

ANNEX III

SUMMARY OF PRODUCT CHARACTERISTICS, LABELLING AND PACKAGE LEAFLET

Note: This SPC, labelling and packages leaflet is the version valid at the time of Commission decision.

After the Commission decision the Member State competent authorities, in liaison with the reference Member State, will update the product information as required. Therefore, this SPC, labelling and package leaflet may not necessarily represent the current text.

SUMMARY OF PRODUCT CHARACTERISTICS

The valid summary of product characteristics is the final version achieved during the Coordination group procedure with the following amendments:

SUMMARY OF PRODUCT CHARACTERISTICS

1. NAME OF THE MEDICINAL PRODUCT

Prevora 100mg/ml Dental Solution.

2. QUALITATIVE AND QUANTITATIVE COMPOSITION

Each 1 ml of Prevora Dental Solution (Stage 1 chlorhexidine coating) contains chlorhexidine diacetate 100mg

For a list of full excipients, see section 6.1

3. PHARMACEUTICAL FORM

Dental Solution

Stage 1 chlorhexidine coating

A clear, slightly brownish solution with a characteristic odour, free of visible particulate matter.

Stage 2 sealant coating

A milky-white liquid of low viscosity with a faint characteristic odour, free of visible particulate matter.

4. CLINICAL PARTICULARS

4.1 *Therapeutic indications*

Prevora 100mg/ml Dental Solution is an antiseptic solution which is applied topically to the dentition of patients for the prevention of ~~root~~ **coronal and root** caries in adult patients at high-risk of dental caries (e.g. xerostomia sufferers **or those with 3 or more caries at the start of the treatment plan**). To be used in dental offices only by dental professionals.

Patients should be advised on the importance of oral hygiene and sugar intake: In the case of patients with poor oral hygiene and/or frequent consumption of sugars, advise the patient that regular and frequent tooth brushing with fluoridated tooth paste, combined with controlling sugar intake, are important to the overall success of treatment.

4.2 *Posology and method of administration*

Posology

An individual dosing in adults is between 300 µl and 600 µl of Prevora 100 mg/ml Dental Solution.

The patient is to receive 5 treatments in the initial year of treatment, of which 4 are administered one week apart in the first month, and the final dose is administered at 6 months. Treatment of the dental patient thereafter is according to professional clinical judgment of the risk of dental caries.

Prevora 100 mg/ml Dental Solution is administered topically to the entire dentition of the patient using a cotton pellet or fine paint brush. The cotton pellet or brush is dipped into the Prevora 100 mg/ml Dental solution and thereafter is applied to the tooth surfaces (Figure 1).

Figure 1 Application of Prevora 100 mg/ml Dental Solution



Dip cotton pellet or fine brush in vial and then apply pellet or brush to tooth surfaces.

The patient should be instructed that:

- The dried Prevora coating will begin coming off the teeth during the next meal.
- They should avoid eating hard foods (e.g. meat, apples) for at least 4 hours after treatment
- Do not chew gum for at least 24 hours.
- Do not brush his/her teeth for 24 hours after treatment. Then to resume brushing with a new brush 2 to 3 times daily with fluoride tooth paste.
- Do not floss for 3 days after treatment. Then to resume flossing daily.
- If dentures are worn, clean and disinfect these dentures regularly prior to use. Disinfect using soap and warm water.

Method of administration

External (oral) topical use in the dental office by a dental professional. This product is not intended to be swallowed.

4.3 Contraindications

Hypersensitivity to chlorhexidine, Sumatra benzoin or ethanol.

4.4 Special warnings and precautions for use

For external (oral) topical use only – keep out of the eyes and ears. If the drug product comes into contact with the eyes, wash out promptly and thoroughly with water.

Prevora 100 mg/ml Dental Solution should be used with caution in patients with a history of asthma or eczema. Avoid application of Prevora 100 mg/ml Dental Solution to the soft tissues. Failure to do so can result in temporary stinging or mild inflammation of the soft tissues.

4.5 Interaction with other medicinal products and other forms of interaction

Prevora 100 mg/ml Dental Solution should not be applied immediately following use of an oil-based prophylactic paste or up to 3 days following application of a fluoride dental varnish.

Chlorhexidine is incompatible with anionic agents.

4.6 Fertility, pregnancy and lactation

No controlled studies have been carried out to ascertain if there are any adverse reactions when Prevora 100 mg/ml Dental Solution is applied to the dentition of women of childbearing potential, or to the dentition of expectant or breast feeding mothers. Therefore, it is recommended that Prevora 100 mg/ml Dental Solution should not be administered during pregnancy. Since many drugs are excreted during lactation and there have not been any studies performed using Prevora 100 mg/ml Dental

Solution in nursing mothers, it is recommended that Prevora 100 mg/ml Dental Solution should not be applied if the mother is nursing.

