The Prevalence of Cancer and its Associated Risk Factors among Patients Visiting Oncology Unit, Tikur Anbessa Specialized Hospital, Addis Ababa- Ethiopia

Minyahil Alebachew Woldu†, Dejenu Aklilu Legese‡, Faysel Ebrahim Abamecha† and Alemseged Beyene Berha†
Department of Pharmacology and Clinical Pharmacy, School of Pharmacy, College of Health Sciences, Addis Ababa University, Addis Ababa, Ethiopia
†Equal contributors

Abstract

Background: The global burden of cancer continues to increase largely because of the aging and growth of the world population alongside an increasing adoption of cancer-causing behaviors, particularly smoking, in economically developing countries. Cancer (CA) is an increasing public health burden for Ethiopia and Sub-Saharan Africa at large. In Ethiopia, hospital records show that there are more than 150,000 cancer cases per year and currently cancer accounts for 4% of all deaths.

Objective: The present prospective cross-sectional study was designed to assess the prevalence of cancer and its associated risk factors in patients visiting Tikur Anbessa Specialized Hospital (TASH).

Methods: A prospective cross-sectional study was carried out in patients of all age groups visiting oncology unit of TASH during the study period of November 2015 to June 2016. Sample size was computed using margin of error 5%, confidence level of 95% and estimating cancer patients visiting TASH per annum equal to 20000. The response distribution was taken 10%, considering the 2012 Cancer Prevalence Report of the WHO in East Africa 5% to 10%. The calculated sample size then was 142.

Result and discussion: The mean age of the participants in our study was 42.27 year with a standard deviation of 16.8, and 2 and 86 years were the age range. More than half of the patients in the study (61.3%, n=87) were females and about (38.7%, n=55) were males. In the study area uterine, breast, cervical, colorectal and gastric CA were more prevalent in the age group between 25-49 years while Acute Lymphoblastic Leukemia (ALL) was more common in 13-18 years of age. Ewing sarcoma was seen in age between 19 and 24 years. In our findings, breast CA (14.8%, n=21) was the most frequent type of CA followed by uterine CA (14.1%, n=20), colorectal CA (11.3%, n=16), gastric cancer CA (7.7%, n=11), cervical and esophageal cancers (7%, n=10). ALL was commonly seen in students, while gastric, lung, AML, osteosarcoma; and squamous cell CA were common in farmers. Furthermore, among 20 patients who had breast CA, 65% (n=13) were used contraceptives at least once in their last 10 years period, while uterine cancer patients (40%, n=8) were used either oral or injectable contraceptives. The prevalence of breast CA was not affected by the duration of contraceptive use rather it was stable across the duration of use. Many patients who were developed colorectal CA 12 (8.5%) and gastric CA 9 (6.3%) were reported that they were regularly consuming uncooked red meat as their favorite meal. In our study, many patients were receiving radiation therapy for cases such as uterine, breast, cervical and colorectal CA while chemotherapy was predominantly used in cases such as colorectal, squamous cell, osteosarcoma, ALL, AML and lung cancer. The multinomial regression analysis result further revealed that there was a significant association between breast CA with age (25-49 year, P. value 0.016), Uterine and Cervical CA with gender (female, P. value 0.001); and eating of uncooked red raw meat with colorectal CA (P. value 0.011).

Conclusion: Cancer prevalence in the study area looks greatly jeopardizing the life of many Ethiopians in association with life styles. Hence, further studies should be done. Awareness creation programs to the people, allocation of public health preventive medicine approaches and in-depth governmental involvement in the area are highly needed.

Keywords: Cancer; Cancer associated risk factors; Oncology unit of Tikur Anbessa specialized hospital; TASH

Abbreviations: AAU: Addis Ababa University; AIDS: Acquired Immunodeficiency Syndrome; ALL: Acute Lymphoblastic Leukemia; AML: Acute Myeloid Leukemia; CA: Cancer; CHS: College of Health Sciences; CML: Chronic Myeloid Leukemia; CRC: Colorectal Cancer; DNA: Deoxyribonucleic Acid; ECA: Ethiopian Cancer Association; ESCC: Esophageal Squamous Cell Carcinoma; ETB: Ethiopian Birr; FAP: Familial Adenomatous Polyposis; GBD: Global Burden of Disease; GERD: Gastroesophageal Reflux Disease; HL: Hodgkin-Lymphoma; NHL: Non-Hodgkin Lymphoma; HPV: Human Papilloma Virus; SNNP: South Nation Nationalities and People Region; SOP: School of Pharmacy; SPSS: Statistical Package for Social Sciences; USA: United States of America; WHO: World Health Organization

