Delay in seeking care for tuberculosis symptoms among adults newly diagnosed with HIV in rural Malawi

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SUMMARY

SETTING: Ten primary health clinics in rural Thyolo District, Malawi.

OBJECTIVE: Tuberculosis (TB) is a common initial presentation of human immunodeficiency virus (HIV) infection. We investigated the time from TB symptom onset to HIV diagnosis to describe TB health-seeking behaviour in adults newly diagnosed with HIV.

DESIGN: We asked adults (≥18 years) about the presence and duration of TB symptoms at the time of receiving a new HIV diagnosis. Associations with delayed health seeking (defined as >30 and >90 days from the onset of TB symptoms) were evaluated using multivariable logistic regression.

RESULTS: TB symptoms were reported by 416 of 1265 participants (33%), of whom 36% (150/416) had been symptomatic for >30 days before HIV testing. Most participants (260/416, 63%) were below the poverty line (US$0.41 per household member per day). Patients who first sought care from informal providers had an increased odds of delay of >30 days (adjusted odds ratio [aOR] 1.6, 95%CI 0.9–2.8) or 90 days (aOR 2.0, 95%CI 1.1–3.8).

CONCLUSIONS: Delayed health seeking for TB-related symptoms was common. Poverty was ubiquitous, but had no clear relationship to diagnostic delay. HIV-positive individuals who first sought care from informal providers were more likely to experience diagnostic delays for TB symptoms.

KEY WORDS: TB; human immunodeficiency virus; health seeking; symptom screening

HUMAN IMMUNODEFICIENCY VIRUS (HIV) and tuberculosis (TB) are the leading causes of adult deaths due to infectious agents worldwide, especially in sub-Saharan Africa. An estimated 1.2 million (12%) of the 10.4 million people who developed TB worldwide in 2015 had HIV infection. The incidence of and mortality due to TB increased steeply with the onset of the HIV epidemic and, despite recent gains, TB remains the leading cause of death among people living with HIV. TB thus still remains a major challenge.

TB symptoms are commonly associated with undiagnosed HIV at all levels of the health system. Compared with TB diagnostics, HIV rapid diagnostic tests are quick, highly accurate, available in point-of-care (POC) format and more widely decentralised. As such, it is common for HIV to be diagnosed before TB investigations have started, even when the presenting complaint is consistent with active TB. Lack of rapid POC diagnostics for TB translates into a lack of rapid decision making, thereby increasing the number of visits required for TB diagnosis, loss to follow-up and patient costs. Being investigated for TB is also associated with substantial costs to patients, both direct (e.g., food and transport) and indirect (loss of wages due to time spent seeking care). Anti-tuberculosis treatment also requires multiple facility visits, and as TB is more stigmatised than HIV, it is often less widely recognised in the community. People with symptoms such as cough may therefore delay seeking care for TB, yet seek a diagnosis for HIV. Individuals with newly diagnosed HIV thus represent a group whose initial steps in the TB care pathway can be investigated.

Socio-economic factors are closely associated with health-seeking behaviour, including HIV and TB.

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Catastrophic costs (defined as totalling >20% of the annual income) occur mostly from the onset of TB symptoms to starting treatment, a fundamental driver being the number of clinic visits. Interventions focused at reducing the time to diagnosis could reduce not only these costs, but also transmission of both TB and HIV.

Several studies have evaluated the time from the first TB symptom to TB diagnosis. Few studies, however, have looked at patient delays from the onset of symptom(s) suggestive of TB to the time of HIV diagnosis in a formal health facility. The primary aim of the present study was to investigate the factors associated with delay from the onset of TB symptoms to HIV diagnosis of >30 days among adults newly diagnosed with HIV in a rural primary health care setting in Malawi. The secondary aim was to investigate risk factors for a delay of >90 days.

