

NUROFEN PLUS Tablets

Ibuprofen 200mg

Codeine Phosphate Hemihydrate 12.8mg

ADVANCE DUAL ACTION POWERFUL RELIEF

Read all of this leaflet carefully because it contains important information for you.

This medicine is available without prescription. However, you will still need to take Nurofen Plus Tablets carefully to get the best results from them.

- Keep this leaflet. You may need to read it again.
- Ask your pharmacist if you need more information or advice
- You must contact a doctor if your symptoms worsen or do not improve after 3 days.
- If any of the side-effects gets serious, or if you notice any side-effect not listed in this leaflet, please tell your doctor or pharmacist.

In this leaflet:

1. What Nurofen Plus Tablets are and what are they used for?
2. What you need to know before you take Nurofen Plus Tablets
3. How to take Nurofen Plus Tablets
4. Possible side-effects
5. How to store Nurofen Plus Tablets
6. Further Information

1. WHAT NUROFEN PLUS TABLETS ARE AND WHAT ARE THEY USED FOR?

Nurofen Plus Tablets contains two active ingredients – each tackles pain in a different way: Ibuprofen belongs to a group of medicines known as non steroidal anti-inflammatory drugs (NSAIDs) and works to relieve inflammation and pain at source. The product also contains Codeine. Codeine belongs to a group of medicines called opioid analgesics which act to relieve pain. It can be used on its own or in combination with other pain killers such as paracetamol. Together they change the body's response to pain and swelling to provide relief. Nurofen Plus Tablets have been specially formulated for use when you need pain relief for example **rheumatic and muscular pain, migraine, cramping period pain, dental pain, backache, fever and symptoms of cold & flu.**

2. WHAT YOU NEED TO KNOW BEFORE YOU TAKE NUROFEN PLUS TABLETS

Do not take Nurofen Plus Tablets if you:

- Are allergic to ibuprofen, codeine, or any of the ingredients of Nurofen Plus listed in Section 6
- Have experienced shortness of breath, worsening of asthma, allergic rash or an itchy, runny nose when taking ibuprofen, codeine, aspirin or other similar medicines
- Are already taking non steroidal anti-inflammatory drugs (NSAIDs)
- Have ever had stomach bleeding or perforation after taking ibuprofen, aspirin or other similar medicines
- Have or have ever had a stomach ulcer, perforation or bleeding
- Having breathing difficulties
- Suffer from severe kidney, liver or heart problems
- Suffer from chronic constipation
- Are under 12 years of age

- For pain relief in children and adolescents (0-18 years of age) after removal of their tonsils or adenoids due to obstructive sleep apnoea syndrome.
- You know that you metabolise very rapidly codeine into morphine.
- Are in the last trimester of pregnancy or breast feeding (see pregnancy and breastfeeding)
- Are already taking monoamine oxidase inhibitors (MAOIs)

Consult your doctor or pharmacist before taking Nurofen Plus Tablets if you

- have an infection – please see heading 'Infections' below
- Have asthma or have suffered from asthma
- Have decreased respiratory reserve, acute respiratory depression or obstructive airways disease
- Have kidney, liver or heart problems
- Have heart problems including heart failure, angina (chest pain), or if you have had a heart attack, bypass surgery, peripheral artery disease (poor circulation in the legs or feet due to narrow or blocked arteries), or any kind of stroke (including "mini-stroke" or transient ischaemic attack "TIA")
- Have stomach or intestinal problems (such as Crohn's disease, ulcerative colitis or paralytic ileus)
- Have low blood pressure
- Have high blood pressure, diabetes, high cholesterol, have a family history of heart disease or stroke, or if you are a smoker
- Have a head injury or raised intracranial pressure
- Suffer from a thyroid disorder
- Suffer from a connective tissue disease such as Systemic Lupus Erythematosus (SLE) (Lupus)
- Are elderly
- Are receiving regular treatment from your doctor
- Have an adrenal gland deficiency
- Have a bowel obstruction
- Have an enlargement of the prostate (prostatic hyperplasia)
- Are taking other codeine medicines
- Have a congenital or acquired muscle weakness
- Have a history of convulsions, drug abuse or acute alcoholism
- Have chicken pox (varicella), as it is advisable to avoid the use of Nurofen Plus tablets

Warnings and precautions

Nurofen Plus Tablets contains codeine, which is an opioid medicine.

