	13	8
	(32.7%)	(30%)	(35%)	(23%)	(26%)	(19%)	(19%)	(18%)	(28%)	(20%)	(11%)
Single	35	26	43	24	46	42	13	25	48	29	26
	(63.6%)	(41%)	(30%)	(32%)	(37%)	(40%)	(13%)	(34%)	(41%)	(45%)	(36%)
Married	20	38	98	50	78	62	88	49	62	35	46
	(36%)	(59%)	(70%)	(68%)	(63%)	(60%)	(87%)	(66%)	(53%)	(55%)	(64%)
Malay	35	45	94	51	88	71	70	52	86	38	55
	(63.6%)	(70%)	(67%)	(69%)	(70%)	(68%)	(69%)	(70%)	(74%)	(59%)	(76%)
Chinese	5	5	7	6	6	5	4	4	5	4	4
	(90.1%)	(8%)	(5%)	(8%)	(5%)	(5%)	(4%)	(5%)	(4%)	(6%)	(6%)
Indian	11	10	35	13	28	25	24	17	22	21	12
	(20%)	(16%)	(25%)	(18%)	(23%)	(24%)	(24%)	(23%)	(19%)	(33%)	(17%)
Others	4	4	5	4	2	3	3	2	3	1	1
	(7%)	(6%)	(4%)	(5%)	(2%)	(3%)	(3%)	(3%)	(3%)	(1.5%)	(1.4%)
Working	10	12	49	26	35	12	24	16	29	15	11
	(18%)	(19%)	(35%)	(35%)	(28%)	(12%)	(24%)	(22%)	(25%)	(23%)	(15%)
Not	45	52	92	48	89	92	77	58	87	49	61
working	(81%)	(81%)	(65%)	(65%)	(72%)	(88%)	(76%)	(78%)	(75%)	(77%)	(85%)
Alcoholic	8	16	37	7	22	29	30	10	30	21	9
	(14.5%)	(25%)	(26%)	(9%)	(18%)	(28%)	(30%)	(14%)	(26%)	(33%)	(13%)
Non-	47	48	104	67	102	75	81	64	78	43	63
alcoholic	(85%)	(75%)	(74%)	(91%)	(82%)	(72%)	(80%)	(86%)	(67%)	(67%)	(88%)
Diabetic	42	55	72	65	78	52	70	48	77	53	60
	(76%)	(86%)	(51%)	(88%)	(63%)	(50%)	(69%)	(65%)	(66%)	(83%)	(83%)

Non-	13	9	69	9	46	52	31	26	39	11	12
diabetic	(24%)	(14%)	(49%)	(12%)	(37%)	(50%)	(31%)	(35%)	(34%)	(17%	(17%)
Non- smoker	4 (7%)	2 (3%)	7 (5%)	4 (5%)	5 (4%)	6 (6%)	5 (5%)	3 (41%	2 (2%)	3 (5%)	4 (6%)
Smoker	51	62	134	70	119	98	96	71	114	61	68
	(92.7%)	(97%)	(95%)	(95%)	(96%)	(94%)	(95%)	(96%)	(98%)	(95%)	(94%)

Note:***Total of (MDR-TB) cases from year 2009 to year 2019:989

Table 2. TB cases from year 2009 to 2019.