Introduction
Cancer is a disease in which abnormal cells grow and reproduce

*Corresponding author: Minyahil Alebachew Woldu, Addis Ababa University, College of Health Sciences, School of Pharmacy, Department of Pharmacology and Clinical Pharmacy, P.O. Box 9086, Addis Ababa, Ethiopia, Tel: +251912648527; E-mail: minwoldu@gmail.com

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uncontrollably and invade nearby tissues by spreading to other parts of the body through blood streams and lymphatic systems hindering the activities of the normal cells [1,2]. The global burden of cancer continues to increase largely because of the aging and growth of the world population alongside an increasing adoption of cancer-causing behaviors, particularly smoking, in economically developing countries [3].

Cancer fall into two categories: Solid and hematological cancers. Solid tumor is an abnormal mass of tissue that usually does not contain cysts or liquid areas. Different types of solid tumors are named for the type of cells that form them. Examples of solid tumors are sarcomas, carcinomas, and lymphomas. Hematologic malignancies are forms of cancer that begin in the cells of blood-forming tissue, such as the bone marrow, or in the cells of the immune system. Examples of hematologic cancer are acute and chronic leukemia [4-6].

The exact causes of cancer are not known and very often no single cause can be found. Findings so far show that cancer causes are related to changes on genes. Such changes on genes are related to living habit such as tobacco smoking, unhealthy diet and physical inactivity, and environmental factors like exposure to infections and carcinogens, and longer life expectancy [7-9].

The burden of cancer is increasing in economically developing countries as a result of population aging and growth as well as, increasingly, an adoption of cancer-associated lifestyle choices including smoking, physical inactivity, and “westernized” diets. Regardless of all this, Cancer has received low priority for health care services in Sub-Saharan Africa [10-12].

Cancer arises through a series of somatic alterations in DNA that result in unrestrained cellular proliferation. Most of these alterations involve actual sequence changes in DNA (i.e. mutations). They may originate as a consequence of random replication errors, exposure to carcinogens (e.g. radiation), or faulty DNA repair processes. While most cancers arise sporadically, familial clustering of cancers occurs in certain families that carry a germline mutation in a cancer gene [13-15].

There are two major types of cancer genes. The first type comprises genes that positively influence tumor formation and are known as oncogenes. The second type of cancer genes negatively impact tumor growth and have been named tumor-suppressor genes. Both oncogenes and tumor-suppressor genes exert their effects on tumor growth through their ability to control cell division (cell birth) or cell death (apoptosis), although the mechanisms can be extremely complex [16-19].

Worldwide, about 25 million people are living with cancer. Each year, 7 million people die from it and 11 million new cases are diagnosed globally. In fact more people die from cancer than from AIDS, malaria and tuberculosis put together where 12.5% of all deaths each year in the world are due to cancer. The bad news is that the incidences of cancer in developing countries like Ethiopia, which are not prepared to control, is rising at an alarming rate where as it is declining in the developed world because they started timely to avert it [20,21].

The World Health Organization (WHO) estimated in 2011 that 34% of Ethiopian population is dying from non-communicable diseases, with a national cardiovascular disease prevalence of 15%, cancer and chronic obstructive pulmonary disease prevalence of 4% each, and diabetes mellitus prevalence of 2% [22,23]. The resulting double burden of non-communicable diseases, with higher prevalence of pre-existing communicable, maternal, perinatal and nutritional conditions, constrains the already-meagre health resources and hinders economic development in Ethiopia [16,24,25].

Similarly, global burden of disease (GBD) studies estimated age-standardized death rates of 800 per 100,000 populations for non-communicable diseases in Ethiopia, of which higher death rates (approximately 450 per 100,000) were attributed to cardiovascular disease and diabetes, 150 per100,000 attributed to cancer, and 100 per 100,000 to chronic obstructive pulmonary disease [25,26].