Repeated use of Nurofen Plus Tablets may result in you becoming accustomed to it (needing to take higher doses). Repeated use of Nurofen Plus Tablets may also lead to dependence, abuse and addiction, which may result in life-threatening overdose. If you are taking <Product name> for longer than the recommended time or at higher than recommended doses you are at risk of serious harms. These include serious harms to the stomach/gut and kidneys, as well as very low levels of potassium in your blood. These can be fatal (see section 4).

If you experience any of the following signs whilst taking Nurofen Plus Tablets talk to your doctor or pharmacist as it could be an indication that you are dependent or addicted.

- You need to take this medicine for longer than advised
- You need to take more than the recommended dose
- You are using this medicine for reasons other than medical reasons, for instance, 'to stay calm' or to 'help you sleep'
- You have made repeated, unsuccessful attempts to quit or control the use of this medicine
- When you stop taking this medicine you feel unwell, and you feel better once taking this medicine again ('withdrawal effects')

Codeine is transformed to morphine in the liver by an enzyme. Morphine is the substance that provides the effects of codeine and relieves pain and symptoms of cough. Some people have a variation of this enzyme and this can affect people in different ways. In some people morphine is not produced or produced in very small quantities and it will not provide enough pain relief or relieve their cough. Other people are more likely to get serious side effects because a very high amount of morphine is produced.

If you notice any of the following side effects, you must stop taking this medicine and seek immediate medical advice: slow or shallow breathing, confusion, sleepiness, small pupils, feeling or being sick, constipation, lack of appetite.

Anti-inflammatory/pain-killer medicines like ibuprofen may be associated with a small increased risk of heart attack or stroke, particularly when used at high doses. Do not exceed the recommended dose or duration of treatment.

Serious skin reactions have been reported in association with Nurofen Plus Tablets. You should stop taking this medicine and seek medical attention immediately if you develop any skin rash, lesions of the mucous membranes, blisters or other signs of allergy since this can be the first signs of a very serious skin reaction. See section 4.

Infections: Nurofen Plus Tablets may hide signs of infections such as fever and pain. It is therefore possible that Nurofen Plus Tablets may delay appropriate treatment of infection which may lead to an increased risk of complications. This has been observed in pneumonia caused by bacteria and

bacterial skin infections related to chicken pox. If you take this medicine while you have an infection and the symptoms of the infection persist or worsen, consult a doctor without delay.

Children and adolescents older than 12 years of age

There is a risk of renal impairment in dehydrated children and adolescents. Use in children and adolescents after surgery: Codeine should not be used for pain relief in children and adolescents after removal of their tonsils or adenoids due to Obstructive Sleep Apnoea Syndrome. Use in children with breathing problems: Codeine is not recommended in children with breathing problems, since the symptoms of morphine toxicity may be worse in these children.

Nurofen Plus is not recommended in adolescents with compromised respiratory function for the treatment of cough and or cold.

Taking other medicines

Please tell your doctor or pharmacist if you are taking or have recently taken any other medicines. In particular, do **NOT** take this product if you are taking any of the following:

- Aspirin or other NSAIDs including cyclooxygenase selective 2 inhibitors
- Corticosteroids (such as prednisolone) since this may increase the risk on gastrointestinal ulceration or bleeding
- Antihypertensives and diuretics: since NSAID may diminish the effect of these drugs
- Hydroxyzine (an antihistamine)
- Antiplatelet agents (such as aspirin) and selective serotonin reuptake inhibitors (a medicine for depression) since these may increase the risk of gastrointestinal side effects
- Medicines for high blood pressure and water tablets since NSAIDs may diminish the effects of these
- Lithium (a medicine for depression) since there is evidence for a potential increase in plasma levels of lithium
- Methotrexate (a medicine for cancer or rheumatism) since there is evidence for a potential increase in plasma levels of methotrexate
- Ciclosporin or tacrolimus (a medicine to suppress the immune reaction) since there is limited evidence on an increased risk for kidney activity
- Zidovudine: there is evidence of an increased risk of haematomas and haematoma in HIV(+) haemophiles receiving concurrent treatment with Zidovudine and ibuprofen
- Monoamine oxidase inhibitors, e.g. Selegiline, as depression or excitation may occur
- If patients on warfarin are prescribed Nurofen Plus by their doctor, they will need more frequent blood tests to monitor INR
- Glycosides, to stimulate your heart or treat high blood pressure
- Mifepristone (now or in the last 12 days)
- Quinolone antibiotics since patients taking NSAIDs and quinolone antibiotics may have an increased risk of developing convulsions
- Quinidine or mexiletine (a medicine used to treat heart rhythm disorder)
- Medicine used to treat diarrhoea, gastrointestinal problems, nausea and vomiting (including cimetidine, metoclopramide and domperidone)
- Naloxone used in drug overdose
- Medicines that are anti-coagulants (i.e. thin blood/prevent clotting e.g. aspirin/acetylsalicylic acid, warfarin, ticlopidine)
- Medicines that reduce high blood pressure (ACE-inhibitors such as captopril, beta-blockers such as atenolol medicines, angiotensin-II receptor antagonists such as losartan)
- Other Opioids
- Abiraterone (a medicine used to treat prostate cancer) as it may reduce the analgesic effect of codeine

