Appropriateness and Factors Associated with Antibiotics Prescription to Outpatients with Respiratory Tract Infection in Tema Polyclinic, Ghana.

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Research

Keywords: Appropriateness, antibiotics, factors, prescriptions, respiratory tract infections, Tema Polyclinic, Ghana

DOI: https://doi.org/10.21203/rs.3.rs-41132/v1

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Abstract

**Background:** Appropriate prescription of antibiotics, consistent with nationally approved guidelines for the management of patients with respiratory tract infections (RTI) is an important step to reduce antibiotics resistance. This study assessed the appropriateness and factors associated with antibiotic prescription among RTI outpatients at the Tema Polyclinic in Ghana.

**Methods:** Records of 600 outpatients diagnosed with at least one RTI condition between 1\textsuperscript{st} July and 31\textsuperscript{st} December, 2018 were manually retrieved and assessed in an analytic cross-sectional study. The outcomes were the proportion of RTI cases appropriately managed with antibiotics using Kunin's modified criteria and antibiotic prescription. Independent variables included sociodemographic factors, clinical factors and prescriber's professional category. Univariable logistic regression was used to estimate crude odds ratios for factors associated with antibiotic prescription. A threshold of p< 0.20 was used to include factors into a multivariable logistic regression model. Adjusted odds ratios were estimated using backward stepwise logistic regression. Statistical significance was set at p< 0.05.

**Results:** The proportion of RTI cases which were prescribed with at least one antibiotic was 59.7% (358/600). Majority of RTI cases (68.2%) had non-specific diagnoses. Of the 32.8% (121/600) with specific diagnoses, (63.3%, 95% CI: 56.2, 69.9) were appropriately prescribed with antibiotics. Inadequate dosage duration accounted for most of the inappropriately prescribed antibiotics. The factors which were independently associated with reduced odds of antibiotic prescription were presenting with catarrh, diagnosis of rhinitis, acute respiratory tract infection, upper respiratory tract infection, and common cold. Requesting laboratory investigations, presenting with breathlessness, headache and sputum production were associated with increased odds of antibiotic prescription in the adjusted model.

**Conclusion:** Majority of patients with specific diagnoses were prescribed antibiotics appropriately. However, most cases could not be evaluated for appropriateness because the diagnoses were non-specific. The frequency of antibiotic prescription was high. Antibiotic stewardship efforts should focus on imparting knowledge on specific diagnoses and corresponding treatment with emphasis on dosage regimen while deemphasizing symptomatic treatment based on sputum production.

**Background**

Respiratory tract infections (RTIs) are among the most common acute illnesses seen by physicians at the outpatients department (OPD) in majority of primary healthcare facilities globally(1–5). The incidence of RTI is higher in children and responsible for most hospital admissions, especially among those younger than 5 years (1, 6, 7). In 2016, RTIs (mainly lower respiratory tract infections, LRTI) accounted for 5.2% (2,964,000) of total deaths from all causes globally (8). Deaths attributable to RTIs in 2016 alone for Ghana was 140.0 per 100,000 population with 1,085,100 disability-adjusted life years (DALYs) (8, 9). There has been a drop in morbidity from 18.3% in 2016 to 12.5% in 2017 in the Ghanaian public sector(10).

To rationalise the use of antibiotics in the management of RTI, the National Institute for Health and Clinical Excellence (NICE) of Great Britain has recommended that
1. serious conditions which may require antibiotics such as bacterial pneumonia as required by available treatment guidelines need to be ruled out
2. the patient or caregiver of a child is educated and reassured, and
3. symptomatic relief is provided those who do not require antibiotics (11, 12).

Other strategies to improve on use of antibiotics in RTI management include avoiding, delaying or deferring the prescribing of antibiotics (12, 13). The treatment recommendations by the Ghana Standard Treatment Guidelines (STG) is similar to the above in managing specified RTIs. Additionally, strategies such as antibiotic stewardship and provision of guidelines, clinician and patient education have been shown to be valuable (14–16). For instance, the existence of functional drugs and therapeutic committees (DTCs) which has responsibility for antibiotic stewardship in hospitals promotes rational prescription of antibiotics and compliance to guidelines (7, 17).

