

PATIENT MEDICATION INFORMATION

READ THIS FOR SAFE AND EFFECTIVE USE OF YOUR MEDICINE

Pr **PLAQUENIL**[®]

Hydroxychloroquine Sulfate Tablets

Read this carefully before you start taking **PLAQUENIL** and each time you get a refill. This leaflet is a summary and will not tell you everything about this drug. Talk to your healthcare professional about your medical condition and treatment and ask if there is any new information about **PLAQUENIL**.

What is **PLAQUENIL** used for?

PLAQUENIL is used in adults to:

- Treat **Rheumatoid Arthritis (RA)**: a disease marked by stiffness, swelling and pain in your joints.
- Treat **Systemic Lupus Erythematosus (SLE)**: a disease where your immune system attacks healthy parts of your body by mistake. It can affect your skin, joints, kidneys, brain, and other organs.
- Treat **Discoid Lupus Erythematosus (DLE)**: a disease similar to SLE. DLE only affects your skin with red rash or scaly patches.

PLAQUENIL is used in patients 6 years of age and older to:

- Prevent and treat certain forms of **Malaria**: an infection caused by parasites in your red blood cells. Symptoms can include high fever, shaking, chills, and extreme sweating.

You can only get **PLAQUENIL** with a doctor's prescription.

How does **PLAQUENIL** work?

It is not known how **PLAQUENIL** works in the body to treat RA, SLE, and DLE. **PLAQUENIL** may take up to six months to take effect. For malaria, **PLAQUENIL** works by killing the parasite that causes the infection.

What are the ingredients in **PLAQUENIL**?

Medicinal ingredient: hydroxychloroquine sulfate.

Each 200 mg tablet contains 155 mg of hydroxychloroquine as the base.

Non-medicinal ingredients: black ink, calcium hydrogenophosphate, carnauba wax, corn starch, magnesium stearate, Opadry White YS-I-7443 and polyethylene glycol 400.

PLAQUENIL comes in the following dosage forms:

Tablets, 200 mg.

Do not use PLAQUENIL if:

- you are allergic to
 - hydroxychloroquine sulfate
 - any of the other ingredients of PLAQUENIL
 - any similar drugs such as chloroquine
- you have retinopathy. This is an eye problem affecting the retina at the back of your eye. PLAQUENIL may cause irreversible damage to your retina. You should tell your doctor right away if you have any **Visual Problems**.
- you are a child below 6 years of age **or** weigh less than 35 kg.

To help avoid side effects and ensure proper use, talk to your healthcare professional before you take PLAQUENIL. Talk about any health conditions or problems you may have, including if you:

- were born with, now have, or have a family history of long QT interval. PLAQUENIL may cause **Heart Rhythm Disorders** in some patients. These **Heart Rhythm Disorders** can be seen on an ECG, or an electrical recording of the heart. Caution should be taken when taking PLAQUENIL if you:
 - have heart disease, which can include heart failure, slow heartbeat, heart palpitations or irregular heartbeat. The risk of heart problems may increase with higher doses of PLAQUENIL.
 - have had a heart attack (myocardial infarction)
 - have a family history of sudden death from heart attack before the age of 50
 - take other drugs that can cause prolonged QT interval or are known to affect the rhythm of your heart
- have mental health problems.
- have a low level of potassium, calcium or magnesium in your blood, or have a condition that may affect the levels of those salts in the blood. Examples are an eating disorder or prolonged vomiting.
- are allergic or sensitive to a drug called to quinine.
- are pregnant, or you are planning to get pregnant. PLAQUENIL may be passed to your unborn baby. PLAQUENIL may harm your unborn baby. Your doctor will evaluate the benefit and risk of using PLAQUENIL during pregnancy.

