Risk Factors for Extra Pulmonary Tuberculosis in the Health District of Thies, Senegal

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Abstract: Tuberculosis (TB) is a national health priority in African countries particularly pulmonary form. But Extrapulmonary TB (EPTB) is increasing with less consideration for the governments. However, it has a negative impact in the social, economic and sanitary context. Few studies on the disease have been conducted in the African continent. The objective of this study was to estimate the incidence of Extrapulmonary TB in the Health District of Thies, a Senegalese province. Through a retrospective study, we collected data of all cases of TB admitted at the TB Division of the Health District of Thies, Senegal from January 2012 to December 2014. Sociodemographic characteristics of the patients, past medical history, lifestyles, the type of TB (pulmonary or extrapulmonary), the treatment regimen and the prognosis was evaluated. Uni, bi and multivariate analysis were done and results expressed with a 95% confidence interval. On average, 824 patients with TB were included. They were mostly male (68.4%), jobless (62.9%), not married (58.9%) with a mean age of 33.3 years (± 14.3). They had in their past medical history a notion of TB contamination (30.1%), BCG vaccination (96.5%), hypertension (4.6%), diabetes (2.7%), HIV infection (1.7%), smoking (11.5%) and alcohol consumption (2.8%). Hundred and Five patients (12.7% with CI 95%: 10.7% - 14.7%) had EPTB of pleural (56.2%), lymphatic (12.4%), bone (9.5%) and gastrointestinal (9.5%) localization mainly. Age, notion of contact with patient, employment and HIV infection were associated independently to EPTB. It is important to take into consideration these results to strengthen primary prevention strategies.

Keywords: Extrapulmonary Tuberculosis, Tuberculosis, Risk Factors, Thies, Senegal

1. Introduction

Tuberculosis (TB) is an infectious disease, endemo-epidemic and a health priority worldwide. With the occurrence of Human Immunodeficiency Virus (HIV) infection, TB is increasing either in developing than developed countries. ExtraPulmonary Tuberculosis (EPTB) characterized by the development of TB outside the lungs is also increasing in term of frequency worldwide [1-8]. However, clinical presentations are various and delaying the diagnosis. Some patients prefer living alone or staying at

home without going to hospital for diagnosis and treatment because of stigma [9, 10]. However, contact with TB patient alone sometimes can be sufficient to start TB treatment. All these factors could delay the treatment and compromise the vital and functional prognosis of EPTB patient [10]. This situation can compromise vital and functional prognosis of patients. Few studies have been conducted in Africa regarding ExtraPulmonary Tuberculosis. Several factors could explain this situation. However it is shared clinically by different specialties, it seems that it is underdiagnosed. Declaration of the disease could be missing. The other factor

could be physician-based because of absence or delay in the screening of TB among patients. This could be related to lack of training, poor knowledge on the disease and lack of lab equipment in the African continent [1]. In another view, sometimes, treatment is proposed empirically based on clinical and radiological arguments. Studies conducted worldwide have identified several risk factors associated with the occurrence of EPTB: age [1, 2, 9], female sex [1, 11, 12], no vaccination [13], contact with TB patient [5, 12, 14], alcohol consumption [11, 15], employment [13, 16], diabetes [15] and HIV infection [1, 11] particularly. Senegal, like many Sub-Saharan African countries, is facing the TB epidemics. Thus, 10 975 TB cases detected countrywide in 2008 and 11 734 cases in 2009. Every year, according to the TB Program Experts, almost 70% of screened cases are expected of whom 80% should be treated to stop the transmission chain. The region of Thies is the third region where TB cases are prevalent (68%) after Ziguinchor (72%) and Dakar (79%) [17]. To fight TB in general, decision-makers need epidemiological studies for better management of dedicated program. The aim of this study was to identify the main risk factors associated to EPTB in the Health District of Thies, Senegal.

