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(54) Title: TOOTH WHITENING STRIP

(57) Abstract: Described herein are tooth whitening strips comprising a hydratable adhesive film with a first side and a second side, the first side having a granular bleaching ingredient attached thereto in an amount effective to whiten teeth, together with methods of making and using the same.



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TOOTH WHITENING STRIP

BACKGROUND

[0001] There exists a need for whitening strips suitable for home use, having reduced total levels of peroxide, yet providing enhanced whitening activity.

SUMMARY

[0002] Some embodiments of the present invention provide whitening strips that deliver a solid peroxide material directly to the teeth, without substantial dilution from formulation excipients, thereby permitting enhanced bleaching with lower total amounts of peroxide. In some embodiments, the strips comprise an adhesive film, which hydrates in water or saliva to stick to teeth. In further embodiments, the strip is applied in such a way to ensure that the bleaching ingredient is placed directly on the teeth (that is, between the teeth and the adhesive layer), permitting the granules to release peroxide by rapidly dissolving in oral cavity environment. The bleaching ingredient can be optionally coated by or incorporated within a matrix comprising a quickly dissolving material, such as hydroxypropylmethyl cellulose, cornstarch or gum arabic.

[0003] In some embodiments, the strip further comprises a perhydrolase (e.g., an enzyme capable of catalyzing the reaction of carboxylic acid and hydrogen peroxide to form a peracid) and a carboxyl donor, e.g., selected from carboxylic acids and acyl compounds, wherein the carboxyl donor reacts with the peroxide source in the strip in the presence of the perhydrolase to form a peracid, which further enhances the bleaching action of the strip. In yet other embodiments, the strips comprise a peroxygen compound in granular form and an orally acceptable ketone which react to provide a dioxirane, thereby enhancing the bleaching action of the strip.

[0004] Further areas of applicability of the present invention will become apparent from the detailed description provided hereinafter. It should be understood that the detailed description and specific examples, while indicating the preferred embodiments of the invention, are intended for purposes of illustration only and are not intended to limit the scope of the invention.

DETAILED DESCRIPTION

[0005] As used throughout, ranges are used as shorthand for describing each and every value that is within the range. Any value within the range can be selected as the terminus of the range.

[0006] All references cited herein are hereby incorporated by reference in their entireties.

[0007] In the event of a conflict in a definition in the present disclosure and that of a cited reference, the present disclosure controls.

[0008] Exemplary embodiments of the invention include for example tooth whitening strips and methods of whitening teeth, e.g.:

1. A tooth whitening strip (Strip 1) comprising a hydratable adhesive film with a first side and a second side, the first side having a granular bleaching ingredient attached thereto in an amount effective to whiten teeth; for example,
 - 1.1. Strip 1 further comprising a backing layer. In some embodiments, the backing layer controls dissolution of the hydratable adhesive film.
 - 1.2. Strip 1 or 1.1 wherein the granular bleaching ingredient is coated with or incorporated within a matrix comprising a quickly dissolving material, e.g., cornstarch or gum arabic. In some embodiments, the matrix dissolves more quickly than the hydratable adhesive film. In some embodiments, the matrix dissolves at a rate substantially similar to the dissolution rate of the hydratable adhesive film.
 - 1.3. Any of the foregoing strips wherein the granular bleaching ingredient is selected from solid peroxides and solid peroxide donors, e.g., selected from peroxide salts or complexes (e.g., such as peroxyphosphate, peroxy carbonate, perborate, peroxy silicate, or persulfate salts; for example calcium peroxyphosphate, sodium perborate, sodium carbonate peroxide, sodium peroxyphosphate, sodium peroxydisulfate, and potassium persulfate); hypochlorites; urea peroxide; hydrogen peroxide polymer complexes such as hydrogen peroxide-polyvinyl pyrrolidone polymer complexes; metal peroxides e.g. zinc peroxide and calcium peroxide; peracids, e.g., 6-phthalimodoperoxyhexanoic acid (P.A.P.); and combinations thereof.
 - 1.4. Any of the foregoing strips wherein the granular bleaching ingredient comprises urea peroxide.