However, use of standard protocols for treatment of RTI, especially in terms of antibiotic prescription, has not been adequate. In spite of the fact that most RTIs are viral in aetiology and thus self-limiting, several studies have identified frequent use of antibiotics in their management across the globe (4, 5, 12, 18, 19). The high rate of antibiotic use in RTI management is often associated with inappropriate prescription. A study in the USA among veterans found that only 11% of antibiotics were optimally prescribed for management of RTI, with 89% being inappropriate (20). Prescription rates of above 50% have been reported in some developed countries (11, 12, 21, 22). In Africa, antibiotic prescription rates between 26% and 70% have been observed (18, 23–25). A study conducted among children in two facilities in Ghana reported that 30–65% of the RTI cases were inappropriately prescribed antibiotics (18). Inappropriately prescribed antibiotics can lead to antibiotic resistance (23, 26, 27). Thus, ensuring the appropriate prescription of antibiotics in the management of RTI is a key step in combatting antimicrobial resistance (12). Reduction in inappropriate antibiotic prescription is also key in reducing needless exposure to potential side effects/adverse reactions from antibiotics, drug interactions and increased medical cost (11, 12, 28).

Patient, physician and institutional factors are associated with antibiotic prescription. While a physician's area of specialty, and perceptions of a patient's desire for antibiotics (31, 34, 35) can influence prescribing behaviour, some diagnoses such as bronchitis, or symptoms including coloured sputum, nasal discharge, fever, breathlessness, bronchial exudates, and requests for laboratory investigations, are associated with antibiotic prescription (29–35).

In Ghana, the Ministry of Health and Ghana Health Service have been organizing trainings and seminars for health workers on appropriate use of antibiotics and rational use of medicines. The national antimicrobial policy was also recently launched as part of broad measures to address antimicrobial resistance in Ghana (36).

Despite all these measures, antibiotic prescription per patient encounter at the OPD of Tema polyclinic has remained above 40% for three consecutive years (i.e. 53.3, 51.9 and 46.0% in 2016, 2017 and 2018 respectively). These rates are above the National and WHO target of less than 30%. Additionally, there has been increased claims-rejection by the national health insurance scheme (NHIS) for patients diagnosed with RTI, majority of whom were prescribed antibiotics (37). These underscore the need for a thorough
assessment of the appropriateness of antibiotic use in the facility. Kunin's criteria modified by Deshmuk and colleagues, is a tool with standardized indicators and as such is reliable for such an assessment (28). Its use requires that treatments are standardized for purposes of benchmarking. In Ghana, the Standard Treatment Guideline (STG) serves this purpose (38). Generally, the STG recommends the identification of signs of suprainfection either by symptom or clinical investigation before antibiotic prescription. Regular audit of prescriptions at the OPD to ascertain their appropriateness is necessary to identify and address inappropriate use at the facility level. It will also contribute to the promotion of practices that will reduce the development of antibiotic resistance. Therefore, this study was designed to assess the appropriateness of antibiotic prescription to RTI outpatients and identify the factors associated with antibiotic prescription at the Tema Polyclinic in Ghana.

Methods

Study location

The study was conducted at the Tema Polyclinic in the Greater Accra region of Ghana. In 2018, the Polyclinic served as the main sub-metropolitan primary health care facility for the Tema West sub-metropolis with an estimated population of 138,410. The facility has 6 medical doctors in general practice, 2 obstetrician/gynaecologists, 2 dental surgeons, 6 physician assistants and 4 nurse prescribers.

The medical laboratory is staffed with 4 biomedical scientists and equipped to undertake the tests for which data were collected in the study, except for culture and sensitivity testing, which patients had to get from elsewhere. The Pharmacy unit supplies various classes of antibiotics once a valid prescription is presented.

Study design

This was a facility-based retrospective analytic cross-sectional study. Data were extracted from folders of patients who had been diagnosed with a specific or non-specific RTI at the out-patient department (OPD) of the polyclinic from 1st July to 31st December, 2018. The specific RTI diagnoses included otitis media, bronchitis, common cold, pharyngitis, pneumonia, rhinitis, tonsillitis and sinusitis. Non-specific RTI were cases diagnosed as acute respiratory tract and upper respiratory tract infections.

