Delays in diagnosis and treatment of pulmonary tuberculosis in Wakiso and Mukono districts, Uganda

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Abstract

Background: Delay in tuberculosis (TB) diagnosis may worsen the disease and increase TB transmission. Therefore, timely diagnosis and treatment is critical in TB control. We aimed to assess the treatment delay of pulmonary TB and its determinants in two Ugandan districts where TB infection control (TBIC) guidelines were formerly implemented.

Methods: A facility based cross-sectional study was conducted in Mukono and Wakiso districts. Adult pulmonary TB patients within three months of initiating treatment were included in the study. Delays were categorized into unacceptable patient delay (more than 3 weeks from the onset of cough and the first consultation with a health care provider), health service (more than one week from the first consultation to the initiation of TB treatment) and total delay (more than 4 weeks since the onset of cough). The prevalences as well as predictors for the three delays were determined.

Results: We enrolled 158 sputum positive patients. Unacceptable patient delay was noted in 91 (58%) patients, a health service delay in 140 (88%) patients and a total delay in 140 (90%) patients. An independent predictor for patient delay was male gender (p < 0.001). First visiting a non-public health facility (p = 0.001) was an independent predictor of health service delay.

Conclusion: There is still a significant TB diagnosis and treatment delay in Uganda. Most of the delay was caused by health system delay in the non-public health care sector. There is need for TB advocacy in the community, training of health workers in TBIC and strengthening public-private partnerships in TB control.

Keywords: TB treatment delay, Patient delay, Health service delay, TB infection control, Uganda

Background

Uganda is among the 22 high tuberculosis (TB) burden countries. It has a treatment success of 67%, default rate of 11% and 50% of TB patients are HIV co-infected [1]. Multi-drug resistant TB (MDR-TB) poses a problem in 1.1% of new cases and 12% of retreatment cases in 2010 [1,2]. One of the contributors to these poor indicators is due to the prolonged delay from the onset of TB disease to the time of diagnosis and initiation of treatment. Delay in diagnosis may lead to progression of disease, increased mortality and TB transmission in the community [3,4]. Therefore, timely diagnosis and treatment is critical in TB control. Studies from Uganda and Tanzania have shown median diagnostic delays varying from 8 to 13 weeks [5-7]). In two systematic reviews, it was found that diagnosis and treatment delays of TB occurs more often in patients with smear negative TB, HIV infection, living in a rural area and or far away from a health unit and in patients consulting with traditional healers [8,9].

In 2009, WHO released TB infection control (TBIC) guidelines [10]. Early detection and prompt effective TB treatment are the hallmark of a successful TBIC. These guidelines recommend minimizing the time spent in health facilities, by ensuring that people with TB symptoms are rapidly identified and if found to have TB, promptly treated. Exposure to people who are infectious
should be minimized by reducing the number of outpatient visits [10]. Furthermore, encouraging people to rapidly seek care if they have symptoms suggestive of TB [10]. In 2008, the Uganda Ministry of Health (MOH) in collaboration with the TB Union, through support from the Tuberculosis Control Assistance Programme implemented a two year TBIC project in 12 districts in Uganda including Mukono district. In 2010, the Ugandan MOH implemented TBIC in Wakiso district and in 2011 released national TBIC guidelines, adopted from the WHO [11]. To our knowledge no other study in Uganda has examined TB treatment delays since the implementation of TBIC. Our study assessed the magnitude of patient, health service and total delay; and their determinants in a predominantly rural setting in Uganda.

Methods
Design and study setting
A cross-sectional survey among patients with pulmonary TB was conducted in Mukono and Wakiso districts. These two central districts surrounding the capital Kampala are partly semi-urban but predominantly rural [12]. The reported case detection rates of new pulmonary positive TB in Mukono and Wakiso districts were 40.1% and 38.5%, respectively in 2008. Mukono has 25 and Wakiso 26 healthcare facilities currently providing TB services. The study was conducted among patients attending selected health centre IV and hospital level facilities; government and one Private-Not-For Profit (PNFPs) facilities. These facilities were purposely selected because they handle most of the TB patients in their geographical locations. In Uganda, TB services are provided at health centre III (sub-county level) and above. TB services are, at least in principle, free of charge in all government facilities, while in PNFPs, patients pay for consultation and investigations, but after diagnosis the TB drugs are provided for free. Diagnosis of pulmonary TB is by smear microscopy. According to TBIC guidelines, patients presenting to the facility with cough of two or more weeks have to submit two sputum samples in the form of a spot-sample and morning sample the following day. Spot specimens are collected on the day the patient suspected to have TB is seen at the facility, morning samples are collected early in the morning of the second day [13]. A patient is diagnosed with TB if one sputum smear is positive. Treatment should be initiated on the same day after receipt of a positive result, ideally within 24 hours after submission of the first sputum specimen [11]. TB/HIV collaborative activities exist in the study districts and so all HIV patients are supposed to be screened for TB and TB patients counselled and tested for HIV.

