EARLY DETECTION OF CHANGES IN BLOOD GLUCOSE LEVELS AS AN EFFORT TO INCREASE SUCCESS FOR THE TREATMENT OF LUNG TUBERCULOSIS

Pariyana¹, Iche Andriyani Liberty¹, Eka Febri Zulissetiana¹

¹Faculty of Medicine, Sriwijaya University, Palembang

Email: pariyana07@gmail.com

ABSTRACT

Tuberculosis (TB) is a pulmonary infectious disease which is the main cause of increased morbidity and mortality worldwide. Diabetes mellitus is one of the factors that can increase the incidence of pulmonary tuberculosis. This study aims to determine changes in glucose levels in pulmonary TB patients at the beginning of being diagnosed with pulmonary TB or before receiving OAT therapy and after receiving OAT therapy. This study is an analytical study with a cross-sectional repeated measurement design, namely measuring blood glucose levels repeatedly for 2 times (the initial measurement of treatment before receiving OAT therapy and measurement after receiving OAT therapy). The sample in this study were 107 pulmonary tuberculosis patients who were treated at the Public Health Center in Palembang City. The sampling technique is proportional random sampling. Data analysis in this study was carried out descriptively and analytically using the t-test (Wilcoxon). The results showed that in TB patients with normal initial glucose level status, there was a significant difference in glucose levels before and after receiving OAT therapy (p-value = 0.000) p $<\alpha$, TB patients experienced binding of glucose levels after receiving OAT therapy while in TB patients with DM status, showed no significant difference in glucose levels before and after receiving OAT therapy (p-value = 0.135) p $>\alpha$.

Keywords: Pulmonary Tuberculosis, Blood Glucose Levels, OAT Therapy

1. INTRODUCTION

Pulmonary tuberculosis a disease caused contagious bv Mycobacterium tuberculosis. Mycobacterium tuberculosis is an aerobic germ that can live mainly in the lungs or other parts of the body that have high Data from the 2016 Annual Tuberculosis pulmonary Report, patients with tuberculosis in 2014 worldwide were 9.6 new cases of pulmonary tuberculosis in 2009 occurred in the Asia region by 50%, where Indonesia ranks fifth

oxygen pressure. This germ is in the form of a slender, straight or slightly bent stem with rounded ends, and has a high-fat content in the cell membrane, thus causing acid resistance and slow growth.¹ TB disease is a worldwide health problem because one-third of the world's population is currently infected with tuberculosis. after India, China, South Africa, and Nigeria. Indonesia is also the second-largest contributor to pulmonary tuberculosis cases after India, amounting to 647,000 cases.^{1,2}

The high number of pulmonary tuberculosis cases is influenced by the body's immune system, and several cases such as HIV, malnutrition, poverty, and population density. Apart from these factors, several studies have shown that diabetes mellitus can also increase the risk pulmonary tuberculosis. Diabetes mellitus is a condition characterized by increased blood sugar levels and has longcomplications involving term vessels, eyes, and kidneys. This disease affects many people who have an unhealthy lifestyle, namely high nutritional intake but low physical activity.

Similar to the prevalence of tuberculosis. the pulmonary annual prevalence of diabetes mellitus is also high. Based on data from the World Health Organization (WHO), Indonesia now ranks 4th in the number of people with diabetes mellitus in the world. In theory, diabetes mellitus affects the body's immune system, so that sufferers are more prone to contracting infections including Mycobacterium Tuberculosis infection. Patients with diabetes mellitus, including tuberculosis, were more common in the low-moderate socio-economic group.

Several studies have shown that diabetes mellitus can cause differences in clinical manifestations and responses to pulmonary tuberculosis treatment. especially when the blood sugar levels in the patient are high.⁴ The condition of hyperglycemia is a marker of disease severity and a good predictor in conditions of infection, sepsis, and critical illness.⁴ In a case study- Incontrast to other studies stated the presence that that hyperglycemia in TB patients did not affect treatment outcomes.⁶ In a study conducted by José Moreira, et al, it was found that TB patients with coinfection had recently started treatment with hyperglycemic TB., has a worse TB treatment result than euglycemia. ⁷

Based on the background description above, as a basis, the

researcher wanted to know the changes in blood glucose levels in pulmonary TB patients as an effort to increase the success of pulmonary tuberculosis treatment.

