A Cross-sectional Study on the Magnitude of undernutrition in Tuberculosis Patients in the Oromia Region of Ethiopia

Bedru Hussien1
Gobena Ameni2–4

1Department of Public Health, Madda Walabu University, Goba Referral Hospital, Robe Bale, Oromia, Ethiopia; 2Aklilu Lemma Institute of Pathobiology, Addis Ababa University, Addis Ababa, Ethiopia; 3Department of Veterinary Medicine, College of Agriculture and Veterinary Medicine, United Arab Emirates University, Al Ain, United Arab Emirates; 4Department of Health Studies, University of South Africa, Pretoria, South Africa

Purpose: To estimate the magnitude of undernutrition in tuberculosis (TB) patients and evaluate its association with selected sociodemographic and clinical characteristics of the patients.

Patients and Methods: A health facility-based cross-sectional design was used. Four hundred and fifty pulmonary TB (PTB) patients were recruited between September 2017 and August 2018. Data were collected by structured questionnaire and anthropometric measurements. Data were analyzed using SPSS 20. Descriptive statistics was used for the analysis and expression of the data. Regression model was used to determine the association between undernutrition and selected factors.

Results: The magnitude of underweight was 51.6%, (95%CI: 47.15–56.2). Binary logistic regression indicated that previous treatment with anti-TB (crude odds ratio, COR=1.68, 95% CI: 1.08–2.63, \(p<0.023\)), duration of illness greater than two months (COR=2.11, 95%CI: 1.26–3.55, \(p<0.005\)), positive HIV serum status (COR=3.83, 95%CI: 1.63–9.02, \(p<0.002\)) and history of resistance to any anti TB drug (COR=2.76, 95%CI: 1.29–5.91, \(p<0.009\)) were associated with underweight. Multiple logistic regression analysis of the association of the aforementioned variables with undernutrition indicated that HIV positivity (adjusted odds ratio, AOR=0.26, 95%CI: 0.104–0.65, \(p<0.004\)) and resistance to any anti-TB drug (AOR=0.39, 95%CI: 0.173–0.90, \(p<0.026\)) were the associated factors.

Conclusion: A significant proportion of TB patients in the Oromia Region were malnourished. Therefore, nutritional counseling and nutritional supplementation are required for the effective treatment of TB patients in the Region.

Keywords: body mass index, undernutrition, tuberculosis, Oromia Region of Ethiopia

Introduction
The association between TB and undernutrition has long been known. TB and undernutrition interrelate in such a way that TB negatively affects the nutritional status of somebody and undernutrition exposes someone to active TB.1–3 Most individuals with active TB are in a consumed state worsening undernutrition and making them weak in defending against infections or allowing them to sustain easily the progression of subclinical cases to clinically apparent cases like latent TB to active TB.4,5 The patient may have poor appetite to eat perhaps as a result of fatigue or may not get enough to eat if they have few resources.4,6 It has also been found that malnourished TB patients have poor bioavailability (low proportion of a drug that can reach its target) of essential drugs like rifampicin. This can
contribute to treatment failure and development of multidrug resistance. There could also be higher risk of hepatotoxicity (which is a major side effect of TB therapy), poor adherence to medical treatment, delayed recovery and hence increased transmission rate, and higher mortality rates than well-nourished patients.\textsuperscript{7–9}

Food and nutritional support as an incentive and enabler are employed by national TB control programs worldwide to make a difference in adherence to anti-TB drug treatment and sputum conversion period, cure and treatment completion rates, gain in body weight and body composition as well as performance status, or to modify associated risk factors for a better TB control program.\textsuperscript{10–12} Closer nutritional monitoring and earlier initiation of nutrition support (before the first two months of treatment are completed) should be considered particularly if the nutritional indicator is approaching the cutoff value for a diagnosis of severe-acute malnutrition.\textsuperscript{13}

For appropriate intervention regarding undernutrition in TB patients for effective control program it is important to have updated information on the magnitude of the problem as well as associated factors. Although previous studies have shown the association between TB and undernutrition, additional studies can help in generating scientific information on the extent of undernutrition in TB patients in different geographic settings so that appropriate correction measures are taken in order to improve the outcome of TB treatment. The Oromia Region is one of the regional states of Ethiopia with the largest human population. Furthermore, similar to the other regions of the country, TB is prevalent in the Oromia Region. But little information is available on the extent of undernutrition in TB patients in the Oromia Region. The present study was formulated with the objective of investigating the magnitude of undernutrition in TB patients and associated risk factors in the Oromia Region and it forms part of a thesis that was done through the University of South Africa.

