Knowledge, Attitudes, and Practices about Trachoma in Rural Communities of Tigray Region, Northern Ethiopia: Implications for Prevention and Control

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Background. Trachoma is a neglected tropical disease which is the leading infectious cause of blindness in the world. Trachoma largely affects people in low socioeconomic conditions in the rural areas and usually communities with poor hygiene and environmental sanitation [2]. In 2019, globally, the number of people at risk of trachoma was 142.2 million and it is responsible for the visual impairment of about 1.9 million people. Ethiopia is one of the most trachoma-affected countries which shares about 50.6% of the burden in the world [4, 5].

In 1998, the World Health Organization (WHO) formally adopted the SAFE strategy (surgery, antibiotics, facial cleanliness, and environmental improvements) for the effective prevention and control of trachoma [3]. The target for eliminating trachoma as a public health problem can be achieved if the SAFE strategy is properly applied. However, trachoma still remains one of the major health problems in the world [4].

Several factors are associated with increased risk of trachoma. These include lack of water, poor personal hygiene, and environmental sanitation [6–8]. Furthermore, poor knowledge [9, 10], and unfavorable sociocultural...
perceptions and poor practices about prevention of trachoma are the main factors in the transmission and sustaining of the infection of trachoma in the communities [11–14].

Trachoma is highly endemic in Tigray Region, Northern Ethiopia, especially in Raya-Alamata and Endamekoni districts (the study area). In 2013, the adjusted prevalence of the clinical sign trachomatous inflammation-follicular (TF) among children aged 1–9 years in the enumeration units encompassing Raya-Alamata and Endamekoni districts was 41.4% and 41.0%, respectively [15]. However, there are no data on knowledge, attitudes, and practices on trachoma prevention and control among rural communities in the two districts so far. Therefore, areas with high burden of trachoma such as Alamata and Endamekoni districts need to assess knowledge, attitudes, and practices and its associated factors of trachoma among rural communities. The findings obtained from the study are of paramount importance for health providers and program managers in planning and implementing to improve prevention and control of trachoma among communities. Moreover, it is useful to policy makers to formulate appropriate interventions for implementation of the SAFE strategy.

2. Methods

2.1. Study Design and Area. A community based cross-sectional study design was conducted to assess knowledge, attitudes, and practices on trachoma among rural communities in two districts (Raya-Alamata and Endamekoni) of Tigray Region, Northern Ethiopia, from May 7–24, 2017. According to the 2007 census, the total estimated population was 96,132 in Raya-Alamata and 94,635 in Endamekoni districts in 2017. From 2014 to 2018; LIGHT FOR THE WORLD in collaboration with the Tigray Regional Health Bureau (TRHB) had a project to eliminate trachoma as a public health problem in the two districts. The focusing activities of the project were mainly on communities’ awareness creation and behavioral change to improve hygiene and environmental sanitation by using different communication channels and health education. The Zithromax MDA campaign was carried out in April 2017 (fourth round) before three-four weeks of the data collection in both districts.

2.2. Sample Size Determination. The sample size was calculated using single population proportion formula with the assumption of proportion of had heard about trachoma among communities in rural Amhara Region, Ethiopia, which was 85.6% [16], using 5% margin of error, 95% confidence level, and nonresponse rate of 10%, and the total sample size was 209.

2.3. Sampling Procedures. Two districts were purposively included in the study. We randomly selected four kebelles (the smallest administrative unit) by the lottery method from the selected districts (2 kebelles from each district). The list of all Women Development Groups (WDGs) with their respective list of households in each selected kebelle was obtained from the health post WDG registration books, and then six WDGs were selected within each selected kebelle using the lottery method. (The WDG is the smallest community organization unit and consists of groups of 25–30 households residing in a neighborhood). Finally, 209 households in the selected WDGs were enrolled using systematic random sampling technique.

2.4. Data Collection Procedures and Data Quality Assurance. A structured questionnaire was adapted from a previous study [16], after reviewed the different related literature. Data were collected by face-to-face interviews and by observations for some practice-related questions through a paper-based questionnaire. We interviewed mothers of households about sociodemographic information, knowledge, attitudes, and practices on trachoma, Zithromax uptake, water, sanitation, and hygiene (WASH), health education, and promotion. Their responses on some practice-related questions were verified by observational technique (availability of latrine and utilization, house compound cleanliness, having separated human and animal dwellings, and solid and liquid waste disposal methods). Six diploma nurses to collect data and two bachelor-degree health officers to supervise the data collection process were recruited. Data collectors and supervisors were trained for two days on the questionnaire, data collection techniques, and the purpose of the study. The pretest was conducted in 5% of respondents (10 households) in two kebelles other than the study sites. Daily completeness and checking for missing data was done by the data collectors and supervisors.