Risk factors/ Years	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019
ТВ	25241	27450	27500	29021	29221	25031	24220	25240	26168	24300	23700
Male	15210	17720	16245	17510	16731	14500	13100	14620	14084	14150	12850
	(60.29%)	(64.5%)	(59%)	(60.3%)	(57.2%)	(58%)	(54%)	(58%)	(54%)	(58.2%)	(54%)
Female	10031	9730	11255	11511	12490	10531	11120	10620	12084	10150	10850
	(39.74%)	(35.4%)	(40.9%)	(39.6%)	(42.7%)	(42%)	(46%)	(42%)	(46%)	(41.7%)	(45.8%)
Single	2241	1675	2408	1825	2135	1772	1670	1665	1872	2438	1782
	(8.87%)	(6.1%)	(8.7%)	(6.2%)	(7.3%)	(7%)	(6.9%)	(6.6%)	(7.2%)	(10%)	(7.5%)
Married	23000	25775	25092	27196	27086	23259	22550	23575	24296	21862	21918
	(91.1%)	(93.8%)	(91.2%)	(93.7%)	(92.6%)	(94%)	(93%)	(93%)	(93%)	(90%)	(92.4%)
Malay	14855	17135	17137	17832	19601	15670	15437	16375	17618	15789	15556
	(58.8%)	(62.4%)	(62.3%)	(61.4%)	(67%)	(62.6%)	(64%)	(65%)	(67%)	(65%)	(65.6%)
Chinese	2996	3702	3501	3224	3102	3017	3275	3070	2972	3021	2752
	(11.8%)	(13.4%)	(12.7%)	(11.1%)	(10.6%)	(12%)	(14%)	(12%)	(11.3%)	(12.4%)	(11.6%)
Indian	5870	4872	5241	6423	5020	4772	4100	4370	4200	4172	4090
	(23.2%)	(17.7%)	(19.0%)	(22.1%)	(17.1%)	(19%)	(17%)	(17%)	(16%)	(17.2%)	(17.2%)
Others	1520	1741	1621	1542	1498	1572	1408	1425	1378	1318	1302
	(6.02%)	(6.3%)	(5.9%)	(5.3%)	(5.1%)	(6.2%)	(6%)	(6%)	(5.3%)	(5.4%)	(5.5%)
Vorking	4527	4383	4272	4307	4772	4326	3966	5271	4788	4624	4613
	(17.9%)	(15.9%)	(15.5%)	(14.8%)	(16.3%)	(17.2%)	(16.4%)	(21%)	(18.2%)	(19%)	(19.4%)
Not Nortsing	20714	23067	23228	24714	24449	20705	20254	19969	21380	19676	19087
Working	(82%)	(84%)	(84.4%)	(85.1%)	(83.6%)	(82%)	(84%)	(79%)	(81.7%)	(81%)	(80.5%)
Alcoholic	5201	3724	4672	4407	4328	4269	4432	4170	3983	4007	4002
	(20.6%)	(13.5%)	(16.9%)	(15.1%)	(14.8%)	(17%)	(18.2%)	(16%)	(15.2%)	(16.5%)	(16.9%)
Non-	20040	23726	22828	24614	24893	20762	19788	21070	22185	20293	19698
Alcoholic	(79.3%)	(86.4%)	(83%)	(84.8%)	(85%)	(83%)	(82%)	(83.4%)	(85%)	(83.5%)	(83%)
Diabetic	15201	17720	16358	15520	15366	15201	15179	15073	14788	14702	14585
	(60.2%)	(64.5%)	(59.4%)	(53.4%)	(52.5%)	(61%)	(63%)	(60%)	(56.5%)	(61%)	(61.5%)
Non-	10040	9730	11142	13501	13855	9830	9041	10167	11380	9598	9115
Diabetic	(39.7%)	(35.4%)	(40.5%)	(46.5%)	(47.4%)	(39%)	(37%)	(40%)	(43.4%)	(39.4%)	(38.4%)

Non- Smoker	3856 (15.2%)	2129 (7.7%)	2277 (8.2%)	2539 (8.7%)	2867 (9.8%)	2729 (11%) 2614 (11%	6) 3729 (15	5%) 1484 (5.7%)	4088 (17%)	5216 (22%)
Smoker	21385 (84.7%)	25321	25321 (92.2%)	26482 (91.2%)	26354 (90%)	22302 (89%)	21606 (89%)	21511 (85%)	24684 (94.3%)	20212 (83%)	18484 (78%)

Note: ***Total of TB cases from year 2009 to year 2019: 287,092

Based on national TB surveillance data for Malaysian TBIS, the overall estimate of the MDR-TB occurrence among TB infected individuals in Malaysia is determined based on the count, (MDR-TB patients/MDR-TB patient's count+non-MDR-TB patients count) as shown in Table 3. Overall prevalence of (MDRT-TB/Total TB) from 2009 to 2019 is shown in Figure 1. As for 2011, the cumulative incidence of MDR-TB over total TB was recorded as the highest compared to other years. In 2012 and 2018, MDR-TB incidence

dropped drastically. The frequency of MDR-TB among low, which patients is still relatively approximately 0.34% between 2009 and at such Independent risk factors, as socio demographic characteristics, are also linked to MDR-TB occurrence in Malaysia

Table 3. Overall incidence of MDR-TB cases from 2009 to 2019.