- 1.5. Any of the foregoing strips where the particle size (D50) of the granular bleaching ingredient is from about 10 to about 500 microns, e.g. from about 60 to about 150 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 15 to about 450 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 20 to about 400 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 25 to about 350 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 30 to about 300 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 35 to about 250 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 40 to about 225 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 45 to about 200 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 50 to about 175 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 55 to about 160 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 65 to about 145 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 70 to about 140 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 75 to about 135 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 80 to about 125 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 85 to about 120 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 90 to about 110 microns. In some embodiments, the particle size (D50) of the granular bleaching ingredient is from about 95 to about 100 microns.
- 1.6. Any of the foregoing strips wherein the granular bleaching ingredient comprises about 1.0% or less, e.g. from about 0.01 to about 1.0%, e.g. from about 0.2 to about 0.8%, of the total weight of the hydratable adhesive film and a granular bleaching ingredient attached thereto.

- 1.7. Any of the foregoing strips wherein the granular bleaching ingredient comprises about 0.1% or less, e.g. from about 0.01 to about 0.1%, e.g. from about 0.02 to about 0.08%, of the total weight of the hydratable adhesive film and a granular bleaching ingredient attached thereto.
- 1.8. Any of the foregoing strips wherein the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.001 to about 1 mg/ cm², e.g., from about 0.001 to about 0.1 mg/ cm², for example from about 0.005 to about 0.015 mg/ cm².
- 1.9. In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.01 to about 0.75 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.05 to about 0.7 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.075 to about 0.65 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.1 to about 0.6 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.15 to about 0.55 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.2 to about 0.5 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.25 to about 0.45 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.3 to about 0.4 mg/ cm².
- 1.10. In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.005 to about 0.1 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.0075 to about 0.095 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.01 to about 0.09 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the

hydratable adhesive film is from about 0.015 to about 0.085 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.02 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.025 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.03 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.035 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.04 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.045 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.05 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.055 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.06 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.065 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.07 to about 0.08 mg/ cm².

- 1.11. In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.075 to about 0.08 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is about 0.078 mg/ cm².
- 1.12. In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.005 to about 0.25 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.005 to about 0.1 mg/ cm². In some

embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.005 to about 0.075 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.005 to about 0.05 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.0075 to about 0.04 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.009 to about 0.035 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.01 to about 0.03 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.01 to about 0.025 mg/ cm². In some embodiments, the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.01 to about 0.02 mg/ cm².

- 1.13. Any of the foregoing strips further comprising (i) a perhydrolase (e.g., an enzyme capable of catalyzing the reaction of carboxylic acid and hydrogen peroxide to form a peracid), for example, a perhydrolase comprising a Ser-His-Asp catalytic triad, e.g., derived from a lipase, serine hydrolase or carbohydrate esterase, and (ii) a carboxyl donor, e.g., selected from carboxylic acids and acyl compounds, wherein upon use, the peroxide released by the granular bleaching ingredient reacts with the carboxyl donor in the presence of the perhydrolase to form a peracid.
- 1.14. The foregoing strip wherein the carboxyl donor is selected from (i) one or more C₂₋₁₈ carboxylic acids, e.g. C₂₋₆ carboxylic acids (e.g., acetic acid), including lower linear or branched alkyl carboxylic acids, optionally substituted with hydroxyl and/or C₁₋₄ alkoxy; (ii) one or more hydrolysable and acceptable esters thereof (e.g. mono-, di-, and tri-glycerides and acylated saccharides) and (iii) mixtures thereof.
- 1.15. The foregoing strip wherein the carboxyl donor is selected from 1,2,3-triacetoxyp propane (sometimes referred to herein as triacetin or glycerin triacetate) and acylated saccharides, e.g. acetylated saccharides.
- 1.16. Any of the foregoing strips comprising a carboxyl donor which comprises an ester compound having solubility in water of at least 5 ppm at 25 °C.

