LETTERS TO THE EDITOR

Paraneoplastic non-caseating granulomatous inflammation of the eyelid

EDITOR,—Periocular granulomatous inflammation is most commonly due to a chalazion or sarcoidosis, and more rarely to allergic granulomatosis (Churg-Strauss), Erdheim-Chester disease, Wegener’s granulomatosis, and necrobiotic xanthogranuloma. When no cause for this inflammation can be found, it may be called idiopathic non-infectious granulomatous inflammation1 or orbital sarcoidosis.2 We report a case in which this idiopathic inflammation appeared to be associated with squamous cell carcinoma of the lung.

CASE REPORT
A 77-year-old man presented with a 2½ year history of non-tender inflammation of the left upper eyelid (Fig 1). There was no history of injury to the eyelid. A chest x-ray showed an abnormality that was thought to be due to overprojection. Systemic steroids reduced the inflammation temporarily. After referral to our hospital, extensive clinical and laboratory investigations were performed. Computed tomography scanning showed the lacrimal gland to be of normal size. Tests for tuberculosis, other infectious diseases, and systemic inflammatory disease were negative. An incisional biopsy of the eyelid revealed a non-caseating, granulomatous inflammation without foreign body material or acid fast bacilli. In a few granulomas some B cells could be demonstrated immunohistochemically (Fig 2). A new chest x-ray revealed a coin lesion of the right upper lobe and a lobectomy was performed. Histopathological examination of the specimen showed a poorly differentiated squamous cell carcinoma of the lung without signs of lymph node metastasis or sarcoidosis. Systemic corticosteroids and non-steroidal anti-inflammatory drugs had no significant effect on the inflammation in the eyelid. Some more tissue was removed for cosmetic reasons, and the histopathology was similar to the first specimen.

COMMENT
The association of granulomatous inflammation (especially in lymph nodes) and malignancy has been noted before.2-4 The pathogenesis of this malignancy associated inflammation may be diverse (sarcoid-like reaction to tumour derived components, sarcoidosis, infection, granulomatous lesions of unknown significance or GLUS).2 Sarcoid-like reactions to pulmonary neoplasms were described to occur in regional lymph nodes in the form of non-caseating granulomas.3 Granulomas in lymph nodes may be divided into two categories on the basis of the presence or absence of B cells in these lesions. B cell positive: toxoplasmosis, tumour related sarcoid reactions, and GLUS; B cell negative: sarcoidosis, mycobacterial infections.5 In our case, a few granulomas showed some admixture with B cells.

To our knowledge, the association of a non-caseating granulomatous inflammation of the eyelid and a squamous cell carcinoma of the lung has not been described before. This case suggests that screening for malignancy might be useful in patients with non-caseating granulomatous inflammation of unknown origin.

TTQ REUSER
Department of Ophthalmology
P WESSELING
Department of Pathology,
University Hospital Nijmegen, the Netherlands

Correspondence to: TTQ Reuser, Department of Ophthalmology, University Hospital Nijmegen, PO Box 9101, 6500 HB Nijmegen, the Netherlands.

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Paraneoplastic retinopathy in association with large cell neuroendocrine bronchial carcinoma

EDITOR,—Cancer associated retinopathy is a rare paraneoplastic manifestation of a variety of tumours, most commonly small cell carcinoma of the lung (SCCL).1-4 Characteristically the syndrome presents with reduced vision, photopsia, progressive ring scotomas in the visual field, and nightblindness. Ocular symptoms may precede the diagnosis of malignancy for several months, especially as the ocular signs (attenuation of retinal arteries, mild optic disc atrophy) are easily missed. Pathologically, there is a severe photoreceptor degeneration,5 and it is thought that the disease is mediated by antibodies that cross react between the tumour and photoreceptor antigens.1,4 We present a 60-year-old ex-smoker with the typical features of the syndrome whose investigations revealed a large cell neuroendocrine carcinoma.