2.5. Operational Definition. Knowledge, attitudes, and practices on trachoma were assessed using 12 knowledge-related, 4 attitude-related, and 8 practice-related questions on trachoma. Correct answers were given a score of one, incorrect answers were scored zero, and those who scoring greater than the mean were considered to have good knowledge, attitudes, and practices, respectively.

2.6. Ethical Approval. The study protocol was reviewed and approved by the institutional ethical review committee of Tigray Health Research Institutes (reference number: THRI/RMO12/2017). The support letter was obtained from the Tigray Regional Health Bureau and the selected district health offices. The respondents were informed about the objective and purpose of the study, and verbal consent was obtained. Confidentiality of the information was assured, and participants were informed that they have the right to withdraw from the interview at any stage.

2.7. Data Processing and Analysis. Data were entered into Epi Info version 3.5.1 and then imported to SPSS version 21 for analysis. Data cleaning and editing was carried out before analysis. Knowledge, attitudes, and practices were computed separately, and its total scores were recorded in to dichotomous categorical variables (good vs poor) based on their
mean score stated in operational definition. Frequency distribution and percentages were performed using frequency tables. Bivariate logistic regression analysis was done for each outcome variables (knowledge, attitudes, and practices). All variables with a p value < 0.25 in the bivariate analysis were included in the multivariable logistic regression model. Finally, variables which had a p value with less than 0.05 were considered statistically significant.

3. Results

3.1. Sociodemographic Characteristics of Study Participants. A total of 194 participants were included in this study, and this made a response rate of 92.8%. The remaining (7.2%) participants were not included due to closed households during the data collection period. From a total respondents, 87 (44.8%) were in the age range of 30–49 years and 112 (57.7%) were currently in the union of marital status. Majority of the respondents, 180 (92.8%), were house wife and 155 (79.9%) were illiterate (Table 1).

3.2. Knowledge on Trachoma. Most (89.2%) of respondents had ever heard about trachoma. Above half (54.6%) of respondents knew that trachoma can be transmitted from person to person and answered correctly that trachoma can be transmitted by dirty fingers (53.1%), flies (35.6%), and by using contaminated towel (24.7%). The majority, 164 (84.5%), of respondents knew trachoma as a preventable disease, and 161 (83%) of respondents knew that trachoma can lead to blindness. The mean score of respondents on knowledge was 6.79 ± 1.40 SD (Table 3). Ninety-nine (51%) of respondents were scored above the mean score and classified as having good knowledge on trachoma (Table 2).

3.3. Attitudes on Trachoma Prevention and Control. Most; 185 (95.4%), of participants agreed availability of adequate water is important for trachoma prevention, and 171 (86.1%) of respondents believed taking mass drug administration is important to prevent and control trachoma. Overall, 96 (49.5%) of respondents were classified as having good attitudes on trachoma prevention and control with a mean score of 3.39 ± 0.69 SD (Table 3).

3.4. Practices on Trachoma Prevention and Control. The majority, 171 (88.1%), of respondents took mass drug administration, and 149 (76.8%) of households had separated human and animal dwellings. However, of the total 194 households, only 72 (37.1%) were utilizing a latrine and 59 (30.4%) had clean house compounds. Out of the total, 69 (35.6%) of respondents were classified as having good practices towards trachoma prevention and control with a mean score of 4.10 ± 1.40 SD (Table 4).

3.5. Factors Associated with Good Knowledge, Attitudes, and Practices towards Trachoma. Factors associated with good knowledge, attitudes, and practices towards trachoma prevention and control were observed. Having ever received health education about trachoma was significantly associated with good knowledge (AOR = 4.10; 95% CI: 1.91–8.79) and good attitudes (AOR = 2.10; 95% CI: 1.02–4.25) (Tables 5 and 6). Moreover, good knowledge on trachoma was significantly associated with good practices towards trachoma prevention and control (AOR = 2.86; 95% CI: 1.46–5.62) after adjusting the other variables (Table 7).

