Infective endocarditis complicating psittacosis: response to rifampicin

Endocarditis rarely complicates Chlamydia psittaci infection. We describe a patient who developed insidious progressive aortic valve regurgitation while the psittacosis/lymphogranuloma venereum complement fixation (LGVCVF) titres were rising. The infection responded to rifampicin after treatment with tetracycline had failed.

Case report
A previously healthy 43-year-old electrician was first admitted to hospital in August 1978 with a seven-day history of generalised aches, sweating, malaise, oliguria, vomiting, and frontal headaches. He kept pigeons but had had no contact with a parrot. There had been no ill health among his pigeons—as verified later by the public health authorities. He was feverish (40°C) with sinus tachycardia of 120/min. There were signs of left upper lobe consolidation. Cardiac auscultation was normal. Investigations: haemoglobin 14·9 g/dl; white cell count 3·4 x 10^9/l (3400/mm³), 68% neutrophils, 22% lymphocytes, 10% monocytes; erythrocyte sedimentation rate 120 mm 1st h (Westergren); arterial oxygen pressure 7·8 kPa (58 mm Hg) and arterial carbon dioxide pressure 3·1 kPa (23 mm Hg) on room air; urine microscopy showed numerous red cells, granular casts, and white cells; serum sodium 133 mmol/ml; potassium 3·8 mmol/ml; blood urea 7·0 mmol/l (42 mg/100 ml); total serum protein 57 g/l; albumin 28 g/l; aspartate aminotransferase (AST) 174 IU/l (normal 5-32); gamma-glutamyl-transpeptidase 49 IU/l (normal up to 45). Protein electrophoresis showed increases in a₁ and a₂ globulins. Chest x-ray examination confirmed left upper lobe consolidation. Tuberulin test, electrocardiogram (ECG), fluorescence antibody test against legionnaire's disease, immunoglobulins, blood cultures, and putum examination were normal.

He was initially thought to have a "bacterial" pneumonia and was given courses of amoxycillin, flucloxacillin, and erythromycin. He failed to improve and deteriorate progressively and died of respiratory failure and disseminated intravascular coagulation. He had developed a pulmonary embolus manifest radiologically as right lower zone collapse/consolidation of the lung. Venography showed extensive thrombus in the right femoral vein in the upper two-thirds of the thigh and the deep veins of the calf. He improved somewhat on anticoagulants. Fever settled and there was radiological resolution of the left upper lobe consolidation. He was discharged home. The results of virus studies were negative. He was readmitted with worsening dyspnoea. A summation gallop was accompanied by signs of aortic incompetence. Chest x-ray examination showed a small right basal effusion and resolving pneumonia of the left upper lobe. Serial blood cultures were again negative. ECG showed first-degree heart block (P-R interval 0·72 s) with lower septal defect over the left precordial leads. The psittacosis/LGVCVF titre had risen to 1/32. Anticoagulation was continued. He was given oxytetracycline 4 g daily for six weeks. There was no clinical response and the psittacosis/LGVCVF titre rose to 1/512 (December 1978). Since he had already had erythromycin he was given rifampicin 600 mg daily for a further 16 weeks. It was then stopped. His condition improved. He is now well although signs of trivial aortic incompetence persist. The psittacosis/LGVCVF titre fell steadily to 1/32 in May 1979.

Comment
Chlamydia are recognised as potential cardiovascular pathogens and are implicated in various acquired valvular heart diseases. Of the 107 cases of psittacosis dealt with by the Regional Virus Laboratory, Ruchill Hospital, Glasgow, between 1953 and 1969 five had "cardiac" features and one had endocarditis. Necropsy in the endocarditis case showed multivalvular lesions, pulmonary oedema, and large aortic vegetations. Our patient had no evidence of pre-existing valvular disease and developed signs of endocarditis and aortic incompetence while the psittacosis/LGVCVF titres were rising. We cannot explain the rather late rise in the psittacosis/LGVCVF titres. He failed to respond to large doses of tetracycline. Various other antibiotics were used without success. We are not aware of any reports of response to rifampicin, which is a bactericidal antibiotic alleged to be active at high concentration against some viruses.

