(19) World Intellectual Property Organization International Bureau





(43) International Publication Date 4 January 2001 (04.01.2001)

(10) International Publication Number WO 01/00171 A1

(51) International Patent Classification7: A61K 7/50, 7/00

(21) International Application Number: PCT/US00/17645

(22) International Filing Date: 27 June 2000 (27.06.2000)

(25) Filing Language: English

English (26) Publication Language:

(30) Priority Data:

9915094.8 28 June 1999 (28.06.1999) GB 9915095.5 28 June 1999 (28.06.1999) GB 60/201,912 4 May 2000 (04.05.2000) US

(71) Applicant (for all designated States except US): THE PROCTER & GAMBLE COMPANY [US/US]; One Procter & Gamble Plaza, Cincinnati, OH 45202 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): EVANS, Erica, Louise [GB/GB]; Plough Cottage, Plough House Court, Hungerford, Berkshire RG17 0ED (GB). VANOOST-HUYZE, Kristina, Emma, Inge [BE/GB]; Lynton, Horsell Birsch, Horsell, Woking, Surrey GU21 4XD (GB). SIVIK, Mark, Robert [US/US]; 2434 Sheffield Court, Ft. Mitchell, KY 41017 (US).

- (74) Agents: REED, T., David et al.: The Procter & Gamble Company, 5299 Spring Grove Avenue, Cincinnati, OH 45217-1087 (US).
- (81) Designated States (national): AE, AL, AM, AT, AT (utility model), AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, CZ (utility model), DE, DE (utility model), DK, DK (utility model), DM, EE, EE (utility model), ES, FI, FI (utility model), GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK (utility model), SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

Published:

- With international search report.
- Before the expiration of the time limit for amending the claims and to be republished in the event of receipt of

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: COSMETIC COMPOSITIONS

(57) Abstract: According to the present invention there is provided a cosmetic composition comprising: (a) at least one readily biodegradable quaternary ammonium agent; and (b) at least 5 %, by weight, of humectant. The compositions of the present invention give good skin care benefits, such as good moisturisation, good hydration, good skin feel, good skin softness and/or good skin smoothness, with low levels of negatives such as greasiness, stickiness or tack. In addition, such compositions are readily degraded once used and are consequently kinder to the environment.

COSMETIC COMPOSITIONS

Technical Field

The present invention relates to cosmetic compositions. In particular, the present invention relates to cosmetic compositions that provide good moisturisation, hydration, skin feel, skin softness and/or skin smoothness benefits.

Background to the Invention

Skin is made up of several layers of cells which coat and protect the keratin and collagen fibrous proteins that form the skeleton of its structure. The outermost of these layers, referred to as the stratum corneum, is known to be composed of 25nm protein bundles surrounded by 8nm thick layers. Anionic surfactants and organic solvents typically penetrate the stratum corneum membrane and, by delipidization (i.e. removal of the lipids from the stratum corneum), destroy its integrity. This destruction of the skin surface topography leads to a rough feel and may eventually permit the surfactant or solvent to interact with the keratin, creating irritation. Many people expose their skin to this type of insult every day. In addition, the skin can be damaged by other factors such as exposure to cold air or wind, mechanical abrasion, immersion in water etc. Thus, there exists a need for a way of mitigating or ameliorating this damage.

In the past compositions have been formulated that are claimed to assist the stratum corneum in maintaining its barrier and water-retention functions at optimum performance in spite of deleterious interactions which the skin may encounter in washing, work, and recreation. Desirable properties for such compositions are that they have good skin feel, water retention, moisturisation, absorption, and/or rub-in characteristics. Prior art compositions have attempted to deliver these properties by the use of one or more 'skin benefit agents'. For example, one way of delivering high moisturisation to the skin is to incorporate polyhydric alcohol-like humectant materials such as glycerine into a composition.

Skin compositions with high levels of polyhydric alcohols and therefore high levels of moisturisation, however, are often perceived by the consumer as unpleasant as such compositions can feel sticky when applied to the skin.

There appears to be a direct relationship between the amount of the skin benefit agent and the effectiveness of the composition at delivering benefits to the skin. However, it is also the case that the higher the level of skin benefit agent the higher the risk of the associated negatives. Therefore, to date, it has been necessary to balance the benefits of compositions comprising high levels of the skin benefit agents against the negatives associated with such high levels. Thus, there remains a need for compositions which contain high levels of skin benefit agents, and therefore provide high levels of the associated benefits such as skin feel, skin softness and skin smoothness, but show low levels of the associated negatives such as greasiness, stickiness or tackiness.

Quaternary ammonium agents are also known for use in cosmetic compositions. See, for example, WO-A-99/27904, WO-A-96/32089, and EP-A-789,076. Also, US-A-5,804,205 which discloses skin care compositions which are claimed to provide a high degree of moisturisation without leaving a "tacky" or "sticky" residue. The compositions contain quaternary ammonium compounds having two alkyl groups of 16-22 carbon atoms, humectant and non-irritating hydrophobic microspheres having an average particle size of less than 50 µm. It is claimed that the hydrophobic polymeric microspheres significantly reduce the "tackiness" associated with high humectant levels. However, the quaternary ammonium agents commonly used in the prior art are often not readily biodegradable and can, therefore, be harmful to the environment.

It has now been unexpectedly found that compositions comprising at least one readily biodegradable quaternary ammonium agent and at least 5%, by weight, humectant provide high levels of the associated benefits such as skin feel, skin softness and skin smoothness, but show low levels of the associated negatives such as greasiness, stickiness or tackiness. In addition, such compositions are readily degraded once used and are consequently kinder to the environment.

While not wishing to be bound by theory, it is believed that the quaternary ammonium agents of the present invention can vesiculate the humectants and drive their deposition onto the skin. This results in a smooth and uniform application of the humectant to the skin with minimal tack/stickiness/greasiness. Even when the compositions comprise extremely high percentages of humectant, for example 30% by weight, there are only low levels of the usual negatives. Also, it is believed that the quaternary ammonium agents help reduce the loss of humectants from the skin due to environmental factors such as water or abrasion. Moreover, it is believed that the quaternary ammonium agents themselves deliver skin care benefits, such as good moisturisation, good skin feel, good skin softness.

