

Prevalence of diabetes mellitus in newly diagnosed pulmonary tuberculosis in Beira, Mozambique.

Damiano Pizzol¹, Francesco Di Gennaro^{1,2}, Kajal D. Chhaganlal³, Claudia Fabrizio²,
Laura Monno², Giovanni Putoto⁴, Annalisa Saracino²

1. Research Section, Doctors with Africa CUAMM; Beira – Mozambique
2. Clinic of Infectious Diseases, University of Bari, Bari, Italy
3. Center for Research in Infectious Diseases, Faculty of Health Sciences, Catholic University of Mozambique; Beira - Mozambique
4. Research Section, Doctors with Africa CUAMM; Padova – Italy

Abstract:

Introduction: Data regarding the association between diabetes mellitus (DM) and tuberculosis (TB) in Africa are scarce. DM screening among TB patients in Mozambique was carried out.

Methods: The study was implemented from January to August 2016 in three Urban Health Centers in Beira, Mozambique and recruited adult (>18 years) patients newly diagnosed with pulmonary TB.

Results: Three hundred and one patients were enrolled (67.4%, males mean age 31.7(SD 11 years). Diabetes was diagnosed in only 3 patients (1%) and impaired glucose tolerance (IGT) in an additional 6 subjects (2%).

Conclusion: A lower than expected prevalence of DM was observed, which could be explained by the lack of traditional risk factors for DM (overweight, age over 45 years, hypertension and smoking) in Mozambique.

Keywords: Diabetes mellitus, pulmonary tuberculosis, Beira, Mozambique.

DOI: <https://dx.doi.org/10.4314/ahs.v17i3.20>

Cite as: Pizzol D, Gennaro FD, Chhaganlal KD, Fabrizio C, Monno L, Putoto G, Saracino A. Prevalence of diabetes mellitus in newly diagnosed pulmonary tuberculosis in Beira, Mozambique. *Afri Health Sci.* 2017;17(3): 773-779. <https://dx.doi.org/10.4314/ahs.v17i3.20>

Introduction

Tuberculosis (TB) is one of the leading causes of death worldwide. Mozambique represents one of the most affected countries in the world, with an estimated prevalence of over 500 per 100,000 population¹. Although great efforts have already been made, the way to defeat TB remains long. The WHO End TB Strategy aims to a 90% reduction in TB deaths and an 80% reduction in the TB incidence rate by 2030¹.

Many factors contribute to the lack of efficacy in containing TB, as social determinants of health (SDH) and concomitance of other co-morbidities, such as co-infections (in particular with HIV) and non-communicable diseases, such as diabetes mellitus (DM)².

SDH, which can be defined as conditions in which people are born, grow, live, work and get old, are strongly influenced by the distribution of wealth, power, resources and they have an immediate impact on health². In fact, it is well known the association between low socio-economic levels and TB³. Furthermore, TB is the most common opportunistic infection among HIV-positive patients and, especially in sub-Saharan Africa, the burden of HIV/TB co-infection is the highest worldwide, with almost 80% of all cases of incident TB in persons with HIV infection⁴.

Growing evidence suggests that TB and DM represent a mutual risk factor of occurrence of each other⁵, causing also a reciprocal worsening, due to both pathogenic mechanisms⁶, and metabolic factors⁷. In fact, if on one hand patients with DM have a higher risk of acquiring TB, on the other hand, patients with DM and TB have a high likelihood of delayed diagnosis, healing, and increased severity of symptoms and mortality for both diseases⁸. Nowadays, sub-Saharan Africa and low income

Corresponding author:

Francesco Di Gennaro,
Clinic of Infectious Diseases,
University of Bari, Bari, Italy
Email: cicciodigennaro@yahoo.it

countries are increasingly affected by DM⁹⁻¹⁰ and, the increasing frequency of DM co-existing with TB and HIV, is an example of interaction between communicable and non-communicable diseases, requiring a multidisciplinary and integrated approach⁸. In Mozambique, data on the prevalence of DM in TB patients are limited. The few data available on DM in this country in the general population shows a national prevalence of 2.9%¹¹⁻¹², and people are often unaware of their condition¹¹⁻¹². The present study aimed to assess the prevalence of DM in patients newly diagnosed with pulmonary TB in Beira, the second largest city in Mozambique, to better define the relationship between TB and DM.

