

*Article***Active case finding for tuberculosis through TOUCH agents in selected high TB burden wards of Kolkata, India: A mixed methods study on outcomes and implementation challenges**

Abhijit Dey^{1*}, Pruthi Thekkur^{2,3}, Ayan Ghosh⁴, Tanusree Dasgupta¹, Soumyajyoti Bandopadhyay⁴, Arista Lahiri⁵, Chidananda Sanju S V⁶, Milan K.Dinda¹, Vivek Sharma¹, Namita Dimari¹, Dibyendu Chatterjee¹, Isita Roy¹, Anuradha Choudhury¹, Parthiban Shanmugam¹, Brojo Kishore Saha⁷, Sanghamitra Ghosh⁸, Sharath Burugina Nagaraja⁹

1. Tuberculosis Health Action Learning Initiative (THALI), Kolkata, West Bengal, Pin- 700107
 2. International Union against Tuberculosis and Lung Diseases. Paris, France, Zip-75006
 3. The Union South East Asia Office, New Delhi, India, Pin-110016
 4. Department of Community Medicine, College of Medicine & JNM Hospital, Kalyani, West Bengal, Pin-741235
 5. Department of Community Medicine, College of Medicine & Sagore Dutta Hospital, Kamarhati, Kolkata-700058
 6. District Tuberculosis Officer, Udupi, Government of Karnataka, India, Pin-576104
 7. State Tuberculosis Officer, Govt of West Bengal, India, Pin-700 091
 8. General Secretary, Indian Public Health Association (IPHA), HQ, Kolkata, Pin-700073
 9. Department of Community Medicine, ESIC Medical College and PGIMS, Bengaluru, India, Pin-560010
- * Correspondence: Dr. Abhijit Dey, 404 Kalikapur, Live Valley Apartment, Mukundapur (PO), Kolkata- 700099. Telephone- +91-8100650578, Email- drabhijitdey@gmail.com

Abstract:

Background: Active case finding for TB was implemented in selected sixty high TB burden wards of Kolkata, India. Community volunteers called TOUCH agents (TAs) identified and referred presumptive TB patients (PTBPs) to health facilities for TB diagnosis and treatment. We aimed to describe the 'care cascade' of PTBPs identified during July to December, 2018 and to explore the reasons for attrition as perceived by TAs and PTBPs.

Methods: An explanatory mixed methods study with quantitative phase of cohort study using routinely collected data followed by descriptive qualitative study with in-depth interviews was conducted.

Results: Of the 3, 86,242 individuals enumerated, 1132 (0.3%) PTBPs were identified. Only 713 (63.0%) PTBPs visited referred facility for TB diagnosis. TB was diagnosed in 177 (24.8%) and the number needed to screen for one TB was 2,183 individuals. The potential reasons for low yield were stigma and apprehension about TB, distrust about TA, wage loss for attending health facilities and substance abuse among PTBPs.

Conclusion: The yield of ACF was suboptimal with low PTBP identification rate and high attrition rate. Interviewing each individual for symptoms of TB and supporting PTBPs for diagnosis through sputum collection and transport can be adopted to improve the yield.

Keywords: Active case findings, Tuberculosis, TOUCH Agent, High TB burden area, TB Surveillance, 4S Screening, THALI Project, SORT IT, Operational Research

1. Introduction

Tuberculosis (TB) remains the major public health problem of concern with an estimated 10 million incident TB patients and 1.6 million deaths globally due to TB in the year 2017[1]. In the same year, only 6.7 million TB patients were notified to national TB programs (NTP)[1]. The rest 3.3 million (33% of the estimated) were either undetected or were detected but not notified to NTP. India alone has an estimated 0.8 million patients not notified to NTP[1].

The TB patients undetected or not treated can be a potential threat to TB control efforts as they contribute to uninterrupted transmission of disease. One of the potential reasons for undetected TB patients is the over reliance on passive case finding for diagnosis of TB in low-and-middle income countries (LMICs)[2,3].

The World Health Organization's (WHO) End TB strategy proposed early diagnosis of TB through Active case findings (ACF) in high risk groups as a key component to end TB epidemic by 2030[4]. ACF requires healthcare providers to actively reach and provide access to communities or population groups that are underserved or at higher risk of TB for TB diagnostic services.

The systematic reviews have reported ACF to be beneficial in increasing the case detection [5,6]. Also, the studies have reported ACF to be beneficial in reducing delay in TB diagnosis, out of pocket expenditure, unsuccessful treatment outcomes and incidence of TB[3,7,8]. The benefits of community based ACF activity depends on selection of high risk group, prevalence of the TB in the selected group, incentives provided to healthcare workers, the diagnostic algorithm adopted, and support provided to patients in completing the cascade of TB diagnosis and treatment.

The ACF activity has sequential stages ('cascade of care') through which patient traverses from diagnosis to treatment. Nevertheless, the studies elsewhere have shown as high as 46% attrition rates at various stages[9,10]. Thus, it is essential for ACF projects to monitor attrition at different stages, yield of TB patients, reasons for attrition and challenges in implementing ACF activity. This information can help programme managers to take informed decisions to optimize ACF activity services and the benefits of ACF.

The National Strategic Plan for TB elimination [2017-25] of India recommended ACF among high risk groups as a strategy to detect the 'missing' TB patients[11]. Since mid-2018, the NTP conducted five rounds of two week long ACF activity among identified marginalized and vulnerable groups in selected districts of country. The NTP also encouraged partner agencies supporting TB control to implement ACF among high risk groups[11].

The United States Agency for International Development (USAID) funded Tuberculosis Health Action Learning Initiative (THALI) project in West Bengal implemented ACF activity in the sixty high TB burden wards of Kolkata Municipal Corporation (KMC)[12-16]. A part-time, salaried community health volunteer known as Targeted Outreach for Upliftment of Community Health (TOUCH) agent was deployed in each ward to engage with community and conduct ACF. The THALI project ACF was different from that conducted by NTP as the activity was conducted throughout the year by a TOUCH agent (TA) who is not part of general health system.

There is limited information on the process, yield and challenges in implementing ACF activities in India. Though THALI project is implementing ACF from 2018, there has been no systematic evaluation of the programme. The information and lessons learnt from the same is

imperative to fix the deficiencies and improve the efficiency of ACF conducted by THALI and also NTP.

Hence, we aimed to determine the proportion identified as presumptive TB patient (PTBP), reached the health facility for diagnosis, diagnosed with TB and initiated on TB treatment among individuals enumerated by TAs during THALI project ACF activity conducted between July and December, 2018. Also, we aimed to explore the challenges in implementing the ACF activity as perceived by TAs and PTBPs.

2. Materials and Methods

2.1. Study Design:

We conducted an explanatory mixed methods study with the quantitative part (cohort study using secondary data collected routinely by the THALI Project) followed by qualitative part (descriptive study)[17].