Summary of the Invention

According to the present invention there is provided a cosmetic composition comprising:

- (a) at least one readily biodegradable quaternary ammonium agent; and
- (b) at least 5%, by weight, of humectant.

The compositions of the present invention give good skin care benefits, such as good moisturisation, good hydration, good skin feel, good skin softness and/or good skin smoothness, with low levels of negatives such as greasiness, stickiness or tack. In addition, such compositions are readily degraded once used and are consequently kinder to the environment.

Detailed Description of the Invention

The compositions of the present invention comprise at least one readily biodegradable quaternary ammonium agent and a humectant. These elements will be described in more detail below.

The present compositions can be used for any suitable purpose. In particular, the present compositions are suitable for topical application to the skin. In particular, the skin care compositions can be in the form of creams, lotions, gels,

and the like. Preferably the cosmetic compositions herein are in the form of an oil-in-water emulsion of one or more oil phases in an aqueous continuous phase, each oil phase comprising a single oily component or a mixture of oily components in miscible or homogeneous form but said different oil phases containing different materials or combinations of materials from each other.

The compositions of the present invention preferably comprise vesicles. Preferably said vesicles comprise the quaternary ammonium compound together with humectant. As used herein the term "vesicle" means one or more bilayers arranged in a closed, usually spherical geometry, said bilayer comprises quaternary ammonium agent as described hereinbelow.

Preferably the compositions of the present invention comprise less than 10%, preferably less than 5%, more preferably less than 3%, even more preferably 0%, by weight, of anionic surfactant.

The compositions of the present invention are preferably formulated so as to have a product viscosity of at least about 1,000 mPa.s and preferably in the range from about 1,000 to about 300,000 mPa.s, more preferably from about 2,500 to about 250,000 mPa.s and especially from about 5,000 to about 200,000 mPa.s (26.8°C, neat, Brookfield DV-II+ Spindle CP52/CP41).

Quaternary Ammonium Agent

The compositions of the present invention must comprise at least one readily biodegradable quaternary ammonium agent. Any readily biodegradable quaternary ammonium agent suitable for use in cosmetic compositions may be used herein. As used herein the term "quaternary ammonium agent' means a compound or mixture of compounds having a quaternary nitrogen atom substituted with one or more, preferably two, moieties containing six or more carbon atoms. As used herein the term "readily biodegradable" means that, according to the OECD 301 test, at least 60% of the compounds mineralises within 28 days. Preferably the quaternary ammonium agents for use herein are selected from those having a quaternary nitrogen substituted with two moieties wherein each moiety comprises ten or more, preferably 12 or more, carbon atoms. Highly preferred quaternary ammonium agents for use herein are

selected from those which are able to form vesicles in polar solvents, as detected by microscopic analysis (polarised light microscopy at a magnification of x60 using a Nikon Eclipse E800 microscope).

Preferably the present compositions comprise at least 0.01%, more preferably at least 0.1%, even more preferably at least 1%, still more preferably at least 3%, by weight, of quaternary ammonium agent.

Preferably the quaternary ammonium agents for use herein are selected from:

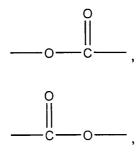
(a) quaternary ammonium compounds according to general formula (I) or (II):

$$\left[(R_5)_{4\text{-m}} - N^+ - \left((CH_2)_n - Q - R_6 \right)_m \right] X^-$$

$$\left[(R_5)_{4\text{-m}} - N^+ - \left((CH_2)_n - CH - CH_2 - Q - R_6 \right)_m \right] X^-$$

$$\left[(II)_{4\text{-m}} - R_6 \right]_{Q - Q}$$

wherein, each R_5 unit is independently selected from hydrogen, branched or straight chain C_1 - C_6 alkyl, branched or straight chain C_1 - C_6 hydroxyalkyl and mixtures thereof, preferably methyl and hydroxyethyl; each R_6 unit is independently linear or branched C_{11} - C_{22} alkyl, linear or branched C_{11} - C_{22} alkenyl, and mixtures thereof; X^- is an anion which is compatible with skin care actives and adjunct ingredients; m is from 1 to 4, preferably 2 and Q is a carbonyl unit selected from:



5

wherein R₇ is hydrogen, C₁-C₄ alkyl, C₁-C₄ hydroxyalkyl, and mixtures thereof.

In the above quaternary ammonium compound example, the unit -QR $_6$ contains a fatty acyl unit which is typically derived from a triglyceride source. The triglyceride source is preferably derived from tallow, partially hydrogenated tallow, lard, partially hydrogenated lard, vegetable oils and/or partially hydrogenated vegetable oils, such as, canola oil, safflower oil, peanut oil, rapeseed oil, sunflower oil, corn oil, soybean oil, tall oil, rice bran oil, etc. and mixtures of these oils.

The counterion, X⁻ in the above compounds, can be any compatible anion, preferably the anion of a strong acid, for example, chloride, bromide, methylsulfate, ethylsulfate, sulfate, nitrate and the like, more preferably chloride or methyl sulfate. The anion can also, but less preferably, carry a double charge in which case X⁻ represents half a group.

The preferred quaternary ammonium compounds of the present invention are the diester and/or diamide Quaternary Ammonium (DEQA) compounds, the diesters and diamides having general formula (II), wherein the carbonyl group Q is selected from:

Tallow, canola and palm oil are convenient and inexpensive sources of fatty acylunits which are suitable for use in the present invention as R_6 units.

The counterion, X⁻, can be chloride, bromide, methylsulfate, formate, sulfate, nitrate, and mixtures thereof. In fact, the anion, X, is merely present as a counterion of the positively charged quaternary ammonium compounds. The scope of this invention is not considered limited to any particular anion.