CASE REPORT
A 60-year-old woman presented with a 1 year history of dimming of vision associated with severe nightblindness. A left cataract had been removed 1 year earlier. There was no relevant family or dietary history and she had taken no potentially retinotoxic drugs. More recently she noticed increasing malaise and weight loss. General examination was normal except for an enlarged right supraventricular node. Ocular examination showed a vision of 6/24 right, 6/6 left with normal colour vision. Pupils reacted poorly and visual fields (Fig 1) showed gross constriction, the right eye being worse than the left. Fundus examination showed minimal arteriolar constriction and

Figure 1A Figure 1B

1 Goldmann visual fields showing marked constriction in the right eye (A) and a dense ring scotoma in the left eye (B).

References
cell neuroendocrine carcinoma. Previous reports have demonstrated SCC1 to be the commonest tumour association,\textsuperscript{1,4} others being related to carcinoma of the breast,\textsuperscript{5} cervix,\textsuperscript{6} and uterus.\textsuperscript{7} There is one previous report of a large cell anaplastic lung tumour\textsuperscript{2} but neuroendocrine features\textsuperscript{8} were not detected. The characteristic clinical picture of progressive nightblindness, ring scotomas, and eventual visual loss point to a process of photoreceptor degeneration which has been confirmed pathologically.\textsuperscript{3,7} The underlying pathogenesis for the association is thought to be due to the production of antibodies that cross react between tumour and retinal tissue. Grunwald et al\textsuperscript{4} have shown that such antibodies are only found in the sera of patients with both the relevant tumour and visual loss. The antibody is not found in those with visual loss and no tumour, nor in those with tumours but no visual loss. The retinal protein with which the sera of these patients react remains controversial. The immunofluorescent staining pattern in our patient is similar to that found by Kornguth\textsuperscript{2} and Grunwald\textsuperscript{9} with antibody largely directed against the ganglion cell and inner nuclear layers, whereas Keltner found antibody predominantly staining photoreceptor inner segments.\textsuperscript{10} The 23 kDa protein, recently acknowledged to be the photoreceptor protein recoverin\textsuperscript{11} and which has been postulated to be specific for the syndrome associated with SCC1\textsuperscript{7} was not revealed by immunoblotting in our patient. This finding agrees with a recent study which found only one in 10 patients with cancer associated retinopathy demonstrating antibodies to this protein.\textsuperscript{12} The most important clinical feature of this syndrome is to consider and then recognise the diagnosis and to realise that symptoms may precede detection of the underlyng tumour for many months or years. Electrodagnostic studies are important and immunological studies in future patients may clarify the relation between the retina and the tumour.

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Comment

This is the first reported case of paraneoplastic retinopathy found in association with large panel 2A: Large cell carcinoma with large nuclei. Note positive granular staining in the extensive cytoplasm of tumour cells for chromogranin A.avidin-biotin method of immunostaining (x200).

Figure 2B: Tumour cell containing large dense core granules within the cytoplasm. Unravel acetoce and lead citrate staining (x18 000).

Cryptococcus presenting as cloudy choroiditis in an AIDS patient

EDITOR

Cryptococcus neoformans is a well known organism which causes opportunistic infection in patients with AIDS. Patients commonly present with meningitis and are referred to the ophthalmologist with extra-ocular muscle paresis and papillodema.\textsuperscript{1} However, other ocular manifestations are rare. Cryptococcus has also been described: the clinical features are cells in the vitreous with focal choroidal lesions; the presence of the fungus in the choroid implies haematoogenous spread and, consequently, is associated with a terrible prognosis.\textsuperscript{3} This paper describes a patient whose cryptococcal infection initially manifested with eye symptoms due to a hitherto unrecognised pattern of choroidal disease.

Case report

A 39-year-old white homosexual man with AIDS presented in May 1992 with a week history of blurred vision in both eyes. He complained, in particular, that everything appeared wavy. He had been HIV positive since 1988 and had developed Kaposi’s sarcoma in April 1992 when his CD4 count was 0-02×10\textsuperscript{9}/ml. Ophthalmic examination revealed visual acuities of 6/5 in each eye, full colour vision, and mild constriction to the 12e and 14e targets in both eyes on Goldmann perimetry. The pupillary reactions were normal. The eyes were white and slit-lamp examination was unremarkable with no
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M R Stanford, C E Edelsten, J D Hughes, M D Sanders, C I Brooks, D Mitchell and M N Sheppard

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