Risk factors/ Years	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	Total	P-value
MDR- TB/Total TB	0.22%	0.23%	0.51%	0.25%	0.42%	0.41%	0.42%	0.29%	0.44%	0.26%	0.30%	0.34%	<0.001
Male	0.24%	0.25%	0.56%	0.32%	0.55%	0.58%	0.62%	0.42%	0.54%	0.36%	0.42%	0.44%	<0.001
Female	0.18%	0.19%	0.43%	0.15%	0.26%	0.19%	0.17%	0.12%	0.27%	0.36%	0.07%	0.20%	_
Single	1.54%	1.53%	1.75%	1.30%	2.11%	2.32%	0.77%	1.48%	2.50%	1.18%	1.44%	1.63%	<0.001
Married	0.09%	0.15%	0.39%	0.18%	0.29%	0.27%	0.39%	0.21%	0.25%	0.16%	0.21%	0.24%	_
Malay	0.24%	0.26%	0.55%	0.29%	0.45%	0.45%	0.45%	0.32%	0.49%	0.24%	0.35%	0.37%	<0.001
Chinese	0.17%	0.13%	0.20%	0.19%	0.19%	0.17%	0.12%	0.13%	0.17%	0.13%	0.15%	0.16%	_
Indian	0.19%	0.20%	0.66%	0.20%	0.55%	0.52%	0.58%	0.39%	0.52%	0.50%	0.29%	0.41%	_
Others	0.26%	0.23%	0.31%	0.26%	0.13%	0.19%	0.21%	0.14%	0.22%	0.08%	0.08%	0.20%	_
Working	0.22%	0.27%	1.13%	0.60%	0.73%	0.28%	0.60%	0.30%	0.60%	0.32%	0.24%	0.48%	<0.001
Not working	0.22%	0.22%	0.39%	0.19%	0.36%	0.44%	0.38%	0.29%	0.41%	0.25%	0.32%	0.32%	_
Alcoholic	0.15%	0.43%	0.79%	0.16%	0.51%	0.67%	0.45%	0.24%	0.75%	0.52%	0.22%	0.44%	<0.001
Non- Alcoholic	0.23%	0.20%	0.45%	0.27%	0.41%	0.36%	0.41%	0.30%	0.35%	0.21%	0.32%	0.32%	_
Diabetic	0.28%	0.31%	0.44%	0.42%	0.51%	0.34%	0.46%	0.32%	0.52%	0.36%	0.41%	0.39%	<0.001
Non- Diabetic	0.13%	0.09%	0.62%	0.07%	0.33%	0.53%	0.34%	0.26%	0.34%	0.11%	0.13%	0.27%	_
Non- Smoker	0.10%	0.09%	0.31%	0.16%	0.17%	0.22%	0.19%	0.08%	0.13%	0.07%	0.08%	0.13%	<0.001
Smoker	0.24%	0.24%	0.53%	0.26%	0.45%	0.44%	0.44%	0.33%	0.46%	0.30%	0.37%	0.37%	_

Note: ***Incidence of (MDR-TB) from year 2009 to 2019: 0.34%

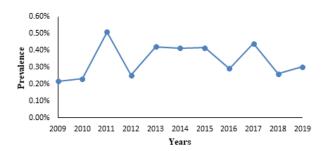


Figure 1. Overall prevalence of MDR-TB/total TB.

Table 4 shows the association of independent factors which have contributed to the incidence of MDR-TB in Malaysia. The

frequencies of MDR-TB cases among male patients are 732 (74%) and for female patients are 241 (24.3%). For TB patients the frequencies for male patients are 166,720 (58%) and followed by female patients are 120,372 (42%). The proportion of males in the MDR-TB group is equal to the proportion of females in the MDR-TB group. The frequencies of MDR-TB cases among single patients are 357 (36%) and for married patients are 626 (63%). The crude odds ratio for married patients is (OR: 7.051 p<0.001) and for single patients is (OR: 0.142 p<0.001). For TB patients the frequencies for single patients are 21,483 (7.5%) and followed by married patients are 256,609 (92.5%). The proportion of singles in the MDR-TB group is equal to the proportion of marrieds in the MDR-TB group. Pearson's chisquare indicated that there is a significance difference in the prevalence of MDR-TB between singles and married patients (1.63% vs. 0.24%, p<0.001) [10].