2.2. Study Setting:

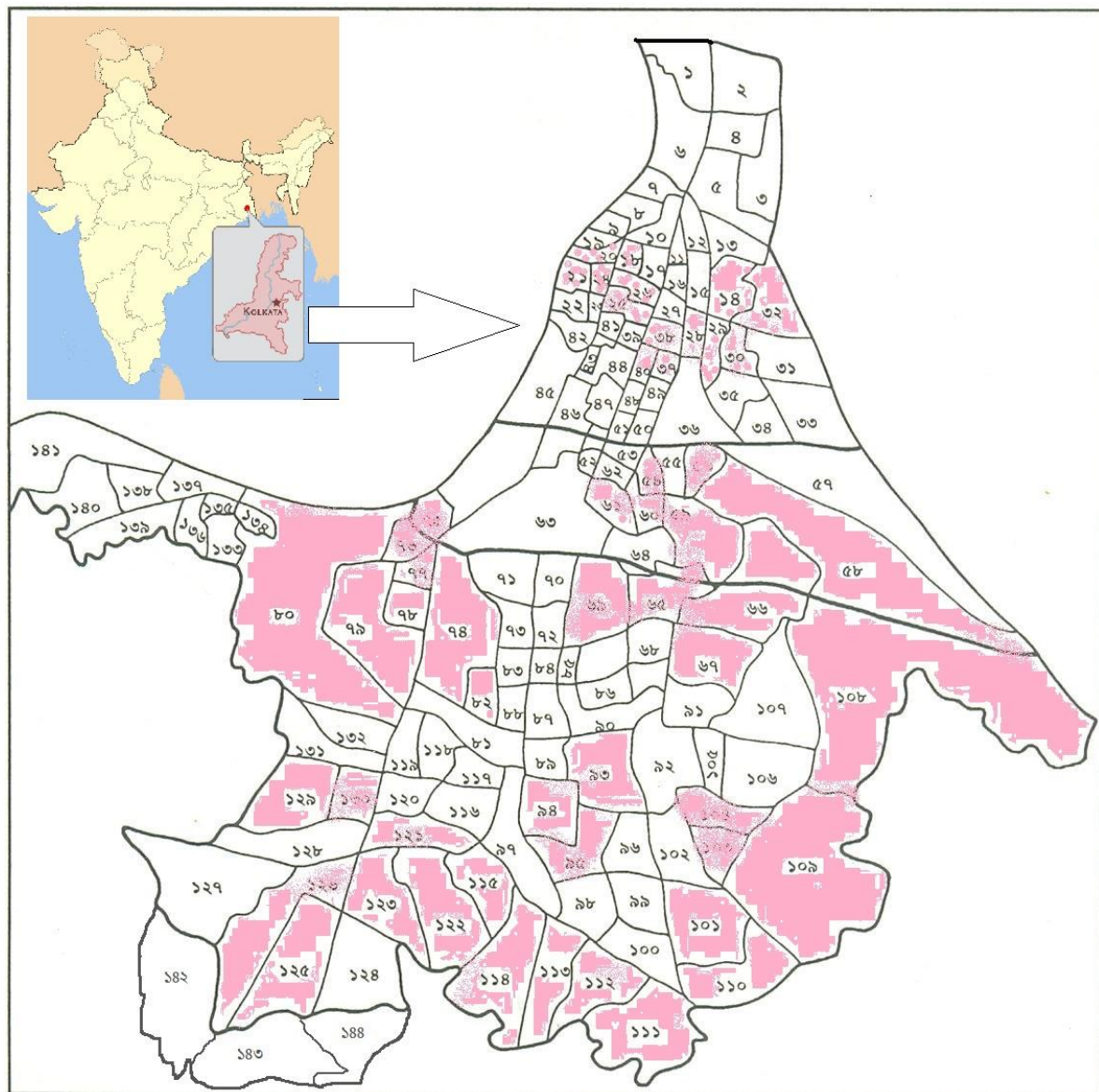
The Kolkata is a capital of West Bengal, India. It is one of the largest metro cities in the country with a population of 4.5 million[18]. The Kolkata Municipal Corporation (KMC) is divided into 144 urban wards as administrative units. The KMC has 5600 recognized slums and about 1.4 million individuals (~33% of total population) reside in slums[19].

The KMC is divided into ten urban health districts and each has one district TB officer who reports to city TB officer, a nodal person for delivery of NTP services in KMC. In total, there are 21 tuberculosis units (TU) and 42 designated microscopy centers (DMCs) in KMC[20]. The USAID supported THALI project implements ACF activity in sixty high TB burden wards of KMC since June-2018 through TAs.

2.2.1. Ward selection:

The wards for ACF were identified based on the burden of TB cases and the percentage of slum population. The number of TB patients notified in each of 144 wards during 1st March-31st May 2018 was calculated. The 54 wards with ten or more cases were considered as high TB burden wards. Other six wards with high proportion of slum population were considered and in total, 60 wards were identified as high priority wards for ACF activity. (Figure-1)

Figure 1: High Tuberculosis (TB) Burden Wards of Kolkata Municipal Corporation (KMC)



2.2.2. TOUCH Agent (TA):

They are mainly female community volunteer residing in the same ward with ability to read and write English language. Those selected as TA are trained on ACF activity. The TA spends at least 3 hours per day, six days a week, in the ward conducting ACF and community sensitization on TB. During each day, the TA visits 20 houses for ACF. TA is paid a fixed compensation of 3000 INR (~43 USD) per month.

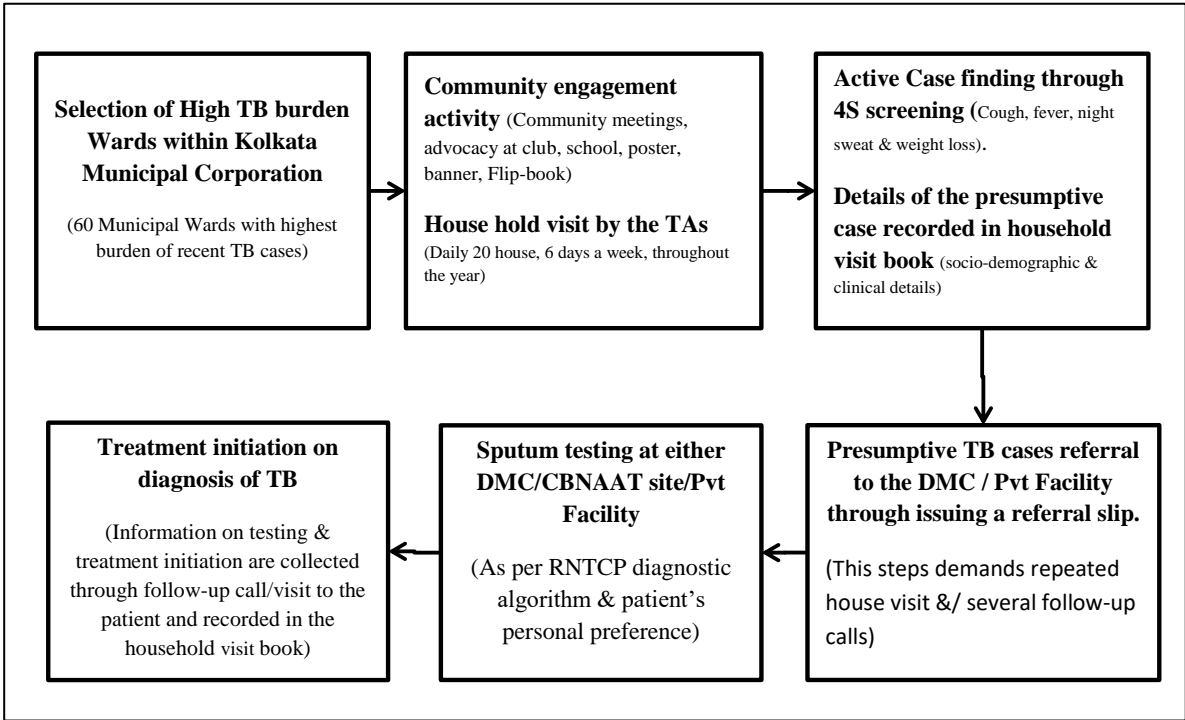
2.2.3. Active Case Finding:

The TA during house visit, enquires available individuals about the presence of symptoms indicative of TB (persistent cough for ≥ 2 weeks, fever for ≥ 2 weeks, significant weight loss ($>5\%$ weight loss over last 3 months), blood in sputum and night sweat ≥ 2 weeks). Also, TA enquires about the presence of symptoms indicative of TB among those not available at the home from the adult informant.

