

**SPUTUM SMEAR POSITIVITY GRADE AND CHEST X-RAY FINDINGS
IN TUBERCULOSIS: A CROSS-SECTIONAL STUDY**

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**ПОЛОЖИТЕЛЬНОСТЬ МАЗКА МОКРОТЫ И РЕНТГЕНОВСКОГО
ОБСЛЕДОВАНИЯ ГРУДНОЙ КЛЕТКИ ПРИ ТУБЕРКУЛЕЗЕ:
ОДНОМОМЕНТНОЕ ИССЛЕДОВАНИЕ**

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Abstract. Despite many advances in the diagnosis, screening, and rapid treatment of tuberculosis, it is still a public health concern in the world. Due to the importance of this issue in diagnosis and reduction of transmission of infection and treatment of the disease especially where this study is conducted due to the high prevalence of tuberculosis, this study was done to determine The relationship between sputum smear positivity grade and chest X-ray findings in pulmonary tuberculosis patients in a hospital in southeast of Iran. This cross-sectional study was performed on all patients with pulmonary TB referencing the health centers in Zabol city, southeast of Iran from 1 January 2015 to 30 December 2020. Sputum smear and radiographic findings of the chest X-ray were evaluated. Data was collected using a form of information and finally analyzed by SPSS 22. Out of 101 patients examined in the present study, 71 were women and 30 were men. The mean age of the patients was 62.68 ± 13.61 years. The frequency of opacity in patients with grades 1, 2, and 3 was 71.4, 78.5, and 76.5%, respectively. Frequency of cavitation in patients with Grade 1, 2 and 3 was 11.5%, 28.5% and 52.9% respectively (P value 0.001). The frequency of reticulonodular presentations in patients with grade 1, 2, and 3 was 24.2, 7.1, and 0%, respectively. In general, the results of this study showed that, with increasing grading of smears (1+, 2+, and 3+), the frequency of cavitation presentation increased significantly and the frequency of reticulonodular presentations decreased significantly. In general, the results of this study showed that, with increasing grading of smears (, the frequency of Cavitation presentation increased significantly and the frequency of reticulonodular presentations decreased significantly. The findings of the present study can help physicians better diagnose TB.

Keywords: Cross-Sectional Studies; Sputum; Iran; X-Rays; Tuberculosis; Radiography

Резюме. Несмотря на многочисленные достижения в диагностике, скрининге и быстром лечении туберкулеза, он по-прежнему вызывает

озабоченность общественного здравоохранения во всем мире. Ввиду важности этого вопроса для диагностики и снижения передачи инфекции и лечения заболевания, особенно там, где проводится настоящее исследование из-за высокой распространенности туберкулеза, мы определяли взаимосвязи между степенью положительного мазка мокроты и рентгенограммой грудной клеткой в виде радиологических находок у больных туберкулезом легких в больнице на юго-востоке Ирана. Это поперечное исследование было проведено на всех пациентах с легочным туберкулезом в медицинских центрах города Забол на юго-востоке Ирана с 1 января 2015 года по 30 декабря 2020 года. Были оценены мазки мокроты и рентгенологические данные грудной клетки, собранные с использованием бланка пациента и проанализированы с помощью программы SPSS 22. Из 101 пациента, обследованного в настоящем исследовании, 71 были женщинами и 30 мужчинами. Средний возраст пациентов составил $62,68 \pm 13,61$ года. Частота помутнения у пациентов 1, 2 и 3 степени составила 71,4, 78,5 и 76,5% соответственно. Частота каверн у пациентов со степенью 1, 2 и 3 составила 11,5%, 28,5% и 52,9% соответственно (значение $P < 0,001$). Частота ретикулонодулярных признаков у пациентов 1, 2 и 3 степени составила 24,2, 7,1 и 0% соответственно. В целом, результаты этого исследования показали, что с увеличением градации мазков (1+, 2+ и 3+) частота появления кавитации значительно увеличивалась, а частота ретикулонодулярных проявлений значительно снижалась. В целом, результаты этого исследования показали, что с увеличением степени градации мазков частота появления кавитации значительно увеличилась, а частота ретикулонодулярных представлений значительно снизилась. Результаты настоящего исследования могут помочь врачам лучше диагностировать ТБ.