Table 4. Independent risk factors related to MDR-TB prevalence

Independent risk factors	Number (Prevalence)	P-value
Gender		
Male	166,720 (0.44%)	<0.001
Female	120,372 (0.20%)	
Marital status		
Single	21,483 (1.63%)	<0.001
Married	256,609 (0.24%)	
Ethnic groups		
Malay	183,008 (0.37%)	
Chinese	34,632 (0.16%)	
Indian	53,130 (0.41%)	
Other	16,325 (0.20%)	
Working status		
Working	49,849 (0.48%)	<0.001
Not working	237,243 (0.32%)	
Alcohol consumption		
Alcoholic	47,195 (0.44%)	<0.001
Non-alcoholic	239,897 (0.32%)	
Diabetic status		
Diabetic	169,693 (0.39%)	<0.001
Non-diabetic	117,399 (0.27%)	_
Smoking status		
Smoking	253,564 (0.13%)	<0.001
Non-smoking	33,528 (0.27%)	_

The frequencies of MDR-TB cases among Malay patients are 685 (69.2%), Chinese patients are 55 (5.6%), Indian patients are 218 (22%) and others are 32 (3.2%). For TB patients the frequencies for Malay patients are 183,005 (63.7%), Chinese patients are 34,632 (12%), Indian patients are 53,130 (19%) and others are 16,325 (5.7%). The proportion of all ethnic's groups in the MDR-TB group is equal. Pearson's chi-square indicated that there is a significance difference in the prevalence of MDR-TB between ethnic's groups (p<0.001). The frequencies of MDR-TB cases among working patients are 239 (24%) and not working are 750 (75.8%) patients. For TB patients the frequencies for working patients are 49,849 (17.4%) and not working patients are 237,243 (82.6%). The proportion of working patients in the MDR-TB group is equal to the proportion of not working patients in the MDR-TB group. Pearson's chi-square indicated that there is a significance difference in the prevalence of MDR-TB between working patients and not working patients (0.48% vs. 0.32%, p<0.001). The frequencies of MDR-TB cases among alcoholic patients are 209 (21%) and non-alcoholic are 772 (78%) patients. For TB patients the frequencies for alcoholic patients are 47,195 (16.4%) and non-alcoholic patients are 239,897 (83.5%). The proportion of alcoholic patients in the MDR-TB group is equal to the proportion of non-alcoholic patients in the MDR-TB group. Pearson's chi-square indicated that there is a significance difference in the prevalence of MDR-TB between alcoholic patients and non-alcoholic patients (0.44% vs. 0.32%, p<0.001) [11].

The frequencies of MDR-TB cases among diabetic patients are 672 (0.67%) and non-diabetic patients are 317 (32%). For TB patients the frequencies for diabetic patients are 169,693 (59.1%) and non-diabetic patients are 117,399 (40.9%). The proportion of diabetic patients in the MDR-TB group is equal to the proportion of non-diabetic patients in the MDR-TB group. Pearson's *chi-square* indicated that there is a significance difference in the prevalence of MDR-TB between diabetic patients and non-diabetic patients (0.39% vs. 0.27%, p<0.001).

The frequencies of MDR-TB cases among smoking patients are 944 (95.4%) and non-smoking patients are 45 (4.6%). For TB patients the frequencies for smoking patients are 253,564 (88.3%) and non-smoking patients are 33,528 (11.7%). The proportion of smoking patients in the MDR-TB group is equal to the proportion of non-smoking patients in the MDR-TB group. Pearson's *chi-square* indicated that there is a significance difference in the prevalence of MDR-TB between smoking patients and non-smoking patients (0.13% vs. 0.27%, p<0.001).