METHODS

Study design and setting of the parent study: CHEPETSA study

The CHEPETSA study was a cluster randomised trial conducted in 12 rural primary care clinics in Thyolo District, Malawi, among adults with newly diagnosed HIV infection (CHEPETSA, clinicaltrials.gov #NCT01450085). In this parent study, clinics (clusters) were randomised to one of two TB screening algorithms: symptom screening plus sputum smear microscopy and symptom screening plus sputum testing using the Xpert® MTB/RIF assay (Cepheid, Sunnyvale, CA, USA). All study participants were screened for TB symptoms (cough of any duration, fever, recent weight loss or night sweats) at enrolment. If at least one TB symptom was reported, participants were asked to provide sputum for TB diagnosis with smear microscopy or Xpert, depending on the study arm. If asymptomatic and eligible, participants were initiated on isoniazid preventive therapy for 6 months. All participants were followed for 1 year after HIV diagnosis. The primary outcome of the parent trial was all-cause mortality at 1 year from enrolment.

Study sample for this analysis

In this analysis, we used enrolment data from participants recruited into the CHEPETSA study. Trial participants were included if they were enrolled on or before 1 April 2015 from 10 clinics (five per study arm) and reported one or more of the four TB symptoms listed above at enrolment. Baseline evaluation used standardised questionnaires to elicit demographics (age and sex), time of onset and duration of TB symptoms, asset ownership, smoking status and transit time to clinic. (See Appendix Table A for the variables used in asset ownership).

Statistical methods

We defined delay a priori as >30 days from the onset of TB symptoms to the time of HIV diagnosis. By design, HIV diagnosis occurred at the same time as enrolment into the CHEPETSA study. Using data collected by self-reporting on asset ownership, recent purchase of sugar, education level of the household head, household cooking over firewood, acreage cultivated, household size, and whether maize and/or tobacco was grown in the household, we created a ‘wealth score’ variable for household wealth that was measured using a proxy means test developed for rural populations from the 1998 Malawi Integrated Household Survey (IHS) (see Appendix). This method estimates household consumption (measured as the wealth score) using proxy measures. The wealth score was then coded as a binary variable using a predefined cut-off point of 10.47 Malawian kwacha (valued in 1998 currency, equal to US$0.41) per person per day. Participants with an estimated household consumption below this cut-off were classified as severely poor. Age was grouped into three categories: <30 years, 30–40 years and >40 years. The initial site of care seeking was dichotomised as clinic/hospital (formal services) vs. other (traditional healer/pharmacist/none). In this rural setting, pharmacists are generally not formally trained and do not dispense prescription drugs, but rather sell non-prescription drugs as well as other grocery items.

We used logistic regression to explore the factors associated with delay, including age group, sex, education, smoking status, employment, time taken to reach the clinic from home, marital status, mode of transport to the clinic, site of first attempted treatment seeking, time taken from home to the clinic and then back home, history of previous anti-tuberculosis treatment, household size, self-reported general health and wealth. We hypothesised that age group, wealth and sex would affect health-seeking behaviour, and therefore included these variables a priori in all adjusted analyses. In addition, other factors from the univariable analysis with P < 0.2 were included in the multivariable analysis and retained if, after adjustment, the P value remained <0.2. Analysis was also repeated using a secondary outcome of delay of >90 days. All analyses were performed using Stata 13 (Stata Corp, College Station, TX, USA).

Ethical considerations

The parent trial was approved by the Malawi College
of Medicine Research, Blantyre, Malawi, the Ethics Committee of the London School of Hygiene & Tropical Medicine, London, UK, and Johns Hopkins Medicine Institutional Review Board, Baltimore, MD, USA. All study participants provided written consent before enrolment.

RESULTS

Overall, 1577 participants from 10 clinics were enrolled into the parent trial on or before 1 April 2015 (Figure). Of these, 312 were excluded from further analysis as not having complete data on socio-economic status collected. During screening, 458 (36%) of the remaining 1307 eligible participants reported one or more TB symptoms at enrolment. Of these, a further 42 participants were excluded due to missing data on delay and/or components of the wealth score, leaving 416 participants for the analysis of delay in seeking care.

The baseline characteristics of the 416 participants are given in Table 1. Overall, 52% of the participants were male; 60% were aged <40 years. Most participants were severely poor based on the 1998 poverty line (n = 260, 63%), had low levels of education (none/primary: n = 329, 81%) and had travelled >1 h from home to get to the clinic (n = 294, 71%).