Patients and methods

Study design and population

This study took place in three urban health centres of Beira city involved in the National Tuberculosis Control Programme (PNCT): Ponta-Gea, Munhava and Macurungo, that are largest in Sofala's Province. The study was implemented from January to August 2016 and we recruited a total of 301 patients who were newly diagnosed with pulmonary TB. Inclusion criteria were: age ≥ 18 years, a confirmed TB diagnosis (positive sputum smear result or GeneXpert positive result or culture positive result) and starting of anti-TB treatment. Exclusion criteria were: previous TB treatment or TB treatment started within the previous two months. Informed consent was requested from all enrolled patients. Anti-TB treatment was prescribed according to guidelines of the National Tuberculosis and Leprosy Control Programme of Mozambique¹³.

Methods

A face-to-face interview conducted by a trained nurse encompassed questions about demographic characteristics (age, residence, education, occupation, marital status, monthly income), possible pregnancy, risk behaviours (sexual behaviour/partnerships, concurrent sex partners, condom use, smoke and alcohol abuse, etc.) and medical history, including TB and diabetes symptoms. A basic physical examination (vital signs, weight, height, waist circumference, blood pressure and general appearance) was performed. The body mass index (BMI) was calculated. Moreover, subjects with unknown HIV status received a pre-HIV test, other suggestion to undergo a rapid HIV test, if not done before. For the diabetes diagnosis, two

consecutive fasting blood glucose tests were performed to each participant. We did a screening and diagnosis for TB infection according to clinical, microbiological and radiological algorithm established by the WHO Guidelines for patients with or without HIV infection¹ and in patients with TB infection, the clinical severity was evaluated based on extension, site and number of localizations of disease (pulmonary/extrapulmonary). According to the WHO guidelines, patients were considered as non-diabetic if both measurements were ≤ 110 mg/dl, and as diabetic if both measurement were above 126 mg/dl. If at least one value was between 110 and 126 mg/dl, the Oral Glucose Tolerance Test (OGTT) was performed: patients were considered diabetic when plasma glucose at 2 hours was ≥ 200 mg/dl¹⁴.

All TB-diabetic patients underwent fundus examination, urinary stick and assessment of diabetic complications.

Statistical analysis

Data obtained from the answers to the questionnaire were imported on Excel 5.0 and analysed using the STATA 13.0 software.

Mean and standard deviation for continuous variables and frequency for categorical variables were calculated as descriptive statistics. Due to the small number of diabetic patients, a comparison between diabetic and non-diabetic subjects was not possible.

Ethical approval

The study was approved by the Comité Nacional de Bioética para a Saúde/ National Bioethics Committee for Health by the protocol Ref: 168/CNBS/15.

Results

Socio-demographic characteristics

The social-demographic characteristics of the study population and patients' life style are reported in Table 1. The 301 patients with a new diagnosis of pulmonary TB included 203 males (67.4%) and 98 females (32.6%), mean age 36.7 years. Nearly a half 141 patients (47.5%) had no educational degree; 157 (52.9%) patients were employed and only 10 (3.8%) had a monthly income higher than 100 Euros. Only 7 patients (2.4%) claimed to have more than one sexual partner and 32 (11.3%) stated to always use condoms. The majority of subjects 274 (93.2%) declared non smoking while alcohol consumption was moderate, and only 2 patients (0.7%) were daily drinkers.