The association of risk factors with MDR-TB and non MDR-TB patients are shown in (Table 5). The proportion of MDR-TB has been found to be higher in male patients 75.23% (n=732) compared to female patients 24.77% (n=241) with (pvalue<0.001). Marital status distribution shows statistically significant between married and single patients whereby the proportion of married patients are higher 63.68% (n=626) than single patients 36.32 (n=357). The crude OR value shows single patients are higher (7.051 at 95% CI) compared to married patients. The association among different ethnicity groups also to be reported as statically significant (p-value<0.001). The proportion of Malay are higher 22.02% (n=218), (n=685) compared to Indian 69.19% followed by Chinese 5.56% (n=55) and others 3.23% (n=32). MDR-TB Those non-working patients shows proportion 75.83% (n=750) than working MDR-TB patients 24.17% Similarly, the proportion of non MDR-TB has also found to be higher in male patients 58.07% (n=166,720) compared to female patients 41.93% (n=120,372). Meanwhile the proportion of married non MDR-TB patients 95.52% (n=265,609) is much higher than single non MDR-TB patients 7.48% (n=21,483). Apart from that, the proportion among different ethnicity groups found that Malay patients are to be higher 63.74% (n=183,005) compared to Indian patients 18.51% (n=53,130), followed Chinese patients 12.06% (n=34,632) and others 5.69% (n=16,325).

Table 5. Association (adjusted OR) of MDR-TB risk factors.

Risk factor	MDR-TB		Non MDR-	ГВ	P-value	Adjusted (OR (95% CI)	
	Number (%) (n=)		Number	· (%) (n=)				
	%	n	%	n		OR	95	% CI
Gender								
Male	75.23	732	58.07	166720	0.001	2.193	1.896	2.537
Female	24.77	241	41.93	120372		Reference	0.394	0.528
Marital status								
Single	36.32	357	7.48	21483	0.001	7.051	6.187	8.036
Married	63.68	626	92.52	265609		Reference	0.124	0.162
Ethnicity								
Malay	69.19	685	63.74	183005	0.001	1.277	1.116	1.462
Chinese	5.56	55	12.06	34632		0.429	0.327	0.563
Indian	22.02	218	18.51	53130		1.243	1.07	1.446
Others	3.23	32	5.69	16325		Reference	0.389	0.788
Employment s	tatus							
Working	24.17	239	17.36	49849	0.001	1.517	1.311	1.755
Non-working	75.83	750	82.64	237243		Reference	0.570	0.763
Alcoholic								
Yes	21.3	209	16.44	47195	0.001	1.376	1.181	1.604
No	78.7	772	83.56	239897		Reference	0.623	0.847
Diabetic								
Yes	67.95	672	59.11	169693	0.001	1.467	1.283	1.676
No	32.05	317	40.89	117399		Reference	0.596	0.779
Smoking								
Yes	95.45	944	50	253564	0.001	2.774	2.056	3.742
No	4.55	45	50	253564		Reference	0.267	0.486

On multivariate analysis, the regression model showed that male patients (OR 2.193, 95% CI=1.896-2.537) and the married patients showed (OR 7.051, 95% CI=6.187-8.036). Following on the ethnicity groups the Malay patients showed higher (OR 1.277, 95% CI=1.116-1.462), Indian (OR 1.243, 95% CI=1.243), Chinese (OR 0.429 95% CI=0.327-0.563) and others (OR 0.554 95% CI=0.389-0.788). Meanwhile working patients (OR 1.517, 95% CI=1.311-1.755) are more likely associated with MDR-TB infections compare to non-working patients (OR 0.659 95% CI=0.570-0.763). On binary logistic regression analysis, the crude odds ratio (OR) showed that alcoholic (OR 1.376, 95%Confidence Interval (CI) 1.181-1.604). For diabetic patients (OR 1.467, 95% CI=1.283-1.676) and smoking patients (OR 2.774, 95%CI=2.056-3.742) were significantly associated with MDR-TB. All risk factors such as gender, marital status, ethnicity groups, employment status, alcoholic, diabetic and smoking has proven that statistically significant associated with MDR-TB infections with (p<0.001).

Discussion

In Malaysia, the MDR-TB frequency is low and can be attributed to patients' positive response to first line antibiotics such as isoniazid and rifampicin. According TB patients with isoniazid resistant can be cured at a risk of relapse of less than 5%. Men are at greater risk of being infected with MDR-TB compared to women identified in this study. One potential explanation, according to is that men primarily smoke, drink alcohol and use drugs compared to According to younger and single individuals tend to be more vulnerable to MDR-TB due to their high participation in includes alcohol social behaviors which intake smoking compared to older and married individuals. Evidence from this study indicates that MDR-TB infections between different ethnicities may rely on their behavioral activities. Malay and Indian ethnic groups have a higher incidence of MDR-TB relative to other ethnic groups in Malaysia and