**Materials and Methods**

**Study Design and Setting**

A health facility (HF)-based cross-sectional design was used to conduct this study in the Oromia Region. This region is the largest state of the Federal Republic of Ethiopia with a population of about 40 million. With regard to health facilities, the region has a total of 108 hospitals, 1405 health centers and 7090 health posts. The health coverage of the region in terms of health centers and health posts were 93% and 94% in 2019, respectively.\textsuperscript{14} Data were collected during the period between September 2017 and August 2018.

**Study Population and Subjects**

Pulmonary TB (PTB) patients who are permanent residents of the Oromia Region, and individuals visiting health centers and hospitals located in the selected zones for ill health and were not on drug treatment for the current episode of TB were the study population.

**Sampling and Sample Size**

For sampling, the study zones and particular HFs were selected on the basis of convenience in terms of geographical accessibility and logistic capacity and patients were enrolled according to their presence for clinical consultation until the proportional numbers of people are obtained from each HF to totally recruit the estimated sample size of 450. The criteria for recruitment of the study participates were, being a case of PTB diagnosed by a clinician according to the National TB and Leprosy Control Program Guideline\textsuperscript{15} and a permanent resident of the region.

**Data Collection Tool and Process**

A structured questionnaire was prepared in English and administered by the interviewer for data collection. Data were collected on sociodemographic characteristics and clinical history of the patient. The questionnaires were pretested on 25 TB patients outside the catchment areas of selected HFs and not included in the analysis. Modifications were made as necessary.

After informed consent, questionnaires were administered by trained data collectors in local languages. The data collectors were skilled laboratory technicians, nurses, and public health workers who can fluently speak, read and write local languages, Afan Oromo and Amharic.

Anthropometric measurements were taken. Weights were recorded using regularly calibrated beam balance (±100 g precision), with the patient wearing light clothing. Heights were recorded to the nearest centimeter with a stadiometer using standard procedures. Body mass index (BMI) was calculated as weight in kilograms divided by squared height in meters (Wt (kg)/Ht (m\(^2\))). For those who were under the age of 18 years, the BMI-for-age percentiles were used for assessing nutritional status of boys and girls differently from BMI of adults.
Statistical Analysis
The statistical analysis was performed using SPSS version 20. Descriptive statistics were used to depict the socio-demographic characteristics, clinical history, and nutritional status. The heights and weights measured were used to calculate the BMI (BMI=weight in kg/height in m²) at registration. A BMI of ≥18.5 and ≤24.99 kg/m² was classified as normal, and undernutrition as BMI <18.5 kg/m², overweight as BMI ≥25 kg/m². Different classes of malnutrition were also defined as follows: mild malnutrition, BMI=17.0–18.4 kg/m²; moderate malnutrition, BMI=16.0–16.9 kg/m²; severe malnutrition, BMI <16.0 kg/m². 16–18 Bivariate and multiple logistic regression model analysis were used to test the statistical association between nutritional status and sociodemographic characteristics as well as clinical parameters. A p-value <0.05 was taken as statistically significant.

Results
Sociodemographic and Clinical Characteristics of the Study Participants
Of the 450 patients who were recruited into the study, 364 (80.9%) of the study participants were within the age group of 18–45 years with the age range of 4–81 years and median age was 26. Male to female ratio was almost one to one and 180 (40%) were from Arsi and East Shewa zones. About one-third of the patients were illiterate with very few first degree graduates. The clinical history of the patients revealed that nearly a quarter of them were previously treated as PTB patients for at least four weeks. There was delay in seeking medical treatment from modern HFs as concluded from the fact that about 70% sought traditional treatment from the beginning 50s after four weeks of symptoms. One hundred and forty-eight (32.9%) of them were users of khat or cigarette or alcohol beverages or combinations of these at least in the last one year prior to the interview. The TB-HIV co-infection rate was below 10%. History of treatment from modern HFs as concluded from the fact that about 70% sought modern treatment after four weeks of symptoms. One hundred and forty-eight (32.9%) of them were users of khat or cigarette or alcohol beverages or combinations of these at least in the last one year prior to the interview. The TB-HIV co-infection rate was below 10%. History of resistance to anti-TB drug was reported in 23 (7.8%) of previously treated patients (Table 1).

