

Effect of smoking and alcohol consumption on pulmonary tuberculosis among Batak ethnic population in Medan, Indonesia

B Y M Sinaga^{1*}, Y Siregar², M Amin³, and S Sarumpaet⁴

¹Department of Pulmonology and Respiratory Medicine, Adam Malik General Hospital, Faculty of Medicine, Universitas Sumatera Utara, Medan, Indonesia

²Department of Biochemistry, Faculty of Medicine, Universitas Sumatera Utara, Medan, Indonesia

³Department of Pulmonary and Respiratory Medicine, Faculty of Medicine, Universitas Airlangga, Surabaya, Indonesia

⁴Department of Epidemiology, Faculty of Public Health, Universitas Sumatera Utara, Medan, Indonesia

*Corresponding author: bintang@usu.ac.id

Abstract. Simultaneous consumption of *tuak*, a traditional alcoholic beverage, and smoking is prevalent among Batak ethnic group in Indonesia. This research was to find out the association between smoking and alcohol consumption with the risk of Pulmonary Tuberculosis (PTB) in Batak ethnic group in Medan, Indonesia. A matched case-control study was conducted on 100 PTB patients and 100 healthy individuals group. Smoking and alcohol consumption was self-reported. Data were analyzed with Epi Info program. Smoking and alcohol consumption habit is a significant difference in case and control group ($p < 0.01$). After conditional logistic regression analysis with non-smoking and non-alcohol consuming as a comparative, the Odds Ratio (OR) for the smoking-only group was 4.08 (95% CI: 1.28-13.05). For the alcohol-only consuming group was 1.83 (95% CI: 0.11-28.95) and for the smoking and alcohol consuming group was 13.7 (95% CI: 4.02-46.94). There is an association between smoking and alcohol consumption and the risk of PTB in Batak ethnic group in Medan, Indonesia.

1. Introduction

Previous studies showed an association between smoking, alcohol and the increased risk of PTB^[1-4]. Smoking may lower the immune system by affecting the function of epithelial permeability, cilia movement and macrophage^[5]. Alcohol consumption weakens the immune system via direct or indirect toxic effect of alcohol through nutrient deficiency and the other alcohol-related medical condition such as cancer and depression^[3].

It is common for Batak ethnic group to consume *tuak*, a traditional beverage containing alcohol. *Tuak* is usually enjoyed in a *lapo*, a traditional coffee shop while smoking cigarettes^[6]. Based on this condition, this study was conducted to find out the association between smoking and consuming alcohol and the risk of PTB in the Batak ethnic group in Medan, Indonesia.



2. Method

2.1. Cases and controls

It is a matched case-control study. Cases were PTB patients who were recruited from several TB centers in Medan, Indonesia, from November 2012 to November 2013. The inclusion criteria in the case group were the newly diagnosed PTB patients, age 16-55 years old, Batak ethnic, have symptoms of PTB, positive sputum smear and chest radiography consistent with the active disease. The exclusion criteria in the case group were HIV positive, have diabetes mellitus and other severe diseases, and taking the immunosuppressive drug.

The sample size used a matched case and control formula. The sample size was 100 for case group and 100 for the control group. Cases were found by continuous sampling and control group matched in sex, age and ethnicity were healthy subjects with ordinary chest X-ray and no prior history of PTB.

All of the subjects were interviewed, and informed consents were obtained. Smoking and alcohol consumption were self-reported. The habits of matter were into smoking-only, alcohol-only consuming, smoking & alcohol drinking and non-smoking & non-alcohol drinking. This research has been approved by the Ethics Committee of the Faculty of Medicine, Universitas Sumatera Utara, Medan, Indonesia.

2.2. Statistical analysis

The association between smoking and alcohol consumption and PTB was through Odds Ratio (OR) calculation with 95% confidence interval (95% CI) with p-value <0.05 to be statistically significant. The conditional logistic regression analysis was conducted by separating the study population into smoking-only, alcohol-only consuming, smoking and alcohol-consuming, with non-smoking and non-consuming alcohol group was used as a control. The data obtained were processed and analyzed by using Epi Info program.

3. Results

Table 1. Characteristic of the subject in case and control group.

Characteristics		TB (Case)		Non-TB (Control)	
		n	%	N	%
Sex	Male	70	70	70	70
	Female	30	30	30	30
Age (Years)	16-25	32	32	32	32
	26-35	39	39	39	39
	36-45	20	20	20	20
	46-55	9	9	9	9
Ethnic	Batak	100	100	100	100

The characteristics of the subjects comprising age, sex, and ethnic group in the case and control groups were in table 1. Gender, age and racial characteristic between case and control group, were matched.

Table 2. Smoking and alcohol consumption habits in case and control group.

Habits	TB (case)		Non-TB (control)		x ²	p-value
	n	%	N	%		
Non-smoking & non-alcohol consuming	37	37	61	61	21.85	< 0.01
Smoking-only	19	19	22	22		
Alcohol-only consuming	1	1	3	3		
Smoking & alcohol consuming	43	43	14	14		
Total	100	100	100	100		

MxN table epi info

There was a significant difference in smoking and alcohol consumption habits in case and control group ($p < 0.01$).