As used herein, when the diester is specified, it will include the monoester and triester that are normally present as a result of the manufacture process.

(b) quaternary ammonium compounds according to general formula (III) or (IV):

$$R_9$$
 N
 R_{10}
 NH
 C
 R_9
 O
(III)

$$R_9 \xrightarrow{N} \\ R_{10} \xrightarrow{O} C \xrightarrow{C} R_9 \\ 0 \qquad (IV)$$

wherein R_9 is an acyclic aliphatic C_{15} - C_{21} hydrocarbon group and R_{10} is a C_1 - C_6 alkyl or alkylene group.

These ammonium compounds, having a pKa value of not greater than about 4, are able to generate a cationic charge in situ when dispersed in an aqueous solution, providing that the pH of the final composition is not greater than about 6.

(c) quaternary ammonium compounds according to general formula (V) or (VI):

$$\begin{bmatrix} R_9 & & & \\ & & &$$

$$\begin{bmatrix} R_9 & & & \\ & & &$$

wherein R_9 & R_{10} are as specified hereinabove and R_{11} is selected from C_1 - C_4 alkyl and hydroxyalkyl groups.

The counterion, X̄, can be chloride, bromide, methylsulfate, formate, sulfate, nitrate, and mixtures thereof. In fact, the anion, X, is merely present as a counterion of the positively charged quaternary ammonium compounds. The scope of this invention is not considered limited to any particular anion.

(d) quaternary ammonium compounds according to general formula (VII) or (VIII):

$$R_{9} \longrightarrow O \longrightarrow n(H_{2}C) \longrightarrow N \longrightarrow (CH_{2})_{n} \longrightarrow N \longrightarrow R_{9}$$

$$(VIII)$$

$$R_{9} \longrightarrow O \longrightarrow n(H_{2}C) \longrightarrow N \longrightarrow (CH_{2})_{n} \longrightarrow N \longrightarrow R_{9}$$

$$R_{12} \longrightarrow N \longrightarrow R_{9}$$

$$R_{13} \longrightarrow N \longrightarrow R_{9}$$

$$R_{14} \longrightarrow N \longrightarrow R_{9}$$

$$R_{15} \longrightarrow N \longrightarrow R_{15}$$

$$R_{15} \longrightarrow N \longrightarrow R_{15$$

wherein, n is from 1 to 6, R_9 is selected from acyclic aliphatic C_{15} - C_{21} hydrocarbon groups and R_{12} is selected from C_1 - C_4 alkyl and hydroxyalkyl groups.

These ammonium compounds, having a pKa value of not greater than about 4, are able to generate a cationic charge in situ when dispersed in an aqueous solution, providing that the pH of the final composition is not greater than about 6.

The counterion, X, can be chloride, bromide, methylsulfate, formate, sulfate, nitarte, and mixtures thereof. In fact, the anion, X, is merely present as a counterion of the positively charged quaternary ammonium compounds. The scope of this invention is not considered limited to any particular anion.

(e) diquaternary ammonium compounds according to general formula (IX) or (X):

$$\begin{bmatrix} & & & & & \\ & & & \\ & & & & \\ &$$

wherein R_5 , R_6 , Q, n & X^- are as defined hereinabove in relation to general formula (I) and (II) and R_{13} is selected from C_1 - C_6 alkylene groups, preferably an ethylene group.

(f) mixtures of the above quaternary ammonium compounds.

The preferred quaternary ammonium agents for use in the present invention are those described in section (b) hereinabove. In particular, diester and/or diamide quaternary ammonium (DEQA) compounds according to general formula (II) hereinabove are preferred. Preferred diesters for use herein are those according to general formula (II) wherein R_5 , R_6 , and X^* are as defined hereinabove and Q is:

Preferred diamides for use herein are those according to general formula (II) wherein R₅, R₆, and X⁻ are as defined hereinabove and Q is:

Preferred examples of quaternary ammonium compounds suitable for use in the compositions of the present invention are N,N-di(canolyl-oxy-ethyl)-N,N-dimethyl ammonium chloride, N,N- di(canolyl-oxy-ethyl)-N-methyl,N-(2-hydroxyethyl) ammonium methyl sulfate, N,N-di(canolyl-oxy-ethyl)-N-methyl, N-(2-hydroxyethyl) ammonium chloride and mixtures thereof. Particularly preferred for use herein is N,N-di(canolyl-oxy-ethyl)-N-methyl,N-(2-hydroxyethyl) ammonium methyl sulfate.

Although quaternary ammonium compounds are derived from "canolyl" fatty acyl groups are preferred, other suitable examples of quaternary ammonium compounds are derived from fatty acyl groups wherein the term "canolyl" in the above examples is replaced by the terms "tallowyl, cocoyl, palmyl, lauryl, oleyl, ricinoleyl, stearyl, palmityl" which correspond to the triglyceride source from which the fatty acyl units are derived. These alternative fatty acyl sources can comprise either fully saturated, or preferably at least partly unsaturated chains.

Humectant

A second essential element of the compositions of the present invention is that they comprise at least 5%, by weight, of humectant. As used herein the term "humectant" means a substance which provides the skin with water-retention benefits. Preferably, the compositions of the present invention comprise at least 7.5%, more preferably at least 10%, even more preferably at least 15%, even more still preferably at least 30%, by weight of humectant.