Those with symptom suggestive of TB are considered as PTBP and referred to either nearest DMC or the private practitioner based on patient’s preference. All the PTBPs referred are provided with referral slips signed by the TA. At the health facilities, PTBPs are evaluated for TB using sputum smear microscopy, chest X-ray and Xpert MTB/Rif assay as per NTP diagnostic algorithm[21].

The TA re-visits the house of PTBP within seven days, to collect the information about their visit to referred health facility. In case if patient has failed to visit health facility, the TA counsels patient and repeat house visit is made. If the patient has made the visit, the results of diagnostic test and details of treatment initiation are collected. Also, those diagnosed with TB are visited at least three times for contact tracing and counselling support during TB treatment. (Figure-2)

Figure 2: Flow Diagram of the ACF activity by the TAs under THALI Project



Abbreviations: TA= Touch Agent; DMC= Designated Microscopy Center; CBNAAT= Cartridge Based Nucleic Acid Amplification Test; RNTCP= Revised National Tuberculosis Control Program

2.2.4. Recording:

The TA maintains a “Household visit book” and documents the number of individuals screened and number of individuals with chest symptoms. The socio-demographic, clinical, referral and treatment details of the PTBP are noted in the “Household visit book”. Field coordinator reviews the “Household visit book” for completeness during the supervision visits.

2.3. Study Population

2.3.1. Quantitative:

We included all the PTBPs identified by TA between July and December, 2018 during ACF activity conducted in the selected wards of KMC.

2.3.2. Qualitative:

We included TAs (n=10) and PTBPs (n=7) identified during the study reference period. 'TA's were purposively selected using maximum variation sampling with selection of those who had referred maximum and minimum number of PTBPs. The PTBPs were purposively selected using the extreme variation sampling based on whether he/she attended the health facility on referral. The final sample size was guided by the saturation of findings.

2.4. Data variables, sources of data and data collection:

2.4.1. Quantitative:

Using structured proforma, principal investigator extracted the details of PTBPs from 'household visit book' during February, 2019. The date of house visit, ward number, age, gender, history of previous TB, family history of TB, tobacco use, alcohol use, diabetes, HIV, presenting symptoms, referral site, visited referred facility, date of visit to referred facility, presence of pulmonary TB, date of TB diagnosis, status of treatment initiation and date of treatment initiation were extracted.

2.4.2. Qualitative:

In-depth interviews were conducted face-to-face in local language (Bengali) by AD (a male medical doctor, MPH, trained in qualitative research) and audio recorded using mobile voice recorder after obtaining consent. The interviewer is working in a THALI project but not involved in implementation of ACF activity. Separate interview schedules were used to interview TAs and PTBPs. The 'TA's were interviewed at their residence or workplace. PTBPs were interviewed at their residence. Interviews lasted for an average of 21 minutes (range 8-42 minutes). At the end of each interview, debriefing was done to ensure member checking.

2.5. Data entry and analysis

2.5.1. Quantitative:

Data was double-entered and validated using EpiData entry version 3.1 (EpiData Association, Odense, Denmark). The analysis was done using Stata 11.0 (StataCorp LP, College Station, TX, USA). Key analytic outputs were the number and proportion of PTBPs at each step of diagnosis- treatment pathway. The percentage with 95% CI was used to summarize the proportion of PTBPs at each step.

To assess the independent association of socio demographic and clinical characteristics of PTBPs with 'not visiting referred health facility for diagnosis', a multivariable model (modified Poisson regression) with cluster adjustment at ward level was used to calculate adjusted relative risk (aRR) with 95% confidence intervals (CI). Similarly, a multivariable model (modified Poisson regression) with cluster adjustment at ward level was used to assess the factors independently associated with 'diagnosis of TB' among those who reached the health facility after referral. A 'p' value of <0.05 was considered to be statistically significant.

2.5.2. Qualitative:

Transcripts were prepared on the same day or one day after the interview using audio-recording and field notes. Manual descriptive thematic analysis of the transcripts was done by

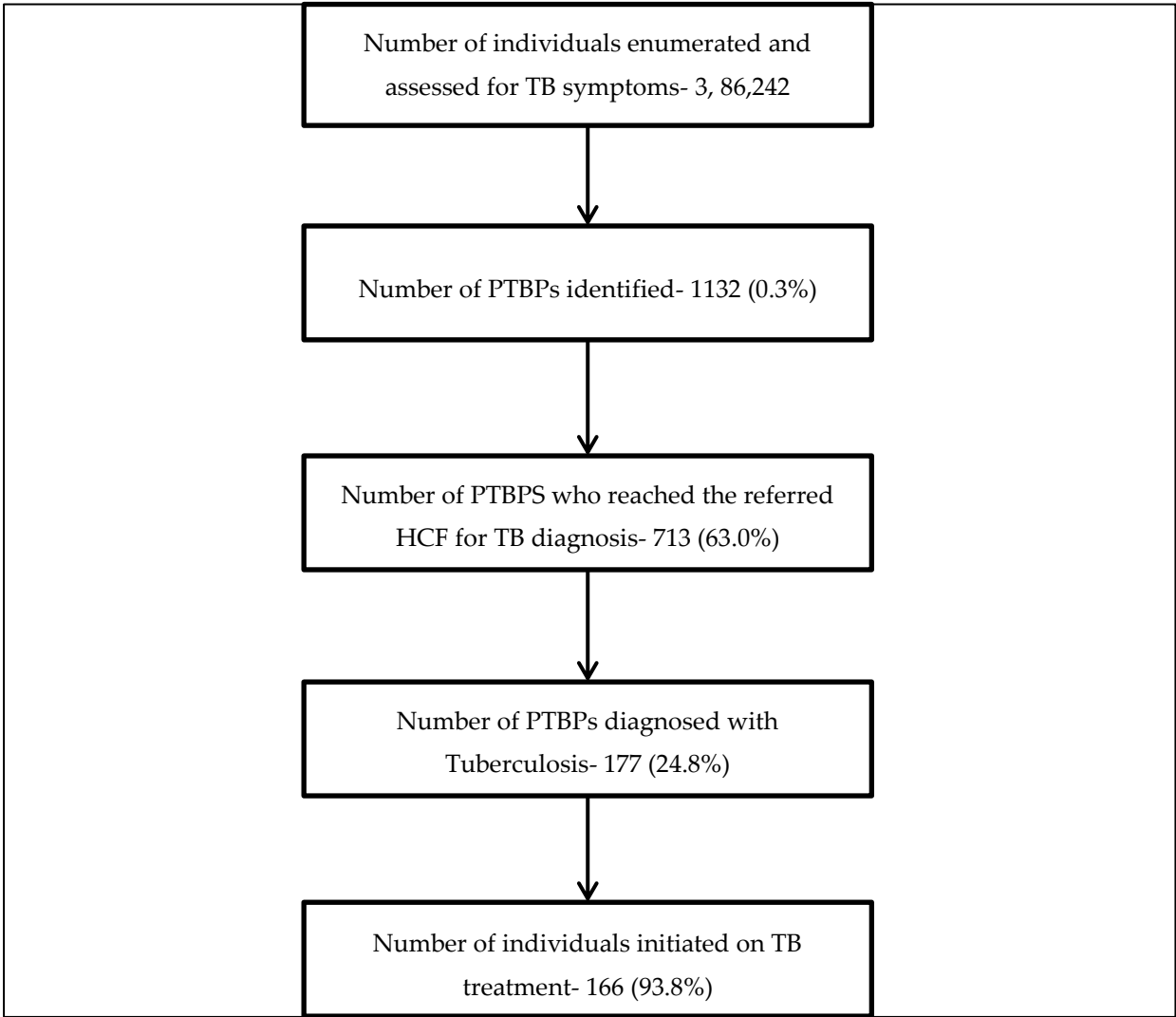
the AD and PT to identify the codes. The decision on the final coding and theme generation was done by using standard procedures and in consensus. The findings were reported by using 'Consolidated Criteria for Reporting Qualitative Research' [COREQ guideline][22].

