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# Extrapulmonary Tuberculosis Scoring System Development. A Design for Android Studio Application Reza Fadilah

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#### 6 Abstract

Tuberculosis is an infectious disease with high incidence and mortality rates in the world. TB 7 not only attacks lung, but other organs called Extrapulmonary Tuberculosis (EPTB) as well. 8 At present, the diagnostic for EPTB is challenging since there are many examinations needed. 9 This study developed a scoring system for EPTB diagnosis based on some parameters. The 10 parameters derived from our studies evaluating AFB stain examination of biopsy tissue 11 associated with the clinical feature of EPTB patients. Parameter included a detailed medical 12 history and clinical examination, radiological, microbiological, molecular, and histopathological 13 investigations. The proposed scoring system used a minimum android version of jellybean 4.1. 14 The application is simple to accommodate users' low ability to operate. The scoring system is 15 proposed for accommodating many parameters important for diagnostic. It differentiates 16 which conditions are highly suspicious and should be included in the differential diagnosis. 17 EPTB Scoring system can be an alternative for EPTB diagnostic tools. 18

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20 Index terms— application, extrapulmonary tuberculosis, scoring system, diagnostic tools.

## <sup>21</sup> 1 Introduction

22 uberculosis is an infectious disease with high incidence and mortality rates in the world. Based on the World Health Organization (WHO) report in 2019, there is no single country free from Tuberculosis (TB).(1) Around 23 24 7 million Tuberculosis (TB) received treatment globally; however, the number of deaths was still hovering at 25 1.5 million patients. (2) Several factors that contribute to the high incidence of TB are sociocultural, including herbal medicines and stigma, adherence to drugs, TB program, and environment. (2)(3)(4)(5)(6) (7) The estimated 26 incidence of TB cases in Indonesia is 845,000 cases per year, with 570,289 notified cases. Approximately 32% of 27 cases have not been counted, either because they are not detected or not reported. (8) (9) Other than infected the 28 lung, TB can also attack other organs such as bones, lymph nodes, and abdominal areas known as extrapulmonary 29 tuberculosis (EPTB). (10) (11) The incidence of EPTB varies across regions ranging from 8% ?34% of all TB 30 cases. Extrapulmonary tuberculosis (EPTB) levels vary widely -depending on their risk factors -including host 31 immunological response, socio-demographics, comorbidities, lifestyle behaviors, genetic factors, and lymph node 32 pathogenesis. (12) (13) It also depends on the previous history of pulmonary TB, non-adherence to taking anti 33 TB drugs, and failure to therapy due to drug resistance.(3) (??4) Drugs of choice and inaccurate diagnoses 34 35 depend on a country's socio-economic level and the resources devoted to the TB program.(2)(15) Direct sputum 36 smear microscopy is the most widely used method for diagnosing pulmonary TB and is available in most primary 37 health-care laboratories at the health-center level. Because TB can present with many different symptoms, the first obstacle in diagnosing smear-negative TB is discerning the varied clinical presentations. (13) (16) It is 38 essential to determine which conditions are highly suspicious and included in the differential diagnosis. (17) (18) 39 Previous studies have found that using a scoring system can support pulmonary TB diagnosis. (19) (20) This 40 paper discussed the proposed scoring system for EPTB diagnostic tools based on several parameter discussed 41 above. The data used for the parameters in the application were based on our study of 1.034 TB cases registered 42 from 2015-2018 in Bandung. 43

### 44 **2** II.

#### 45 3 Approach

A diagnostic approach to an AFB smearnegative patient with possible TB includes, where available, a detailed 46 medical history and clinical examination, as well as radiological, microbiological, molecular, and histological 47 48 investigations. (21) Diagnosis of pulmonary tuberculosis in adults with clinical manifestations of chronic cough 49 and sputum can be quickly made using the production of sputum smears with the Ziehl Neelsen AFB staining method. However, this method is challenging for the diagnose in patients with a non-productive cough that 50 cannot expel phlegm and TB patients with HIV/AIDS or other immunecompromised conditions. (?? Below 51 is the diagram showing the overall phase used in this study. Figure 1 taken initially from (21) described the 52 approach used for the base of this study. F results. The results were presented according to the number of 53 systemic manifestations and combined local manifestations and AFB and histopathological results. From 1,034 54 biopsied TB sputum smear using the AFB staining method and other supporting examination, we found 44 55 patients with EPTB, 3.4% of the total, which less than the average figure from the literature. 56

Positivity degree of EPTB recorded by AFB staining from paraffin block test preparations, using International
 Union Against Tuberculosis and Lung Disease (UATLD) scale modification on sputum smears.