- 1.17. Any of the foregoing strips comprising an orally acceptable ketone, e.g., a C₃₋₈ alkyl ketone compound, for example methyl ethyl ketone, wherein the ketone is oxidized to the corresponding dioxirane when the strip is used.
- 1.18. Any of the foregoing strips which comprise a peracid or which generates a peracid upon use, e.g., wherein the peracid is selected from peracetic acid or phthalimidoperoxyhexanoic acid (P.A.P.).
- 1.19. Any of the foregoing strips wherein the ingredients are present in amounts sufficient to provide, upon mixing, a bleaching agent in an amount and concentration effective to whiten teeth.
- 1.20. Any of the foregoing strips wherein the hydratable adhesive film comprises one or more water-soluble polymers selected from hydrophilic cellulose ethers (e.g. carboxymethyl cellulose, hydroxypropyl cellulose, hydroxypropyl methylcellulose,), polyvinyl acetates, carbomers (e.g., Carbopol 971P), polysaccharide gums (e.g. xanthan gum), modified food starches, gelatin (e.g. animal or fish-based gelatin), cross-linked carboxyvinyl copolymers, cross-linked polyvinylpyrrolidones, polyethylene oxide (ak.a. Polyox), polyacrylic acids and polyacrylates, polyvinyl alcohols, alginate, casein, pullulan, and combinations thereof.
- 1.21. Any of the foregoing strips wherein the hydratable adhesive film comprises one or more water-soluble polymers selected from hydrophilic cellulose ethers (e.g. hydroxypropylmethyl cellulose), polyvinyl acetates, and carbomers (e.g., Carbopol 971P), and combinations thereof.
- 1.22. Any of the foregoing strips wherein the hydratable adhesive film comprises hydroxypropylmethyl cellulose, polyvinyl acetates, and a carbomer, e.g., in a dry weight ratio of 10 to 20 HPMC : 2 to 10 PVAc : 1 carbomer.
- 1.23. Any of the foregoing strips wherein the hydratable adhesive film further comprises a plasticizer, e.g. propylene glycol.
- 1.24. Any of the foregoing strips wherein the first side of the hydratable adhesive film is covered by a protective cover prior to use.
- 1.25. Any of the foregoing strips wherein the hydratable adhesive film is substantially dry prior to application.

- 1.26. Any of the foregoing strips wherein the thickness of the hydratable adhesive film is from about 0.5 to about 15 mil, wherein 1 mil = 0.001 inches.
- 1.27. Any of the foregoing strips wherein the approximate overall dimensions are from about 3 to about 9 cm long X from about 0.5 to about 2.5 cm wide X from about 0.5 to about 15 mil thick, for example a strip wherein the surface area of one side is from about 5 to about 15 cm², e.g., from about 10 to about 12 cm².
- 1.28. Any of the foregoing strips wherein the approximate overall dimensions are from about 5 to about 7 cm long X from about 1.5 to about 2.5 cm wide X from about 2 to about 7 mil thick, for example a strip wherein the surface area of one side is from about 7.5 to about 16 cm², e.g. from about 10 to about 12 cm².
2. Some embodiments of the present invention provide a method of whitening teeth comprising applying the first side of a strip as hereinbefore described, e.g. Strip 1 et seq. directly to the teeth, and leaving it on for a sufficient time, e.g., at least 5 minutes, for example 10-30 minutes, to whiten the teeth.
3. Other embodiments provide a method of making a strip for tooth whitening, e.g., a strip as hereinbefore described, according to Strip 1 et seq., comprising providing a semi-dry hydratable adhesive film, e.g., as hereinbefore described, e.g., which film has been cast from water and not fully dried, or which film has been moistened, adding to one surface of the film granules of a granular bleaching ingredient, e.g., as hereinbefore described, and drying the film with the granules added to one surface.

[0009] For example, the strips may be made by first making the hydratable adhesive film using conventional means, then adding the granulated whitening ingredient to one surface. The hydratable adhesive film strips can be cast from water in a variety of ways known in the art, such as by extrusion, or by casting from a water suspension (for example at a solids level of 10-30%) onto a heated belt, from which the water is evaporated. Alternatively, the film is dried, but then remoistened. The granules can be added to the surface of this film while the film is semi-dry, i.e. just moist enough to be tacky, so that the granules stick to the surface of the film. Alternatively, the granules themselves can be moistened to improve adhesion, either as the primary means or in conjunction with one of the aforementioned approaches. Once the film is fully dry and cooled to room temperature, the granules continue to adhere to the surface of the film. Prior to use, therefore, the hydratable adhesive film and the strip as a whole are substantially dry. Because

the peroxide is on the surface of the film only, a relatively small quantity of granules are required to provide an effective concentration at the surface. For 0.1 % hydrogen peroxide equivalent in the single layer strip, a base film with a surface area of 10 cm² and a dry weight of 7.75 mg/cm² would require only about 0.078 mg of bleach granules.

[0010] When exposed to saliva or other sources of water (such as from a sink), the granules become active. The hydratable adhesive film also is activated and sticks to the teeth effectively.