- Selective Serotonin Reuptake Inhibitors (SSRIs): a type of medicine used to treat depression
- Neuroleptics or Antipsychotics (medications used to treat psychosis)
- Ritonavir (a medicine used in the treatment of HIV/AIDS)

Some other medicines may also affect or be affected by the treatment of Nurofen Plus Tablets. You should therefore always seek the advice of your doctor or pharmacist before you use Nurofen Plus Tablets with other medicines.

This medicine contains less than 1 mmol of sodium per dose, that is to say essential 'sodium free'.

Pregnancy and breastfeeding

Consult your doctor before use if you are in the first six months of pregnancy. Do not use Nurofen Plus tablets in the third trimester of pregnancy. Do not take Nurofen Plus while you are breastfeeding.

Codeine and morphine passes into breast milk.

Driving and using machines

Nurofen Plus Tablets could make you feel dizzy or drowsy. If affected, do not drive or use any tools or machinery.

3. HOW TO TAKE NUROFEN PLUS TABLETS

Adults and children over 12 years:

This medicine should not be taken for more than 3 days. The lowest effective dose should be used for the shortest duration necessary to relieve symptoms. If you have an infection, consult a doctor without delay if symptoms (such as fever and pain) persist or worsen (see Section 2). If the pain does not improve after 3 days, talk to your doctor for advice. Swallow 2 tablets with water, then if necessary, 1-2 tablets every 6 hours. Do not exceed 6 tablets in 24 hours. Nurofen Plus Tablets should not be taken by children below the age of 12 years, due to the risk of severe breathing problems.

This medicine contains codeine and can cause addiction if you take it continuously for more than 3 days. When you stop taking it you may get withdrawal symptoms. You should talk to your doctor or pharmacist if you think you are suffering withdrawal symptoms.

If you take more Nurofen Plus Tablets than you should:

If you have taken more Nurofen Plus Tablets than you should, or if children have taken this medicine by accident always contact a doctor or nearest hospital to get an opinion of the risk and advice on action to be taken. The symptoms can include nausea, stomach pain, vomiting (may be blood streaked), headache, ringing in the ears, confusion and shaky eye movement. At high doses, drowsiness, chest pain, palpitations, loss of consciousness, convulsions (mainly in children), weakness and dizziness, blood in urine, cold body feeling, and breathing problems have been reported.

If you forget to take Nurofen Plus Tablets

Take your tablets as usual. Do not take a double dose to make up for forgotten tablets.

4. POSSIBLE SIDE EFFECTS:

Like all medicines, Nurofen Plus Tablets can cause side-effects, although not everybody gets them. Tell your doctor or pharmacist if you notice any of the following:

- Stomach problems such as unexplained stomach pain, decreased appetite, dry mouth, indigestion, feeling sick and/or vomiting, diarrhoea or constipation, flatulence
- Any sign of bleeding from the stomach or bowels (vomiting blood and/or passing black stools) or worsening of colitis or Crohn's disease
- Stomach ulcers
- Liver, kidney problems or difficulty urinating
- Severe sore throat with a high fever and flu-like symptoms, severe exhaustion, nose or skin bleeding, mouth ulcers
- Skin flushing, severe skin reactions such as skin peeling
- A severe headache, stiff neck, nausea, vomiting, fever or confusion
- Allergic reactions such as unexplained wheezing, shortness of breath, swelling of the face, tongue or throat, palpitations, skin rash or itching ('Nettle rash')
- Asthma, aggravation of asthma or wheezing
- Drowsiness, difficulty coughing or breathing
- Blood disorders, swelling, high blood pressure, heart failure, blurred or double vision
- Depression, hallucination, confusion, dependence, altered mood, restlessness, nightmares
- Vertigo
- Hypothermia, excessive sweating, irritability, fatigue, malaise
- severe skin infections and soft tissue complications during chicken pox (varicella) infection
- A severe skin reaction known as DRESS syndrome can occur. Symptoms of DRESS include: skin rash, fever, swelling of lymph nodes and an increase of eosinophils (a type of white blood cells).
- A red, scaly widespread rash with bumps under the skin and blisters mainly localised on the skin folds, trunk and upper extremities accompanied by fever at the initiation of treatment (acute generalised exanthematous pustulosis). Stop using Nurofen Plus Tablets if you develop these symptoms and seek medical attention immediately. See also section 2.
- Skin becomes sensitive to light.