The reported incidence of psittacosis is rising. In many of the pathological cases there is no relevant "bird-contact his" history. Fowls are a major reservoir of the infecting agent. Valve damage is found in birds dying from the infection. It is rarely described in man and in the context of fatal overwhelming infections. We suggest that infection in relation to contact with birds be included in the differential diagnosis of "blood-culture-negative" endocarditis.

Problems with intravenous chloromethiazole (Heminevrin) in status epilepticus

Intravenous diazepam is a drug of first choice for status epilepticus. Administration is by either continuous intravenous infusion or bolus injection, and we have also used the rectal route successfully. For very resistant seizures, however, we have used instead a continuous intravenous infusion of chloromethiazole edisylate (Heminevrin) 8 g/l in an aqueous solution of 4%, dextrose, achieving control of fits in both major and minor status epilepticus with doses of 5-10 mg/kg/h by titrating fits with appropriate infusion rates. Respiratory depression and hypotension have not been serious hazards. Nevertheless, we report here several problems that we have encountered with this drug.

Recorded complications

Reaction with plastics—Intravenous chloromethiazole reacts with the plastic of giving sets and may soften the burette sets to four to six hours' contact. No plastic interaction occurs with continuous intravenous infusion set (Heminevrin) 8 g/l in an aqueous solution of 4%, dextrose, achieving control of minor and status epilepticus with doses of 5-10 mg/kg/h by intravenous infusion rates. Respiratory depression and hypotension have not been serious hazards. Nevertheless, we report here several problems that we have encountered with this drug.

Effects of Chloromethiazole on five children with seizures resistant to diazepam

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We thank Dr A D Evans, consultant virologist, University Hospital of Wales, Cardiff, and his department for help in investigating this patient.


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Comment

Continuous intravenous chlormethiazole is an effective anti-convulsant for status epilepticus, but because of the complications mentioned should be used carefully, preferably for short periods only.

We thank the scientific and technical branch of the DHSS and the Government Chemist for their interest and chemical analysis, and the manufacturers of Heminervin (Astra Chemicals), for their analysis and pyrogen tests.


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Late displacement of central venous catheter resulting in vascular obstruction

The introduction of large-calibre plastic catheters into the central veins of patients has facilitated total parenteral nutrition and drug administration. These two procedures may be of critical importance in managing very sick people with bacterial endocarditis, acute leukaemia during initial induction chemotherapy, or after bone marrow transplantation. In each of these the peripheral vessels are unsatisfactory routes for intravenous infusion, since large volumes need to be given and also the high viscosity or hyperosmolality of some of the materials may result in thrombophlebitis and thus render small veins especially prone to occlusion. A number of techniques are available for achieving access to the high-flow veins within the thorax. To date these procedures have been relatively free from serious problems, but the catheters may become dislocated from the original desired site and then follow an abnormal route within the thoracic vessels, leading to complications.1 In most reported cases the unsatisfactory results arise during the introduction process and we are not aware of late catheter migration, particularly against venous flow, with resulting vascular occlusion.

Case report

A 49-year-old woman was hospitalised for cytotoxic chemotherapy for stage IV lymphocytic lymphoma. In view of her poor physical state a 10-gauge Silastic catheter was introduced via the left subclavian line, using a previously described technique.2 Postoperative chest x-ray examination confirmed that the end of the catheter was lying in the superior vena cava with its end 1 cm above the right atrium. For the next three weeks the patient received her intravenous drugs, hyperalimentation, and daily fluid requirements by this route without any problems. During the same period blood samples were collected from the catheter, since measurements in these correlate well with those in samples simultaneously collected from peripheral vessels.3 On day 23 after introduction of the catheter the patient’s left hand and arm started to swell and this progressed slowly for two days. Careful serial examination showed increasing tenderness and swelling of the left arm, supraclavicular space, and axilla. Radiological studies showed that the radio-opaque catheter had looped up inside the superior vena cava and wedged into the axillary vein, obstructing blood flow. The catheter was simply repositioned under x-ray control, and within two days all symptoms had disappeared. No late complications have occurred.