Table 1: Socio-demographic and lifestyle characteristics

Variable	N. (%)
Number of cases examined	301
Gender	
Female	98 (32.6%)
Male	203 (67.4%)
Age	
Mean age (years)	36.7
Range (years)	18 – 83
Educational degree	
None	141 (47.5%)
Primary school	125 (42.1%)
Secondary school	25 (8.4%)
University degree	6 (2.0%)
Occupation	
Unemployed	95 (32%)
Housewife	18 (6%)
Student	27 (9.1%)
Worker	157 (52.9%)
Monthly income (Euro)	
None	106 (40.8%)
< 25	29 (11.2%)
25- 50	53 (20.4%)
50 – 100	62 (23.8%)
> 100	10 (3.8%)
BMI	
Severe thinness	59 (19.6%)
Moderate thinness	32 (10.6%)
Light Thinness	78 (26%)
Healthy	128 (42.5%)
Overweight	4 (1.3%)
Sexual partner/s	
No	110 (37.4%)
One	177 (60.2%)
More than one	7 (2.4%)
Condom Use	
Never	131 (46.1%)
Sometimes	121 (42.6%)
Always	32 (11.3%)
Smoking	
No	274 (93.2%)
1-5/day	15 (5.1%)
20/day	5 (1.7%)
Alcohol drinking	
No	260 (87.8%)
Once a week	34 (11.5%)
Every day	2 (0.7%)

Clinical and laboratory characteristics

The main laboratory and clinical characteristics are shown in Table 2. The most sensitive test for TB diagnosis was GeneXpert, which was positive in 56 of 58 (96.6%). A chest X-ray was performed for 155/301 subjects and 148 of 155 (95.5%) showed TB-compatible lesions.

The HIV status was known for 282 patients: of them, 161 (57.1%) were positive. Sixty-six patients (26.7%) were on anti-retroviral treatment (ART). The BMI was below healthy range in 169 (55.6%) whereas 4 subjects (1.3%) were overweight. One individual (0.3%) had hypertension.

Table 2 Clinical and laboratorial characteristics

Variable	N. (%)
Sputum Examination	
Negative	50 (16.9%)
Positive	246 (83.1%)
Sputum culture	
Negative	3 (75%)
Positive	1 (25%)
GeneXpert	
Negative	2 (3.4%)
Positive	56 (96.6%)
Chest X-ray	
Negative	7 (4.5%)
Positive	148 (95.5%)
HIV	
Negative	121 (42.9%)
Positive	161 (57.1%)
ART	
Yes	66 (26.7%)
No	181 (73.3%)
Diabetes	
Yes	3 (1%)
No	290 (97%)
IGT	6 (2%)
High Blood Pressure	
Yes	1 (0.3%)
No	293 (99.7%)

Diabetes was diagnosed in only 3 patients (1%) and IGT in additional 6 subjects (2%). Nobody was aware of their diabetic status.

Reported clinical signs and symptoms are summarized in

Table 3. The most frequent TB symptoms were: cough (90.9%), weight loss (88.2%), asthenia (58.8%) and fever (58.8%). The patients with DM and IGT reported weight loss, polyuria and asthenia.

Table 3 Main symptoms reported from patients with pulmonary TB

Symptoms	N. (%)
No symptoms	3 (1%)
Cough	269 (90.9%)
Fever	174 (58.8%)
Asthenia	215 (72.6%)
Dyspnea	95 (32.1%)
Hemoptysis	4 (1.4%)
Night sweats	118 (39.9%)
Weight loss	261 (88.2%)
Polyuria	13 (4.4%)
Polydipsia	7 (2.4%)
Polyphagia	11 (3.7%)