The formulation of the problem in this monograph is whether there is a change in blood glucose levels in pulmonary TB patients before and after OAT therapy as an effort to increase the success of pulmonary tuberculosis treatment.

The research objectives of early detection of changes in blood glucose levels as an effort to increase the success of pulmonary tuberculosis treatment in this monograph were to determine the blood glucose levels in the early phase of treatment before receiving OAT therapy in pulmonary TB patients, to determine the blood glucose levels of pulmonary TB patients after receiving OAT therapy and to find out changes blood glucose levels of pulmonary TB patients before and after receiving OAT therapy.

2. METHODS

The research design used a crosssectional repeated measurement design, namely measuring blood glucose levels repeatedly for two measurements. In this study, the initial measurements were carried out in pulmonary TB patients before receiving OAT therapy and three months after receiving OAT therapy. The study was conducted in the Palembang City Health Center Work Area in 2019. The study population was all pulmonary tuberculosis patients in Palembang City. The sample in this study were some pulmonary tuberculosis patients who were treated at the Public Health Center in Palembang City. There were as many as 107 samples. Descriptive analysis in this study is presented in the form of numbers percentages. Meanwhile. and inferential analysis used Wilcoxon statistical test to determine differences in blood glucose levels before and after

receiving OAT therapy in pulmonary TB patients.

3. RESULTS

The results of the study were collected as many as 107 pulmonary TB patients with pulmonary TB patient age, namely 45 years with a minimum age of 10 years and a maximum age of 85 years.

The most age found in this study occurred at the age of 20-59 years, as many as 78 people (72.9%), while those aged> 60 years were 21 people (19.6%) and at least at the age <20 years 8 people (7.5%). Based on gender, pulmonary TB patients in this study were mostly found to be male, as many as 65 people (60.7%) compared to 42 people (39.3%) (Table 1).

Table 1. Frequency Distribution Based on Age and Gender

Characteristics	Total (n)	Percent (%)
Age		
- < 20 year	8	7,5
- 20 – 59 year	78	72,9
$- \ge 60 \text{ year}$	21	19,6
Gender		
- Male	65	60,7
- Female	42	39,3

Results of Measurement of Glucose Levels in Pulmonary TB Patients

Distribution based on blood glucose levels in patients with pulmonary tuberculosis. The results of measuring the initial glucose level of TB patients or before receiving OAT therapy were divided into two categories, namely normal and diagnosed diabetes mellitus. The measurement results of pulmonary TB patients who had normal

glucose levels were 86 patients (80.4%) while those diagnosed with diabetes mellitus (blood sugar levels> 200mg / dL) were 21 patients (19.6%). Whereas in the last measurement in pulmonary TB patients with normal initial blood sugar levels and becoming uncontrolled, it increased by 23 people (21.5%) (Table 2).

Table 2. Frequency Distribution Based on the Results of Measurement of Glucose Levels in Pulmonary TB Patients

Measurement Results	Total (n)	Percent (%)
Initial Measurement	86	80,4
- Normal		
Mean \pm SD =110 \pm 33,17, min=36, max=198		
- DM	21	19,6
Mean \pm SD = 296 \pm 75,43, min=209, max=449		
Final Measurement		
- Controlled	84	78,5
Mean \pm SD = 121,75 \pm 27,64, min = 61, max = 200		
- Non-Controlled	23	21,5
Mean \pm SD = 261,72 \pm 53,28, min= 201, max= 387		

Differences in glucose levels before drug therapy and after receiving drug therapy in TB patients

The results of glucose research before and after receiving anti-tuberculosis drug therapy (OAT) were analyzed using the Wilcoxon test. Pulmonary TB patients with initial glucose level status in the normal category were 86 patients and 21 TB patients were diagnosed with DM. The results of the analysis of glucose levels before and after receiving OAT therapy in the group of TB patients who had initial glucose levels showed that there were

differences in glucose levels before and after receiving OAT therapy (p value = 0.000) p $<\alpha$. Pulmonary TB patients who were diagnosed with TB at the start of normal glucose level measurements after 3 months of receiving OAT therapy experienced an increase in glucose levels. Whereas in pulmonary TB patients who had initial glucose levels in the DM category as many as 21 patients, the results of the analysis were no differences in sugar levels before and after receiving OAT therapy (p value = 0.135)p> α .