- are breastfeeding. PLAQUENIL passes into breast milk in small amounts. Infants can be very sensitive to the toxic effects of drugs like PLAQUENIL. There is not enough PLAQUENIL in breast milk to protect an infant against malaria. The infant should receive their own malaria treatment if necessary. Talk to your doctor about the risks PLAQUENIL can have on your baby. These risks depend on:
 - why you are taking PLAQUENIL;
 - how long you will be taking PLAQUENIL for.
- have diabetes or symptoms of **low blood sugar**. PLAQUENIL can cause low blood sugar, and sometimes, low blood sugar can be very dangerous. You may pass out or need to go to the hospital.
- have liver or kidney disease.
- have alcoholism.
- have a blood disease, including a rare blood disease called porphyria. PLAQUENIL can make this worse.
- have skin problems or diseases such as psoriasis. PLAQUENIL can make these worse.
- have nervous system problems or disease.
- Have bone marrow problems that cause low counts of blood platelets, and white and red blood cells
- have a genetic red blood cell disease known as “glucose-6-phosphate dehydrogenase deficiency”.
- have gastrointestinal disorders. These are problems in the intestines, stomach, or gut.
- have decreased vision.
- have muscle, tendon or nerve problems, including muscle weakness.
- have mood problems, including thoughts of suicide or depression.
- are 65 years old or older. There is a higher chance of side effects when this age group takes PLAQUENIL.

Other warnings you should know about:

PLAQUENIL can cause **long QT interval** or **torsade de pointes**. This is a dangerously fast heart rate. It can lead to cardiac arrest, sudden collapse and death.

Heart problems or failure, cardiomyopathy, an enlarged or weak heart can occur if you take PLAQUENIL for long periods of time. These are serious and can result in death. Your doctor will check your heart regularly.

PLAQUENIL can cause **mental health problems** such as abnormal thoughts, anxiety, hallucinations, confusion, depression, thoughts of **self-harm** or **suicide**. This can happen to

people who have never had mental health problems before. These can occur within the first month of treatment with PLAQUENIL.

PLAQUENIL can cause permanent **eye damage**. To help prevent this, you should have an eye exam before you start taking PLAQUENIL. You will need more eye exams while you are taking PLAQUENIL.

When you go outside, protect your skin from the sun by:

- wearing appropriate clothing, and
- using sunscreen cream with a minimum SPF 30 rating.

It is unclear whether PLAQUENIL may affect male fertility. Talk to your doctor if you would like to father a child in the future.

Severe Skin Reactions / Rashes: PLAQUENIL can cause **severe cutaneous adverse drug reactions (SCARs)** such as drug reaction with eosinophilia and systemic symptoms (DRESS), acute generalized exanthematous pustulosis (AGEP), Stevens-Johnson syndrome (SJS) and toxic epidermal necrolysis (TEN) and maybe fatal. These skin reactions and rashes can involve ulcers of the mouth, throat, nose, genitals and conjunctivitis (red and swollen eyes). Flu-like symptoms such as fever, headache and body ache can happen before these reactions start. The skin reactions and rashes may progress to blisters and peeling skin. If you develop these symptoms, stop taking PLAQUENIL and tell your doctor right away.

Driving and Using Machines: You may have blurry vision when taking PLAQUENIL. Do not drive or do things that require you to be alert. Wait until you know how you respond to PLAQUENIL and can see well. If you continue to have difficulty, your doctor may reduce your dose.

Before and during your treatment with PLAQUENIL your doctor may do some tests. These may include:

- blood tests
- an electrocardiogram (ECG)
- a periodic exam of your muscles and tendon reflexes
- complete eye exams

Tell your healthcare professional about all the medicines you take, including any drugs, vitamins, minerals, natural supplements or alternative medicines.

The following may interact with PLAQUENIL:

- Drugs for depression (tricyclic antidepressants) and psychiatric disorders (antipsychotics).
- Digoxin. If you are taking both PLAQUENIL and digoxin, your doctor may decide to

check the level of digoxin in your blood, as the dose may need to be reduced.