Of the 416 participants with TB symptoms, 150 (36%) reported a delay of >30 days from the onset of these symptoms to HIV diagnosis. Seventy-eight (19%) participants reported a delay of >90 days.

In multivariable analysis, patients who first sought care from informal (traditional healers and pharmacists) or no other services, as opposed to formal services (clinics or hospitals), had increased odds of delay; this finding was not statistically significant (adjusted odds ratio [aOR] 1.61, 95% confidence interval [CI] 0.9–2.8, P = 0.09; Table 2). When delay was defined as >90 days from symptom onset to HIV diagnosis, patients who first sought care from settings other than formal services (clinics or hospitals) had increased odds of delay (aOR 2.0, 95%CI 1.1–3.8, P = 0.03; Table 2). Patients from households with ≥5 members (compared with those with <5 members) had lower odds of delay (aOR 0.7, 95%CI 0.4–1.2, P = 0.16; Table 2), although this finding was not significant. Delay—whether measured as 30 or 90 days—did not vary significantly according to age group, sex, smoking status, time taken to reach the clinic from home, wealth, mode of transport or having previous anti-tuberculosis treatment (Table 2).

DISCUSSION

This analysis of 416 rural Malawian adults newly diagnosed with HIV and reporting TB symptoms suggests that patients who first seek care from traditional healers and pharmacists may have increased odds of a prolonged delay in diagnosis. We also highlight the high prevalence of both extreme poverty (63%) and delay in diagnosis (36% with >30 days delay) in this rural population. We report the duration of TB symptoms before HIV diagnosis, and provide support for the measurement of TB symptoms at HIV diagnosis as a potentially useful approach for assessing delayed care seeking for HIV-TB.

Despite careful consideration of multiple risk factors, we found no significant associations with delayed care seeking, which was defined using our a priori cut-off of >30 days between the self-reported onset of TB symptoms and date of HIV diagnosis. When considering prolonged delays (>90 days), seeking assistance from a traditional healer or pharmacist was associated with a doubling of the odds of diagnostic delay. While only 15% of patients first sought a diagnosis from traditional healers and pharmacists (as opposed to visiting clinics or hospitals), these individuals accounted for nearly half of all patients who experienced prolonged delays. We did not find evidence of an association between first seeking treatment from traditional healers/pharmacists and any other risk factors in this cross-sectional study.

Our findings are consistent with work in Malawi by Brouwer et al., who found that 37% of TB patients...
Table 1  Characteristics of HIV-positive adults attending primary care clinics, and association with the delay from TB symptom onset to HIV diagnosis of >30 days (n=416)