Study participant selection
We aimed to recruit patients 18 years and above, who started treatment in the 3 months preceding the start of the survey. Eligible participants were asked to participate in the study as they visited the TB clinics for drug refill. Patients were consecutively enrolled and interviewed using a structured questionnaire. Patients previously treated for TB and those with extra-pulmonary TB were excluded from the study. Only sputum smear positive patients were included in the study. All of them were out-patients.

An interviewer-administered structured questionnaire was used, adapted from a previous study on TB diagnosis delay conducted in public primary health facilities in Kampala city, Uganda [7]. The questionnaire was translated to Luganda. The following information was collected: demographic and socio-economic characteristics, TB symptoms and duration in number of weeks before seeking care outside home, the date of patient’s first consultation with a health care provider and any other consultations with subsequent providers, if any, before TB treatment was started, distance to the health facility where TB diagnosis was made, and date of starting treatment. In addition patients were asked whether TB is curable and if they had tested for HIV before being diagnosed of TB. Participants were enrolled between December 2012 and March 2013.

Definitions and measures
Patient delay was defined in a similar way as in previous studies in Uganda, as the time interval in weeks between onset of cough and the first consultation with any health care provider [5-7]. A health care provider was defined as any person consulted by the patient about his/her sickness who prescribed any form of medication (Table 1). These included dispensers, pharmacists, medical staff and herbalists/traditional healers. Unacceptable patient delay was defined as a delay of more than 3 weeks since the onset of cough (i.e. one week after the 2 weeks of cough when one is considered a TB suspect) [14,15]. Health service delay was defined as the time in weeks from the first consultation to the initiation of the TB treatment. Unacceptable health service delay was defined as a delay of more than 1 week. The total delay was the sum of both, patient and health service delay [6,7]. Unacceptable total delay was defined as a delay of more than 4 weeks [5,14]. Cough was chosen to define delay because it is the cardinal symptom of pulmonary TB. The estimated sample size of 158 patients was obtained assuming, from prior local study that 91% would have a total delay of more than 4 weeks [5]. This took into consideration a power of 80%,
95% confidence interval and a 20% adjustment for missing and non response.

**Quality control**

The questionnaire used was pre-tested. Research assistants were trained and supervised during data collection. Duration of symptoms, date of initiation of treatment were counterchecked in the medical records. Data was double entered into a computer, and the two copies of the data verified.

**Ethical considerations**

The research protocol was approved by the institutional review board of the Makerere University School of Public

