

Figure 1 Relation between onset of rheumatoid arthritis and menopause in 191 postmenopausal women in a case-control study.

have arisen by chance, we tentatively suggested that other unknown genetic factors might have been responsible. We would be grateful for any other interpretations of the data. Unlike Dr McHugh we are unaware of the conclusive clinical evidence that pregnancy loss ameliorates existing disease, though this remains a possibility.

Dr McHugh suggests that the effect of the menopause should be given wider consideration owing to the age of onset in women. Dr McHugh's study showed a peak age of onset around the menopause, as did our larger case-control study (fig 1). There are a number of problems that have to be considered before a direct correlation can be postulated. Firstly, the age of onset distribution seen in hospital outpatient studies is not representative of all women with RA. This is because hospital case groups tend to contain those with more severe, longstanding disease, and patients developing RA at an older age will be underrepresented (fig 2). Population surveys are the only accurate way of determining ages of onset. In the United Kingdom these have confirmed that prevalence does indeed increase at the menopause, but remains at this level thereafter.¹ Another study of incidence related to age at the Mayo clinic found that this increased steadily in women and men.² If the menopause were a major factor one might expect differences in prior age of menopause or incidence of artificial menopause and these have not been found in studies.³⁻⁵ As well as the mentioned bias due to recall of disease onset, it should be remembered that surveys have shown that up to 50% of women with

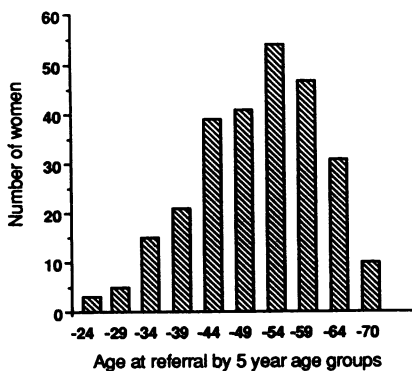


Figure 2 Distribution of age at onset of women with rheumatoid arthritis from a case-control study. (Source: Spector TD. Hormonal and reproductive factors in the aetiology of RA. MD thesis, University of London, 1989.)

menopausal symptoms have joint pain,⁶ and this is likely to be confused with early RA.

A number of studies have looked at hormonal status of women with RA, and, unfortunately, no consistent abnormalities have been detected.⁷ We are just completing a prospective study examining the hypothesis that hormone replacement therapy (HRT) is protective against the development of RA. The data on the possible protective effects of HRT are patchy and contradictory at present and suffer from many of the methodological problems outlined above. Although hormonal change at the menopause may be a factor in RA, current evidence suggests that prior pregnancy and oral contraceptive use at an earlier age may be of more importance.⁸

TIM D SPECTOR

ALAN J SILMAN

Departments of Environmental
and Preventive Medicine and Rheumatology
Medical College of St Bartholomew's Hospital
London EC1M 6BQ
and
ARC Epidemiology Unit
Manchester M13 9PT

- Lawrence J S. *Rheumatism in populations*. London: Heinemann, 1977.
- Linos A, Worthington J W, O'Fallon W M, Kurland L T. The epidemiology of rheumatoid arthritis in Rochester, Minnesota: a study of its incidence, prevalence and mortality. *Am J Epidemiol* 1980; 111: 87-98.
- Vandenbroucke J P, Wittman J C, Valkenburg H A, et al. Noncontraceptive hormones and rheumatoid arthritis in perimenopausal and postmenopausal women. *JAMA* 1986; 255: 1299-303.
- Short C L, Bauer W, Reynolds W E. *Rheumatoid arthritis*. Cambridge, Mass: Harvard University Press, 1957.
- Spector T D, Brown G C, Silman A J. Increased rates of previous hysterectomy and gynaecological operations in women with osteoarthritis. *Br Med J* 1988; 297: 899-900.
- Neugarten B L, Kraines R J. "Menopausal symptoms" in women of various ages. *Psychosom Med* 1965; 27: 266-73.
- Spector T D. Sex hormone measurements in RA. *Br J Rheumatol* 1989; 28 (suppl 1): 62-9.
- Spector T D, Roman E, Silman A J. The pill, parity and rheumatoid arthritis. *Arthritis Rheum* (in press).

