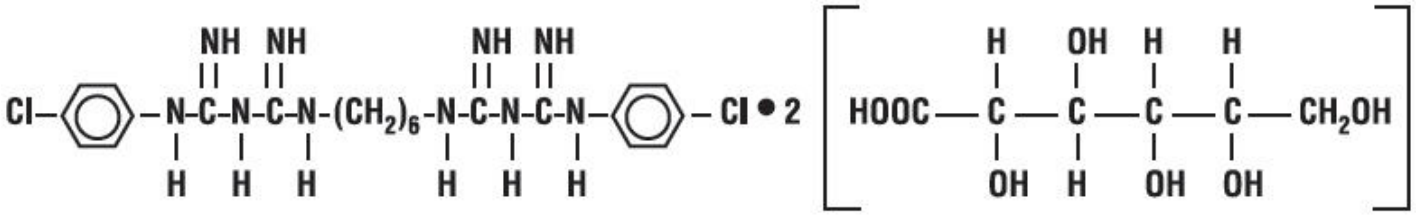
PERIOGARD (CHLORHEXIDINE GLUCONATE)- chlorhexidine gluconate rinse
Colgate Oral Pharmaceuticals, Inc.

Periogard® (Chlorhexidine Gluconate Oral Rinse, 0.12%)

DESCRIPTION

PerioGard® (Chlorhexidine Gluconate Oral Rinse USP, 0.12%) is an oral rinse containing (1,1'-hexamethylene bis [5-(p-chlorophenyl) biguanide] di-D-gluconate) in a base containing water, 11.6% alcohol, glycerin, PEG-40 sorbitan diisostearate, flavor, sodium saccharin and FD&C blue no. 1. PerioGard® (Chlorhexidine Gluconate Oral Rinse USP, 0.12%) is a near-neutral solution (pH range 5-7). Chlorhexidine gluconate is a salt of chlorhexidine and gluconic acid. Its chemical structure is:



CLINICAL PHARMACOLOGY

PerioGard® (Chlorhexidine Gluconate Oral Rinse USP, 0.12%) provides antimicrobial activity during oral rinsing. The clinical significance of chlorhexidine gluconate oral rinse's antimicrobial activities is not clear. Microbiological sampling of plaque has shown a general reduction of counts of certain assayed bacteria, both aerobic and anaerobic, ranging from 54-97% through six months use. Use of chlorhexidine gluconate oral rinse USP, 0.12% in a six month clinical study did not result in any significant changes in bacterial resistance, overgrowth of potentially opportunistic organisms or other adverse changes in the oral microbial ecosystem. Three months after chlorhexidine gluconate oral rinse USP, 0.12% use was discontinued, the number of bacteria in plaque had returned to baseline levels and resistance of plaque bacteria to chlorhexidine gluconate was equal to that at baseline.

PHARMACOKINETICS

Pharmacokinetic studies with chlorhexidine gluconate oral rinse USP, 0.12% indicate approximately 30% of the active ingredient, chlorhexidine gluconate, is retained in the oral cavity following rinsing. This retained drug is slowly released in the oral fluids. Studies conducted on human subjects and animals demonstrate chlorhexidine gluconate is poorly absorbed from the gastrointestinal tract. The mean plasma level of chlorhexidine gluconate reached a peak of 0.206 mcg/g in humans 30 minutes after they ingested a 300-mg dose of the drug. Detectable levels of chlorhexidine gluconate were not present in the plasma of these subjects 12 hours after the compound was administered. Excretion of chlorhexidine gluconate occurred primarily through the feces (~90%). Less than 1% of the chlorhexidine gluconate ingested by these subjects was excreted in the urine.

INDICATIONS AND USAGE

PerioGard® (Chlorhexidine Gluconate Oral Rinse USP, 0.12%) is indicated for use between dental visits as part of a professional program for the treatment of gingivitis as characterized by redness and swelling of the gingivae, including gingival bleeding upon probing. PerioGard® has not been tested among patients with acute necrotizing ulcerative gingivitis (ANUG). For patients having coexisting gingivitis and periodontitis, see PRECAUTIONS.

CONTRAINDICATIONS

PerioGard® should not used by persons who are known to be hypersensitive to chlorhexidine gluconate or other formula ingredients.