### Table 1 Patient, health service and total delays in published studies and current study

| Reference | Definitions used | Health service/facility delay | Total delay = Patient + health facility delay | Prevalence of delay | Median/mean total delay |
|-----------|------------------|------------------------------|---------------------------------------------|---------------------|------------------------|
| Kiwuwa et al. (2005) Uganda [5] | - The time interval between symptom onset and the first medical consultation. - A health provider was defined as any person consulted by the patient about his/her sickness, who prescribed any form of medication. These included dispensers, pharmacists, medical staff and herbalists or traditional healers. - Interval of > 2 weeks was considered as long patient delay. | - The time taken from the first medical consultation to when the diagnosis was confirmed and treatment started. - Health service delay was when there was a delay of > 4 weeks between the initial contact with the health provider and the start of TB treatment. | > 4 weeks was considered as prolonged/ unacceptable total delay. | Patient delay: not specified | Total delay: 91% |
| Ngadaya et al. (2009) [6] Tanzania | - The time interval between the day of experiencing for the first time one of the pulmonary symptoms to the day the patient sought medical advice for the first time. - Interval of > 30 days was considered as patient delay. | - A time interval between first consultations at a health facility to the day the treatment was initiated. - A time interval > 35 days. | The sum of the patient and health facility delay. | Health facility: 59%. Patient and total delay not specified. | Mean: 125.5 (SD98.5) days |
| Basnet et al. (2009) Nepal [16] | - The time interval from the appearance of the first symptoms of tuberculosis until the first visit to any formal health care facility (health centres, hospitals or DOTS centres). - Interval of > 30 days was considered as patient delay. | - The time interval from the first consultation at any formal health facility until the date of diagnosis. | Not specified. | Median: 60 days, IQR not specified |
| Sendagire et al. (2010) [7] | - The time in weeks from the onset of cough to a first consultation with any health care provider. - Prolonged patient delay was defined as a delay of > 8 weeks. | - The time in weeks from the first consultation to initiation of treatment. - Prolonged health facility was defined as a delay of > 6 weeks. | Unacceptable total delay was defined as a delay of > 14 weeks. | Prolonged patient delay: 19% | Median: 8 (IQR 4-12) weeks |
| Current study | - The time interval in weeks between onset of cough and the first consultation with any health care provider (dispensers, pharmacists, medical staff and herbalists/traditional healers). | - The time in weeks from the first consultation to the initiation of the TB treatment. | Unacceptable total delay was defined as > 4 weeks. | Unacceptable patient delay: 58% | Median total delay of 16 (IQR 9-30) weeks |

≠IQR = Interquartile range.
*SD = Standard deviation.
Health and the Uganda National Council for Science and Technology (Ref.nr. HS 880). Permission from the in-
charges of the health facilities was obtained. Informed
written consent was obtained from the participants at the
time of data collection. Respondents were assured of
confidentiality.

Data management and analysis
Data were entered into Epi-info software version 6 and
exported to Stata software version 10 for analysis. Health
service delay (in days) was computed from the date of
first seeking care with cough symptom to when treat-
ment was initiated. It was later converted into weeks by
diving by 7. Descriptive statistics were computed, such
as frequency distributions, median and interquartile
ranges (IQRs). The normality of the delays was checked
using Shapiro test and were all found not be normally
distributed. Bivariate analysis using Chi-square test and
prevalence ratios (using generalized linear model with a
Poisson link) were performed. Multivariate analysis was
used to explore the predictors of the outcome variables
(patient, health service and total delay). Unacceptable
patient delay, health service delay and total delay were
compared for different sub-groups using prevalence ra-
tios and 95% CI. To identify factors independently asso-
ciated with unacceptable patient, health service and total
delays, a multivariate analysis with delay time dichoto-
mized was performed. Possible predictors (age, sex, resi-
dence, type of health provider first visited, education
level and distance to the facility of diagnosis) were eval-
uated in bivariate associations with each type of delay.
Variables associated with delays in the bivariate analysis
(p ≤ 0.2) were included in the model. Statistical signifi-
cance was taken as p < 0.05. We adjusted for the cluster-
ing effect at both bivariate and multivariate analysis.

Results
A total of 229 TB patients were prospectively enrolled
from 6 health facilities (3 hospitals and 3 health centre
IVs). Seventy one patients were excluded from further
analysis because they were sputum negative (59) and did
not have cough as one of their presenting symptoms
(12), leaving 158 patients (Figure 1). The median age
was 30 (IQR 25.5-38) years. Ninety six (61%) patients
were males, 78 (49%) were married and 13 (8%) had no
formal education, (Table 2). Patients’ median family size
was 4 (IQR3-7) and they were generally living in one liv-
ing room. More than half (102; 65%) of the patients lived
less than 60 minutes walking distance to the facility
where the TB was diagnosed. The majority of the pa-
tients, 104/157 (65%) had ever tested for HIV before TB
diagnosis (Table 2). One hundred forty three (97%)

![Figure 1 Flowchart showing the patients included in the analysis of treatment delay in Mukono and Wakiso districts, Uganda.](http://www.biomedcentral.com/1471-2458/14/586)
respondents accepted to share their results, with 31% (45/143) reporting being HIV positive. In addition to cough, at the time of diagnosis, patients had the following symptoms: chest pain 94/157 (60%), fever 85/157 (54%), weight loss 80/157 (51%), loss of appetite 67/156 (43%) and haemoptysis 28/158 (18%).

