The Prevalence of Latent Tuberculosis Infection and Smear Positive Pulmonary Tuberculosis in People with Household Close Contact with Tuberculosis in North of Iran

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Abstract
One of the recommended strategies for preventing tuberculosis is to screen high-risk populations with respect to Mycobacterium tuberculosis (TB) infection. The aim of the present study was to investigate latent infection and active tuberculosis in people with close household contact. It was a cross-sectional descriptive, analytical study with the sample size of 668 people from homes with one infected resident. In order to diagnose tuberculosis latent infection, the PPD test was done. To determine patients with smear-positive pulmonary tuberculosis, three sputum samples were taken from every patient and were examined using direct microscopy and culture. Data was analyzed by SPSS20 software. The prevalence of latent tuberculosis infection and smear-positive pulmonary tuberculosis were 42.8% and 0.9% respectively. The prevalence of latent tuberculosis infection and smear-positive pulmonary tuberculosis in people with close household contact were less than that of other studies. However, smear-positive pulmonary tuberculosis in people with close household contact was 199.5 times more than that of the general population.

Keywords
- Tuberculosis
- Household close contact
- Iran
- Prevalence

Introduction
Mycobacterium tuberculosis (TB) is mainly an airborne pathogen that is transferred from one person to another through respiration. About one third of the world’s population is estimated to be infected with Mycobacterium TB. Factors such as the source case, organism, environment and the person who has had contact with the source case determines the transfer of TB and the development of a new case.

It has been proven that almost 30% of people with patient contact show some symptoms of this infection and half of them will develop the disease in the first two years. Previous studies have reported that latent infection and tuberculosis is more prevalent in home residents who have had contact with smear-positive pulmonary tuberculosis patients. In a study carried out in one of the rural areas of Pakistan, the prevalence of positive tuberculin skin test was 49.4% among people exposed to TB patients. One of the recommended strategies for preventing tuberculosis...
is to screen high-risk populations regarding Mycobacterium TB infection in order to identify people in need for chemoprophylaxis or treatment. Among those, people with close contact with patients are one of the high-risk groups regarding Mycobacterium TB infection. Hence, to offer suitable recommendations and guidelines to develop and improve the objectives of the tuberculosis control program, relatives of such patients have to be examined. The aim of this study was to determine the prevalence of latent tuberculosis infection and smear-positive pulmonary tuberculosis in people with household close contact with tuberculosis in the north of Iran.

**Materials and Methods**

This cross-sectional study was carried out during 2010 and 2011 and the sampling was based on census. The populations under study were family members older than 6 years of age, having had contact with patients with smear-positive pulmonary tuberculosis in Mazandaran from 2006 to 2009. A "household close contact" was defined as someone who has lived for more than 30 days in the same house as an index case.

The number of enrolled people participating in this study was 668. The data collection tool was based on a checklist that included variables such as age, sex, skin induration test, coughing more than two weeks, coughing with sputum, weight loss, and the result of sputum smear microscopic examination.

After obtaining the agreement from the regional Health Deputy, an investigative team (including a physician, a disease expert, and a laboratory science expert) was formed and trained by our researchers to execute the investigation. Following the consent from the participants, demographic and field variables were inquired. Three sputum samples were taken from the individuals followed by direct sputum smear test at the Tuberculosis Laboratory of the City Health Department. The sputum culture was carried out at the Reference Laboratory of Mazandaran University of Medical Sciences. In order to ensure data accuracy of the sputum tests, 10% of negative sputum smear samples and 100% of positive sputum smear samples were re-examined at the Reference Laboratory. To evaluate smear-positive pulmonary tuberculosis, if one sputum smear sample and one sputum culture sample or at least two sputum smear samples of a person were positive, then that person was considered as a patient with smear-positive pulmonary tuberculosis. Moreover, all participants underwent a PPD test and their results were examined after 72 hours. If the induration diameter of the PPD test was 10 mm or more, this was considered as a latent tuberculosis infection.

Data was entered into the SPSS20 software and the results were analyzed using descriptive statistics including mean, standard deviation, frequency, and analytical methods, including Chi-square or Fisher’s exact test (for comparison with fewer than 5 expected observations per cell) and t test. Crude and adjusted odds ratios were also calculated.

**Results**

In total, 668 people being in close contact with smear-positive pulmonary tuberculosis patients were reviewed and examined; among which 44.5% (297 individuals) were men. The average age was 36.9±19.6. In men, the mean age was 33.9±18.8 and it was 39.3±19.9 in women. 1.5% of the participants had continuous cough for more than two weeks and 0.6% had weight loss. The prevalence of latent tuberculosis infection and smear-positive pulmonary tuberculosis was 42.8% and 0.9% respectively (table 1).