Nurofen Plus Tablets, especially when taken at higher than recommended doses or for a prolonged period of time, can cause damage to your kidneys and affect them removing acids properly from your blood into the urine (renal tubular acidosis). It can also cause very low levels of potassium in your blood (see section 2). This is a very serious condition and will require immediate treatment. Signs and symptoms include muscle weakness and light-headedness.

Undesirable effects may be minimised by using the minimum effective dose for the shortest duration. Medicines such as Nurofen Plus Tablets may be associated with a small risk of heart attack ("myocardial infarction") or stroke. Any risk is more likely with high doses and prolonged treatment.

Do not exceed the recommended dose or duration of treatment (3 days). If you have heart problems, previous stroke or think you might be at risk of these conditions (for example if you have high blood pressure, diabetes or high cholesterol or are a smoker) you should discuss your treatment with your doctor or pharmacist.

Nurofen Plus Tablets belong to a group of medicines which may impair fertility in women. This is reversible on stopping the medicine. Patients should be aware that this product may make it more difficult to become pregnant and should inform their doctor if planning to become pregnant or having problems becoming pregnant. The elderly are at an increased risk of side effects. Shortness of breath may occur if you suffer from or have a history of allergic disease.

Tell your doctor if you notice any unusual symptoms – especially dark bowel motions, vomiting blood or stomach pains. Serious skin reactions (such as Steven-Johnson syndrome) have been

reported rarely in association with the use of NSAIDs. The use of Nurofen Plus Tablets should be stopped immediately at the first appearance of skin rash, mouth ulcers or any signs of allergic reactions.

There is increased risk of GI bleeding, ulceration, perforation with increasing NSAID doses, in patients with a history of ulcers, and the elderly. It is recommended to commence treatment at the lowest dose. The option of adding a protective agent should also be discussed with your doctor.

There is an increased risk of developing side effects of opioid toxicity in a small number of individuals who take this product even at low doses (estimated 1-2% of Caucasians), who are referred to as ultra metabolisers.

If any of the side effects gets serious, or if you notice any side effects not listed in the leaflet, please tell your doctor or pharmacist.

If you get any side effects, talk to your doctor or pharmacist. This includes any possible side effects not listed in this leaflet. You can also report side effects directly (see details below). By reporting side effects you can help provide more information on the safety of this medicine. HPR
Pharmacovigilance, Earlsfort Terrace, IRL-Dublin 2. Tel: +353 1 6764971.
Fax: + 353 1 6762517. Website: www.hpra.ie, e-mail: medsafety@hpra.ie.

5. HOW TO STORE NUROFEN PLUS TABLETS?

KEEP OUT OF SIGHT AND REACH OF CHILDREN

Do not use after the expiry date printed on the carton and the blister packaging.
Do not store above 25°C. Store in the original packaging in order to protect from light.

6. FURTHER INFORMATION

What Nurofen Plus Tablets contain

The active substances are Ibuprofen 200mg, Codeine Phosphate Hemihydrate 12.8mg. Codeine Phosphate Hemihydrate is the new name for Codeine Phosphate. The ingredient itself has not changed. The other ingredients are Microcrystalline Cellulose, Sodium Starch Glycolate, Pregelatinised Starch, Hypermellose, Colour (titanium dioxide (E171))

What Nurofen Plus Tablets look like and contents of the pack

Nurofen Plus Tablets are white film coated tablets, embossed with the logo "N+" on one side. Available in packs of 12 or 24 tablets.

Marketing Authorisation Holder and Manufacturers.

PA Holder: Reckitt Benckiser Ireland Ltd, 7 Riverwalk, Citywest Business Campus, Dublin 24, Ireland.
Manufacturers: Reckitt Benckiser Healthcare International Ltd. Thane Road, Nottingham, NG90 2DB, UK and RB NL Brands B.V. Schiphol Blvd 207, 1118 BH Schiphol, NL. For product enquiries, please telephone (01) 6305429

The leaflet was updated in upon approval