Ключевые слова: кросс-секционные исследования; Мокрота; Иран; Рентгеновские лучи; Туберкулез; Рентгенография

1 **Introduction**

2 Despite many advances in the diagnosis, screening, and rapid treatment of
3 tuberculosis, it is still a public health concern in the world. According to the latest
4 WHO report, more than 10 million people worldwide are infected with tuberculosis.
5 Geographically, most TB patients are in Africa and EMRO [1].
6 According to the latest meta-analysis reports, the prevalence of TB in Iran is 23% [2]
7 to 27% [3]. TB is the biggest cause of death among single-agent infectious diseases
8 (even more so than AIDS, malaria, and measles) and has a tenth-highest global
9 burden of disease, and is expected to continue to maintain its present status until
10 2020 [4]. The basis of the diagnosis of pulmonary tuberculosis is a direct and simple
11 screening of susceptible patients. In the best of cases, the sensitivity of the sputum
12 test to detect pulmonary tuberculosis is fifty to sixty percent [5]. By the standard
13 definition, patients who experience at least two positive sputum smear tests, or have
14 only a positive sputum smear test for bacilli acid-fast associated with radiographic
15 changes in the chest X-ray, or a positive smear for acid bacilli in addition to a
16 positive culture are considered as positive for active tuberculosis [6-8]. The grade of
17 the smear is determined by the bacillary load in each microscopic field. Some studies
18 have found that the grade of primary smear can be considered as a predictive factor
19 of patient's morbidity and mortality, which, in the case of a higher grade of
20 positivity, it is more likely to be a failure in treatment and cause death [9,10]. In
21 some studies, the relationship between the grade of primary positive smear and
22 increased clinical manifestations has been stated [11]. Chest X-ray is also a suitable
23 and sensitive diagnostic tool for detecting pulmonary lesions, including in
24 tuberculosis, so that in the case of a normal chest X-ray, the diagnosis of tuberculosis
25 is partially excluded [6,12]. On the other hand, in cases where this disease is actively
26 sought, and when it is diagnosed at an early stage, pulmonary involvement can be a
27 sign of our success in the early detection of these patients, resulting from
28 radiographic findings [9]. Based on the researcher's best knowledge there is no study
29 has been conducted to investigate the relationship between the findings of chest X-

30 ray radiography and the grade of positivity of sputum smear in Iran and especially
31 Southeast of Iran as an area with a high prevalence of tuberculosis. According to the
32 statistics of the Ministry of Health of Iran, Sistan and Baluchestan province and
33 Zabol city are the most common cities for tuberculosis in Iran [13,14].

34 Some studies have been done in this regard, and due to the importance of this issue
35 in diagnosis and reduction of transmission of infection and treatment of the disease
36 especially where this study is conducted due to the high prevalence of tuberculosis,
37 this study was done to determine The relationship between sputum smear positivity
38 grade and chest X-ray findings in pulmonary tuberculosis patients in a hospital in
39 southeast of Iran.

40

41

42 **Methods**

43 This cross-sectional study was performed on all patients with pulmonary TB
44 referencing the health centers in Zabol city, southeast of Iran from 1 January 2015
45 to 30 December 2020.

46 In this study, the national TB diagnosis protocol based on the WHO guidelines was
47 used to diagnose TB in included patients. Patients over 18 years of age were
48 included. Patients without smear grading that had chest radiographs were excluded.
49 A researcher-made checklist for collecting information. The checklist were
50 containing demographic information, smear positivity grading, and chest
51 radiographs. The study protocol approved has in the Ethics Committee of Zabol
52 University of Medical Sciences. Written consent was obtained from all participants
53 prior to the study. Participants were assured that their information would be kept
54 confidential. STROBE checklist was used to report the study.

55 The patient's characteristics were described using descriptive tests including mean,
56 standard deviation, frequency, and percentage. The Kolmogorov-Smirnov test was
57 used to evaluate data normality. SPSS Version 22 for Windows (SPSS Inc., Chicago,

58 IL, USA) was used to analyze the data. The confidence interval of 95% and a
59 significance level of P-value less than 0.05 was considered significant.

