Patient leaflet in accordance with the Pharmacists' Regulations (Preparations) - 1986

This medicine is dispensed with a doctor's prescription only

Etopan XL 400 Extended-Release Tablets	**Etopan XL 600 Extended-Release Tablets**
Active ingredient and its quantity: Each extended-release tablet contains: etodolac 400 mg	**Active ingredient and its quantity:** Each extended-release tablet contains: etodolac 600 mg

Inactive ingredients and allergens in this medicine: see section 2 'Important information about some of this medicine's ingredients', and section 6 'Additional information'.

Read the entire leaflet carefully before you start using this medicine. This leaflet contains concise information about this medicine. If you have any further questions, consult your doctor or pharmacist.

This medicine has been prescribed to treat your illness. Do not pass it on to others. It may harm them, even if it seems to you that their illness is similar to yours.

1. What is this medicine intended for?

For the management of signs and symptoms of osteoarthritis and rheumatoid arthritis.

Therapeutic group:

Non-steroidal anti-inflammatory drugs (NSAIDs).

2. Before using this medicine

Do not use this medicine if:

- You are sensitive (allergic) to the active ingredient (etodolac) or to any of the other ingredients in this medicine (see section 6).
- You have **severe heart failure.**
- You have an ulcer (a peptic ulcer, a small lesion or hole in the stomach or duodenum) or bleeding in the stomach, or have had two or more episodes of ulcers, stomach bleeding or perforation.
- You have had an **allergic reaction or asthmatic reaction** (wheezing, itching or skin rash) after taking aspirin, Etopan or another NSAID.
- You have severe liver failure and kidney failure.
- **You are in the last trimester of pregnancy**.
- You have previously had bleeding from the stomach or bowel due to taking a NSAID.

Special warnings about using this medicine

Before using Etopan XL, tell your doctor if:

- You have a kidney, heart or liver disease, or a blood disorder, especially if you are also taking diuretics. The dose should be as low as possible and you should have periodic tests.
- You are already on long-term therapy with a medicine other than Etopan XL, since your doctor will want to perform regular tests, especially if you are elderly.
- You suffer from fluid retention (swelling of legs, ankles or feet).
- You suffer from high blood pressure or heart failure.
- You suffer or have ever suffered from asthma or breathing difficulties.
- You have heart problems, have previously had a stroke or think that you might be at risk of these conditions (for example, if you have high blood pressure, diabetes, high cholesterol or are a smoker).

Medicines such as Etopan XL may slightly increase the risk of heart attack or stroke. The risk is higher with high doses and prolonged treatment. Do not exceed the recommended dose or treatment duration.

Serious gastrointestinal side effects such as bleeding, ulceration and perforation may occur at any time during treatment, with or without warning symptoms, in patients treated with NSAIDs. **If** you experience any sign of gastrointestinal bleeding, **stop using** Etopan XL **immediately**.

If you have a blood or urine test, tell your doctor that you are taking Etopan XL, as the medicine may affect the results.

Children and adolescents

Etopan XL is not recommended for use in children.

Interactions with other medicines

If you are taking or have recently taken other medicines, including nonprescription medicines and dietary supplements, tell your doctor or pharmacist. Particularly if you are taking:

- anti-thrombotic medicines such as warfarin (blood thinner) and aspirin (prevention of platelet aggregation)
- ciclosporin or tacrolimus (used following organ transplantations)
- digoxin - for heart problems
- lithium - for mental illnesses
- methotrexate - to treat conditions such as psoriasis or rheumatoid arthritis
- corticosteroids (such as prednisolone)
- quinolone antibiotics e.g. ciprofloxacin
- medicines for treatment of hypertension
- mifepristone (for pregnancy termination) within the last 12 days
- other NSAIDs e.g. aspirin, ibuprofen, naproxen, diclofenac
- selective serotonin reuptake inhibitor (SSRI) antidepressants
- diuretics
- zidovudine (for treatment of HIV)

Using this medicine and food

Take the medicine with or after a meal.

Pregnancy and breastfeeding
If you are pregnant, breastfeeding, think you are pregnant or are trying to become pregnant, ask your doctor or pharmacist for advice before using this medicine.