2. Materials and Methods

Through a retrospective study, we collected charts of all TB patients who were followed at the TB Program Division of the Health District of Thies, Senegal from January 1st 20012 to December 31st 20014. The Health District of Thies covered a population of 632214 inhabitants. It had a health center and several health posts. The TB Program Division is located in the health center. It ensures the treatment of TB patients following the World Health Organization (WHO) Guidelines [18]. However, TB treatment is decentralized to health post for a better management of TB patients, they all have a medical/record at the TB Program Division of the health center of Thies. In the record, all required and useful informations are collected. The TB Program is run by a team coordinated by a Medical Doctor with nurse-assistant and community health workers trained for the task. During our study, all TB patients followed during the study period and who had finished their TB treatment were included in the study population. However, those who did not finish their treatment in 2014 were excluded. We collected data related to the sociodemographic characteristics of the patients (age, sex, employment, education, marital status), the past medical history (High Blood Pressure, Diabetes, HIV infection, BCG vaccination, Cancer, immunosuppressive treatment and others diseases), the lifestyle (smoking, alcohol), the notion of contact with TB patient, the localization of TB (Lung, Extrapulmonary TB) and prognostic (recovering, relapse, loss of follow-up, death). Data processing was done using SPSS 20 package for Windows. Uni, bi and multivariate analysis were realized. So, we computed frequencies, mean with standard deviation first. Secondly, we compared percentage of EPTB according to sociodemographic

characteristics of the patients, their past medical history, lifestyle to select independent variables to put in the final model for multivariate analysis. Odds ratios were computed and expressed with a 95% Confidence interval (CI).

3. Results and Discussions

3.1. Results

Eight hundred and twenty four patients were included during our study. They were mostly male (68.4%), married (58.9%), unemployed (62.9%) with a mean age of 33.3 years (± 14.3). Contact with TB patient (69.9%), BCG vaccination (96.5%), Hypertension (4.6%), diabetes (2.7%) and HIV infection (1.7%) were the main past medical history. Smoking (11.5 %), alcohol consumption (2.8%) and drug use (0.60 %) were the main lifestyle observed in our population (Table 1).

Table 1. Characteristics of the patients.

Variables	Frequency	Percentage (%)
Age		
Mean: 33.3 years (± 14.3 , 1-85 years)		
Median: 31 years		
≤ 19 years	114	13.8
20-39 years	474	57.5
40-59 years	186	22.6
60 years and plus	50	6.1
Male Sex	564	68.4
Employment (yes)	306	37.4
Married (yes)	339	41.1
Past medical history		
High Blood pressure	38	4.6
Diabetes	22	2.7
HIV infection	14	1.7
Contact with TB patient	578	69.9
Immunization with BCG	795	96.5
Lifestyles		
Smoking	95	11.5
Alcohol	23	2.8
Drug use	5	0.6

During treatment, prognosis was characterized by remission in 72% of the patients while a lethality rate of 5.5% was observed. Among our population of patients, 105 had extrapulmonary tuberculosis [12.7%; IC_{95%}: 10, 7-14, 7] of whom 99 had pure EPTB and 6 EPTB with lung TB. The mean number of cases was 35 per year. The main localization was predominantly pleural (56.19%), lymph nodes (12.40%), bone and gastrointestinal (9.52%) (Table 2).

Table 2. Localization of ExtraPulmonary Tuberculosis (N=105).

Localization of the EPTB	Frequency	Percentage (%)
Pleural	59	56,2
Lymph nodes	13	12,4
Bone-articulation	10	9,5
Digestive system	10	9,5
Miliary	5	4,8
Pericardia	4	3,8
Skin	2	1,9
Urogenital system	1	0,9
Nervous System	1	0,9

In the bivariate analysis, proportion of EPTB varied with age of the patients ($p \leq 0.002$), contact with TB patient ($p \leq 0.046$), employment ($p \leq 0.005$) and HIV infection ($p \leq 0.002$) (Table 3).

Table 3. Proportion of EPTB according to characteristics of the patients.

