Pyrophosphate heart scan in patients with progressive systemic sclerosis

FRANTIŠEK DUŠKA, PETR BRADNA, JAN NOVÁK, JIŘÍ KUBÍČEK, JAROSLAV VIŽĎA, PETR KAFKA, YVONA MAZUROVÁ, VLADISLAV BLAHA

From the Department of Normal and Pathological Physiology, Division of Nuclear Medicine, Department of Radiology, and Chair of the 2nd Department of Medicine, Charles University Faculty of Medicine, Hradec Králové, Czechoslovakia

SUMMARY Scintigraphic examination of the myocardium, using \(^{99m}\)Tc-labelled pyrophosphate, was carried out in 17 patients suffering from systemic sclerosis. This connective tissue disorder very often affects the myocardium secondarily. The results of the cardiac scan were compared with the information obtained from the electrocardiogram of systolic time intervals. In addition, spirometry was undertaken to detect a potential relation between cardiac and pulmonary involvement. The scan was found to be positive in seven patients and electrocardiographic findings were pathological in five patients only. The systolic time intervals were abnormal in three patients only. A ventilation disturbance was recorded in 10 patients. No clear relation was found between the results of the individual examinations. It is concluded that pyrophosphate heart scintigraphy may detect myocardial impairment in some cases of systemic sclerosis before it is manifested by heart failure. Examination of systolic time intervals is of little importance.

Scintigraphic myocardial imaging using \(^{99m}\)Tc-labelled pyrophosphate has recently become a standard procedure in some hospitals. This diagnostic method has proved useful especially in the detection of acute myocardial infarction.\(^1\) The accumulation of the radiopharmaceutical in heart tissue, however, is not specific for fresh necrosis. Positive scintigraphic findings have been reported in a number of other heart diseases.\(^2\)\(^-\)\(^5\) Hitherto, there has been very little experience with pyrophosphate heart scanning in cardiomyopathies. Only a few papers concerning this have been published to date, with conflicting results.\(^6\)\(^-\)\(^8\) The introduction of new diagnostic methods in cardiomyopathies is important, as the recognition of these using present techniques is often difficult. Our group, therefore, has studied pyrophosphate cardiac scanning in various cardiomyopathies. The results obtained from 17 patients suffering from progressive systemic sclerosis are presented. It is well known that diffuse scarring of the myocardium is a late finding in patients with this disease. This change leads to low cardiac output and death resulting from sudden arrhythmias or congestive heart failure.\(^9\) The results of the pyrophosphate heart scan were compared with the findings obtained from a study of systolic time intervals. As progressive pulmonary fibrosis, with obliteration of the pulmonary capillary bed, may occur, and often leads to pulmonary hypertension and cor pulmonale, ventilation disturbances were also sought.

Subjects and methods

Our series consisted of 17 patients in whom progressive systemic sclerosis was diagnosed using histological examination of a sample of affected skin. There were 13 female and four male patients aged from 20 to 77 years, average 52 years. All patients underwent physical examination; temperature was measured and urine and blood were routinely tested.

Scintigraphic examination was carried out using a Pho/Gamma HP scintillation camera (Nuclear Chicago) with a high resolution collimator. During scintigraphic observation anteroposterior, left anterior oblique, and left lateral projections were used. Two hours before the start of the scintigraphic observation, \(^{99m}\)Tc-labelled pyrophosphate (370 MBq/10 mCi) was given intravenously. An analogue image was obtained by recording 200 000 to 300 000 counts on a film of Polaroid type. The digital image was obtained by recording the data on a CDS 4096

Accepted for publication 6 April 1981
channel analyser. The intensity of the $^{99m}$Tc-phosphate uptake was graded from 0 to 4+ according to the method of Parkey et al. 10

Heart rate, the total duration of systole, pre-ejection as well as ejection phases, mechanical systole, and QT interval were obtained. The quotient was calculated using the formula: pre-ejection phase divided by ejection phase. The systolic time intervals were obtained from a simultaneous electrocardiogram (lead II), phonocardiogram, and carotid pulse trace on a direct writing EK 21 Hellige multiscriptor. The paper speed was 50mm/s. Systole was measured from the Q-wave on the electrocardiogram to the second sound on the phonocardiogram. The ejection phase was determined as the distance from the upstroke of the carotid pulse to the lowest point in its incisura. The pre-ejection phase was calculated as the difference between total systole and ejection phase. Patients were studied, fasting, always at the same time in the morning. After one hour’s rest in a lying position the measurements were taken and a mean of 10 cardiac contractions calculated. To eliminate the influence of rate the values obtained were correlated using Weisser’s indices. The results of the systolic time interval examination were summarised and labelled as “normal” or “pathological”.