Eligibility criteria

OPD folders with at least one specific or non-specific RTI diagnosis during the study period were considered to have met the inclusion criteria. Of these, those excluded were folders of pregnant women, and patients who had HIV or TB. In addition, patients who visited the dental unit on the same day RTI diagnosis was made or visited the OPD for review of previously diagnosed RTI. They were excluded because ideal antibiotic choices for such patients may rightly vary from STG recommendations which are for the general population.

Sampling
A sampling frame consisting of all 1,331 patient folders with specific or non-specific RTI diagnoses within the study period, grouped month by month, was generated. Systematic random sampling was used to select 600 folders. For each month, every third folder was selected and the process repeated if necessary, until the sample size for the month was reached.

Data collection

Ashaiman polyclinic, located in the Ashaiman Municipality, served as site for pretesting of the data collection tool. The tool was designed based on the STG and the Kunin's modified criteria (28). The tool was used to collect patients' biodata, presenting signs and symptoms, laboratory investigations, diagnoses, medicines prescribed, their doses, frequency of administration and duration of treatment, and professional category of the prescriber. The laboratory investigations included comprised blood film for malaria parasites or malaria rapid diagnostic test, full blood count, blood urea electrolytes, stool examination for parasitic infections (stool routine examination), simple urine examination for protein and haematuria (urine routine examination), urine pregnancy test, fasting blood sugar, Widal test, and sickling test. Presence or absence of the following symptoms and signs were recorded: abdominal pain, breathlessness, chest pain, cough, diarrhoea, fever, headache, malaise, nasal congestion, rapid breathing, rhinorrhea, sore throat, sneezing, wheezing, catarrh, and sputum.

Antibiotic prescription, the primary outcome, was defined as a primary visit having at least one (1) antibiotic prescribed. A prescribed medicine was considered an antibiotic if it is classified under the WHO Anatomical Therapeutic Chemical classification J01 (Antibacterials for systemic use) or P01AB (nitroimidazole derivatives) (39). Antibiotics for topical use were not included. The independent variables were grouped as sociodemographic factors, symptoms, diagnoses, laboratory investigations and prescriber's professional category.

Two pharmacists including a specialist clinical pharmacist assessed the appropriateness of antibiotics prescribed using the STG recommendations and the Kunin's modified criteria (28) as benchmark.

1. Agree with the use of antibiotic therapy given as in the prescription. The treatment is appropriate in terms of choice of antibiotic prescription or no prescription of antibiotic, dose, dosage regimen, and duration of therapy.
2. Agree with the use of antibiotic therapy because a potentially fatal infection of bacterial origin which will require antibiotic cannot be ruled out. Priority is given to saving life compared to antibiotic misuse.
3. Agree with the use of the antibiotic therapy but a different (usually less expensive and toxic) combination of therapy is preferred.
4. Agree with the use of antibiotic therapy but a modified dose, dosage regimen, and duration would be recommended.
5. Disagree with the use of antibiotic therapy; administration is unjustified or unnecessary use of drugs.

Category I and II were considered as “Appropriate” antibiotic therapy while category III, IV and V were considered as “Inappropriate” antibiotic therapy. Appropriateness of antibiotic prescription was defined as
the proportion of all antibiotics with specific diagnoses (Categories I, II, III, IV and V) categorized as appropriate (categories I and II).

Information on drug use training was obtained from in-service training reports using a data extraction tool. A checklist was used to assess the availability of STGs in the consulting rooms and medicines in the pharmacy stores.

Data management and analysis

Data were double entered into Epi Info version 7.2.2.6 and checked for completeness and discrepancies. A frequency table was used to present biodata of patients and professional category of prescribers. Frequencies were used to describe the types of antibiotics prescribed. Appropriateness of antibiotic prescriptions were estimated using proportions with 95% confidence Interval (CI). Chi-square test (or fisher’s exact test, where appropriate) was used to examine the association between antibiotic prescription and the independent variables. Crude odds ratio of the association between antibiotic prescription and the independent variables were estimated using univariable logistic regression. The multivariable logistic regression model was fitted using backward elimination. All variables with p<0.20 in the univariable model were put together in the multivariable regression model. Using a cut-off of p<0.20 proved useful as some variables which were not significant at p<0.05 in the univariable model turned out to be significant in the multivariable model. From the analysis of collinearity, the highest variance inflation factors were 1.94 and 1.88 for ARTI and URTI, respectively. Robust standard errors were estimated using the “robust” option for all the models in Stata 14 (40). Statistical significance in the multivariable model was set at p<0.05. All analyses were done using Stata 14. Due to the large number of variables assessed, results are presented for only variables which were significant at p<0.05 in the multivariable model.

