Risk Factors Associated with Breast Cancer among Women in Addis Ababa, Ethiopia: Unmatched Case–Control Study

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Background: Breast cancer is a common public health problem and the main cause of cancer-related death worldwide. There is a paucity of evidence on the risk factors of breast cancer in Ethiopia. Therefore, we aimed to identify the risk factors of breast cancer among women in Addis Ababa, Ethiopia.

Methods: We conducted an institutional-based unmatched case–control study with a sample of 348 women (116 cases and 232 controls). Participants were selected by a systematic random sampling technique. Data were collected using an interviewer-administered questionnaire. Data were entered using EpiData version 4.6 and analyzed using SPSS version 25. Multivariable analysis was carried out using the adjusted odds ratio (AOR) with a 95% confidence interval (CI). P-value of less than 0.05 was considered statistically significant.

Results: The mean age (+SD) of the participants was 42.7 (+11.3) and 40.7 (+14.6) for the cases and controls, respectively. Early onset of menarche (AOR= 4.10; 95% CI: 1.84, 9.15), rural women (AOR= 3.64; 95% CI:1.38, 9.57), utilization of packed foods or drinks (AOR= 2.80; 95% CI:1.52, 5.15), and smoke-dried meat (AOR= 2.41; 95% CI:1.36, 4.27), family history of cancer (AOR= 2.11; 95% CI:1.04, 4.26), overweight and/or obesity (AOR= 2.38; 95% CI:1.31, 4.31), and women with one or less children (AOR= 1.86; 95% CI:1.01, 3.41) were associated factors with breast cancer risk.

Conclusion: In this study, early onset of menarche, rural women, utilization of packed foods or drinks and smoke-dried meat, family history of cancer, overweight and/or obesity, and women with one or fewer children were factors that increased breast cancer risk. Therefore, focusing on modifiable risk factors and increasing awareness of the community such as a healthy diet, promotion of breast self-examination, and creation of programs to increase women’s knowledge is important to reduce the increasing burden of breast cancer.

Keywords: breast cancer, risk factors, women, Ethiopia

Background
Breast cancer is the most common public health problem and the main cause of cancer-related death worldwide.1–4 Breast cancer is a type of cancer characterized by uncontrolled development and the spread of abnormal breast cells.5,6 Globally, approximately 2.1 million women, which account for 24.2% of new cancer cases and 627,000 (15%) deaths in 2018.1,7 More than half of the incidence of breast cancer and 60% of deaths occur in low- and middle-income countries (LMICs).1,8 For example, African countries had the highest age-standardized mortality rate (17.3 deaths per 100,000 annually) associated with breast cancer.9,10
In Ethiopia, breast cancer incidence is rising and become the foremost common cancer, causing high rates of morbidity and mortality.11 The incidence of breast cancer accounts for 15,244 (22.6%) all cases of cancer and 8,159 (17%) cancer mortality annually.7,12 The age-standardized incidence rate of breast cancer was 40.6 per 100,000 females.11 According to the Addis Ababa cancer registry, breast cancer incidence increased from 18 to 160 new cases between 2012 and 2018 in Addis Ababa.13

Although breast cancer overwhelmingly occurs in high-income countries, recently, the burden has extended in LMICs due to factors such as a westernized lifestyle and urbanization.1-4 Evidence indicates that breast cancer has several influences, including psychological distress to the patients, family members, reduced productivity, and increased cost of the health care system.14-17 In African countries, including Ethiopia, the health care system cannot handle care and treatment for the expanding incidence rates of breast cancer due to inappropriate treatment strategies in place and economic barriers. Therefore, focusing on preventive strategies through appropriate identification and exposure reduction to established risk factors is important.10,18

According to the World Health Organization (WHO) and American Cancer society, they recognized several risk factors for breast cancer, including early menarche, late menopause, null parity, late age at first pregnancy (>30 years), non-breastfeeding mothers, hormonal contraceptives, hormone therapy after menopause, alcohol, cigarette smoking, obesity, physical inactivity, diet, and family history of breast cancer.6,19-21 Several risk factors are assumed to increase disease occurrence, which incorporates modifiable and non-modifiable risk factors.6,22 However, several epidemiological studies conducted in different populations indicate that the incidence and mortality of breast cancer greatly vary between countries. This indicates that environmental and lifestyle factors may contribute to the development of breast cancer.7,22-25