[0011] In some embodiments, the hydratable adhesive film comprises one or more water-soluble, orally acceptable polymers, e.g. selected from hydrophilic cellulose ethers (e.g. carboxymethyl cellulose, hydroxypropyl cellulose, hydroxypropyl methylcellulose,), polyvinyl acetates, carbomers (e.g., Carbopol 971P), polysaccharide gums (e.g. xanthan gum), modified food starches, gelatin (e.g. animal or fish-based gelatin), cross-linked carboxyvinyl copolymers, cross-linked polyvinylpyrrolidones, polyethylene oxide (ak.a. Polyox), polyacrylic acids and polyacrylates, polyvinyl alcohols, alginate, casein, pullulan, and combinations thereof. Adhesive gel formulations for use with tooth whitening agents are known in the art, e.g. as described in US Patents 7,862,801; 5,746,598; 6,730,316; 7,128,899. The hydratable adhesive film allows the bleaching agent to stay in contact with the teeth for extended periods of time and protects soft tissues.

[0012] Where a second film layer (e.g. carrier or backing layer) is used to protect the hydratable adhesive film from rapid degradation or dissolution, the carrier or backing layer may be made from textiles, cloth, wood composite, resin, elastomer, paper, insoluble or less soluble cellulose derivatives such as ethyl cellulose and cellulose acetate, polyvinyl chloride, wax, ParafilmsTM, polyethylene, polyvinyl alcohol, TeflonTM, polyvinyl chloride, polyvinyl acetate and their derivatives.

[0013] The granular bleaching ingredient may be a solid peroxide or solid peroxide donor or other active oxygen donor, e.g., selected from peroxide salts or complexes (e.g., such as peroxyphosphate, peroxycarbonate, perborate, peroxy silicate, peroxy monosulphate or peroxydisulphate salts; for example calcium peroxyphosphate, sodium perborate, sodium carbonate peroxide, sodium peroxyphosphate, sodium peroxydisulphate, and potassium persulfate), hypochlorites; urea peroxide; hydrogen peroxide polymer complexes such as hydrogen peroxide-polyvinyl pyrrolidone polymer complexes, and metal peroxides e.g. zinc peroxide and calcium peroxide; a solid peracid, e.g., phthalimidoperoxyhexanoic acid (P.A.P.);

and combinations thereof. In particular embodiments, the granular bleaching ingredient is urea peroxide.

[0014] Peroxycarboxylic acids (“peracids”) useful in the present invention are known as effective antimicrobial and bleaching agents. U.S. Patent 5,302,375 to Viscio, D., discloses oral compositions for whitening comprising peracetic acid dissolved in a vehicle, wherein the peracetic acid is generated within the vehicle in situ by combining water, acetylsalicylic acid, and a water soluble alkali metal percarbonate. U.S. Patent 5,279,816 to Church et al. discloses the use of a composition comprising peracetic acid to whiten stained or discolored teeth. U.S. Patents 6,221,341 and 7,189,385 to Montgomery, R., disclose peroxy acid tooth-whitening compositions suitable for use in a method to whiten teeth. More specifically, a peracetic acid composition may be produced by combining a hydrogen peroxide precursor, an acetic acid ester of glycerin, and water to generate, via chemical perhydrolysis, peracetic acid. Perhydrolase enzymes are also reported, for example, in WO 2005/056782.

[0015] Many hydrolases and esterases, for example, lipases, serine hydrolases and carbohydrate esterases, catalyze perhydrolysis, the reversible formation of peracids from carboxylic acids and hydrogen peroxide. Perhydrolases, esterases, and lipases generally contain a catalytic triad consisting of a serine (Ser), a glutamate (Glu) or aspartate (Asp), and a histidine (His). Many perhydrolases (e.g. metal-free haloperoxidases) contain a Ser-His-Asp catalytic triad and catalyze the reversible formation of peracid from hydrogen peroxide and carboxylic acids. Without being bound by theory, it is believed that perhydrolysis takes place with an esterase-like mechanism in which a carboxylic acid reacts with the active site serine to form an acyl enzyme intermediate, which then reacts with hydrogen peroxide to form a peracid.

[0016] Numerous perhydrolases have been described in the art. The inclusion of specific variant subtilisin Carlsberg proteases having perhydrolytic activity in a body care product is disclosed in U.S. Patent 7,510,859 to Wieland et al. Perhydrolytic enzymes beyond the specific variant proteases are not described nor are there any working examples demonstrating the enzymatic production of peracid as a personal care benefit agent.

[0017] Carboxyl donors for use in the present invention, e.g., to form peracids upon reaction with peroxide, are selected from one or more of (i) C₂₋₁₈ carboxylic acids, e.g. C₂₋₆ carboxylic acids (e.g., acetic acid), including lower linear or branched alkyl carboxylic acids, optionally substituted with hydroxyl and/or C₁₋₄ alkoxy; (ii) hydrolysable and acceptable esters thereof

(e.g. mono-, di-, and tri-glycerides and acylated saccharides) and (iii) mixtures thereof. For example, carboxyl donors include 1,2,3-triacetoxyp propane (sometimes referred to herein as triacetin or glycerin triacetate) and acylated saccharides, e.g. acetylated saccharides. In a particular embodiment, esters for this use may, for example, be esters having solubility in water of at least 5 ppm at 25 °C.