- Anti-diabetic drugs, including insulin and metformin. If you take PLAQUENIL and a drug for diabetes or high blood sugar, there is a risk of having very low blood sugar. This can be life-threatening. Your doctor may decide to reduce the doses of the drug or insulin to control diabetes.
- Antiepileptic drugs (e.g. carbamazepine).
- Some antibiotics used for bacterial infections (e.g. aminoglycoside antibiotics, erythromycin, azithromycin, moxifloxacin, clarithromycin, spiramycin). Taking these antibiotics at the same time as hydroxychloroquine may increase the chance of you getting side effects that affect your heart which could be life-threatening.
- Neostigmine and pyridostigmine (medicines used to treat muscle disorders).
- Cimetidine (medicine used to treat heartburn).
- Cyclosporine (an immunosuppressant medication).
- Drugs known as "CYP2C8 and CYP3A4" inhibitors
 - Medicines used to treat fungal infections like ketoconazole, itraconazole, fluconazole
 - Medicines used to treat infections like erythromycin, clarithromycin.
 - Medicines used to treat nausea and vomiting like aprepitant.
 - Medicines used to treat multiple sclerosis like teriflunomide.
 - Medicines used to treat viral infections in immunocompromised patients like letermovir, ritonavir.
- Drugs known as CYP2C8 and CYP3A4 inducers
 - Medicines used to treat epilepsy like carbamazepine, phenobarbital.
 - Medicines used to treat Tuberculosis like rifampin.
 - Medicine used to treat depression like St. John's Wort.
- Medicines that are known to cause cardiac arrhythmias (irregular heartbeats).
- Medicines used to treat heart problems like digoxin, flecainide, propafenone
- Medicines used to treat high blood pressure like metoprolol
- Medicines used to treat blood clots like dabigatran, clopidogrel
- Medicines used to treat blood lipid problems, like statins (e.g. simvastatin), or gemfibrozil
- Medicines used for sedation, like midazolam

- Halofantrine (a medicine used to treat malaria). If you are taking halofantrine, you should not be taking PLAQUENIL at the same time.
- Antacids. You should take antacids at least 2 hours before or 2 hours after taking PLAQUENIL.
- Rabies vaccine.
- Medicines that may affect the liver, the kidney, the skin or the eye.
- Medicines that may increase the risk of convulsions (e.g. antimalarials (mefloquine)).
- Medicines that decrease blood salt levels (e.g. water pills, laxatives, amphotericin B, high dose corticosteroids, and proton pump inhibitors).
- Agalsidase (a medicine used to treat a rare genetic disease called Fabry disease).
- Medicines that may increase risk of retinal toxicity. An example is tamoxifen, which is used to treat breast cancer. When taken alone, both PLAQUENIL and tamoxifen can cause damage to your retina at in the eye. Taking both drugs at the same time can increase your risk of retinal damage.
- Praziquantel (a medicine used to treat some infestations).

Do NOT eat grapefruit or drink grapefruit juice while taking PLAQUENIL.

PLAQUENIL has been used safely with salicylates (Aspirin[®]), non-steroidal anti-inflammatory medications, methotrexate and corticosteroids.

How to take PLAQUENIL:

Take PLAQUENIL exactly as your doctor told you to. Never take more PLAQUENIL than your doctor has prescribed.

To help avoid an upset stomach, take PLAQUENIL with a meal or a glass of milk.

Usual dose:

Your doctor will decide on the best dose for you. It may be based on your weight, physical health and other factors such as what other medications you are taking. The dose may need to be stopped or temporarily reduced due to side effects. The dose may then be re-started or increased to an optimum level by your doctor. Your dose will likely be lowered during treatment, after your Initial Dose. You may take the lower dose for a lengthy amount of time. This is called a Maintenance Dose.

Condition	Recommended dose	Number of tablets a day
Rheumatoid Arthritis	Initial: 400 – 600 mg a day	2 - 3
	Maintenance: 200 – 400 mg a day	1 - 2
Lupus Erythematosus	Initial: 400 mg, once or twice a day	2 - 4
	Maintenance: 200 – 400 mg a day	1 - 2
Malaria (adults)	Prevention: 400 mg a week, on the same day of each week, starting 2 weeks before exposure.	2
	Treatment: 800 mg initially, followed by 400 mg 6-8 hours later, and then 400 mg daily for the next two days.	4 2 2
Malaria (children)	Your dose will be calculated by your doctor based on each child's body weight.	

