

CLONOTRIL PLUS

For the use of a Registered Medical Practitioner or Hospital or a Laboratory only.

Abbreviated Prescribing information for CLONOTRIL PLUS (Escitalopram Oxalate and Clonazepam Tablets IP)

[Please refer the complete prescribing information available at www.torrentpharma.com]

PHARMACOLOGICAL PROPERTIES:

Mechanism of action: Clonazepam exhibits pharmacological properties, which are common to benzodiazepines and include anticonvulsive, sedative, muscle relaxing and anxiolytic effects. Animal data and electroencephalographic investigations in man have shown that clonazepam rapidly suppresses many types of paroxysmal activity including the spike and wave discharge in absence seizures (petit mal), slow spike wave, generalised spike wave, spikes with temporal or other locations as well as irregular spikes and waves. Escitalopram is a selective inhibitor of serotonin (5-HT) re-uptake with high affinity for the primary binding site. It also binds to an allosteric site on the serotonin transporter, with a 1000 fold lower affinity.

DOSAGE AND ADMINISTRATION: As directed by physician.

CONTRAINDICATION: Hypersensitivity to the active substance (Clonazepam and Escitalopram) or any of the excipients. History of sensitivity to benzodiazepines. Clinical or biochemical evidence of significant liver disease. Acute narrow angle glaucoma (it may be used in patients with open angle glaucoma who are receiving appropriate therapy). Concomitant treatment with non-selective, irreversible monoamine oxidase inhibitors (MAO-inhibitors) is contraindicated due to the risk of serotonin syndrome with agitation, tremor, hyperthermia etc. The combination of CLONOTRIL PLUS with reversible MAO-A inhibitors (e.g. moclobemide) or the reversible non-selective MAO-inhibitor linezolid is contraindicated due to the risk of onset of a serotonin syndrome. CLONOTRIL PLUS is contraindicated in patients with known Q_Ts that are known to prolong the QT interval, Acute pulmonary insufficiency; severe respiratory T interval prolongation or congenital long QT syndrome, together with medicinal product insufficiency, sleep apnoea syndrome, myasthenia gravis, severe hepatic insufficiency. CLONOTRIL PLUS must not be used in patients in a coma, or in patients known to be abusing pharmaceuticals, drugs or alcohol.

WARNINGS & PRECAUTIONS: Warning: Clinical Worsening and Suicide Risk, Risks from Concomitant Use with Opioids may result in in profound sedation, respiratory depression, coma, and death. Precautions: Worsening of Seizures, Laboratory Testing During Long-Term Therapy, Psychiatric and Paradoxical Reactions, Risks of Abrupt Withdrawal, Hypersalivation, Respiratory Depression, Porphyria. Paradoxical anxiety, Seizures, Mania, Diabetes, Akathisia/psychomotor restlessness, Hyponatraemia, Haemorrhage, ECT (electroconvulsive therapy), Serotonin syndrome, St. John's wort, Discontinuation symptoms seen when stopping treatment, Coronary heart disease, QT interval prolongation, Angle-Closure Glaucoma.

DRUG INTERACTION: Clonazepam: with opioids: increases the risk of sedation, respiratory depression, coma and death because of additive CNS depressant effect, Alcohol: modify the effects of the drug, Antiepileptic drugs: sedation and apathy, and toxicity, Escitalopram: may increase side effects such as dizziness, drowsiness, confusion, & difficulty concentrating. Escitalopram: *Pharmacodynamic interactions:* Irreversible non-selective MAOIs, Reversible, selective MAO-A inhibitor (moclobemide), Reversible, non-selective MAO-inhibitor (linezolid), Irreversible, selective MAO-B inhibitor (selegiline), QT interval prolongation. Combinations requiring precautions for use: Serotonergic medicinal products,

Medicinal products lowering the seizure threshold, Lithium, tryptophan, St. John's wort, Haemorrhage, Medicinal products inducing hypokalaemia/hypomagnesaemia, Etizolam. *Pharmacokinetic interactions: Influence of other medicinal products on the pharmacokinetics of escitalopram:* Omeprazole, cimetidine, esomeprazole, fluvoxamine, lansoprazole, ticlopidine. *Effect of escitalopram on the pharmacokinetics of other medicinal products:* Flecainide, propafenone and metoprolol, desipramine.

ADVERSE REACTIONS: Clonazepam: Blood dyscrasias, anaphylaxis and angioedema, Anterograde amnesia, Depression, loss of libido, impotence, aggressiveness, excitability, nervousness, hostility, anxiety, sleep disturbances, nightmares, vivid dreams, irritability, agitation, psychotic disorders, Dizziness, light-headedness, somnolence, fatigue, co-ordination disturbances, poor concentration, restlessness, confusion and disorientation, headache, Dysarthria and ataxia, Cardiac failure including cardiac arrest, nausea, gastrointestinal symptoms, urticaria, pruritus, transient hair loss, pigmentation changes, Muscle weakness, occasional muscular hypotonia, urinary incontinence, tremor, sweating, headaches, muscle pain, extreme anxiety, tension, restlessness, confusion, irritability and epileptic seizures.

Escitalopram: Thrombocytopenia, Anaphylactic reaction, Decreased appetite, increased appetite, weight increased/decreased, Hyponatraemia, anorexia Anxiety, restlessness, abnormal dreams, libido decreased, Female: anorgasmia, Bruxism, agitation, nervousness, panic attack, confusional state, Aggression, depersonalisation, hallucination, Mania, suicidal ideation, suicidal behaviour, Headache, Insomnia, somnolence, dizziness, paraesthesia, tremor, Taste disturbance, sleep disorder, syncope, Serotonin syndrome, Dyskinesia, movement disorder, convulsion, psychomotor restlessness/akathisia, Mydriasis, visual disturbance, Tinnitus, Tachycardia, Bradycardia, Electrocardiogram QT prolonged Ventricular arrhythmia including torsade de pointes, Orthostatic hypotension, Sinusitis, yawning, Epistaxis, Nausea, Diarrhoea, constipation, vomiting, dry mouth, Gastrointestinal haemorrhages (including rectal haemorrhage), Hepatitis, liver function test abnormal, Sweating increased, Urticaria, alopecia, rash, pruritus, Ecchymosis, angioedemas, Arthralgia, myalgia, Urinary retention, Male: ejaculation disorder, impotence, Female: metrorrhagia, menorrhagia, Galactorrhoea, Male: priapism, Fatigue, pyrexia, Oedema.

Manufactured BY:

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OR

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IN/CLONOTRIL PLUS 10, 0.5 mg/Mar-21/04/ABPI

(Additional information is available on request)