Results

Six hundred (600) patient folders with specific or non-specific RTI at the OPD of the Tema Polyclinic were sampled for the study. The largest proportion of the patients were children younger than 5 years (n = 252, 42.1%). The proportion of females was 57.3%. More than half (52.3%) of patients were seen by medical doctors while the rest (47.7%) were seen by physician assistants. A detailed description of background characteristics of patients is presented (Table 1).
Table 1
Background characteristics of patients

| Characteristics   | Frequency, N = 600 | %  |
|-------------------|-------------------|----|
| **Age (in years)**|                   |    |
| < 5               | 252               | 42.1|
| ≥ 5               | 347               | 57.9|
| **Sex**           |                   |    |
| Male              | 256               | 42.7|
| Female            | 344               | 57.3|
| **NHIS**          |                   |    |
| Not registered    | 157               | 29.5|
| Registered        | 376               | 70.5|
| **Occupation**    |                   |    |
| Pupil/student     | 156               | 26.0|
| Formal employee   | 16                | 2.7 |
| Informal employee | 72                | 12.0|
| Not known         | 356               | 59.3|

Of the 600 prescriptions assessed, 358 (59.7%) had at least one (1) antibiotic. The total number of antibiotics prescribed was 379. As shown in Table 2, majority of the prescribed antibiotics were penicillins (n = 270, 71.2%). The other classes of antibiotics prescribed were cephalosporins (n = 57, 15.0%), macrolides (n = 27, 7.1%), quinolones (n = 10, 2.6%), nitroimidazole (n = 8, 2.1%), and other less frequently prescribed classes (n = 7, 1.8%). The penicillins prescribed were amoxicillin, amoxicillin-clavulanic acid and flucloxacillin. Cefuroxime, ceftriaxone and cefixime were the cephalosporins prescribed, whereas azithromycin and erythromycin were the macrolides prescribed.
Table 2
Distribution of prescribed antibiotic class and age group

| Antibiotic class | Age Group (years) | < 5 | ≥ 5 | Total |
|------------------|-------------------|-----|-----|-------|
| Penicillins      |                   | 97  | 173 | 270   |
| Cephalosporins   |                   | 27  | 30  | 57    |
| Macrolides       |                   | 4   | 23  | 27    |
| Quinolones       |                   | 1   | 9   | 10    |
| Nitroimidazole   |                   | 2   | 6   | 8     |
| Others*          |                   | 3   | 4   | 7     |
| Total            |                   | 134 | 245 | 379   |

| Category | Frequency | % of Total Prescription | % of Prescriptions with specific diagnoses | Remarks |
|----------|-----------|-------------------------|------------------------------------------|---------|
| I        | 107       | 17.8                    | 56.0                                     | Appropriate |
| II       | 14        | 2.3                     | 7.3                                      | Appropriate |
| III      | 1         | 0.2                     | 0.5                                      | Inappropriate |
| IV       | 60        | 10.0                    | 31.4                                     | Inappropriate |
| V        | 9         | 1.5                     | 4.7                                      | Inappropriate |
| N/A      | 409       | 68.2                    | -                                        | -         |

N/A – cases which could not be evaluated for appropriateness because of non-specific diagnoses

The distribution of the categorization using Kunin's modified criteria for antibiotic use is provided (Table 3). The majority (68.2%, 409 out of 600) of the RTI cases could not be classified because the diagnoses were non-specific. Of the 31.8% (n = 191) of folders which had specific diagnosis indicated, 121 had appropriate antibiotic prescription (63.3%, 95% CI: 56.2, 69.9). Regarding appropriateness in respect of choice of antibiotic, dose, and duration of therapy, 68.8% of inappropriate antibiotic prescription was because of wrong duration of therapy while 23.4% resulted from improper dose.

