

C-PRAM S PLUS

For the use of a Registered Medical Practitioner or a Hospital or a Laboratory Only

WARNINGS: SUICIDALITY AND ANTIDEPRESSANT DRUGS

Abbreviated Prescribing information for C-PRAM S PLUS (Escitalopram and Clonazepam Tablets I.P.)
[Please refer the complete prescribing information available at www.torrentpharma.com]

PHARMACOLOGICAL PROPERTIES:

Mechanism of Action: *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. Escitalopram has no or low affinity for a number of receptors including 5-HT_{1A}, 5-HT₂, DA D₁ and D₂ receptors, α ₁-, α ₂-, β -adrenoceptors, histamine H₁, muscarine cholinergic, benzodiazepine, and opioid receptors. The inhibition of 5-HT re-uptake is the only likely mechanism of action explaining the pharmacological and clinical effects of escitalopram. *Clonazepam*, is believed to act via its ability to enhance the activity of gamma aminobutyric acid (GABA), the major inhibitory neurotransmitter in the central nervous system.

INDICATIONS: It is indicated for treatment of comorbid depression and anxiety disorder.

DOSAGE AND ADMINISTRATION: As directed by the physician.

CONTRAINDICATION: Hypersensitivity to the active substance (Escitalopram and Clonazepam). History of sensitivity to benzodiazepines. Hypersensitivity to any of the excipients. 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 Escitalopram and Clonazepam tablets with reversible MAO-an inhibitors (e.g. moclobemide) or the reversible non-selective MAO-inhibitor linezolid is contraindicated due to the risk of onset of a serotonin syndrome. Escitalopram and Clonazepam tablets is contraindicated in patients with known QT interval prolongation or congenital long QT syndrome. Escitalopram and Clonazepam tablets is contraindicated together with medicinal products that are known to prolong the QT interval. Acute pulmonary insufficiency; severe respiratory insufficiency, sleep apnoea syndrome, myasthenia gravis, severe hepatic insufficiency. Escitalopram and Clonazepam tablets must not be used in patients in a coma, or in patients known to be abusing pharmaceuticals, drugs or alcohol.

WARNINGS & PRECAUTIONS: Some patients with manic-depressive illness may enter into a manic phase. This is characterized by unusual and rapidly changing ideas, inappropriate happiness and excessive physical activity. If you experience this, contact your doctor. Symptoms such as restlessness or difficulty to in sitting or standing still can also occur during the first weeks of the treatment. Tell your doctor immediately if you experience these symptoms. Thoughts of suicide and worsening of your depression or anxiety disorder If you are depressed and/or have anxiety disorders you can sometimes have thoughts of harming or killing yourself. These may be increased when first starting antidepressants, since these medicines all take time to work, usually about two weeks but sometimes longer. You may be more likely to think like this: If you have previously had thoughts about killing or harming yourself. If you are a young adult. Information from clinical trials has shown an increased risk of suicidal behaviour in adults aged less than 25 years with psychiatric conditions who were treated with an antidepressant. If you have thoughts of harming or killing yourself at any time, contact your doctor or go to a hospital straight away. You may find it helpful to tell a relative or close friend that you are depressed or have an anxiety disorder, and ask them to read this leaflet. You might ask them to tell you if they think your depression or anxiety is getting worse, or if they are worried about changes in your behaviour.

DRUG INTERACTIONS: Tell your doctor if you are taking: Non-selective monoamine oxidase inhibitors (MAOIs)", containing phenelzine, isoniazid, isocarboxazid, nialamide, and tranylcypromine as active ingredients. "Irreversible MAO-B inhibitors", containing selegiline. These increase the risk of side effects. The antibiotic linezolid. Lithium and tryptophan. Imipramine and desipramine . Sumatriptan and similar medicines and tramadol . Cimetidine and omeprazole , fluvoxamine and ticlopidine . These may cause increased blood levels of escitalopram. Acetylsalicylic acid and non-steroidal anti-inflammatory drugs. These may increase bleeding-tendency. Warfarin, dipyridamole, and phenprocoumon. Mefloquine , bupropion and tramadol due to a possible risk of a lowered threshold for seizures. Neuroleptics and antidepressants due to a possible risk of a lowered threshold for seizures. Flecainide, propafenone, and metoprolol clomipramine, and nortriptyline and risperidone, thioridazine, and haloperidol . The dosage of C-PRAM S PLUS may need to be adjusted. Medicines that decrease blood levels of potassium or magnesium, as these conditions increase the risk of life-threatening heart rhythm disorders. Do not take C-PRAM S PLUS if you take medicines for heart rhythm problems or medicines that may affect the heart's rhythm, such as Class IA and III antiarrhythmics, antipsychotics, tricyclic antidepressants, certain antimicrobial agents, certain antihistamines. If you have any further questions about this, you should speak to your doctor. Do not take C-PRAM S PLUS if you take medicines for heart rhythm problems or medicines that may affect the heart's rhythm, such as Class IA and III antiarrhythmics, antipsychotics, tricyclic antidepressants, certain antimicrobial agents, certain antihistamines. If you have any further questions about this you should speak to your doctor.

ADVERSE REACTIONS: *Escitalopram:* Headache, Nausea, Decreased appetite, Anxiety, Insomnia, Sinusitis, Diarrhoea, Sweating increased, Arthralgia, Ejaculation disorder, Fatigue, Weight decreased, Bruxism, Taste disturbance, Mydriasis, Tinnitus, Tachycardia, Epistaxis, Gastrointestinal haemorrhages, Urticaria, metrorrhagia, Oedema, Thrombocytopenia, Inappropriate ADH secretion, Hyponatraemia, Mania, Dyskinesia, Electrocardiogram QT prolonged, Orthostatic hypotension, Hepatitis, Ecchymosis, Urinary retention, Postpartum haemorrhage. *Clonazepam:* Blood dyscrasias, Anaphylaxis and Angioedema, Incomplete precocious puberty, Anterograde amnesia, Dizziness, Double vision, Cardiac failure, Respiratory depression, Nausea, Abnormal liver function tests, Urticaria, Muscle weakness, Urinary incontinence, Erectile dysfunction, Physical dependence develop, Falls and fractures, Decreased platelet.

MARKETED BY:



TORRENT PHARMACEUTICALS LTD.

IN/C-PRAM S PLUS 10mg, 0.5mg/0.25mg /Mar-22/02/ABPI

(Additional information is available on request)