Brucellar arthritis: a study of 39 Peruvian families

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SUMMARY A study was conducted to characterise the articular manifestation of *Brucella melitensis* within a family in Peru. From January 1981 to June 1986, 39 families with 232 individuals were evaluated. Brucellosis was diagnosed in 118 family members (attack rate of 50.9%). A lower attack rate was observed in children less than 10 years' old compared with other age groups ($p < 0.02$). Complete clinical data were available in 92 of the 118 affected members. Moderate and severe forms of the diseases were more prevalent in women than in men (41.8% v 13.5%; $p < 0.001$). Twenty eight of the 92 patients developed some brucellar complications; the articular involvement was the most prevalent (23.9%). Arthritis was also more common in women than in men (34.5% v 8.1%; $p < 0.01$). Children appeared to have less articular involvement. Overall, the following pattern was observed: peripheral arthritis (54.5%); unilateral sacroiliitis (23.0%); mixed arthritis (4.5%), and spondylitis (9.1%). Spondylitis was seen only in the elderly with chronic brucellosis. Four patients developed extra-articular rheumatism. Within members of family groups, brucellar arthritis occurred less frequently than in individual patients from the same hospital. This suggests that many family cases were diagnosed in the early stages.

Key words: brucellosis, family studies.

Brucellosis is a zoonosis endemic in certain areas of the world, where it is considered a public health problem.¹ Four species of the *Brucella* genus, each one with a different animal reservoir, are recognised as capable of causing human disease: *B melitensis* (goats), *B abortus* (cows), *B suis* (hogs), and *B canis* (dogs).²

Humans acquire brucellosis by handling contaminated animal products or by eating dairy products made of unpasteurised milk.^{3,4} The clinical manifestations of brucellosis are protean and include rheumatic complaints in 20-85% of the patients. The prevalence and pattern of musculoskeletal involvement depend on the strain of *Brucella* infecting the individual (it is more common with *B melitensis*), the age of the patient, and the duration of the disease.⁵ Arthritis of the sacroiliac and

peripheral joints is more common in younger individuals and in acute disease, whereas spinal involvement usually occurs in the elderly and in those with chronic brucellosis.

Several outbreaks of brucellosis have been reported in individuals infected through a common contaminated edible product (*B melitensis*),⁶⁻⁹ but brucellosis occurring within family groups has been the subject of very few reports.¹⁰⁻¹²

The present study was conducted to determine the clinical characteristics of *B melitensis* in a group of Peruvian families. Special emphasis was placed on the characterisation of the articular manifestations in these individuals.

Patients and methods

This prospective study was conducted through the collaborative efforts of the divisions of infectious diseases and tropical medicine and rheumatology of

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the Cayetano Heredia university hospital in Lima, Perú. Family groups were included if they fulfilled the following criteria: (a) proband with a definite diagnosis of brucellosis (positive blood, bone marrow and/or synovial fluid cultures and/or brucellar febrile agglutinins at a titre $\geq 1/160$); (b) at least two affected family members within the same household; (c) complete pedigree data; (d) interval between brucellosis in the proband and other family member(s) not longer than 120 days.

All afflicted individuals were evaluated by a rheumatologist for the presence or absence of musculoskeletal involvement: peripheral arthritis, sacroiliitis, spondylitis, extra-articular rheumatism, or combinations, as previously described.^{5 13}

The severity of brucellosis was graded according to the presence or absence of complications as mild (without complications), moderate (a non-fatal complication), and severe (a potentially fatal complication).

The course of the disease was defined by the duration of symptoms as acute (eight weeks' duration), subacute or undulant (more than eight weeks and less than one year with intervening periods of remission during the course of the disease), and chronic (more than one year).⁴

Results

From January 1981 to June 1986 47 family groups were studied; eight did not meet the inclusion criteria and were excluded. A total of 232 individuals from 39 families was available for evaluation. Age distribution is shown in Table 1. Brucellosis was diagnosed in 118 family members, giving an attack rate of 50.9%. The attack rate was similar in men and women (51.1% v 50.7%), but it was lower in individuals younger than 10 when compared with the other age groups (Fig. 1, $p<0.02$).

Complete clinical data were available in 92 (78%) of the 118 affected family members. The prevalence of acute, subacute, and chronic brucellosis was similar in both sexes and in all age groups (data not shown); moderate and severe disease was more prevalent in women than in men (41.8% v 13.5%, $p<0.001$) and in patients with subacute or chronic brucellosis compared with those with acute disease (53.1% v 18.3%, $p<0.01$).

Twenty eight (30%) of the 92 patients developed 33 complications, musculoskeletal involvement being the most frequent (Table 2). Articular manifestations were more common in women than in men (34.5% v 8.1%, $p<0.01$). Children appeared to have articular involvement less often than adults,

Table 1 Attack rate in brucellosis by age groups in 232 members of 39 families

Age group (years)	Affected	Unaffected	Attack rate (%)
Under 10	11	31	26.2
10-19	36	27	57.1
20-29	25	19	56.8
30-39	17	18	48.6
40-49	9	6	60.0
50-59	11	8	57.9
Over 60	9	5	64.3
Total	118	114

Fig. 1 Attack rate of brucellosis by age group in 39 Peruvian families.

