Evaluation of Drug Resistance and Treatment Outcomes among TB Patients with Diabetes Mellitus at Tertiary Care Hospital of Sindh, Pakistan

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Authors’ contributions

This work was carried out in collaboration among all authors. Author KK designed the study, performed the statistical analysis, wrote the protocol and wrote the first draft of the manuscript. Authors AAU, KU and FQR managed the analyses of the study. Author MTB managed the literature searches. All authors read and approved the final manuscript.

ABSTRACT

The objective of the study is to evaluate of drug resistance and treatment outcomes among tuberculosis patients with diabetes mellitus at tertiary care hospital of Sindh, Pakistan. Tuberculosis (TB) is considered as very dangerous infectious disease caused by Mycobacterium tuberculosis other tubercle bacilli pathogen affects the lungs and it can be spread from person to person through very minute droplet released by infected person via coughing or sneezing. Descriptive cross-sectional study was carried out at different Tuberculosis setting located at tertiary
care hospital of Sindh, Pakistan for the period of 10 months. Tuberculosis OPD is considered as major health care facility for the TB, MDR-TB and XDR-TB patients for the local population and peripheries. Total 389 samples were collected through purposive sampling techniques. From the given data, there were 209 Males and 180 females, all participants belong to different areas, so 216 patients belong to rural areas, 121 study participants were house wives, 134 were employed and 41 were have their own business. From the clinical data, 229 participants shown positive response with TB smear test, whereas; 156 participants had positive results with TB culture test. 247 patients had previous history of Tuberculosis, 120 had developed drug resistance and 24 had developed multi-drug resistance (MDR). 312 patients were recently diagnosed with Diabetes mellitus and they had one year of duration of diabetes. Complication of tuberculosis developed among 113 patients and complication of diabetes developed among 194 participants. 39 people were using oral therapy for the management of diabetes and 341 participants had successfully completed their therapy and cured whereas 2 patients were died due to complication. It was concluded from the current research that there were many chances for developing drug resistance and multi-drug resistance among the patients suffering from co-morbid including tuberculosis along with Diabetes mellitus. Proper counseling should be conducted, in order to reduce the complication of either type of disease.

Keywords: Diabetes mellitus; tuberculosis; Multi-Drug Resistance (MDR); oral therapy.

1. INTRODUCTION

Internationally, approximately 9 million new cases of Tuberculosis (TB) reported every year and according to WHO reports there were almost 350 million people, who had diagnosed with diabetes mellitus [1,2]. Diabetes Mellitus is considered as major hazardous aspect for the development of tuberculosis cases [3,4] and almost 18% of active tuberculosis patients had diabetes mellitus [5]. There was strong correlation between Tuberculosis and Diabetes Mellitus as both disease possess high dominance globally and since last few decades, mortality and morbidity rates of diabetes increased enormously [2,6,7]. The cases of multi drug resistant tuberculosis (MDR-TB) are also increased very fast [8], so the significance of understanding about the correlation of diabetes and tuberculosis is also developing [9]. Literatures from India, Spain, Turkey and Pakistan elaborate that diabetes with active cases of tuberculosis was very much common among these countries. 25% of MDR-TB patients were having diabetes mellitus [10-13]. According to various studies there was strong correlation between diabetes and MDR-TB [14,15]. Due to lack of medical facilities and diagnostic criteria, there was very low data available for the MDR-TB patients with diabetes. Moreover, patients of MDR-TB had lack of knowledge and attitude towards the relation of Diabetes with other diseases and their control along with clinical impact [16,17]. Among the low socio-economic nations, the burden of MDR-TB is almost very high and frequency of diabetes patients is expected to grow in coming 20 years [16,7]. In 2020, the incidence of MDR-TB in Pakistan was 230 per 1 lac individual that is quite greater among south Asian countries [18]. On national level the patients of MDR-TB were never treated when previously diagnosed with tuberculosis that was considered as major risk factor for developing Multi Drug Resistance [8]. In the same case, the burden of Diabetes was also raised in Pakistan. The frequency of diabetic patients among the urban areas was quite greater than the local residents of rural areas [13]. As per national epidemic report of diabetes, the rate of cases of new diagnosed diabetes was high among the adults [19] and it is expected to enhance by 9% up to 2030 [20-23]. Diabetes and Tuberculosis routinely increase the burden on the population of Pakistan so it becomes very difficult to manage such types of co-morbidities simultaneously. [24-27] Although, it is necessary to understand the relationship between diabetes and tuberculosis. If tuberculosis is properly managed without creating any hindrance to management of diabetes so the chances of developing MDR-TB were also increased. [28-30] So, the aim of the study was to evaluate the characteristics of Tuberculosis patients’ assessment for MDR-TB along with and without Diabetes Mellitus and clinical aspect of Diabetes with medical care correlate with drug resistance among the patients with Diabetes and Tuberculosis and also determine the management of diabetes associated with the best TB regimen. [31-32]