Health care seeking behaviour
Only 35/158 (22%) patients first sought care from a public health facility. Patients who first sought care from non-public health facilities were similar to those who first visited a public health facility regarding sex, marital status, residence and having been tested for HIV before TB diagnosis. Patients of different age category and education level tended to seek care differently. Non-public health facilities were visited by 123 (78%) of the 158 patients: private clinics by 65 patients (41%), drug shops by 17 patients (11%), a private pharmacy by 16 patients (10%), traditional healers/herbalists by 15 patients (9%) and PNFPs by 10 patients (6%). Reasons for not seeking care from public health facilities were available for 119 patients. These included: long waiting hours in 33/119 (28%), the facility being far away in 35/119 (29%), a bad previous experience in 20/119 (17%) and thinking that their illness was not that serious 31/119 (26%). Majority of the respondents (143/158 (91%)), made the first consultation because of cough. The services provided by the health care provider during this first visit were: offering non-TB medical treatment in 128/156 (82%) patients, sputum examination in 24/156 (15%) patients, x-ray in 2/156 (1%) patients, and referring the remaining patients.

Delays and determinants
Patient delay
The median patient delay was 4 (IQR 2-8) weeks. In 91 respondents (58%) there was unacceptable patient delay. The majority of the patients (81/91) who delayed seeking care, thought that their symptoms would spontaneously go away, while 11/91 (9%) did not perceive a time span of more than 3 weeks as problematic. The remaining patients delayed to seek care due to economic constraints and lack of family support. In our bivariate analysis, patient delay was significantly associated with being male (p < 0.001), but not with age, education level, salary, distance to the facility of diagnosis, residence, marital status or having tested for HIV before the current TB diagnosis (Table 3). In our multivariate analysis, after controlling for age, sex, education level, and marital status, male patients were more likely to delay seeking care (adjusted Prevalence ratio [aPR] 1.61(1.43-1.82).

Health service delay
The median reported health system delay was 11 (5-24) weeks. In the majority of the patients 140/158 (89%) there was an unacceptable health service delay. The median number of visits to all sorts of health care providers before being diagnosed was 3 (IQR3-9). Almost half of the patients who delayed 60/140 (43%), lived beyond 5 km from the facility where diagnosis was made (Table 4). In our bivariate analysis, health service delay was significantly associated with first seeking care in a non-public health facility (p < 0.001). In multivariate analysis, after controlling for age, sex, distance to the facility of diagnosis, education level and type of facility visited first, patients who first sought care in the public health facility were significantly less likely to have unacceptable health service delay, (aPR 0.62; 0.47-0.83).

Table 2 Socio-demographic and clinical characteristics of the study patients

| Variable                                      | Total n (%) |
|-----------------------------------------------|-------------|
| **Sex (n = 158)**                             |             |
| Female                                        | 62 (39)     |
| Male                                          | 96 (61)     |
| **Age years (n = 158)**                       |             |
| 15-24                                         | 36 (23)     |
| 25-34                                         | 65 (41)     |
| 35-44                                         | 38 (24)     |
| >44                                          | 19 (12)     |
| **Monthly salary income in Ugandan shillings (n = 156)** |         |
| ≤20,000                                       | 39 (25)     |
| >20,000                                       | 117 (75)    |
| **Cost for one way journey to the facility of diagnosis (n = 158)** |       |
| ≤2,000                                        | 104 (66)    |
| >2,000                                        | 54 (34)     |
| **Marital status (n = 158)**                  |             |
| Married                                       | 78 (49)     |
| Single                                        | 38 (24)     |
| Separated/widowed                             | 42 (27)     |
| **Residence (n = 158)**                       |             |
| Rural                                         | 57 (36)     |
| Semi-urban                                    | 101 (64)    |
| **Education level (n = 158)**                 |             |
| No formal education                           | 13 (8)      |
| Primary                                       | 73 (46)     |
| Secondary and above                           | 72 (46)     |
| **Patient knows TB can be cured (n = 158)**    |             |
| Yes                                           | 82 (52)     |
| No                                            | 76 (48)     |
| **Tested for HIV before TB diagnosis (n = 158)** |           |
| Yes                                           | 104 (66)    |
| No                                            | 54 (34)     |

