High-resolution computed tomography in silicosis: correlation with chest radiography and pulmonary function tests*

Tomografia computadorizada de alta resolução na silicose: correlação com radiografia e testes de função pulmonar

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Abstract

Objective: To correlate tomographic findings with pulmonary function findings, as well as to compare chest X-ray findings with high-resolution computed tomography (HRCT) findings, in patients with silicosis. Methods: A cross-sectional study was conducted in 44 non-smoking patients without a history of tuberculosis. Chest X-ray findings were classified according to the International Labour Organization recommendations. Using a semiquantitative system, the following HRCT findings were measured: the full extent of pulmonary involvement; parenchymal opacities; and emphysema. Spirometry and forced oscillation were performed. Pulmonary volumes were evaluated using the helium dilution method, and diffusing capacity of the lung for carbon monoxide was assessed. Results: Of the 44 patients studied, 41 were male. The mean age was 48.4 years. There were 4 patients who were classified as category 0 based on X-ray findings and as category 1 based on HRCT findings. Using HRCT scans, we identified progressive massive fibrosis in 33 patients, compared with only 23 patients when X-rays were used. Opacity score was found to correlate most closely with airflow, DLCO and compliance. Emphysema score correlated inversely with volume, DLCO and airflow. In this sample of patients presenting a predominance of large opacities (75% of the individuals), the deterioration of pulmonary function was associated with the extent of structural changes. Conclusions: In the early detection of silicosis and the identification of progressive massive fibrosis, HRCT scans are superior to X-rays.

Keywords: Silicosis; Occupational diseases; Tomography, x-ray computed; Respiratory function tests.

Resumo

Objetivo: Correlacionar os parâmetros tomográficos com os de função pulmonar em portadores de silicose, bem como comparar os resultados da tomografia computadorizada de alta resolução (TCAR) com os da radiografia de tórax. Métodos: Foi realizado um estudo de corte transversal, em que foram avaliados 44 pacientes não-tuberculícos e sem história pregressa de tuberculose. As radiografias foram classificadas de acordo com a proposta da Organização Internacional do Trabalho. Utilizando um sistema de escore semiquantitativo, os seguintes achados na TCAR foram quantificados: extensão total do envolvimento pulmonar, opacidades parenquimatosas e enfisema. Foram realizados espirometria, oscilações forçadas, avaliação dos volumes pulmonares pela técnica de diluição com hélio e medida da diffusing capacity of the lung for carbon monoxide (DLCO, capacidade de difusão do monóxido de carbono). Resultados: Dos 44 pacientes estudados, 41 eram homens, com média de idade de 48,4 anos. Na análise comparativa, 4 pacientes classificados na categoria 0 pela radiografia foram reclassificados na categoria 1 pela TCAR e, enquanto a radiografia diagnosticou 23 casos de fibrose maciça progressiva, a TCAR estabeleceu esse diagnóstico em 33 doentes. Para o escore de opacidades, as maiores correlações foram observadas com as medidas de fluxo, DLCO e complacência. Já o enfisema correlacionou-se negativamente com as medidas de volume, DLCO e fluxo. Nesta amostra com predomínio de grandes opacidades (75% dos indivíduos), a deterioração da função pulmonar associou-se com a extensão das alterações estruturais. Conclusões: Na silicose, a TCAR é superior à radiografia tanto na detecção precoce da doença quanto na identificação de fibrose maciça progressiva.

Descritores: Silicose; Doenças ocupacionais; Tomografia computadorizada por raios X; Testes de função respiratória.

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Introduction

According to the current guidelines of the International Labour Organization (ILO),(1) the principal method of diagnosing silicosis is the analysis of chest X-rays and of the history of occupational exposure to free silica. However, high-resolution computed tomography (HRCT) has increasingly taken a prominent position in the evaluation of silicosis, offering relevant additional information for the early detection of small opacities and emphysema and identification of complications.(2-4) Pulmonary function tests, although not used as diagnostic tools, are widely employed in longitudinally studies of individuals with silicosis.(3,4)

Since imaging and function tests are the most widely used diagnostic resources in the follow-up evaluation of individuals with silicosis, it is fundamental to establish the correlation between these two methods. Due to its higher sensitivity, HRCT has been the imaging test most often used for this purpose. However, in studies correlating imaging tests with respiratory function tests, quantification of the severity of the disease by function tests is affected by history of smoking and associated pulmonary emphysema, a fact highlighted by most investigators.(5-10)

The objective of the present study was to correlate tomographic parameters with respiratory function parameters in nonsmokers with silicosis and to compare the HRCT findings with the chest X-ray findings.