All patients were examined spirometrically. The vital capacity, timed expiration curve, and percentage vital capacity in the first second were measured. We also determined whether the ventilation of the lungs was normal or whether an obstructive, restrictive, or mixed ventilation disturbance was involved.

Results

The results are summarised in Table 1. It can be seen that the most frequent positive finding in the examination of the heart was that found on the myocardial scan. A pathological scan was recorded in seven patients; in 10 the scintigraphic image was normal.

![Fig. 1](image1.png)

**Fig. 1** Pyrophosphate myocardial scan of case 7 with systemic scleroderma. Focal pathological activity 2+. Left lateral projection.

![Fig. 2](image2.png)

**Fig. 2** Pyrophosphate myocardial scan of case 4 with systemic scleroderma. Diffuse pathological activity 3+. Left lateral projection.

<table>
<thead>
<tr>
<th>Case No.</th>
<th>Sex</th>
<th>Age</th>
<th>Scan</th>
<th>Electrocardiogram</th>
<th>STI</th>
<th>Abnormality of Ventilation</th>
</tr>
</thead>
<tbody>
<tr>
<td>1</td>
<td>F</td>
<td>63</td>
<td>Normal</td>
<td>Ventricular extrasystoles</td>
<td>Normal</td>
<td>Mixed</td>
</tr>
<tr>
<td>2</td>
<td>F</td>
<td>71</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
</tr>
<tr>
<td>3</td>
<td>M</td>
<td>68</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
</tr>
<tr>
<td>4</td>
<td>M</td>
<td>64</td>
<td>3+ diffuse</td>
<td>Normal</td>
<td>Normal</td>
<td>Restrictive</td>
</tr>
<tr>
<td>5</td>
<td>F</td>
<td>57</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
</tr>
<tr>
<td>6</td>
<td>F</td>
<td>46</td>
<td>Normal</td>
<td>Incomplete right bundle-branch block</td>
<td>Normal</td>
<td>Normal</td>
</tr>
<tr>
<td>7</td>
<td>F</td>
<td>72</td>
<td>2+ focal</td>
<td>Normal</td>
<td>Normal</td>
<td>Mixed</td>
</tr>
<tr>
<td>8</td>
<td>F</td>
<td>54</td>
<td>Normal</td>
<td>Atrial extrasystoles</td>
<td>Normal</td>
<td>Mixed</td>
</tr>
<tr>
<td>9</td>
<td>F</td>
<td>49</td>
<td>2+ diffuse</td>
<td>Ventricular extrasystole inversion T in V,-V,</td>
<td>Pathological</td>
<td>Mixed</td>
</tr>
<tr>
<td>10</td>
<td>M</td>
<td>25</td>
<td>Normal</td>
<td>Normal</td>
<td>Pathological</td>
<td>Mixed</td>
</tr>
<tr>
<td>11</td>
<td>F</td>
<td>38</td>
<td>2+ focal</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
</tr>
<tr>
<td>12</td>
<td>F</td>
<td>39</td>
<td>2+ focal</td>
<td>Normal</td>
<td>Normal</td>
<td>Mixed</td>
</tr>
<tr>
<td>13</td>
<td>F</td>
<td>56</td>
<td>Normal</td>
<td>Normal</td>
<td>Pathological</td>
<td>Mixed</td>
</tr>
<tr>
<td>14</td>
<td>M</td>
<td>20</td>
<td>Normal</td>
<td>Complete right bundle-branch block</td>
<td>Normal</td>
<td>Restrictive</td>
</tr>
<tr>
<td>15</td>
<td>F</td>
<td>77</td>
<td>3+ focal</td>
<td>Normal</td>
<td>Normal</td>
<td>Mixed</td>
</tr>
<tr>
<td>16</td>
<td>F</td>
<td>50</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
</tr>
<tr>
<td>17</td>
<td>F</td>
<td>33</td>
<td>2+ focal</td>
<td>Normal</td>
<td>Normal</td>
<td>Normal</td>
</tr>
</tbody>
</table>

STI, systolic time intervals.
The focus of increased activity was 3+ (diffuse) in case 4 and 2+ (circumscribed) in cases 7, 9, 11, and 12 (Table 1). Examples of positive scintigraphic findings can be seen in Fig. 1 and 2. The electrocardiogram disclosed pathological findings in five patients, but these were not specific. In 12 cases the electrocardiogram was normal. Systolic time intervals contributed least; only three positive results were recorded, and in 14 cases results were normal. Spirometry showed a disturbance of ventilation in 10 patients, which was either of a mixed or restrictive type. The number of positive and negative findings in all examinations carried out is compared in Table 2.