Primary prevention offers the greatest public health potential and most cost-effective long-term cancer control program. Approximately 50% of breast cancers could be prevented when a maximal benefit is achieved through prevention programs and lifestyle modification.2,20,26 However, adequate information is not available to validate specific lifestyle modification programs, especially in LMICs.7,20,26 Additionally, effective programs to prevent the onset of breast cancer have not been a priority for the health systems in LMICs, including Ethiopia, and there is a lack of awareness of the risk factors of cancer.16

A study conducted in Africa on the management of breast cancer suggests that there is a research gap in LMICs.27 The population of women in Ethiopia is diverse, with different lifestyle factors, cultures, economic statuses, and reproductive and breastfeeding practices. These factors affect the overall health conditions and make it an area of interest for investigating risk factors of breast cancer. Moreover, studies focused on identifying the risk factors in a local context play an important role in the prevention of breast cancer and would contribute to the prioritization of interventions.10,18 Despite the abovementioned geographic and behavioral variation in breast cancer risk and increased incidence rate in low-income countries, including Ethiopia. There is a paucity of studies on risk factors of breast cancer in Ethiopia. Therefore, this study aimed to identify the risk factors associated with breast cancer among women attending Tikur Anbessa Specialized Hospital (TASH) and St. Paul’s Hospital Millennium Medical College (SPHMMC), Addis Ababa, Ethiopia, 2020.

Materials and Methods
Study Setting and Population
The study participants were recruited from the two most common referral public hospitals (TASH and SPHMMC) in Addis Ababa, Ethiopia. We conducted an institutional-based unmatched case–control study from February 1st to March 30, 2020. Women who have histologically confirmed breast cancer cases and unmatched noncancerous clients visiting the same hospital in the study period were included. However, women who had cancer from another site and disseminated to the breast, critically ill, and mentally handicapped patients were excluded. All women diagnosed clinically and histologically confirmed breast cancer were eligible for inclusion as cases. A potential control was a woman aged 20 years or older who had no prior history of breast cancer and who was unrelated to cases. Controls were relatively healthy women recruited from maternal and child health unit, family planning, and outpatient units for any illness other than breast cancer in two hospitals.

Sample Size and Procedure
The sample size was calculated using Epi-Info version 7.2 based on the following assumptions: 95% confidence
interval, 80% power, and 1:2 ratio of cases to controls, 24.3% proportion of family history of breast cancer in Senegal Dakar with an odds ratio of 2.1228 and a 10% non-response rate. This yields a maximum sample size of 116 cases and 232 controls. The hospitals were selected purposively because more than 90% of breast cancer patients were attending the oncology unit in these Hospitals in Addis Ababa, Ethiopia. The sample size was allocated using probability proportional to size based on the proportion of average monthly client flow reviewed from the registration book. Cases were selected for all women with confirmed breast cancer attending two hospitals during the two-month data collection period until the final sample size was reached. For every case, two controls were selected among healthy women between ages 20 to 74 years using systematic random sampling methods.

Data Collection Procedure
All participants were interviewed by trained interviewers at the hospitals. Data were collected by 6 nurses using a pretested, interviewer-administered questionnaire that was adapted and modified from published literature.29,30 The questionnaires were translated into Amharic version, and then translated back to English to ensure its consistency. The questionnaire includes sociodemographic characteristics and behavioral, reproductive, and biological related factors. We defined women who had sisters or mothers or daughters with breast cancer were categorized as having a first-degree family history of breast cancer. Post-menopausal status was defined as no menses for the past one year. The participants were categorized as parous if they had at least one full-term pregnancy (live or stillbirth). The weight scale was calibrated at 0 with no object on it and placed on the level surface before the measurement was carried out. Height was measured with a stadiometer attached to the wall. A continuous checkup of the scale was performed for reliability. Body mass index (BMI) was calculated by BMI = weight (kg)/height square (M2). A BMI of 25.0 or more is overweight, more than 30 is obese, while the healthy range is 18.5 to 24.9.

Data Quality Control
Two days training was given to all the data collectors and supervisors. A pretest was carried out on 5% (in Zewuditu Hospital) of the total sample size before the actual data collection. Based on the findings of the pretest, some modifications were undertaken. The data collection process was closely supervised, and the completeness of each questionnaire was checked daily. During data cleaning, a logical checking technique was employed to identify errors. Finally, double data entry was performed to check the consistency of the data.