Table 2 Socio-demographic and clinical characteristics of the study patients
Table 3 Predictors of unacceptable patient delay (>3 weeks)

| Variable                | Unacceptable patient delay | Univariate analysis | Multivariate analysis | P       |
|-------------------------|-----------------------------|---------------------|-----------------------|---------|
|                         | Yes                         | No                  | Crude PR 95% CI       | Adjusted PR 95% CI |
| **Sex**                 |                             |                     |                       |         |
| Female                  | 26/62 (42)                  | 36/62 (58)          | 1                     | 1       |
| Male                    | 65/94 (69)                  | 29/94 (31)          | 1.64 (1.41-1.91)      | 1.61 (1.43-1.82) | <0.001* |
| **Age group**           |                             |                     |                       |         |
| 15-34                   | 52/100 (52)                 | 48/100 (48)         | 1                     | 1       |
| 35 & above              | 39/56 (70)                  | 17/56 (30)          | 1.33 (0.98-1.82)      | 1.00 (0.85-1.18) | 0.97    |
| **Education level**     |                             |                     |                       |         |
| No formal education     | 9/13 (69)                   | 4/13 (31)           | 1                     | 1       |
| Primary                 | 49/72 (68)                  | 23/72 (32)          | 0.98 (0.63-1.52)      | 0.92 (0.66-1.27) | 0.61    |
| Secondary & above       | 33/71 (46)                  | 38/71 (54)          | 0.67 (0.42-1.06)      | 0.70 (0.45-1.09) | 0.12    |
| **Income in Ugandan shillings** |                     |                     |                       |         |
| 0-20,000                | 21/38 (55)                  | 17/38 (45)          | 1                     | -       |
| >20,000                 | 69/116 (59)                 | 47/116 (59)         | 1.07 (0.85-1.34)      | -       |
| **Distance to facility of diagnosis** |                     |                     |                       |         |
| Within 5km              | 48/85 (56)                  | 37/85 (44)          | 1                     | -       |
| Over 5Km                | 43/71 (61)                  | 28/71 (39)          | 1.07 (0.83-1.38)      | -       |
| **Type of facility first visited** |                     |                     |                       |         |
| Non-Public/private      | 69/122 (57)                 | 53/122 (43)         | 1                     | -       |
| Public                  | 22/34 (65)                  | 12/34 (35)          | 1.14 (0.83-1.57)      | -       |
| **Patient knows TB can be cured** |                     |                     |                       |         |
| No                      | 48/81 (59)                  | 33/81 (41)          | 1                     | -       |
| Yes                     | 43/75 (57)                  | 32/75 (43)          | 0.96 (0.86-1.08)      | -       |
| **Tested for HIV before** |                             |                     |                       |         |
| Yes                     | 59/102 (58)                 | 43/102 (42)         | 1                     | -       |
| No                      | 32/54 (59)                  | 22/54 (41)          | 1.02 (0.81-1.29)      | -       |
| **Residence**           |                             |                     |                       |         |
| Rural                   | 35/57 (61)                  | 22/57 (39)          | 1                     | -       |
| Semi-urban              | 56/99 (57)                  | 43/99 (43)          | 0.92 (0.68-1.23)      | -       |

*Statistically significant.

Total delay

Data for total delay was available for only 154 patients. The median total delay from onset of cough until start of anti-TB treatment was 16 (9-30) weeks. In the majority of the patients 140/154 (91%) there was an unacceptable total delay. In a chi-square test unacceptable total delay was significantly associated with first visiting a non-public health facility (p = 0.05) (Table 5). Health service delay was the greatest contributor (75%) to total delay. In our multivariate analysis being male and first seeking care from non-public health facility were more likely to have unacceptable total delay, though not significant (Table 5).

Discussion

This study performed in two Ugandan districts highlights the unacceptable delay from onset of cough to initiation of TB treatment. More than half of the patients delayed to seek medical care in the presence of cough for more than 3 weeks. Health service delay was the biggest contributor to the total delay. Patient delay was significantly associated with being male. Health service delay was significantly associated with first seeking care from a non-public health facility.