Methods

A descriptive study evaluating 53 nonsmokers with history of exposure to silica was carried out. Participants were informed regarding the objective of the study, after which they all gave written informed consent. The protocol was approved by the ethics in research committee of the institution.

The inclusion criteria were having an ≥10-year history of exposure to silica and presenting X-ray or HRCT images consistent with silicosis. Smokers and former smokers were excluded, as were individuals in whom the radiological findings indicated pulmonary tuberculosis. Based on these criteria, we excluded 9 patients, 4 of whom presented tomographic findings consistent with pulmonary tuberculosis.

All tests were performed within one month of each and in the following order: pulmonary function test, chest X-ray and HRCT.

The chest X-ray was performed in the antero-posterior position using a Siemens X-ray unit (model LX30; Siemens AG, Erlangen, Germany). The following technique was used: distance of 180 cm between focal plane and the film; device at 80 kVp; exposure time of 0.04 ms. Readings were performed independently by three specialists certified by the Jorge Duprat Figueiredo Foundation for Occupational Safety and Medicine and by the Brazilian Ministry of Labor and Employment. For the sake of simplification, the classification of the profusion of small opacities was summarized: 0/−, 0/0 or 0/1 = 0; 1/0, 1/1 or 1/2 = 1/2/1; 2/2 or 2/3 = 2; 3/2; and 3/3 or 3/+ = 3. Large opacities were classified as types A, B or C, according to the standards provided by the ILO.(1) The results were summarized as the medians of the readings.

The pulmonary function tests consisted of the forced oscillation technique (FOT), spirometry, evaluation of lung volumes using the helium dilution method and assessment of diffusing capacity of the lung for carbon monoxide (DLCO). The FOT was performed using an impedance analyzer, and the following parameters were evaluated: total respiratory resistance; airway resistance; and dynamic respiratory compliance (Crs,dyn). The remaining tests were performed using the Collins Plus Pulmonary Function Testing System (Warren E. Collins, Inc., Braintree, MA, USA), standardized and interpreted in accordance with the guidelines established by the Brazilian Thoracic Society.(11) Equations devised by Pereira (for spirometry) and Neder (for static lung volumes and DLCO) were used in the interpretation of the following parameters(12-14):

- spirometry: forced vital capacity (FVC); forced expiratory volume in one second (FEV1); FEV1/FVC ratio; forced expiratory flow between 25 and 75% of FVC (FEF25-75%); and FEF25-75%/FVC ratio
- helium dilution method: residual volume; total lung capacity; and the residual volume/total lung capacity ratio
- determination of single-breath DLCO

The HRCT was performed using a GE HiSpeed scanner (General Electric Medical Systems, Milwaukee, WI, USA). The following technique was used: 1-mm
slices, at 1.5-s intervals and increased by 10 mm; image reconstruction with a 512 x 512 pixel matrix, using a high-resolution algorithm; 1000 Hounsfield unit (HU)-width window; −700 HU medium window level. The tomographic findings were interpreted by consensus among four radiologists who had been previously trained for one month in how to apply the classification system adopted. Small opacities were classified into four categories, according to profusion, and in a manner similar to that used to classify the chest X-ray findings:

0) absence of micronodules
1) a small number of micronodules without vascular blurring
2) a large number of micronodules, with or without vascular blurring but with no confluence
3) confluence of nodules <10 mm, typically accompanied by pronounced vascular blurring

Large opacities were classified as type A or B (respectively, one or more opacities > 10 mm having the sum of the diameters ≤50 mm, and one or more opacities having the sum of the diameters <50 mm). Subsequently, HRCT was evaluated as to the extent of pulmonary involvement, considering five cut-off levels: origin of major vessels; aortic arch; carina; confluence of pulmonary veins; and 1 cm above the right diaphragm. Using a semiquantitative evaluation system, each of these levels (right and left, separately, totaling 10 levels) was analyzed as to the following aspects:

- score of total interstitial disease (TID, including nodules, masses, emphysema and other parenchymal alterations):
  0) no alteration
  1) pulmonary involvement ≤5% of the area
  2) pulmonary involvement from >5 to ≤25% of the area
  3) pulmonary involvement from >25 to 49% of the area
  4) pulmonary involvement from 50 to 75% of the area
  5) pulmonary involvement >75% of the area

- score of total extent of parenchymal opacities, including nodules and masses:
  0) no parenchymal opacities
  1) parenchymal opacities involving ≤5% of the area
  2) parenchymal opacities involving >5 to ≤25% of the area
  3) parenchymal opacities involving >25 to 49% of the area

