

Calcium Folate 10 mg/ml

Solution for injection or infusion

Folinic acid

Read all of this leaflet carefully before you start using this medicine because it contains important information for you.

- Keep this leaflet. You may need to read it again.
- If you have any further questions, ask your doctor, pharmacist or nurse.
- This medicine has been prescribed for you only. Do not pass it on to others. It may harm them, even if their signs of illness are the same as yours.
- If you get any side effects, talk to your doctor, pharmacist or nurse. This includes any possible side effects not listed in this leaflet. See section 4.

What is in this leaflet

1. What Calcium Folate is and what it is used for
2. What you need to know before you use Calcium Folate
3. How to use Calcium Folate
4. Possible side effects
5. How to store Calcium Folate
6. Contents of the pack and other information

1. WHAT CALCIUM FOLINATE IS AND WHAT IT IS USED FOR

Calcium Folate solution for injection or infusion contains calcium folinate, which is one of a group of medicine called detoxifying agents. It is a calcium salt of folic acid, which is related to the vitamin folic acid.

Calcium Folate is used to:

- reduce the harmful effects and treat overdose of certain types of anti-cancer medicines for instance methotrexate and other folic acid antagonists. This is known as "calcium folinate rescue".
- treat cancer in combination with 5-fluorouracil (an anti-cancer medicine). 5-fluorouracil works better when it is given together with Calcium Folate.

2. WHAT YOU NEED TO KNOW BEFORE YOU USE CALCIUM FOLINATE

Calcium Folate must not be injected intrathecally (into the spine).

Do not use Calcium Folate:

- if you are allergic to calcium folinate or any of the other ingredients of this medicine (listed in section 6).
- if you have a type of anaemia caused by too little vitamin B12

Warnings and precautions

Talk to your doctor, pharmacist or nurse before using Calcium Folate.

If you are to receive calcium folinate and fluorouracil treatment at the same time take special care if:

- you have had radiotherapy
- you have stomach or bowel trouble

Special care is also needed if you are elderly and you are to receive calcium folinate and fluorouracil treatment at the same time.

Calcium Folate contains sodium. This medicine contains less than 1 mmol sodium (23 mg) per dose, i.e. essentially "sodium free".

Other medicines and Calcium Folate

Tell your doctor if you are using, have recently used or might use any other medicines.

Special care is needed if you are taking/using other medicines as some could interact with Calcium Folate, for example:

- folic acid antagonists (see section 1) – the effectiveness of these medicines will be reduced by calcium folinate;
- fluorouracil (anti-cancer medicine) – the effectiveness and side effects of this medicine will be increased by calcium folinate;
- medicines used to treat epilepsy (phenobarbitone, phenytoin, primidone or succinimides) – the effectiveness of these medicines may be reduced by calcium folinate. Your doctor may check blood levels of these medicines and change your dose to prevent increased convulsions (fits).

Pregnancy and breast-feeding

If you are pregnant or breast-feeding, think you may be pregnant or are planning to have a baby, ask your doctor or pharmacist for advice before taking this medicine.

It is unlikely that your doctor will ask you to take/use a folic acid antagonist or fluorouracil whilst you are pregnant or breast-feeding. However, if you have taken/used a folic acid antagonist whilst pregnant or breast-feeding, this medicine (calcium folinate) may be used to reduce its side effects.

3. HOW TO USE CALCIUM FOLINATE

This medicine may be given by injection (using a syringe) into muscle. Alternatively it may be given by injection or infusion (drip) into a vein. If it is given by infusion Calcium Folate will be diluted first.

Your doctor will work out the correct dose of Calcium Folate for you and how often it must be given. It will depend upon the medical condition which is being treated.

If you use more Calcium Folate than you should

This medicine will be given to you in a hospital, under the supervision of a doctor. It is unlikely that you will be given too much or too little, however, tell your doctor or nurse if you have any concerns.

4. POSSIBLE SIDE EFFECTS

Like all medicines, this medicine can cause side effects, although not everybody gets them.

Very rare: may affect up to 1 in 10,000 people

- severe allergic reaction - you may experience a sudden itchy rash (hives), swelling of the hands, feet, ankles, face, lips, mouth or throat (which may cause difficulty in swallowing or breathing), and you may feel you are going to faint. This is a serious side effect. You may need urgent medical attention.



The following information is intended for healthcare professionals only:

Calcium Folate 10 mg/ml

Solution for injection or infusion

Folinic acid

Further to the information included in section 3, practical information on the preparation/handling of the medicinal product is provided here.

Incompatibilities

Incompatibilities have been reported between injectable forms of calcium folinate and injectable forms of droperidol, fluorouracil, foscarnet and methotrexate.

