## Studies of reactivity of some Sri Lankan population groups to antigens of *Mycobacterium leprae*. III. The post-lepromin test scar in healthy populations in Sri Lanka

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Summary The occurrence of the post-lepromin scar was investigated in two general population groups, in two geographically different areas, in whom reactivity to lepromin A and SPA of *Mycobacterium leprae* were also studied, earlier. The incidence of the scar was more or less similar to that reported in a patient group in Burma. The incidence of scar was not related to age, sex, race or geographic area (even though examination for the scar was carried out in 3 months at one area and 7 months at the other). The occurrence of the scar increased with BCG vaccination, and was directly related to Mitsuda, Fernandez and SPA reactivity—being most correlated to Mitsuda reactivity and least to SPA reactivity.

#### Introduction

It has been known for a long time that the lepromin test may lead to the formation of a scar, or an area of atrophy. However, no systematic investigation of the latter was made, until 1977.<sup>1</sup> The latter concluded, from an evaluation of the scar in a series of leprosy patients, that 'the post-lepromin scar may be considered as an indicator for a stabilized immune situation...'. The above study was carried out using lepromin of human origin with a bacillary content of 160 million bacilli per ml. The same workers suggested that further studies be carried out using weaker lepromins and non-leprosy affected groups. In this paper we report a study of the scar following lepromin testing in two population groups in geographically different areas of Sri Lanka.

#### Materials and methods

The test populations, the method of the tests, etc., were as described earlier.<sup>2,3</sup> One study<sup>1</sup> considered the post-lepromin scar to be 'best read' at least 3–4 months after the late lepromin reaction, and is perfectly visible after many years. The populations studied were examined for the presence and size

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| Area                              | BCG not vaccinated |    |   |    |       |    | BCG vaccinated |                 |             |   |    |   | Total of |       |    |       |                        |                       |                       |
|-----------------------------------|--------------------|----|---|----|-------|----|----------------|-----------------|-------------|---|----|---|----------|-------|----|-------|------------------------|-----------------------|-----------------------|
|                                   | S                  |    | Т |    | Total |    |                | Total<br>of all | S           |   | -  | Т |          | Total |    | Total | both BCG<br>vaccinated |                       |                       |
|                                   | М                  | F  | М | F  | S     | Т  | М              | F               | individuals | М | F  | М | F        | s     | Т  | М     | F                      | of all<br>individuals | and non<br>vaccinated |
| Mahagastota<br>Scar+              | _                  | 1  | 5 | 10 | 1     | 15 | 5              | 11              | 16<br>(36)  | 1 | 1  | 5 | 11       | 2     | 16 | 6     | 12                     | 18<br>(60)            | 34<br>(46)            |
| Scar –                            | -                  | 2  | 9 | 17 | 2     | 26 | 9              | 19              | 28<br>(64)  | _ | 1  | 2 | 9        | 1     | 11 | 2     | 10                     | 12<br>(40)            | 40<br>(54)            |
| Galagedera <sup>.</sup><br>Scar + | 7                  | 19 | 5 | 6  | 26    | 11 | 12             | 25              | 37<br>(42)  | 9 | 20 | 9 | 14       | 29    | 23 | 18    | 34                     | 52<br>(70)            | 89<br>(55)            |
| Scar –                            | 12                 | 23 | 3 | 13 | 35    | 16 | 15             | 36              | 51<br>(58)  | 2 | 10 | 2 | 8        | 12    | 10 | 4     | 18                     | 22<br>(30)            | 73<br>(45)            |

## Table 1. Occurrence of the post-lepromin test scar

M, male; F, female; S, Sinhala; T, Tamil (percentages within parenthesis).

of the scar at 7 months (at Mahagastota) and at 3 months (at Galagedera). Thus a total of 74 individuals at Mahagastota, and 162 at Galagedera, were examined.