| Characteristic                          | Total (n=416) | Delay (n=150) | Unadjusted OR (95%CI) | P value | Adjusted OR (95%CI)* | P value |
|----------------------------------------|---------------|---------------|-----------------------|---------|----------------------|---------|
| **Age group, years**                   |               |               |                       |         |                      |         |
| <30                                    | 89 (21.4)     | 33 (37.1)     | 1 (reference)         | 0.35    | 1 (reference)        | 0.3     |
| 30–39                                  | 160 (38.5)    | 51 (31.9)     | 0.79 (0.46–1.37)      | 0.74    | 0.42–1.30            |         |
| ≥40                                    | 167 (40.1)    | 66 (39.5)     | 1.11 (0.65–1.88)      | 1.05    | 0.61–1.81            |         |
| **Sex**                                |               |               |                       |         |                      |         |
| Male                                   | 218 (52.4)    | 86 (39.5)     | 1 (reference)         | 0.13    | 1 (reference)        | 0.82    |
| Female                                 | 198 (47.6)    | 64 (32.3)     | 0.73 (0.49–1.10)      | 1.17    | 0.30–4.50            |         |
| **Marital status**                     |               |               |                       |         |                      |         |
| Ever married                           | 398 (95.7)    | 142 (35.7)    | 1 (reference)         | 0.45    |                      |         |
| Never married                          | 18 (4.3)      | 8 (44.4)      | 1.43 (0.55–3.73)      |         |                      |         |
| **Wealth†**                            |               |               |                       |         |                      |         |
| Not severely poor                      | 156 (37.5)    | 57 (36.5)     | 1 (reference)         | 0.87    | 1 (reference)        | 0.42    |
| Severe poor                            | 260 (62.5)    | 93 (35.8)     | 1.03 (0.68–1.56)      | 1.25    | 0.74–2.08            |         |
| **Household size, people**             |               |               |                       |         |                      |         |
| <5                                     | 215 (51.7)    | 80 (37.2)     | 1 (reference)         | 0.61    |                      |         |
| ≥5                                     | 201 (48.3)    | 70 (34.8)     | 0.9 (0.60–1.34)       |         |                      |         |
| **Smoking**                            |               |               |                       |         |                      |         |
| Never smoked                           | 297 (71.4)    | 100 (33.7)    | 1 (reference)         | 0.11    |                      |         |
| Smoker/ever smoked                     | 119 (28.6)    | 50 (42.0)     | 0.7 (0.45–1.08)       |         |                      |         |
| **Employment**                         |               |               |                       |         |                      |         |
| Non-formal                             | 256 (61.5)    | 89 (34.8)     | 1 (reference)         | 0.6     |                      |         |
| Formal                                 | 108 (26.0)    | 39 (36.1)     | 1.06 (0.66–1.70)      |         |                      |         |
| Student/unemployed                     | 52 (12.5)     | 22 (43.1)     | 1.42 (0.77–2.62)      |         |                      |         |
| **Previous anti-tuberculosis treatment**|             |               |                       |         |                      |         |
| No                                     | 397 (95.4)    | 141 (35.5)    | 1 (reference)         | 0.29    |                      |         |
| Yes                                    | 19 (4.6)      | 9 (47.4)      | 1.63 (0.65–4.12)      |         |                      |         |
| **Self-reported general health‡**      |               |               |                       |         |                      |         |
| Excellent/good                         | 122 (29.3)    | 38 (31.2)     | 1 (reference)         | 0.18    |                      |         |
| Fair/poor                              | 294 (70.7)    | 112 (38.1)    | 1.36 (0.87–2.13)      |         |                      |         |
| **Education level§**                   |               |               |                       |         |                      |         |
| None                                   | 60 (14.8)     | 21 (35.0)     | 1 (reference)         | 0.85    |                      |         |
| Primary (1–8)                          | 269 (66.4)    | 98 (36.4)     | 1.06 (0.59–1.91)      |         |                      |         |
| At least secondary                     | 76 (18.8)     | 25 (32.9)     | 0.91 (0.45–1.86)      |         |                      |         |
| **Time to clinic from home, h**        |               |               |                       |         |                      |         |
| >2                                     | 151 (36.3)    | 58 (38.4)     | 1 (reference)         | 0.69    |                      |         |
| 1–2                                    | 143 (34.4)    | 48 (33.6)     | 0.81 (0.50–1.31)      |         |                      |         |
| <1                                     | 122 (29.3)    | 44 (36.1)     | 0.9 (0.55–1.48)       |         |                      |         |
| **Mode of transport¶**                 |               |               |                       |         |                      |         |
| Walking                                | 262 (63.4)    | 89 (34.0)     | 1 (reference)         | 0.55    |                      |         |
| Bicycle                                | 100 (24.2)    | 40 (40.0)     | 1.3 (0.81–2.08)       |         |                      |         |
| Vehicle                                | 51 (12.4)     | 19 (37.3)     | 1.15 (0.62–2.15)      |         |                      |         |
| **First care seeking#**                |               |               |                       |         |                      |         |
| Formal services                        | 354 (85.3)    | 121 (34.2)    | 1 (reference)         | 0.08    | 1 (reference)        | 0.09    |
| Traditional/pharmacy/none              | 61 (14.7)     | 28 (45.9)     | 1.63 (0.94–2.83)      | 1.61    | 0.93–2.80            |         |
| **Turnaround time, h**                 |               |               |                       |         |                      |         |
| Last clinic visit²**                   |               |               |                       |         |                      |         |
| >3                                     | 248 (60.1)    | 96 (38.7)     | 1 (reference)         | 0.38    |                      |         |
| 2–3                                    | 72 (17.4)     | 24 (33.3)     | 0.79 (0.46–1.38)      |         |                      |         |
| <2                                     | 93 (22.5)     | 29 (31.2)     | 0.72 (0.43–1.19)      |         |                      |         |
| Recruitment visit²**                   |               |               |                       |         |                      |         |
| >3                                     | 356 (85.6)    | 133 (37.4)    | 1 (reference)         | 0.37    |                      |         |
| 2–3                                    | 36 (8.7)      | 11 (30.6)     | 0.74 (0.35–1.55)      |         |                      |         |
| <2                                     | 24 (5.8)      | 6 (25.0)      | 0.56 (0.22–1.44)      |         |                      |         |