Data Processing Analysis
The data were entered into Epi Data version 4.6 and analyzed using Statistical Package for the Social Sciences (SPSS) Version 25. Descriptive statistics including frequencies and proportions were employed. The findings were presented using tables, figures, and graphs. Multivariable logistic regression was used to measure the association between covariates and outcome variable. Adjusting for potential confounders factors, the models were adjusted for age, anthropometric, lifestyle variables, age of menarche, age of menopause, and family history of breast cancer, were added when appropriate. Multicollinearity was checked to see the linear correlation among the independent variables by using variance inflation factor (>10) and standard error (>2) and goodness-of-fit was checked by Hosmer-Lemeshow test (>0.05). Adjusted odds ratio (AOR) with 95% confidence intervals (CI) using a p-value<0.05 was considered a statistically significant association with the outcome variable.

Results
Sociodemographic Characteristics
A total of 116 cases and 232 controls participated in the study, with a response rate of 100.0%. The mean (±SD) age of the participants was 42.7 (±11.3) and 40.7 (±14.6) for cases and controls, respectively. Of the participants, 67 (57.4%) cases and 96 (41.2%) controls were recruited from TASH. Ninety-eight (84.5%) cases and 223 (96.1%) controls were urban dwellers. Of these, more than half 98 (84.5%) of the cases and 162 (69.8%) controls were from Addis Ababa. Thirty-seven (31.9%) cases and 84 (36.2%) controls had college and above educational status. Regarding occupational status, 50 (43.1%) cases, and 91 (39.2%) controls were employed. More than half (55.5%) of the participants earn greater than 2000 Ethiopian birr (ETB) per month (Table 1).

Behavioral and Biological-Related Characteristics
Among the total participants, 37 (31.9%) cases and 58 (25.0%) controls had overweight and/or obesity. Among the study participants, 23 (19.8%) cases and 29 (12.5%)
controls had a family history of cancers. Regarding physical activities, in most cases, 98 (84.5%) and 198 (85.3%) controls did not participate in regular physical exercise. More than half of the cases 79 (68.1%) and 132 (56.9%) controls eat less than seven servings of fruits and vegetables per week. Regarding exposure to smoked meat, 58 (50.0%) cases, and 70 (30.2%) controls were exposed. Thirty-nine (33.6%) cases and 46 (19.8%) controls used packed food or drinks. The majority of the participants, 84 (72.4%) cases and 181 (78.0%) control had no history of breast trauma (Table 2).

**Table 1** Sociodemographic Characteristics of Women among Cases and Controls at TASH and SPHMMC Addis Ababa, Ethiopia, 2020

| Variables                      | Controls (n=232) | Cases (n=116) | Total (n=348) |
|-------------------------------|-----------------|---------------|---------------|
| Place of residence            |                 |               |               |
| Urban                         | 223 (96.1)      | 98 (84.5)     | 321 (92.2)    |
| Rural                         | 9 (3.9)         | 18 (15.5)     | 27 (7.8)      |
| Region                        |                 |               |               |
| Addis Ababa                   | 162 (69.8)      | 98 (84.5)     | 260 (76.1)    |
| Oromia                        | 45 (19.4)       | 10 (8.6)      | 55 (14.9)     |
| Amhara                        | 20 (8.6)        | 5 (4.3)       | 25 (7.2)      |
| Others                        | 5 (2.2)         | 3 (2.6)       | 8 (2.3)       |
| Educational status            |                 |               |               |
| No formal education           | 39 (16.8)       | 27 (23.3)     | 66 (19.0)     |
| Primary education             | 76 (32.8)       | 33 (28.4)     | 109 (31.3)    |
| Secondary education           | 33 (14.2)       | 19 (16.4)     | 52 (14.9)     |
| College and above             | 84 (36.2)       | 37 (31.9)     | 121 (34.8)    |
| Marital status                |                 |               |               |
| Married                       | 135 (58.2)      | 84 (72.4)     | 219 (62.9)    |
| Widowed                       | 28 (12.1)       | 8 (6.9)       | 36 (10.3)     |
| Divorced                      | 11 (4.7)        | 10 (8.6)      | 21 (6.0)      |
| Single                        | 58 (25.0)       | 14 (12.1)     | 72 (20.7)     |
| Occupation                    |                 |               |               |
| House wife                    | 106 (45.7)      | 55 (47.4)     | 161 (46.3)    |
| Employed                      | 91 (39.2)       | 50 (43.1)     | 141 (40.5)    |
| Merchant                      | 12 (5.2)        | 3 (2.6)       | 15 (4.2)      |
| Farmer                        | 23 (9.9)        | 8 (6.9)       | 31 (9.0)      |
| Family monthly income         |                 |               |               |
| ≤ 1000                        | 62 (26.7)       | 37 (31.9)     | 99 (28.4)     |
| 1001–1999                     | 36 (15.5)       | 20 (17.2)     | 56 (16.1)     |
| ≥ 2000 ETB                    | 134 (57.8)      | 59 (50.9)     | 193 (55.5)    |