2. METHODOLOGY

Descriptive cross-sectional study was carried out at different Tuberculosis setting located at tertiary...
care hospital of Sindh, Pakistan for the period of 10 months. Tuberculosis OPD is considered as major health care facility for the TB, MDR-TB and XDR-TB patients for the local population and peripheries. Total 389 samples were collected through purposive sampling techniques. All participants were facilitated with questionnaire and guided regarding its filling and data was collected and Statistical analysis was done by using SPSS version 20.00

3. RESULTS AND DISCUSSION

Descriptive cross-sectional study was carried out at different Tuberculosis setting located at tertiary care hospital of Sindh, Pakistan for the period of 10 months. Total 389 samples were collected through purposive sampling techniques; from them 209 were males and 180 females as mentioned in Table 1.

From all study subjects, 209 were belongs to rural areas where as 173 participants belongs to urban areas of Sindh as mentioned in Table.

All the study subjects were divided in accordance with different age groups as described in Table 3.

All the participants were also divided in the different groups according to their marital status as mentioned in Table 4.

Job status of study subjects were also described in Table 5.

Qualification of study subjects were also mentioned in Table 6.

| Table 1. Gender wise distribution of study subjects |
|----------------------------------|
| Frequency | Percent | Valid Percent | Cumulative Percent |
| Valid   | Male     | 209 | 53.7 | 53.7 | 53.7 |
|         | Female   | 180 | 46.3 | 46.3 | 100.0 |
| Total   |          | 389 | 100.0 | 100.0 | |

| Table 2. Area wise distribution of study subjects |
|----------------------------------|
| Frequency | Percent | Valid Percent | Cumulative Percent |
| Valid   | Rural    | 216 | 55.5 | 55.5 | 55.5 |
|         | Urban    | 173 | 44.5 | 44.5 | 100.0 |
| Total   |          | 389 | 100.0 | 100.0 | |

| Table 3. Age wise distribution of study subjects |
|----------------------------------|
| Frequency | Percent | Valid Percent | Cumulative Percent |
| Valid   | 15-24 Years | 79 | 20.3 | 20.3 | 20.3 |
| 24-34 Years | 134 | 34.4 | 34.4 | 54.8 |
| 35-44 Years | 86 | 22.1 | 22.1 | 76.9 |
| 45-54 Years | 40 | 10.3 | 10.3 | 87.1 |
| 55 and above | 50 | 12.9 | 12.9 | 100.0 |
| Total   |          | 389 | 100.0 | 100.0 | |

| Table 4. Marital Status wise distribution of study subjects |
|----------------------------------|
| Frequency | Percent | Valid Percent | Cumulative Percent |
| Valid   | Married  | 202 | 51.9 | 51.9 | 51.9 |
|         | Single   | 146 | 37.5 | 37.5 | 89.5 |
|         | Divorced/Separated | 41 | 10.5 | 10.5 | 100.0 |
| Total   |          | 389 | 100.0 | 100.0 | |

| Table 5. Job Status of study subjects |
|----------------------------------|
| Frequency | Percent | Valid Percent | Cumulative Percent |
| Valid   | House wives | 121 | 31.1 | 31.1 | 31.1 |
|         | Employed    | 134 | 34.4 | 34.4 | 65.6 |
|         | Un-employed | 93 | 23.9 | 23.9 | 89.5 |
|         | Landlord/Self Business | 41 | 10.5 | 10.5 | 100.0 |
| Total   |          | 389 | 100.0 | 100.0 | |
Tuberculosis was measured through TB Smear test and the results of TB smear test was mentioned in Table 7.