* Adjusted for all variables listed, n=415.
† Measured using 1998 proxy means score.
‡ Measured by self-report as excellent, good, fair or poor.
§ Missing for 11 participants.
¶ Missing for 3 participants.
# Missing for 1 participant, 6 participants did not go anywhere.
** Measured by the time from home to clinic and then back home and waiting time at the clinic.
HIV = human immunodeficiency virus; TB = tuberculosis; OR = odds ratio; CI = confidence interval.
Table 2  Characteristics of HIV-positive adults attending primary care clinics, and association with the delay from TB symptom onset to HIV diagnosis of >90 days (n = 416)

| Characteristic                        | Delay (n = 78) | Unadjusted OR (95%CI) | Unadjusted P value | Adjusted OR (95%CI) | Adjusted P value |
|---------------------------------------|----------------|-----------------------|--------------------|---------------------|------------------|
|                                       |                |                       |                    |                     |                  |
|                                       | (n = 416)      | n (row %)             |                    |                     |                  |
|                                       |                | Unadjusted           | Adjusted           |                     |                  |
|                                       |                | OR (95%CI)            | P value            | OR (95%CI)*         | P value          |
| Age group, years                      |                |                       |                    |                     |                  |
| <30                                   | 16 (18.0)      | 1 (reference)         | 0.11               | 1 (reference)       | 0.48             |
| 30–39                                 | 23 (14.4)      | 0.77 (0.38–1.54)      | 0.77 (0.37–1.60)   |                     |                  |
| ≥ 40                                  | 39 (23.4)      | 1.39 (0.73–2.66)      | 1.43 (0.73–2.82)   |                     |                  |
| Sex                                   |                |                       |                    |                     |                  |
| Male                                  | 45 (20.6)      | 1 (reference)         | 0.3                | 1 (reference)       | 0.39             |
| Female                                | 33 (16.7)      | 0.77 (0.47–1.26)      | 0.49 (0.09–2.53)   |                     |                  |
| Marital status                         |                |                       |                    |                     |                  |
| Ever married                          | 74 (18.6)      | 1 (reference)         | 0.7                |                     |                  |
| Never married                         | 4 (22.2)       | 0.8 (0.26–2.50)       | 0.5                |                     |                  |
| Wealth†                               |                |                       |                    |                     |                  |
| Not severely poor                     | 33 (21.2)      | 1 (reference)         | 0.33               | 1 (reference)       | 0.61             |
| Severely poor                         | 45 (17.3)      | 0.78 (0.47–1.29)      | 0.84 (0.45–1.60)   |                     |                  |
| Household size, people                |                |                       |                    |                     |                  |
| <5                                    | 46 (21.4)      | 1 (reference)         | 0.15               | 1 (reference)       | 0.16             |
| ≥ 5                                   | 32 (15.9)      | 0.7 (0.42–1.15)       | 0.69 (0.41–1.16)   |                     |                  |
| Smoking                               |                |                       |                    |                     |                  |
| Never smoked                          | 54 (18.2)      | 1 (reference)         | 0.64               |                     |                  |
| Smoker/ever smoked                    | 24 (20.2)      | 1.14 (0.67–1.94)      | 0.7                |                     |                  |
| Employment                            |                |                       |                    |                     |                  |
| Non-formal                            | 47 (18.4)      | 1 (reference)         | 0.56               |                     |                  |
| Formal                                | 18 (16.7)      | 0.89 (0.49–1.62)      | 0.84 (0.45–1.60)   |                     |                  |
| Student/unemployed                    | 13 (25.5)      | 1.52 (0.75–3.08)      | 0.84 (0.45–1.60)   |                     |                  |
| Previous anti-tuberculosis treatment  |                |                       |                    |                     |                  |
| No                                    | 76 (19.1)      | 1 (reference)         | 0.35               |                     |                  |