#### Results

In Table 1 the occurrence of individuals with post-lepromin scars is presented. In both areas, the BCG vaccinated showed a greater incidence of the occurrence of scars than those not so vaccinated. There seemed also to be a slight increase in the occurrence of scars at Galagedera (at lower altitude) than at Mahagastota (at higher altitude), but whether this is due to geographic or other variations in the test population, or due to differences in time of examination for scar, are uncertain. However, statistical analysis showed that these differences are not significant.

The incidence of scars with different types of reactivity to antigens of M. leprae (namely, Mitsuda and Fernandez reactivity and that to Soluble Protein Antigen (SPA) of M. leprae) is presented in Tables 2, 3 and 4 respectively.

Table 5 gives a comparative analysis of the percentage incidence of scars in relation to the 3 types of reactivity to *M. leprae* antigens. Linear regression analysis revealed that the occurrence of the scar was highly correlated (at 1% level) with all 3 types of reactivity, the highest degree of relationship being with Mitsuda reactivity and lowest with SPA reactivity. Similarly, statistical analysis was carried out to evaluate the possible variability of the occurrence of the scar with age, sex and race. These variables were not found to effect its occurrence. Another variable, whose effect was obvious is that of BCG vaccination status. The occurrence of the scar is significantly higher in those who have been vaccinated in contrast to those not so vaccinated.

On examination of Mitsuda reactivity, it was found that in a few individuals, Mitsuda reactions had a different quality from what is described<sup>2</sup>—instead of being of the well defined, circumscribed

|                               |                     |                        |                   | Total of I             |                    |                    |  |
|-------------------------------|---------------------|------------------------|-------------------|------------------------|--------------------|--------------------|--|
| Mitsuda reaction<br>size (mm) | Population<br>group | BCG non-<br>vaccinated | BCG<br>vaccinated | BCG non-<br>vaccinated | BCG<br>vaccinated  | Grand total        |  |
| 0–2                           | Mahagastota         | 2 (14)<br>12 (86)      | 2 (40)<br>3 (60)  | 6 (15)<br>35 (85)      | 3 (25)<br>9 (75)   | 9 (17)<br>44 (83)  |  |
|                               | Galagedera          | 4 (15)<br>23 (85)      | 1 (14)<br>6 (86)  |                        |                    |                    |  |
| 3–5                           | Mahagastota         | 4 (36)<br>7 (64)       | 4 (57)<br>3 (43)  | 10 (33)<br>20 (67)     | 12 (55)<br>10 (45) | 22 (42)<br>30 (58) |  |
|                               | Galagedera          | 6 (32)<br>13 (68)      | 8 (53)<br>7 (47)  |                        |                    |                    |  |
| 6–9                           | Mahagastota         | 4 (50)<br>4 (50)       | 7 (70)<br>3 (30)  | 22 (69)<br>10 (31)     | 34 (76)<br>11 (24) | 56 (73)<br>21 (27) |  |
|                               | Galagedera          | 18 (75)<br>6 (25)      | 27 (77)<br>8 (23) |                        |                    |                    |  |
| ≥10                           | Mahagastota         | 3                      | 4                 | 10 (100)               | 19 (100)           | 29 (100)           |  |
|                               | Galagedera          | 7                      | 15                |                        |                    |                    |  |

Table 2. Occurrence of the post-lepromin test scar in relation to Mitsuda reaction size

With scar-upper row; without scar-lower row; percentages within parenthesis.