Figure 1 - Example of extent of disease scored using high-resolution computerized tomography in a 42-year-old man with silicosis. At cut-off levels 1, 4 and 5, the test was normal. At level 2 (aortic arch, A), a score of 2 (opacities involving >5 to ≤25% of the area) was assigned to both lungs (total = 4). At level 3 (carina, B), a score of 1 (opacities involving ≤5% of the area) was assigned to the right lung and a score of 2 was assigned to the left lung (total = 3). After applying correction factors for each cut-off level, the scores obtained were 0.76 (4 x 0.190) for level 2 and 0.67 (3 x 0.222) for level 3. Therefore, summing the scores, we obtained a total score of 1.43 for the extent of parenchymal opacities.
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...functional indices and tomographic scores were studied using Pearson’s parametric test. Values of $p \leq 0.05$ were considered significant.

Results

Of the 44 patients studied, 41 were male. The mean age was 48.4 years. The mean duration of silica exposure and the mean time since withdrawal were 16.1 and 16.9 years, respectively. Sandblasting and stone cutting were the professional activities most often cited (45.5% and 34.1%, respectively). Other professional activities cited were marble quar-...
in 13 cases and moderate in 1); 13 had mixed respiratory disorder (mild in 8 cases, moderate in 3 and severe in 2); 11 had restrictive defect (mild in 8 cases, moderate in 2 and severe in 1); and 6 presented spirometric indices and static volumes that were within the normal ranges. In this study, 29 individuals presented a DLCO reduction, which was characterized as mild in 15 cases, moderate in 12 and severe in 2.

In relation to the HRCT scores, the mean values of TID, parenchymal opacities and emphysema were 3.7, 3.3 and 1.5, respectively. The coefficients of the correlations between functional parameters and values obtained in the HRCT analysis are given in Table 4. The opacities score correlated most strongly with airflow, DLCO and Crs,dyn. However, emphysema correlated inversely with lung volume, DLCO and airflow.

**Discussion**

In any study of morphofunctional correlations, it is fundamental that the investigators eliminate the influence of any factors other than the one analyzed. Most previously studies have included smokers and former smokers, as well as patients with a history of tuberculosis. However, in the present study, we were careful to exclude smokers, former smokers and individuals with a history of tuberculosis. Consequently, it was necessary to recruit patients from various referral centers for silicosis in order to reach the ideal sample size.

Conventionally, silicosis-related lesions are assessed radiologically according to the ILO classification of pneumoconiosis, which was created for epidemiological purposes. However, its usefulness in clinical diagnosis is debatable, particularly in relation to two aspects: the high inter-rater variability in chest X-ray analysis, especially to low profusion categories; and the fact that it can underestimate the presence of pulmonary disease. In the present study, greater discrepancy among specialists was observed for the small opacity category. Inter-rater variability was lower for large opacities than for small opacities. This finding is in accordance with those of other studies, which also showed higher kappa values for large opacities. Despite these limitations, chest X-ray is still the most efficient tool for the follow-up evaluation of workers exposed...
to silica, since it is an inexpensive procedure and exposes subjects to only a low dose of radiation.\(^{[3]}\)

Many studies have been carried out to establish the importance of HRCT in the early detection of small opacities. One study showed that 13 out of 32 individuals with a history of exposure to silica and normal chest X-ray results presented evidence of silicosis on HRCT scans.\(^{[16]}\) In a study carried out in Brazil and including 68 former miners, concordance of chest X-ray findings and HRCT findings was observed in 55 cases. Divergence was observed in 13 cases; HRCT diagnosed 5 new cases of silicosis and ruled out 8 cases.\(^{[17]}\) In the present study, 4 patients classified as category 0 based on the chest X-ray findings were reclassified as category 1 based on the HRCT findings, underscoring the value of the latter exam.

In terms of the capacity to detect large opacities, the superiority of HRCT over chest X-ray is well established. In a study assessing the HRCT results of 76 patients, 10 cases of PMF were observed among the 26 classified as simple silicosis based on chest X-rays. In the present study, 10 new cases of PMF were diagnosed based on the HRCT findings, confirming the higher sensitivity of the method for detecting accelerated silicosis. Another interesting finding was the detection of emphysema, which was diagnosed in 27 patients (61.4%). Considering the fact that the sample consisted exclusively of nonsmokers, the detection of emphysema, in this study, can only be attrib-