Estimation of Body Mass Index
Two hundred and thirty-two (51.6%, 95% CI: 47.15–56.2) of all the subjects had fallen under the undernutrition range of BMI, and very few were found to be overweight (Table 2). The maximum, average, and minimum BMI of the patients were 27.11 kg/m², 18.27 kg/m², and 12.5 kg/m², respectively.

Association Between Undernutrition and Different Factors
Bivariate regression model analysis has shown that age group <18 years of age compared to age group 29–39; educational status of below secondary school compared to secondary school complete; and family size >4 compared to those with family size ≤4 are associated with undernutrition (p<0.25) as shown in Table 3. Multiple logistic regression analysis indicated that age group <18 was associated (AOR= 0.028, 95% CI: 0.006–0.125) with undernutrition.

In addition, the selected clinical parameters were statistically (p<0.05) associated with undernutrition as computed by binary logistic regression model. Further analysis of the association of these risk factors with undernutrition using multiple logistic regression indicated that positive HIV serum status (adjusted odds ratio, AOR=0.259, 95% CI: 0.104–0.647) and history of resistance to anti-TB drug (AOR=0.394, 95% CI: 0.173–0.897) were statistically associated with subnormal nutritional status (Table 4).

Discussion
The study involved 450 PTB patients in Oromia region of Ethiopia with the intention to determine the magnitude of undernutrition and the association of undernutrition with selected socio-demographic characteristics and clinical history.

The prevalence of undernutrition recorded by the present study was 51.6% and was a bit less than that reported from the Amhara Region of Ethiopia (57.17%),19 but similarly in the beginning 50s with that reported from the Adama Town of Ethiopia (53%).20 The finding of systematic review and meta-analysis on the prevalence of undernutrition in TB patients in Ethiopia also reported a pooled prevalence of 50.8% (95% CI: 43.97–57.63)21 and the report from Kenya, a neighboring country, as the institution-based study done in Turkana, showed the proportion of TB patients with undernutrition of 50.12%22 which are also in agreement with the report of the present study. However, lower prevalence (35.8%) of undernutrition was reported from Burkina Faso23 while a significantly higher prevalence (85%) was reported from central rural India.24 This could possibly be due to differences in socioeconomic standards among others between the populations.

According to this study, those previously treated and who had duration of illness >1 month before starting...
Table 1 Sociodemographic and Clinical Characteristics of the Study Participants, Oromia, 2020

| Characteristics                                      | N (%)        |
|------------------------------------------------------|--------------|
| Age (years)                                          |              |
| <18                                                  | 52 (11.6)    |
| 18–28                                                | 203 (45.1)   |
| 29–39                                                | 106 (23.6)   |
| 40–45                                                | 55 (12.2)    |
| >45                                                  | 34 (7.6)     |
| Sex                                                  |              |
| Male                                                 | 240 (53.3)   |
| Female                                               | 210 (46.7)   |
| Address (zones)                                      |              |
| North Shewa                                          | 37 (8.2)     |
| South West Shewa                                     | 28 (6.2)     |
| East Shewa                                           | 92 (20.4)    |
| Arsi                                                 | 88 (19.6)    |
| West Arsi                                            | 24 (5.3)     |
| Bale                                                 | 64 (14.2)    |
| Guji                                                 | 60 (13.3)    |
| Borena                                               | 25 (5.6)     |
| West Harerge                                         | 32 (7.1)     |
| Education                                            |              |
| Cannot read and write                                | 169 (37.6)   |
| Read and write                                       | 88 (19.6)    |
| Elementary                                           | 146 (32.4)   |
| Secondary                                            | 31 (6.9)     |
| College and above                                    | 16 (3.6)     |
| Family size                                          |              |
| Lives alone                                          | 10 (2.2)     |
| 1–4                                                  | 298 (66.2)   |
| >4                                                   | 142 (31.6)   |
| Category of tuberculosis treatment history           |              |
| New                                                  | 343 (76.2)   |
| Previously treated                                   | 107 (23.8)   |
| History of anti-TB drug resistance (N=107)           |              |
| No                                                   | 58 (54.2)    |
| Yes                                                  | 23 (21.5)    |
| Unknown                                              | 26 (24.3)    |
| Duration of illness before medical treatment         |              |
| <1 month                                              | 137 (30.4)   |
| 1–2 months                                           | 225 (50.0)   |
| >2 months                                            | 88 (19.6)    |
| Currently smoking                                    |              |
| No                                                   | 421 (93.6)   |
| Yes                                                  | 29 (6.4)     |
| Currently chewing khat                               |              |
| No                                                   | 382 (84.9)   |
| Yes                                                  | 68 (15.1)    |
| Drink alcohol beverages                              |              |
| No                                                   | 399 (88.7)   |
| Yes                                                  | 51 (11.3)    |
| HIV serum status                                     |              |
| Negative                                             | 404 (89.8)   |
| Positive                                             | 34 (7.6)     |
| Unknown                                              | 12 (2.6)     |
treatment have shown a tendency to become undernourished probably because of the longstanding and recurrent existence of infection that negatively affects dietary intake, and overall metabolism whereby lipolysis and proteolysis may cause surplus anabolic activities.\textsuperscript{25} Patients with positive HIV serum status have also shown greater tendency to become undernourished; this could be due to additional stress of infection and also possibility of opportunistic diseases on top of the underlying common factors with HIV negative TB cases.