Variables	Frequency	Percentage (%)	P Value
Age			0.002 *
≤19 years	15	13.2	
20-39 years	47	9.9	
40-59 years	29	15.6	
60 years and over	14	28	
Sex			NS
Male	65	11.5	
Female	40	13.4	
Employment			0.046 *
Yes	52	10.8	
No	53	15.5	
Married			NS
Yes	52	15.3	
No	53	10.9	
Past Medical History			
High Blood pressure			NS
Yes	3	7.9	
No	102	13	
Diabetes			NS
Yes	1	4.5	
No	104	13	
HIV Infection			0.002 *
Yes	6	42.9	
No	99	12.2	
Contact with TB patient			0.004 *
Yes	86	14.9	
No	19	7.7	
Immunization with BCG			NS
Yes	101	12.7	
No	4	13.8	
Lifestyles			
Smoking			NS
Yes	10	10.5	
No	95	13	
Alcohol			NS
Yes	3	13	
No	102	12.7	

Younger age ‘1-19 years [OR=3.74; CI_{95%}: 1.55-7.02], 20-39 years [OR=3.58; CI_{95%}: 1.75-7.20]’, notion of TB contamination [OR=2.22; CI_{95%}: 1.30-3.78], absence of HIV infection [OR=0.19; CI_{95%}: 0.06-0.57] and having an employment [OR=0.52; CI_{95%}: 0.33-0.82] were independently associated with Extra Pulmonary Tuberculosis at the multivariate logistic regression analysis (Table 4).

Table 4. Results of the multivariate analysis.

Variables	OR	95% IC	P value
Age			
1-19 years	3.74	[1.55 – 9.02]	0.003
20-39 years	3.55	[1.75 – 7.20]	0.000
40-59 years	2.02	[0.95 – 4.28]	0.66
60 years and over	1	-	-
Employment			
Yes	0.52	[0.33 – 0.82]	0.005
No	-	-	-
Contact with TB patient			

Variables	OR	95% IC	P value
Yes	2.22	[1.30 – 3.78]	0.004
No	1	-	-
HIV infection			
No	0.19	[0.06 – 0.57]	0.003
Yes	1	-	-

3.2. Discussions

Through a retrospective study, we have identified that younger age, employment, contact with TB patient and HIV infection were the main risk factors for ExtraPulmonary Tuberculosis in the Health District of Thies, Senegal. The age 0-19 years and 20-40 years were associated with a high risk for EPTB in our population of patients compared to 60 years and over. This means that the younger the patient is the higher the risk. This result confirms what was already found worldwide. The role of age as risk factors for EPTB has been highlighted by several studies worldwide. In Dakar, Senegal, a mean age of 36 years [19], 37 years [20] and 39 years [21] were found respectively with the age range of 30-44 years and 14-45 years being the most affected. In Benin [1] and Madagascar [22], similar results were almost observed with the age range of 25-34 years being the most affected. This was also the case in China [23], Saudi Arabia [24], Brazil [15], Germany [6] and USA [8]. According to Sandgren et al. [25], the European Union adult population in the range age of 25-44 years (36.7 %) is the most affected by EPTB during a study among 167652 patients. The role of age could be explained by several factors. Young subjects are more exposed to be in contact with TB patient. They are more mobile and more at risk to meet TB patients in workplace, public transport and at home. Precarity with jobless and poverty are more frequent among youth exposing the young subjects to contact with Mycobacterium tuberculosis. The third factor could be related to HIV infection which is more common in young subjects exposing to TB. Employment was associated with EPTB in our study population. This result confirmed the role of employment in the occurrence of TB in general and EPTB in particular through several studies conducted worldwide. Indeed, in Africa, Oni et al. [13] showed that being employed was associated with a low risk [0.46; 95% CI: 0.31-0.69] for the occurrence of tuberculosis. In Europe, same results were observed in Denmark [26] and in Croatia [7]. In the USA, unemployment was associated with EPTB [12, 16]. This is the case too in China [27] and Japan too [28]. Indeed, having a job provides income, improves standard of living, prevent from promiscuity and thus protects against tuberculosis. These results confirm the hypothesis that tuberculosis is usually a disease of poverty. However, among the jobs, those related to the health environment were the most at risk. Indeed, according to some authors, health care professionals are those who are at greater risk of developing tuberculosis. This could be related to the fact that health professionals are more in contact with patients and also pathological products (sputum, cerebrospinal fluid, bones, pleural liquid and so on) than no health professionals. Contact with TB patient was associated