Table 3. Differences in glucose levels before and after receiving OAT therapy based on glucose status

Normal	Therapy	n	Mean ±SD	p value
Glucose Status			(Min-Max)	
Normal	Before OAT therapy	86	110,80±33,17 (36-198)	0,000
	After OAT therapy	86	124,95±37,14 (61-329)	
DM	Before OAT therapy	21	296,19±75,43 (209-449)	0,135
	After OAT therapy	21	255,28±55,80 (180-387)	

4. **DISCUSSION**

Characteristics of pulmonary TB patients mostly occurred in men (60.7%) than women (39.3%). This is consistent with several studies which state that men are indeed more susceptible to infection with M. tuberculosis. This can be related to a greater smoking habit in men, which causes disorders of the immune system of the respiratory tract, making them more susceptible to infection. Disorders of the immune system of the respiratory tract can

be in the form of damage to mucociliary clearance due to toxins in inhaled cigarette smoke. Cigarette smoke can also damage phagocyte cells in the respiratory tract and decrease the response to antigens, thereby increasing the susceptibility of pulmonary tuberculosis.¹⁷ WHO (2012) also states that according to several studies, the incidence of pulmonary TB in women is lower than in men because less than women with pulmonary TB go to health care facilities. This happens because

women tend to find it more difficult to access health service facilities. community's negative stigma about pulmonary TB affects the psychosocial aspects of women more than men so that women tend to choose not to go to health service facilities. 16 The increasing age factor also affects the incidence of TB. In this study, it was found that almost all TB patients were in the age range of 20-59 years (72.9%). This is also in accordance with several other studies. According to Tabrani (2010), patients over 40 years of age are more susceptible to infection with M. tuberculosis. This is due to the biological changes that occur in the patient's body, especially in lung tissue, associated with aging. These changes can damage the barrier system and the microbial clearance mechanism of the respiratory system.¹⁸ The study, which was conducted in Semarang, showed that the results of patients with TB-DM were more common in patients aged 16-50 years, married, middle income. This vast age range suggests that pulmonary TB patients can occur at any age. This is because TB infection is also influenced by the environment and low immunity status. Women of reproductive age are more prone to developing TB than men in the same age range.²⁵

The results of the study on differences in blood glucose levels before and after receiving OAT therapy found that pulmonary TB patients with initial glucose level status in the normal category were 86 patients, the analysis showed that there was a significant difference in glucose levels before and after receiving OAT therapy (p value = 0,000). $<\alpha$,

glucose levels in TB patients have increased after receiving OAT therapy.

This research is in line with the results of the Gopathi kk study in 2015 which stated that there was an effect of OAT and tuberculosis on increasing blood sugar levels.²³ TB patients are at risk of experiencing DM, one of which is due to the impact of TB drugs in the pancreas which causes a decrease in insulin levels, which can increase blood sugar. Antituberculosis drugs commonly used are bactericides such as rifampin, isoniazid, pyrazinamide, streptomycin, and bacteriostatic ethambutol.²⁴ such as Isoniazid and rifampin have hyperglycemic effect. Pyrazinamide can cause difficult DM control. Rifampin induces metabolism and lowers blood sulfonylurea levels. 12 This drug can induce hyperglycemia the early in associated with increased absorption in the intestine, but there have been no apparent cases of diabetes.²³ Apart from the antituberculosis drugs themselves, TB disease can make TB patients susceptible to DM because the inflammation caused by cytokines such as IL6 and TNFa in response to TB infection can lead to increased insulin resistance and decreased insulin production, leading hyperglycemia.¹² The mechanism damage to the pancreas due to tuberculosis is probably caused by a mycobacterial attack directly to the pancreas through the spread of bacterial tubercles in the blood or through the penetration of the tissue of the abdominal lymph nodes around the pancreas. It was reported that when the pancreas is calcified there is a 23-50% incidence of DM.²³

5. CONCLUSION

Based on the results of research on early detection of changes in blood glucose levels as an effort to increase the of pulmonary tuberculosis success treatment, it can be concluded that in patients with normal blood sugar status before OAT therapy there statistically significant difference (p value = 0.000) p $<\alpha$, the patient's glucose level. TB has increased after receiving OAT therapy, while TB patients with DM status showed no significant difference in glucose levels before and after receiving OAT therapy (p value = 0.135) p> α .