4. Discussion

This study assessed knowledge, attitudes, and practices on trachoma in two districts of Tigray Region, Northern Ethiopia. Having ever received health education about trachoma was significantly associated with good knowledge and attitudes about trachoma prevention and control. Furthermore, good knowledge was significantly associated with good practices towards trachoma prevention and control.

In this study, 89.2% of respondents had ever heard about trachoma. This result was almost the same with the reports from Ethiopia (92.6%) and Bangladesh (86%), [17, 18]. However, it is higher than the report in Kenya (65.7%) of participants had heard about trachoma [10]. The discrepancy could be due to the differences in social mobilization activities in different sites and studies methods.

The current study shows that 54.6% of respondents knew trachoma can be transmitted from person to person. The respondents mentioned that trachoma can be transmitted by contaminated fingers, flies, and contaminated towels. Similar studies from Kenya and Ethiopia reported that the most reported mode of trachoma transmission was contact with flies and dirty [10, 19]. The current study results also indicate

| Variables                           | Numbers | Percent |
|------------------------------------|---------|---------|
| Age                                |         |         |
| 18–29 Years                        | 69      | 35.6    |
| 30–49 Years                        | 87      | 44.8    |
| ≥50 Years                          | 38      | 19.6    |
| Marital status                     |         |         |
| Currently in union                 | 112     | 57.7    |
| Currently not in union**           | 82      | 42.3    |
| Occupation                         |         |         |
| House wife                         | 180     | 92.8    |
| Others*                            | 14      | 7.2     |
| Educational status                 |         |         |
| Illiterate                         | 155     | 79.9    |
| Literate                           | 39      | 20.1    |
| Family size                        |         |         |
| ≤5                                 | 132     | 68      |
| >5                                 | 62      | 32      |
| Monthly family income in US dollar |         |         |
| <22 USD                            | 84      | 43.3    |
| 22–44 USD                          | 87      | 44.8    |
| >44 USD                            | 23      | 11.9    |

*Others (government employed, merchant, and daily laborer). ** Currently not in union (single, widowed, divorce, and separated).
that the majority (84.5%) of respondents knew trachoma as preventable disease. The participants responded that trachoma can be prevented by using latrine, improve environmental sanitation, not using common towel, and washing hand and face with soap. Having knowledge on washing face, not sharing towels, and environmental sanitation was important on prevention of trachoma infection [11].

Regarding attitudes and practices of respondents in the current study, the majority, 86.1%, of respondents agreed that taking mass drug administration is important to prevent and control trachoma and 88.1% of the respondents took mass drug administration. This result is almost comparable with the other reports from Amhara and Tigray regions, and the coverage of mass drug administration ranged from 76.8% to 93.3% [20–22]. On the other hand, 88.1% of respondents agreed that trachoma can be prevented by utilizing latrine. However, only 37.1% of households were utilizing a latrine. Our finding on latrine utilization was

Table 2: Knowledge of study participants on trachoma in two rural districts of Tigray Region, Northern Ethiopia, 2017 (N = 194).

| Variables                                      | Numbers | Percent |
|------------------------------------------------|---------|---------|
| Heard about trachoma disease                   |         |         |
| Yes                                            | 173     | 89.2    |
| No                                             | 21      | 10.8    |
| Knew trachoma can be transmitted from person to person |         |         |
| Yes                                            | 106     | 54.6    |
| No                                             | 88      | 45.4    |
| Knew trachoma can be transmitted by contaminated fingers |         |         |
| Yes                                            | 103     | 53.1    |
| No                                             | 91      | 46.9    |
| Knew trachoma can be transmitted by flies       |         |         |
| Yes                                            | 69      | 35.6    |
| No                                             | 125     | 64.4    |
| Knew trachoma can be transmitted by contaminated towels |         |         |
| Yes                                            | 48      | 24.7    |
| No                                             | 146     | 75.3    |
| Knew trachoma as preventable disease           |         |         |
| Yes                                            | 164     | 84.5    |
| No                                             | 35      | 15.5    |
| Knew trachoma can lead to blindness            |         |         |
| Yes                                            | 161     | 83      |
| No                                             | 33      | 17      |
| Knew trachoma can be prevented by washing hands with soap |         |         |
| Yes                                            | 118     | 60.8    |
| No                                             | 76      | 39.2    |
| Knew trachoma can be prevented by washing face |         |         |
| Yes                                            | 131     | 67.5    |
| No                                             | 63      | 32.5    |
| Knew trachoma can be prevented by using latrine |         |         |
| Yes                                            | 37      | 19.1    |
| No                                             | 157     | 80.9    |
| Knew trachoma can be prevented by improving environmental sanitation |         |         |
| Yes                                            | 159     | 82      |
| No                                             | 30      | 15.5    |
| Knew trachoma can be prevented by not using common towel |         |         |
| Yes                                            | 48      | 24.7    |
| No                                             | 146     | 75.3    |
| Mean +SD 6.79 ± 3.29                           |         |         |
| Median + IQR 7 ± 5.25                          |         |         |