60

61 **Results**

62 Of the 101 participants, 71 (70.3%) were male and the rest were women. The mean
63 age of patients was 62.68 years with a standard deviation of 13.61. The youngest
64 and oldest patients were 18 and 86 years old respectively. The Women with a
65 sputum positivity grade of 1, 2, and 3 were 73.3%, 50%, and 70.6%, respectively,
66 and the prevalence of men in Grade 1, 2, and 3 was 25.7%, 50%, and 29.4%. There
67 was no significant difference between the two sexes in terms of smear grade ($p =$
68 0.192). The following table shows that the frequency of consolidation in 3 chest X-
69 rays of patients with smear grade of 1, 2, 3 and was 71.4, 78.5, and 76.5%,
70 respectively. This difference in the size of consolidation in patients with different
71 grades was not statistically significant (Table 1). ($p = 0.833$)

72 The following table shows that the frequency of cavitation in patients with Grade
73 1, 2, and 3 was 11.5%, 28.5%, and 52.9% respectively. This difference in the
74 frequency of cavity was statistically significant in three groups ($p = 0.001$) (table 2)

75 The following table shows that **nodular** presentations in patients with grades 1, 2,
76 and 3 were 18.6, 42.8, and 35.3%, respectively. This difference was not statistically
77 significant in the three groups ($p = 0.086$). (table 3)

78

79 The following table shows that the prevalence of reticulonodular involvement in
80 patients with Grade 1, 2, and 3 was 24.2%, 7.1%, and 0.0%, respectively. The
81 difference between the frequency of reticulonodular involvement in the three groups
82 was statistically significant. ($p = 0.022$). (Table 4)

83

84 **Discussion**

85 In this study, 101 patients with tuberculosis were examined. Among them, 71
86 (70.3%) were women and the rest were men. In the studied patients, 70 cases had

87 Grade 1 (74.3% female and 25.7% male), 14 had Grade 2 (male = female) and 17
88 Grade 3 (70.6% female and 35.7% male). There was no significant difference
89 between the sexes in terms of smear grade. The findings of this study cannot be
90 compared to any other studies because of the lack of similar research on the
91 relationship between sex and grading of the smear. The mean age of patients was
92 62.68 years with a standard deviation of 13.61. The youngest and oldest patients
93 were 18 and 86 years old. The mean age of patients with grades 1, 2, and 3 was
94 64.47, 62.07, and 55.82 years, respectively. The age difference of patients in
95 different grades was not statistically significant. There was also no relationship
96 between age and grading of the smear. The frequency of consolidation in patients
97 with grades 1, 2, and 3 was 71.4, 78.5, and 76.5%, respectively. The difference in
98 the degree of opacity in patients with different grades was not statistically
99 significant. Although Grade 2 patients were more frequent than grade 1 and grade 3
100 patients, the difference between the three groups was not significant. There does not
101 seem to be a relationship between the degree of smear and the consolidation in the
102 graph. This finding is not consistent with other studies. In the study of BJ Parcell
103 (2017), with increasing the degree of smear (+1, +2, +3 or +4), the frequency of
104 consolidation increased significantly(in degrees +1, +2, +3, the frequency of
105 consolidation was +4 81%, 95%, 100% respectively) [15]. In the study of KP
106 Brahmapurkar, (2017), with increasing the grade of smear positivity, the number of
107 cases also increased significantly [16] which can be the reason for this inconsistency.
108 F Bisognin et al. (2019) also showed that the frequency of opacity increased with
109 increasing the number of acid bacilli [17]. The difference between the findings of
110 the present and other studies could be attributed to the fact that the present study
111 focused on investigating chest radiographs based on smear grading, while other
112 studies examined the relationship between CT scan and HRCT with smear grading.
113 The frequency of cavitation in patients with grades 1, 2, and 3 was 11.5%, 28.5%,
114 and 52.9% respectively. This difference in the frequency of cavity was statistically
115 significant in the three groups. Different types of patients had different cavitation