Cancer is an increasing public health burden for Ethiopia and Sub-Saharan Africa at large. In Ethiopia, hospital records show that there are more than 150,000 cancer cases per year and currently cancer accounts for 4% of all deaths. However, Cancer is the most neglected and least prioritized health issue in Ethiopia. On the other hand, the WHO report which updated on February 2015, shows Cancers figure is among the leading causes of morbidity and mortality worldwide, with approximately 14 million new cases and 8.2 million cancer related deaths in 2012, with 5.3 million deaths a year in low income countries. According to this report, the number of new cases is expected to rise by about 70% over the next 2 decades [27,28]. Hence, the present prospective cross-sectional study is designed to assess the prevalence of cancer and its associated risk factors in patients vesting Tikur Anbessa Specialized Hospital (TASH)-the only referral cancer center in Ethiopia until this time.

Methodology

Study setting and period

The study was carried out in oncology unit of TASH from November 2015 to June 2016. TASH is a referral and teaching hospital located in Arada sub city of Addis Ababa, Ethiopia. Until now, it has been the only site where cancer cases are treated within the country. The unit was giving services for the community with the workforce of four oncologists, three palliative care specialists, six pharmacists of which, three were working for adult age groups, two were for pediatrics and one of them who hold a master degree serves as manager, and there were 26 nurses working actively in the unit. This oncology unit was giving services with these workforces on both inpatient and outpatient basis. The inpatient ward has about 18 beds.

Study design and study population

A prospective cross-sectional study was carried out. Patients of all age groups visited oncology unit of TASH during the study period and those who were volunteer to participate in the study were the study population.

Sample size determination and sampling method

Sample size was computed using margin of error 5%, confidence level of 95% and estimating cancer patients visiting TASH per annum equal to 20000. The response distribution was taken 10%, considering the 2012 Cancer Prevalence Report of the WHO in East Africa 5% to 10% [12]. The calculated sample size then was 142. All patients willing to participate was included in the study.

Inclusion and exclusion criteria

Inclusion criteria: All Patients visiting oncology unit of TASH during study period.
Exclusion criteria: Patients who were unwilling to participate were excluded.

Study variables

- **Dependent variable:** Type of cancer
- **Independent variables:** Risk factors

1. Socio-demographic factors such as age, sex, address, ethnicity, religion and marital status
2. Living habits such as source of drinking water, alcohol, exposure to carcinogens (occupation, contamination and accidents), diet, hormones, smoking, obesity, radiation, sunlight and immunosuppression.

Data collection methods and tools

Data was collected from November 2015 to June 2016 through reviewing the patients’ medical records and interviewing either of patients, patients’ parent or care giver. Structured questionnaires and checklists were used focusing on the assessment of risk factors. It contains three parts. The first part contains questions about the socio-demographic characteristics of the patients. The second part contains information about patient medical history and the third part contains questions related to living habits and risk factors associated with corresponding cancer types. The designed questionnaires were translated to local Amharic language and administered to willing study participants.

Data management and analysis

Daily collected data was checked for completeness and compiled and entered in to SPSS version 21. Tables, charts and graphs were used to present the result. Subgroup analysis was done to compare effects of independent variables using Chi-squared or Fishers exact test for categorical variables and Mann-Whitney tests for continuous variables. Logistic regression was done to assess the association of type of cancer with independent variables. A 95% CI and p-value of <0.05 was considered statistically significant.

Ethical clearance

Ethical clearance was obtained from Addis Ababa University, College of Health Science, School of Pharmacy Research and Ethical review board (ERB) and supportive letter from Department of Pharmacology and Clinical Pharmacy was received to conduct the study. Permission was granted by the Head of TASH Oncology Department. 142 patients were interviewed using the structured questionnaire and their medical charts were reviewed after a verbal consent was secured.

Quality assurance mechanisms

Data collection was done by clinical pharmacy oriented bachelor of pharmacy final year students. Two students were given appropriate orientation on how to collect the data and interact with patients and other health care professionals. All steps in data collection and recording were closely supervised by the principal investigator. Daily taken data was scrutinized for accuracy, adequacy and consistency. Discussion among the research team was be done every week on the challenges, progress and success of the study.

Result and Discussion

A total of 142 patients were involved in the study. They were interviewed to identify the risk factors associated with corresponding cancer types and their medical charts were reviewed.

