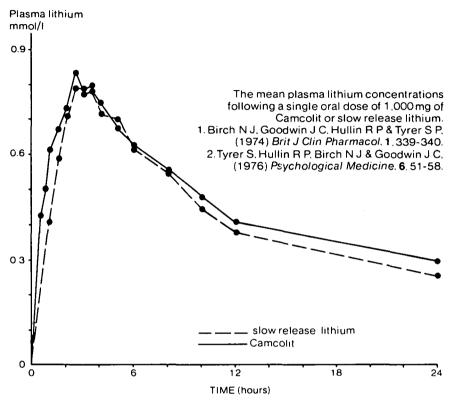
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Clomipramine (Anafranil) in the treatment of chronic intractable depression.

Paper read at the Fifth World Congress of Psychiatry, Mexico D.F. 1971.

"The difference between the proportion of patients in hospital who improved when treated with electroconvulsive therapy, conventional antidepressant drug therapy and intravenous infusion of clomipramine was statistically significant in favour of the last mentioned treatment. Patients on clomipramine as a group needed fewer treatments and returned to work more rapidly than did their counterparts having electroconvulsive therapy."

A new adjunct to the treatment and management of depression: intravenous infusion of clomipramine (Anafranil). S. Afr. med. J., 45, 168 (1971)

"72% (of 57 patients) showed a very good or good response and 96% made some improvement. This compares very favourably with the response of similar groups of severely depressed patients to E.C.T., and it is postulated that intravenous chlorimipramine can be offered as an alternative form of treatment."

"Oral group: 78 per cent showed a very good or good response and 96 per cent improved to some extent. This also compared favourably with the results obtained with other antidepressant drugs in similar groups of patients."

Parenteral and oral chlorimipramine treatment of depressive states. Brit. J. Psychiat., 122, 189 (1973)

Anafranil® in obsessional/phobic disorders

"Clomipramine has the two distinct properties of being an anti-depressive and an anti-obsessional drug."

Clomipramine (Anafranil) in the treatment of obsessional states: A psychiatrists view.
J. Int. Med. Res., 3 (Supp. 1) 83 (1975)

"Obsessional illnesses have always been notorious for their resistance to treatment and phobic states, especially, when they are diffuse and polysymptomatic, do not respond always to deconditioning or flooding techniques A treatment which offers brevity with a 70% chance of disappearance or considerable reduction in symptoms is worth offering to patients as a first choice of therapies."

Clomipramine (Anafranil) in the treatment of obsessional illnesses and phobic anxiety states. J. Int. Med. Res., 1, 403 (1973)

"It is our view that clomipramine not only gives good results in severe and moderate depressive states, but it is emerging as the treatment of choice in obsessive compulsive disorders and phobic states."

Letter, Treating phobias. World Medicine, 7, 11: 15 (1972)

"The mode of action of Anafranil is unknown but without doubt it appears to exert a beneficial effect on neurotic responses in general and phobic and obsessional states in particular."

An investigation into the use of Anafranil in phobic and obsessional disorders. Scot. med. J., 20 (Supp), 61 (1975)

Indications

ANGLEATIONS
Tryptizol' is recommended in the treatment of depression including that accompanied by anxiety.
Tryptizol' may be used successfully in depression that is a manifestation of psychosis or neurosis, whether endogenous or reactive in nature. Endogenous depression is more likely to be alleviated than are other depressive states.
Tryptizol' has an anxiety-reducing and sedative component to its action which is particularly helpful in alleviating anxiety or agitation that often accompanies depression.

It has been used with benefit in depressions of long or short duration, and with a wide range of intensity. As with other psychotherapeutic agents, all patients do not respond to the same degree. Some patients respond promptly, while others may require up to 30 days to obtain benefit. Lack of response may occur occasionally.

Depressive reactions and associated anxiety accompanying chronic illness may be relieved by 'Tryptizol'.

Dosage and administration

Start with one 75 mg capsule at bedtime and increase, if necessary, to two capsules at bedtime or one in the morning and one at night. The 75 mg capsule may also be used as maintenance therapy.

The anxiety-reducing and sedative effect is usually rapidly apparent. The anxiety-reducing and sedative effect is usually rapidly apparent. The antidepressant activity may be seen within 3 or 4 days or may take up to 30 days to develop adequately.