WARNINGS

The effect of PerioGard® on periodontitis has not been determined. An increase in supragingival calculus was noted in clinical testing in chlorhexidine gluconate oral rinse USP, 0.12% users compared with control users. It is not known if chlorhexidine gluconate oral rinse use results in an increase in subgingival calculus. Calculus deposits should be removed by a dental prophylaxis at intervals not greater than six months. Anaphylaxis, as well as serious allergic reactions, have been reported during postmarketing use with dental products containing chlorhexidine. SEE CONTRAINDICATIONS.

PRECAUTIONS

General

1. For patients having coexisting gingivitis and periodontitis, the presence or absence of gingival inflammation following treatment with PerioGard® (Chlorhexidine Gluconate Oral Rinse USP, 0.12%) should not be used as a major indicator of underlying periodontitis.
2. PerioGard® can cause staining of oral surfaces, such as tooth surfaces, restorations, and the dorsum of the tongue. Not all patients will experience a visually significant increase in toothstaining. In clinical testing, 56% of the chlorhexidine gluconate oral rinse USP, 0.12% users exhibited a measurable increase in facial anterior stain, compared to 35% of control users after six months; 15% of the chlorhexidine gluconate oral rinse USP, 0.12% users developed what was judged to be heavy stain, compared to 1% of control users after six months. Stain will be more pronounced in patients who have heavier accumulations of unremoved plaque. Stain resulting from the use of PerioGard® does not adversely affect health of the gingivae or other oral tissues. Stain can be removed from most tooth surfaces by conventional professional prophylactic techniques. Additional time may be required to complete the prophylaxis.
   Discretion should be used when prescribing to patients with anterior facial restorations with rough surfaces or margins. If natural stain cannot be removed from these surfaces by a dental prophylaxis, patients should be excluded from the

PerioGard® treatment if permanent discoloration is unacceptable. Stain in these areas may be difficult to remove by dental prophylaxis and on rare occasions may necessitate replacement of these restorations.

3. Some patients may experience an alteration in taste perception while undergoing treatment with chlorhexidine gluconate oral rinse USP, 0.12%. Rare instances of permanent taste alteration following chlorhexidine gluconate oral rinse USP, 0.12% use have been reported via post-marketing product surveillance.

Pregnancy

Teratogenic Effects

Pregnancy Category B

Reproduction studies have been performed in rats and rabbits at chlorhexidine gluconate doses up to 300mg/kg/day and 40mg/kg/day respectively, and have not revealed evidence of harm to fetus. However, adequate and well-controlled studies in pregnant women have not been done. Because animal reproduction studies are not always predictive of human response, this drug should be used during pregnancy only if clearly needed.

Nursing Mothers

It is not known whether this drug is excreted in human milk. Because many drugs are excreted in human milk, caution should be exercised when PerioGard® (Chlorhexidine Gluconate Oral Rinse USP, 0.12%) is administered to nursing women.

In parturition and lactation studies with rats, no evidence of impaired parturition or of toxic effects to suckling pups was observed when chlorhexidine gluconate was administered to dams at doses that were over 100 times greater than that which would result from a person's ingesting 30 ml (2 doses) of PerioGard® per day.

Pediatric Use

Clinical effectiveness and safety of PerioGard® have not been established in children under the age of 18.

Carcinogenesis, Mutagenesis, Impairment of Fertility

In a drinking water study in rats, carcinogenic effects were not observed at doses up to 38mg/kg/day. Mutagenic effects were not observed in two mammalian in vivo mutagenesis studies with chlorhexidine gluconate. The highest doses of chlorhexidine used in a mouse dominant-lethal assay and a hamster cytogenetics test were 1000mg/kg/day and 250mg/kg/day, respectively. No evidence of impaired fertility was observed in rats at doses up to 100mg/kg/day.

ADVERSE REACTIONS

The most common side effects associated with chlorhexidine gluconate oral rinse USP, 0.12% are: 1) an increase in staining of teeth and other oral surfaces; 2) an increase in calculus formation; and 3) an alteration in taste perception; see WARNINGS and PRECAUTIONS. Oral irritation and local allergy-type symptoms have been spontaneously

reported as side effects associated with use of chlorhexidine gluconate oral rinse. The following oral mucosal side effects were reported during placebo-controlled adult clinical trials: aphthous ulcer, grossly obvious gingivitis, trauma, ulceration, erythema, desquamation, coated tongue, keratinization, geographic tongue, mucocele, and short frenum. Each occurred at frequency of less than 1%.