Table 2 Comparison of number of positive and negative findings

<table>
<thead>
<tr>
<th></th>
<th>Positive</th>
<th>Negative</th>
</tr>
</thead>
<tbody>
<tr>
<td>Scan</td>
<td>7 (41%)</td>
<td>10 (59%)</td>
</tr>
<tr>
<td>Electrocardiogram</td>
<td>5 (29%)</td>
<td>12 (71%)</td>
</tr>
<tr>
<td>Systolic time intervals</td>
<td>3 (18%)</td>
<td>14 (82%)</td>
</tr>
<tr>
<td>Spirometry</td>
<td>10 (59%)</td>
<td>7 (41%)</td>
</tr>
</tbody>
</table>

A mutual comparison of positive findings in individuals is more important that their absolute numbers. All four examinations yielded a positive result in one woman only (case 9). In nine patients at least two examinations were positive. Most frequently the results of the scan and spirometry concurred, both examinations yielding positive results in four patients. Electrocardiography and spirometry showed positive findings in three patients. Agreement between systolic time intervals and spirometry was seen in two cases. There was no agreement between the scan and the electrocardiogram or between the scan and systolic time intervals in any patient. In summary, each of the methods of study yielded some positive findings, but there was no clear relation between them.

Discussion

The interpretation of the findings is difficult. Necropsy or biopsy of the myocardium was not carried out in any patients, so we do not know how many had myocardial involvement by scleroderma. Nevertheless, it can be assumed on the basis of the results that, if a positive scintigraphic pyrophosphate scan is found, involvement of the heart in scleroderma can be presumed in patients where this is not clinically evident. In patients with heart failure such studies add nothing, because the involvement of the heart is already evident. Systolic time intervals seem of little help. Unfortunately, echocardiography could not be carried out; a comparison of this with the scintigraphic and electrocardiographic findings would be interesting. Spirometry, indicating possible involvement of the lungs, did not correlate closely with the cardiac results and is no guide to cardiac involvement; unfortunately we did not measure the diffusing capacity of these patients. Thus, of all these, only the pyrophosphate scan and the electrocardiography proved useful in the early detection of cardiac involvement in scleroderma. In this context the discrepancy between the two is surprising.

The interpretation of scintigraphic findings is difficult because we do not know enough about the behaviour of 99mTc-labelled pyrophosphate in damaged myocardium. According to current thinking, the pyrophosphate accumulates in the damaged cardiac muscle if the following three conditions are fulfilled:

1. (1) at least some fresh necrotic tissue must be present;
2. (2) the blood flow through the region affected must be partially preserved; and (3) intramitochondrial calcium phosphate inclusions must be present. A bond of the radiopharmaceutical to these inclusions is assumed. Other authors, however, do not agree with the above hypothesis and assume a bond of the radiopharmaceutical to denatured proteins of damaged myocardial cells. The fact that positive scintigraphic findings occur in fresh lesions only is of fundamental importance. This implies that myocardial involvement in scleroderma can be diagnosed by scintigraphy only while the disease is active, but not retrospectively. This could account for the discrepancy between the electrocardiographic and scintigraphic findings in our patients. Of course, all this is based on theoretical studies dealing with acute myocardial infarction, not with cardiomyopathies, but we believe these observations on the pyrophosphate cardiac scan in scleroderma to be original.

References

6. Duška F, VIžka J, Kubíček J, et al. The sensitivity of scintigraphic myocardial imaging by the use of 99mTc-labelled pyrophosphate in the diagnosis of cardiomyo-
Pyrophosphate heart scan in scleroderma


Requests for reprints to Dr František Duška, Department of Normal and Pathological Physiology, Charles University Faculty of Medicine, Šímkova 870, 500 38 Hradec Králové, Czechoslovakia.
Pyrophosphate heart scan in patients with progressive systemic sclerosis.

F Duska, P Bradna, J Novák, J Kubícek, J Vizda, P Kafka, Y Mazurová and V Blaha

Br Heart J 1982 47: 90-93
doi: 10.1136/hrt.47.1.90

Updated information and services can be found at:
http://heart.bmj.com/content/47/1/90

These include:

Email alerting service
Receive free email alerts when new articles cite this article.
Sign up in the box at the top right corner of the online article.

Notes

To request permissions go to:
http://group.bmj.com/group/rights-licensing/permissions

To order reprints go to:
http://journals.bmj.com/cgi/reprintform

To subscribe to BMJ go to:
http://group.bmj.com/subscribe/