| Yes                                   | 2 (10.5)       | 0.5 (0.11–2.20)       | 1.14 (0.67–1.94)   |                     |                  |
| Self-reported general health‡         |                |                       |                    |                     |                  |
| Excellent/good                        | 23 (18.9)      | 1 (reference)         | 0.97               |                     |                  |
| Fair/poor                             | 55 (18.7)      | 1.0 (0.58–1.70)       | 0.78               |                     |                  |
| Education level§                      |                |                       |                    |                     |                  |
| None                                  | 11 (18.3)      | 1 (reference)         | 0.78               |                     |                  |
| Primary (1–8)                         | 52 (19.3)      | 1.07 (0.52–2.19)      | 0.78               |                     |                  |
| At least secondary                    | 12 (15.8)      | 0.84 (0.34–2.05)      | 0.78               |                     |                  |
| Time to clinic from home, h           |                |                       |                    |                     |                  |
| >2                                    | 32 (21.2)      | 1 (reference)         | 0.38               |                     |                  |
| 1–2                                   | 28 (19.6)      | 0.91 (0.51–1.60)      | 0.38               |                     |                  |
| <1                                    | 18 (14.8)      | 0.64 (0.34–1.21)      | 0.38               |                     |                  |
| Mode of transport¶                    |                |                       |                    |                     |                  |
| Walking                               | 46 (17.6)      | 1 (reference)         | 0.61               |                     |                  |
| Bicycle                               | 22 (22.0)      | 1.32 (0.75–2.34)      | 0.61               |                     |                  |
| Vehicle                               | 9 (17.7)       | 1.01 (0.46–2.21)      | 0.61               |                     |                  |
| First care seeking¶                   |                |                       |                    |                     |                  |
| Formal services                       | 60 (17.0)      | 1 (reference)         | 0.02               | 1 (reference)       | 0.03             |
| Traditional/pharmacy/none             | 18 (29.5)      | 2.05 (1.11–3.80)      | 2.04 (1.09–3.81)   |                     |                  |
| Turnaround time, h                    |                |                       |                    |                     |                  |
| Last clinic visit¶                    |                |                       |                    |                     |                  |
| >3                                    | 52 (21.0)      | 1 (reference)         | 0.33               |                     |                  |
| 2–3                                   | 11 (15.3)      | 0.68 (0.33–1.38)      | 0.33               |                     |                  |
| <2                                    | 14 (15.1)      | 0.67 (0.35–1.27)      | 0.33               |                     |                  |
| Recruitment visit¶                    |                |                       |                    |                     |                  |
| >3                                    | 71 (19.9)      | 1 (reference)         | 0.31               |                     |                  |
| 2–3                                   | 4 (11.1)       | 0.50 (0.17–1.46)      | 0.31               |                     |                  |
| <2                                    | 3 (12.5)       | 0.57 (0.17–1.98)      | 0.31               |                     |                  |

* Adjusted for all variables listed, n = 415.
† Measured using 1998 proxy means score.
‡ Measured by self-report as excellent, good, fair or poor.
§ Missing for 11 participants.
¶ Missing for 3 participants.
# Missing for 1 participant, 6 participants did not go anywhere.
** Measured by the time from home to clinic and then back home and waiting time at the clinic.
HIV = human immunodeficiency virus; TB = tuberculosis; OR = odds ratio; CI = confidence interval.
visited a traditional healer before seeking formal medical care, that these patients spent an average of 4 weeks with traditional healers and that none of the traditional healers referred patients to the formal health care system.\textsuperscript{19} These findings suggest that individuals seeking care from traditional healers and pharmacists may be an important group for targeted case-finding interventions, and that further engagement of such informal providers may be required to reduce diagnostic delays in the rural sub-Saharan African setting. Future studies are needed to confirm this finding and also to explore the potential impact of private-sector care on population-level transmission of TB in Malawi and other similar settings.