Comment

A survey of central venous catheters used in our hospital over a two-year period confirmed the finding of Fischer and his colleagues1 that initial misplacement could occur and that intrathoracic complications may then arise. Nevertheless, once securely in position none of the catheters that we studied became displaced. Our case was the first example of late dislocation we encountered over the same two-year period, when catheters were routinely employed for hyperalimentation, induction of cytotoxic chemotherapy in patients with leukaemia, and long-term support after bone marrow transplantation. The final resting place of the catheter is all the more surprising in view of the retrograde course followed by the large diameter and relative inflexibility of the plastic tubes. This complication should be recognised as soon as it occurs since the hypertonic and irritating solutions infused may lead to local endothelial damage and thrombus formation. One way to check the patency is by regular flushing, while direct determination of venous pressure is an easy method of monitoring freedom from obstruction. While radiological control is essential when introducing the catheter its subsequent use is of lower priority in the complex management of a critically ill patient. It is therefore not surprising that the symptoms of mechanical obstruction may not be immediately evident. Since delay in diagnosis is potentially serious we suggest that these patients should be regularly examined, with particular emphasis on catheter efficiency, and that x-ray studies should be undertaken whenever anxiety arises. A dislocation may then be promptly corrected and x-ray control will thus avoid venous complications.


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Hydralazine-induced cutaneous vasculitis

The commonest features of the hydralazine-associated lupus syndrome are arthralgia and malaise. Brythematosus rashes may occur but vasculitis has not been reported. We have recently seen two patients with ulcerating cutaneous vasculitis associated with hydralazine treatment.

Case reports

(1) A 63-year-old woman had been treated for hypertension and stable renal impairment for 15 years. Hydralazine was started in September 1977 and the daily dosage increased to 250 mg in addition to diuretic and beta-blocking drugs and methylprip. In July 1978 her serum antinuclear antibody (ANA) was positive at a titre of 1/40 and her erythrocyte sedimentation rate (ESR) 33 mm in 1 h. In July 1979 the ANA titre was 1/160 and the ESR 70 mm in 1 h. In August 1979 she was admitted to hospital with a 10 × 5 cm ulcer on the lateral aspect of her left leg, which had developed over two weeks. She gave a six-month history of arthralgia, myalgia, and weight loss of 3 kg. Two days after admission she developed multiple vasculitic lesions over the buttocks, thighs, and arms. Hydralazine was withdrawn and the lesions began to resolve. When one of them ulcerated, however, prednison was given for five days, with rapid disappearance of the lesions and healing of the ulcer. Results of investigations included ESR 90 mm in 1 h; creatinine clearance 25 ml/min; no proteinuria; haemoglobin 11·4 g/dl; white cell count (WBC) 4·1 × 106/l (4100/mm3), normal differential and platelets; immunoglobulins IgG 13·0 g/l, IgA 1·75 g/l, IgM 2·60 g/l (normal IgM 0·5-1·8 g/l). IgG and IgM ANA titres were 1/2560, DNA binding 20%. No other autoantibodies were found, complement screen was normal, Cq binding negative, and cryoglobulins were present in trace only. Sulphonamidase, phenotyping showed her to be a slow acetylator and her total intake of hydralazine was estimated to be 120 g. Histology of a lesion from her thigh showed allergic vasculitis with fibrinoid necrosis of small vessels. Immunofluorescence microscopy showed deposits of IgM, fibrin, and C3 in small blood vessels.

(2) A 61-year-old woman with essential hypertension had received hydralazine, propranolol, and cyclopenthiazide K for two years. During this time her maximum daily dose of hydralazine was 100 mg and her estimated total intake 95 g. In 1979 she presented with a six-month history of pain and stiffness in her hands, wrists, elbows, shoulders, and knees...