[0018] The carboxyl donors and/or enzymes may optionally be encapsulated. There are a variety of encapsulation options well-known to the art, both natural and synthetic. Modified celluloses, modified starches and gum arabic are particularly well-suited since they are food grade, relatively inexpensive, quick to dissolve, and can adsorb fairly high levels of liquid oils.

[0019] All ingredients for use in the strips described herein should be orally acceptable. By “orally acceptable” as the term is used herein is meant an ingredient which is present in a strip as described in an amount and form which does not render the strip unsafe for use in the oral cavity.

[0020] Unless otherwise specified, all percentages and amounts expressed herein and elsewhere in the specification should be understood to refer to percentages by weight. The amounts given are based on the active weight of the material.

EXAMPLES

Example 1

[0021] A strip is prepared as described above, forming the hydratable adhesive film and then while the film is still tacky, adding the granulated whitening agent to the surface of one side, using the ingredients in Table 1. The strip will erode slowly in the mouth upon application, and so does not need to be removed.

TABLE 1

Ingredients in film	Weight % of dry film
Hydroxypropylmethylcellulose (HPMC)	59
Polyvinyl acetate (PVAc)	30
Carbomer (CARBOPOL® 971)	5
Propylene glycol	5
Titanium dioxide	1
Total	100

Ingredients in bleach granules	Concentration (%)
Urea peroxide	100

Example 2

[0022] A strip is prepared as described above, forming the hydratable adhesive film and then while the strip is still tacky, adding the granulated whitening agent to one side and the protective backing layer to the other side, using the ingredients in Table 2. Because the backing layer will not dissolve, the user should remove it after a sufficient period has passed to permit whitening to take place, typically about 10-30 minutes. The two layers can also be produced simultaneously by extrusion or solvent-based casting, then the granulated whitening agent can be added to the surface of the hydratable adhesive film.

TABLE 2

Ingredients in backing layer	Weight % of dry strip
Ethyl cellulose	94
Propylene glycol	5
Titanium dioxide	1
Ingredients in hydratable adhesive strip	Weight % of dry strip
Hydroxypropylmethylcellulose (HPMC)	69
Polyvinyl acetate (PVAc)	15
Carbomer (CARBOPOL® 971)	5
Propylene glycol	5
Titanium dioxide	1
Total	100
Ingredients in bleach granules	Concentration (%)
Urea peroxide	100

[0023] As those skilled in the art will appreciate, numerous changes and modifications may be made to the embodiments described herein without departing from the spirit of the invention. It is intended that all such variations fall within the scope of the appended claims.

CLAIMS

1. A tooth whitening strip comprising a hydratable adhesive film with a first side and a second side, the first side having a granular bleaching ingredient attached thereto in an amount effective to whiten teeth.
2. The tooth whitening strip according to claim 1, further comprising a backing layer.
3. The tooth whitening strip according to any of the foregoing claims wherein the granular bleaching ingredient is coated with or incorporated within a matrix comprising a quickly dissolving material.
4. The tooth whitening strip according to any of the foregoing claims wherein the granular bleaching ingredient is selected from solid peroxides, solid peroxide donors, and solid oxygen donors.
5. The tooth whitening strip according to any of the foregoing claims wherein the granular bleaching ingredient is selected from peroxide salts or complexes, peroxyphosphate, peroxycarbonate, perborate, peroxysilicate, persulphate salts, calcium peroxyphosphate, sodium perborate, sodium carbonate peroxide, sodium peroxyphosphate, sodium peroxydisulphate, potassium persulfate, hypochlorites, urea peroxide, hydrogen peroxide polymer complexes, hydrogen peroxide-polyvinyl pyrrolidone polymer complexes, metal peroxides, zinc peroxide, calcium peroxide, peracids, phthalimidoperoxyhexanoic acid (P.A.P.), and combinations thereof.
6. The tooth whitening strip according to any of the foregoing claims wherein the granular bleaching ingredient comprises urea peroxide.
7. The tooth whitening strip according to any of the foregoing claims wherein the particle size (D50) of the granular bleaching ingredient is from about 10 to about 500 microns.
8. The tooth whitening strip according to any of the foregoing claims wherein the granular bleaching ingredient comprises 1.0% or less of the total weight of the hydratable adhesive film and a granular bleaching ingredient attached thereto.
9. The tooth whitening strip according to any of the foregoing claims wherein the granular bleaching ingredient comprises 0.1% or less of the total weight of the hydratable adhesive film and a granular bleaching ingredient attached thereto.