Some time smear test didn’t get positive results of TB so for the confirmation of TB, culture test was also conducted and the resultant of Culture test was mentioned in Table 8

History of TB among the patients was mentioned in Table 9.

Patients who had already taken the treatment of TB and their results were mentioned in Table 10.

With irregular drug therapy, number of participants had developed various drug resistant including Pan susceptible, drug resistant, Multi Drug Resistant and XDR and their data is mentioned in Table 11.

TB patients had developed many symptoms such as weight loss, dyspnea, blood in sputum, dry coughing, yellowish eyes, dry mouth, abdominal cramps and results for each symptoms were mentioned in Table 12.

Various risk factors were also observed among the TB patients and the results of each risk factor is mentioned in Table 13.

Each participants had different diabetes types such as Type I and Type II and the results for Types of diabetes is mentioned in Table 14.

### Table 6. Qualification wise distribution of study subjects

|                | Frequency | Percent | Valid percent | Cumulative percent |
|----------------|-----------|---------|---------------|--------------------|
| Valid Primary  | 203       | 52.2    | 52.2          | 52.2               |
| Secondary      | 90        | 23.1    | 23.1          | 75.3               |
| Graduation     | 88        | 22.6    | 22.6          | 97.9               |
| Post graduation| 8         | 2.1     | 2.1           | 100.0              |
| Total          | 389       | 100.0   | 100.0         |                    |

### Table 7. Tuberculosis Smear Test results of study subjects

|                | Frequency | Percent | Valid Percent | Cumulative Percent |
|----------------|-----------|---------|---------------|--------------------|
| Valid Positive | 229       | 58.9    | 58.9          | 58.9               |
| Negative       | 160       | 41.1    | 41.1          | 100.0              |
| Total          | 389       | 100.0   | 100.0         |                    |

### Table 8. Tuberculosis Culture Test results of study subjects

|                | Frequency | Percent | Valid Percent | Cumulative Percent |
|----------------|-----------|---------|---------------|--------------------|
| Valid Positive | 156       | 40.1    | 40.1          | 40.1               |
| Negative       | 233       | 59.9    | 59.9          | 100.0              |
| Total          | 389       | 100.0   | 100.0         |                    |

### Table 9. Previous history of Tuberculosis of study subjects

|                | Frequency | Percent | Valid Percent | Cumulative Percent |
|----------------|-----------|---------|---------------|--------------------|
| Valid Yes      | 247       | 63.5    | 63.5          | 63.5               |
| NO             | 142       | 36.5    | 36.5          | 100.0              |
| Total          | 389       | 100.0   | 100.0         |                    |

### Table 10. Previously taken Tuberculosis treatment by study subjects

|                | Frequency | Percent | Valid Percent | Cumulative Percent |
|----------------|-----------|---------|---------------|--------------------|
| Valid Yes      | 201       | 51.7    | 51.7          | 51.7               |
| No             | 188       | 48.3    | 48.3          | 100.0              |
| Total          | 389       | 100.0   | 100.0         |                    |
Table 11. Drug Susceptibility of study subjects

| Frequency | Percent | Valid Percent | Cumulative Percent |
|-----------|---------|---------------|--------------------|
| Valid Pan Susceptible | 243 | 62.5 | 62.5 | 62.5 |
| Drug Resistance | 120 | 30.8 | 30.8 | 93.3 |
| Multi Drug Resistance | 24 | 6.2 | 6.2 | 99.5 |
| Extended Drug Resistance | 2 | .5 | .5 | 100.0 |
| Total | 389 | 100.0 | 100.0 | 100.0 |

Table 12. Symptoms of Tuberculosis found among study subjects

| Frequency | Percent | Valid Percent | Cumulative Percent |
|-----------|---------|---------------|--------------------|
| Valid weight loss | 22 | 5.7 | 5.7 | 5.7 |
| Dyspnea | 88 | 22.6 | 22.6 | 28.3 |
| Blood in Sputum | 83 | 21.3 | 21.3 | 49.6 |
| Dry Coughing | 118 | 30.3 | 30.3 | 79.9 |
| Yellowish Eyes | 19 | 4.9 | 4.9 | 84.8 |
| Dry Mouth | 40 | 10.3 | 10.3 | 95.1 |
| Abdominal Cramps | 19 | 4.9 | 4.9 | 100.0 |
| Total | 389 | 100.0 | 100.0 | 100.0 |

