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Abstract

Objective: To determine the association between food insecurity and HIV infection with depression and anxiety among new tuberculosis (TB) patients.

Design: Our cross-sectional study assessed depression, anxiety and food insecurity with Patient Health Questionnaire (PHQ-9), Zung Anxiety Self-Assessment Scale (ZUNG) and Household Food Insecurity Access Scale, respectively. Poisson regression models with robust variance were used to examine correlates of depression (PHQ-9 ≥ 10) and anxiety (ZUNG ≥ 36).

Setting: Gaborone, Botswana.

Participants: Patients who were newly diagnosed with TB.

Results: Between January and December 2019, we enrolled 180 TB patients from primary health clinics in Botswana. Overall, 99 (55·0%) were HIV positive, 47 (26·1%), 85 (47·2%) and 69 (38·5%) indicated depression, anxiety and moderate to severe food insecurity, respectively. After adjusting for potential confounders, food insecurity was associated with a higher prevalence of depression (adjusted prevalence ratio (aPR) = 2·30; 95% CI 1·40, 3·78) and anxiety (aPR = 1·41; 95% CI 1·05, 1·91). Prevalence of depression and anxiety was similar between HIV-infected and HIV-uninfected participants. Estimates remained comparable when restricted to HIV-infected participants.

Conclusions: Mental disorders may be affected by food insecurity among new TB patients, regardless of HIV status.

Keywords
HIV infection
HIV/tuberculosis co-morbidity
Mental illness
Common mental disorders
Epidemic

Mental illness is a leading cause of disability, with approximately 970 million people being affected worldwide⁽¹⁾. Compared with the general population, individuals affected by tuberculosis (TB) are significantly more likely to have poor mental health due to biological and social complexities⁽²⁾. Common mental illness, including depression and anxiety disorders, imposes immense costs on healthcare systems and poor quality of life on patients and can be important barriers for TB diagnosis and treatment. Moreover, depression may lead to medication non-adherence resulting in unfavourable TB treatment outcomes and ongoing community transmission⁽³⁾, which has been cited as the fundamental obstacle in effective TB control⁽⁴⁾. Therefore, mental health is an integral part

of achieving WHO's The End TB Strategy and eliminating the global TB epidemic⁽⁵⁾. Unfortunately, the majority of TB and mental illness occur in low- and middle-income countries⁽⁶⁾, where mental health is often overlooked due to a myriad of challenges such as lack of mental health specialists, limited treatment options, perceived stigma and other competing public health priorities⁽⁶⁾.

Meanwhile, individuals with TB often experience malnutrition⁽⁷⁾, which may be exacerbated by food insecurity. Globally, about 2 billion people suffer from moderate to severe levels of food insecurity⁽⁸⁾. According to The United States Department of Agriculture, a household is defined as food insecure if at least one member experiences insufficient or inconsistent access to nutritionally

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adequate and safe food⁽⁹⁾. A survey of patients with TB/HIV co-infection in multiple African cities reported that only 10 % TB patients feel they are food secure⁽¹⁰⁾. Food insecurity not only negatively affects general physical health but also has important ramifications for adverse mental health. For instance, studies have shown that adults with food insecurity are more likely to have anxiety disorder, depression and suicidal ideation compared with those who are food secure⁽¹¹⁾.

Globally, Botswana has one of the highest TB incidence - 300 cases per 100 000 population in 2017⁽¹²⁾. About 60 % of all TB patients also have co-morbid HIV infection⁽¹²⁾, which has been suggested as a risk factor in psychiatric populations^(13,14). People living with HIV/AIDS may have an increased risk of developing mood disorders and clinical depressive symptoms triggered by distress associated with initial HIV diagnosis, disease progression, difficult life circumstances, hospitalisation, HIV-induced brain injury and adverse effects of antiretroviral medications⁽¹⁴⁾. The most recent data indicate that psychiatric morbidity in the country contributes an estimated 4.6% of the global burden of disease⁽¹⁵⁾. There are 0.5 mental health outpatient facilities, 2.5 psychiatrists and 15.2 psychologists available per million population in spite of the heavy burden of psychological distress⁽¹⁵⁾.

For the past few decades, Botswana has been experiencing rapid urbanisation and positive economic growth with a significant decline in poverty incidence⁽¹⁶⁾. At the same time, many challenges remain, including food insecurity. The Food and Agricultural Organisation (FAO) indicated that the average prevalence of severe food insecurity was 41.3 % between 2016 and 2018⁽¹⁷⁾. It is crucial to understand the relationship between food insecurity and mental health among the socially vulnerable TB patients because, while the expansion and integration of mental health care can be a challenging and gradual task in resource-limited settings, social support programmes, particularly food supplementation, may be relatively low intensity and straightforward for policy delivery and implementation. The objectives of our study were to examine the prevalence of depression and anxiety among newly diagnosed TB patients in Botswana and to explore the association between those conditions with food insecurity and HIV co-infection.