4.7 *Effects on ability to drive and use machines*

No studies on the effects on the ability to drive and use machines have been performed.

4.8 *Undesirable effects*

Within each frequency grouping, undesirable effects are presented in order of decreasing seriousness.

	Uncommon ($\geq 1/1,000$ to $< 1/100$)	Not known (cannot be estimated from the available data)
Skin and subcutaneous tissues disorders	Redness and/or temporary stinging sensation of the oral mucosa Objectionable, bitter taste when Prevora 100 mg/ml Dental Solution comes into contact with the saliva or oral mucosa	Immediate hypersensitivity reactions to chlorhexidine (urticaria or anaphylaxis)
General disorders and administration site conditions		Transient tooth sensitivity and loss of taste Discoloration of the teeth and silicate or composite restorations

4.9 *Overdose*

There is no experience with over-dosage with Prevora 100 mg/ml Dental Solution. Consequently, the signs and symptoms have not been identified. If overdose should occur, treat symptomatically.

5. PHARMACOLOGICAL PROPERTIES

5.1 *Pharmacodynamic properties*

Pharmacotherapeutic group: A01AB Anti-infectives and antiseptics for local oral treatment.

ATC Code: A01AB03 chlorhexidine.

Chlorhexidine is effective against a wide range of important oral microorganisms associated with dental caries. Chlorhexidine in the drug product has been found at bactericidal levels to *Streptococcus mutans* for between 24 hours and 48 hours on the surface of adult dental patients after its application, as measured by HPLC.

There have been no published reports of permanent resistance by *Streptococcus mutans* to the repeated use of chlorhexidine for up to 2 years **and no significant resistance to *Streptococcus mutans* or opportunistic infections with *Candida albicans* were observed after treatment with Prevora over one year in adult patients.** The cumulative monthly mean dose of chlorhexidine delivered by Prevora 100 mg/ml Dental Solution is approximately equal to that of 1.0% w/w chlorhexidine dental gel and approximately half that of 0.2% w/v chlorhexidine oral rinse.

5.2 *Pharmacokinetic properties*

Chlorhexidine binds strongly to the oral mucosa and the dentition and thus has very poor systemic absorption. No detectable blood levels of chlorhexidine have been found after oral use.

5.3 *Preclinical safety data*

Preclinical data reveal no special hazard for humans.

6. PHARMACEUTICAL PARTICULARS

6.1 *List of excipients*

Stage 1 chlorhexidine coating:

Sumatra benzoin

Ethanol

Stage 2 sealant coating:

Ammonio methacrylate copolymer dispersion, Type B

Triethyl citrate

6.2 *Incompatibilities*

In the absence of compatibility studies, this topical medicinal product must not be mixed with other topical medicinal products.

6.3 *Shelf life*

18 months.

Discard any remaining solution immediately after use.

6.4 *Special precautions for storage*

Store in refrigerator (2° to 8°C)

6.5 *Nature and contents of container*

Prevora 100 mg/ml Dental Solution contains:

Stage 1 chlorhexidine coating:

Chlorhexidine diacetate

Sumatra benzoin

Ethanol

Stage 2 sealant coating:

Ammonio methacrylate copolymer dispersion, Type B

Triethyl citrate

One treatment box of Prevora Dental solution contains 6 Type 1 glass vials of Stage 1 chlorhexidine coating along with 6 Type 1 glass vials of Stage 2 sealant coating.

6.6 *Special precautions for disposal and other handling*

Step #1. Preparation: Ensure that the dentition contains no open caries lesions or restorations with imperfect margins. Prepare for the application with a tray (Figure 2), consisting of cotton rolls, cotton pellets or fine brushes, forceps, air syringe, and a vial of Stage 1 chlorhexidine coating and a vial of Stage 2 sealant coating.



Figure 2: Tray set up for Stage

One Stage 1 chlorhexidine coating and Stage 2 sealant coating treatments.

Step #2. Prophylaxis: Give a rubber cup prophylaxis using flour of pumice and water. Avoid using a non-oil based prophylactic paste.

Step #3. Floss: Thoroughly rinse and floss the patient's teeth with un-waxed floss to remove pumice and residual dental plaque. Ensure the cleanliness of the distal surface of the last tooth in each arch by wiping it with a cotton pellet held in a pair of forceps.