Droperidol

1. Droperidol 1.25 mg/0.5 ml with calcium folinate 5 mg/0.5 ml, immediate precipitation in direct admixture in syringe for 5 minutes at 25°C followed by 8 minutes of centrifugation.
2. Droperidol 2.5 mg/0.5 ml with calcium folinate 10 mg/0.5 ml, immediate precipitation when the drugs were injected sequentially into a Y-site without flushing the Y-side arm between injections.

Fluorouracil

Calcium folinate must not be mixed in the same infusion as 5-fluorouracil because a precipitate may form. Fluorouracil 50 mg/ml with calcium folinate 20 mg/ml, with or without dextrose 5% in water, has been shown to be incompatible when mixed in different amounts and stored at 4°C, 23°C, or 32°C in polyvinyl chloride containers.

Foscarnet

Foscarnet 24 mg/ml with calcium folinate 20 mg/ml; formation of a cloudy yellow solution reported.

Instructions for use and handling, and disposal

Calcium Folate is intended only for single use. Any unused portion of the solution should be disposed of in accordance with the local requirements.

Prior to administration, calcium folinate should be inspected visually. The solution for injection should be a clear and yellowish solution. If cloudy in appearance or particles are observed, the solution should be discarded.

Uncommon: may affect up to 1 in 100 people

- fever

Rare: may affect up to 1 in 1,000 people

- an increase in convulsions (fits) in patients with epilepsy
- depression
- agitation
- problems with the digestive system
- difficulty sleeping (insomnia)

Combination therapy with 5-fluorouracil only:

If you receive calcium folinate in combination with an anticancer medicine containing fluoropyrimidines, it is more likely that you experience the following side effects of this other medicine:

Very common: may affect more than 1 in 10 people

- nausea
- vomiting
- severe diarrhea
- drying out which may be due to diarrhea
- inflammation of the lining of the intestine and mouth (life-threatening conditions have occurred)
- reduction in the number of blood cells (including life-threatening conditions)

Common: may affect up to 1 in 10 people

- redness and swelling of the palms of the hands or the soles of the feet which may cause the skin to peel (hand-foot syndrome)

Not known: frequency cannot be estimated from the available data

- elevated ammonia level in the blood

Reporting of side effects

If you get any side effects, talk to your doctor, nurse or pharmacist. This includes any possible side effects not listed in this leaflet. You can also report side effects directly via the Yellow Card Scheme, website: www.mhra.gov.uk/yellowcard.

By reporting side effects you can help provide more information on the safety of this medicine.

5. HOW TO STORE CALCIUM FOLINATE

Store in a refrigerator (2°C-8°C).

Keep this medicine out of the sight and reach of children.

Do not use this medicine after the expiry date which is stated on the label and carton after EXP.

The expiry date refers to the last day of that month.

The vials should be stored in the outer carton, in order to protect from light.

Unopened vials: 18 months.

After first opening: 28 days at both room temperature and 2-8° C.

After dilution: chemical and physical in-use stability after dilution to 0.5 mg/ml and 4 mg/ml with either 0.9% sodium chloride solution or 5% glucose solution was demonstrated for up to 28 days at both room temperature and 2-8°C.

From a microbial point of view, the product should be used immediately. If not used immediately, in-use storage times and conditions prior to use are the responsibility of the user and would normally not be longer than 24 hours at 2 to 8°C, unless reconstitution/dilution has taken place in controlled and validated aseptic conditions.

6. CONTENTS OF THE PACK AND OTHER INFORMATION**What Calcium Folate contains**

- The active substance is calcium folinate. Each ml contains 10 mg of folic acid provided as calcium folinate.
Each 5, 10, 30 and 50 ml solution contains 50, 100, 300 and 500 mg of folic acid respectively (provided as calcium folinate).
- The other ingredients are trometamol, hydrochloric acid, sodium hydroxide and water for injections.

What Calcium Folate looks like and contents of the pack

Calcium Folate is a clear, pale yellow solution for injection which comes in glass containers called vials.

It is supplied in packs containing:

- 1 or 5 x 5 ml of Calcium Folate in vials of 10 ml
- 1 or 5 x 10 ml of Calcium Folate in vials of 10 ml
- 1 or 5 x 30 ml of Calcium Folate in vials of 50 ml
- 1 or 5 x 50 ml of Calcium Folate in vials of 50 ml

Marketing Authorisation Holder and Manufacturer**Marketing Authorisation Holder**

Hikma Farmacêutica (Portugal) S.A.
Estrada do Rio da M6, n°8, 8A e 8B,
Fervença, 2705-906 Terrugem SNT
Portugal

Manufacturer

Hikma Italia S.p.A.
Viale Certosa, 10,
27100 - Pavia
Italia

Distributed by:

Consilient Health (UK) Ltd.
No. 1 Church Road,
Richmond upon Thames, Surrey,
TW9 2QE

This leaflet was last revised in March 2017.

