Acute renal failure from the use of acetazolamide (Diamox)

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Summary
The clinical histories of two patients are presented, who both developed haemorrhagic anuria after a short course of therapy with the carbonic anhydrase inhibitor, acetazolamide (Diamox). The clinical findings, in particular the radiological changes, are illustrated in this potentially fatal condition. The therapeutic success of immediate relief of the ureteric obstruction is emphasized and it is hoped that this report will act as a reminder of the importance of this agent as a cause of anuria.

Introduction
The carbonic anhydrase inhibitor, acetazolamide (Diamox), is widely used to reduce intra-ocular pressure in acute glaucoma and after cataract extraction. In addition, it has been recently advocated as a prophylactic in acute mountain sickness (Hackett, Drummond and Levine, 1976) and has occasionally been used in the treatment of refractory epilepsy.

Acute haemorrhagic anuria occurs uncommonly as a complication of this therapy but has resulted in fatalities (Leopold, 1968; Howlett, 1975). The case histories are now reported of two such patients exhibiting the characteristic radiographic appearances of ureteric obstruction, who responded completely to relief of this obstruction.

Case no. 1
A 62-year-old man of previous good health developed backache and vomiting four days after a successful unilateral cataract extraction. He had received acetazolamide, 250 mg 4 times/day, post-operatively. Within two days he was delirious and haematuria was noted. The serum urea was 72 mmol/l; bicarbonate 13 mmol/l; serum sodium 129 mmol/l and serum potassium 5.5 mmol/l.

Although his condition improved with intravenous saline (0.57 mmol/l), he became anuric. The high dose urogram revealed a dilated renal pelvis and calyces which contained radiolucent filling defects (Fig. 1), which suggested retained clots of blood; little contrast entered the ureters. The cystoscopy was normal, ureteric catheters enabled a diuresis to occur which lasted two days. The patient then made a full recovery with a normal intravenous urogram one month later.

Case no. 2
A 46-year-old man, epileptic, had required

Fig. 1. The high dose intravenous urogram of patient 1 showing the left kidney with a radiolucent filling defect in the renal pelvis.
ureters, dilated lateral acetazolamide, 0·5 g daily for three weeks, in addition to his usual anticonvulsant therapy of phenobarbitone and phenytoin to control his refractory epilepsy.

After a seven-day history of loin pain and haematuria, he presented with anuria. He was delirious and serum urea was 32 mmol/l; bicarbonate 18 mmol/l; sodium 132 mmol/l, and potassium 5 mmol/l.

The high-dose intravenous urogram revealed bilateral dilated calyces with little contrast in the ureters. The cystoscopy was normal, but it was impossible to pass ureteric catheters. A left-sided needle nephrostomy was performed to enable urinary drainage. Contrast injected through the catheter showed ureteric mucosal irregularity (Fig. 2). A diuresis via the nephrostomy catheter was followed by complete recovery, and with a normal intravenous urogram after one month.

Discussion

After a short course of acetazolamide both patients developed anuria which was preceded by the symptoms of back pain and haematuria. The radiographic appearances in both suggest that clot retention and/or mucosal swelling led to an obstructive anuria. This is supported by the complete recovery that both patients made after the diuresis, which followed the early relief of the obstruction, using ureteric catheters in one patient and a unilateral needle nephrostomy in the other.

A similar clinical picture is well described by Arneil (1958) in a case of acute sulphonamide crystalluria, indeed, acetazolamide is a heterocyclic sulphonamide which is excreted unchanged in the urine. Furthermore, with a pKa of 7.2, it is rendered particularly insoluble in acid urine. Acetazolamide-like crystals have also been found in the renal tubules in one fatal case of anuria following therapy (Bertino, Rodman and Myerson, 1957).

The authors would like to emphasize, therefore, the need to ensure a high fluid intake in patients on acetazolamide to reduce the chance of crystalluria, particularly in circumstances associated with an acidic urine and the need for the early relief of ureteric obstruction, if this complication occurs. Above all they remember that acetazolamide (Diamox) remains the cause of haemorrhagic anuria.

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References


FIG. 2. The antegrade urogram through the nephrostomy tube in patient 2, showing the right ureter with marked irregularity, suggesting mucosal swelling.