Step #4. Isolate one quadrant: Isolate one quadrant of the dentition with cotton rolls and a saliva ejector.

Step #5. Dry teeth: Dry all teeth in that quadrant with an air syringe.

Step #6. Apply Stage 1 chlorhexidine coating inter-proximally: Using a cotton pellet held in forceps, or a fine brush suitable for reaching inter-proximal areas, apply Stage 1 Chlorhexidine coating to the inter-proximal areas of all posterior teeth in the quadrant, ensuring not to apply the coating to the soft tissues. Then dry these tooth surfaces with an air syringe.



Figure 3: Apply Prevora Stage 1 chlorhexidine coating with a fine brush to the gingival margin, followed by Prevora Stage 2 sealant coating using the same techniques.

Step #7. Apply Stage 1 chlorhexidine coating to other tooth surfaces: Apply this coating to all other tooth surfaces (Figure 3) in this same quadrant, and then air dry. Be careful to avoid applying Stage 1 chlorhexidine coating to the soft tissues as the patient may experience stinging or burning of the gums or tongue.

Step #8. Apply Stage 2 sealant coating: Apply this second coating (with white cap), using a second cotton pellet or with another fine brush, to this same quadrant. Then dry this second coating with an air syringe.

Step #9. Repeat coating of other quadrants: Repeat steps 4 through 8 in the remaining quadrants of the dentition.

Step #10. Advise the patient:

Instruct the patient:

- The dried Prevora coating will begin coming off the teeth during the next meal.
- To avoid eating hard foods (e.g. meat, apples) for at least 4 hours after treatment.
- Do not chew gum for at least 24 hours.
- Do not brush his/her teeth for 24 hours after treatment. Then to resume brushing with a new brush 2 to 3 times daily with fluoride tooth paste.
- Do not floss for 3 days after treatment. Then to resume flossing daily.
- If dentures are worn, clean and disinfect these dentures regularly prior to use. Disinfect using soap and warm water.

Step #11. Schedule repeat treatments: Repeat this initial Prevora application every week for 3 more weeks after the initial application, followed by a single application at six months and thereafter according to clinical judgment.

Instruments and clothing in contact with Stage 1 chlorhexidine coating may be cleaned with alcohol. Instruments and clothing in contact with Stage 2 sealant coating may be cleaned with water.

Any unused product or waste material should be disposed of in accordance with local requirements.

7. MARKETING AUTHORISATION HOLDER

CHX Technologies Europe Limited
Guinness Enterprise Centre
Taylor's Lane
Dublin 8
Ireland

8. MARKETING AUTHORISATION NUMBER(S)

PA 1205/1/1

9. DATE OF FIRST AUTHORISATION/RENEWAL OF THE AUTHORISATION

Date of First Authorisation: 5th May 2006.

10. DATE OF REVISION OF THE TEXT

September 2010

Detailed information on this medicinal product is available on the website of

LABELLING

The valid labelling is the final version achieved during the Coordination group procedure:

**PARTICULARS TO APPEAR ON THE OUTER PACKAGING
OUTER CARTON**

1. NAME OF THE MEDICINAL PRODUCT

Prevora 100 mg/ml Dental Solution
Chlorhexidine diacetate

2. STATEMENT OF ACTIVE SUBSTANCE(S)

Each 1 ml of Prevora Dental Solution (Stage 1 chlorhexidine coating) contains chlorhexidine diacetate 100 mg.

3. LIST OF EXCIPIENTS

Stage 1: sumatra benzoin and ethanol
Stage 2: ammonio methacrylate copolymer dispersion, Type B and triethyl citrate

4. PHARMACEUTICAL FORM AND CONTENTS

Dental solution
Treatment sets consist of:
6 vials of Stage 1 chlorhexidine coating in a 2 ml Type 1 glass vial
6 vials of Stage 2 sealant coating in a 2 ml Type 1 glass vial

5. METHOD AND ROUTE(S) OF ADMINISTRATION

Read the package leaflet before use. For external (oral) topical use only.

**6. SPECIAL WARNING THAT THE MEDICINAL PRODUCT MUST BE STORED OUT
OF THE REACH AND SIGHT OF CHILDREN**

Keep out of the reach and sight of children.

7. OTHER SPECIAL WARNING(S), IF NECESSARY

Not applicable

8. EXPIRY DATE

EXP

9. SPECIAL STORAGE CONDITIONS

Store upright in a refrigerator at 2° - 8° C.