Table 3. It is an association between smoking and alcohol consumption with PTB.

Habits	TB (Case)		Non-TB (Control)		OR	95% CI	p-value
	N	%	n	%			
Non-smoking & non-alcohol consuming	37	37	61	61	1	1	1
Smoking-only	19	19	22	22	4.08	1.28-13.05	0.01
Alcohol-only consuming	1	1	3	3	1.83	0.11-28.95	0.66
Smoking and alcohol consuming	43	43	14	14	13.7	4.02-46.94	< 0.01

Conditional Regression Logistic

The conditional logistic regression test was conducted by separating the study population into four different group such as smoking-only, alcohol-only consuming, smoking and alcohol consuming, with the non-smoking and non-alcohol consuming group as the control. The Odds Ratio (OR) for the smoking-only group was 4.08 (95% CI: 1.28-13.05); for the alcohol-only consuming group was 1.83 (95% CI: 0.11-28.95), and for the smoking and alcohol consuming group was 13.7 (95% CI: 4.02-46.94) (Table 3).

4. Discussion

This study found a difference in smoking and alcohol consumption habits between pulmonary tuberculosis patients compared to healthy subjects in the Batak population. Similar to previous studies, this study also showed an association between smoking and alcohol consumption and the risk of PTB. Review of 34 previous studies from various countries, it was found out that smoking increased the risk of infection caused by *Mycobacterium tuberculosis*, tuberculosis development, and tuberculosis-caused mortality^[1]. Meta-analysis study of 24 previous studies was in a separate analysis: TB infection (6 studies), TB development (13 studies), and tuberculosis-caused mortality (5 studies) - showed a relationship between smoking and TB infection and TB development after sex, age, and the other confounding factors were adjusted. The relationship between smoking and TB development was also found in the passive smokers^[7].

Regarding alcohol consumption, in a meta-analysis study, Lönnroth concluded that Relative Risk (RR) for heavy drinkers consuming more than 40 grams of alcohol per day was 2.94 (95% CI: 1.89-4.59)^[3]. Rehm reported similar results in a meta-analysis study and found that alcohol influences the incident of TB, TB treatment outcomes, increased risk of re-infection, and drug-resistant TB^[4]. A meta-analysis by Imtiaz found alcohol increased TB incident, death and TB risk rose as ethanol intake increased^[8].

The specific mechanism of cigarette smoke on respiratory infections and inflammation is further study. Smoking directly affects the integrity and permeability of the respiratory alveolar macrophages to produce the inflammatory mediators, reactive oxygen species and proteolytic enzymes^[9]. Cigarette smoke also minimizes alveolar macrophages phagocytic ability^[10-11]. Mice exposed to cigarette smoke during experiment has more *M. tuberculosis* in their lungs and spleens compared to the control group. An increased number of intracellular *M. tuberculosis* in human macrophages was in smokers compared to non-smokers^[12].

Alcohol can cause the direct toxic effect on the human immune system causing susceptibility to TB. Alcohol lowers the response of Nitric Oxide system against mycobacterial infection and inhibits phagocytosis^[13]. The result of the experiments carried out on mice indicates that alcohol may inhibit granuloma formation, produce IL-2, IFN-gamma, and proliferation of CD4^[14]. Migration Inhibition Factor (MIF) is a cytokine whose production is inhibited by the presence of alcohol. Alcohol also lowers the production of cytokines such as TNF- α , IL-1, and IL-6. Alcohol can also affect the activation of T cells causing Th2 population (humoral immunity) to dominate the Th1 population (cellular immunity responsible for TB infection). These changes disrupt the balance between the two

basic types of immune system, and therefore, lowering the immune defense and increasing the susceptibility to TB^[13].

This study was explicitly carried out on Batak ethnic group based on the consideration of their habit of drinking *tuak*, a traditional alcoholic beverage. Most subjects said that they drink *tuak* in the *lapo*, traditional coffee shop while smoking. *Tuak* is from *mayang enau* also known as *aren* (*Arengapinnata*). The *enau* or *aren* tree is tapped to collect its juice known as *nira*. After this *nira* is mixed with *raru*, a kind of wood to add more taste and alcohol content. *Raru* is the main ingredient in the fermentation process. Since *enau* or *aren* is not in Medan, *tuak* can also be made from the tapped coconut tree^[6]. The alcohol content found in *tuak* is quite high compared to other alcoholic beverages, especially the *tuak* made from the mixture of *nira* and *raru*. The previous study revealed that the content of ethanol in pure *tuak* ranges from 12.07 to 21.88%, and in *tuak* made from the mixture of *nira* and *raru* the alcohol content can be up to 47.04%^[15].