Methods

Study design, population and eligibility

We conducted a cross-sectional study and recruited participants with newly diagnosed pulmonary or extrapulmonary TB from primary healthcare clinics in Gaborone, Botswana. Participants were deemed eligible if they were at least 18 years old with a verified TB diagnosis and were recruited by our research staff at the time of receiving TB diagnosis. According to national guidelines during the study period, the first-line diagnostic test for TB was sputum smear microscopy or GeneXpert MTB/RIF for patients

who experienced prolonged general pulmonary TB symptoms such as cough, fever, shortness of breath, night sweats, weight loss and haemoptysis⁽¹⁸⁾. Patients with indications for culture testing, such as symptomatic individuals at high risk of multidrug-resistant TB, also underwent examination by mycobacterial culture and drug susceptibility testing with the Mycobacteria Growth Indicator Tube method. A chest radiography was performed in symptomatic individuals who had negative sputum smears or negative Xpert MTB/RIF. For patients who had negative results in all testing, TB diagnosis was made by a physician based on clinical presentations and/or lymph node examinations. A rapid HIV test was conducted among participants who have not had an HIV test done before, or for those who have had a negative result in previous testing.

Data collection

Socio-demographic data were collected via face-to-face interview and included age, gender, education, monthly income, marital status, smoking and alcohol history. We also collected clinical information that included TB symptoms, duration of the symptoms, HIV test result and HIV diagnosis date if any, antiretroviral treatment (ART) history, and results of smear microscopy, Xpert MTB/RIF, mycobacterial culture and chest X-ray. CD4 T-cell data closest to the time of enrolment were extracted and integrated from the national HIV database for HIV-infected participants.

We administered standardised questionnaires to assess the presence and severity of depressive symptoms, anxiety symptoms and household food insecurity of enrolled participants. For simplicity, we will use depression and anxiety to describe depressive symptoms and anxiety symptoms, respectively, hereinafter.

Depression was assessed by the Patient Health Questionnaire (PHQ-9), a nine-item self-report tool that screened depression as participants identified how much each item statement applied to them within the past 2 weeks⁽¹⁹⁾. Examples of item statements include 'feeling down, depressed or hopeless' and 'little interest or pleasure in doing things'. Answer choices were scored on a Likert-type scale that ranged from 'not at all', 'several days', 'more than half the days', to 'nearly every day'. PHQ-9 has been validated in multiple sub-Saharan African countries to assess depression, including Botswana⁽²⁰⁾.

Anxiety was assessed by Zung Self-Rating Anxiety Scale (ZUNG), a twenty-item self-report scale that measured anxiety levels⁽²¹⁾. Similar to PHQ-9, participants indicated how much each of the twenty statements applied to them within the past several days and replies included 'none or a little of the time', 'some of the time', 'good part of the time' and 'most or all of the time'. Examples of item statements include 'I feel afraid for no reason at all' and 'I get upset easily or feel panicky'.

Finally, food insecurity was evaluated by the Household Food Insecurity Access Scale, a nine-item questionnaire



Food access and mental health in tuberculosis patients

widely used across different cultures, including Botswana⁽¹⁶⁾, to distinguish food-insecure households and to determine the degree of food insecurity⁽²²⁾. Examples on this questionnaire include 'In the past four weeks, did you worry that your home would not have enough food?' and 'In the past four weeks, did you or anyone in the house go to sleep at night hungry because there was not enough food?' Answer choices were scored on a Likert-type scale that ranged from 'never', 'rarely (1 or 2 times)', 'sometimes (3–10 times)', to 'often (more than 10 times)'.

All questionnaires were prepared in English and Setswana, both official languages of Botswana. Study interview was conducted in the preferred language of the participant. We restate that data collection tools (i.e. PHQ-9 and ZUNG) used in this study were suggestive of symptoms of depression and anxiety, and not used for clinical diagnoses. All participants with PHQ-9 scores indicative of depression and suicidal ideation were provided with appropriate psychiatric referrals.

