Objective measurement of rheumatoid arthritis using technetium index

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The technetium index, based on the sum of the count rates over several joints, has been shown to correlate with some clinical measurements of disease severity (Oka, Rekonen, and Ruotsi, 1971; Huskisson and others, 1973) but has not been found to be responsive to anti-inflammatory drug therapy, not even to corticosteroids (Berry and Huskisson, 1974).

Methods

Twenty-two patients with classical or definite rheumatoid arthritis (RA) by the A.R.A. criteria (Committee of the American Rheumatism Association, 1959) were studied; all had active disease which had failed to respond to, or progressed despite, optimal anti-inflammatory drug therapy. Eleven patients were treated with penicillamine (doses increasing by 250 mg fortnightly up to a maintenance dose of 1 g daily) and anti-inflammatory drugs, and 11 patients continued with anti-inflammatory drugs alone.

The technetium index was measured before the start of treatment and again after 6 months' treatment. Approximately 200 μCi technetium were injected intravenously and count rates recorded 20 minutes later over both knees and both wrists. Each joint was counted for 20 seconds. The syringe was counted before and after injection, the difference between the two counts representing the dose given. Technetium index was calculated as the sum of the count rates over the four joints divided by the dose and multiplied by 1000 for convenience.

Concurrently with the technetium studies a series of clinical measurements was made by an observer unaware of the patient's treatment and with whom the side effects of the treatment were not discussed. Measurements included pain, using a visual analogue scale, duration of morning stiffness, proximal interphalangeal joint size (Boardman and Hart, 1967), and articular index (Ritchie and others, 1968). Latex tests were carried out at the beginning and end of the study.

Changes in measurements within each group were analysed by Student's 't' test applied to paired data. Correlation coefficients were calculated for changes in technetium index and changes in other measurements. Student's 't' test was also used to test the comparability of the two groups at the start of the study.

Results

There was no significant difference between the two groups in any of the measurements made at the start of the study. There was a significant reduction in technetium index in patients receiving penicillamine but not in those receiving anti-inflammatory drugs (Table). There was also a significant reduction in latex test titres in the penicillamine group, but not in the anti-inflammatory group (t = 3.46; P < 0.01).

Changes in technetium index showed a significant correlation with improvement in pain (r = 0.81; P < 0.01) and joint circumference (r = 0.60; P < 0.05), but not with articular index, duration of morning stiffness, or changes in latex test titre.

Discussion

Objective measurements of the severity and course of rheumatoid arthritis are badly needed especially in view of the unpredictable course of the disease and the striking effects which symptomatic therapy including placebos can have on the subjective manifestations. The technetium index seems to offer such a
Table  Mean technetium index (± standard error) before and after 6 months' treatment with either penicillamine or anti-inflammatory drugs

<table>
<thead>
<tr>
<th>Treatment group</th>
<th>Mean Tc index Before</th>
<th>Mean Tc index After</th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td>Penicillamine</td>
<td>321-3 ± 15-97</td>
<td>243-6 ± 18-58</td>
<td>&lt;0-005</td>
</tr>
<tr>
<td>Anti-inflammatory drugs</td>
<td>308-6 ± 32-48</td>
<td>270-7 ± 19-94</td>
<td>&gt;0-1</td>
</tr>
</tbody>
</table>

measurement, being proportional to the severity of the disease process and responsive at least to one type of therapy.

Penicillamine is classified as a drug with a specific effect in rheumatoid arthritis (Huskisson, 1974). It differs from anti-inflammatory drugs in having effects on extra-articular manifestations of the disease, such as nodules, and on titres of rheumatoid factor. Its effects have been shown to be greater than those of anti-inflammatory drugs alone (Multicentre Trial Group, 1973) including corticosteroids (Ott and Schmidt, 1976). Penicillamine therapy may reduce the technetium index either because it has a more fundamental effect on the disease process, accompanied by reduction in the volume of inflamed synovium, or simply because it has a much greater effect than anti-inflammatory drugs.

Clinical trials of drugs with this type of activity should now include measurement of technetium index.

References


MULTICENTRE TRIAL GROUP (1973) Lancet, 1, 275 (Controlled trial of D-penicillamine in severe rheumatoid arthritis)