Sternoclavicular erosions in polymyalgia rheumatica

Sir: We read with interest the article by Kyle *et al* about the rarity of synovitis in polymyalgia rheumatica.¹ In their work sternoclavicular erosions were seen in only 2/19 of the tomograms obtained. Our experience contradicts these results.²

From 1984 to 1987 tomography of sternoclavicular joints was carried out in 21 patients (12 female, nine male) with active untreated polymyalgia rheumatica, seven of whom also had giant cell arteritis. Their mean age was 68.8 years. All had a negative Rose-Waaler test. In addition, control tomograms were obtained from 18 volunteers (mean age 69.2 years) who had no clinical evidence of inflammatory arthropathy. The films were all examined by one consultant radiologist and two rheumatologists with no knowledge of clinical data.

The incidence of erosions was significantly higher ($p < 0.05$) in patients with polymyalgia rheumatica (9/21) than in control subjects (1/18). Erosions were found more often in patients who had had symptoms for more than six months. There was no correlation between local pain elicited by pressure and the presence of articular erosions.

Our findings support the existence of an erosive arthropathy in polymyalgia rheumatica and are in agreement with the study of Paice *et al* previously published in the *Annals*.³

J M NOLLA*

J VALVERDE

Department of Rheumatology
Hospital of Bellvitge
University of Barcelona
Barcelona, Spain

*Correspondence to: Dr J M Nolla, Department of Rheumatology (10-2), Hospital of Bellvitge, 08907 L'Hospitalet de Llobregat, Barcelona, Spain.

- Kyle V, Tudor J, Wraight E P, Gresham G A, Hazleman B L. Rarity of synovitis in polymyalgia rheumatica. *Ann Rheum Dis* 1990; 49: 155-7.
- Nolla J M, Valverde J, Andia E, et al. Erosiones esternoclaviculares en la polimialgia reumática. *Revista Española de Reumatología* 1989; 16: 41-3.
- Paice E W, Wright F W, Hill G S. Sternoclavicular erosions in polymyalgia rheumatica. *Ann Rheum Dis* 1983; 42: 379-83.

Immunological indices of patients with rheumatoid arthritis after methylprednisolone pulse therapy

Sir: In a previous letter to the *Annals* we reported the favourable clinical results obtained by high dose intravenous methylprednisolone treatment in active rheumatoid arthritis.¹ We now present the changes in the immunological indices for these same patients.

The discovery of monoclonal antibodies has enabled the determination of different lymphocyte subgroups.²⁻⁹ We investigated the cellular and humoral immune response of 10 patients with active, classical rheumatoid arthritis. The total lymphocyte count (OKT3), the inducer/helper (OKT4) and suppressor/cytotoxic (OKT8) subgroups were determined with Orthomune monoclonal antibodies by the immunoperoxidase method according to Erber.¹⁰ Concentrations of C3, C4, and the immunoglobulins were measured by Mancini's radial immunodiffusion. Determination of immune complexes was by polyethylene glycol precipitation and cross electrophoresis. Blood samples for determinations were taken before infusion and after 2, 6, 24, and 48 hours.

Before infusion we found a high level of helper cells: mean 69.1% (range 59-78%), absolute number 1921/ μ l (1120-3321), and a low concentration of suppressor cells: mean 25.8% (21-34%), absolute number 808/ μ l (352-1700), in agreement with published data.^{2 4 6 7 11-15} The so-called immunoregulatory ratio—the helper/suppressor quotient—was increased to 2.65 (normal 1.55), as found by others.^{2-4 6 11} This disequilibrium of the immune response was favourably influenced by high dose intravenous methylprednisolone pulse therapy, which decreased the centile proportion: mean 45.3% (24-63%) and the absolute number: mean 418/ μ l (135-715) of T4 cells, thus leading to a normal helper/suppressor ratio. According to Bertouch the cause of selective diminution of helper cells is the remigration of the circulating lymphocytes into the peripheral lymph nodes and bone marrow.^{2 16 17} As the lymphopenia was only transitory and the excretion of uric acid did not rise after the pulse treatment the decrease of cell numbers is probably due to a change in the division of circulating cells and not to cell lysis.¹⁸