Statistical analysis

The outcomes of interest for this study were the prevalence of depression and anxiety among newly diagnosed TB patients, and the roles of food insecurity and HIV status on (a) depression, and (b) anxiety. Patients were categorised as having depression if they scored 10 points or more on PHQ-9⁽¹⁹⁾, and having anxiety if they scored 36 points (raw score) or more on ZUNG⁽²¹⁾. Prevalence estimates of depression and anxiety were calculated as the number of observed participants with depression and anxiety, respectively, divided by the total number of enrolled participants, and the 95% confidence limits were obtained by the Wilson score method. Based on Household Food Insecurity Access Scale scores, participants were categorised as 'food secure', 'mildly insecure', 'moderately insecure' or 'severely insecure' households (22). For the purpose of analysis, we dichotomised participants as experiencing 'food security or mild insecurity' or 'moderate to severe insecurity'(23). Monthly income was categorised as having 'no income' or 'some income'. As recommended for estimating prevalence ratios in cross-sectional studies where the outcome of interest was common, Poisson regression models with robust variance were used to examine the bivariate associations between selected covariates and each of the mental disorder outcomes (24). Independent covariates were selected based on a priori knowledge of predictors of adverse mental health, which include age, gender, HIV status, monthly income, education, marital status and food insecurity. We excluded education and marital status in the multivariable model due to the relatively small overall sample size of our study. Subsequently, we analysed the bivariate associations between age, gender, monthly income, ART status, CD4 T-cell count, food insecurity and each outcome among participants infected with HIV. Similarly, multivariable model was constructed to determine the independent effect of each covariate, regardless of their bivariate associations.

However, because this is a smaller subset of participants, we present the model including only food insecurity, ART status and CD4 T-cell count in this paper. Estimates were similar between the smaller model and the full model.

Data analysis was carried out in SAS version 9.4 (SAS Institute). In accordance with recent guidelines, no alpha cut-off was specified for statistical significance⁽²⁵⁾.

Results

Between January and December 2019, 180 participants were recruited into the study. Overall, 99 (55·0%) were HIV positive, 64 (35·6%) were female, 155 (86·1%) were single, 137 (76·1%) were diagnosed with pulmonary TB, 17 (9·4%) had no formal education and 65 (36·1%) had no income (Table 1). Among those who were HIV co-infected, the median duration since HIV diagnosis was 43 months, median CD4 T-cell count was 335 and 31 (31·3%) had never taken ART, of whom 90% were newly diagnosed with HIV at or within one month of study enrolment. Over half of all participants reported being food secure, and 15 (8·4%), 17 (9·5%) and 52 (29·0%) reported experiencing mild insecurity, moderate insecurity and severe insecurity, respectively (Table 1).

The overall prevalence of depression, anxiety and co-existence of both conditions was $26\cdot1\%$, $47\cdot2\%$ and $20\cdot6\%$, respectively. Among those who were depressed, $26(14\cdot4\%)$, $14(7\cdot8\%)$ and $7(3\cdot9\%)$ had moderate, moderately severe and severe depression, respectively (Table 1). Twenty-seven (15·1%) participants had PHQ-9 scores indicative of suicidal ideation. Additionally, 71 (39·4%), $11(6\cdot1\%)$ and $3(1\cdot7\%)$ had minimal to moderate, marked to severe and most extreme anxiety, respectively.

Table 2 shows the results of Poisson regression analysis of covariates associated with depression. In bivariate analysis, moderate to severe food insecurity was associated with a higher prevalence of depression (crude prevalence ratio (PR) = 2.37; 95 % CI 1.44, 3.91). After adjusting for age, gender, monthly income and HIV co-infection status, the association between moderate to severe food insecurity and depression remained similar (adjusted PR = 2.30; 95 % CI 1.40, 3.78).

Similar findings were observed for the outcome of anxiety; moderate to severe food insecurity was associated with a higher prevalence of anxiety (crude PR = 1.43; 95 % CI 1.06, 1.93; Table 3). The association remained robust in multivariable Poisson analysis after adjusting for age, gender, monthly income and HIV co-infection status (adjusted PR = 1.41; 95 % CI 1.05, 1.91; Table 3).

