Lessons from practice

Enlafaxine is described as a safe and effective antidepressant that is chemically distinct from other antidepressants. It is a non-selective inhibitor of the reuptake of serotonin, norepinephrine and dopamine and has no anticholinergic activity in vitro. Because of their relative lack of anticholinergic effects, enlafaxine and similarly acting selective serotonin reuptake inhibitors (SSRIs) are preferred over tri- and tetracyclic antidepressants for patients who are at risk of angle closure. We report the case of a young patient taking enlafaxine who developed simultaneous bilateral acute angle closure secondary to supraciliary effusions.

**Clinical record**

A 35-year-old man presented to the emergency department complaining of right visual blurring and discomfort overnight. Over the previous 2 years, similar episodes had occurred, mostly at night or in dim lighting, affecting one eye at a time and resolving spontaneously each time. The patient was not using any type of medication at the time when the episodes began to occur. Although infrequent initially, these episodes had increased to two or three times a week after mirtazapine, a tetracyclic antidepressant, had been prescribed for depression and anxiety 5 months previously. Symptoms persisted despite treatment being changed to sertraline, a selective serotonin reuptake inhibitor (SSRI). Ten days before presentation, the sertraline was replaced by enlafaxine 75 mg once a day. Symptoms were occurring about every other day.

There was no other significant ocular or family history. Spectacle correction revealed compound hypermetropic astigmatism (right eye, +1.00 + 1.50 3 75°; left eye, +1.75 + 2.25 3 110°).

On presentation, the patient's visual acuity was 6/24, improving, with a pinhole before the eye, to 6/9 (right eye) and 6/6 (left eye). The right pupil was fixed and mid-dilated and the cornea was mildly oedematous. Both anterior chambers were axially shallow, with forward displacement of the iris–lens diaphragm (ie, the plane formed by the iris and the anterior surface of the lens at the pupil). Intraocular pressures were 69 mmHg (right eye) and 62 mmHg (left eye) (intraocular pressure > 21 mmHg is generally considered as being elevated). Gonioscopy revealed bilateral completely closed angles.

The patient was given intravenous mannitol 100 g over 40 minutes, oral acetazolamide 500 mg, topical timolol 0.5%, apraclonidine 1% and pilocarpine 2%. Intraocular pressures fell to 13 mmHg and 5 mmHg in the right and left eyes, respectively. Gonioscopically, the angles could now be opened with indentation of the cornea; however, the anterior chambers remained shallow.

Bilateral laser peripheral iridotomies were performed on the day of presentation (left eye) and on the next day (right eye) to eliminate pupil block. Post-iridotomy gonioscopy showed no change in the angle configuration or anterior chamber depth. The patient was sent home with instructions to take oral acetazolamide 250 mg three times a day, topical timolol 0.5% once a day, pilocarpine 2% four times a day, and brimonidine 2% twice a day. Venlafaxine was discontinued. The patient refused alternative antidepressant medication and requested referral to a psychiatrist or psychologist for counselling.

On Day 4 after the patient presented with acute angle closure, ultrasound biomicroscopy showed bilateral supraciliary effusions and anterior chamber shallowing (Box 1). Manifest refraction (ie, manual subjective measurement of refractive error) at this time revealed a myopic shift of about 3 dioptres in each eye.

Over several weeks, medications to lower intraocular pressure were withdrawn one by one. Follow-up ultrasound biomicroscopy showed gradual resolution of the supraciliary effusion, which was complete some 5 weeks after the acute attack. The anterior chambers deepened slightly and the patient's usual hypermetropic refraction returned. Gonioscopy at this time still showed easily occludable angles. Bilateral laser peripheral iridoplasties were performed to reduce the risk of future angle closure.

Venlafaxine is described as a safe and effective antidepressant that is chemically distinct from other antidepressants. It is a non-selective inhibitor of the reuptake of serotonin, norepinephrine and dopamine and has no anticholinergic activity in vitro. Because of their relative lack of anticholinergic effects, venlafaxine and similarly acting selective serotonin reuptake inhibitors (SSRIs) are preferred over tri- and tetracyclic antidepressants for patients who are at risk of angle closure. We report the case of a young patient taking venlafaxine who developed simultaneous bilateral acute angle closure secondary to supraciliary effusions.

Angle closure and/or acute transient myopia possibly caused by supraciliary effusion has been reported for many drugs, including sulfonamides, tetracycline and some diuretics.
Lessons from practice

- Use venlafaxine (and antidepressants in general) with caution in patients who are at risk of angle-closure glaucoma.
- Patients at risk are those with hypermetropic refraction (i.e., whose distance spectacles magnify objects) and those with symptoms of angle closure (intermittent blurring of vision associated with seeing coloured rings around lights, eye redness, or eye pain). Such symptoms should not be dismissed as “migrainous”.
- Patients at risk of angle closure should undergo ophthalmological screening, particularly gonioscopy, before starting antidepressant drugs.
- Because any patient could develop a supraciliary effusion in response to various drugs (especially antidepressants), it is prudent to include symptoms of angle closure when educating patients about possible side effects of these drugs. They should seek ophthalmological care if they experience symptoms of angle closure or a myopic shift in their vision.

Effusions causing secondary angle closure in patients taking topiramate and sulfonamides have been documented by ultrasound biomicroscopy by various authors. The postulated mechanisms by which supraciliary effusions produce angle-closure glaucoma and transient myopia are illustrated in Box 2.

There have been reports in the literature of angle closure or a myopic shift in their vision.

- Topiramate
- Mirtazapine
- Sertraline

Serotonin and serotonin receptors have been found in the human ciliary body, and serotonin, its agonists and antagonists do affect intraocular pressure. The supraciliary effusions documented here are evidence of the serotonergic mechanism similar to that of the cyclic antidepressants. The serotonergic effects of these drugs may also have a role in angle closure. Serotonin and serotonin receptors have been found in the human ciliary body, and serotonin, its agonists and antagonists do affect intraocular pressure. The supraciliary effusions documented here are evidence of the serotonergic effects of venlafaxine causing angle closure, although the precise cause for the effusions is unknown.

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Neurological sequelae of chronic profound hypocalcaemia

A 66-year-old man presented with a 12-month history of progressive gait disturbance with cerebellar ataxia and extrapyramidal features. Computed tomography of the head showed calcification of the basal ganglia and dentate nuclei of the cerebellum, and periventricular calcification (Box).

Biochemical testing showed serum level of calcium, 1.19 mmol/L (reference range [RR], 2.15–2.55 mmol/L); ionised calcium, 0.61 mmol/L (RR, 1.14–1.29 mmol/L); phosphate, 1.8 mmol/L (RR, 0.8–1.5 mmol/L); and parathyroid hormone, 0.9 pmol/L (RR, 1.5–8.0 pmol/L). The patient was diagnosed with hypocalcaemia caused by idiopathic hypoparathyroidism. He was treated with calcitriol (2 µg daily) and calcium carbonate (6.0 g daily). His gait showed some improvement, and serum calcium levels became normal over several weeks.

The pathophysiology of intracerebral calcification in hypoparathyroidism, which appears paradoxical, is unknown. It is usually asymptomatic. When neurological effects occur, they are thought to be caused by microvascular degeneration from massive perivascular calcium deposition in high-metabolic areas.

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Computed tomography of the head, showing cerebral calcification

A: Calcification of the basal ganglia. B: Calcification of the dentate nuclei of the cerebellum. C: Periventricular calcification.


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