Socio-demographic data

Different studies have shown that, advancing age is the most potent of all carcinogens. In humans, the incidence of cancer rises exponentially in the final decades of life, culminating in a lifetime risk of 1 in 2 for men and 1 in 3 for women. This dramatic age-dependent escalation in cancer risk is fuelled largely by a marked increase in epithelial carcinomas from ages 40 to 80 years, as opposed to cancers of mesenchymal or haematopoietic origin [29]. Similarly, the mean age of the participants in our study was 42.27 years with a standard deviation of 16.75 years and 2 years and 86 years were the age range.

More than half of the patients in our study (61.3%, n=87) were females and about (38.7%, n=55) were males (Table 1). Most of the patients were from Addis Ababa (32.4%, n=46) and Oromia region (31.7%, n=45). This could be better explained by urbanization in case of Addis Ababa, and with respect of the Oromiya region population size could be the factor. The others were (16.2%, n=23) from South Nation Nationalities and People Region (SNNP), (13.4%, n=19) from Amhara, (4.2%, n=6) from Tigray and about (2.1%, n=3) from others.

Of the patients, 59.9% (n=85) were Orthodox Christians, Muslims (22.5%, n=32), Protestant Christians (14.1%, n=20), Adventists (2.1%, n=3) and Catholic Christians (1.4%, n=2). Of the study participants 66 (46.5%) were illiterates, (25.4%, n=36) were attended primary school, (19%, n=27) were attended secondary school, (5.6%, n=8) were attended college while (2.8%, n=4) were attended University level (Table 1).

Cancer type by age distribution

In our study uterine, breast, cervical, colorectal and gastric CA were more common in age group between 25-49 years while ALL was more common in 13-18 years of age, Ewing sarcoma was most prevalent in age between 19 and 24 years (Figure 1).

Cancer type by sex distribution

In our finding, breast CA (14.8%, n=21) was the most frequent type

<table>
<thead>
<tr>
<th>Socio-demographic characteristics</th>
<th>Frequency (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Socio-demographic characteristics</strong></td>
<td>Frequency (%)</td>
</tr>
<tr>
<td><strong>Sex</strong></td>
<td></td>
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<tr>
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<td>87 (61.3)</td>
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<tr>
<td>Primary</td>
<td>36 (25.4)</td>
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<tr>
<td>Secondary</td>
<td>27 (19)</td>
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<tr>
<td>Diploma and above</td>
<td>12 (8.45)</td>
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<tr>
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<tr>
<td>Amhara</td>
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<tr>
<td>SNNP</td>
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<tr>
<td>Tigray</td>
<td>6 (4.2)</td>
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<tr>
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<tr>
<td>Muslim</td>
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<td>Protestant, Christians</td>
<td>20 (14.1)</td>
</tr>
<tr>
<td>Others</td>
<td>5 (3.5)</td>
</tr>
<tr>
<td><strong>Marital status</strong></td>
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<tr>
<td>Single</td>
<td>33 (23.2)</td>
</tr>
<tr>
<td>Married</td>
<td>107 (75.4)</td>
</tr>
<tr>
<td>Others</td>
<td>2 (1.4)</td>
</tr>
</tbody>
</table>

Table 1: The Socio-demographic characteristics of the patients, Addis Ababa, Ethiopia 2016.
of CA followed by uterine CA (14.1%, n= 20), colorectal CA (11.3%, n=16), gastric cancer CA (7.7%, n=11), cervical and esophageal cancers (7%, n=10). The result is in line with other findings across the globe [30,31]. For example, many studies reported that breast cancer was the most prevalent cancer in the vast majority of countries globally; and cervical cancer was the most prevalent cancer in much of Sub-Saharan Africa and Southern Asia while prostate cancer dominates in North America, Oceania and Northern and Western Europe [22,32].

Studies showed that cancer of the lung and bronchus, prostate, and colorectum was relatively common in men as compared to female where as cancers of the lung and bronchus, breast, and colorectum was common among female patients [33]. In our study female were commonly presented with breast CA (22.98%, n=20) and uterine cancer CA (22.98%, n=20) while colorectal CA (18.18%, n=10) and gastric cancer (10.9%, n=6) were common among male patients (Figure 2).