**10. SPECIAL PRECAUTIONS FOR DISPOSAL OF UNUSED MEDICINAL PRODUCTS
OR WASTE MATERIALS DERIVED FROM SUCH MEDICINAL PRODUCTS, IF
APPROPRIATE**

11. NAME AND ADDRESS OF THE MARKETING AUTHORISATION HOLDER

CHX Technologies Europe Limited
Guinness Enterprise Centre
Taylor's Lane
Dublin 8, Ireland

12. MARKETING AUTHORISATION NUMBER(S)

<[To be completed nationally]>

13. BATCH NUMBER

Lot

14. GENERAL CLASSIFICATION FOR SUPPLY

<[To be completed nationally]>

15. INSTRUCTIONS ON USE

Read the package leaflet before use.

16. INFORMATION IN BRAILLE

<Justification for not including Braille accepted>

MINIMUM PARTICULARS TO APPEAR ON SMALL IMMEDIATE PACKAGING UNITS

VIAL – 2ml

1. NAME OF THE MEDICINAL PRODUCT AND ROUTE(S) OF ADMINISTRATION

Stage 1 Chlorhexidine coating
100 mg/ml Dental solution, chlorhexidine diacetate

Stage 2 Sealant coating

2. METHOD OF ADMINISTRATION

Dental use

3. EXPIRY DATE

MM/YYYY

4. BATCH NUMBER

LOT

5. CONTENTS BY WEIGHT, BY VOLUME OR BY UNIT

Each vial contains 1 ml of stage 1 chlorhexidine coating

6. OTHER

MA #

Read the package leaflet before use.

PACKAGE LEAFLET

The valid package leaflet is the final version achieved during the Coordination group procedure with the following amendments:

PACKAGE LEAFLET: INFORMATION FOR THE USER

Prevora 100 mg/ml Dental Solution
Chlorhexidine diacetate

Read all of this leaflet carefully before you start using this medicine.

- Keep this leaflet. You may need to read it again.
- If you have further questions, please ask your dental professional.
- This coating has been recommended for your teeth by your dental professional and will be applied to your teeth by your dental professional on several occasions.
- If any of the side effects get serious, or if you notice any side effects not listed in this leaflet, please tell your dental professional.

In this leaflet

1. What Prevora is and what it is used for.
2. Before you are treated with Prevora.
3. How you are treated with Prevora.
4. Possible side effects.
5. How to store Prevora.
6. Further information.

1. WHAT PREVORA IS AND WHAT IT IS USED FOR

This dental coating is a topical treatment for the prevention of cavities on the crown and the root of the tooth of **adult patients at high-risk of tooth decay (e.g. dry mouth sufferers or those with a number of cavities when you visit the dentist)**. This coating temporarily covers your teeth to reduce the bacteria on your teeth which cause tooth decay.

Oral hygiene and sugar intake: If you have poor oral hygiene and/or you frequently consume sugars you need to make sure you brush your teeth with fluoridated tooth paste and control your sugar intake, as they are important contributors to the overall success of Prevora.

2. BEFORE YOU ARE TREATED WITH PREVORA

Do not be treated with Prevora

- If you are allergic to chlorhexidine, Sumatra benzoin or ethanol.
- If you are allergic to the ingredients of the Stage 2 sealant, which is a secondary coating applied immediately over Stage 1. The ingredients of this second coating are methacrylate, triethyl citrate and purified water.
- If you have been treated with a fluoride dental varnish in the past 3 days.

Take special care with Prevora

- before undergoing treatment with Prevora, to report your medical condition including asthma, eczema and other allergies, to your dentist.

Taking other medicines

Please inform your dental professional if you are taking or have recently taken any other medicines, including medicines obtained without a prescription.

Pregnancy and breast-feeding

Ask your dentist for advice before taking or receiving any medicine.

Driving and using machines

Prevora has not been shown to affect driving.

Prevora has not been shown to affect the use of machines.

Important information about some of the ingredients of Prevora

This dental coating can cause a temporary irritation or stinging of the gums, lips or tongue and can also have a bitter taste. Chlorhexidine, the active substance in Prevora, can also cause staining of the teeth; however, when chlorhexidine is applied as a topical, temporary dental coating, staining is rare.

3. HOW YOU ARE TREATED WITH PREVORA

This topical, temporary coating on your teeth is applied by your dental professional in a short appointment in the dental office. The dental professional will first clean your teeth, and then apply Stage 1 to all tooth surfaces (Figure 1), followed immediately by a second coating of Stage 2. This second coating temporarily protects Stage 1 from your saliva and from food abrasion.