AFB staining from the test material's sputum smear can be homogeneous, while the preparations from paraffin 59 60 blocks cannot be homogenized because the test preparations come from different tissues. To accommodate those, 61 the analysis of the data developed using two types of interpretation, some using the degree of positivity, and others used positive and negative findings only. EPTB based on the BTA count results and its relationship 62 with systemic manifestations and local manifestations in EPTB patients is in the table below. Detected clinical 63 manifestations in EPTB patients are divided into systemic manifestations and local manifestations. Systemic 64 manifestations consist of fever, night sweats, fatigue, and weight loss. It appears that the number of patients 65 who complain of fever is the same as those who complain of fatigue. Interestingly, the number of patients who 66 67 complained of night sweats was the same as patients complained of weight loss. In addition to systemic clinical 68 manifestations, there are also local clinical manifestations.

On these local manifestations, the symptoms depend on the origin of the organ where EPTB is located. Patients diagnosed with lymph node EPTB complain of enlarged lymph nodes accompanied by pain, while patients diagnosed with EPTB have lumps and pain. Overall, all EPTB patients complained of symptoms related to the origin of the organ where EPTB locate, although the number of complaints differed from patient to patient.

Most of EPTB occurred in the lymph nodes (48.6%), while the rest were evenly distributed among other 74 organs, including bones, breasts, perianal, ileum. The findings were consistent with studies in other developing 75 76 countries that have previously reported that the most frequent EPTB incidence location is in the lymph nodes 77 (12)(13)(14)(15)(16). Histopathological readings from EPTB infection paraffin blocks found 97% of cases with 78 granuloma (+); however, the AFB (+) was only 8.5%. It can be explained since granuloma collects several inflammatory cells, especially mature macrophages that form aggregates in response to an antigen. An antigen 79 80 can come from a bacterium, a fungus, a foreign object, or an immune complex. The purpose of granuloma formation is to isolate the host body's antigen and facilitate the eradication of the antigen. Early in granuloma 81 formation, antigen-presenting cells express a wide variety of proinflammatory and chemoattractant cytokines. 82 The AFB staining results on the extrapulmonary TB patients' biopsy tissue based on the degree of positivity 83 were mostly with a value of +1 (8.1%). There was no correlation between the degree of AFB's degree of positivity 84 from biopsy tissue with clinical symptoms in this study. 85

86 Most EPTB sufferers do not experience systemic symptoms such as intermittent fever, night sweats, weight 87 loss as experienced by TB patients in general. It is different from previous studies, which explained that systemic symptoms in EPTB patients were around 52.2%. (16) In this study, it was found that systemic fever complaints 88 only occurred in about 16% of cases, night sweats in 5%, 5% experienced weight loss, 16% fatigue. In contrast, in 89 the previous study, fever complaints were found in 37-80% of cases. (??2) This study found that 100% of local 90 clinical symptoms were statistically significant in patients with extrapulmonary TB based on the histopathological 91 features and AFT staining with various positivity degrees. In contrast with previous studies, which explained 92 that local symptoms in EPTB patients were only around 72%. (17) The histopathological examination results 93 on the biopsy tissue were almost entirely with (+) granulomas since granuloma image is a characteristic feature 94 of TB infected tissue. (15,18,19) There were no systemic clinical symptoms (97%) such as fever, night sweats, 95 weight loss, and fatigue in patients with granuloma (+) features found on the histopathological result. In contrast 96 97 with the other studies, which stated that the characteristic of TB infection histopathological is associated with 98 the clinical manifestations of EPTB patients (15,(18)(19)(20)(21)). Based on the above results and considering 99 the scoring system that was previously used to enforce TB in children, a new scoring system is planned as an 100 innovation to assist in diagnosing EPTB. This design expected to capture EPTB cases, taking into account several longused laboratory parameters found in this study. Other laboratory parameters will also support it as predictors 101 of TB, such as erythrocyte sedimentation rates, monocyte/lymphocyte ratio, and neutrophil/lymphocyte ratio. 102 (22)(23) ????4) This scoring system's design uses a simple application that can be used using Android Studio since 103 the aims are to use a broader community for EPTB case finding networking. (25) The simple design is chosen 104 considering user ability on operating apps, which is relatively low. The "Extra-pulmonary TB Score Calculation 105

System (EPTBScore)" application is an android based mobile application run on an android smartphone with
 a minimum android version of jelly bean 4.1. This system works by calculating the total score based on 22
 parameters, as described in Table 3. In the application there are seven application pages, as follows:

Opening Page; The opening page in the application only displays information of the the application name and originator. Figure 3 illustrates the opening page. The following pages will be for language preferences followed by pages for personal information such as name, age, and gender of the patient. The pages are illustrated in Figure The above proposed application designed to accommodate the need for several parameter inclusion. We developed a very simple application with regard to the user ability in operating application. This application is still in the development process with future plan of accomodataing user experience and needs.