Should you have a serious change of health at any point while taking PLAQUENIL, see your doctor.

For patients with RA, SLE or DLE, if PLAQUENIL makes your symptoms completely better, talk to your doctor. They may want to bring down your daily dose. Never change your dose without talking with your doctor first.

Overdose:

Taking too much PLAQUENIL is dangerous and can lead to death. You could have symptoms of overdose within 30 minutes after taking it.

Taking too much PLAQUENIL is also dangerous for children. Children have died by taking too much PLAQUENIL. If you think an infant or small child has swallowed even one pill, immediately take them to the nearest hospital emergency room or dial "911" on your telephone.

The symptoms of overdose can also be side effects of PLAQUENIL. These include:

- headache

- feeling drowsy
- vision problems, such as blurry or double vision
- heart problems, such as uneven heartbeats or rapid heartbeats
- fainting
- muscle weakness
- convulsions
- serious trouble breathing

If you think you, or a person you are caring for, have taken too much PLAQUENIL, contact a healthcare professional, hospital emergency department, or regional poison control centre immediately, even if there are no symptoms.

Missed Dose:

If you forget to take a dose, take it as soon as you remember. But if it is within twelve hours of your next dose, skip the one you missed and take only the regularly scheduled dose. **Never take a double dose.**

What are possible side effects from using PLAQUENIL?

These are not all the possible side effects you may feel when taking PLAQUENIL. If you experience any side effects not listed here, contact your healthcare professional.

Before and during your treatment with PLAQUENIL your doctor may do some tests. These may include:

- blood tests
- an electrocardiogram (ECG)
- a periodic exam of your muscles and tendon reflexes
- complete eye exams

PLAQUENIL can cause permanent **eye damage**. To help prevent this, you should have an eye exam before you start taking PLAQUENIL. You will need more eye exams while you are taking PLAQUENIL.

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
VERY COMMON			
Nausea, stomach pain, stomach cramps	✓		

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
COMMON			
Diarrhea		✓	
Vomiting		✓	
Anorexia: loss or lack of appetite		✓	
Visual problems and damage to the retina of the eye: blurred vision, seeing halos around lights, especially at night. Seeing light flashes and streaks. Night blindness with difficulty seeing at night or in poor light. Visual field loss including blind spots or blind areas in your vision. Change in eye colour. Difficulty focusing your eyes, or skipping words when reading.		✓	
Headache	✓		
Rash, itchy rash with raised red bumps		✓	
Nervousness, quick changes in mood (emotional lability)		✓	
RARE			
Dizziness or vertigo: feel as if you or the objects around you are moving when they are not.	✓		
Change in colour of skin, mucous membranes and hair: bleaching of hair. Loss or increase in skin pigment (bluish-black colour).		✓	
Alopecia: hair loss from your head or any part of your body.		✓	
Hearing problems: ringing in the ears. Hearing loss.		✓	

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
Nerve and muscle problems: tingling, numbness, burning pain, weakness, cramps, spasms, restlessness, rigidity, tremors, twitches, difficulty walking		✓	
UNKNOWN FREQUENCY			
Allergic reaction or angioedema: rash, hives, swelling of the face, lips, tongue or throat, difficulty swallowing or breathing			✓
Heart rhythm disorders including long QT interval, torsade de pointes and heart block: abnormal heartbeat, life-threatening irregular heartbeat, palpitations			✓