Cancer type by occupation

In our study, ALL was common in students, while gastric, lung, AML, osteosarcoma; and squamous cell CA were common in farmers (Figure 3). Other studies showed that rates of retinoblastoma, Wilms’ tumor, non-Hodgkin’s lymphoma, neuroblastoma, Hodgkin’s disease, acute lymphoid leukemia, acute myeloid leukemia, and osteosarcoma

Figure 1: Pattern of cancer type by age category at Oncology Unit of Tikur Anbessa Specialized Hospital, Addis Ababa, 2016.

Figure 2: Pattern of cancer type by sex at Oncology Unit of Tikur Anbessa Specialized Hospital, Addis Ababa, 2016.
were often higher among young children. Studies also revealed that rates of white children were generally higher than those of black children, especially during the first 5 years of life [34,35].

In most of the studies, a slight excess risk of prostate cancer incidence or mortality was observed among farmers. It is as yet unclear whether this excess risk is caused by particular occupational exposures or by risk factors in their personal lifestyle (e.g. dietary habits) [36].

Many studies also associated rectal cancer with several retail industries; prostate cancer with ministers, farmers, plumbers, and coal miners; malignant melanoma with school teachers; and invasive cervical cancer with women working in hotels and restaurants [37]. In our study, bladder CA was seen in daily laborers, osteosarcoma in industry workers, and gastric cancer in drivers.

Other risk factors

Smoking has been identified as the single most commonly associated risk factor in majority CA types for a number of studies done globally [38-40]. One study reported that cigarette smoking was more strongly associated with death from lung adenocarcinoma [41]. Smoking habit has large impact on causing cancer in males and respiratory cancers and gastrointestinal cancers were most common in males [42]. In our study three lung CA patients were a cigarette smoker for more than 10 years. Two from each AML, gastric and colorectal CA patients were also cigarette smoker (Tables 2 and 3). One study reported that, the dose response curve for intensity of smoking was steeper in women. The findings of this study provided additional evidence for a causal relationship between smoking and all histologic types of lung cancer [43].

Even if it was difficult to associate the number of cigarette use with respect to cancer type in our study, one AML and one lung cancer patient were using more than 15 pieces of cigarette per day. Two gastric CA and one Lung CA and one AML patients were also using 10-15 pieces of cigarette per day (Figure 4). One study also reported that Lifelong nonfilter smokers were at nearly twice the risk of lung cancer compared to lifelong filter smokers after controlling for duration of cigarette use and number smoked per day. Lung cancer risks for filter, nonfilter and mixed smokers increased in proportion to intensity and duration of smoking and decreased with years since stopping smoking. The findings indicate that prevention activities should continue to emphasize smoking cessation, although switching to low-tar cigarettes may also yield some reductions in lung cancer risk [44].

An oral contraceptive agent has been revealed as a possible preventive treatment for breast CA [45]. In our study, among 20 patients who had breast CA, 65% (n=13) were used contraceptives at least once in their last 10 years period. Among uterine cancer patients 40% (n=8) were used either oral or injectable contraceptives. On the other hand, patients who had cervical CA were used either oral or injectable contraceptives (30%, n=3) (Figure 5). In One case control study a multivariate relative risk estimate decreased prevalence of

Figure 3: Distribution of cancer type by occupation among cancer patients at oncology unit of Tikur Anbessa Specilized Hospital, Addis Ababa, Ethiopia; 2016.

![Graph showing distribution of cancer type by occupation among cancer patients at oncololgy unit of Tikur Anbessa Specilized Hospital, Addis Ababa, Ethiopia; 2016.](image-url)

Figure 4: Duration of smoking Type of cancer Number of patients

<table>
<thead>
<tr>
<th>Duration of smoking</th>
<th>Type of cancer</th>
<th>Number of patients</th>
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</thead>
<tbody>
<tr>
<td>5-10 years</td>
<td>breast cancer</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td>nasopharyngeal cancer</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td>AML</td>
<td>2</td>
</tr>
<tr>
<td></td>
<td>gastric cancer</td>
<td>2</td>
</tr>
<tr>
<td></td>
<td>colorectal cancer</td>
<td>2</td>
</tr>
<tr>
<td></td>
<td>esophageal cancer</td>
<td>1</td>
</tr>
<tr>
<td></td>
<td>lung cancer</td>
<td>3</td>
</tr>
</tbody>
</table>

Table 2: A cross tabulation showing type of cancer versus cigarette smoking and duration of smoking, patients at oncology unit of Tikur Anbessa Specilized Hospital, Addis Ababa, Ethiopia; 2016.
Figure 4: A cross tabulation showing type of cancer versus cigarette smoking and number of cigarette pieces used; at oncology unit of Tikur Anbessa Specialized Hospital, Addis Ababa, Ethiopia; 2016.