Table 3
Categorization of antibiotic prescription appropriateness using Kunin's modified criteria

| Category | Frequency | % of Total Prescription | % of Prescriptions with specific diagnoses | Remarks |
|----------|-----------|-------------------------|------------------------------------------|---------|
| I        | 107       | 17.8                    | 56.0                                     | Appropriate |
| II       | 14        | 2.3                     | 7.3                                      | Appropriate |
| III      | 1         | 0.2                     | 0.5                                      | Inappropriate |
| IV       | 60        | 10.0                    | 31.4                                     | Inappropriate |
| V        | 9         | 1.5                     | 4.7                                      | Inappropriate |
| N/A      | 409       | 68.2                    | -                                        | -         |

N/A – cases which could not be evaluated for appropriateness because of non-specific diagnoses

The proportion of patients who were prescribed at least one (1) antibiotic was 59.7% (95% CI: 55.7, 63.5) (Table 4). The frequency of antibiotic prescription did not differ significantly between males and females (p = 0.644). Based on age group the proportion of antibiotics prescribed was higher among those aged 5 years or older compared to those younger than 5 years. The frequency of prescribing antibiotics by Physician
Assistants was higher compared to Medical Doctors (65.4% vs 54.5%, p = 0.006). Four (4) training / meeting sessions on drug-use were organized in 2018, while fifteen (15) copies of the 2017 Edition of the STG were purchased. However, none of the consulting rooms where prescribers attended to patients had copies available during the data collection period.

Table 4
Bivariate analysis of patient and prescriber factors by antibiotics prescription

| Antibiotics | Not Prescribed n (%) | Prescribed n (%) | Total, N = 600 | Chi square | p-value |
|-------------|----------------------|------------------|---------------|------------|---------|
| Patient factors |                      |                  |               |            |         |
| Age-group (in years) |                  |                  |               | 17.255     | < 0.001* |
| < 5         | 126 (50.0)           | 126 (50.0)       | 252 (100.0)   |            |         |
| ≥ 5         | 115 (33.1)           | 232 (66.9)       | 347 (100.0)   |            |         |
| Sex         |                      |                  |               | 0.214      | 0.644   |
| Male        | 106 (41.4)           | 150 (58.6)       | 256 (100.0)   |            |         |
| Female      | 136 (39.5)           | 208 (60.5)       | 344 (100.0)   |            |         |
| NHIS        |                      |                  |               | 0.600      | 0.438   |
| No          | 57 (36.3)            | 100 (63.7)       | 157 (100.0)   |            |         |
| Yes         | 150 (39.9)           | 226 (60.1)       | 376 (100.0)   |            |         |
| Occupation  |                      |                  |               | 9.038      | 0.029*  |
| Pupil/Student | 56 (35.9)           | 100 (64.1)       | 156 (100.0)   |            |         |
| Formal employee | 8 (50.0)            | 8 (50.0)         | 16 (100.0)    |            |         |
| Informal employee | 20 (27.8)     | 52 (72.2)        | 72 (100.0)    |            |         |
| Not known   | 158 (44.4)           | 198 (55.6)       | 356 (100.0)   |            |         |
| Prescriber Factor |                 |                  |               | 7.425      | 0.006*  |
| Prescriber's category |             |                  |               |            |         |
| Medical Officer | 143 (45.5)          | 171 (54.5)       | 314 (100.0)   |            |         |
| Physician Assistant | 99 (34.6)          | 187 (65.4)       | 286 (100.0)   |            |         |
| Total       | 242 (40.3)           | 358 (59.7)       | 600 (100.0)   |            |         |

*statistically significant association relative to p < 0.05
Other diagnoses made which were not related to RTI included malaria, allergies, dysentery, hypertension, enteric fever among others. The average number of drugs prescribed per prescription was 3.59. Drugs prescribed other than antibiotics included analgesics, haematinics, cough remedies, anti-hypertensives and anti-allergic drugs.