3. Results

3.1. Quantitative

Of the total 3, 86,242 individuals were enumerated by TAs from 92,294 houses, 1132 (0.3%) were identified as PTBPs (**Figure-3**).

Figure 3: Flowchart depicting number and percentage of participants at different stages of diagnostic and treatment cascade test among PTBPs identified by ACF activity in Kolkata West Bengal India during July-December 2018



Abbreviations: ACF= Active Case Findings; PTBP= Presumptive TB Patient; HCF= Health Care Facility

The mean age of the PTBPs was 43.1 (standard deviation: 17.1) years and 647 (57.2%) were males. Tobacco use in last one month was reported by 175 (15.5%) PTBPs and 45 (3.9%) reported use of alcohol in past one month. Known diabetes was reported by 244 (21.5%) and 13 (1.2%) reported to have HIV infection. Of the total, 84 (7.4%) had previous history of TB and 118 (10.4%) gave history of TB among family members in last two years. Cough of ≥ 2 weeks without any other symptoms indicative of TB was seen in 687 (60.7%). (**Table 1**)

Table 1: Socio-demographic & clinical profile of the PTBPs identified by ACF activity in Kolkata Municipal Corporation, West Bengal, India during Jul- Dec 2018, N=1132

Characteristics	n	(%)
Age in years		
0-14	36	(3.2)
15-29	206	(18.2)
30-44	323	(28.5)
45-59	280	(24.7)
60-74	184	(16.3)
75 & above	32	(2.8)
Not recorded	71	(6.3)
Gender		
Male	647	(57.2)
Female	485	(42.8)
Alcohol use*		
Yes	45	(3.9)
No	1087	(96)
Tobacco user*	26	(2.3)
Yes	175	(15.5)
No	957	(84.5)
Presenting Symptoms		
Cough with other symptoms	101	(8.9)
Only Cough	687	(60.7)
No cough but other symptoms	344	(30.4)
Previous history of TB		
Yes	84	(7.4)
No	1048	(92.6)
History of TB of other family#		
Yes	118	(10.4)
No	1014	(89.6)
Diabetes		
Yes	244	(21.5)
No/Unknown	888	(78.5)
HIV		
Yes	13	(1.2)
No/Unknown	1119	(98.8)

Abbreviation: TB- Tuberculosis; PTBP- Presumptive TB patient; ACF- Active Case Finding, Diabetes= Diabetes Mellitus, HIV= Human Immunodeficiency Virus, * use in last one month, # History of TB among any other house hold member within last two years.

Of the 1132 PTBPs, 713 (63.0%, 95% CI: 60.1%-65.8%) visited the referred health facility for diagnosis of TB. Among those who visited referred health facility, the median duration between referral and reaching the facility was 2 (inter quartile range 1-6). **Table 2** shows the factors associated with not visiting the referred health facility among PTBPs. Female gender (aRR-1.3 (95% CI: 1.1-1.5)), alcohol use (aRR-1.7 (95% CI: 1.3-2.2)), cough as the only symptom (aRR-1.8 (95% CI: 1.5-2.2)) and family history of TB (aRR-1.3 (95% CI: 1.1-1.5)) were independently associated with not visiting the referred health facility.

Table 2: Association of socio-demographic & clinical characteristics with not getting TB diagnostic test (Sputum microscopy ,CXR, CBNAAT) among the **PTBPs** identified by ACF activity in Kolkata Municipal Corporation, West Bengal, India during Jul- Dec 2018, N= 1132

Variable	Total	Not visiting health facility, n (%)*	Unadjusted RR (95% CI)	Adjusted RR (95% CI)\$
Total	1132	419 (37.0)		
Age in years				
0-14	36	10 (27.8)	1	1
15-29	206	63 (30.6)	1.1 (0.6-1.9)	1.3 (0.7-2.2)
30-44	323	117 (36.2)	1.3 (0.8-2.3)	1.5 (0.8-2.5)
45-59	280	93 (33.2)	1.2 (0.7-2.1)	1.4 (0.8-2.3)
60-74	184	64 (34.8)	1.3 (0.7-2.2)	1.4 (0.8-2.5)
75 & above	32	4 (12.5)	0.5 (0.2-1.3)	0.5 (0.2-1.5)
Not recorded	71	68 (95.8)	3.4 (2.0-5.9)	3.3 (1.9-5.6)
Gender				
Male	647	211 (32.6)	1	1
Female	485	208 (42.9)	1.3 (1.1-1.5)	1.3 (1.2-1.6)
Alcohol use				
Yes	45	28 (62.2)	1.7 (1.4-2.2)	1.7 (1.3-2.2)
No	1087	391 (36.0)	1	1
Tobacco use				
Yes	175	74 (42.3)	1.2 (1.0-1.4)	1.2 (1.0-1.4)
No	957	345 (36.1)	1	1
Presenting Symptom				
Cough with other symptoms	101	26 (25.7)	1.1 (0.7-1.6)	1.1 (0.8-1.6)
Only Cough	687	312 (45.4)	1.9 (1.6-2.4)	1.8 (1.5-2.2)
No cough but other symptoms	344	81 (23.5)	1	1
Previous history of TB				
Yes	84	27 (32.1)	1	1
No	1048	392 (37.4)	1.2 (0.8-1.6)	1.0 (0.8-1.4)
Family history of TB				
Yes	118	70 (59.3)	1.7 (1.5-2.0)	1.5 (1.3-1.8)
No	1014	349 (34.4)	1	1
Diabetes				
Yes	244	63 (25.8)	1	1
No/Unknown	888	356 (40.1)	1.6 (1.2-1.9)	1.3 (1.0-1.6)
HIV				
Yes	13	1 (7.7)	1	1
No/Unknown	1119	418 (37.4)	4.9 (0.7-32.0)	6.0 (0.9-39.8)

* Row percentage, \$ cluster adjusted (wards) generalized linear (Poisson) model

Abbreviation: TB- Tuberculosis; PTBP- Presumptive TB patient; RR- Relative Risk; aRR- Adjusted Relative Risk; CI- Confidence Interval; ACF- Active Case Finding

Of the 713 PTBPs visiting health facility for diagnosis, 177 (24.8%) were diagnosed with TB. Thus, the number needed to screen (NNS) one TB was 2,183 (386242/177) individuals. Aged less than 30 years (aRR-2.4 (95% CI: 1.6-3.5)), presence of cough along with other symptoms (aRR-3.3 (95% CI: 2.4-4.4)), no previous history of TB (aRR-3.6 (95% CI: 1.7-7.6)) and family history of TB (aRR-2.7 (95% CI: 2.0-3.6)) were independently associated with presence of TB (**Table-3**).