Among HIV-co-infected participants, the estimates were similar in the bivariate analysis for depression (crude PR = 2.41; 95 % CI 1.28, 4.55; Table 4) and anxiety (crude PR = 1.54; 95 % CI 1.04, 2.27). Accounting for ART status and CD4 count closest to the time of TB diagnosis, food insecurity continued to be associated with increased





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Table 1 Characteristics of study participants in Botswana, 2019 (n 180)

	HIV	positive	HIV negative		
Characteristics	n	%	n	%	
Gender					
Male	62	62⋅6	54	66.7	
Female	37	37⋅4	27	33.3	
Age Median IQR		41 1–47		29 –39	
Marital status	3	1-47	20	-39	
Married	9	9⋅1	9	11.1	
Single	86	86⋅9	69	85.2	
Divorced/separated/widow	4	4.0	3	3.7	
Education None	11	11.1	6	7.4	
Secondary or less	74	74·7	49	60·5	
Tertiary or certificate	7	7·1	12	14.8	
Diploma or degree	7	7.1	13	16.1	
Masters or PhD	0		1	1.2	
Income (in Botswana pula*)	0.5	05.4	00	07.0	
No income <5000	35 53	35⋅4 53⋅5	30 45	37·0 55·6	
<5000 5000–10 000	9	9·1	45 5	6.2	
>10 000	2	2.0	1	1.2	
TB diagnosis methods					
Culture	0		2	2.5	
XPERT	34	34.3	38	46.9	
AFB	8	8.1	15	18.5	
Chest X-ray Clinical	27 30	27·3 30·3	11 15	13·6 18·5	
TB symptoms length	30	30.3	15	10.5	
None	6	6.1	0		
<1 month	47	47.5	42	51.8	
1–2 months	19	19⋅2	22	27.2	
2–3 months	12	12.1	8	9.9	
>3 months	15	15⋅1	9	11.1	
Smoking No	75	75.8	60	74.1	
Yes	24	24.2	21	25.9	
Alcohol					
No	57	57⋅6	44	54.3	
Yes	42	42.4	37	45.7	
Disease classification†	74	70.5	00	01.5	
Pulmonary Extrapulmonary	71 27	72⋅5 27⋅5	66 15	81·5 18·5	
Time since HIV diagnosis (in months)‡	21	21.5	13	10.5	
Median		43	N	I/A	
IQR	1-	-119			
CD4 T-cell count at time closest to TB diagnosis (cells/mm³)§			_		
Median		335	N	I/A	
IQR ART history	120	0–469			
Never taken ART	31	31.3	N/A		
Taking ART	59	59.6	14//		
Took ART but stopped	9	9⋅1			
Depression					
No depression (0)	3	3.0	5	6.2	
Minimal depression (1–4)	27	27·3	20	24.7	
Mild depression (5–9) Moderate depression (10–14)	40 15	40·4 15·2	38 11	46·9 13·6	
Moderately severe depression (15–19)	11	11.1	3	3.7	
Severe depression (20–27)	3	3.0	4	4.9	
Anxiety					
Normal (20–35)	50	50.5	45	55.6	
Minimal to moderate anxiety (36–47)	39	39·4	32	39.5	
Marked to severe anxiety (48–59) Most extreme anxiety (≥60)	7 3	7·1 3·0	4 0	4.9	
Depression–Anxiety (≥60)	S	3.0	U		
No depression, no anxiety	44	44.4	41	50.6	
INO depression, no anxiety	77				



Table 1 Continued

	HIV	HIV n	HIV negative	
Characteristics	n	%	n	%
Anxiety, no depression	26	26.3	22	27.2
Depression and anxiety	23	23.2	14	17.3
Household food insecurity				
Secure	51	51.5	45	55.6
Mildly insecure	8	8.1	7	8.6
Moderately insecure	11	11.1	6	7.4
Severely insecure	29	29.3	23	28.4

IQR, interquartile range; TB, tuberculosis; XPERT, GeneXpert MTB/RIF; AFB, acid-fast Bacilli smear; ART, antiretroviral therapy.

Table 2 Correlates of depression* among study participants in Botswana, 2019 (n 180)

	Preval	ence				
Variables	n/N	%	Crude PR	95 % CI	Adjusted PR†	95 % CI
Age	N/A		1.01	0.99, 1.03	1.01	0.99, 1.02
Gender						
Male	26/116	22.4	1.00		1.00	
Female	21/64	32.8	1.46	0.90, 2.38	1.41	0.88, 2.27
HIV						
Negative	18/81	22.2	1.00		1.00	
Positive	29/99	29.3	1.32	0.79, 2.19	1.20	0.73, 1.98
Income						
No	19/65	29.2	1.00		1.00	
Yes	28/115	24.4	0.83	0.51, 1.37	0.95	0.59, 1.52
Education						
No	5/17	29.4	1.00		Not included	
Yes	42/163	25.8	0.88	0.40, 1.91		
Marital status				·		
Single/divorced or separated/widow	3/18	16.7	1.00		Not included	
Married	44/162	27.2	0.61	0.21, 1.78		
Food insecurity						
None or mildly insecure	19/111	17.1	1.00		1.00	
Moderate to severe insecure	28/69	40-6	2.37	1.44, 3.91	2.30	1.40, 3.78

PR, prevalence ratio

symptoms of depression (adjusted PR = 2.33; 95 % CI 1.24, 4.38) and anxiety (adjusted PR = 1.53; 95 % CI 1.03, 2.26) in the multivariable model.

