BEPSAR *Plus* Tablets

(Losartan Potassium and Hydrochlorothiazide Tablet USP)

COMPOSITION:
Each film coated tablet contains:
Losartan Potassium USP ..... 50mg.
Hydrochlorothiazide USP … 12.5mg.
USP Specs.

DESCRIPTION: Losartan and its principal active metabolite act at the final stage of renin angiotesin pathway. They inhibit the binding of angiotensin II with angiotensin type I (AT_1 receptor), thus blocking the vasoconstrictive and aldosterone secreting effects of the system. Both Losartan and its principal active metabolite have an affinity for the AT_1 receptor about 1000 fold greater than for the AT_2 receptor. Hydrochlorothiazide is a thiazide diuretic. Thiazides affect the renal tubular mechanisms of electrolyte reabsorption directly and indirectly. This action reduces plasma volume, with consequent results in an increase in plasma renin activity, aldosterone secretion, in urinary potassium loss, and decrease in serum potassium. The mechanism of the antihypertensive effect of thiazides is unknown.

PHARMACOKINETICS: Losartan Potassium: Following oral administration, Losartan is well absorbed and undergoes substantial first pass metabolism by cytochrome P450, about 14% of an orally administered dose of Losartan is converted to the active carboxylic acid metabolite that is responsible for most of its pharmacodynamic properties. Plasma clearance of Losartan and its active metabolite is about 600mL/min and 50mL/min, respectively. Following oral administration, plasma concentrations of losartan and its active metabolite decline polyexponentially with a terminal half-life of about 2 hours and 6-9 hours, respectively. There is no pharmacokinetic interaction between Losartan and Hydrochlorothiazide. **Hydrochlorothiazide:** It is rapidly eliminated by kidneys after oral administration without being metabolized. When plasma levels have been followed for at least 24 hours, the plasma half-life has been observed to vary between 5.6 and 14.8 hours. At least 61% of the oral dose is eliminated unchanged within 24 hours.

INDICATIONS: Bepsar Plus Tablet is indicated for the treatment of essential hypertension, for patients in whom combination therapy is appropriate.

CONTRA-INDICATIONS: Hypersensitivity to losartan, sulphonamide-derived substances (as hydrochlorothiazide) or to any of the excipients of the product. Therapy resistant hypokalemia or hypercalcemia.
Severe hepatic impairment; cholestasis and biliary obstructive disorders
Refractory hypnonatremia
Symptomatic hyperuricemia/ GOUT
Pregnancy
Severe renal impairement
Anuria
The concomitant use of Losartan potassium and Hydrochlorthiazide tablet with aliskiren-containing products is contra-indicated in patients with diabetes mellitus or renal impairment.

USUAL DOSAGE: To minimize dose-independent side effects, it is usually appropriate to begin combination therapy only after a patient has not achieved the desired effect with monotherapy. Dosage may require modifications to adjust for individual sensitivities and associated medical conditions. The usual starting and maintenance dose of **Bepsar Plus** is 1 tablet once daily, which may be increased to 2 tablets once daily for patients who do not respond adequately.

SIDE EFFECTS: Bepsar Plus may cause following side effects, Dizziness, runny nose, sore throat, and back pain. The following symptoms are uncommon, but if you experience any of them, consult your doctor immediately, swelling of the face, throat, tongue, lips, eyes, hands, feet, ankles, or lower legs; hoarseness, difficulty breathing or swallowing, hives, dry mouth, thirst, weakness, drowsiness, restlessness, confusion, seizures, muscle pains or cramps, infrequent urination, upset stomach, vomiting, fainting, rapid pounding, or irregular heartbeat.

SPECIAL WARNINGS AND PRECAUTIONS:
Losartan: Patients with a history of angioedema should be closely monitored. The concomitant use of potassium-sparing diuretics, potassium supplements and potassium containing salt substitutes with Losartan potassium and Hydrochlorthiazide tablet. Losartan potassium and Hydrochlorthiazide tablet is conta-indicated in patients with severe hepatic impairment. Losartan should be used with caution in patients with bilateral renal artery stenosis or stenosis of the artery to a solitary kidney.

Hydrochlorothiazide: As with all antihypertensive therapy, symptomatic hypotension may occur in some patients. Thiazide therapy may impair glucose tolerance.