Three quarters of the total delay was due to health service delay. This proportion of the total delay is similar to what was found in a study among patients at the national referral hospital in Kampala [5]. Sendagire et al., on the other hand, in a study in Kampala City Councils clinics observed that patient and health service delay equally contributed to the delay [7]. The long health service delay in our study shows that there are many missed chances for TB diagnosis. While according to the Ugandan National
TBIC guidelines, a patient with cough for 2 weeks or more should be investigated for TB [11], in our study only 15% of such patients had a sputum examination done on their first visit to the health care provider. The unacceptable health service delay is an indication of a weak performing health system. This may be because of poor knowledge of health workers about TB, resulting into a low index of suspicion for TB, and/or lack of staff, infrastructure to diagnose and treat TB. This long health service delay means that a significant number of patients remain infectious. This puts other patients in the health facilities and health care workers (particularly HIV infected) at risk of getting nosocomial TB, in addition to transmitting TB to the household members. The finding of unacceptable health service delay being associated with first seeking care from non-public health facilities (non-TB health care providers) is similar to what has been reported in other studies [9,17]. Only public health facilities and a few private providers provide TB treatment in Uganda [7]. Therefore, non-public health facilities like pharmacies, drug shops, some private clinics and traditional healers/herbalist which do not regularly treat TB patients, may have a lower index of suspicion for TB or even no knowledge at all, compared to public facilities. Training of health workers on new guidelines like the TBIC guidelines, are mainly organized in public facilities. Thus seeking care first with a private care provider results in delays in diagnosis and treatment of TB due the longer path from the initial visit to finally accessing a TB service provider (which is usually a public health facility) [17].

### Table 4 Predictors of unacceptable health service delay (>1 week)

| Variable                        | Unacceptable health service delay | Univariate analysis | Multivariate analysis | P    |
|---------------------------------|----------------------------------|---------------------|-----------------------|------|
|                                 | Yes/No                           | Crude PR 95% CI     | Adjusted PR 95% CI    |      |
| Sex                             |                                  |                     |                       |      |
| Female                          | 57/62 (92)                       | 1                   | 1                     |      |
| Male                            | 83/96 (86)                       | 0.94 (0.85-1.03)    | 0.98 (0.91-1.05)      | 0.63 |
| Age group                       |                                  |                     |                       |      |
| <=34                            | 92/101 (91)                      | 1                   | 1                     |      |
| >=35                            | 48/57 (84)                       | 0.92 (0.84-1.00)    | 1.02 (0.96-1.09)      | 0.44 |
| Monthly income in Ugandan shillings |                       |                     |                       |      |
| ≤20,000                         | 35/39 (90)                       | 1                   | -                     | -    |
| >20,000                         | 104/117 (89)                     | 1.01 (0.93-1.09)    |                       |      |
| Education level                 |                                  |                     |                       |      |
| No formal education             | 10/13 (77)                       | 1                   | 1                     |      |
| Primary                         | 63/73 (86)                       | 1.12 (0.82-1.51)    | 0.99 (0.77-1.28)      | 0.99 |
| Secondary and above             | 67/72 (93)                       | 1.12 (0.92-1.57)    | 1.02 (0.80-1.30)      | 0.83 |
| Marital status                  |                                  |                     |                       |      |
| Married                         | 68/78 (87)                       | 1                   | -                     | -    |
| Single                          | 34/38 (89)                       | 1.02 (0.90-1.16)    |                       |      |
| Separated/Widow/Divorce         | 38/42 (90)                       | 1.03 (0.94-1.13)    |                       |      |
| Distance to facility of diagnosis |                                  |                     |                       |      |
| Within 5km                      | 80/86 (93)                       | 1                   | 1                     |      |
| Over 5km                        | 60/72 (83)                       | 0.89 (0.788-1.01)   | 0.93 (0.82-1.06)      | 0.29 |
| Type of facility first visited  |                                  |                     |                       |      |
| Non-Public (private)            | 119/123 (97)                     | 1                   | -                     | -    |
| Public                          | 21/35 (60)                       | 0.62 (0.47-0.81)    | 0.62 (0.47-0.83)      |      |
| Tested for HIV before           |                                  |                     |                       |      |
| Yes                             | 91/104 (88)                      | 1                   | -                     | -    |
| No                              | 49/54 (91)                       | 1.03 (0.94-1.13)    |                       |      |
| Residence                       |                                  |                     |                       |      |
| Rural                           | 50/57 (88)                       | 1                   | -                     | -    |
| Semi-urban                      | 90/101 (89)                      | 1.01 (0.89-1.15)    |                       |      |