Discussion

In our population of TB patients, we showed that household food insecurity was the most important factor associated with symptoms of depression and anxiety, independent of HIV co-infection status, age, monthly income and gender. Our findings confirmed previous research that showed a linkage between food insecurity and poor mental health^(26,27) and extended this finding in TB patients. Several reasons may explain this association. First, as a consequence of food insecurity, deficiency in critical micronutrient could lead to depression and anxiety via biological pathways⁽²⁸⁾. Additionally, food insecurity encompasses not only

hunger and under-nutrition but also uncertainty and distress over the access of food. Thus, food insecurity may compromise mental health by generating instability and unpredictability into lives of the already socio-economically vulnerable TB population⁽²⁶⁾. This pathway could be independent of the degree of social and economic deprivation, as a global analysis of 149 countries concluded an association between food insecurity and mental illness in a dose-response fashion, after controlling for socio-economic status⁽²⁷⁾.

To our knowledge, this is the first study that examined the association between food insecurity and mental illness among new TB patients in a high burden setting. In compliance with The End TB Strategy, Botswana has social (e.g. food support) programmes to help TB-affected individuals to improve nutritional status and treatment adherence and alleviate the steep costs associated with the disease and treatment. Specifically, TB patients with



^{*5000} Botswana pula is approximately 465 US dollars.

 $[\]dagger n = 1$ missing data on disease classification.

 $[\]ddagger n = 2$ missing data on HIV diagnosis date.

 $[\]S{n} = 14$ missing data on CD4 T-cell count.

^{*}The presence of depression is defined as scoring equal to or more than 10 points on the Patient Health Questionnaire-9.

[†]The multivariable Poisson regression analysis adjusted for age, gender, HIV status, income and food insecurity.



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Table 3 Correlates of anxiety* among study participants in Botswana, 2019 (n 180)

Variables	Prevalence					
	n/N	%	Crude PR	95 % CI	Adjusted PR [†]	95 % CI
Age	N/A		1.01	0.99, 1.02	1.01	0.99, 1.02
Gender						
Male	52/116	44.8	1.00		1.00	
Female	33/64	51.6	1.15	0.84, 1.57	1.14	0.84, 1.56
HIV						
Negative	36/81	44.4	1.00		1.00	
Positive	49/99	49.5	1.11	0.81, 1.53	1.05	0.77, 1.42
Income						
No	32/65	49.2	1.00		1.00	
Yes	53/115	46.1	0.94	0.69, 1.30	0.98	0.72, 1.34
Education						
No	8/17	47.1	1.00		Not included	
Yes	77/163	47.2	1.00	0.59, 1.70		
Marital status						
Single/divorced or separated/widow	8/18	44.4	1.00		Not included	
Married	77/162	47.5	0.94	0.54, 1.61		
Food insecurity						
None or mildly insecure	45/111	40.5	1.00		1.00	
Moderate to severe insecure	40/69	58.0	1.43	1.06, 1.93	1.41	1.05, 1.91

PR, prevalence ratio.

Table 4 Correlates of depression* and anxiety† among HIV-infected participants in Botswana, 2019 (n 99)

	Depression				Anxiety			
Variables	Crude PR	95 % CI	Adjusted PR	95 % CI	Crude PR	95 % CI	Adjusted PR	95 % CI
Age	1.01	0.99, 1.04	Not included		1.01	0.99, 1.03	Not included	
Gender								
Male	1.00		Not included		1.00		Not included	
Female	1.56	0.86, 2.86			1.16	0.78, 1.72		
Income								
No	1.00		Not included		1.00		Not included	
Yes	0.67	0.37, 1.23			0.86	0.58, 1.29		
ART								
No	1.00		1.00		1.00		1.00	
Yes	1.29	0.67, 2.47	1.14	0.58, 2.21	1.17	0.77, 1.78	1.12	0.73, 1.73
CD4 T-cell count (cells/mm3)								
≤200	1.00		1.00		1.00		1.00	
>200	1.48	0.70, 3.11	1.26	0.59, 2.72	1.17	0.72, 1.91	1.06	0.64, 1.75
Missing	0.92	0.28, 3.03	0.87	0.26, 2.94	1.32	0.72, 2.43	1.27	0.68, 2.35
Food insecurity								
None or mildly insecure	1.00		1.00		1.00		1.00	
Moderate to severe insecure	2.41	1.28, 4.55	2.33	1.24, 4.38	1.54	1.04, 2.27	1.53	1.03, 2.26