Table 13. Risk factors of Tuberculosis

| Frequency | Percent | Valid Percent | Cumulative Percent |
|-----------|---------|---------------|--------------------|
| Valid HIV Positive | 100 | 25.7 | 25.7 | 25.7 |
| TB Contact at Home | 138 | 35.5 | 35.5 | 61.2 |
| MDR-TB contacts at Home | 28 | 7.2 | 7.2 | 68.4 |
| Tobacco consumption | 80 | 20.6 | 20.6 | 88.9 |
| Alcohol Consumption | 43 | 11.1 | 11.1 | 100.0 |
| Total | 389 | 100.0 | 100.0 | 100.0 |

Table 14. Types of Diabetes among study subjects

| Frequency | Percent | Valid Percent | Cumulative Percent |
|-----------|---------|---------------|--------------------|
| Valid TYPE-I | 77 | 19.8 | 19.8 | 19.8 |
| TYPE-II | 312 | 80.2 | 80.2 | 100.0 |
| Total | 389 | 100.0 | 100.0 | 100.0 |

Table 15. Duration of Diabetes mellitus among study subjects

| Frequency | Percent | Valid Percent | Cumulative Percent |
|-----------|---------|---------------|--------------------|
| Valid Below one year | 192 | 49.4 | 49.4 | 49.4 |
| Below 05 years | 182 | 46.8 | 46.8 | 96.1 |
| Above 05 years | 15 | 3.9 | 3.9 | 100.0 |
| Total | 389 | 100.0 | 100.0 | 100.0 |

Table 16. Management of Diabetes by study subjects

| Frequency | Percent | Valid Percent | Cumulative Percent |
|-----------|---------|---------------|--------------------|
| Valid Oral Medicine | 39 | 10.0 | 10.0 | 10.0 |
| Insulin therapy | 350 | 90.0 | 90.0 | 100.0 |
| Total | 389 | 100.0 | 100.0 | 100.0 |
Table 17. Characteristics of study subjects after using proper management of Tuberculosis and Diabetes Mellitus

|                          | Frequency | Percent | Valid Percent | Cumulative Percent |
|--------------------------|-----------|---------|---------------|--------------------|
| Valid Cure and complete therapy | 341       | 87.7    | 87.7          | 87.7               |
| Failed                   | 46        | 11.8    | 11.8          | 99.5               |
| Adequate and Died        | 2         | .5      | .5            | 100.0              |
| Total                    | 389       | 100.0   | 100.0         |                    |

Table 18. Cross tabulation of Gender along with Tuberculosis Smear Test

| Gender | TB Smear test | Total |
|--------|--------------|-------|
|        | Positive     | Negative |   |
| Male   | 153          | 56      | 209 |
| Female | 76           | 104     | 180 |
| Total  | 229          | 160     | 389 |

Each participant had different diabetes history as mentioned in Table 15.

Management of Diabetes among the study subjects were controlled through oral therapy and Insulin and the results for the management of Diabetes were mentioned in Table 16.

Characteristics of study subjects after using proper management of Tuberculosis and Diabetes Mellitus were mentioned in Table 17.

4. CONCLUSION

It was concluded from the current study that there were many chances for developing drug resistance and multi-drug resistance among the patients suffering from co-morbid including tuberculosis along with Diabetes mellitus (DM). Proper counseling and seminar should be conducted, for the patients, in order to reduce the complication of either type of disease. People from remote areas were unaware of complexities of symptoms and their proper management, so they were properly guided for dosing. Oral therapy for DM had developed number of drug-drug interaction, so the patients were facilitated regarding proper insulin therapy.

CONSENT

As per international standard or university standard, respondents' written consent has been collected and preserved by the author(s).

ETHICAL APPROVAL

As per international standard or university standard written ethical approval has been collected and preserved by the author(s).

COMPETING INTERESTS

Authors have declared that no competing interests exist.

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