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# 116 **4 III.**

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#### 117 5 Conclusion

This paper presented the proposed EPTB Scoring Application used to diagnose Extrapulmonary Tuberculosis. The scoring system is ideal for accommodating many parameters from either systemic and local manifestation. Parameters chose using detailed medical history and clinical examination and radiological, microbiological, molecular, and histological investigations. It can also differentiate which conditions are highly suspicious and should be included in the differential diagnosis. EPTB Scoring system can be an alternative for EPTB diagnostic tools.

 $<sup>^1 \</sup>odot$  2020 Global Journals Extrapulmonary Tuberculosis Scoring System Development. A Design for Android Studio Application

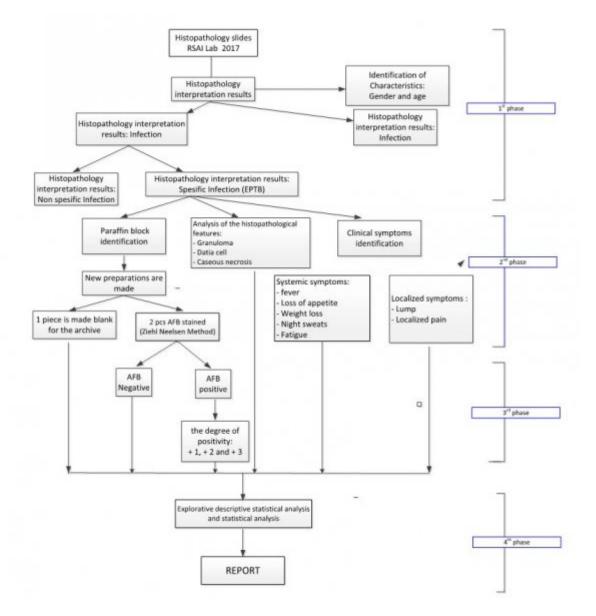


Figure 1:

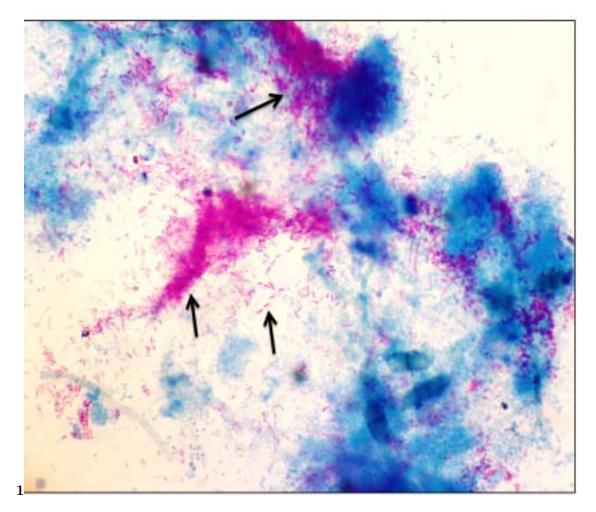


Figure 2: Figure 1 :

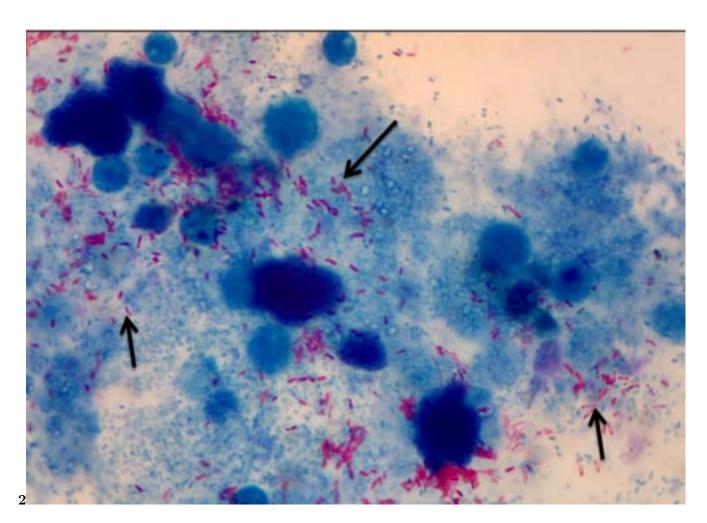


Figure 3: Figure 2 :