PR, prevalence ratio; ART, antiretroviral therapy.

confirmed diagnosis may receive food allocations that include cooking oil, beans and maize flour with additive vitamins. However, the food rations are usually not enough to support every member of the household. In addition, TB renders many patients unable to work and contribute to household expenditures, and caregivers may still experience household food insecurity as they face challenges of providing for their family due to income loss⁽²⁹⁾. The total financial costs of TB can be catastrophic for patients – up to

58% of annual individual income and 39% of household income in low- and middle-income countries, and income loss constitutes the largest portion of all TB-related cost at 60%⁽²⁹⁾. The cyclical nature of the relationship between food insecurity and TB fosters sustained impoverishment and makes it difficult for household to break the cycle without outside intervention. Malnutrition, a consequence of food insecurity, may predispose patients for risks of TB infection and TB disease reactivation⁽³⁰⁾.

^{*}The presence of anxiety is defined as scoring equal to or more than 36 points (raw score) on the Zung Anxiety Self-Assessment Scale.

[†]The multivariable Poisson regression analysis adjusted for age, gender, HIV status, income and food insecurity.

^{*}The presence of depression is defined as scoring equal to or more than 10 points on the Patient Health Questionnaire-9.

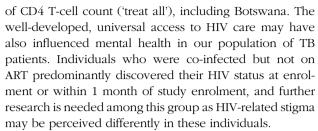
[†]The presence of anxiety is defined as scoring equal to or more than 36 points (raw score) on the Zung Anxiety Self-Assessment Scale.

Comparable with national data released by FAO⁽¹⁷⁾, our survey indicated 38.5 % of the participants had experienced moderate to severe food insecurity prior to receiving their TB diagnosis. Further, a separate survey estimated that only 12 % of households report to be 'food secure' in urban parts of Gaborone⁽¹⁶⁾. These findings highlight the pervasive food-insecure households among TB patients and those at risk for TB in Botswana, which lend support for food assistance programmes and social protection interventions to target populations who are at high risks for TB. Other research suggests members of households who participate in food supplementation programmes have lower rates of psychological distress; by reducing food insecurity, these programmes can be effective in improving mental health and overall security among vulnerable populations⁽³¹⁾.

In this study, we revealed that anxiety and depression were highly prevalent among patients newly diagnosed with TB in Botswana. Approximately 26 % of enrolled TB patients had PHQ-9 scores indicative of moderate to severe depression and 15% indicative of suicidal ideation. Our estimate was comparable to recent studies in several countries that examined depression (moderate to severe) among TB patients, including Ethiopia (17·6–54·0 %)(32–34), South Africa (32.9%)(13), Angola (49.4%)(35) and China (18·1%)(36). They were, however, considerably lower than those found in Ethiopia using PHO-9 by Ambaw et al. (32); these differences may have arisen from variations in study location and population. For example, roughly 45 % of the participants in the Ethiopian study had no formal education and over half were married.

Approximately 46 % of our participants reported anxiety symptoms, comparable with published estimates of other settings: Ethiopia (41.5 %)(34), Angola (38.3 %)(35) and China (18.4%)⁽³⁶⁾. None of those studies utilised the ZUNG scale; instead, Generalized Anxiety Disorder Questionnaire and Hospital Anxiety and Depression Scale were used to assess the presence of anxiety symptoms, which might be an explanation for the discrepant results. Our estimate was much higher than those published by Wang et al. of Chinese TB patients⁽³⁶⁾. One explanation for this discrepancy could be patient characteristics: in the Chinese study, over half of the participants had tertiary or higher education levels; majority were married and patients with extrapulmonary TB were excluded.