Score of knowledge on trachoma

| Good | Poor |
|------|------|
| 99   | 95   |
| 51   | 49   |

Table 3: Attitudes of study participants on trachoma in two rural districts of Tigray Region, Northern Ethiopia, 2017 (N = 194).

| Variables                                      | Numbers | Percent |
|------------------------------------------------|---------|---------|
| Availability of adequate water is important for trachoma prevention and control |         |         |
| Agree                                        | 185     | 95.4    |
| Disagree                                     | 9       | 4.6     |
| Personal hygiene is important for trachoma prevention |         |         |
| Agree                                        | 134     | 69.1    |
| Disagree                                     | 60      | 30.9    |
| Latrine utilization is important for trachoma prevention |         |         |
| Agree                                        | 171     | 88.1    |
| Disagree                                     | 23      | 11.9    |
| Taking mass drug administration is important to prevent and control trachoma |         |         |
| Agree                                        | 167     | 86.1    |
| Disagree                                     | 27      | 13.9    |
| Mean +SD 3.39 ± 0.69                         |         |         |
| Median + IQR 3 ± 1                           |         |         |

Score of attitudes on trachoma prevention and control

| Good | Poor |
|------|------|
| 96   | 98   |
| 49.5 | 50.5 |

Table 4: Practice of community towards trachoma prevention and control in two rural districts of Tigray Region, Northern Ethiopia, 2017 (N = 194).

| Variables                                      | Numbers | Percent |
|------------------------------------------------|---------|---------|
| Took mass drug administration for prevention of trachoma |         |         |
| Yes                                            | 171     | 88.1    |
| No                                             | 23      | 11.9    |
| Utilization of adequate water for bathing       |         |         |
| Yes                                            | 150     | 77.3    |
| No                                             | 44      | 22.7    |
| Having separated human and animal dwellings    |         |         |
| Yes                                            | 149     | 76.8    |
| No                                             | 45      | 23.2    |
| Latrine utilization                            |         |         |
| Yes                                            | 72      | 37.1    |
| No                                             | 122     | 62.9    |
| Handwashing practice after using latrine       |         |         |
| Yes                                            | 95      | 49      |
| No                                             | 99      | 51      |
| Having proper solid waste disposal management  |         |         |
| Yes                                            | 83      | 42.8    |
| No                                             | 111     | 57.2    |
| Having proper liquid waste disposal management |         |         |
| Yes                                            | 15      | 7.7     |
| No                                             | 179     | 92.3    |
| Having clean house compound                    |         |         |
| Yes                                            | 59      | 30.4    |
| No                                             | 135     | 69.6    |
| Mean +SD 4.10 ± 1.40                          |         |         |
| Median + IQR 4 ± 2                            |         |         |

Score of practices on trachoma prevention and control

| Good | Poor |
|------|------|
| 69   | 125  |
| 35.6 | 64.4 |
lower than from previous studies in Ethiopia [9, 17]. This difference could be due to differences in the study areas because our study was not included urban communities. Latrine utilization has been considered an important intervention for trachoma prevention and control [23]. It is important to reduce burden of flies and improve environmental sanitation through proper disposal of human excreta [24].

In the current study, health education about trachoma was significantly associated with knowledge on trachoma. Previous interventional studies from Ethiopia and Egypt reported that mothers' knowledge was significantly improved after the implementation of the program based health education [16, 25]. Providing health education about trachoma can increase knowledge on trachoma dramatically [26]. With adequate and appropriate health education, communities may increase knowledge on the cause of trachoma, how it is transmitted, and how the disease is prevented [2].