REFERENCES

- [1]. Hapsari, Pantaria NF, Muhammad AI. Hubungan Sosioekonomi dan Gizi dengan Risiko Tuberkulosis pada Penderita DM Tipe 2.JurnalBerkalaEpidemiologi. 2017. 5(2):185-194
- [2]. Pande T, Huddart S, Xavier W, et al. Prevalence of diabetes mellitus amongst hospitalized tuberculosis paru patients at an Indian tertiary care center: A descriptive analysis. PLos ONE. 2018. 13(7):1–13.
- [3]. Gadallah M, Amin W, Fawzy M, et al. Screening for diabetes among tuberculosis paru patients: a nationwide population-based study in Egypt. African Health Science. 2018. 18(4):884–890.
- [4]. Eur RJ (ed). Prevalence and factor associated with diabetes mellitus among tuberculosis paru patients: a nationwide cohort. 2016. 48:264 268.
- [5]. Boillat-Blanco N, Ramaiya KL, Mganga M., et al. Transient Hyperglycemia in Patients with Tuberculosis paru in Tanzania: Implications for Diabetes

- Screening Algorithms. 2016. J Infect Dis 213(7): 1163-1172.
- [6]. Tabarsi P, Baghaei P, Marjani M, Vollmer WM., et al. Changes in Glycosylated Haemoglobin and Treatment Outcomes in Patients with Tuberculosis paru in Iran: A Cohort Study. 2014. J Diabetes MetabDisord 13(1): 123.
- [7]. José Moreira., et al. Hyperglycemia During Tuberculosis paru Treatment Increases Morbidity and Mortality in Contemporary Cohort of HIV-Infected Patients in Rio de Janeiro, Brazil. International Journal of Infectious Diseases. 2017. 69(2018): 11-19.
- [8]. Soegondo S. Konsensus Pengelolaan dan Pencegahan Diabetes Mellitus tipe 2 di Indonesia 2011. Jakarta : PERKENI.
- [9]. Giovanni D, M. Sali, F. Giovanni. 2013. The Biology of Mycobacterium tuberculosis Infection. Rome: Mediterranean Journal of Hematology and Infection Disease.
- [10]. WHO. 2018. Global Tuberculosis Report. WHO-int 2018.
- [11]. Hu F. Globalization of diabetes: the role of diet, lifestyle, and genes. Diabetes Care. 2011; 34:1249–1257. [PubMed: 21617109]
- [12]. Reed GW, Choi H, Lee SY, Lee M, Kim Y, Park H, Lee J, Zhan X, Kang H, Hwang S, Carroll M, Cai Y, Cho SN, Barry CE III, Via LE, Kornfeld H. 2013. Impact of diabetes and smoking on mortality in tuberculosis. PLoS ONE. 8:e58044. [PubMed: 23469139]
- [13]. Lonnroth K, Jaramillo E, Williams BG, Dye C, Raviglione M. 2013. Drivers of tuberculosis epidemics: the role of risk factors and social determinants. Soc Sci Med.