Among post-marketing reports, the most frequently reported oral mucosal symptoms associated with chlorhexidine gluconate oral rinse USP, 0.12% are stomatitis, gingivitis, glossitis, ulcer, dry mouth, hypesthesia, glossal edema, and paresthesia.

Minor irritation and superficial desquamation of the oral mucosa have been noted in patients using chlorhexidine gluconate oral rinse.

There have been cases of parotid gland swelling and inflammation of the salivary glands (sialadenitis) reported in patients using chlorhexidine gluconate oral rinse.

OVERDOSAGE

Ingestion of 1 or 2 ounces of PerioGard® (Chlorhexidine Gluconate Oral Rinse USP, 0.12%) by a small child (~10kg body weight) might result in gastric distress, including nausea, or signs of alcohol intoxication. Medical attention should be sought if more than 4 ounces of PerioGard® Oral Rinse is ingested by a small child or if signs of alcohol intoxication develop.

DOSAGE AND ADMINISTRATION

PerioGard® (Chlorhexidine Gluconate Oral Rinse USP, 0.12%) therapy should be initiated directly following a dental prophylaxis. Patients using PerioGard® should be reevaluated and given a thorough prophylaxis at intervals no longer than six months. Recommended use is twice daily rinsing for 30 seconds, morning and evening after tooth brushing. Usual dosage is 15 ml (marked in cap) of undiluted PerioGard®. Patients should be instructed to not rinse with water or other mouthwashes, brush teeth, or eat immediately after using PerioGard®. PerioGard® is not intended for ingestion and should be expectorated after rinsing.

HOW SUPPLIED

PerioGard® is supplied as a blue liquid in a 1-pint (473 ml) amber plastic bottle with child-resistant dispensing closure. (NDC 0126-0035-16).

Store at 20° to 25°C (68° to 77°F); excursions permitted between 15° to 30°C (59° to 86°F) [see USP Controlled Room Temperature].

R_x Only. Keep out of reach of children.

Revised: September 2017

Distributed By:
Colgate Oral Pharmaceuticals, Inc.,
a subsidiary of Colgate-Palmolive Company
New York, NY 10022 USA