Health education on trachoma was also a predictor of attitudes on trachoma prevention and control. The odds of good attitudes on trachoma prevention and control among participants who had ever received health education were 2.1 times higher than participants who had not ever received health education about trachoma. Health education can help reducing misconceptions and changing cultural perceptions of communities on trachoma prevention and control [11, 27]. Effective health education is the key to building the favorable attitudes on trachoma prevention and control among communities to achieve the goal of eliminating trachoma as a public health problem [28, 29]. In Ethiopia, health extension workers and WDG leaders are giving health education to communities on personal hygiene, environmental sanitation, and mass drug administration for prevention and control of trachoma [22]. This could be the possible explanation that if mothers have received health education on trachoma, they may feel responsibility and accountability to prevent their children and families from trachoma infection. This could be resulted to have good attitudes on prevention and control of trachoma.

The current study revealed that mothers who have good knowledge on trachoma was significantly associated with good practices on trachoma prevention and control. This is supported by a study from Egypt, reported that knowledge was significantly associated with trachoma prevention and control practices [25]. Another study from Ethiopia also revealed that good knowledge was a predictor of good practices towards childhood blindness prevention [19]. Increasing knowledge of trachoma transmission and prevention (e.g., F&E-related preventive behaviors) is important to improve practices of trachoma prevention and control [30]. This might be due to the reason that if someone

| Variables | Knowledge | COR (95% CI) | AOR (95% CI) |
|-----------|-----------|--------------|--------------|
|           | Good (%)  | Poor (%)     |              |
|           | N=194     | N=194        |              |
|           |            |              |              |
| Marital status |          |              |              |
| Currently in union | 53 (53.5) | 59 (62.1) | 1 | 1 |
| Currently not in union | 46 (46.5) | 36 (37.9) | 1.42 (0.80–2.52) | 1.39 (0.77–2.53) |
| Availability of radio/television in the households | | | | |
| Yes | 19 (19.2) | 12 (12.6) | 1.64 (0.75–3.60) | 1.35 (0.60–3.03) |
| No | 80 (80.8) | 83 (87.4) | 1 | 1 |
| Ever received health education about trachoma | | | | |
| Yes | 88 (88.9) | 62 (65.3) | 4.26 (2.00–9.07) | 4.10 (1.91–8.79) |
| No | 11 (11.1) | 33 (34.7) | 1 | 1 |

*Statistically significant at 0.05 < p < 0.01. **Statistically significant at 0.01 < p < 0.001. ***Statistically significant at p < 0.001.

| Variables | Attitude | COR (95% CI) | AOR (95% CI) |
|-----------|----------|--------------|--------------|
|           | Good (%)  | Poor (%)     |              |
|           | N=194     | N=194        |              |
|           |            |              |              |
| Age       |           |              |              |
| 18–29 years | 30 (31.2) | 39 (39.8) | 1 | 1 |
| 30–49 years | 49 (51)   | 38 (38.8) | 1.68 (0.89–3.17) | 1.46 (0.75–2.84) |
| ≥50 years | 17 (17.3) | 21 (21.4) | 1.10 (0.47–2.34) | 1.20 (0.53–2.72) |
| Family size |           |              |              |
| ≤5 | 59 (61.5) | 73 (74.5) | 1 | 1 |
| >5 | 37 (38.5) | 25 (25.5) | 1.83 (0.99–3.38) | 1.52 (0.78–2.93) |
| Ever received health education about trachoma | | | | |
| Yes | 81 (84.4) | 69 (70.4) | 2.27 (1.13–4.58) | 2.10 (1.02–4.25) |
| No | 15 (15.6) | 29 (29.6) | 1 | 1 |

*Statistically significant at 0.05 < p < 0.01. **Statistically significant at 0.01 < p < 0.001. ***Statistically significant at p < 0.001.
knew benefit of personal hygiene and environmental sanitation, she/he could practice to prevent and control trachoma [1].

4.1. Limitation of the Study. The sample size of this study was relatively small and did not consider the design effect to account for the multilevel nature of the sampling design. This may have affected both the precision of the prevalence estimates as well as the statistical inferences of the modeling. Responses on knowledge-, attitude-, and practice-related questions might be influenced by social desirability biases. Correct responses for those are socially acceptable opinions and practices, and incorrect responses for those are socially undesirable opinions and practices. There might be missed some factors that could influence knowledge, attitudes, and practices that were not considered in the model.