Table 3: Association of socio-demographic & clinical characteristics with getting positive TB diagnosis among who undergone TB testing during the ACF activity in Kolkata Municipal Corporation, West Bengal, India during Jul- Dec 2018, N= 713

Variable	Total	Diagnosed as TB, n (%) *	Unadjusted RR (95% CI)	Adjusted RR (95% CI) ^{\$}
Total	713	177 (24.8)		
Age in years				
0-14	26	10 (38.5)	2.7 (1.5-4.8)	2.2 (1.2-3.9)
15-29	143	70 (49.0)	3.4 (2.3-5.0)	2.4 (1.6-3.5)
30-44	206	47 (22.8)	1.6 (1.0-2.4)	1.3 (0.9-2.0)
45-59	187	27 (14.4)	1	1
60-74	120	19 (15.8)	1.1 (0.6-1.9)	1.0 (0.6-1.6)
75 & above	28	3 (10.7)	0.7 (0.2-2.3)	0.7 (0.2-1.8)
Not recorded	3	1 (33.3)	2.3 (0.4-11.9)	2.4 (0.4-14.2)
Gender				
Male	436	95 (21.8)	1	1
Female	277	82 (29.6)	1.4 (1.1-1.8)	1.1 (0.9-1.4)
Alcohol user				
Yes	17	4 (23.5)	1	1
No	696	173 (24.9)	1.1 (0.4-2.5)	1.3 (0.7-2.5)
Tobacco user				
Yes	101	9 (8.9)	1	1
No	612	168 (27.5)	3.1 (1.6-5.8)	2.2 (1.1-4.2)
Presenting Symptoms				
Cough with other symptoms	75	50 (66.7)	4.4 (3.3-5.8)	3.3 (2.4-4.4)
Only Cough	375	57 (15.2)	1	1
No cough but other symptoms	263	70 (26.6)	1.8 (1.3-2.4)	1.4 (1.1-2.0)
Previous history of TB				
Yes	57	7 (12.3)	1	1
No	656	170 (25.9)	2.1 (1.0-4.3)	3.6 (1.7-7.6)
Family history of TB				
Yes	48	29 (60.4)	2.7 (2.1-3.6)	2.7 (2.0-3.6)
No	665	148 (22.3)	1	1
Diabetes				
Yes	181	54 (29.8)	1.3 (1.0-1.7)	1.2 (0.9-1.5)
No	532	123 (23.1)	1	1
HIV				
Yes	12	1 (8.3)	1	0.8 (0.1-5.8)
No	701	176 (25.1)	3.0 (0.5-19.8)	1

* Row percentage

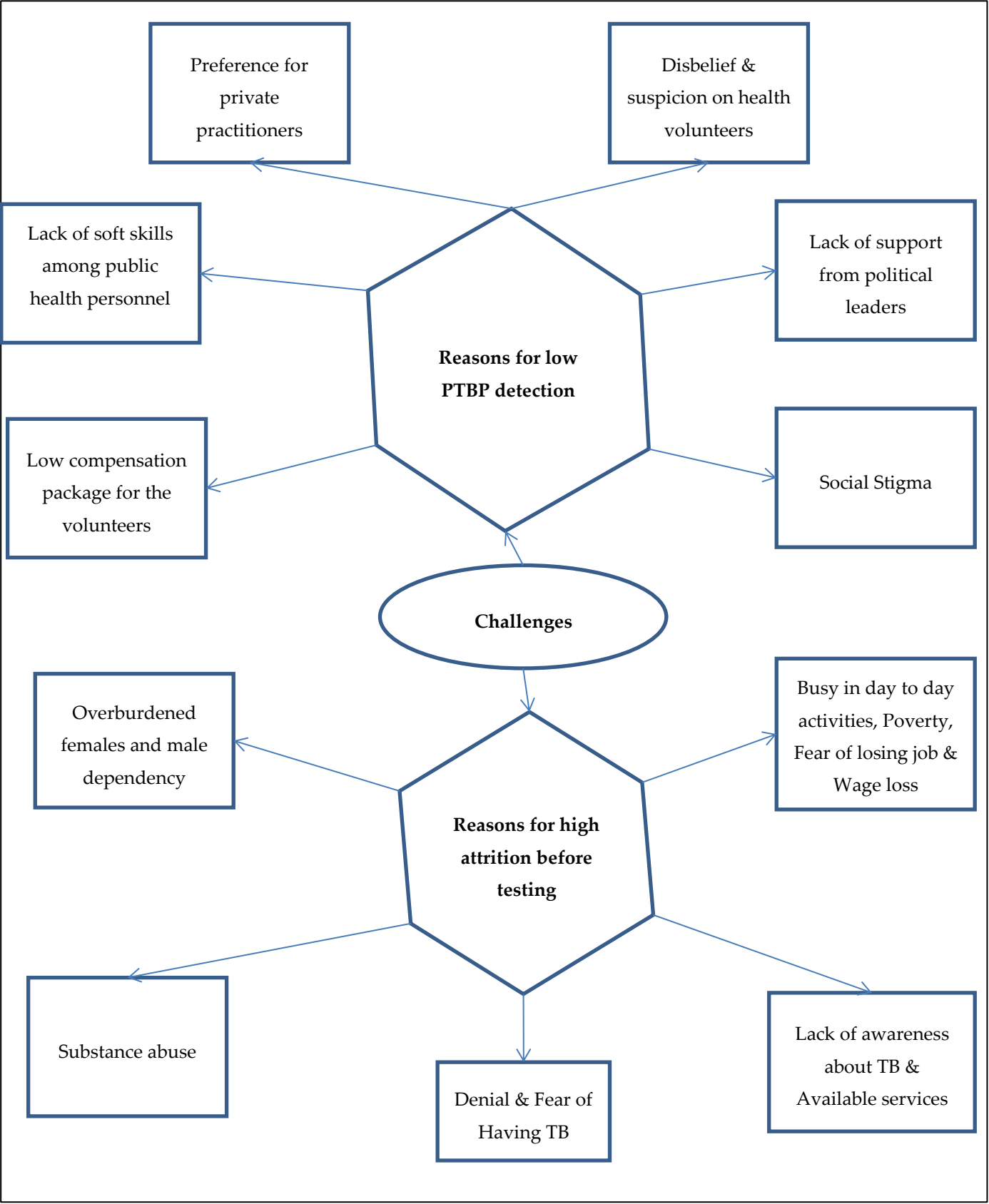
\$ cluster adjusted (wards) generalized linear (Poisson) model

Abbreviation: TB- Tuberculosis; RR- Relative Risk; aRR- Adjusted Relative Risk; CI- Confidence Interval; ACF- Active Case Finding; r= row percentage

3.2. Qualitative

We deduced 38 codes which were clubbed into eleven categories. The categories are described below under the two broad themes, 1) Challenges in house to house visit and identification of PTBP and 2) Challenges in visiting referred health facility for TB diagnosis. **(Figure-4)**

Figure 4: Themes and categories related to challenges in implementation of ACF activity in high TB burden wards of Kolkata, 2018



3.2.1. Challenges in house to house visit and identification of PTBP:

The codes related to this theme were grouped and summarized into six categories.

