The ESR in the diagnosis and management of the polymyalgia rheumatica/giant cell arteritis syndrome

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SUMMARY A review of 80 patients with the polymyalgia rheumatica/giant cell arteritis (PMR/GCA) syndrome has revealed that the erythrocyte sedimentation rate (ESR) is normal in 22.5% of cases at presentation. Too much reliance on the ESR as a screening test led to delay in diagnosis in 10 patients. Serious complications occurred in 4 patients with a normal or minimally raised ESR. Analysis of follow-up data at 4 weeks showed no significant correlation between change in the ESR and change in symptoms, and at subsequent clinical attendances symptoms did not consistently parallel the ESR. Our results cast serious doubts on the use of the ESR as an entirely reliable diagnostic and therapeutic aid in the PMR/GCA syndrome.

Considerable emphasis has been placed on the ESR in the polymyalgia rheumatica/giant cell arteritis (PMR/GCA) syndrome. Most workers consider a markedly raised ESR to be essential for the diagnosis.¹⁻³ The raised ESR is also accepted as a reflection of disease activity and consequently it is used to regulate corticosteroid dosage.⁴ Conversely a normal ESR has been taken to exclude the diagnosis³ or to imply quiescence of disease activity, despite reported cases of PMR/GCA with a normal ESR.⁵

We have undertaken a retrospective study of the ESR in patients admitted to the Aberdeen Hospitals with a diagnosis of PMR/GCA. In this study the relationship of the ESR to clinical findings at various times in the disease process is reported and an assessment of the role of the ESR in diagnosis and management is presented.

Patients and methods

Records were obtained by using the International Classification of Diseases index number for PMR/GCA over the period 1968-78. Of the 97 case records requested 6 were rejected because of an error in diagnostic classification and a further 11 cases because they did not satisfy fully our diagnostic criteria. Both the typed and hand-written entries in the remaining 80 case notes were scrutinised, and data were obtained from a total of 758 hospital

attendances. Duration of follow-up varied from 4 weeks to 10 years.

The diagnostic criteria are shown in Table 1. Patients were accepted if criteria 1-3 were present plus any 3 of criteria 5-10 or criterion 4. Patients with evidence of myeloma, rheumatoid arthritis, systemic lupus erythematosus, polymyositis, and disseminated malignancy were excluded. The ESR in all cases was performed by the Westergren method.⁶

We made particular note of initial presentation, the time at which a trial of steroid therapy was thought to be indicated (diagnosis), and follow-up after 4 weeks on corticosteroid therapy. In assessing response to treatment the patients were grouped by symptom type as follows: (PMR), proximal muscle pain and stiffness; (GCA), headache, jaw claudication, scalp tenderness, and systemic weight loss, anorexia, malaise. A graded clinical score was em-

Table 1 *Diagnostic criteria*

1.	Age greater than 55 years
2.	Positive response within 48 hours to corticosteroid therapy
3.	Length of history greater than 2 weeks
4.	Positive temporal artery biopsy
5.	Proximal, symmetrical girdle or upper arm muscle pain $\pm$ stiffness $\pm$ tenderness
6.	Jaw claudication
7.	Clinical abnormality of a temporal artery (tenderness, thickening, redness)
8.	Systemic symptoms or signs (malaise, anorexia, weight loss, anaemia, pyrexia)
9.	Temporal headache
10.	Visual disturbance (loss, diplopia, blurring)

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ployed to assess disease activity (0: asymptomatic to 4: increasingly severe). Correlation coefficients were obtained by means of Pearson's product moment correlation method.

Results

Clinical features at diagnosis. These are summarised in Table 2.

ESR at presentation. A total of 18 patients (22.5%) presented with an ESR of <30 mm/h (Fig. 1). Of this group 8 were diagnosed immediately on clinical grounds alone and therapy with corticosteroids was begun.

In the 10 patients with an ESR <30 mm/h not diagnosed immediately on clinical grounds there was a mean interval of 106 days from presentation or institution of corticosteroid therapy. This compares with an interval of 35 days in those presenting with an ESR >30 mm/h.

ESR and clinical features. In Table 3 correlation coefficients are shown between ESR at presentation and other variables. Statistical analysis confirmed a highly significant correlation between ESR and anaemia at both presentation and diagnosis ($p<0.001$). There was no significant correlation between presenting ESR and subsequent development of visual loss or any other variable. Four patients in the series developed visual loss with either a normal or slightly raised ESR at the time of visual loss (ESR = 11, 21, 30, 40).

ESR and management of patients. On initiation of corticosteroid therapy all the patients had rapid relief of symptoms and the ESR also tended to fall (Fig. 1). When the degree of symptomatic improvement at 4 weeks as assessed by clinical score was compared with the numerical change in the ESR, however, we found no significant correlation ($r = 0.088$). On analysis of follow-up data on a further 678 atten-

* ESR on one occasion was not repeated at diagnosis.