We did not find an association between HIV comorbidity and mental disorders in our sample. This finding is in agreement with some studies (32,33), but not others (13,34). Studies that reported an association between co-morbidity and mental disorders have posited HIV-associated stigma as the plausible driving force (13,34). Among people with TB, factors such as malnutrition, poor physical health, socio-economic adversities and interaction between these factors may have a stronger influence on depression and anxiety than HIV-related stigma. Furthermore, following WHO's guidelines in 2015, nearly all sub-Saharan African countries adopted universal HIV care policies irrespective



We note several limitations associated with our research. First, depression and anxiety assessment relied on selfreporting using screening questionnaires. This may lead to self-report bias, particularly under-reporting owing to stigmatisation of psychological disorders. However, we tried to minimise this by training the research staff on administering the questionnaires with consistency and without judgement, and by asking the participants to record their answer on the tablet directly. Second, the PHQ-9 and ZUNG screening tools have not been validated in Botswana, nor in TB patients in Botswana. The English version of these questionnaires was translated by a native speaker of Setswana. Some items on the questionnaires overlap with TB symptoms, including fatigue, loss of appetite and insomnia, which may lead to misclassification of depressive and anxiety cases. As a consequence, our estimate of associations between depression, anxiety and independent variables may be biased towards the null. Third, due to the cross-sectional study design, statistical associations may not be representative of casual relationships between exposure and outcome variables. Finally, confounding by unmeasured variables is a possibility in our study.

Conclusion

In conclusion, we have shown that common mental disorders are highly prevalent and food insecurity is associated with poor mental health among new TB patients. This study highlights the critical need for integration of TB, mental health and social services (i.e. food assistance) to address the TB-mental health co-morbidity in Botswana. Mental illness is increasingly being recognised as an important driver of the TB epidemic and negatively impacts effective TB control⁽²⁾. Individually, poor mental health is associated with elevated morbidity and mortality, and greater risk of drug resistance⁽³⁾. Despite the need, mental health counselling and psychiatric services are severely underfunded in low-resource settings where numerous public health priorities compete⁽⁶⁾. Social assistance programmes that mitigate costs of TB care and improve food access may ameliorate symptoms of common mental disorders. Current programmes intended to support individuals struggling with food security may require additional measures to identify patients who experience moderate to severe food insecurity and ensure their access to and adequacy of nutritious and sufficient food for the household. Given the global goal





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of ending TB and rising interest in mental health, longitudinal research is needed to determine the temporal relationship between food insecurity and mental disorders and to identify potential pathways through which food insecurity may elevate the risk of common mental disorders among TB patients.

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References

- GBD 2017 Disease and Injury Incidence and Prevalence Collaborators (2018) Global, regional, and national incidence, prevalence, and years lived with disability for 354 diseases and injuries for 195 countries and territories, 1990–2017: a systematic analysis for the Global Burden of Disease Study 2017. *Lancet* 392, 1789–1858.
- Sweetland AC, Kritski A, Oquendo MA et al. (2017) Addressing the tuberculosis-depression syndemic to end the tuberculosis epidemic. Int J Tuberc Lung Dis 21, 852–861.
- Pachi A, Bratis D, Moussas G et al. (2013) Psychiatric morbidity and other factors affecting treatment adherence in pulmonary tuberculosis patients. Tuberc Res Treat 2013, 489865.
- Suryanarayana L, Rajalakshmi R, Jaigopal MV et al. (1999)
 Performance of national tuberculosis programme during
 1996—a report. Indian J Tuberc 46, 11–20.
- World Health Organization (2015) Implementing the End TB Strategy: the essentials. https://www.who.int/tb/publications/ 2015/The_Essentials_to_End_TB/en/ (accessed February 2020).
- Rathod S, Pinninti N, Irfan M et al. (2017) Mental health service provision in low- and middle-income countries. Health Serv Insights 10, 1178632917694350.