Table 2 Complications in the course of brucellosis in 92 members of 39 Peruvian families

	No	%
Musculoskeletal	22	23.9
Neurological	3	3.3
Hepatic	3	3.3
Dermatological	2	2.2
Haematological	2	2.2
Ocular	1	1.1

Table 3 Age distribution of brucellosis patients with articular involvement

Age group (years)	No of cases	Articular involvement	
		n	%
Under 15	22	3	13.6
15-35	45	12	26.7
35-55	15	4	26.7
Over 55	10	3	30.0
Total 92		22	23.9

Table 4 Pattern of articular involvement in 22 patients with brucellosis from 39 Peruvian families

	Total No (patients)
Peripheral arthritis*	12
Monoarthritis (knee 6, ankle, MTP PIP-hand, 1 each)	9
Polyarticular (pauciarticular)	3
Sacroiliitis (all unilateral)	5
Spondylitis	2
Mixed arthritis	1
Extra-articular rheumatism*	4

*Two patients had both peripheral arthritis and extra-articular rheumatism.

MTP=metatarsophalangeal; PIP=proximal interphalangeal.

but the differences were not statistically significant (Table 3). All patterns of articular involvement, as described by Gotuzzo *et al.*,⁵ were seen (Table 4). Peripheral arthritis (54.5%), usually affecting joints of the lower extremities, and unilateral sacroiliitis (23%) were the most common patterns of joint involvement. Spondylitis was seen in the elderly with chronic brucellosis, and simultaneous involvement of the sacroiliac and peripheral joints in one patient. Four patients developed extra-articular rheumatism, two concomitantly with peripheral arthritis.

Discussion

There are few reports on the characteristics of

brucellosis occurring in families. Spink studied nine family groups over a period of 16 years in an endemic area for *B abortus*, emphasising the relatively low rate of symptomatic to infected individuals in these family groups, but the clinical characteristics of the symptomatic patients were not the thrust of his communication.¹⁰ Feiz *et al* put particular emphasis on the clinical features of this infection in children but did not present actual family data.¹¹

Since brucellosis by *B melitensis* biotype 1 is epidemic in the coastal cities of Perú, particularly in Lima,¹⁵ a unique opportunity for the study of the clinical characteristics of this disease when occurring in family groups was available to us. In 39 families afflicted with brucellosis, and followed up for 6 months, half of the individuals became symptomatic, confirming the high degree of infectivity of *B melitensis*.²⁻³ Subclinical infection such as that described with *B abortus*¹⁰⁻¹⁶ was unlikely as serological testing conducted in four of these families did not demonstrate seroconversion (data not shown).

There were clear differences in the attack rate among the various age groups, with the lowest in the under 10 and the highest in the over 55 year group. More than one factor is likely to explain these differences: less exposure to contaminated food, more efficient clearance of *Brucella* by the oropharyngeal lymphoid tissue of children, and the relatively low gastric pH among youngsters. On the other hand, a higher attack rate in older adults may also reflect a relatively deficient immune system in these individuals as part of the normal aging process.¹⁷

In this study the prevalence of brucellosis was similar in both sexes, but the clinical manifestations, including articular involvement, were more severe in women. This observation has not been reported before to our knowledge, probably because brucellosis in many places is an occupational disease¹⁸⁻¹⁹ occurring mainly in men. It may also be explained by the high prevalence of iron deficiency anaemia in our female population,²⁰ since microorganisms of the *Brucella* genus show a greater degree of invasiveness and more rapid multiplication when inoculated into iron deficient hosts.²¹ Genetically determined susceptibility factor(s) could also play a part in predisposing women to more severe disease and arthritis, but most immunogenetic studies have not shown an association between a given HLA specificity and osteoarticular involvement in brucellosis.¹²⁻²²⁻²⁴

All previously described musculoskeletal manifestations of brucellosis were seen in these families, with peripheral arthritis and sacroiliitis occurring mainly in the younger age groups and spondylitis

more frequently in older adults, as seen by others.^{25 26} The lower prevalence of articular involvement in family members as compared with our patients in general may be due to earlier diagnosis in these families and the consequently reduced likelihood that afflicted individuals will go on to develop subacute or chronic disease.

In summary, the study of these families in a highly endemic area for brucellosis demonstrates the infectivity of *B melitensis* compared with that of *B abortus*. A lower attack rate in children and more severe disease in women were also observed. Finally, arthritis occurred less frequently within these families compared with patients with brucellosis from the same geographical area, suggesting that many cases were diagnosed earlier than usual. Exposure to a highly contaminated food product(s) is the most likely explanation for the clustering of cases within families.

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