Contra-indications

Hypersensitivity to amiltriptyline, concomitant use with a monoamine oxidase inhibitor (see "Precautions"), during the acute recovery phase following myocardial infarction. See "Usage in Pregnancy" under Precautions".

Precautions

Combined use of antidepressants having varying modes of activity requires thorough knowledge of pharmacology of all agents. Allow minimum of 14 days to elapse following discontinuation of MAO inhibitors and introduction of Tryptizol'. (Hyperpyretic crises, severe convulsions and deaths have occurred when tricyclic antidepressants and MAOI drugs were given simultaneously.) Introduce 'Tryptizol' cautiously and gradually increase until optimum response is

achieved.

When amitriptyline is used to treat the depressive component of schizophrenia, psychotic symptoms may be aggravated. If used in manic-depressive psychosis a shift towards the manic phase may occur. Paranoid delusions, with or without associated hostility may be exaggerated. (In such circumstances either reduce dose of amitriptyline or add major tranquillising drug.)

The possibility of suicide in depressed patients remains during treatment and until significant remission occurs. This type of patients should not have easy access to large quantities of the drug.

Concurrent administration of amitriptyline and electroconvulsive therapy may increase the hazards of therapy and therefore should be limited to patients for whom it is essential.

Liee with caution in patients with history of seizures, urinary retention, or

Whom It is essential.

Use with caution in patients with history of seizures, urinary retention, or increased intra-ocular pressure. With narrow-angle glaucoma, even average doses may precipitate an attack. Closely supervise treatment in hyperthyroid patients, those receiving thyroid medication, anticholinergic or sympathomimetic drugs, including adrenaline combined with local anaesthetics (careful adjustments of dosage are required) and those who receive large doses of ethchlorvynol concurrently.

It may enhance the response to alcohol and the effects of barbiturates or other CNS depressants. Alertness may be impaired in some patients (activities made hazardous by this diminished alertness should be avoided).

Patients with cardiovascular disorders should be watched closely. Tricyclic anti-depressant drugs have been reported to produce arrhythmias, sinus tachycardia, and prolongation of the conduction time, myocardial infarction and stroke. Amitriptyline may block the antihypertensive action of guanethidine or similarly acting compounds.

Discontinue amitriptyline several days before elective surgery, if possible. Both elevation and lowering of blood sugar levels have been reported. Not recommended for depressed patients under 12 years of age.

Safe use during pregnancy has not been established. Weigh benefits against possible hazards to mother and child when administered to pregnant or possibly

pregnant women and to nursing mothers. Side effects

Note: Included in the listing which follows are a few adverse reactions which have not been reported with this specific drug. However, pharmacological similarities among the tricyclic antidepressant drugs require that each of the reactions be considered when amitriptyline is administered.

Cardiovascular: Hypotension, hypertension, tachycardia, palpitations, myocardial infarction, arrhythmias, heart block, stroke.

CNOS and neuromuscular: Condusional states, disturbed concentration, disorientation, delusions, hallucinations, excitement, anxiety, restlessness, insomnia, nightmares, numbness, tingling, and paraesthesiae of the extremities, peripheral neuropathy, incoordination, ataxia, tremors, scizures, alteration of EEG patterns, extrapyramidal symptoms, tinnitus.

Anticholinergic: Dry mouth, blurred vision, disturbance of accommodation, constipation, paralytic ileus, urinary retention, dilatation of urinary tract. Allergic: Skin rash, urticaria, photosensitisation, oedema of face and tongue. Haematological: Bone-marrow depression including agranulocytosis,

Gastro-Intestinal: Nausea, epigastric distress, vomiting, anorexia, stomatitis, peculiar taste, diarrhoea, parotid swelling, black tongue, rarely hepatitis (including altered liver function and jaundice).

Endocrine: Testicular swelling and gynaccomastia in the male, breast enlarge-ment and galactorrhoea in the female, increased or decreased libido, elevation or lowering of blood sugar levels.

Other: Dizziness, weakness, fatigue, headache, weight loss or gain, increased perspiration, urinary frequency, mydriasis, drowsiness, alopecia. Wthdrawal symptoms: Abrupt cessation of treatment after prolonged admini-stration may produce nausea, headache, and malaise. These are not indicative of addiction.

Presentation

Tryptizol' 75, slow-release capsules marked 'MSD 649' containing 75 mg ami-triptyline hydrochloride BP as a pelleted formulation in bottles of 100. Product licence number: 0025/01120

Detailed information is available to physicians on request. @ denotes registered





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