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
<p>Severe skin reactions such as:</p> <ul style="list-style-type: none"> • Drug Reaction with Eosinophilia and Systemic Symptoms (DRESS): rash with a fever and flu-like symptoms and enlarged lymph nodes. • Acute Generalized Exanthematous Pustulosis (AGEP): blistering, widespread scaly skin, pus-filled spots together with fever. • Stevens-Johnson syndrome (SJS): blistering or peeling of the skin around the lips, eyes, mouth, nose, genitals, hands or feet, flu-like symptoms and fever. • Toxic Epidermal Necrolysis (TEN): multiple skin lesions, itching of the skin, joint aches, fever and a general ill feeling. • Sweet's syndrome: skin reaction including reddish-purple color, raised, painful sores, particularly on your arms, hands, fingers, face and neck, fever. • Erythema Multiforme: skin reaction/rash characterised by raised red or purple skin patches, possibly with blister or crust in the centre with or without mild itching, joint pain, fatigue and fever. 			✓
Severe breathing problems including bronchospasm, angioedema: sudden shortness of breath			✓
Increased sensitivity to sunlight. Skin rash due to sunlight can be reduced by appropriate use of sunscreen creams.		✓	
Muscle weakness		✓	
Permanent damage to vision		✓	

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
Fainting spells or loss of consciousness		✓	
Heart problems or heart failure, cardiomyopathy, an enlarged or weak heart: shortness of breath with exercise or even at rest. Swelling of the legs, ankles and feet. Irregular heartbeats that feel rapid or pounding. Chest pain. Sudden fainting or feeling tired, light-headed and dizzy. You can have a seizure or fit.			✓
Liver problems: unusual tiredness, nausea, vomiting, abdominal pain, jaundice (yellow discoloration of the eyes or skin), dark urine, weakness, poor appetite.		✓	
Bone Marrow Depression or a decrease in production of cells: Low White Blood cells (leukocytes): Fever and chills. Infections. Anemia or low red blood cells (erythrocytes): Fatigue, extreme tiredness that does not get better with rest. Paleness of skin, lips, and nail beds. Low platelets used for blood clotting (thrombocytes): Bleeding: nose bleeds, gums, or mouth. Tiny red spots on the skin.		✓	
Convulsions, seizures or fits			✓
Psychosis: hallucinations, loss of contact with reality		✓	
Mental health problems: irrational/abnormal thoughts, irritability, anxiety, hallucinations, feeling confused or depressed, agitation, difficulty sleeping, delusions (feelings of distrust and false beliefs), changes in mood, feeling elated or overexcited or abnormally happy		✓	
Thoughts or actions of suicide or self-harm			✓

Serious side effects and what to do about them			
Symptom / effect	Talk to your healthcare professional		Stop taking drug and get immediate medical help
	Only if severe	In all cases	
Hypoglycemia or low blood sugar: hunger pains, sweating, shakiness, weakness, dizziness, fast heartbeat, nausea, irritability, blurred vision, confusion, loss of consciousness		✓	
Muscle, nerve and tendon problems: long-lasting involuntary muscle contraction, impairment of voluntary movements, tremor. Weakness. Decreased reflexes or feeling by nerves.			✓

If you have a troublesome symptom or side effect that is not listed here or becomes bad enough to interfere with your daily activities, tell your healthcare professional.

Reporting Side Effects

You can report any suspected side effects associated with the use of health products to Health Canada by:

- Visiting the Web page on Adverse Reaction Reporting (<https://www.canada.ca/en/health-canada/services/drugs-health-products/medeffect-canada/adverse-reaction-reporting.html>) for information on how to report online, by mail or by fax; or
- Calling toll-free at 1-866-234-2345.

NOTE: Contact your health professional if you need information about how to manage your side effects. The Canada Vigilance Program does not provide medical advice.

Storage:

- Keep out of reach and sight of infants and small children.
- Store at room temperature (15°C - 30°C).
- Do not use PLAQUENIL after the expiry date.

If you want more information about PLAQUENIL:

- Talk to your healthcare professional.
- Find the full product monograph that is prepared for healthcare professionals and includes this Patient Medication Information by visiting the Health Canada website: (<https://www.canada.ca/en/health-canada/services/drugs-health-products/drug-products/drug-product-database.html>); the manufacturer's website (www.sanofi.ca) or by

calling 1-800-265-7927.

This leaflet was prepared by sanofi-aventis Canada Inc.

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