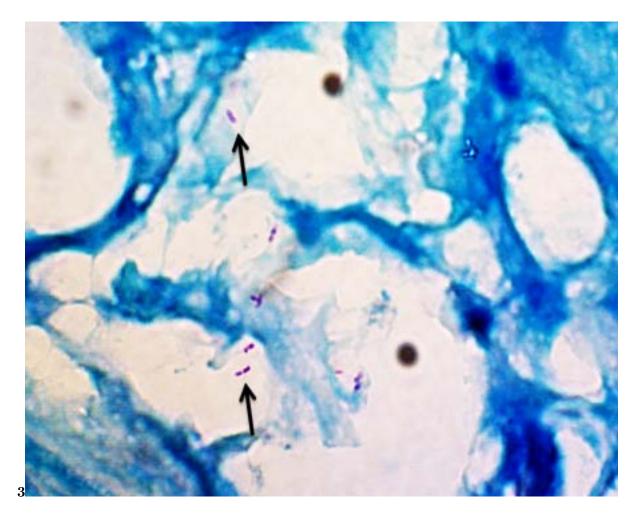


Figure 4: Figure 3 :

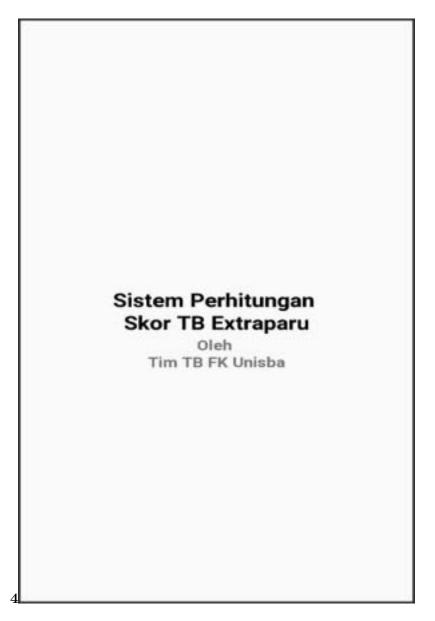


Figure 5: Figure 4 :





## 

Clinical				AFB stained	d			
Manifestation		NA		?	+			Total
	n	%	n	%	n	%	n	%
Systemic								
Manifestation								
Fever								
Yes	3	10	21	70	6	20	30	100
No	2	$28,\! 6$	5	71,4	0	0	7	100
Total	5	$13,\!5$	26	70,3	6	$^{8,1}$	37	100
Weight Loss								
Yes	4	$11,\!4$	25	71,4	6	17,2	35	100
No	1	50	1	50	0	0	2	100
Total	5	$13,\!5$	26	70,3	6	$^{8,1}$	37	100
Night Sweat								
Yes	4	$11,\!4$	25	71,4	6	17,2	35	100
No	1	50	1	50	0	0	2	100
Total	5	$13,\!5$	26	70,3	6	$^{8,1}$	37	100
Fatique								
Yes	4	$11,\!4$	25	71,4	6	17,2	35	100
No	1	50	1	50	0	0	2	100
Total	5	$13,\!5$	26	70,3	6	$^{8,1}$	37	100
Local								
Manifestation								
Yes	0	0	0	0	0	0	0	100
No	5	$13,\!5$	26	70,3	6	16,2	37	0
Total	5	$13,\!5$	26	70,3	6	16,2	37	100

Figure 8: Table 1 :

 $\mathbf{2}$ 

Year 2020 34 Volume XX Issue XII Version I D D D D ) Global Jour- nal of Medi- cal Research (	Clinical Manifestation Systemic Manifestation Fever Yes No Total Weight Loss Yes No To- tal Night Sweat Yes	No Granuloma n % 1	3,3 0 (	) 1 2	,7 1 2,9 0 0 1 2,7 1 2,9 Histopatho					
	No	0	0	2	100					
	Total	1	$^{2,7}$	36	97,3					
	Fatique		,		,					
	Yes	1	$_{3,2}$	30	96,8					
	No	0	0	6	100					
	Total	1	2,7	36	97,3					
	Local Manifestation									
	Yes	0	0	00	0					
	No	1	2,7	36	97,3					

2,7

36

97,3

 $[Note: F @ 2020 \ Global \ Journals \ Extrapulmonary \ Tuberculosis \ Scoring \ System \ Development. \ A \ Design \ for \ Android \ Studio \ Application]$ 

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Figure 9: Table 2 :

3

PARAMETER

Total

Figure 10: Table 3 :

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