**Table 2** Behavioral and Biological Related Characteristics of Women among Cases and Controls at TASH and SPHMMC Addis Ababa, Ethiopia, 2020

| Variables                      | Controls (n=232) | Cases (n=116) | Total (n=348) |
|-------------------------------|-----------------|---------------|---------------|
| Physical activity             |                 |               |               |
| Yes                           | 34 (14.7)       | 18 (15.5)     | 52 (14.9)     |
| No                            | 198 (85.3)      | 98 (84.5)     | 296 (85.1)    |
| Intensity of activity         |                 |               |               |
| Light                         | 17 (50.0)       | 10 (55.6)     | 27 (51.9)     |
| Moderate                      | 14 (41.2)       | 6 (33.3)      | 20 (38.5)     |
| Heavy                         | 3 (8.8)         | 5 (11.1)      | 8 (9.6)       |
| Vegetable or fruit intake per week |             |               |               |
| ≤ 7                           | 132 (56.9)      | 79 (68.1)     | 211 (60.6)    |
| > 7                           | 100 (43.1)      | 37 (31.9)     | 137 (39.4)    |
| Exposure to smoked meat       |                 |               |               |
| Yes                           | 70 (30.2)       | 58 (50.0)     | 128 (36.8)    |
| No                            | 162 (69.8)      | 58 (50.0)     | 220 (63.2)    |
| Packed food consumption       |                 |               |               |
| Yes                           | 46 (19.8)       | 39 (33.6)     | 85 (24.4)     |
| No                            | 186 (80.2)      | 77 (66.4)     | 263 (75.6)    |
| History of trauma to the breast |             |               |               |
| Yes                           | 51 (22.0)       | 32 (27.6)     | 83 (23.9)     |
| No                            | 181 (78.0)      | 84 (72.4)     | 265 (76.1)    |
| Ever had a mammogram          |                 |               |               |
| Yes                           | 5 (2.2)         | 7 (6.0)       | 12 (3.4)      |
| No                            | 227 (97.8)      | 109 (94.0)    | 336 (96.6)    |
| Breast self-examination       |                 |               |               |
| Yes                           | 94 (40.5)       | 46 (39.7)     | 140 (40.2)    |
| No                            | 138 (59.5)      | 70 (60.3)     | 208 (59.8)    |

**Reproductive Health-Related Characteristics**

Among participants, 40 (34.5%) cases and 27 (11.6%) controls had seen their first menstrual period before the age of 12 years (Figure 1). Sixty (44.1%) cases and 53
(39.0%) controls had a history of using contraceptives, nearly half (47.5%) of the cases and 21 (39.6%) controls used Oral contraceptives. Seventy-eight (67.2%) cases 181 (78.0%) controls had no history of abortion. From a total of women, the majority of cases 94 (81.0%) and 180 (77.6%) controls had ever breastfed their infants (Table 3).

Factors Associated with Breast Cancer Risk

In the multivariable logistic regression analysis, place of residence, use of packed foods or drinks, exposure to smoked-dried meat, age at menarche, BMI, family history of cancer, and having less than one child were independently associated with breast cancer risk. The risk of breast cancer was higher among rural women (AOR=3.64; 95% CI: 1.38, 9.57) than women who lived in urban areas. Women who used packed foods or drinks were nearly 3 times (AOR= 2.80; 95% CI: 1.52, 5.15) more likely to have breast cancer risk compared with their counterparts. Similarly, women who utilized smoke-dried meat were 2.41 times (AOR= 2.41; 95% CI: 1.36, 4.27) more likely to have breast cancer risk compared with their counterparts. Women with a family history of cancer were 2 times AOR= 2.11; 95% CI: 1.04, 4.26) more likely to be affected with breast cancer than their counterparts. Furthermore, women who had seen their first menstrual period before the age of 12 were almost 4 times (AOR= 4.10; 95% CI: 1.84, 9.15) more likely to have breast cancer risk compared with those women who had seen after the age of 12 years. Moreover, women who were overweight or obese were 2.38 times (AOR= 2.38; 95% CI: 1.31, 4.31) higher chance of getting the risk of breast cancer compared with the normal-weight women. Also, women with one or less child were 1.86 times (AOR=1.86; 95% CI: 1.01, 3.41) more likely to get the risk of breast cancer (Table 4).