this may be attributed to social behavioral factors such as smoking habits, use and abuse of substance and alcohol consumption. Jobless TB infected individuals are more vulnerable to transmission of MDR-TB than those working due to their poverty line, which prevents them from follow up with routine health checks with clinicians in TB centers and purchasing of medicines. A study has revealed that there is a likelihood of a 3.54 fold increase in MDR-TB transmission between alcoholics compared to non-alcoholics, indicating the possibility of disease recurrence. The substantial proportion of MDR-TB multidrug resistant tuberculosis among diabetic patients appears to be growing steadily compared to non-diabetic patients. Smoking is associated with the cessation of TB/MDR-TB treatment and it has been found that this association is independent of alcohol or illicit drug usage.

Conclusion

To date, MDR-TB cases in Malaysia is relatively low (<0.34%) compared to the cases reported globally. In Malaysia, some MDR-TB patients go through heavy financial burdens for their diagnosis and treatments. Multi drug resistance to TB is developed either due to infection with a resistant strain, or as a result of inadequate treatment and poor patient compliance. Hence, proper patient care and commitment to the completion of anti-TB therapy are recommended. The results of the study revealed that all contributing risk factors including gender, ethnicity, marital status, smoking, job, diabetic and intake are associated significantly with MDR-TB alcohol prevalence and it is relatively low (<2.0%) in Malavsia. MDR-TB prevalence estimates will theoretically Reliable guide successful improve help to prevention measures, up interventions in vulnerable intervention and follow populations and assist in overall clinical decision making. Inadequate treatment for Tuberculosis (TB) patients and transmission are two main ways for MDR-TB infection and MDR-TB prevalence was higher in persons with previously treated TB patients than that is persons without, proven in many studies. In terms of improving public health related to the condition, development of a national guideline for surveillance and monitoring of MDR-TB and engagement with non-profit organization, civil society and communities for treatment support should be prioritized and strengthened.

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Conflict of Interest

The authors declare no conflict of interest, financial or otherwise.

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Ethical Approval

National institutes of health and medical research ethics committee. NMRR-19-1460-46943

References

 Ahuja, S.D., et al. "Multidrug resistant pulmonary tuberculosis treatment regimens and patient outcomes: An individual patient data metaanalysis of 9,153 patients." PLoS Med. 9 (2012):1001300.

- Caminero, J.A. "Multidrug resistant tuberculosis: Epidemiology, Epidemiology, risk factors and case finding." J Int Union Against Tuberculosis Lung Disease. 14 (2010):382-390.
- Garcia, F., et al. "Prevalence of diabetes mellitus and related risk factors in an urban population." Revista de la Soc Peruana de Med Int. 20 (2007):90-94
- Cherkaoui, Í., et al. "Treatment default amongst patients with tuberculosis in urban Morocco predicting and explaining default and post default sputum smear and drug susceptibility results." PLoS One. 9 (2014):93574-93574.
- Chien, J.Y., & Wang, J.Y. "Isoniazid resistant tuberculosis treatment with first line drugs." *Lancet Infect Dis*. 17 (2017):259-260.
- Elmi, O.S., et al. "Multidrug resistant tuberculosis and risk factors associated with its development: A retrospective study." J Infect Dev Ctries. 9 (2015):1076-1085.
- Faustini, A., et al. "Risk factors for multidrug resistant tuberculosis in Europe: A systematic review." Thorax. 61 (2006): 158-163.
- Imtiaz, S., et al. "Alcohol consumption as a risk factor for tuberculosis: Meta-analyses and burden of disease." European Resp J. 50 (2017):1700216.
- Rajendran, M., et al. "Contributing risk factors towards the prevalence of multidrug resistant tuberculosis in Malaysia: A systematic review" *Tuberculosis* (*Edinb*). 122 (2020):101925.
- Sharma, S.K., et al. "Prevalence of multidrug resistant tuberculosis among category II pulmonary tuberculosis patients." *Indian J Med Res.* 133(2011):312-315.
- Suarez-Garcia, I., et al. "Risk factors for multidrug resistant tuberculosis in a tuberculosis unit in Madrid, Spain." Eur J Clin Microbiol Infect Dis. 28 (2009):325-330.