Social stigma:

The TAs reported that there is social stigma about TB and is found to be deterrent for the activity. They felt people don't disclose the presence of PTBP in their house if neighbors are around during the interview.

"When first time we visit any community or household & asked about whether anybody having symptoms like TB, usually none responds on spot. But after coming back they call us that there is a TB suspect in their family.

At the time of my visit there were many people so they didn't tell that time." - 38 years old female TA

Lack of support from political leaders:

The TAs reported interference of local political leaders in their activity. The political leaders deny the presence of disease in their wards and ask the TAs not to conduct ACF activity.

"They ask 'why you people come to my house often? I don't have time to entertain your queries. There is no TB in my Ward', despite having lots TB patients in and around the party office!" - 27 year old female TA

Suspicion about TAs:

The TAs felt that people were suspicious about them. The people were not aware of the ACF activity and assumed that the TAs visited the houses with some hidden agenda.

"The community dwellers & the local club initially did not believe that there are many TB patients in their area & house to house visit is needed to detect more TB case. They were thinking that we have some other motive!" -

55 year old female TA

Preference for private practitioners:

The TAs felt that the community has preference for private healthcare providers. Thus, they don't want disclose the details of their symptoms with TAs as they relate them to be from public health system.

"For any illness, here, people prefer to go to Private clinic if they can afford. If cant spent money then prefer to go to homeopath, unani or even quacks. For maintaining privacy also sometime they prefer private clinic. They don't tell us anything." - 28 year old female TA

Lack of soft skills among public health personnel's:

The TAs felt inappropriate way of handling patients at the public health facilities further dampen their relationship with the community. They refer the patient to health facilities and patients are treated badly

"This is actually the biggest challenges for us to get the report timely. When parents visit TU for CBNAAT report, the person sitting at TU sometime behaves rudely & sends back the parents. Sometime they say this test cost 1 lakh so you have to wait!" —38 years old female TA

Unjustified pay to the job done:

The TAs complained that they are paid less for the amount of efforts they put in the field. They also felt, low salary as one of the reason for high attrition among the TAs.

"What can we do with this 3000? People in other project with similar job profile getting three four times the salary we are getting.. That's why many people are leaving this project... I've also applied for.." - 45 years old female TA

3.2.2. Challenges in PTBPs visit to referred health facility for TB diagnosis

The codes related to this theme were grouped and summarized into five categories.

Loss of wages and fear of losing jobs:

The TAs felt the loss of wages on missing a work-day among daily wage labors and non-availability of leaves for salaried employees hinders the PTBPs from visiting the referred health facility.

"..... she (a PTBP who had not gone for testing) said that she couldn't go DMC due to fear of wage loss. As sudden leave can trigger deduction of pay and employer may become angry! So, she planned for testing on any suitable next day." - 60 yrs female TA

Lack of awareness about the available TB services at health facilities:

The TAs reported that a few of the PTBPs they had identified had difficulties in identifying health facility in their locality to avail TB services.

"Till now people are not getting government services in my area , that is ward no. XX, they are not aware of the available government services." —45 years female TA

Denial and apprehension of having TB disease:

The TAs reported that those PTBPs who have awareness about TB are apprehensive to get themselves tested for the disease. Such patients try to deny that they may have disease and avoid going to health facility.

"I was unable to arrange for sputum test for all identified presumptive case. Because I couldn't convince few patients that sputum test is needed for them. Patients actually don't want to believe that they may have TB."—53 years old female TA

Substance abuse:

The TAs reported that the substance abuse among PTBPs as one of the potential reason for them not visiting the health facilities. They felt it was difficult to get the investigation done for drug addicts and alcoholics.

"Not all patients have gone for testing. Especially who are addicted with hashish (Marijuana), have not gone to DMC, not even given the phone number. I have at least three such case. Even after detection of TB many people, especially the rickshaw-puller & the ganja addicts don't wear mask (Govt used to provide the mask after AFB +ve)."- 45 years old female TA

Overburdened females and male dependency:

The TAs felt it is difficult to mobilize female patients for health facility as they are pre-occupied with household works during the day time. Also, they are dependent on males to take them to health facilities.

"I have told many times to my husband but he said that he will arrange but he is yet to arrange. I don't know where to go so I did not go alone."- 29 year old PTBP

4. Discussion

This is one of the first mixed methods study to assess the 'care cascade' and explore the challenges in implementation of ACF for TB in India. Our study has a few key findings. First, the ACF activity had low PTBP detection rate with only three out of thousand apparently healthy individuals enumerated during house-to-house visit having symptoms suggestive of TB. Second, there was high attrition of PTBP prior to diagnosis with only two out of three PTBP referred visiting the health facility. Third, among PTBPs who undergo diagnostic test, the sputum positivity rate was high with one out four having diagnosed with TB. Fourth, the NNS for one TB patient was relatively high. Fifth, the stigma about TB, suspicion about TA, lack of support from political leaders and low salary for TAs were the major challenges in conducting house-to-house visit for identifying the PTBPs. Sixth, the wage loss, apprehension about the diagnosis, substance abuse and male dependency of the females for seeking care at health facility were the major challenges for PTBPs to visit the referred health facility.

The study has several strengths. First, it was conducted under programmatic settings and reflects the field realities. Second, we used mixed methods design which provided considerable insights on challenges in implementation of ACF and enabled us to interpret the quantitative results. Third, the study has a relatively large sample size and there was no selection bias as we included all the individuals enumerated during study reference period in sixty wards where ACF is implemented. Fourth, we used double data entry and validation using EpiData software and thus data entry errors were minimized. Fifth, we adhered to STrengthening the Reporting of Observational studies in Epidemiology (STROBE) and COREQ guidelines for reporting the study findings.

The study has a few limitations. First, as details of diagnostic tests that PTBPs underwent were not recorded routinely, we failed to determine the uptake of various diagnostic tests for TB. Second, as it was as a retrospective study using routinely collected data, we failed to capture and adjust in our model for potential confounders like socioeconomic status, distance from referred health facility, employment status, education level and marital status. Thus, the factors associated with 'not visiting health facility' among PTBPs have to be interpreted with caution. Third, the qualitative exploration was limited only to TAs & PTBPs. Thus, we failed to capture experiences of other stakeholders involved in ACF activity like field supervisor, project leads & programme managers of NTP.

We further discuss our study findings in detail. First, the study had low rate of PTBP detection, whereas the previous studies on ACF have consistently shown high rates of PTBP detection[23–25]. The TAs failed to enquire about symptoms suggestive of TB individually from all those enumerated. As the ACF activity was conducted during the daytime, the symptoms of individuals away from home were enquired from adult informants. The studies in the past have shown that the detection rate of PTBP could be low if the symptoms are not elicited individually[26]. The informants might not be aware of the TB symptoms and thus, could have failed to identify the PTBPs among those absent during interview. As explored in qualitative study, the other potential reasons for low rate of PTBP detection were hesitancy to disclose the details of TB symptoms due to stigma and distrust about TAs.