- Sinha P, Davis J, Saag L et al. (2019) Undernutrition and tuberculosis: public health implications. J Infect Dis 219, 1356–1363
- 8. Food and Agriculture Organization of the United Nations (2019) The state of food security and nutrition in the world 2019. http://www.fao.org/state-of-food-security-nutrition/en/ (accessed February 2020).
- United States Department of Agriculture (2019) Food security in the U.S. https://www.ers.usda.gov/topics/food-nutritionassistance/food-security-in-the-us/(accessed February 2020).
- Caesar M & Crush J (2017) The triple burden of HIV, TB and food insecurity. In *Food and Nutrition Security in Southern African Cities*, pp. 135–155 [B Frayne, J Crush & C McCordic, editors]. London: Taylor & Francis.
- Leung CW, Epel ES, Willett WC et al. (2015) Household food insecurity is positively associated with depression among lowincome supplemental nutrition assistance program participants and income-eligible nonparticipants. J Nutr 145, 622–627.
- 12. World Health Organization (2018) Global tuberculosis report 2018. https://www.who.int/tb/publications/global_report/en/ (accessed February 2020).
- Peltzer K, Naidoo P, Matseke G et al. (2012) Prevalence of psychological distress and associated factors in tuberculosis patients in public primary care clinics in South Africa. BMC Psychiatry 12, 89.
- American Psychiatric Association: Work Group on HIV/AIDS (2000) Practice guideline for the treatment of patients with HIV/AIDS. Am J Psychiatry 157, 1–62.
- World Health Organization (2011) Mental Health Atlas 2011. https://www.who.int/mental_health/evidence/atlas/profiles/ bwa_mh_profile.pdf?ua=1 (accessed February 2020).
- Acquah B, Kapunda S, Legwegoh A et al. (2013) The State of Food Insecurity in Gaborone, Botswana. Botswana: African Food Security Urban Network (AFSUN).
- 17. Food and Agriculture Organization of the Union Nations (2019) FAO statistical yearbook. http://www.fao.org/faostat/en/-country/20 (accessed February 2020).
- Botswana Ministry of Health (2007) The Botswana National Tuberculosis Programme Manual. https://www.who.int/hiv/ pub/guidelines/botswana_tb.pdf (accessed February 2020).
- 19. Kroenke K, Spitzer RL & Williams JB (2001) The PHQ-9: validity of a brief depression severity measure. *J Gen Intern Med* **16**, 606–613.
- 20. Mothhatlhedi K, Setlhare V, Ganiyu A *et al.* (2017) Association between depression in carers and malnutrition in children aged 6 months to 5 years. *Afr J Prim Health Care Fam Med* **9**, e1–e6.
- 21. Zung WW (1971) A rating instrument for anxiety disorders. *Psychosomatics* **12**, 371–379.
- Coates J, Swindale A & Bilinsky P (2007) Household Food Insecurity Access Scale (HFIAS) for Measurement of Household Food Access: Indicator Guide (v.3).
 Washington, D.C.: Food and Nutrition Technical Assistance Project (FANTA).
- 23. Maxwell D, Coates J & Vaitla B (2013) How Do Different Indicators of Household Food Security Compare? Empirical Evidence from Tigray. Medford, USA: Feinstein International Center, Tufts University.
- Zou G (2004) A modified Poisson regression approach to prospective studies with binary data. Am J Epidemiol 159, 702–706.
- Greenland S, Senn SJ, Rothman KJ et al. (2016) Statistical tests, P values, confidence intervals, and power: a guide to misinterpretations. Eur J Epidemiol 31, 337–350.
- Weaver LJ & Hadley C (2009) Moving beyond hunger and nutrition: a systematic review of the evidence linking food insecurity and mental health in developing countries. *Ecol Food Nutr* 48, 263–284.
- 27. Jones AD (2017) Food insecurity and mental health status: a global analysis of 149 countries. *Am J Prev Med* **53**, 264–273.





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- Rao TS, Asha MR, Ramesh BN et al. (2008) Understanding nutrition, depression and mental illnesses. *Indian J Psychiatry* 50, 77–82.
- Tanimura T, Jaramillo E, Weil D et al. (2014) Financial burden for tuberculosis patients in low- and middleincome countries: a systematic review. Eur Respir J 43, 1763–1775.
- Chandrasekaran P, Saravanan N, Bethunaickan R et al. (2017) Malnutrition: modulator of immune responses in tuberculosis. Front Immunol 8, 1316.
- Oddo VM & Mabli J (2015) Association of participation in the supplemental nutrition assistance program and psychological distress. *Am J Public Health* 105, e30–e35.
- Ambaw F, Mayston R, Hanlon C et al. (2017) Burden and presentation of depression among newly diagnosed individuals with TB in primary care settings in Ethiopia. BMC Psychiatry 17, 57.

- Molla A, Mekuriaw B & Kerebih H (2019) Depression and associated factors among patients with tuberculosis in Ethiopia: a cross-sectional study. *Neuropsychiatry Dis Treat* 15, 1887–1893.
- Duko B, Gebeyehu A & Ayano G (2015) Prevalence and correlates of depression and anxiety among patients with tuberculosis at WolaitaSodo University Hospital and Sodo Health Center, WolaitaSodo, South Ethiopia, Cross sectional study. BMC Psychiatry 15, 214.
- Paulo BX & Peixoto B (2016) Emotional distress patients with several types of tuberculosis. A pilot study with patients from the Sanatorium Hospital of Huambo. *Int J Mycobacteriol* 5, Suppl. 1, S58.
- Wang XB, Li XL, Zhang Q et al. (2018) A survey of anxiety and depressive symptoms in pulmonary tuberculosis patients with and without tracheobronchial tuberculosis. Front Psychiatr 9, 308.