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Dosage adjustment of antidiabetic agents, including insulin, may be required. Thiazides should be used with caution in patients with impaired hepatic function or progressive liver disease, as it may cause intrahepatic cholestasis, and since minor alterations of fluid and electrolyte balance may precipitate hepatic coma.
Pregnancy: Drugs that act directly on the renin-angiotensin system can cause fetal and neonatal morbidity and death when administered to pregnant women. When pregnancy is detected, **Bepsar Plus** should be discontinued as soon as possible.
Nursing Mothers: Because of the risk of potential harm to the newborn, avoid using this drug.
Adults Over 60 Years: Losartan Potassium should be used with caution in this age group, as older patients are more likely to experience the drug's adverse reactions.

Effects on ability to drive and use machines: When driving vehicles or operating machinery it must be borne in mind that dizziness or drowsiness may occasionally occur when taking antihypertensive therapy, in particular during initiation of treatment or when the dose is increased.

IMPORTANT INSTRUCTIONS: Volume depleted patients should be given low initial dose. It is not recommended for patients with severe renal impairment (Creatinine Clearance 30mL/min) or for patients with hepatic impairment. May be administered before or after meals. May be administered with other antihypertensive agents. Patient should not stop this medicine without the consent of the physician.

DRUG INTERACTIONS: No significant drug interactions were seen with the concomitant use of Losartan Potassium and digoxin, warfarin, cimetidine and phenobarbital, but concomitant use of Losartan Potassium and potassium sparing diuretics (e.g, spironolactone, triamterene & amiloride), potassium supplements or salt substitutes containing potassium may lead to increase in serum potassium. Thiazide diuretics may interact with following drugs: Narcotics, Alcohol, Corticosteroids, ACTH, Antidiabetic drugs, Skeletal muscle relaxants, NSAIDs & Lithium.

OVERDOSAGE: Symptoms of overdose may include: Dizziness, light headedness, fainting, rapid or pounding heartbeat. No specific information is available on the treatment of overdose. Treatment is symptomatic and supportive. Suggested measures include induction of emesis if ingestion is recent and correction of dehydration, electrolyte imbalance, hepatic coma and hypotension by established procedures.

INSTRUCTIONS: Store below 30°C. Protect from heat, light and moisture. Keep out of the reach of children.

PRESENTATION: Bepsar Plus Tablets are available in Alu Alu Blister pack of 10'sx2.

بیپ سار پلس گولیاں

(لوسارٹن پوٹاشیم اور ہائیڈروکلوروتھایازائڈ)

اجزاءترکیب: بیپ سارپلس کی ہر فلم کوٹڈ گولی میں بالترتیب ۵۰ اور ۱۲٫۵ ملی گرام لوسارٹن پوٹاشیم اور ہائیڈروکلوروتھایازائڈ موجود ہے۔

استعمال: بیپ سارپلس گولیاں بلند فشارِخون (ہائی بلڈ پریشر) کے ان مریضوں میں استعمال کی جاتی ہے جن میں دو دوائیاں زیادہ اثر کرتی ہیں۔

عمومی خوراک: مریض کی حساسیت اور طبی کیفیت کو مدنظر رکھتے ہوئے بیپ سارپلس کی خوراک میں ردوبدل کیا جاسکتا ہے۔ عام طور پر شروع میں اور بعد میں جاری رکھنے کے لئے بیپ سارپلس کی ایک گولی روزانہ استعمال کریں۔ وہ مریض جن کو ایک گولی روزانہ سے افاقہ نہ ہو ان کی خوراک دو گولیوں تک بڑھائی جاسکتی ہے۔

ممانعت: حاملہ خواتین، رضاعی مائیں، اینیوریا کے مریض اور ایسے مریض جو دوا کے اجزاء سے حساسیت رکھتے ہوں۔

احتیاط: ایسے مریض جن کے جگر کی کارکردگی نارمل نہ ہو ایسے مریض جن میں پانی کی کمی ہو، ۶۰ سال سے زائد عمر کے مریض اور گاڑی یا بھاری مشینری چلانے والوں کو دوا کی احتیاط کو ملحوظ خاطر رکھنا چاہیئے۔

ہدایات: ۳۰°ڈگری سینٹی گریڈ سے کم درجہ حرارت پر رکھیں۔ گرمی، روشنی اور نمی سے بچائیں۔ بچوں کی پہنچ سے دور رکھیں۔

پیشکش: بیپ سارپلس 2 x 10's گولیوں کے ایلو ایلو بلسٹر پیک میں دستیاب ہیں۔

PHARMA
Manufactured by:
NABIQASIM INDUSTRIES (PVT) LTD.
17/24, Korangi Industrial Area,
Karachi-Pakistan.
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