In general, it is an established fact that undernutrition has been found to be associated with adverse TB treatment outcomes and its presence at the initial diagnosis of active TB has been reported to be a predictor of increased risk of death and TB relapse.\textsuperscript{5,26–29} In support of this fact, the study done in Beira (Mozambique) had shown that low BMI, among other social determinants including: being male, low socioeconomic status (education and income), and HIV serum positivity increases the likelihood of therapy failure in newly diagnosed PTB patients.\textsuperscript{30} Studies have also indicated that undernutrition may affect responses to BCG vaccination which is an important approach in the prevention of active TB in addition to impairment of innate and adaptive immune responses needed to control \textit{Mycobacterium tuberculosis} infection.\textsuperscript{31} It is also possible to extrapolate that malnutrition could have a role in the development of drug resistance which could be due to impaired immunity as a result of poor nutrition that may allow organisms of \textit{M. tuberculosis} to stand against the effect of a drug or drugs.

\textbf{Table 3} Association of Sociodemographic Characteristics of TB Patients with Undernutrition in TB Patients, Oromia, 2020 (N=445)

| Characteristics       | Nutritional Status | COR (95% CI) | \(p\)-value | AOR(95% CI) | \(p\)-value |
|-----------------------|--------------------|--------------|-------------|-------------|-------------|
| Age group             |                    | \(<N\) | \(N\) | \(95\%\)  | \(95\%\)  |
| <18                   |                    | 50 | 2 | 0.032 (0.007–0.140) | 0.000 | 0.028 (0.006–0.125) | 0.000 |
| 18–28                 | 92 | 108 | 0.951 (0.952–0.529) | 0.837 | 0.893 (0.547–1.468) | 0.651 |
| 29–39                 | 47 | 58 | 1.047 (0.542–2.020) | 0.892 | 1.031 (0.530–2.007) | 0.928 |
| 40–50                 | 24 | 31 | 1.079 (0.571–1.136) | 0.201 | 0.624 (0.280–1.388) | 0.247 |
| >50                   | 19 | 14 | 1.056 (0.627–1.234) | 0.893 | 0.286 (0.072–1.188) | 0.659 |
| Education             |                    | 86 | 81 | 0.680 (0.313–1.477) | 0.330 | 0.730 (0.327–1.630) | 0.442 |
| Illiterate            | 96 | 62 | 0.659 (0.288–1.508) | 0.324 | 0.786 (0.337–1.835) | 0.578 |
| Read and write        | 81 | 63 | 0.562 (0.256–1.232) | 0.150 | 0.910 (0.404–2.051) | 0.821 |
| Elementary            | 13 | 18 | 1.083 (0.309–3.802) | 0.901 | 1.230 (0.334–4.533) | 0.756 |
| Secondary             | 6 | 9 | 0.653 (0.181–2.363) | 0.516 | 0.531 (0.142–1.983) | 0.347 |
| College +             | 4 | 6 | 0.738 (0.493–1.105) | 0.140 | 1.125 (0.717–1.767) | 0.608 |

| Family size           |                    | \(<N\) | \(N\) | \(95\%\)  | \(95\%\)  |
|-----------------------|--------------------| 6 | 4 | 0.653 (0.181–2.363) | 0.516 | 0.531 (0.142–1.983) | 0.347 |
| 1–4                   | 145 | 148 | 0.738 (0.493–1.105) | 0.140 | 1.125 (0.717–1.767) | 0.608 |
| >4                    | 81 | 61 | 0.653 (0.181–2.363) | 0.516 | 0.531 (0.142–1.983) | 0.347 |