We did not detect an association between extreme poverty and delayed care seeking for TB symptoms in this population. In most societies, the greatest burden of TB (and increasingly HIV) is experienced by the poorest populations.\textsuperscript{20,21} HIV in itself is also a powerful risk factor for progression from tuberculous infection to TB disease, and is a driver of poverty.\textsuperscript{22} However, despite the very high levels of poverty in our study (with 63\% of the study population classified as extremely poor based on a proxy means test benchmarked to the 1998 poverty line), we found no evidence of a difference in diagnostic delays between severely poor and non-severely poor participants. In practice, the majority of participants in this study classified as not severely poor would still be classified as extremely poor by most international standards. Thus we may have failed to observe an association between wealth and diagnostic delay, in part due to the relatively ubiquitous poverty in our study setting.\textsuperscript{23}

Different approaches have been recommended for integrating HIV and TB services so as to provide universal access even to the poorest in all societies.\textsuperscript{24} Systematic screening of people living with HIV and prompt treatment are the principal tools for reducing transmission and controlling the spread of TB disease.\textsuperscript{25} However, despite these longstanding recommendations, the integration of HIV and TB services has been slow,\textsuperscript{26} and many studies have indicated that much still needs to be done.\textsuperscript{27,28} The novel approach discussed here—monitoring and reporting of duration of TB symptoms before HIV diagnosis—may provide a useful and readily obtainable metric that could be used to investigate both TB and HIV programme performance relating to a prompt offer of HIV testing to all patients reporting TB symptoms. This metric can also provide an indicator of health care-seeking behaviour among people with symptoms suggestive of TB, who may be more likely to present for an HIV diagnosis than for evaluation of their TB symptoms.

The present study had five important limitations. The first was our inability to explicitly measure integration of HIV and TB services and potential recall bias in participants' self-reported time from onset of TB symptoms to HIV diagnosis. Second, data on the total number of visits made to the formal health sector before HIV diagnosis were not collected. Third, our epidemiological setting of rural Malawi—while important—is not likely to be generalisable to urban settings or to those outside of sub-Saharan Africa. Fourth, our proxy means test of household wealth benchmarked against a poverty line (i.e., an absolute measure of poverty) may not accurately reflect poverty as actually experienced in this population. Not only was this measure designed to reflect a specific context and time (i.e., Malawi in 1998), but it may be sensitive to participants’ conceptualisation of household size or structure. Future studies might consider evaluating relative measures of poverty (in which wealth is compared against that of other members in the same society) to provide a different perspective. Finally, we did not include some variables (for example, seasonality) that may have important effects on the delay in seeking care.

In conclusion, we found that about two thirds of this rural Malawian population newly diagnosed with HIV and reporting TB symptoms met the criteria for extreme poverty, and one third reported delays of >30 days from symptom onset to the time of HIV diagnosis. Seeking care with informal providers was associated with an extreme delay in care seeking. These data highlight the challenges faced in diagnosing TB among people living with HIV in this setting, provide a metric (duration of TB symptoms at the time of HIV diagnosis) that can be used to evaluate programme performance, and underscore the importance of engaging informal providers if global targets for HIV and TB control are to be met in rural sub-Saharan Africa.

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Conflicts of interest: none declared.