Discussion

In sub-Saharan Africa, there is an emerging strong correlation between communicable and non-communicable diseases due to the long lasting presence of epidemic infections such as HIV and TB and the increasing incidence of chronic diseases as DM and hypertension. In 2030, people living with diabetes in Africa are estimated to rise from 21.1 million in 2010 to up to 23.9 million¹⁵. In the same area, DM causes about 4.9 million deaths per year and approximately 76% of these deaths occur in people aged less than 60 years⁹. Diabetes is considered among priorities in the National Strategic Plan for the prevention and control of non communicable diseases also in Mozambique¹³. However, the prevalence of DM seems to be quite low, around 3% in the general population aged from 25 to 64 years old, and it is higher among people living in rural areas¹¹. Our study showed a low prevalence of DM in newly TB-diagnosed patients (1%) compared to data regarding the general population, although a disorder in glycometabolic control was found in 3% of subjects.

This finding in our population could be explained by the lack of some of the risk factors recognised for the onset of DM, including overweight, age over 45 years, hypertension and smoking¹⁴. In fact, in our population the mean age was 36.7 with only few patients overweight and affected by hypertension. Indeed, two out of three diabetic patients in our study were older than 45 years and

were smokers, whereby one subject was HIV positive. Among the six patients with IGT the median age was 32, there was no smoking, three were HIV positive and one consumed alcohol.

In low-income countries, the association between TB and DM is variable, ranging from 1.9% to 35%¹⁴. Other authors reported a higher incidence of DM in TB patients from African countries or low income settings¹⁰⁻¹⁶. On the contrary, other studies showed a low association of DM and TB in low-income settings¹⁷, which was explained by the young age of enrolled patients, similarly to the results of the present study¹⁸⁻¹⁹.

Furthermore, this study underlines the importance of other factors associated with the onset of TB, including SDH. The phenotype of newly diagnosed patients with pulmonary TB in Beira was outlined: patients are mainly young, underweight, with a low educational background and low income, without smoke nor alcohol habits, basically without hypertension and diabetes mellitus but in most cases living with HIV infection. According to our data, SDH appear to influence the onset of TB but also the frequency of MDR and the adherence to therapy³. HIV is the most well known disease associated with TB; and our data confirms this correlation as almost 60% of patients with a known HIV status were seropositive. On the other hand, additional risk factors for TB such as smoking and alcohol use, are almost inexistent in our

study. A limitation of our study is that, our data could be biased because they were not verified and were based on personal interview. Another drawback of the study is the small sample size and lack of a control group.

Conclusion

Our study shows a low prevalence of DM in newly diagnosed patients with pulmonary TB in Mozambique. The lower than expected DM prevalence, however, does not clash with the hypothesis of a mutual interaction existing between these two diseases. Future studies enrolling a greater number of diabetic patients would be crucial in order to assess definitive epidemiological data to contribute to the TB End Strategy in one of the most affected countries like Mozambique.

Conflict of interest

All authors have no conflict of interest

References

1. WHO. Global Tuberculosis Report 2016 <http://apps.who.int/iris/bitstream/10665/250441/1/9789241565394-eng.pdf?ua=1> (Accessed December 2016)
2. Rasanathan K, Sivasankara Kurup, Jaramillo E, Lönnroth K. The social determinants of health: key to global tuberculosis control. *Int J Tuberc Lung Dis* 2011; 15 Suppl 2: S30–6.
3. Di Gennaro F et al Social determinants of therapy failure and multi drug resistance among people with tuberculosis: A review Tuberculosis • March 2017
4. Kwan CK, Ernst JD. HIV and tuberculosis: a deadly human syndemic. *Clin Microbiol Rev.* 24(2):351–76.
5. Faurholt-Jepsen D, et al (2012) The role of diabetes co-morbidity for tuberculosis treatment outcomes: a prospective cohort study from Mwanza, Tanzania. *BMC Infect Dis*, 27;12:165
6. Geerlings SE et al (1999) Immune dysfunction in patients with diabetes mellitus (DM). *FEMS Immunology & Medical Microbiology*, 26, no. 3-4, pp. 259-65
7. Hong JY, Kim SY, Chung KS, Kim EY, Jung JY, Park MS, Kim YS, Kim SK, Chang J, Kang YA. (2014) Association between vitamin D deficiency and tuberculosis in a Korean population. *Int J Tuberc Lung Dis*, 18(1):73-8
8. Pizzol D, Di Gennaro F, Chhaganlal KD, Fabrizio C, Monno L, Putoto G, Saracino A. Tuberculosis and diabetes: current state and future perspectives. *Trop Med Int Health.* 2016 Jun;21(6):694-702.