Any humectant suitable for use in cosmetic compositions may be used herein. Non-limiting examples of suitable humectants for use in the present invention are described in WO98/22085, WO98/18444 and WO97/01326. Preferably the humectants for use herein are selected from, but not limited to; amino acids and derivatives thereof such as proline and arginine aspartate, 1,3-butylene glycol, propylene glycol and water and codium tomentosum extract, collagen amino acids or peptides, creatinine, diglycerol, biosaccharide gum-1, glucamine salts, glucuronic acid salts, glutamic acid salts, polyethylene glycol ethers of glycerine (e.g. glycereth 20), glycerine, glycerol monopropoxylate, glycogen, hexylene glycol, honey, and extracts or derivatives thereof, hydrogenated starch hydrolysates, hydrolyzed mucopolysaccharides, inositol, keratin amino acids, urea, LAREX A-200 (available from Larex), glycosaminoglycans, methoxy PEG 10, methyl gluceth-10 and -20 (both commercially available from Amerchol located in Edison, NJ), methyl glucose, 3-methyl-1,3-butanediol, N-acetyl glucosamine salts, polyethylene glycol and derivatives thereof (such as PEG 15 butanediol, PEG 4, PEG 5 pentaerythitol, PEG 6, PEG 8, PEG 9), pentaerythitol, 1.2 pentanediol, PPG-1 glyceryl ether, PPG-9, 2-pyrrolidone-5-carboxylic acid and its salts such as glyceryl pca, saccharide isomerate, SEACARE (available from Secma), sericin, silk amino acids, sodium acetylhyaluronate, sodium hyaluronate, sodium poly-aspartate, sodium polyglutamate, sorbeth 20, sorbeth 6, sugar and sugar alcohols and derivatives thereof such as glucose, mannose polyglycerol sorbitol, trehalose, triglycerol, trimethyolpropane, (hydroxymethyl) amino methane salts, and yeast extract, and mixtures thereof.

More preferably, the humectants for use herein are selected from glycerine, urea, butylene glycol, polyethylene glycol and derivatives thereof, and mixtures thereof. Even more preferably, the humectants for use herein are selected from glycerine, urea and mixtures thereof, especially glycerine.

Optional Ingredients

The compositions herein can contain a variety of optional components suitable for rendering the present compositions more cosmetically or aesthetically acceptable or to provide them with additional usage benefits. Such conventional optional ingredients are well-known to those skilled in the art. These include any cosmetically acceptable ingredients such as those found in the CTFA International Cosmetic Ingredient Dictionary and Handbook, 7th edition, edited by Wenninger and McEwen, (The Cosmetic, Toiletry, and Fragrance Association, Inc., Washington, D.C., 1997). Some non-limiting examples of these optional ingredients are given below.

Emollients

A highly preferred optional ingredient of the compositions of the present invention is emollient. Emollients tend to lubricate the skin, increase the smoothness and suppleness of the skin, prevent or relieve dryness of the skin, and/or protect the skin. A wide variety of suitable emollients are known and may be used herein. Sagarin, Cosmetics, Science and Technology, 2nd Edition, Vol. 1, pp. 32-43 (1972) contains numerous examples of materials suitable for use as emollients. Preferably the compositions of the present invention comprise greater than 1%, more preferably at least 5%, even more preferably at least 10%, by weight, of emollient.

Preferably the emollients for use herein are selected from:

- i) Straight and branched chain hydrocarbons having from about 7 to about 40 carbon atoms, such as dodecane, squalane, petrolatum, cholesterol and derivatives thereof, hydrogenated polyisobutylene, isohexadecane and the C₇-C₄₀ isoparaffins, which are C₇-C₄₀ branched hydrocarbons.
- ii) C₁-C₃₀ alcohol esters of C₁-C₃₀ carboxylic acids and of C₂-C₃₀ dicarboxylic acids, e.g. isononyl isononanoate, isopropyl myristate, myristyl propionate, isopropyl stearate, behenyl behenate, dioctyl maleate, diisopropyl adipate, and diisopropyl dilinoleate.
- iii) mono-, di- and tri- glycerides of C₁-C₃₀ carboxylic acids and ethoxylated derivatives thereof. Suitable polyethylene glycol derivatives of glycerides

include PEG-20 almond glycerides, PEG-60 almond glycerides, PEG-11 avocado glycerides, PEG-6 capric/caprylic glycerides, PEG-8 capric/caprylic glycerides, PEG-20 corn glycerides, PEG-60 corn glycerides, PEG-60 evening primose glycerides, PEG-7 glyceryl cocoate, PEG-30 glyceryl cocoate, PEG-40 glyceryl cocoate, PEG-78 glyceryl cocoate, PEG-80 glyceryl cocoate, PEG-12 glyceryl dioleate, PEG-15 glyceryl isostearate, PEG-20 glyceryl isostearate, PEG-30 glyceryl isostearate, PEG-75 cocoa butter glycerides, PEG-20 hydrogenated palm oil glycerides, PEG-70 mango glycerides, PEG-13 mink glycerides, PEG-75 shorea butter glycerides, PEG-10 olive glycerides, PEG-12 palm kernal glycerides, PEG-45 palm kernal glycerides, PEG-8 glyceryl laurate and PEG-30 glyceryl laurate. Mixtures of polyethylene glycol derivatives of glycerides can also be used herein.

- iv) alkylene glycol esters of C_1 - C_{30} carboxylic acids, e.g. ethylene glycol monoand di- esters, and propylene glycol mono- and di- esters of C_1 - C_{30} carboxylic acids e.g., ethylene glycol distearate.
- v) Organopolysiloxane oils. The organopolysiloxane oil may be volatile, nonvolatile, or a mixture of volatile and non-volatile silicones. The term "nonvolatile" as used in this context refers to those silicones that are liquid under ambient conditions and have a flash point (under one atmospheric of pressure) of or greater than about 100°C. The term "volatile" as used in this context refers to all other silicone oils. Suitable organopolysiloxanes can be selected from a wide variety of silicones spanning a broad range of Non-volatile polysiloxanes are preferred. volatilities and viscosities. Suitable silicones are disclosed in U.S. Patent No. 5,069,897, issued December 3, 1991. Preferred for use herein are organopolysiloxanes polyalkylsiloxanes, dimethicones, alkyl substituted selected from dimethiconols, polyalkylaryl siloxanes, and mixtures thereof. More preferred for use herein are polyalkylsiloxanes and cyclomethicones. Preferred among the polyalkylsiloxanes are dimethicones.
- vi) Vegetable oils and hydrogenated vegetable oils. Examples of vegetable oils and hydrogenated vegetable oils include safflower oil, castor oil, coconut oil, cottonseed oil, menhaden oil, palm kernel oil, palm oil, peanut oil, soybean oil, rapeseed oil, linseed oil, rice bran oil, pine oil, sesame oil, sunflower seed oil, partially and fully hydrogenated oils from the foregoing sources, and mixtures thereof.

vii) animal fats and oils, e.g. cod liver oil, lanolin and derivatives thereof such as acetylated lanolin and isopropyl lanolate. Lanolin oil is preferred.