10. The tooth whitening strip according to any of the foregoing claims wherein the amount of granular bleaching agent on the first side of the hydratable adhesive film is from about 0.001 to about 1 mg/ cm².
11. The tooth whitening strip according to any of the foregoing claims further comprising (i) a perhydrolase and (ii) a carboxyl donor, wherein upon use, the peroxide released by the granular bleaching ingredient reacts with the carboxyl donor in the presence of the perhydrolase to form a peracid.
12. The tooth whitening strip of claim 10 wherein the carboxyl donor is selected from (i) one or more C₂₋₁₈ carboxylic acids, optionally substituted with hydroxyl and/or C₁₋₄ alkoxy; (ii) one or more hydrolysable and acceptable esters thereof, and (iii) mixtures thereof.
13. The tooth whitening strip of claim 11 wherein the carboxyl donor is 1,2,3-triacetoxyp propane.
14. The tooth whitening strip according to any of the foregoing claims further comprising an orally acceptable ketone, wherein the ketone is oxidized to the corresponding dioxirane when the strip is used.
15. The tooth whitening strip according to any of the foregoing claims which comprises a peracid or which generates a peracid upon use.
16. The tooth whitening strip according to any of the foregoing claims wherein the ingredients are present in amounts sufficient to provide, upon mixing, a bleaching agent in an amount and concentration effective to whiten teeth.
17. The tooth whitening strip according to any of the foregoing claims wherein the hydratable adhesive film comprises one or more water-soluble polymers selected from hydrophilic cellulose ethers, carboxymethyl cellulose, hydroxypropyl cellulose, hydroxypropyl methylcellulose, polyvinyl acetates, carbomers, polysaccharide gums, xanthan gum, modified food starches, gelatin, animal or fish-based gelatin, cross-linked carboxyvinyl copolymers, cross-linked polyvinylpyrrolidones, polyethylene oxide, polyacrylic acids, polyacrylates, polyvinyl alcohols, alginate, casein, pullulan, and combinations thereof.
18. The tooth whitening strip according to any of the foregoing claims wherein the hydratable adhesive film comprises one or more water-soluble polymers selected from hydrophilic cellulose ethers, polyvinyl acetates, carbomers, and combinations thereof.

19. The tooth whitening strip according to any of the foregoing claims wherein the hydratable adhesive film comprises hydroxypropylmethyl cellulose (HPMC), polyvinyl acetates (PVAc), and a carbomer in a dry weight ratio for HPMC : PVAc : carbomer of 10- 20 : 2- 10 : 1.
20. The tooth whitening strip according to any of the foregoing claims wherein the hydratable adhesive film further comprises a plasticizer.
21. The tooth whitening strip according to any of the foregoing claims further comprising propylene glycol.
22. A method of whitening teeth comprising applying a strip according to any of the foregoing claims directly to the teeth, and leaving it on for a sufficient time to whiten the teeth.
23. A method of making a strip according to any of the foregoing comprising providing a semi-dry hydratable adhesive film, adding to one surface of the film granules of a granular bleaching ingredient, and drying the film so that the granules adhere to the surface of the film.

INTERNATIONAL SEARCH REPORT

International application No
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A. CLASSIFICATION OF SUBJECT MATTER		
INV. A61K8/02	A61K8/22	A61K8/73 A61K8/81 A61Q11/00
ADD.		
According to International Patent Classification (IPC) or to both national classification and IPC		
B. FIELDS SEARCHED		
Minimum documentation searched (classification system followed by classification symbols) A61Q A61K		
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched		
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) EPO-Internal, WPI Data		
C. DOCUMENTS CONSIDERED TO BE RELEVANT		
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<input checked="" type="checkbox"/> Further documents are listed in the continuation of Box C. <input checked="" type="checkbox"/> See patent family annex.		
* Special categories of cited documents : "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier application or patent but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art "&" document member of the same patent family		
Date of the actual completion of the international search 4 July 2012		Date of mailing of the international search report 16/07/2012
Name and mailing address of the ISA/ European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Fax: (+31-70) 340-3016		Authorized officer Verrucci, Marinella

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