| Characteristic | N   | %     |
|---------------|-----|-------|
| Sex           |     |       |
| Male          | 297 | 44.5  |
| Female        | 371 | 55.5  |
| Age group     |     |       |
| 6-24          | 214 | 32    |
| 25-44         | 225 | 33.7  |
| ≥45           | 229 | 34.3  |
| Cough more than two weeks | | |
| Yes           | 10  | 1.5   |
| No            | 658 | 98.5  |
| Weight loss   |     |       |
| Yes           | 4   | 0.6   |
| No            | 664 | 99.4  |
| PPD test      |     |       |
| <10 mm        | 382 | 57.2  |
| ≥10 mm        | 286 | 42.8  |
| Result of sputum smear | | |
| Yes           | 6   | 0.9   |
| No            | 662 | 99.1  |
Table 2 shows that among men and women, the prevalence of latent tuberculosis (41.8% vs. 43.7%) and smear-positive pulmonary tuberculosis (0.7% vs. 1.1%) had no significant difference (P=0.3 and P=0.4 respectively). The maximum prevalence of latent tuberculosis infection (47.7%) and smear-positive pulmonary tuberculosis (1.7%) was found in people aged 6-24 years and ≥45 years respectively, but these differences were not significant. People with continuous coughs for more than two weeks were meaningfully more than those without it in people with latent tuberculosis infection (90% vs. 42.1%) (P=0.003) and in patients with smear-positive pulmonary tuberculosis (30% vs. 0.5%) (P<0.001). Moreover, people with weight loss were more than people without weight loss in the latent infection group (100% vs. 42.5%) and was also higher in people with smear-positive pulmonary tuberculosis (75% vs. 0.5%) (P=0.03 and P<0.001 respectively).

Discussion

Our findings show that the prevalence of both latent tuberculosis infection and smear-positive pulmonary tuberculosis among people with household close contacts was high in spite of the development of tuberculosis control program strategies.

Khalilzadeh et al., at Masih Daneshvari Pulmonary Diseases Research Center of Shahid Beheshti University (Iran),12 showed that the prevalence of latent tuberculosis infection and smear-positive pulmonary tuberculosis in people with close household contact was 38.2% and 4.8% respectively. Also, Khalilzadeh et al. in another study which was carried out during a two-year period (2002-2004) at the National Research Institute of Tuberculosis and Lung Disease of Iran, showed that among 224 close contacts, the initial Tuberculin Skin Test (TST) was positive (indurations>10 mm) in 16.5% of contacts and 7.6% had active positive smear.11

It is worth mentioning that the prevalence of smear-positive pulmonary tuberculosis among the general population of Mazandaran province was 4.5 in 100,000 people, 22.8 in 100,000 people in Golestan province (east of Mazandaran) and 7.7 in 100,000 people in Gilan Province (west of Mazandaran). Such incidence rates are less than the rate in people with household contacts.5,13

In one of the rural areas of Pakistan, the prevalence of positive tuberculin skin test (TST) was 49.4% among the people with close contact. In addition, the prevalence of Mycobacterium tuberculosis infection among the people with close household contacts in Spain and Peru was 44% and 55% respectively.2,3

In a systematic review that was carried out to determine the yield of household contact,14 the yield for all tuberculosis (bacteriologically confirmed and clinically diagnosed) was 4.5% (95% CI 4.3-4.8, I^2=95.5%) of the investigated contacts and for cases with bacteriological confirmation the yield was 2.3% (95% CI 2.1-2.5, I^2=96.6%). Latent tuberculosis infection was found in 51.4% (95% CI 50.6-52.2, I^2=99.4%) of the investigated contacts.14

One reason for this lower prevalence can be related to the year where other studies are carried out which were around the previous six years or prior to 2008. More recently, governments have made improvements in procedures and strategies in the tuberculosis control program in order to achieve the millennium development objectives.14 Another probable reason is the heterogeneity of people who participated in these studies. It is proved in various studies1,4,7,8 that the contact duration, age of exposed people, relative’s sleeping site, severity of sputum smear examination of patients and BCG Scar status of contact people were variables that predict tuberculosis infection among contact people. These variables, however, were not accounted for in some studies, including the present study.

In this research, the prevalence of tuberculosis

| Characteristic | PPD test with result of ≥10 N (%) | Positive sputum smear N (%) |
|---------------|---------------------------------|-----------------------------|
|              | Yes | No | P value | Yes | No | P value |
| Sex          |     |    |         |     |    |         |
| Male         | 124 (41.8) | 173 (58.2) | 0.3 | (2) | 0.7 | 295 (99.3) | 0.4 |
| Female       | 162 (43.7) | 209 (56.3) |     | (4) | 1.1 | 367 (98.9) |     |
| Age group    |     |    |         |     |    |         |
| 6-24         | 102 (47.7) | 112 (52.3) | 0.09 | (2) | 0.9 | 223 (99.1) | 0.1 |
| ≥45          | 100 (43.7) | 129 (56.3) |     | (4) | 1.7 | 225 (98.3) |     |
| Cough more than two weeks |     |    |         |     |    |         |
| Yes          | 9 (90) | 1 (10) | 0.003 | (3) | 0.5 | 655 (99.5) | <0.001 |
| No           | 277 (42.1) | 381 (57.9) |     | (3) | 0.5 | 1 (25) |     |
| Weight loss  |     |    |         |     |    |         |
| Yes          | 4 (100) | 0 |     | (3) | 0.5 | 661 (99.5) | <0.001 |
| No           | 282 (42.5) | 382 (57.5) | 0.03 |     | (3) | 0.5 |     |
infection had no difference in men and women and matched the results of a study done by Rathi et al. Moreover, in another study, the prevalence of latent tuberculosis infection and smear-positive pulmonary tuberculosis increased as age increased; which was in line with our study.

One of the limitations in the current study was in examining people with household close contact, where we did not study the effect of the exposure time, contact duration, continuous exposure or sleeping in one place while inquiring infection. Another limitation was its cross-sectional nature where temporality (cause-and-effect relationship) could not be recognized.

Conclusion

In this research, the prevalence of smear-positive pulmonary tuberculosis in people with household close contact was 898 out of 100,000 people. This is 199.5 times more than the general population of this region in Iran. Therefore, it can be declared that investigating household close contacts is necessary and of great importance in the national tuberculosis control program.

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Conflict of Interest: None declared.

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