- viii) Also useful are C_4 - C_{20} alkyl ethers of polypropylene glycols, C_1 - C_{20} carboxylic acid esters of polypropylene glycols, and di- C_8 - C_{30} alkyl ethers, examples of which include PPG-14 butyl ether, PPG-15 stearyl ether, dioctyl ether, dodecyl octyl ether, and mixtures thereof.
- ix) polyol carboxylic acid esters.
- x) mixtures of the above.

Preferred emollients for use in the compositions herein are selected from dodecane, squalane, cholesterol and derivatives thereof, isohexadecane, isononyl isononanoate, petrolatum, lanolin and derivatives thereof, safflower oil, castor oil, coconut oil, cottonseed oil, palm kernel oil, palm oil, peanut oil, soybean oil, polyol carboxylic acid esters and mixtures thereof. More preferred emollients for use herein are selected from polyol carboxylic acid esters, petrolatum and mixtures thereof.

These esters are derived from a sugar or polyol moiety and one or more carboxylic acid moieties. Depending on the constituent acid and sugar, these esters can be in either liquid or solid form at room temperature. Examples of liquid esters include: glucose tetraoleate, the glucose tetraesters of soybean oil fatty acids (unsaturated), the mannose tetraesters of mixed soybean oil fatty acids, the galactose tetraesters of oleic acid, the arabinose tetraesters of linoleic acid, xylose tetralinoleate, galactose pentaoleate, sorbitol tetraoleate, the sorbitol hexaesters of unsaturated soybean oil fatty acids, xylitol pentaoleate, sucrose tetraoleate, sucrose pentaoletate, sucrose hexaoleate, sucrose hepatoleate, sucrose octaoleate, and mixtures thereof. Examples of solid esters include: sorbitol hexaester in which the carboxylic acid ester moieties are palmitoleate and arachidate in a 1:2 molar ratio; the octaester of raffinose in which the carboxylic acid ester moieties are linoleate and behenate in a 1:3 molar ratio; the heptaester of maltose wherein the esterifying carboxylic acid moieties are sunflower seed oil fatty acids and lignocerate in a 3:4 molar ratio; the octaester of sucrose wherein the esterifying carboxylic acid moieties are oleate and behenate in a 2:6 molar ratio; and the octaester of sucrose wherein the esterifying carboxylic acid moieties are laurate, linoleate and behenate in a

1:3:4 molar ratio. A preferred solid material is sucrose polyester in which the degree of esterification is 7-8, and in which the fatty acid moieties are C18 mono- and/or di-unsaturated and behenic, in a molar ratio of unsaturates: behenic of 1:7 to 3:5. A particularly preferred solid sugar polyester is the octaester of sucrose in which there are about 7 behenic fatty acid moieties and about 1 oleic acid moiety in the molecule. Other materials include cottonseed oil or soybean oil fatty acid esters of sucrose. The ester materials are further described in, U. S. Patent No. 2,831,854, U. S. Patent No. 4,005,196, to Jandacek, issued January 25, 1977; U. S. Patent No. 4,005,195, to Jandacek, issued January 25, 1977, U. S. Patent No. 5,306,516, to Letton et al., issued April 26, 1994; U. S. Patent No. 5,306,515, to Letton et al., issued April 26, 1994; U. S. Patent No. 5,305,514, to Letton et al., issued April 26, 1994; U. S. Patent No. 4,797,300, to Jandacek et al., issued January 10, 1989; U. S. Patent No. 3,963,699, to Rizzi et al, issued June 15, 1976; U. S. Patent No. 4,518,772, to Volpenhein, issued May 21, 1985; and U. S. Patent No. 4,517,360, to Volpenhein, issued May 21, 1985.

The polyol fatty acid polyesters suitable for use herein can be prepared by a variety of methods well known to those skilled in the art. These methods include: transesterification of the polyol with methyl, ethyl or glycerol fatty acid esters using a variety of catalysts; acylation of the polyol with a fatty acid chloride; acylation of the polyol with a fatty acid anhydride; and acylation of the polyol with a fatty acid, per se. See, for example, U.S. Patent No. 2,831,854; U.S. Patent No. 4,005,196, to Jandacek, issued January 25, 1977.

An especially preferred material is known by the INCl name sucrose polycottonseedate.

Other Skin Benefit Agents

Other skin benefit agents may be useful in the compositions of the present invention. Examples of other skin benefit agents that may be used in the present compositions include:

(a) Vitamin Compounds

The present compositions may comprise vitamin compounds, precursors, and derivatives thereof. These vitamin compounds may be in either natural or synthetic form. Suitable vitamin compounds include, Vitamin A (e.g., beta carotene, retinoic acid, retinol, retinoids, retinyl palmitate, retinyl proprionate, etc.), Vitamin B (e.g., niacin, niacinamide, riboflavin, pantothenic acid, etc.), Vitamin C (e.g., ascorbic acid, etc.), Vitamin D (e.g., ergosterol, ergocalciferol, cholecalciferol, etc.), Vitamin E (e.g., tocopherol acetate, etc.), and Vitamin K (e.g., phytonadione, menadione, phthiocol, etc.) compounds. Preferred vitamin compounds for use in the compositions of the present invention are vitamin B₃ compounds. Vitamin B₃ compounds are particularly useful for regulating skin condition as described WO-A-97/39733. When present, the compositions of the present invention preferably comprise from about 0.01% to about 50%, more preferably from about 0.1% to about 10%, even more preferably from about 0.5% to about 5%, by weight, of the vitamin B₃ compound.