5. Conclusion and Recommendation

Our study implies that areas with high burden of trachoma need to improve communities’ knowledge, attitudes, and practices on trachoma prevention and control in order to eliminate trachoma as a public health problem. Therefore, health education focused on the SAFE strategy should be provided at community level, during mass drug administration, and at health facilities while people are waiting to receive treatment to increase good knowledge and changing cultural perceptions that contribute to behavioral changing and increase good practices towards trachoma prevention and control among communities.

Abbreviations

AOR: Adjusted odds ratio  
CI: Confidence interval  
COR: Crude odds ratio  
SD: Standard deviation  
WDG: Women developmental group.

Data Availability

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

Conflicts of Interest

All authors declare that they have no conflicts of interest.

Authors’ Contributions

HG, NM, TT, KF, TG, AB, FG, and KD contributed to the conception, development of data collection tools, design, data collection, analysis, and interpretation of the data. HG

Table 7: Logistic regression analysis of selected variables with practice of communities on trachoma in two rural districts of Tigray Region, Northern Ethiopia, 2017 (N = 194).

| Variables                                | Practice |          |          |          |          |
|------------------------------------------|----------|----------|----------|----------|----------|
|                                           | Good (%) | Poor (%) | COR (95% CI) | AOR (95% CI) |
| Marital status                           |          |          |          |          |          |
| Currently in union                       | 33 (47.8)| 79 (63.2)| 1        | 1        |
| Currently not in union                   | 36 (52.2)| 46 (36.8)| 1.87 (1.03–3.40)* | 1.71 (0.88–3.23) |
| Educational status                       |          |          |          |          |          |
| Illiterate                               | 52 (75.4)| 103 (82.4)| 1        | 1        |
| Literate                                 | 17 (24.6)| 22 (17.6)| 1.53 (0.75–3.13) | 1.01 (0.43–2.40) |
| Occupation                               |          |          |          |          |          |
| House wife                               | 61 (88.4)| 119 (95.2)| 1        | 1        |
| Others                                   | 8 (11.6)| 6 (4.8)| 2.60 (0.86–7.83) | 2.10 (0.57–7.64) |
| Availability of radio/television in the household |          |          |          |          |          |
| Yes                                      | 18 (26.1)| 13 (10.4)| 3.04 (1.39–6.68)** | 2.20 (0.93–5.20) |
| No                                       | 51 (73.9)| 112 (89.6)| 1        | 1        |
| Time taken to fetch water from its source |          |          |          |          |          |
| ≤30                                      | 57 (82.6)| 94 (75.2)| 1.57 (0.75–3.29) | 0.97 (0.40–2.40) |
| >30 minutes                              | 12 (17.4)| 31 (24.8)| 1        | 1        |
| Access to sustainable sufficient water supply |          |          |          |          |          |
| Yes                                      | 63 (91.3)| 104 (83.2)| 2.12 (0.81–5.54) | 2.02 (0.66–6.18) |
| No                                       | 6 (8.7)| 21 (16.8)| 1        | 1        |
| Ever received health education about trachoma |          |          |          |          |          |
| Yes                                      | 60 (87)| 2.59 (1.16–5.78)* | 1.55 (0.63–3.83) | 1        |
| No                                       | 9 (13)| 1        | 1        | 1        |
| Total score of knowledge on trachoma     |          |          |          |          |          |
| Good                                     | 22 (31.9)| 73 (58.4)| 3.0 (1.62–5.57)** | 2.86 (1.46–5.62)** |
| Poor                                     | 47 (68.1)| 52 (41.6)| 1        | 1        |
| Total score of attitudes on trachoma     |          |          |          |          |          |
| Good                                     | 30 (43.5)| 68 (54.4)| 1.55 (0.86–2.80) | 1.51 (0.79–2.87) |
| Poor                                     | 39 (56.5)| 57 (45.6)| 1        | 1        |

*Statistically significant at 0.05 < p < 0.01. **Statistically significant at 0.01 < p < 0.001. ***Statistically significant at p < 0.001.
did the major writing work of the manuscript and revised the draft and final manuscript. NM, TT, KF, TG, AB, FG, and KD revised the draft of the manuscript. All authors read and approved the final manuscript.

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