Study about smoking and alcohol consumption as a risk factor for tuberculosis based on ethnic has been done in India and the results were similar to this study. Among Tribal population in India, a person who was both smoker and alcohol-consuming increased the risk of pulmonary tuberculosis than smoking and alcohol consumption only^[16].

There was some limitation when conducting this study. The amount of the alcohol consumed, alcohol content and smoking severity index could not be determined. Though sex, age, and ethnicity of the case and control groups were matched, other confounding factors such as education level, nutrition, drug abuse, air pollution, occupation and socio-economic status remain unmatched. The limitation of the analysis in this study was related to the sample size especially the sample size for the alcohol-only consuming group.

Further study on the association between smoking, alcohol consumption and the risk of PTB using a large sample size is needed to be conducted with a careful assessment of the confounding factors to obtain a better understanding of the possible causal pathways between smoking and alcohol consumption and pulmonary tuberculosis.

5. Conclusion

Smoking and alcohol consumption are risk factors for pulmonary tuberculosis occurring in Batak ethnic group in Indonesia. This information is essential for TB control and the strategy to prevent the incidence of tuberculosis in Indonesia whose population comprising various ethnic backgrounds and cultures.

References

- [1] Hasmy K M 2006 The association between smoking and tuberculosis *Salud Publica Mex* **48(1)** S201-16
- [2] Bates M N, Khalakdina A, Pai M, Chang L, Lessa F and Smith K R 2007 Risk of tuberculosis from exposure to tobacco smoke. A systematic review and meta-analysis *Arch. Intern. Med.* **167(4)** 335-42
- [3] Lönnroth K, Williams B G, Stadlin S, Jaramillo E and Dye C 2008 Alcohol use as a risk factor for tuberculosis – a systematic review *BMC Pub. Health* **8** 289
- [4] Rehm J, Samokhalov AV, Neuman MG, Room R, Parry C and Lönnroth K 2009 The association between alcohol use, alcohol use disorders and tuberculosis (TB): A systematic review *BMC Pub. Health* **9** 450
- [5] Stämpfli M R and Anderson G P 2009 How cigarette smoke skews immune responses to promote infection, lung disease, and cancer *Nat. Rev. Immunol.* **9(5)** 378-84
- [6] IKEGAMI and Shigehiro 2014 *Tuak* in the Toba Batak society: A preliminary report on the socio-cultural aspect of palm wine consumption (Annual report of the University of Shizuoka, Hamamatsu College) **11-3(1997)** Part 5
- [7] Leung C C, Lam T H, Ho K S, Yew W W, Tam C M, Chan W M, Law W S, Chan C K, Chang K C and Au K F 2010 Passive smoking and tuberculosis *Arch. Intern. Med.* **170(3)** 287-92

- [8] Imtiaz S, Shield K D, Roerecke M, Samokhalov A V, Lönnroth K and Rehm J 2017 Alcohol consumption as risk factor for tuberculosis: meta-analyses and burden of disease *Eur. Respir. J.* **50(1)** 1700216
- [9] Dye J A and Adler K B 1994 Effects of cigarette smoke on epithelial cells of the respiratory tract *Thorax* **49(8)** 825-34
- [10] Berenson C S, Garlipp M A, Grove L J, Maloney J and Sethi S Impaired phagocytosis of nontypeable *Haemophilus influenzae* by human alveolar macrophages in chronic obstructive pulmonary disease *JID* **194(10)** 1375-84
- [11] Hodge S, Hodge G, Ahern J, Jersmann H, Holmes M and Reynolds P N 2007 Smoking alters alveolar macrophage recognition and phagocytic ability: implications in chronic obstructive pulmonary disease *Am. J. Respir. Cell. Mol. Biol.* **37(6)** 748-55
- [12] Shang S, Ordway D, Henao-Tamayo M, Bai X, Oberley-Deegan R and Shanley C 2011 Cigarette smoke increases susceptibility to tuberculosis-evidence from in vivo and in vitro models *JID* **203(9)** 1240-8
- [13] Gamble L, Mason C M and Nelson S 2006 The effect of alcohol on immunity and bacterial infection in the lung *Méd Maladies Infect.* **36(2)** 72-7
- [14] Mason C M, Dobard E, Zhang P and Nelson S 2004 Alcohol exacerbates murine pulmonary tuberculosis *Infect. Immun.* **72(5)** 2556-63
- [15] Sembiring S 2011 Pengaruh pemberian vitamin E terhadap perubahan bobot dan gambaran mikroskopis tubulus proksimal ginjal mencit (*Mus musculus*, L.) jantan dewasa yang dipapari *tuak* (alkohol) [Tesis] (Medan, Indonesia: Program Pasca Sarjana Ilmu Biomedik Fakultas Kedokteran Universitas Sumatera Utara)
- [16] Rao V G, Bhat J, Yadav R, Muniyandi M, Bhondeley M K and Wares D F 2017 Smoking and alcohol consumption: risk factors for pulmonary tuberculosis among the tribal community in Central India *Indian J. Tuberc.* **64(1)** 40-3