### Table 3 - Functional indices according to the tomographic categories of small and large opacities.\(^{a}\)

<table>
<thead>
<tr>
<th>Functional index</th>
<th>Total sample n = 44</th>
<th>Small opacities</th>
<th>Large opacities</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>1 (n = 17)</td>
<td>2 (n = 21)</td>
<td>3 (n = 6)</td>
<td></td>
</tr>
<tr>
<td>FVC (%T)</td>
<td>82.9 ± 21.6</td>
<td>81.9 ± 20.8</td>
<td>88.5 ± 21.1</td>
<td>0.796</td>
</tr>
<tr>
<td>FEV(_1) (%T)</td>
<td>70.4 ± 21.6</td>
<td>69.1 ± 22.0</td>
<td>78.5 ± 15.6</td>
<td>0.623</td>
</tr>
<tr>
<td>FEV(_1)/FVC (%T)</td>
<td>84.9 ± 14.2</td>
<td>83.8 ± 16.6</td>
<td>90.7 ± 8.5</td>
<td>0.576</td>
</tr>
<tr>
<td>FEF(_{25-75})% (%T)</td>
<td>52.3 ± 26.2</td>
<td>49.2 ± 26.4</td>
<td>58.8 ± 15.9</td>
<td>0.744</td>
</tr>
<tr>
<td>FEF(_{25-75})/FVC (%T)</td>
<td>63.1 ± 29.3</td>
<td>61.0 ± 31.6</td>
<td>69.3 ± 30.8</td>
<td>0.842</td>
</tr>
<tr>
<td>R0 (cmH(_2)O/L/s)</td>
<td>3.3 ± 1.4</td>
<td>3.3 ± 1.3</td>
<td>3.5 ± 0.8</td>
<td>0.557</td>
</tr>
<tr>
<td>mR (cmH(_2)O/L/s)</td>
<td>3.0 ± 1.1</td>
<td>3.0 ± 0.8</td>
<td>3.1 ± 0.6</td>
<td>0.537</td>
</tr>
<tr>
<td>Crs,dyn (L/cm/ H(_2)O)</td>
<td>0.014 ± 0.006</td>
<td>0.014 ± 0.007</td>
<td>0.014 ± 0.005</td>
<td>0.016 ± 0.003</td>
</tr>
<tr>
<td>TLC (%T)</td>
<td>78.4 ± 16.4</td>
<td>76.4 ± 18.5</td>
<td>79.9 ± 16.0</td>
<td>0.815</td>
</tr>
<tr>
<td>RV (%T)</td>
<td>93.2 ± 32.9</td>
<td>91.8 ± 28.3</td>
<td>95.2 ± 40.2</td>
<td>0.927</td>
</tr>
<tr>
<td>RV/TLC (%T)</td>
<td>117.0 ± 29.3</td>
<td>116.9 ± 23.2</td>
<td>117.3 ± 35.5</td>
<td>116.5 ± 24.9</td>
</tr>
<tr>
<td>DLCO (%T)</td>
<td>67.2 ± 17.2</td>
<td>67.8 ± 18.7</td>
<td>64.4 ± 17.0</td>
<td>0.418</td>
</tr>
</tbody>
</table>

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FVC: forced vital capacity; %T: percentage of theoretical value; FEV\(_1\): forced expiratory volume in one second; FEF\(_{25-75}\)/FVC: forced expiratory flow between 25 and 75% of FVC; FEF\(_{25-75}\)/FVC ratio; R0: resistance at the intercept; mR: mean resistance; Crs,dyn: dynamic respiratory compliance; TLC: total lung capacity; RV: residual volume; and DLCO: diffusing capacity of the lung for carbon monoxide. *Values expressed as mean ± standard deviation. **Significant difference.
related emphysema be the principal mechanisms responsible for the obstructive phenomenon, since 75% of the patients presented PMF in the HRCT results. These masses of PMF, composed of clusters of fibrotic nodules, can distort the lung parenchyma and cause an irregular increase in the area of the adjacent air spaces, ultimately limiting airflow.\(^{10,22,23,24}\)

Comparing the means of the pulmonary function parameters between the different HRCT categories, we observed that, for small opacities, there was no significant difference in any isolated functional parameter for any HRCT category evaluated. This finding is in accordance with those of a previous study in which the micronodules detected on the HRCT scan, \textit{per se}, were found to have no influence on functional deterioration.\(^{15}\) However, in the present study, we observed that, in patients presenting large opacities, there was a progressive decrease in DLCO, Crs,dyn and airflow decreased in parallel with increases in the extent of damage (HRCT classification). These findings are in accordance with those of other studies also showing that functional damage correlates more strongly with the extent of large opacities than with that of small opacities.\(^{5,9-10,23}\) Therefore, an HRCT finding of large opacities can be an important indicator of the severity of silicosis.\(^{8,9,10,25}\)

In our study, as well as in others, parenchymal opacity correlated negatively with a reduction in DLCO.\(^{13,5,10,23}\) Some investigators have attributed this correlation to the presence of PMF, which tends to diminish as the disease advances. That results in the appearance of scar-related emphysema and a decrease in the surface area available for gas exchange, which reduces DLCO.