Abbreviations: \(<N\), undernutrition; CI, confidence interval.
It is not only the socioeconomic support that is observed to be in deficit, but also the nutritional assessment and counseling were not as they are supposed to be as manifested by missing information regarding nutritional status of previous TB patients and absences of nutritional counseling for newly presenting patients during data collection. In support of this fact, the qualitative study done in Tigray Region (a preprint of manuscript) has indicated the existence of related problems and that factors associated with these inappropriate conditions in regard to nutrition include: lack of supportive supervision, shortage of staff, phasing out of supporting partners; lack of commitment from health workers; and very low economic status of patients, sharing and selling of provided supplies, and defaulting from treatment.

In line with the socioeconomic aspect of TB, people with TB commonly face reduction of income in two ways. That is, they can become too sick to work and get income; or, they or their families have to cover other expenses related to the treatment even if diagnosis and drug treatment is free. Therefore, they economically go downhill.

Out of the 450 TB patients participated in the study only 30 (7.8%) could possibly benefit from the existing nutritional support system because of their positive HIV status. Supplementary feeding in food insecure and vulnerable people is proven to be associated with reduced number of patients dropping out, increased recovery rate, increased treatment completion and sputum conversion rate during active TB. Food and nutritional support optimize the nutritional value and adequacy of the diet, improve quality of life and improve various health parameters of disadvantaged families.

The socioeconomic situation of TB patients demands multisectoral action to address the social and economic determinants including material support. This can be food or financial support like meals, food baskets, food supplements, food vouchers, transport subsidies, living allowance, housing incentives, or financial bonus as recommended by the WHO in order to access health services and, possibly, to mitigate consequences of income loss related to the disease and consequences of TB as part of the strategies of ending TB.

Conclusion
Quite a significant proportion of TB patients in Oromia region is undernourished. Medical treatment alone is not adequate for effective TB control program. Therefore, nutritional assessment, counseling and need based socioeconomic support for TB patients demand attention of the program holders.

Table 4 Association of Clinical History with Undernutrition in TB Patients, Oromia, 2020

| Characteristic       | Nutritional Status | COR (95%CI)       | p-value | AOR (95%CI)      | p-value |
|----------------------|--------------------|-------------------|---------|------------------|---------|
| Treat. Cat           | <N                 | 1                 |         | 1.41 (0.628–2.075) | 0.665   |
|                      | Normal             |                   |         |                   |         |
| Repeat               | 167                | 173               | 1       | 1.683 (1.076–2.634) | 0.023   |
|                      | 65                 | 40                |         |                   |         |
| Dol                  | <1 month           | 1                 |         | 1.994 (1.141–3.487) | 0.015   |
|                      | 1–2 months         | 108               | 115     | 2.112 (1.257–3.549) | 0.005   |
|                      | >2 months          | 57                | 29      |                   |         |
| HIV status           | Negative           | 200               | 199     | 3.838 (1.634–9.017) | 0.002   |
|                      | Positive           | 27                | 7       |                   |         |
| Khat                 | No                 | 200               | 179     | 0.842 (0.499–1.421) | 0.520   |
|                      | Yes                | 32                | 34      |                   |         |
| Alcohol              | No                 | 207               | 188     | 0.908 (0.504–1.636) | 0.748   |
|                      | Yes                | 25                | 25      |                   |         |
| Cigarette            | No                 | 217               | 199     | 0.983 (0.463–2.087) | 0.963   |
|                      | Yes                | 15                | 14      |                   |         |
| History of DR        | NR                 | 166               | 183     | 2.756 (1.285–5.910) | 0.009   |
|                      | R                  | 25                | 10      |                   |         |

Abbreviations: Dol, Duration of illness; DR, Drug Resistance; NR, No resistance; R, Resistance.
Abbreviations
BMI, body mass index; HF, health facility; PTB, pulmonary tuberculosis; TB, tuberculosis.

Data Sharing Statement
The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

Ethics Approval and Consent to Participate
Ethical clearance was obtained from Health Studies Higher Degree Committee of University of South Africa (Ref No: REC-012714-039, HSHDC/454/2015). The Oromia Regional Health Bureau gave permission and has written a letter of support to respective Health Departments. The purpose of the study was explained to the study participants and written consent was obtained from each. For those aged less than 18 years as well, written informed consent was obtained from their parents/legal guardians. All in accordance with the Declaration of Helsinki. 42

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Disclosure
The authors report no conflicts of interest in this work.

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