Discussion

The identification of modifiable factors in the local context may contribute to the development of prevention strategies that decrease breast cancer incidence and mortality.\textsuperscript{31,32} To our knowledge, this study was the first to investigate breast cancer risk in Ethiopian women. The present study identified that rural residence, early onset of menarche, use of packed foods or drinks, smoked dried meat, BMI, family history of cancer, and less than one child were independently associated with the breast cancer risk.

In this finding, the place of residence had a significant association with breast cancer risk. The odds of breast cancer were approximately 4 times higher among women who were living in rural areas. This is consistent with a study conducted in the Central African Republic that showed that decreased odds of breast cancer were associated with living in urban areas.\textsuperscript{31} However, it is inconsistent with the systematic review and meta-analysis indicating that residing in urban areas is associated with higher breast cancer incidence.\textsuperscript{33} This might be due to differences in the economic condition of the population, educational status, and better awareness of risk factors in urban areas.

The risk of breast cancer may be reduced to the extent that one can make lifestyle changes consistent...
Table 3 Reproductive Health-Related Characteristics of Women among Cases and Controls at TASH and SPHMMC Addis Ababa, Ethiopia, 2020

| Variables                  | Controls (n=232) | Cases (n=116) | Total (n=348) | Count (%) | Count (%) | Count (%) |
|----------------------------|------------------|--------------|--------------|-----------|-----------|-----------|
| Menstrual period regularity|                  |              |              |           |           |           |
| Yes                        | 208 (89.7)       | 93 (80.2)    | 301 (86.5)   |           |           |           |
| No                         | 24 (10.3)        | 23 (19.8)    | 47 (13.5)    |           |           |           |
| Menopausal status          |                  |              |              |           |           |           |
| Premenopausal              | 148 (63.8%)      | 60 (51.7)    | 208 (59.8)   |           |           |           |
| Menopausal                 | 84 (36.2%)       | 56 (48.3)    | 140 (40.2)   |           |           |           |
| Age at menopause           |                  |              |              |           |           |           |
| ≤ 45 years                 | 58 (69.0)        | 37 (66.1)    | 95 (67.9)    |           |           |           |
| 46–54 years                | 24 (28.6)        | 19 (33.9)    | 43 (30.7)    |           |           |           |
| ≥ 55 years                 | 2 (2.4)          | 0 (0.0)      | 2 (1.4)      |           |           |           |
| Duration of contraceptive use|                  |              |              |           |           |           |
| ≤ 4 years                  | 50 (54.9)        | 25 (52.1)    | 75 (54.0)    |           |           |           |
| > 4 years                  | 41 (45.1)        | 23 (47.9)    | 64 (46.0)    |           |           |           |
| Age at first live birth    |                  |              |              |           |           |           |
| ≤ 20 years                 | 67 (46.2)        | 33 (39.3)    | 100 (43.7)   |           |           |           |
| 21–29 years                | 58 (40.0)        | 38 (45.2)    | 96 (41.9)    |           |           |           |
| ≥ 30 years                 | 20 (13.8)        | 13 (15.5)    | 33 (14.4)    |           |           |           |
| History of abortion        |                  |              |              |           |           |           |
| Yes                        | 51 (22.0)        | 38 (32.8)    | 89 (25.6)    |           |           |           |
| No                         | 181 (78.0)       | 78 (67.2)    | 259 (74.4)   |           |           |           |
| Breastfeeding              |                  |              |              |           |           |           |
| Yes                        | 180 (77.6)       | 94 (81.0)    | 274 (78.7)   |           |           |           |
| No                         | 52 (22.4)        | 22 (19.0)    | 74 (21.3)    |           |           |           |

with modifiable risk factors.44 According to this study, women who have used packed foods or drinks were nearly 3 times more likely to have breast cancer risk. This finding is supported by studies conducted in Iran that identified that the consumption of soft drinks and industrially produced juices was associated with a significantly increased risk of breast cancer.35,36 Highly processed foods such as packaged foods, instant soups, reconstituted meats, frozen meals, and shelf-stable snacks also contain substances that may significantly increase the overall risk for cancer and breast cancer.37 Moreover, exposure to smoke-dried meat increases breast cancer risk by 2 times. This finding is supported by studies that indicated processed meat intake was associated with increased breast cancer risk and statistically associated with an elevated risk of all-cause mortality.38–40