The analysis of the association obtained between parenchymal opacity and Crs,dyn is interesting, since, to our knowledge, there have been no studies correlating compliance values with HRCT findings. In practice, Crs,dyn represents how easily the respiratory system, including the lungs and chest wall, reaches total lung capacity.\(^{26}\) In accelerated silicosis, the intensification of pleuropulmonary involvement causes a reduction in Crs,dyn. Therefore, since the method of measurement is noninvasive, Crs,dyn might eventually become another functional parameter evaluated in the follow-up treatment of patients with silicosis. In addition, our results indicate that, in nonsmoking silicosis patients presenting large opacities (as in 75% of our sample), deterioration

<table>
<thead>
<tr>
<th>Functional index</th>
<th>Total score</th>
<th>pacO score</th>
<th>Emphysema score</th>
</tr>
</thead>
<tbody>
<tr>
<td>FVC (%T)</td>
<td>-0.42**</td>
<td>-0.33*</td>
<td>-0.44**</td>
</tr>
<tr>
<td>FEV(_1) (%T)</td>
<td>-0.61**</td>
<td>-0.55**</td>
<td>-0.44**</td>
</tr>
<tr>
<td>FEV(_1)/FVC (%T)</td>
<td>-0.46**</td>
<td>-0.51**</td>
<td>-0.10</td>
</tr>
<tr>
<td>FEF(_25-75) (%T)</td>
<td>-0.59**</td>
<td>-0.58**</td>
<td>-0.35*</td>
</tr>
<tr>
<td>FEF(_25-75)/FVC (%T)</td>
<td>-0.41**</td>
<td>-0.46**</td>
<td>-0.12</td>
</tr>
<tr>
<td>R0 (cmH(_2)O/L/s)</td>
<td>+0.20</td>
<td>+0.19</td>
<td>+0.10</td>
</tr>
<tr>
<td>Rm (cmH(_2)O/L/s)</td>
<td>+0.13</td>
<td>+0.11</td>
<td>+0.13</td>
</tr>
<tr>
<td>Crs,dyn (L/cm/H(_2)O)</td>
<td>-0.48**</td>
<td>-0.45**</td>
<td>-0.25</td>
</tr>
<tr>
<td>TLC (%T)</td>
<td>-0.42**</td>
<td>-0.32*</td>
<td>-0.51**</td>
</tr>
<tr>
<td>RV (%T)</td>
<td>-0.06</td>
<td>-0.01</td>
<td>-0.30*</td>
</tr>
<tr>
<td>RV/TLC (%T)</td>
<td>+0.31*</td>
<td>+0.28</td>
<td>+0.06</td>
</tr>
<tr>
<td>DLCO (%T)</td>
<td>-0.63**</td>
<td>-0.57**</td>
<td>-0.43**</td>
</tr>
</tbody>
</table>

pacO: parenchymal opacities; FVC: forced vital capacity; %T: percentage of theoretical value; FEV\(_1\): forced expiratory volume in one second; FEF\(_25-75\): forced expiratory flow between 25 and 75% of FVC; R0: resistance at the intercept; mR: mean resistance (mR); Crs,dyn: dynamic respiratory compliance; TLC: total lung capacity; RV: residual volume; e DLCO: diffusing capacity of the lung for carbon monoxide. *p < 0.05. **p < 0.01.
of pulmonary function correlates with the extent of the structural alterations.

As has been reported in previous studies, we found that emphysema presented significant correlations with airflow, lung volume and DLCO. However, the correlation coefficients were slightly inferior to those obtained in studies that included smokers in their samples. Therefore, again, the difference can be attributed to smoking, which is known to reduce airflow and DLCO.

A critical analysis of the results of the present study and its limitations is called for. One potential limitation of this study was that the dose and incidence of the occupational exposure were not quantified. Such data might have furthered the understanding of morphofunctional correlations. Another limitation is that the coefficient of concordance was not evaluated for the HRCT readings. Therefore, the interpretation of the tomographic findings by consensus implies possible acceptance of the analysis of the most experienced specialists by the less experienced specialists, which constitutes a methodological bias.

In conclusion, our study shows that, in the diagnosis of silicosis HRCT scans are superior to chest X-rays, both for the early detection of the initial phases of the disease and for the identification of PMF.

References