|                                    |                     |                                 |                    | Total of               |                    |                        |  |
|------------------------------------|---------------------|---------------------------------|--------------------|------------------------|--------------------|------------------------|--|
| Fernandez<br>reaction size<br>(mm) | Population<br>group | BCG non-<br>vaccinated          | BCG<br>vaccinated  | BCG non-<br>vaccinated | BCG<br>vaccinated  | Grand total            |  |
| 02                                 | Mahagastota         | 5 (20)<br>20 (80)               | 7 (44)<br>9 (56)   | 15 (28)<br>38 (72)     | 15 (44)<br>19 (56) | 30 (35)<br>57 (65)     |  |
|                                    | Galagedera          | 10 (36)<br>18 (64)              | 8 (44)<br>10 (56)  |                        |                    |                        |  |
| 3–9                                | Mahagastota         | 5 (39)<br>8 (61)                | 6 (67)<br>3 (33)   | 22 (38)<br>36 (62)     | 29 (67)<br>14 (33) | 51 (50·5)<br>50 (49·5) |  |
|                                    | Galagedera          | 17 (38)<br>28 (62) <sup>,</sup> | 23 (68)<br>11 (32) |                        |                    |                        |  |
| 10                                 | Mahagastota         | 4 (80)<br>1 (20)                | 5 (100)            | 12 (67)<br>6 (33)      | 27 (100)           | 39 (87)<br>6 (13)      |  |
| ≥                                  | Galagedera          | 8 (62)<br>5 (38)                | 22 (100)           |                        |                    |                        |  |

Table 3. Occurrence of the post-lepromin test scar in relation to Fernandez reaction size

With scar-upper row; without scar-lower row; percentages within parenthesis.

|                           |                     |                        |                   | Total of I             |                   |                    |  |
|---------------------------|---------------------|------------------------|-------------------|------------------------|-------------------|--------------------|--|
| SPA reaction<br>size (mm) | Population<br>group | BCG non-<br>vaccinated | BCG<br>vaccinated | BCG non-<br>vaccinated | BCG<br>vaccinated | Grand total        |  |
| 0–2                       | Mahagastota         | 7 (26)<br>20 (74)      | 2 (25)<br>6 (75)  | 14 (24)<br>44 (76)     | 4 (21)<br>15 (79) | 18 (23)<br>59 (77) |  |
|                           | Galagedera          | 7 (23)<br>24 (77)      | 2 (18)<br>9 (82)  |                        |                   |                    |  |
| *3_9                      | Mahagastota         | 3 (100)                | 3 (60)<br>2 (40)  | 14 (48)<br>15 (52)     | 19 (73)<br>7 (27) | 33 (60)<br>22 (40) |  |
|                           | Galagedera          | 11 (42)<br>15 (58)     | 16 (76)<br>5 (24) |                        |                   |                    |  |
| ≥10                       | Mahagastota         | 9 (64)<br>5 (36)       | 14 (88)<br>2 (12) | 26 (62)<br>16 (38)     | 44 (85)<br>8 (15) | 70 (75)<br>24 (25) |  |
|                           | Galagedera          | 17 (61)<br>11 (39)     | 30 (83)<br>6 (17) |                        |                   |                    |  |

Table 4. Occurrence of the post-lepromin test scar in relation to SPA reaction size

With scar—upper row; without scar—lower row; percentages within parenthesis. \* Note: Though 5 mm was considered a 'positive' reaction for SPA reactivity,<sup>3</sup> it was thought that there would be no serious error in considering 3 mm as the demarcation for 'positive'. Therefore in order to have similar class intervals for comparative purposes (Table 5) 3-9 mm is included as the second class interval in this Table.

| Reaction size | BCG     | not vaccinat | ed  | BCG vaccinated |           |     |  |
|---------------|---------|--------------|-----|----------------|-----------|-----|--|
| (mm)          | Mitsuda | Fernandez    | SPA | Mitsuda        | Fernandez | SPA |  |
| 0-2           | 15      | 28           | 24  | 25             | 44        | 21  |  |
| 3–9           | 52      | 38           | 48  | 69             | 67        | 73  |  |
| ≥10           | 100     | 67           | 62  | 100            | 100       | 85  |  |

Table 5. Occurrence of the post-lepromin test scar with different reaction sizes of different types of reactivity to antigens of M. *leprae* (percentages of individuals showing scars)

nodule type—yielding a soft and some times plaque-like reaction. The post-lepromin scar was seen in 2 (of 10 such reactions, all of 3 mm or more) at Galagedera, and 3 (of 11) at Mahagastota—24% of the total, whereas 67% of the total with typical reactions showed scars.