Furthermore, the risk of breast cancer was positively associated with overweight and/or obesity. This finding is in agreement with many findings.31–44 Even though, the exact mechanism behind the association between BMI and breast cancer risk is uncertain, there are some potential hypotheses. The positive association between BMI and breast cancer risk in women was speculated to result from the increased estrogens and inflammatory mediators within the larger fat reserves of women of higher BMI and contribute to the aggressive breast cancer phenotype in overweight and obesity.42,45,46

Most of the well-known risk factors for breast cancer are related to the reproductive life of women: Women with early menarche (before age 12 years) have a higher breast cancer risk.47 The present study found that the odds of breast cancer increase by 4 times among women with early menarche, which is consistent with studies conducted in India and Morocco.48,49 This might be due to a woman’s risk of breast cancer being related to the estrogen and progesterone made by her ovaries (known as endogenous estrogen and progesterone). Being exposed for a long time and/or to high levels of these hormones has been linked to an increased risk of breast cancer. In the present findings, contraceptive use did not show a significant association with breast cancer. However, a study in Saudi Arabia50 and Kazakhstan51 showed that the odds of breast cancer were higher among women with long-term use of hormonal contraception. Furthermore, this study revealed that women with fewer children (none or one) increases breast cancer risk. This finding is supported by a similar study conducted in Vietnam.52 We found a history of abortion as a risk factor for breast cancer in crude analysis. This is in line with a study that justified abortion disrupts the maturation process of the breast, it has increased breast cancer risk.34 Besides, null or low parity was related to a higher risk of breast cancer in young women in southern Iran53 and Morocco.49 However, in the current study, parity, age at first live birth, and history of breastfeeding did not show a significant association with breast cancer. This difference might be due to the smaller sample size and poor age recording keeping system in the country.

Family history is an important risk factor for breast cancer and is significantly greater in women with a family history of the disease.24,28,54 Similarly, in this study, family history has significantly associated with the risk of breast cancer. However, this finding is in contrast with
the results of a previous similar study conducted in Vietnam, which indicated that there was no significant association between family history and breast cancer.\cite{55}

Further studies with high-level methodology may be needed to solicit the association between family history and risk of breast cancer.

Limitation of the study: This case–control study cannot determine cause and effect relationships. The findings of this study were based on self-report, as it was not possible to validate claims obtained from study subjects that could be subject to recall bias. A wide confidence interval is noted in some of the variables as a result of a small sample size.

### Conclusion

In conclusion, factors such as the early onset of menarche, rural women, utilization of packed foods or drinks and smoke-dried meat, family history of cancer, overweight and/or obesity, and women with one or less child were factors which increased the risk of breast cancer. Therefore, focusing on modifiable risk factors and increasing awareness of the community such as a healthy diet and promotion of breast self-examination, and the creation of programs to increase women’s knowledge is important to reduce the increasing burden of breast cancer in the study setting. Further studies with larger sample sizes and high-
level study designs are recommended to better understand the local risk factors.

**Data Sharing Statement**
All the data of this study are available from the corresponding author upon reasonable request.

**Ethics Approval and Consent to Participate**
This study was conducted in accordance with the declaration of Helsinki. Ethical clearance was obtained from the Institutional Health Research Ethics Review Committee of the College of Health Sciences, Addis Ababa University (Ref. No. 047/20/SNM). Following approval, a written official letter of cooperation was given to the administrative health bureau and facilities. Informed written consent was obtained from all participants. Furthermore, confidentiality was assured throughout the process.

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**Author Contributions**
All authors made a significant contribution in the conception, study design, execution, acquisition of data, analysis, and interpretation; took part in drafting, revising, or critically for important intellectual content; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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**Disclosure**
The authors declare that they have no competing interests.

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