As used herein, "vitamin B₃ compound" means a compound having the formula:

$$\bigcap_{N}$$
 R

wherein R is -CONH₂ (i.e., niacinamide), -COOH (i.e., nicotinic acid) or -CH₂OH (i.e., nicotinyl alcohol); derivatives thereof; and salts of any of the foregoing.

Exemplary derivatives of the foregoing vitamin B_3 compounds include nicotinic acid esters, including non-vasodilating esters of nicotinic acid, nicotinyl amino acids, nicotinyl alcohol esters of carboxylic acids, nicotinic acid N-oxide and niacinamide N-oxide.

Examples of suitable vitamin B₃ compounds are well known in the art and are commercially available from a number of sources, e.g., the Sigma Chemical Company (St. Louis, MO); ICN Biomedicals, Inc. (Irvin, CA) and Aldrich Chemical Company (Milwaukee, WI).

The vitamin compounds may be included as the substantially pure material, or as an extract obtained by suitable physical and/or chemical isolation from natural (e.g., plant) sources.

(b) Anti-Wrinkle and Anti-Skin Atrophy Actives

Examples of anti-wrinkle and anti-skin atrophy actives that may be used in the compositions of the present invention include, but are not limited to, lactic acid and derivatives thereof, retinoic acid and its derivatives (e.g., cis and trans); retinol; retinyl esters; niacinamide, salicylic acid and derivatives thereof; sulfurcontaining D and L amino acids and their derivatives and salts, particularly the N-acetyl derivatives, a preferred example of which is N-acetyl-L-cysteine; thiols, e.g., ethane thiol; hydroxy acids, phytic acid, lipoic acid; lysophosphatidic acid, and skin peel agents (e.g., phenol and the like).

(c) Antimicrobial and Antifungal Actives

Examples of antimicrobial and antifungal actives that may be used in the compositions of the present invention include, but are not limited to, ß-lactam drugs, guinolone drugs, ciprofloxacin, norfloxacin, tetracycline, erythromycin, amikacin, 2,4,4'-trichloro-2'-hydroxy diphenyl ether, 3,4,4'-trichlorocarbanilide, phenoxyethanol, phenoxy propanol, phenoxyisopropanol, doxycycline, capreomycin, chlorhexidine, chlortetracycline, oxytetracycline, clindamycin, ethambutol, hexamidine isethionate, metronidazole, pentamidine, gentamicin, kanamycin, lineomycin, methacycline, methenamine, minocycline, neomycin, netilmicin, paromomycin, streptomycin, tobramycin, miconazole, tetracycline erythromycin, hydrochloride, erythromycin, zinc erythromycin erythromycin stearate, amikacin sulfate, doxycycline hydrochloride, capreomycin sulfate, chlorhexidine gluconate, chlorhexidine hydrochloride, chlortetracycline oxytetracycline hydrochloride, clindamycin hydrochloride, hydrochloride, metronidazole hydrochloride. pentamidine ethambutol hydrochloride, hydrochloride, gentamicin sulfate, kanamycin sulfate, lineomycin hydrochloride, methacycline hydrochloride, methenamine hippurate, methenamine mandelate, minocycline hydrochloride, neomycin sulfate, netilmicin sulfate, paromomycin sulfate, streptomycin sulfate, tobramycin sulfate, miconazole hydrochloride, amanfadine hydrochloride, amanfadine sulfate, octopirox, parachlorometa xylenol, nystatin, tolnaftate, zinc pyrithione and clotrimazole.

(d) Sunscreen Actives

The compositions herein may also comprise sunscreening actives. A wide variety of sunscreen agents are useful herein. These sunscreen agents include both organic compounds and their salts as well as inorganic particulate materials. Without being limited by theory, it is believed that sunscreen agents provide protection from ultraviolet radiation by one or more of the following mechanisms including absorption, scattering, and reflection of the ultraviolet radiation. Nonlimiting examples of these sunscreen agents are described in U.S. Patent No. 5,087,445, to Haffey et al., issued February 11, 1992; U.S. Patent No. 5,073,372, to Turner et al., issued December 17, 1991; U.S. Patent No. 5,073,371, to Turner et al. issued December 17, 1991; U.S. Patent No. 5.160.731, to Sabatelli et al., issued November 3, 1992; U.S. Patent No. 5,138,089, to Sabatelli, issued August 11, 1992; U.S. Patent No. 5,041,282, to Sabatelli, issued August 20, 1991; U.S. Patent No. 4,999,186, to Sabatelli et al., issued March 12, 1991; U.S. Patent No. 4,937,370, to Sabatelli, issued June 26, 1990; and Segarin, et al., at Chapter VIII, pages 189 et seq., of Cosmetics Science and Technology. Preferred among the sunscreen agents are those selected from the group consisting of 2-ethylhexyl p-methoxycinnamate, octyl salicylate, octocrylene, oxybenzone, 2-ethylhexyl N,N-dimethylaminobenzoate, p-aminobenzoic acid, 2-phenyl-benzimidazole-5-sulfonic acid, homomenthyl salicylate. DEA p-methoxycinnamate, 4,4'methoxy-t-butyldibenzoylmethane, 4isopropyldibenzoylmethane, 3-(4-methylbenzylidene) camphor, 3-benzylidene ester with 2.4-4-N,N-dimethylaminobenzoic acid camphor. dihydroxybenzophenone, 4-N,N-dimethylaminobenzoic acid ester with 2hydroxy-4-(2-hydroxyethoxy)benzophenone, 4-N,N-dimethylaminobenzoic acid ester with 4-hydroxydibenzoyl- methane, 4-N,N-dimethylaminobenzoic acid ester 4-(2-hydroxyeth-oxy)dibenzoylmethane, 4-N,N-di(2-ethylhexyl)with 4-N, N-di(2ester with 2,4-dihydroxybenzophenone, aminobenzoic acid 2-hydroxy-4-(2ester with ethylhexyl)aminobenzoic acid hydroxyethoxy)benzophenone, 4-N,N-di(2-ethylhexyl)aminobenzoic acid ester with 4-hy-droxydibenzoylmethane, 4-N,N-di(2-ethylhexyl)aminobenzoic acid 4-(2-hydroxyeth-oxy)dibenzoylmethane, 4-N,N-(2with ester ethylhexyl)methylaminobenzoic acid ester with 2,4-dihydroxybenzophenone, 4-2-hvdroxy-4-(2acid ester with N,N-(2-ethylhexyl)methylaminobenzoic