Figure 5: Type of cancer versus type of family planning methods used; at oncology unit of Tikur Anbessa Specialized hospital, Addis Ababa, Ethiopia; 2016.

invasive epithelial ovarian CA with the increasing duration of oral contraceptive use [46].

In our study prevalence of breast CA was not affected by the duration of contraceptive use rather it was stable across the duration of use. However, the prevalence of uterine cancer was slightly increased by duration of contraceptive use (Figure 6).

Many patients who were developed colorectal CA 12 (8.5%) and gastric CA 9 (6.3%) were reported that they were regularly eating uncooked red meat as their favorite meal. Also, 6 (4.2%) patients who were developed colorectal CA were also eating factory processed meal including tomato, fish, juices etc. regularly (Table 3).

Cancer management approaches in the study area

Access to cancer treatment has been a challenge for third world countries. Considering the case of TASH oncology unit, until recently that was the only referral site for CA cases. Many patients die before getting the right treatment. Among the treatment modalities chemotherapy, radiation, surgery and immunotherapy have been mentioned and the choice is dependent on the specific CA treatment protocol. Surgery has been promising in early stage solid CA cases with adjuvant or neo-adjuvant radiation therapy. Therefore, the increasing the service of this approach may need priority so as to decrease mortality from CA.

In our study, many patients were receiving radiation therapy for cases such as uterine, breast, cervical and colorectal CA while chemotherapy were predominantly used in cases such as colorectal, squamous cell, osteosarcoma, ALL, AML and lung cancer. Many patients were received surgery for gastric and esophageal CA. Chemotherapy and surgery was considered in case of breast and gastric CAs. Chemotherapy and radiation was done for Uterine, breast, colorect, neck and lung CA. Combination of radiation, chemotherapy and surgery also done for cases such as breast and gastric CA (Figure 7).

Multinomial regresional analysis further revealed that there was a significant association between breast CA with age b/n 25-49 year (P. value 0.016), uterine and cervical CA with female gender (P. value 0.001); and eating of uncooked red raw meat with colorectal CA (P. value 0.011). Since, the most common type of cancers reported were cervical and uterine CA then its association with sex as well as age is expected to be associated with female gender. Some studies already reported that, the presence of an increased risk for colorectal cancer followed by consumption of cooked and processed red meat consistently for long-time [47]. However, the association of eating uncooked raw meat and
its association with colorectal CA may need additional study and close observation.

Conclusions
In our study uterine, breast, cervical, colorectal and gastric CA were more common in age group between 25-49 years while ALL was more common in 13-18 years of age, and Ewing sarcoma was most prevalent in age between 19 and 24 years. Breast CA was the most frequent type of CA followed by uterine CA. ALL was common in students, while gastric, lung, AML, osteosarcoma; and squamous cell CA were common in farmers. Many patients who were developed colorectal CA and gastric CA were reported that they were regularly eating uncooked red meat as their favorite meal. Our study revealed that there was a significant association between breast CA with age b/n 25-49 year (P. value 0.016), and eating of uncooked red raw meat with colorectal CA (P. value 0.011).

Competing Interests
The authors declare that they have no competing interests.

Authors’ Contributions
MAW discovered the area, designed the study and revised the whole manuscript. DAL and FEA collected the data and wrote the draft including analysis; and ABB reviewed the whole document. All authors read and approved the final manuscript.

Acknowledgment
We would also like to acknowledge Addis Ababa University, College of Health Sciences, School of Pharmacy for giving us this chance to conduct the study on this topic and also like to extend our gratitude to Tikur Anbessa Specialized Hospital, Oncology Unit staff as well as patients for their cooperation during data collection. Finally it is our pleasure to acknowledge all those who have directly and/or indirectly provided us their supports.

References