9. Levitt NS. (2008) Diabetes in Africa: epidemiology, management and healthcare challenges. *Heart* 94 (11): 376-382
10. Workneh MH Bjune GA, Yimer SA. Prevalence and Associated Factors of Diabetes Mellitus among Tuberculosis Patients in South-Eastern Amhara Region, Ethiopia: A Cross Sectional Study. *PLoS One.* 2016 Jan 25;11(1):e0147621
11. Silva-Matos C, Gomes A, Azevedo A, Damasceno A, Prista A, Lunet N. Diabetes in Mozambique: prevalence, management and healthcare challenges. *Diabetes Metab.* 2011 Jun;37(3):237-44.
12. INSIDA 2010, Instituto Nacional de Saúde (INS). Instituto Nacional de Estatística (INE). ICF MACRO. Inquérito Nacional de Prevalência, Riscos Comportamentais e Informação sobre o HIV e SIDA em Moçambique 2009. Calverton, Maryland, EUA: INS, INE e ICF Macro. <http://www.measuredhs.com/pubs/pdf/AIS8/AIS8.pdf>. (Accessed December 2016)
13. MISAU: National Strategic Plan for the Prevention and Control of Non Communicable Diseases: 2008–2014. Maputo: Ministério da Saúde; 2008
14. American Diabetes Association. (2015) Classification and Diagnosis of Diabetes Diabetes Care; 38(Suppl. 1):S8–S16
15. Victoria Hall, Reimar W Thomsen, Ole Henriksen and Nicolai Lohse. Diabetes in Sub Saharan Africa 1999–2011: Epidemiology and public health implications. a systematic review. *BMC Public Health* 2011, 11:564
16. Lee EH, Lee JM, Kang YA, Kim EY et al. Prevalence and Impact of Diabetes Mellitus Among Patients with Active Pulmonary Tuberculosis in South Korea. *Lung* 2017 Feb 9 lung).
17. M Senkoro, A M V Kumar, P Chinnakali, S G Mfinanga, S Egwaga, V Kamara, F van Leth, SG Hinderaker. Population impact of factors associated with prevalent pulmonary tuberculosis in Tanzania *INT J TUBERC LUNG DIS* 2016 20(10):1326–1333)
18. Thorny L Haraldsdottira, b, Frauke Rudolfa, Morten Bjerregaard-Andersena, Luis Carlos Joaquínma, Kirstine Stochholmf, Victor F. Gomesa, Henning Beck-Nielsend, Lars Ostergaardb, Peter Aabya, and Christian Wejsea. Diabetes mellitus prevalence in tuberculosis patients and the background population in Guinea-Bissau: a disease burden study from the capital Bissau. *Trans R Soc Trop Med Hyg* 2015; 109: 400–407)
19. S Ade, D Affolabi, G Agodokpessi, P Wachinou, F

Faihun, N Toundoh, W Békou, A Makpenon, G Ade, S Anagonou, A D Harries. Low prevalence of diabetes mellitus in patients with tuberculosis in Cotonou, Benin, or if was self-reported, were misclassified with underestimation of this association. *Public Health Action* Vol 5 n2 June 2015

20. Harries AD, Kumar AM, Satyanarayana S, Lin Y, Zachariah R, Lönnroth K, Kapur A. Addressing diabetes mellitus as part of the strategy for ending TB. *Trans R Soc Trop Med Hyg.* 2016 Mar;110(3):173-9.