Second, the high attrition rate among PTBPs referred to health facilities for TB diagnosis. The previous studies have also reported similar findings with attrition rates ranging from 39% to 80%[9,10,27,28]. The rate was high in spite the repeated house visits by TAs to counsel and

encourage PTBPs to visit health facility. The qualitative interviews provided good insight about potential reasons for such high attrition. The wage or work loss due to travel and visit to health facilities during day time was one of the major reasons. Among the females, the attrition rate was high as they were dependent on male family members to visit health facility. The difficulties in counselling PTBPs with alcohol and other substance abuse also contributed to attrition. The other important issue was general distrust and unhappiness about public health system among public. Thus, these reasons warrants for provision of financial support and assistance to PTBPs to undergo diagnostic tests.

Third, the rate of TB diagnosis among those PTBPs visiting health facility was high. This could be due to the fact that only those with perceived need reached the health facility. Though PTBPs were identified by ACF activity, the decision to visit largely depended on severity of symptoms and thus mimicked the passive case finding. Also, the TAs might have given more preference to referral and linkage of PTBPs with severe and multiple symptoms rather than those with mild symptoms.

Fourth, the overall yield of ACF was low and NNS for TB was high. The previous studies have reported lower NNS on mass screening in high burden TB settings. The potential reason for high NNS might be the low detection and high attrition rate of PTBPs. Though, the NNS is high, the activity is still worth as it indirectly sensitizes the households on TB symptoms and thus creates awareness in community. The regular visit to the same community (six days a week) is gradually leading to good rapport building with the community[16]. This might improve the overall case detection in the community and contribute in TB control efforts.

Based on the study findings there are few implications and recommendations for ACF activities. This model can be replicated in other high TB burden area considering the following recommendations.

First, the house visits needs to be scheduled in the non-working hours of the days to ensure presence of majority of the household members during visit. The feasibility of visits by healthcare workers during early morning or late evening hours needs to be explored. Also, revisits to houses on holidays can be made to probe for symptoms among those not present during initial visit.

Second, the sputum collection and transportation mechanisms need to be established. The healthcare workers can collect the sputum samples from the PTBPs and transport it to diagnostic facilities. Also, the evening clinics can be set up to provide chest x-ray services to working PTBPs. The mobile voice call or SMS reminders to PTBPs can be tried to improve uptake of diagnostic tests.

Third, there is need for establishing robust database management system for tracking of individual PTBPs and programme monitoring. The healthcare workers can enter the PTBPs details directly in the hand held data capture devices instead of documenting it in registers. This will reduce the efforts in digitalizing these registers and data loss. Provision of internet connectivity with the device can be made for real time data sharing & prompt corrective actions. The database also has to include details on investigations to assess the yield with combination of various diagnostic tests. The indicators like proportion of PTBPs detected and proportion of PTBPs undergoing diagnostic tests for TB can be included in the programme monitoring. The healthcare workers can be incentivized based on these indicators.

5. Conclusions

The yield of ACF was suboptimal with low PTBP identification rate and high attrition rate. Interviewing each & every individual of the households for symptoms of TB and supporting PTBPs for diagnosis through sputum collection and transport system can be adopted to improve the yield.

Supplementary Materials: The dataset is available on

https://www.dropbox.com/home?preview=ACF+study+Data+files_Abhijit.rar

Author Contributions:

Conceptualization (CO-PIs)- Abhijit Dey, Pruthu Thekkur, Sharath BN and Ayan Ghosh;
Data collection -Quantitative- Abhijit Dey, Tanusree Dasgupta, Isita Roy, Anuradha Choudhury, Dibyendu Chatterjee and Namita Dimari;
Data collection –Qualitative- Abhijit Dey and Tanusree Dasgupta;
Data entry- Abhijit Dey, Namita Dimari and Dibyendu Chatterjee;
Data curation- Abhijit Dey, Pruthu Thekkur, Ayan Ghosh, Tanusree Dasgupta, Vivek Sharma, Namita Dimari, Dibyendu Chatterjee, Isita Roy, Anuradha Chanda, Brojokishore Saha and Sharath Nagaraja;
Formal analysis- Abhijit Dey, Pruthu Thekkur, Chidananda SV, Vivek Sharma, Namita Dimari and Sharath Nagaraja;
Funding acquisition-Pruthu Thekkur;
Investigation-Abhijit Dey, Ayan Ghosh, Tanusree Dasgupta, Soumyajyoti Bandopadhyay, Arista Lahiri, Milan Dinda, Vivek Sharma and Anuradha Chanda;
Methodology-Abhijit Dey, Pruthu Thekkur, Soumyajyoti Bandopadhyay, Arista Lahiri, Chidananda SV, Sanghamitra Ghosh and Sharath Nagaraja;
Project administration-Milan Dinda, Isita Roy, Parthiban Shanmugam and Brojokishore Saha;
Resources- Tanusree Dasgupta, Milan Dinda, Vivek Sharma, Isita Roy, Parthiban Shanmugam and Brojokishore Saha;
Software-Abhijit Dey, Pruthu Thekkur, Chidananda SV, Vivek Sharma, Namita Dimari, Dibyendu Chatterjee and Sharath Nagaraja;
Supervision-Ayan Ghosh, Tanusree Dasgupta, Soumyajyoti Bandopadhyay, Milan Dinda, Anuradha Chanda, Parthiban Shanmugam, Brojokishore Saha, Sanghamitra Ghosh and Sharath Nagaraja;
Validation-Abhijit Dey, Pruthu Thekkur, Tanusree Dasgupta, Milan Dinda, Vivek Sharma, Anuradha Chanda, Parthiban Shanmugam and Sharath Nagaraja;
Visualization-Abhijit Dey, Pruthu Thekkur, Ayan Ghosh, Soumyajyoti Bandopadhyay, Arista Lahiri, Chidananda SV, Vivek Sharma, Dibyendu Chatterjee, Parthiban Shanmugam and Sharath Nagaraja;
Writing – original draft, Abhijit Dey, Pruthu Thekkur and Sharath Nagaraja;
Writing – review & editing-Ayan Ghosh, Tanusree Dasgupta, Soumyajyoti Bandopadhyay, Arista Lahiri, Chidananda SV, Milan Dinda, Vivek Sharma, Namita Dimari, Dibyendu Chatterjee, Isita Roy, Anuradha Chanda, Parthiban Shanmugam, Brojokishore Saha and Sanghamitra Ghosh.
Appearing before the local ethics committee- Ayan Ghosh, Soumyajyoti Bandopadhyay and Arista Lahiri
Mentoring- Pruthu Thekkur, Sharath BN, Milan Dinda, Sanghamitra Ghosh and Brojo Kishore Saha