The variables which met the cut-off criterion of $p < 0.20$ in the crude analyses were age group, laboratory investigations, breathlessness, occupation, professional category of prescriber, chest pain, headache, rhinorrhea, sore throat, sneezing, catarrh, sputum production, otitis media, ARTI, common cold, pharyngitis, pneumonia, rhinitis, URTI, tonsillitis, sinusitis and nasal congestion. Of the 19 people diagnosed with pneumonia, all were not prescribed with antibiotics. Similarly, all 38 tonsillitis cases were not prescribed with antibiotics. These 2 diagnoses were not included in the regression analyses because each did not vary with regards to antibiotic prescription.

The variables which were significant in the adjusted analysis are presented in Table 5. Requesting laboratory investigations, presenting with breathlessness, headache and sputum production were associated with increased odds of antibiotic prescription in the adjusted model. Cases for which laboratory investigations were requested were 80.0% more likely to be prescribed with antibiotics after controlling for other significant variables.

The factors which were associated with reduced odds of antibiotic prescription were presenting with catarrh, a diagnosis of rhinitis, ARTI, URTI and common cold. After accounting for confounding factors, cases presenting with catarrh were 58.0% less likely to be prescribed with antibiotics ($AOR = 0.42, 95\% CI: 0.27–0.66, p < 0.001$). Those diagnosed with common cold had 87.0% lower odds of antibiotic prescription, after adjusting for factors of interest ($AOR = 0.13, 95\% CI: 0.05–0.31, p < 0.001$). While in the crude model ARTI was associated with 1.9 times increased odds of antibiotic prescription ($AOR = 1.92, 95\% CI: 1.37–2.68, p < 0.001$), in the adjusted model it was associated with 59% reduced odds of antibiotics prescription ($AOR = 0.41, 95\% CI: 0.23–0.74, p = 0.003$).
## Table 5

Crude and Adjusted Odds Ratios of Factors Associated with Antibiotic Prescription

| Variables                  | Crude OR (95% CI) | P-value | Adjusted OR (95% CI) | P-value |
|----------------------------|-------------------|---------|----------------------|---------|
| **Laboratory investigation (Ref: No)** |                   |         |                      |         |
| Yes                        | 1.98 (1.42, 2.76) | < 0.001 | 1.80 (1.22, 2.64)    | 0.003   |
| **Breathlessness (Ref: No)** |                   |         |                      |         |
| Yes                        | 6.93 (0.88, 54.45)| 0.066   | 9.01 (1.11, 73.09)   | 0.04    |
| **Catarrh (Ref: No)**      |                   |         |                      |         |
| Yes                        | 0.50 (0.33, 0.77) | < 0.001 | 0.42 (0.27, 0.66)    | < 0.001 |
| **Headache (Ref: No)**     |                   |         |                      |         |
| Yes                        | 2.09 (1.47, 2.99) | < 0.001 | 1.77 (1.16, 2.71)    | 0.009   |
| **Sputum (Ref: No)**       |                   |         |                      |         |
| Yes                        | 3.36 (2.04, 5.53) | < 0.001 | 3.49 (1.96, 6.22)    | < 0.001 |
| **Rhinitis (Ref: No)**     |                   |         |                      |         |
| Yes                        | 0.40 (0.14, 1.10) | 0.076   | 0.20 (0.06, 0.73)    | 0.015   |
| **ARTI (Ref: No)**         |                   |         |                      |         |
| Yes                        | 1.92 (1.37, 2.68) | < 0.001 | 0.41 (0.23, 0.74)    | 0.003   |
| **URTI (Ref: No)**         |                   |         |                      |         |
| Yes                        | 0.25 (0.18, 0.37) | < 0.001 | 0.11 (0.06, 0.20)    | < 0.001 |
| **Common cold (Ref: No)** |                   |         |                      |         |
| Yes                        | 0.23 (0.12, 0.43) | < 0.001 | 0.13 (0.05, 0.31)    | < 0.001 |

**ABBREVIATION:** ARTI, Acute Respiratory Tract Infection; CI, Confidence Interval; OR, Odds Ratio; URTI, Upper Respiratory Tract Infection

## Discussion

This study presents the outcome of audit of prescriptions of antibiotics in the management of RTIs at Tema Polyclinic (a primary healthcare facility) in Ghana. It also sought to identify the factors that are associated with such prescriptions.