** ESR result unavailable on six follow up attendances; one patient died between diagnosis and follow up.

Fig. 1 ESR at presentation, diagnosis, and follow-up

Table 3 Correlation coefficients

	Presenting ESR
Age	0.073
Sex	0.104
Duration of symptoms	-0.092
PMR symptoms	-0.111
GCA symptoms	-0.066
Systemic symptoms	0.129
Haemoglobin	-0.637*
Visual loss	0.158
Visual disturbance	0.132
Positive biopsy	0.194

* $p<0.001$.

Table 2 Clinical features at diagnosis

	Number of patients	
	Male	Female
Total	19	61
Predominately PMR	0	4
Predominantly GCA	8	31
Mixed	11	26
Systemic symptoms	11	33
Number of biopsies	16	46
Positive biopsies	5	28
Anaemia (Hb <10.5 g/dl)	5	16
Jaw claudication	6	9
Pain on swallowing	3	6
Visual loss	4	13
Visual disturbance	3	8
Average age of patients	74 (SD = 8.5)	69 (SD = 7.6)

dances at outpatient clinics there did not appear to be a universally accepted policy for the adjustment of corticosteroid therapy, and little emphasis was given to other medical conditions known to affect the ESR.⁷ Adjustments in corticosteroid therapy were made on 191 (28.1%) of the above attendances owing to changes in the patients' symptoms, the ESR, or both. Symptoms and ESR paralleled each other on only 44 (23%) of these occasions. In the remainder physicians varied in their treatment policy when an asymptomatic patient presented with a raised ESR or vice versa. On 77 (40%) occasions therapy was altered on the basis of the ESR alone. There were 47 documented instances in 37 symptomatic patients of intercurrent illness—mainly urinary and respiratory

infections. The ESR was raised (>30 mm/h) in 41 of these instances but only on one occasion was the rise attributed to the intercurrent illnesses. In the remainder the raised ESR was thought to reflect an exacerbation of the PMR/GCA syndrome, and therapy was adjusted on this basis.

Discussion

Standard textbooks^{8,9} and clinical reviews¹⁻⁴ emphasise that the ESR is almost invariably raised in the PMR/GCA syndrome, values in excess of 50 mm/h being commonly found,¹⁻⁴ and this level is said by some to be a sine qua non for the diagnosis.¹ However, an 'elevated' ESR is not clearly or consistently defined in the literature. Westergren's original work put the upper limits of normal as 3 mm/h for males and 7 mm/h for females. The accepted limit of normal has risen considerably since then. In a large series of 2500 healthy men and women it was found that the upper limit of normal for the ESR above the age of 50 years was 20 mm/h for men and 30 mm/h for women.¹⁰ Values between 20 and 40 are difficult to interpret,¹¹ but we have adopted a value of >30 mm/h as 'elevated'. Recently others have used this value of >30 mm/h as a diagnostic criterion for PMR/GCA.¹² By this definition a normal ESR was found in 22.5% of the patients at presentation. This is a much higher incidence than in other series.^{4,13-15} It may be argued that the heterogeneous selection of patients in our series may have influenced the ESR, as it has been claimed that patients with GCA syndrome alone tend to have a higher ESR.¹³ We have failed to show any significant difference in the ESR for different symptom types, however (Table 3).

A normal ESR in patients with otherwise classical PMR/GCA tended to mislead physicians, resulting in a serious delay in diagnosis, consistent with the commonly held belief that the diagnosis of PMR/GCA is untenable in the absence of a raised ESR. There was no significant correlation between ESR at presentation and the subsequent development of visual loss. The ESR cannot therefore be used to predict which patients will go on to develop this complication. Furthermore 3 patients in our series had a normal ESR at the time of visual loss, and in one the ESR was only slightly raised at 40 mm/h.

On occasions when an asymptomatic patient presented with a raised ESR, or the converse, the physician's treatment policy varied between treating the patient's symptoms irrespective of the ESR and directing treatment at the ESR alone. The literature is also divided on this point, however. Wadman and Werner¹⁶ noted that, although nonsteroidal anti-inflammatory drugs were successful in relieving symptoms of polymyalgia rheumatica, they did not prevent the development of visual complications.

The same hazard was noted with doses of steroids insufficient to suppress the ESR.¹⁷ More recently, however, it has been reported that after the initial period steroid dosage may be kept at the minimum required for symptomatic relief, irrespective of the ESR, without the development of complications.¹⁵ None of the patients in our series developed visual loss after the initial 48 hours on corticosteroid therapy. Doubts have been cast on the usefulness of the ESR in the diagnosis of other disease processes.¹⁸ Perhaps the difficulties associated with performing the test uniformly and the presence of several different factors affecting the ESR at any one time¹¹ tend to make it an inherently inaccurate diagnostic test.

From our data it is evident that the ESR is not an entirely reliable guide in the diagnosis or monitoring of disease activity in the PMR/GCA syndrome, nor can it predict the development of complications.

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