#### Discussion

The results reported here are of a lepromin test with a lepromin of a lower bacillary content  $(4 \times 10^7/ \text{ ml})$ , and of armadillo origin; carried out on a general population group without any evidence of leprous disease. Hence the results of this study may not be directly comparable with those reported in another<sup>1</sup>. It is noteworthy that broadly, the findings of this study appear to be similar to those of the latter workers, with leprosy patients. For example, in the 0–2 mm and 3–5 mm Mitsuda reaction sizes, 17% and 32% respectively of patients in the Walter *et al.* study<sup>1</sup> showed scars (with first or subsequent testing), while in this investigation, 17% and 42% respectively of members of the general population also showed scars (with only 1 test).

The other situation in which scars are presently examined for after an inoculation, is in the assessment of 'coverage' of BCG vaccination programmes—by looking for the presence of scars in those given BCG vaccination.<sup>4</sup> Here it is said that with a potent vaccine, a scar is seen in almost 100% of those so vaccinated,<sup>4</sup> though not invariably so.<sup>5</sup> In Sri Lanka, however, a scar was not found after confirmed BCG vaccination in 3.4% of 750 children between the ages of 3 months and 11 years (at the time of examination), given BCG vaccination at birth.<sup>6</sup> In comparison, the occurrence of scars following lepromin testing is of much lower incidence. If as considered earlier, the post-lepromin scar is an indicator of immunological status,<sup>1</sup> then, as could be expected, lepromin may be considered to be a much weaker vaccine than BCG.

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## NEWS AND NOTES

#### Slide/script and audiotape sets on leprosy (English)

American Leprosy Missions now has available two slide-script and tape sets produced by Roy Pfaltzgraff. Details are as follows:

Leprosy in general practice—this set of 80 slides briefly covers the essentials of the diagnosis and management of leprosy for a medical practitioner who has had little or no previous experience with leprosy. It gives guidelines on the basic features of the disease, the treatment and the main complications that may arise during the active course of the disease, and what to do in case complications arise. (length, 32 minutes).

'Differential diagnosis of leprosy'—consists of 80 slides chiefly of African dermatological conditions that may be confused with leprosy. The emphasis is on making a definite diagnosis by clinical methods in a 'field' situation. It provides a good overview of tropical dermatology. (length, 22 minutes).

These sets are available at the subsidized price of US\$10.00 per set, including a tape and printed script. Apply to: American Leprosy Missions, One Broadway, Elmwood Park, New Jersey 07407, USA.

# Questions and Answers on the Implementation of Multiple Drug Therapy (MDT) in Leprosy. Portuguese translation

This booklet of 35 pages is available, in English and Portuguese, from the Health Unit, OXFAM 274 Banbury Road, Oxford OX2 7DZ, England. It includes a number of questions, and *some* of the answers, relating to problems which may arise in the practical implementation of multiple drug therapy, using the regimens recommended by WHO. Attention has however been drawn, notably by readers in South America, to a number of defects in the Portuguese translation and these include the following: Certain words are inadequately chosen to describe the real meaning, for example: 'exame dos testes microscopicos do corte da pele,' should be 'baciloscopia de espregaço de corte de pele'. For 'pauci-bacilar' substitute 'paucibacilifero'; for 'multi-bacilar' substitute 'multibacilifero'; for 'reincidencia' substitute 'recidiva o recaida'; for 'sitio' substitute 'local o region'; for 'beneficial' substitute 'benéfico'. 'Leprosos' is a derogatory expression that should not be used, the correct term is 'paciente de lepra'.

Further comments on the quality of translation are being collected and a correction slip will be included in copies of the present edition, pending the possibility of a full revision.