### Patient Demographics

The highest proportion of patients were less than 5 years old (42.1%). This is consistent with the general trend identified in several studies that have identified higher incidence of RTIs in children, especially those...
below the age of 5 years (1,6,7). This is often attributed to their immature respiratory tissues and organs to adequately deal with such infections. It also point to the need to pay particular attention to this age group in the implementation of interventions aimed at controlling and managing RTIs.

Female patients with RTIs were more compared to males. This may be attributed to the general view that females have better health-seeking behaviour compared to males (41). It may also be due to the fact that the incidence of RTIs are higher in females than in males (42).

Most of the patients were NHIS card holders (70.5%). This can be attributed to the fact that Tema Polyclinic is the major public health facility in the municipality that offers full service (i.e. consultation and medicines) to NHIS clients.

**Prescription of Antibiotics**

More than half (59.7%) of the RTI patients in the present study received at least one antibiotic. This is comparable to the 59.0% antibiotic prescription in a study in NANTES University hospital (43) and 63% from the National Ambulatory Medical Care Survey (44). Conversely, our estimate is lower than 67.5% and 82.5%, obtained in the study at Ridge Hospital and Adabraka Polyclinic, Accra, Ghana, respectively (18). It must be noted that Apenteng and colleagues’ study was conducted among children under less than 5 years. Compared to antibiotic prescription among the children under five in our study, our estimate of 50.0% was lower. Since most RTIs are of viral origin (4,11), the high proportion of antibiotic prescription use in the management of RTI needs to be addressed.

The three topmost classes of antibiotics from which prescriptions were made for RTI cases in the study were penicillins, cephalosporins and macrolides. Other studies have also identified same for the management of RTIs (45,46). These are also the 1\textsuperscript{st} and 2\textsuperscript{nd} line options recommended by the STG when an antibiotic is indicated for a specific RTI management. The pattern is also similar to that in the study by Sumaila and Tabong (2018) where the percentage prescription of penicillins, cephalosporins and macrolides for URTI management were 58%, 19% and 10% respectively. In addition, similar specific antibiotics within these classes were identified in both studies with amoxicillin, amoxicillin + clavulanic acid and cefuroxime as the most prescribed, except that the latter was in URTIs management. The STG recommends amoxicillin as the first line antibiotic in most cases of specified RTIs where it is indicated with amoxicillin + clavulanic acid and cefuroxime as the second line options. The choice of specific agents for RTI management is therefore consistent with the STG requirements.

**Appropriateness of Antibiotic Prescription**

The proportion of patients appropriately prescribed antibiotics, from the study, was (63.3%, 95% CI: 56.2, 69.9). This is comparable to the 70.4% found at a secondary referral hospital in Ghana but at variance with the 34.6% found at a primary referral hospital also in Ghana (18). Unlike the present study, the other studies were conducted in children under 5 years. Most of the inappropriateness resulted from inappropriate duration of antibiotic use. A longer duration than is recommended may mean exposing bacteria excessively to antibiotics thereby facilitating the emergence of resistant strains. On the other hand, a shorter duration
may imply a higher chance of treatment failure. This may necessitate retreatment leading to a cumulatively longer duration of bacterial exposure (47). Further studies to describe inappropriate duration of antibiotic use and to ascertain the associated factors are required.

**Associated factors with antibiotic prescription**

The treatment recommendation for RTIs by the STG like all other disease conditions, are based on specific diagnosis. This prevents blind treatment and ensures rational use of medicines. The majority (68.2%) of the RTI cases from this study could not be classified because they were not specific. The 31.8% with specific RTI diagnoses observed in this study is far below 67.5% and 65.0% obtained in the study at Ridge hospital and Adabraka polyclinic, respectively (18). This raises issues with clinical diagnosing of RTIs at Tema polyclinic and may point to the fact that prescribers are more comfortable making a broad diagnosis of RTI or URTI instead of distinguishing the types. This also gives room for empirical use of antibiotics with its public health implications. With the NHIS using the STG recommendations in vetting claims, the facility stands to lose revenue and that has implications for quality and sustainability of service provision. It is particularly worrying since most of the patients access care through their NHIS membership.