hydroxyethoxy)benzophenone, 4-N,N-(2-ethylhexyl)methylaminobenzoic acid ester with 4-hy-droxydibenzoylmethane, 4-N,N-(2-ethylhexyl)methylaminobenzoic acid ester with 4-(2-hydroxyethoxy)dibenzoylmethane, titanium dioxide, zinc oxide, iron oxide, and mixtures thereof.

More preferred for use in the compositions described herein are the sunscreen agents selected from the group consisting of 2-ethylhexyl N,N-dimethyl-p-aminobenzoate, 2-ethylhexyl p-methoxycinnamate, octocrylene, octyl salicylate, homomenthyl salicylate, p-aminobenzoic acid, oxybenzone, 2-phenylbenzimidazole-5-sulfonic acid, DEA p-methoxycinnamate, 4,4'-methoxy-t-butyldibenzoylmethane, 4-isopropyl dibenzoylmethane, 3-(4-methylbenzylidene) camphor, 3-benzylidene camphor, 4-N,N-(2-ethylhexyl)methylaminobenzoic acid ester with 4-(2-hydroxyeth-oxy)dibenzoylmethane, titanium dioxide, zinc oxide, iron oxide, and mixtures thereof.

Exact amounts of sunscreens which can be employed will vary depending upon the sunscreen chosen and the desired Sun Protection Factor (SPF) to be achieved. SPF is a commonly used measure of photoprotection of a sunscreen against erythema. See Federal Register, Vol. 43, No. 166, pp. 38206-38269, August 25, 1978.

Thickeners

The compositions of the present invention preferably comprise thickeners. Any thickener suitable for use in cosmetic compositions can be used herein. Preferred thickeners are selected from nonionic water-soluble polymers, fatty alcohols and mixtures thereof. Suitable nonionic polymers include such water soluble polymers as cellulose ethers (e.g. hydroxybutyl methyl cellulose, hydroxypropylcellulose, hydroxypropyl methyl cellulose, ethylhydroxy ethyl modified hydroxyethyl cellulose and cellulose, hydrophobically alcohol, hydroxyethylcellulose), poly(ethylene oxide), polyvinyl polyvinylpyrrolidone, hydroxypropyl guar gum, amulose, hydroxyethyl amylose, starch, and starch derivatives. Suitable fatty alcohols are higher molecular weight, nonvolatile, primary alcohols having the general formula

RCH2OH

wherein R is a C₈₋₂₀ alkyl. They can be produced from natural fats or oils by reduction of the fatty acid COOH-grouping to the hydroxyl function. Alternatively, identical or similarly structured fatty alcohols can be produced according to conventional synthetic methods known in the art. Suitable fatty alcohols include, but are not limited to behenylalcohol, C₉-C₁₁ alcohols, C₁₂-C₁₃ alcohols, C₁₂-C₁₅ alcohols, C₁₂-C₁₆ alcohols, C₁₄-C₁₅ alcohols, caprylic alcohol, cetearyl alcohol, coconut alcohol, decyl alcohol, isocetyl alcohol, isostearyl alcohol, lauryl alcohol, oleyl alcohol, palm kernel alcohol, stearyl alcohol, cetyl alcohol, tallow alcohol, tridecyl alcohol or myristyl alcohol.

Polar Solvent

The compositions of the present invention may also comprise polar solvent. Any polar solvent suitable for use in cosmetic compositions may be used herein. However, the polar solvent must be sufficiently polar to drive the formation of vesicles in the present invention. Preferably the polar solvent used in the compositions of the present invention is water.

Preferably comprise the present compositions will comprise from 10% to 90%, more preferably from 20% to 80%, even more preferably from 30% to 60%, by weight, of polar solvent.

Other Optional Ingredients

The compositions of the present invention can comprise a wide range of other optional components. These additional components should be pharmaceutically acceptable. Non-limiting examples of functional classes of ingredients suitable for use in the compositions of the present invention include: abrasives, absorbents, anti-acne actives, anticaking agents, anti-dandruff agents, anti-perspirant agents, antioxidants, anti-viral actives, artificial tanning actives and accelerators, biological additives, bleach, bleach activators, brighteners, builders, buffering agents, chelating agents, chemical additives, colorants, cosmetics, cleansers, cosmetic astringents, cosmetic biocides, denaturants, deodorants, desquamation actives, depilatories, drug astringents, dyes, dye transfer agents, enzymes, external analgesics, foam generators, flavours, film

formers, fragrance components, insect repellents, mildewcides, non-steroidal anti-inflammatory active, opacifying agents, oxidative dyes, oxidising agents, pest control ingredients, pH adjusters such as citric acid, pH buffers, pharmaceutical actives, plasticizers, preservatives, radical scavengers, skin, hair or nail bleaching agents, skin, hair or nail conditioners, skin, hair or nail penetration enhancers, stabilisers, surfactants, surface conditioners, reducing agents, temperature depressors, viscosity modifiers, and warmth generators such as exothermic zeolites. Also useful herein are aesthetic components such as colourings, essential oils, and skin healing agents. Other optional materials herein include pigments. Pigments suitable for use in the compositions of the present invention can be organic and/or inorganic. Also included within the term pigment are materials having a low colour or lustre such as matte finishing agents, and also light scattering agents. Examples of suitable pigments are iron oxides, acyglutamate iron oxides, titanium dioxide, ultramarine blue, D&C dyes, carmine, and mixtures thereof.