Figure 1 – Stage 1 followed by a second coating will be applied to your teeth so as to reduce your risk of tooth decay.

At the start of your treatment with Prevora , you will require 4 weekly treatments followed by another single treatment in 6 months. Thereafter, your need for more treatments with Prevora will be evaluated by your dental professional.

You will be told by your Dental Professional that:

- The dried Prevora coating will begin coming off the teeth during the next meal
- You should avoid eating hard foods (e.g. meat, apples) for at least 4 hours after treatment.
- You should not chew gum for at least 24 hours.
- You should not brush your teeth for 24 hours after treatment. Then you resume brushing with a new brush 2 to 3 times daily with fluoride tooth paste.
- You should not floss for 3 days after treatment. Then you resume flossing daily.
- If dentures are worn, clean and disinfect these dentures regularly prior to use. Disinfect using soap and warm water.

Effects when the Prevora treatment is finished

Temporarily, you may have a bitter taste, a sensation of a coating on your teeth and/or a stinging or burning along your gum line or tongue. The stinging will last for a few minutes, the coating for a few hours. The coating will be invisible on your teeth. Chlorhexidine, the active substance in Prevora , can stain the teeth when used in a mouth rinse or gel; however, in controlled clinical studies and dental office use, the occurrence of tooth staining by the Prevora dental coating is rare.

If staining occurs it is not permanent and can largely be eliminated by brushing with a conventional fluoridated toothpaste, once tooth brushing re-commences 24 hours after treatment with Prevora. A professional cleaning will remove any staining but if conducted within 7 days of treatment will also minimize the effect of Prevora. If you receive a professional cleaning, it should be done 7 days after the Prevora treatment.

What to do after you are treated with Prevora

To preserve this coating on your teeth for as long as possible, eat soft foods at your next meal (e.g. soup). Do not eat hard foods (e.g. meat, apples) for at least 4 hours after treatment. Do not chew gum for at least 24 hours.

Do not brush your teeth for 24 hours after this treatment. Then resume brushing with a new tooth brush and brush 2 to 3 times daily with fluoride toothpaste.

Do not floss your teeth for 3 days following this treatment. Then resume flossing daily.

If dentures are worn, clean and disinfect at home prior to use. Disinfect using soap and warm water.

Make sure you receive all the treatments of Prevora, recommended by your dentist.

It is an important contribution to the overall success of this treatment, that you regularly brush your teeth with a fluoridated tooth paste, and that you control your consumption of foods and drinks which have a high amount of sugar and acid.

If you have any further questions on the use of this product, ask your dental professional.

4. POSSIBLE SIDE EFFECTS

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

If you have prolonged stinging or burning of the gums, lips or tongue, you should see your dental professional as soon as possible. If your teeth appear stained after treatment with Prevora, please inform your dental professional.

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

5. HOW TO STORE PREVORA

Keep out of the reach and sight of children.

Do not use Prevora after the expiry date which is stated on the carton and single vial labels after EXP. The expiry date refers to the last day of that month.

Discard any remaining solution immediately after use.

Prevora should be stored in a refrigerator (2°C - 8°C) at your dental professional's office.

Medicines should not be disposed of via wastewater or household waste. These measures will help to protect the environment.

6. FURTHER INFORMATION

What Prevora contains:

Stage 1 chlorhexidine coating:

100 mg/ml Dental Solution

The active substance is Chlorhexidine diacetate.

The other ingredients are Sumatra benzoin and Ethanol

Stage 2 sealant coating:

The ingredients are Ammonio methacrylate copolymer dispersion, Type B and Triethyl citrate

What Prevora looks like and contents of the pack:

Dental Solution

The chlorhexidine coating is a clear, slightly brownish solution with a noticeable scent, free of any floating particles in the solution.

The sealant coating is a milky-white liquid with a faint scent

Each treatment set of Prevora 100 mg/ml Dental Solution consists of:

6 vials of Stage 1 chlorhexidine coating in a 2 ml Type 1 glass vial

6 vials of Stage 2 sealant coating in a 2 ml Type 1 glass vial

Marketing Authorization Holder and Manufacturer

Marketing Authorization Holder:

CHX Technologies Europe Limited

Guinness Enterprise Centre

Taylor's Lane

Dublin 8, Ireland

Tel: (01) 4100600

Manufacturer:

United Drug Plc

United Drug House

Magna Drive, Magna Business Park

Citywest Road

Dublin 24, Ireland

Tel (353) 1 463 2300

This leaflet was last approved in

Detailed information on this medicinal product is available on the website of