Pregnancy
You should inform your doctor if you have problems becoming pregnant. NSAIDs may make it more difficult to become pregnant.

Do not use the medicine if you are in the last 3 months of pregnancy, as it could harm your foetus or cause problems at delivery.
The medicine may cause kidney and heart problems in the foetus. The medicine may affect the maternal or foetal tendency to bleed and cause labour to be later or longer than expected.
Do not take Etopan XL during the first six months of pregnancy, unless absolutely necessary **and advised by your doctor**. If you need treatment during this period or while you are trying to become pregnant, take the lowest dose for the shortest period possible.
Taking Etopan XL for more than a few days from 20 weeks of pregnancy onward can cause kidney problems in the foetus, which may lead to low levels of amniotic fluid (oligohydramnios) or narrowing of a blood vessel (ductus arteriosus) in the foetal heart. If you need treatment for longer than a few days, your doctor may recommend additional monitoring.

Breastfeeding
Do not use Etopan XL during breastfeeding. It is not known if this medicine passes into breast milk. It is not recommended for use during breastfeeding, unless considered essential by your doctor.

Driving and using machines
Etopan XL may cause dizziness, drowsiness or abnormal vision. If you experience these symptoms, do not drive or operate dangerous machines while taking Etopan XL.

Important information about some of this medicine's ingredients
This medicine contains lactose. If you have been told by your doctor that you have an intolerance to certain sugars, contact your doctor before starting treatment with this medicine.
This medicine contains less than 1 mmol (23 mg) sodium per dose, therefore it is considered essentially sodium-free.

3. How to use this medicine?

Always use this medicine according to your doctor's instructions. Check with your doctor or pharmacist if you are not sure about your dose or about how to take this medicine.
Only your doctor will determine your dose and how you should take this medicine. The recommended dosage is usually:

400 mg – one to up to 2 tablets twice a day. You may take up to 3 tables per day, but do not exceed this dosage.
600 mg – one tablet once or twice a day.

Do not exceed the overall dosage of 1200 mg per day!

If you are an elderly person, your doctor will make sure that you take the lowest dose for the shortest period of time due to the risk of severe side effects.

Do not exceed the recommended dose.
- Swallow the medicine whole with a glass of water.
- Do not chew, split or crush!

Use in children
Etopan XL is not recommended for use in children.

If you have accidentally taken a higher dose, or if a child has accidentally swallowed some medicine, immediately see a doctor or go to a hospital emergency room and bring the medicine package with you**.** Symptoms of overdose include headache, nausea and vomiting, abdominal pain, blood in faeces or black tarry stool. On rare occasions, diarrhoea, disorientation, excitation, coma, drowsiness, dizziness, ringing in the ears, fainting and seizures may occur. In cases of significant overdose, kidney failure and liver damage are also possible

If you forget to take the medicine at the scheduled time, do not take a double dose. Take the next dose at the usual time and consult your doctor.

Adhere to the treatment as recommended by your doctor.

Do not take medicines in the dark! Check the label and dose <u>every time</u> you take medicine. Wear glasses if you need them.

If you have any further questions about using this medicine, consult your doctor or pharmacist.

4. <u>Side effects</u>

As with any medicine, using Etopan XL may cause side effects in some users. Do not be alarmed by this list of side effects; you may not experience any of them.

The most serious side effects are allergic reactions, heart failure, stroke, kidney failure, liver failure, inflammation of the pancreas and aseptic meningitis. If you suffer from any of the effects described below, **stop taking Etopan XL and contact your doctor immediately.**

<u>Symptoms of allergic reactions:</u>
- Wheezing, difficulty breathing or shortness of breath,
- Swelling of the face, lips, mouth or tongue,
- Extensive rash, peeling or blistering of the skin, continuous itching.

<u>Symptoms of heart and blood circulation disorders:</u>
- Chest pain, high blood pressure, swelling of the ankles, palpitations, several types of anaemia or other blood disorders, unexpected bruising and bleeding.

<u>Symptoms of stomach and bowel (gastrointestinal) problems:</u>
- Blood in faeces,
- Black tarry stool,
- Vomiting blood or dark particles that look like coffee grounds.