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APPENDIX

The proxy means score was estimated for each participant based on the linear regression model, as follows:

\[ \log(y_i) = \beta_0 + \beta_1 X_{1i} + \beta_2 X_{2i} + \ldots + \beta_k X_{ki} + \epsilon, \]

where \( \beta_0, \beta_1, \beta_2, \ldots, \beta_k \) (\( k = 14 \)) are coefficients provided for in the Malawi Integrated Household Survey 1997–1998 proxy means test model. The coefficient estimates and description of each explanatory variable for the rural proxy means test model are shown in Table A. In this Table, unless stated otherwise, variables are coded as binary variables, where 0 denotes a ‘no’ response and 1 a ‘yes’ response.

From this regression model a logarithm (proxy means test score) was estimated for each participant and converted back to the original scale using the anti-logarithm (exponential) function to give a predicted proxy means test score for each study participant. The predicted wealth scores (on the original scale) were grouped into a binary variable, poor/non-poor, based on a predefined cut-off point using the 1998 proxy means test model score. The cut-off point was based on the weighted poverty line for Malawi as a whole (as of March 1998), which was at Malawian kwacha 10.47 per person per day, or US$0.41.17 All participants with a predicted wealth score of ≤10.47 were classified as poor and those with a score of >10.47 as non-poor.

| Explanatory variable                                      | Coefficient estimates |
|-----------------------------------------------------------|-----------------------|
| 1  Purchased sugar in the past 2 weeks                    | 0.152                 |
| 2  Education level of household head: categorical variable on the level of educational attainment: 1 = standards 1–4; 2 = standards 5–8; 3 = forms 1–2; 4 = forms 3 or 4; 5 = university or higher | 0.078                 |
| 3  Household cooks over collected firewood                | -0.174                |
| 4  Total acreage cultivated (per acre)                    | 0.029                 |
| 5  Household size (per person)                            | -0.283                |
| 6  Household size (per person) squared                    | 0.015                 |
| 7  Number of salaried household members (per person)      | 0.098                 |
| 8  Household owns a bicycle                                | 0.153                 |
| 9  Household owns a fridge                                | 0.591                 |
| 10 Household grows tobacco                                | 0.105                 |
| 11 Household owns a bed                                   | 0.263                 |
| 12 Number of cattle owned (per cow)                       | 0.013                 |
| 13 Household grows hybrid maize                           | 0.076                 |
| 14 Household owns a car or motorcycle                     | 0.693                 |
| Thyolo District                                            | -0.296                |
| Constant term                                             | 2.826                 |
Résumé

Contexte : Dix centres de santé primaires dans le district rural de Thyolo, Malawi.

Objectif : La tuberculose (TB) est une présentation initiale fréquente de l’infection au virus de l’immunodéficience humaine (VIH). Nous avons étudié le délai entre l’apparition des symptômes de TB et le diagnostic de VIH pour décrire le comportement en matière de recherche de soins de TB chez des adultes ayant récemment eu un diagnostic de VIH.

Schéma : Nous avons interrogé des adultes (âge ≥18 ans) à propos de la présence et de la durée des symptômes de TB au moment où ils ont eu un diagnostic de VIH. Les associations avec le retard à la demande de soins (défini comme >30 et >90 jours après l’apparition des symptômes de TB) ont été évaluées à l’aide d’une régression logistique multivariée.

Résultats : Des symptômes de TB ont été rapportés par 416 participants sur 1265 participants (33%), dont 36% (150/416) avaient eu des symptômes pendant >30 jours avant le test VIH. La majorité des participants vivait sous le seuil de pauvreté (260/416, 63%), c’est-à-dire, 0,41 SUS par membre du foyer par jour. Les patients qui ont d’abord sollicité des soins auprès de prestataires informels ont eu davantage de risques de délais >30 jours (OR ajusté [ORa] 1,6 ; IC95% 0,9–2,8) ou >90 jours (ORa 2,0 ; IC95% 1,1–3,8).

Conclusion : Le retard à la recherche de soins pour des symptômes liés à la TB a été fréquent. La pauvreté a été générale, mais n’a pas eu de rapport clair avec le retard au diagnostic. Les individus VIH positifs qui ont d’abord sollicité des prestataires de soins informels ont été plus susceptibles de connaître un retard au diagnostic des symptômes de TB.