116 levels; in patients with grade 3 cavitation, there was a significant increase in grade
117 3 and grade 1 patients. Therefore, there seems to be a relationship between the degree
118 of smear and the presence of the cavity. In the M Saffari study, with the increase in
119 the degree of smear (+1, +2, +3, or +4), the frequency of CT scan findings including
120 cavitation also increased significantly, so that the frequency of cavitation cases in
121 degrees +1, +2, +3, and +4 was 33%, 68%, 94% and 100% respectively [18]. In the
122 study of A Penn-Nicholson (2019), with increasing the degree of smear, cavitation
123 also increased significantly [19]. Matsuka (2004) also showed that the frequency of
124 covariation increased with increasing the number of acid bacilli [20]. In the study of
125 M Hassanzad et al., cavitation had a significant correlation with smear gradation
126 [21]. This study showed that nodular facial abnormalities in patients with Grade 1,
127 2, and 3 were 18.6, 42.8, and 35.3%, respectively. Nodular features were not
128 significantly different in the three groups. Although grade 2 patients had more
129 nodular features in comparison with grade 1 and grade 3 patients, the difference
130 between the three groups was not significant; therefore, there is no significant
131 relationship between the degree of smear and nodular feature abnormalities.
132 Matsuka (2004) also showed that the incidence of nodular presentation increased
133 with increasing degree of smear, but their differences were not statistically
134 significant [6].

135 The incidence of reticulonodular involvement in patients with grades 1, 2, and 3 was
136 24.2%, 7.1%, and 0%. This difference in the frequency of reticulonodular
137 involvement in the three groups was statistically significant. On the other hand,
138 patients with reticulonodular involvement were significantly more likely to have a
139 grade 1 smear. The lowest frequency of reticulonodular appearance belonged to
140 grade 3. These results showed that there is a significant relationship between the
141 degree of smear and reticulonodular involvement in a way that an increase in the
142 grade of smear (1+, 2+, 3+) decreases the frequency of reticulonodular appearance.
143 The findings of this study cannot be compared to any other studies because of the
144 lack of similar research on the relationship between sex and grading of the smear.

145 The most important limitations of the present study were: This is a cross-sectional
146 study. When interpreting the results, the specific limitations of this type of study
147 should be considered. The most important strength of this study was that this is the
148 first report in this long period of this region as the most common area of the
149 tuberculosis outbreak.

150

151 **Conclusion**

152 In general, the results of this study showed that, with increasing grading of smears
153 (, the frequency of Cavitation presentation increased significantly and the frequency
154 of reticulonodular presentations decreased significantly. The findings of the present
155 study can help physicians better diagnose TB.

156

157 **Conflict of interest**

158 All authors declare that they have no conflict of interest.

159

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163

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167

168 **Author contributions**

169 RB and EN designed study. EN collected, analyzed, interpreted data and wrote the
170 manuscript. RB analyzed data, reviewed and revised the manuscript. All authors
171 approved the final version of manuscript.

TABLES

Table 1: Frequency of consolidation in chest X-ray in association with the grading of sputum smear

| Grade of smear positivity | | Grade 1 | Grade 2 | Grade 3 | P-value |
|----------------------------------|-----|----------------|----------------|----------------|----------------|
| Consolidation | Yes | 50 71.4% | 11 78.5% | 13 76.5% | 0.833 |
| | No | 20 28.5% | 3 21.4% | 4 23.5% | |

Table 2: Frequency of cavitation by the degree of smear

| Grade of smear positivity | | Grade1 | Grade2 | Grade3 | P-value |
|---------------------------|-----|-------------|-------------|-------------|---------|
| Cavitation | Yes | 8 11.5% | 4 28.5% | 9 52.9% | 0.001 |
| | No | 62 88.5% | 10 71.4% | 8 47.05% | |

Table 3: Frequency of nodular presentation by the grade of sputum smear positivity

| Grade of smear positivity | | Grade1 | Grade2 | Grade3 | P-value |
|-----------------------------|-----|-------------|------------|-------------|---------|
| Nodular presentation | Yes | 13 18.6% | 6 42.8% | 6 35.3% | 0.086 |
| | No | 57 81.4% | 8 57.1% | 11 64.7% | |

Table 4: Frequency of reticulonodular presentations by the grade of smear positivity

| | | Grade1 | Grade2 | Grade3 | P-value |
|-----------------|-----|-------------|-------------|------------|---------|
| reticulonodular | Yes | 17 24.2% | 1 7.1% | 0 0% | 0.022 |
| | No | 53 75.7% | 13 92.8% | 17 100% | |

TITLE PAGE_METADATA

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