Kidney failure symptoms:
- Difficulty or pain when passing urine, discolouration of urine or urinating more or less often than usual.

Symptoms of liver failure and inflammation of the pancreas:
- Jaundice (yellowing of the eyes or skin), abdominal pain, abnormal liver function test results.

Aseptic meningitis symptoms:
A serious rare condition known as aseptic meningitis may occur in patients with other autoimmune conditions such as systemic lupus erythematosus or mixed connective tissue disease.
The symptoms of aseptic meningitis are:
- A very high fever, vomiting, headaches, a blotchy rash that does not fade upon applying pressure to the skin (this may not develop), neck stiffness, photosensitivity, drowsiness and seizures.

Additional side effects
- Sensory disorders such as headaches, ringing or buzzing in the ears, dizziness, abnormal vision, hallucinations, tingling, pricking and burning of the skin (pins and needles), vertigo (a sensation that all the objects around you are moving or spinning).
- Gastrointestinal problems such as mouth ulcers, sore mouth, nausea, vomiting, abdominal discomfort, diarrhoea, constipation, flatulence (gas), heartburn, indigestion.
- Skin disorders such as swelling of tissues, itching of the skin, rash, redness.
- General disorders such as fever, drowsiness, tiredness, weakness, sleeplessness, shaking, nervousness, depression, confusion.

If you experience any side effect, if any side effect gets worse, or if you experience a side effect not mentioned in this leaflet, consult your doctor.

Reporting side effects
You can report side effects to the Ministry of Health by following the 'Reporting Side Effects of Drug Treatment' link on the Ministry of Health home page (www.health.gov.il), which opens an online form for reporting side effects, or you can also use this link: https://sideeffects.health.gov.il

5. How to store the medicine?

- Prevent poisoning! To prevent poisoning, keep this and all other medicines in a closed place, out of the reach and sight of children and/or infants. Do not induce vomiting unless explicitly instructed to do so by your doctor.
- Do not use the medicine after the expiry date (exp. date) which is stated on the package. The expiry date refers to the last day of that month.

Storage conditions
- Store below 25˚C.
- Do not throw away medicines via wastewater or household waste. Ask the pharmacist how to dispose of medicines you no longer use. These measures will help protect the environment.

6. Additional information

In addition to the active ingredient, this medicine also contains:
Microcrystalline cellulose, hypromellose, lactose anhydrous, povidone, magnesium stearate.

Coating: HPMC 2910, titanium dioxide, polydextrose, triacetin, macrogol, FD&C Red #40 Aluminum Lake (E129), FD&C Yellow #6 Aluminum Lake (E110).

In addition, each Etopan XL 600 tablet contains:
FD&C Blue #2 Aluminum Lake (E132), Black iron oxide (E172), Yellow iron oxide (E172).

What the medicine looks like and contents of the pack:

Etopan XL 400:
Pink film-coated, round, biconvex tablet, debossed with 'T400' on one side. The tablets are packed in blisters in a carton. Each pack contains 20 or 30 tablets.

Etopan XL 600:
Grey film-coated, elliptical, biconvex tablet, debossed with 'T600' on one side. The tablets are packed in blisters in a carton. Each pack contains 10, 12, 18, 20, or 24 tablets, or in a white plastic bottle with a childproof cap, with internal aluminum seal and cotton wool filling. Each pack contains 30, 100, or 1000 tablets.

Not all pack sizes may be marketed.

Manufacturer's and registration holder's name and address:
Taro Pharmaceutical Industries Ltd., 14 Hakitor St., Haifa Bay 2624761

Registration numbers of the medicine in the Ministry of Health National Drug Registry:
Etopan XL 400 130.03.30728
Etopan XL 600 122.59.30274

Revised in September 2023 according to Ministry of Health guidelines.

For further information about this medicinal product and for updated patient leaflets in Hebrew, Arabic and English, please scan the code:

Etopan XL 400	**Etopan XL 600**

https://israeldrugs.health.gov.il/#!/medDetails/130%2003%2030728%2000
https://israeldrugs.health.gov.il/#!/medDetails/122%2059%2030274%2000

For a printed copy of the patient information leaflet in English, please contact the registration holder by email Info@taro.com or by phone 1-800-464-664.