with the occurrence with EPTB with a risk of 2.22 [95% CI: 1.30 - 3.78]. This result confirms what has been observed elsewhere in the world. The same risk was observed during studies conducted in Africa especially in Zambia [14], Malawi [29], Ethiopia [5] and Gambia [30]. In the USA, the risk was 1.7 [11]. In Europe, it was 2.19 in Croatia [7] and 2.06 in Georgia [31]. In Asia, same results were observed in China [27] and Japan [28]. For some authors, the risk is higher among health personnel. In China [27], the risk among this group was 2.47 while in Japan it was 25 among laboratory technicians [28]. These results confirm that contact with TB patient is an important risk factor for the occurrence of tuberculosis in general and EPTB is not an exception. A person with untreated pulmonary tuberculosis with positive sputum exam can contaminate his entourage with Pflügge droplets when coughing or sneezing. This is related to the fact that the main route of TB transmission in human being is airborne. However, the notion of TB patient contact remains underestimated, especially in Africa where TB patients are not so isolated even during hospital admission. With the visits of the family during hospitalization, contamination remains common. This is the case also in public transportation, working place and public stations where promiscuity is also common. Otherwise, some TB patients hide their illness because of stigmatization which increases the risk of TB contamination. This is another reason to search for TB contact in case of ExtraPulmonary Tuberculosis observed in our daily consultation. Absence of past medical history of HIV infection was associated with the occurrence of Extrapulmonary Tuberculosis with a risk of 0.19. HIV infection is a significant risk factor for EPT as observed during studies conducted worldwide. The association HIV and Tuberculosis is common so that it is recommended to screen for HIV whenever TB occurs and vice versa. However in Senegal, few studies focused on the association between HIV infection and Extrapulmonary Tuberculosis [19, 25]. Otherwise, this association is almost common in Zambia [14], Ethiopia [21], South Africa [32] and Burkina Faso [33] in the African continent where HIV infection is very frequent. In Europe, the same risk was observed in Portugal [24], France [34] and Spain [35]. This is the case too in USA [11, 36, 37] and Brazil [15] in the American continent. In Thailand, Kingkaew et al. [38] during a study on the epidemiology and risk factors for death followed 308 patients presenting Extrapulmonary Tuberculosis over 1 year. HIV infection was associated with EPTB tuberculosis with a risk of 1.70. These results confirm the role of HIV infection on the occurrence of EPTB worldwide. In fact, HIV infection leads to immunosuppression. This situation promotes the reactivation of quiescent mycobacterial bacilli, their multiplication and also the destruction of immune/defense mechanisms. Thus, the body become vulnerable and more easily compromised leading to TB infection. It is important to screen for TB whatever the localization when a patient is HIV positive through sputum test or other immunological tests like Genexpert or Quantiferon. By the way, whenever TB

infection is suspected, it is recommended to start antimycobacterial treatment to tackle infection before its spread which can compromise the functional and vital prognosis. However, it presented some limits related to many factors: identification of the cases which is sometimes based on presumptive signs and symptoms leading to misclassification of cases and under or upper estimation of the cases. The second factor is related to the identification of risk factors which could have been missing for some patients.

4. Conclusion

This study has helped to identify risk factors associated with ExtraPulmonary Tuberculosis among TB patients followed at the TB Program Division of the Health District of Thies, Senegal. Younger age, employment, contact with TB patient and HIV infection were independently associated with ExtraPulmonary Tuberculosis in the Health District of Thies, Thies, Senegal. It is important to take into consideration these results to strengthen preventive strategies to tackle this endemic disease in the Health District of Thies, Senegal.

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