*Statistically significant.
The median total delay of 16 weeks observed in our study is close to what was found in other studies from Pakistan and Tanzania [18,19] though lower than what was reported in Kampala City Councils clinics in Uganda [7]. The longer total delay observed in our study compared to previous studies conducted in Uganda [5,7] can be explained by better access to health facilities in the urban setting. Comparing treatment delay between different studies is difficult because they often used different definitions (Table 1). According to the WHO guidelines and the Ugandan MOH TBIC guidelines, patients with cough for 2 weeks and above should be investigated for TB within 24 hours [10,11]. Therefore, in our study we considered a delay of 3 weeks as unacceptable patient delay and a delay of more than one week as unacceptable health service delay. Adhering to these shorter delays will result in low TB transmission and thus improved TB control.

Almost half of the patients sought care within the first three weeks of cough. This is lower than what was observed in a national Ugandan referral hospital [5], where 70% of patients contacted a health care provider within two weeks of cough. This could also be explained by better access to health services in an urban setting, compared to the rural setting where our study was conducted. Almost 10% of our patients first sought care from traditional healers/herbalists. This is consistent with the findings in South Africa [20]. This highlights the need to involve traditional healers in TB control, since a substantial proportion of patients seek care from them. The patient delay observed in this study is comparable to those reported by other studies in sub-Saharan Africa [5,7,21,22].

Table 5 Predictors of unacceptable total delay (>4 weeks)

| Variable                      | Unacceptable Total delay | Univariate analysis | Multivariate analysis |
|-------------------------------|--------------------------|---------------------|-----------------------|
|                               | Yes                      | No                  | Crude PR 95% CI       | Adjusted PR 95% CI  |
| Sex                           |                          |                     |                       |
| Female                        | 53/61 (87)               | 8/61 (13)           | 1                     | 1                     |
| Male                          | 87/93 (94)               | 6/93 (6)            | 1.07 (0.96-1.20)      | 1.09 (0.99-1.20)      | 0.073    |
| Age group                     |                          |                     |                       |
| ≤ 34                          | 89/99 (90)               | 10/99 (10)          | 1                     | _                     | _        |
| ≥35                           | 51/55 (93)               | 4/55 (7)            | 1.03 (0.94-1.12)      | _                     | _        |
| Salary income in Ugandan shillings |                    |                     |                       |
| ≤20,000                       | 35/37 (95)               | 2/37 (5)            | 1                     | _                     | _        |
| >20,000                       | 104/115 (90)             | 11/115 (10)         | 0.95 (0.88-1.03)      | _                     | _        |
| Marital status                |                          |                     |                       |
| Married                       | 68/75 (91)               | 7/75 (9)            | 1                     | _                     | _        |
| Single                        | 34/38 (89)               | 4/38 (11)           | 0.98 (0.92-1.05)      | _                     | _        |
| Separated/Widowed             | 38/41 (93)               | 3/41 (7)            | 1.02 (0.88-1.17)      | _                     | _        |
| Education level               |                          |                     |                       |
| No formal education           | 12/13 (92)               | 1/13 (8)            | 1                     | _                     | _        |
| Primary                       | 66/71 (93)               | 5/71 (7)            | 1.00 (0.90-1.12)      | _                     | _        |
| Secondary and above           | 62/70 (89)               | 8/70 (11)           | 0.95 (0.78-1.17)      | _                     | _        |
| Distance to facility of diagnosis |                      |                     |                       |
| Within 5km                    | 77/85 (91)               | 8/85 (9)            | 1                     | _                     | _        |
| Over 5km                      | 63/69 (91)               | 6/69 (9)            | 1.00 (0.89-1.12)      | _                     | _        |
| Type of facility first visited |                          |                     |                       |
| Non-Public (private)          | 112/120 (93)             | 8/120 (7)           | 1                     | _                     | _        |
| Public                        | 28/34 (82)               | 6/34 (18)           | 0.88 (0.74-1.05)      | 0.86 (0.74-1.01)      | 0.078    |
| Tested for HIV before         |                          |                     |                       |
| Yes                           | 91/104 (88)              | 13/104 (12)         | 1                     | _                     | _        |
| No                            | 46/54 (91)               | 5/54 (9)            | 1.02 (0.91-1.15)      | _                     | _        |
| Residence                     |                          |                     |                       |
| Rural                         | 50/55 (91)               | 5/55 (9)            | 1                     | _                     | _        |
| Semi-urban                    | 90/99 (91)               | 9/99 (9)            | 1.00 (0.88-1.13)      | _                     | _        |
delay in this study contributed least towards total delay. This is contrary to what was observed in a study from Tanzania where patient delay contributed to 90% of the total delay [23]. However the Tanzanian study used the following definition of patient delay “the period between onset of symptoms and reporting to a health facility.” In our study we defined patient delay as the period between onset of symptoms and seeking care from any health care provider. We used “any health care provider” because of the WHO recommendation to involve all health care providers in TB control [2]. Patient delay was significantly associated with being male, unlike what was observed by Mfinanga in Tanzania [24]. On the other hand, Johansson et al. in Vietnam reported that men were more likely to neglect symptoms and to seek care only when the disease gets into an advanced stage [25]. This has also been reported in HIV care, where men have been found to present with advanced disease and were more likely to be lost to follow-up [26,27].