In this study, all patients diagnosed with pneumonia or tonsillitis were not prescribed with antibiotics suggesting that prescribers may be thinking that these infections were not caused by bacteria. This position is corroborated by the finding that all the significant diagnoses were associated with reduced odds of antibiotic prescription. Contrary to the finding that in different geographic locations bronchitis is consistently associated with higher odds of antibiotic prescription, our study did not find enough evidence to support this (31). As have been reported, uncomplicated bronchitis does not require treatment with antibiotics (48). Our findings suggest that in line with this, prescribers do not suspect any underlying bacterial infection when treating these conditions.

A study in the USA found sputum and breathlessness to be associated with antibiotic prescription (32). These observations are consistent with findings from our study. Coloured sputum production have also been observed to be associated with antibiotic prescription in a multi-country study too (33). The prescribers may be perceiving these symptoms as being indicative of bacterial infection. Since evidence from the literature does not support this antibiotic prescribing behaviour, it must be addressed to help control high and irrational prescribing of antibiotics.

Our findings that laboratory investigations is linked to high antibiotic prescription is comparable to a study in Switzerland (30). In the latter study, elevated white blood cell count was associated with high antibiotic prescription. In a related study in Uganda, the investigators found that patients who underwent laboratory investigations and tested negative for malaria were more likely to be prescribed antibiotics (29). Furthermore, among those who did not undergo laboratory testing, diagnostic uncertainty is likely to predispose them to higher antibiotic prescription. In a population where the white cell count and negative malaria testing results may be relatively high, as it may be among those who underwent diagnostic testing in our study, higher antibiotic prescription may be expected. These findings may explain the observation that laboratory testing was significantly associated with antibiotic prescription in this study.
One strength of our study is that it narrows down to RTIs which have been shown to be the most common infection treated with antibiotics by several studies. Regarding limitations, the factors assessed were limited by the records review design. Also, we could not verify if recorded diagnoses by prescribers were wholly consistent with the international classification of diseases.

**Conclusion**

Although many patients with specific diagnoses were prescribed antibiotics appropriately, some RTI cases received prescriptions of antibiotics with inadequate dosage regimen. Majority of RTI cases were prescribed antibiotics. Commonly prescribed antibiotics were amoxicillin, amoxicillin plus clavulanic acid (penicillins), cefuroxime (cephalosporin) and azithromycin (macrolide). Factors influencing high antibiotic prescription were presenting with sputum production, breathlessness, headache and having undergone laboratory investigations.

Antibiotic stewardship efforts should focus on imparting knowledge on specific diagnoses and corresponding treatment with emphasis on dosage regimen while deemphasizing symptomatic treatment based on sputum production.

**List Of Abbreviations**

ARTI Acute Respiratory Tract Infection  
DTC Drugs and Therapeutics Committee  
HIV Human Immunodeficiency Virus  
NHIS National Health Insurance Scheme  
OPD Outpatient Department  
RTI Respiratory Tract Infection  
STG Standard Treatment Guidelines  
TB Tuberculosis  
URTI Upper Respiratory Tract Infection

**Declarations**

**Ethics Approval and Consent to Participate**

All the included folders were de-identified to preserve the privacy of patients and prescribers. Ethical clearance was obtained from the Ethics Review Committee of the Ghana Health Service. In addition, the Medical superintendent of the Polyclinic granted permission for the study to be conducted.
Consent for publication

Consent for publication was obtained from the Management of Tema Polyclinic.

Availability of Data and Material

The dataset is not publicly available. However, it can be accessed after a reasonable request has been sent to the corresponding author.

Competing Interests

The authors declare that they have no competing interests.

Funding

Funding for the original study was provided by Joel Jeffrey Idun-Acquah.

Authors' Contributions

JJIA conceptualised and designed this research which formed a component of his training requirements for the award of Fellowship at the West African Postgraduate College of Pharmacists. HAB supervised the entire research project. MM and EK contributed to data management, analyses and interpretation. JJIA and MM produced the first draft of the manuscript. All authors reviewed and approved the final manuscript for submission.

Acknowledgements

We would like to acknowledge Dr. Sally D. Quartey and the Drugs and Therapeutics Committee of Tema Polyclinic for their technical advice and support throughout the data collection period. Special thanks to Miss Irene A. Boateng and Mr. Edwin Martey for assisting with data collection. We are grateful to Dr. Adolf Awua, and Mr. Daniel Afriyie for reviewing and editing the manuscript.

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