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References

1. World Health Organization *GLOBAL TUBERCULOSIS REPORT 2018*; WHO, Geneva, Switzerland, 2018;
2. Ho, J.; Fox, G.J.; Marais, B.J. Passive case finding for tuberculosis is not enough. *Int. J. mycobacteriology* **2016**, *5*, 374–378.
3. Golub, J.E.; Mohan, C.I.; Comstock, G.W.; Chaisson, R.E. Active case finding of tuberculosis: historical perspective and future prospects. *INT J TUBERC LUNG DIS* 9(11)1183–1203 © 2005 *Union Rev.* **2005**.
4. World Health Organization The End TB Strategy. (2015). WHO, Geneva, Switzerland. http://www.who.int/tb/post2015_TBstrategy.pdf (Accessed 23rd November 2018).
5. Lönnroth, K.; Tomlin, K.; Afnan-Holmes, H.; Schaap, A.; Golub, J.E.; Shapiro, A.E.; Glynn, J.R.; Kranzer, K.; Corbett, E.L.; Afnan-Holmes, H.; et al. The benefits to communities and individuals of screening for active tuberculosis disease: a systematic review [State of the art series. Case finding/screening. Number 2 in the series]. *Int. J. Tuberc. Lung Dis.* **2013**, *17*, 432–446.
6. Mhimbira, F.A.; Cuevas, L.E.; Dacombe, R.; Mkopi, A.; Sinclair, D. Interventions to increase tuberculosis case detection at primary healthcare or community level services. *Cochrane Database Syst. Rev.* **2015**, 2015.
7. Shewade, H.D.; Gupta, V.; Satyanarayana, S.; Kharate, A.; Sahai, K.N.N.; Murali, L.; Kamble, S.; Deshpande, M.; Kumar, N.; Kumar, S.; et al. Active case finding among marginalised and vulnerable populations reduces catastrophic costs due to tuberculosis diagnosis. *Glob. Health Action* **2018**, *11*, 1494897.
8. Shewade, H.D.; Gupta, V.; Satyanarayana, S.; Pandey, P.; Bajpai, U.N.; Tripathy, J.P.; Kathirvel, S.; Pandurangan, S.; Mohanty, S.; Ghule, V.H.; et al. Patient characteristics, health seeking and delays among new sputum smear positive TB patients identified through active case finding when compared to passive case finding in India. *PLoS One* **2019**, *14*, e0213345.
9. B. M. Prasad, S. Satyanarayana, S. S. Chadha, A. Das, B. Thapa, S. Mohanty, S. Pandurangan, E. R. Babu, J. Tonsing, K.S.S. Experience of active tuberculosis case finding in nearly 5 million households in India. *pu* **2016**, *6*.
10. Myint, O.; Saw, S.; Isaakidis, P.; Khogali, M.; Reid, A.; Hoa, N.B.; Kyaw, T.T.; Zaw, K.K.; Khaing, T.M.M.; Aung, S.T. Active case-finding for tuberculosis by mobile teams in Myanmar: Yield and treatment outcomes. *Infect. Dis. Poverty* **2017**, *6*.
11. Central TB Division Directorate General of Health Services *Revised National Tuberculosis Control Programme: National strategic plan for tuberculosis elimination 2017–2025*; 2017;
12. USAID U.S. Announces \$21 Million in Awards to Address and Treat TB in India | U.S. Embassy & Consulates in India Available online:

<https://in.usembassy.gov/u-s-announces-21-million-in-awards-to-address-and-treat-tb-in-india/> (accessed on Aug 11, 2019).

13. The Union *Abstract Book-47th World Conference on Lung Health of the International Union Against Tuberculosis and Lung Disease*; 2016;
14. Saha, I.; Paul, B. Private sector involvement envisaged in the National Strategic Plan for Tuberculosis Elimination 2017–2025: Can Tuberculosis Health Action Learning Initiative model act as a road map? *Med. J. Armed Forces India* **2019**, *75*, 25–27.
15. Official Website of Kolkata Municipal Corporation KMC signs MOU with USAID funded project THALI Available online: https://www.kmcgov.in/KMCPortal/outside_jsp/THALI_18_07_2017.jsp (accessed on Dec 10, 2018).
16. Nari-O Sishu-Kalyan Kendra Tuberculosis Health Action Learning initiative (THALI) | www.noskk.in Available online: http://www.noskk.in/?post_causes=tuberculosis-health-action-learning-initiative-thali (accessed on Aug 10, 2019).
17. Creswell, J.W.; Plano Clark, V.L. *Designing and conducting mixed methods research*; SAGE Publications, 2007; ISBN 1412927927.
18. Census 2011; Government of India Kolkata City Population Census 2011 | West Bengal.
19. Department of Municipal Affair Government of West Bengal Total number of slum Pockets , Slum population percentage of Slum population , total number of CDS , NHC and NHG formed in the urban local bodies. 1–3.
20. Kolkata Municipal Corporation Official Website of Kolkata Municipal Corporation Available online: <https://www.kmcgov.in/KMCPortal/jsp/KMCHealthServiceHome.jsp> (accessed on Aug 10, 2019).
21. Central TB Division TOG-Chapter 3-Case finding & diagnosis strategy :: Ministry of Health and Family Welfare Available online: <https://tbcindia.gov.in/showfile.php?lid=3216> (accessed on Aug 10, 2019).
22. Tong, A.; Sainsbury, P.; Craig, J. Consolidated criteria for reporting qualitative research (COREQ): a 32-item checklist for interviews and focus groups. *Int. J. Qual. Heal. Care* **2007**, *19*, 349–357.
23. Aye, S.; Majumdar, S.S.; Minn Oo, M.; Tripathy, J.P.; Satyanarayana, S.; Thu, N.; Kyaw, T.D.; Wut, K.; Kyaw, Y.; Lynn Oo, N.; et al. Evaluation of a tuberculosis active case finding project in peri-urban areas, Myanmar: 2014–2016. *Int. J. Infect. Dis.* **2018**, *70*, 93–100.
24. Lorent, N.; Choun, K.; Malhotra, S.; Koeut, P.; Thai, S.; Eam Khun, K.; Colebunders, R.; Lynen, L. Challenges from Tuberculosis Diagnosis to Care in Community-Based Active Case Finding among the Urban Poor in Cambodia: A Mixed-Methods Study. **2015**.

25. Oshi, D.; Omeje, J.; Oshi, S.; Alobu, I.; Chukwu, N.; Nwokocha, C.; Emelumadu, O.; Ogbudebe, C.; Meka, A.; Ukwaja, K. An evaluation of innovative community-based approaches and systematic tuberculosis screening to improve tuberculosis case detection in Ebonyi State, Nigeria. *Int. J. Mycobacteriology* **2017**, *6*, 246.
26. Prasad, B.M.; Satyanarayana, S.; Chadha, S.S. Lessons learnt from active tuberculosis case finding in an urban slum setting of Agra city, India. *Indian J. Tuberc.* **2016**, *63*, 199–202.
27. Delva, G.J.; Francois, I.; Claassen, C.W.; Dorestan, D.; Bastien, B.; Medina-Moreno, S.; Fort, D. St.; Redfield, R.R.; Buchwald, U.K.; St Fort, D.; et al. Active Tuberculosis Case Finding in Port-au-Prince, Haiti: Experiences, Results, and Implications for Tuberculosis Control Programs. *Tuberc. Res. Treat.* **2016**, *2016*, 1–11.
28. Armstrong-Hough, M.; Turimumahoro, P.; Meyer, A.J.; Ochom, E.; Babirye, D.; Ayakaka, I.; Mark, D.; Ggita, J.; Cattamanchi, A.; Dowdy, D.; et al. Drop-out from the tuberculosis contact investigation cascade in a routine public health setting in urban Uganda: A prospective, multi-center study. *PLoS One* **2017**, *12*, e0187145.