- 68:2240–2246. [PubMed: 19394122]
- [14]. Kuo MC, Lin SH, Lin CH, Mao IC, Chang SJ, Hsieh MC. 2013. Type 2 diabetes: an independent risk factor for tuberculosis: a nationwide population-based study. PLoS one
- [15]. Cahyadi A, Venty. Tuberkulosis Paru pada Pasien Diabetes Mellitus. J Indon Med Assoc. 2011;61(4).
- [16]. Gustaviani, Reno. Diabetes Mellitus di Indonesia : Buku Ajar Ilmu Penyakit Dalam Jilid III. Edisi IV : hal 1857-1881.Pusat Penerbitan Departemen Ilmu Penyakit Dalam FKUI : Jakarta;2010.
- [17]. Setiati S,ed et al. 2014. Buku Ajar Ilmu Penyakit Dalam. Jakarta: Interna Publishing.
- [18]. Rab, Tabrani. 2010. IlmuPenyakit Paru. Jakarta: TIM
- [19]. Kapur A, Harries AD, Lonnroth K, Bygbjerg C, Lefebvre P. Diabetes and tuberculosis-old associates posing a renewal public health challenge. US Endrocinology. 2009;5(1):12-14.
- [20]. Guptan A, Shah A. Tuberculosis and diabetes: an appraisal. Ind J Tuberc. 2000;47:3-8.
- [21]. Abdelbary BE, Garcia-Viveros M, Ramirez-Oropesa H, Rahbar MH, Restrepo BI. Tuberculosis-diabetes epidemiology in the border and non-border regions of Tamaulipas, Mexico. Tuberculosis (Edinb). 2016 doi:10.1016/j.tube.2016.09.024.
- [22]. Nadliroh zahrotun, Fathur Nur Kholis, Dwi Ngestiningsih. 2015. Prevalensi Terjadinya Tuberkulosis pada Pasien Diabetes Mellitus di RSUP Dr. Kariadi Semarang. Media Medika Muda. Vol 4 (4). hal 1714-25.

- [23]. Gopathi NR, Venu Mandava dan Sravani Makala. 2015. Tuberculosis and Diabetes: the deadly duo. International Journal of Advances in Medicine. Vol 2(3), hal 241-245.
- [24]. McMahon MM, Bistrian Bruce R. Host defences and susceptibility to infection in patients with diabetes mellitus. Infect Dis Clin North Am. 1995;9:1-9.
- [25]. Koziel H, Koziel MJ. Pulmonary complications of diabetes mellitus. Infect Dis Clin North Am. 1995;9:65-96.
- [26]. Dooley KE, Chaisson RE. Tuberculosis and diabetes mellitus : convergence of two epidemics. Lancet Infect Dis. 2009;9(12):737-46.
- [27]. Broxmeyer L. Diabetes mellitus, tuberculosis and the mycobacteria: two millennia of enigma. Med Hypotheses. 2005;65:433–9.
- [28]. Broxmeyer L. Diabetes mellitus, tuberculosis and the mycobacteria: two millennia of enigma. Med Hypotheses. 2005;65:433–9.
- [29]. Jabbar A, Hussain SF, Khan AA.
 Clinical characteristics of
 pulmonary tuberculosis in adult
 Pakistani patients with co-existing
 diabetes mellitus. Eastern
 Mediterranean Health Journal.
 2006;12:522-7.
- [30]. Patel AK, Rami KC, Ghanchi FD. Radiological presentation of patients of pulmonary tuberculosis with diabetes mellitus. Lung India. 2011;28:70.
- [31]. Sulaiman SA, Mohd Zain FA, Abdul Majid S, Munyin N, Mohd Tajuddin NS, Khairuddin Z, et al. Tuberculosis among diabetic patient. Webmed Central Infectious Diseases. 2011;2(12):1-13.

- [32]. Kapur A, Harries AD, Lonnroth K, Bygbjerg C, Lefebvre P. Diabetes and tuberculosis-old associates posing a renewal public health challenge. US Endrocinology. 2009;5(1):12-14.
- [33]. Guptan A, Shah A. Tuberculosis and diabetes: an appraisal. Ind J Tuberc. 2000;47:3-8.
- [34]. Nijland HM. R, Ruslami Stalenhoef JE, Nelwan EJ, Alisjahbana B, Nelwan RHH, et al. Exposure to rifampicin is strongly reduced in patients with tuberculosis and type 2 diabetes. Clin Infect Dis. 2006;43:848-54.
- [35]. Niazi AK, Kalra S. Diabetes and tuberculosis: a review of the role of optimal glycemic control. Journal of diabetes & metabolic disorders. 2012;11(28):1-4