Questions/Comments: 1-800-962-2345
www.colgateprofessional.com

PRINCIPAL DISPLAY PANEL - 473 ml Bottle Label

NDC 0126-0035-16

Colgate® PerioGard®

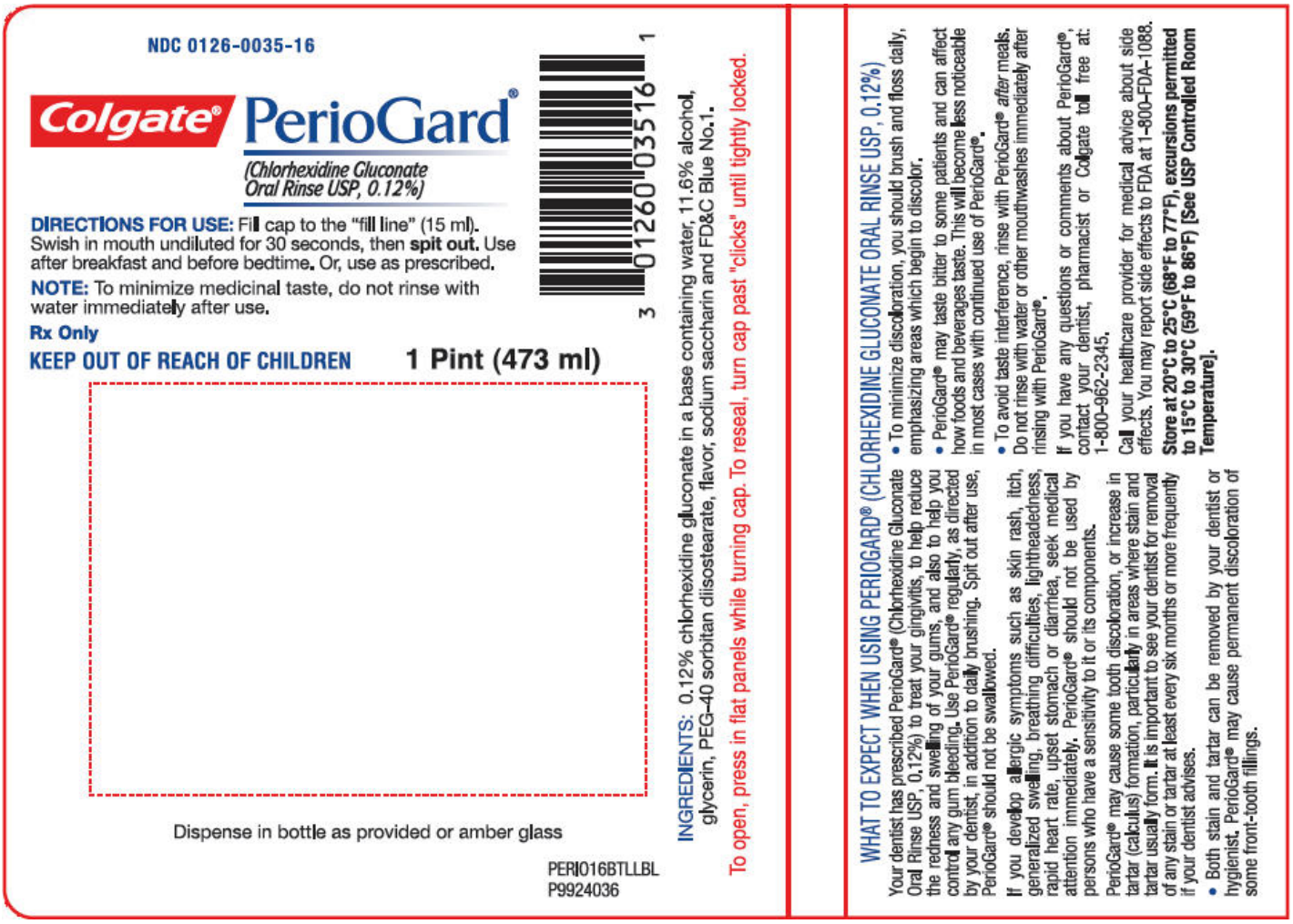
(Chlorhexidine Gluconate
Oral Rinse USP, 0.12%)

DIRECTIONS FOR USE: Fill cap to the "fill line" (15 ml). Swish in mouth undiluted for 30 seconds, then spit out. Use after breakfast and before bedtime. Or, use as prescribed.

NOTE: To minimize medicinal taste, do not rinse with water immediately after use.

Rx Only

KEEP OUT OF REACH OF CHILDREN
1 Pint (473 ml)



PERIOGARD (CHLORHEXIDINE GLUCONATE)

chlorhexidine gluconate rinse

Product Information

Product Type	HUMAN PRESCRIPTION DRUG	**Item Code (Source)**	NDC:0126-0035
Route of Administration	BUCCAL		

Active Ingredient/Active Moiety

Ingredient Name	Basis of Strength	Strength
CHLORHEXIDINE GLUCONATE (UNII: MOR84MUD8E) (CHLORHEXIDINE - UNII:R4KO0DY52L)	CHLORHEXIDINE GLUCONATE	1.2 mg in 1 mL

Inactive Ingredients

Ingredient Name	Strength
WATER (UNII: 059QF0KO0R)	
POLYSORBATE 20 (UNII: 7T1F30V5YH)	
ALCOHOL (UNII: 3K9958V90M)	
SACCHARIN SODIUM (UNII: SB8ZUX40TY)	
GLYCERIN (UNII: PDC6A3C0OX)	
FD&C BLUE NO. 1 (UNII: H3R47K3TBD)	

Product Characteristics

Color	BLUE (Clear, light blue fluid)	**Score**	
Shape		**Size**	
Flavor		**Imprint Code**	
Contains			

Packaging

#	Item Code	Package Description	Marketing Start Date	Marketing End Date
1	NDC:0126-0035-16	473 mL in 1 BOTTLE, PLASTIC; Type 0: Not a Combination Product	01/25/2018	11/30/2023

Marketing Information

Marketing Category	Application Number or Monograph Citation	Marketing Start Date	Marketing End Date
ANDA	ANDA077789	01/25/2018	11/30/2023

Labeler - Colgate Oral Pharmaceuticals, Inc. (968801118)