Unacceptable health service and total delay were associated with visiting non-public health facilities. To reduce these delays, coordination between the National TB and Leprosy Programme (NTLKP) and the private sector is critical for TB control [2,28]. Efforts to engage with private providers have already started in Uganda and this resulted in improved case detection in an urban setting [29]. But pharmacies, drug shops and traditional healers need to be brought on board. This can be achieved by the use of incentives to encourage private care providers to refer all TB suspects to a diagnostic facility [2]. However, there is a need to explore more sustainable ways on how to involve them, more particularly the traditional healers. A public-private mix in TB control is in line with the Stop TB Strategy [2]. Furthermore, there is also a need to train all health care providers to raise their level of suspicion for TB. In addition, community TB advocacy and sensitization, which is advocated in TBIC is necessary to improve TB awareness among the people and thus result in early care seeking for TB symptoms. Rolling out rapid diagnostics would also help in reducing diagnostic delay. This could for instance be done by use of the gene X-pert [2,30,31], but also by taking two spot sputum specimens for smear microscopy on the same day rather than on different days [32], so as to reduce patients failing to return in time or even being lost to follow up.

Our study has limitations. Being a retrospective study, imprecise estimates of delay due to recall bias among the patients could have influenced our results. This was minimised by the use of TB treatment cards and case notes. In addition, we asked patients if they thought TB was curable, before they started on treatment. This could have introduced a bias considering the health education patients are given before they start and also in the course of the treatment. We may have also introduced a selection bias by only including outpatients. The study was carried out in a few facilities (health centre IVs and hospitals) and mainly government, so this limits the generalizability of our findings. The picture could be different in lower level facilities and in private facilities. Last but not least, length of cough was used as an indicator for patient delay. However, it can start in an insidious manner and so the timing of onset of cough can vary among respondents depending on the manner they gauge the seriousness of the symptom. Moreover, we did not assess treatment delay of patients with smear negative pulmonary TB and patients with extrapulmonary TB.

Conclusions
This study reveals that there is still a significant TB treatment delay problem in Uganda. Our study underscores the need to involve all health care providers in TB control. TB treatment delay can be seen as a proxy indicator of health system performance. Our results point to the fact that the public sector seems to be doing better than the various private providers that operate in rural Uganda. The range of relevant actions to be considered could include: health education on TB and TBIC, further decentralization of health services, appropriate training of health care providers in TBIC and public-private partnership in TB control. Research is needed on why there is prolonged health service delay and how private care providers including traditional healers can be effectively engaged by national TB control programmes.

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

Authors’ contributions
EB, BC, FN, RC were involved in the development of the proposal. EB carried out field work. EB, FN, and RC undertook data analysis. EB, FN, RC, and BC drafted the manuscript and approved the final draft.

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