Formulation Process

Preferably the compositions of the present invention are preferably prepared in such a way that the quaternary ammonium compound forms vesicles. It is preferred that said vesicles also comprise humectant. Preferably said vesicles also comprise emollient. In order to ensure optimal performance characteristics it is preferred that the compositions of the present invention are prepared as follows:

- (i) all or part of the quaternary ammonium agent is mixed with humectant, water soluble skin care actives (where included), and, preferably, polar solvent at a temperature which is higher than the melting point of the quaternary ammonium agent;
- (ii) optionally, the mixture is vigorously agitated;
- (iii) In a separate vessel the emulsion is prepared as follows; the oil phase containing the emollients, the relevant thickener in case the said thickener is oil soluble, and any remaining quaternary ammonium agent are mixed together at a temperature which is higher than the melting point of the quaternary ammonium compound. The aqueous phase is prepared separately. The water, the relevant thickener in case the said thickener is water soluble, and any remaining water

soluble ingredients are heated to the same temperature as the oil phase.

- (iv) the temperature of the oil and aqueous phases of the emulsion are then approximately equalised and the aqueous phase is combined with the oil phase with agitation.
- (v) On production of the emulsion the mixture formed in step (i) is added to the aforementioned emulsion with agitation.

Method of Use

The cosmetic compositions of the present invention may be used in a conventional manner for the treatment of skin. An effective amount of the composition, typically from about 0.1 grams to about 50 grams, preferably from about 1 gram to about 20 grams, is applied to wet or dry, preferably wet, skin. Application of the composition typically includes working the composition into the skin, generally with the hands and fingers. The composition is then left on the skin or, preferably, the skin is rinsed.

The preferred method of treating the skin, therefore, comprises the steps of:

- (a) applying an effective amount of the cosmetic composition to the skin,
- (b) rinsing the skin.

A preferred aspect of the present invention involves the above method with an application of the composition on dry skin before an application on wet skin. Therefore, a preferred method comprises:

- (i) applying to dry skin an effective amount of the cosmetic composition;
- (ii) rinsing the skin under a shower;
- (iii) further application of said composition; and
- (iv) further rinsing.

Much of the damage to human skin is caused by repeated exposure to surfactant containing compositions during washing routines. It has been found that this damage can be mitigated using the present compositions. Therefore, another preferred method comprises:

- (i) washing the skin using a composition comprising surfactants;
- (ii) rinsing the skin;
- (iii) applying to the wet skin a composition according to the present invention;
- (iv) rinsing the skin.

It has also been found that the present compositions are particularly useful when incorporated as part of a regular routine. Therefore, another preferred method comprises:

- (i) applying to the skin a composition comprising:
 - (a) at least one quaternary ammonium compound;
 - (b) humectant; and
- (ii) rinsing the skin;
- (iii) repeating steps (i) and (ii) within 48 hours.

The present compositions can also be useful in mitigating damage caused by exposure of the skin to ultra violet radiation, damage caused by exposure of the skin to water during swimming or similar water based exercise, damage caused by shaving or exfoliation or damage caused by exposure of the skin to water during bathing.

Examples

The following examples further illustrate the preferred embodiments within the scope of the present invention. The examples are given solely for the purposes of illustration and are not to be construed as limitations of the present invention as many variations of the invention are possible without departing from its spirit or scope. Unless otherwise indicated, all ingredients are expressed on a weight percentage of the active ingredient.

Example	1	2	3	4	5	6	7	8	9	10	11	12	13	
							Weig	ht %						
Quat. Amm. Agt	3	3	3	3	3	3	3	3	3	3	3	3	3	А
Glycerine	32	32	32	32	32	32	32	-	-	-	_	-	-	Α
Urea	-	-	-	-	_	-	-	22	22	22	22	22	22	А

											r:			$\overline{}$
Petrolatum			10	-	12	15_	-	-	-	10	-	12	15	В
Lanolin 1	-	-		7.5	5	-	15	-	-	-	10	10	-	В
Coronet Lanolin 1			-		-	-	5		-	-	-		-	В
Super Sterol	-	-	-	-	-	-	5	-	-	-	-	-	-	В
Esters ¹														<u> </u>
Sefa Cottonate	21	21	12	12	-	-	-	21	21	12	12		-	В
Dimethicone	-	0.5	0.5	0.5	-	0.5	0.5	0.5	-	-	-	0.5	0.5	В
10Cst														<u> </u>
Dimethicone	-	1.0	1.0	1.0	-	1.0	1.0	1.0	-	-	-	1.0	1.0	В
1000Cst														<u> </u>
Cyclomethicone	-	2.5	2.5	2.5	-	2.5	2.5	2.5	-	<u>-</u>	-	2.5	2.5	В
Florasun PEG-10	-	2	-	2	-	3	-	1	-	-	3	-	-	В
2														L
Methyl Paraben	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	A
Propyl Paraben	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	В
Disodium EDTA	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	Α
Sodium Benzoate	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	В
Niacinamide	5_	5	3		5	5	5	3	5			5	5	A
Panthenol	3	3	1_		3	3	3	1	3			3	3	Α
Tocopherol	2	2	0.5	ļ	2	2	2	0.5	2			2	2	В
Acetate														
Retinol Palmitate	0.1	-	-				0.05		-	0.1	<u>-</u>	-	-	В
Parsol 1789 ³		1	-	1	0.5		0.1		1		1	<u>-</u>	1	В
Parsol MCX 3		5		5	2.5		0.5	-	5		5	-	5	В
Citric acid	1	0.5	1	0.5	0.5	1	0.5	11	0.3	0.75	1	1_	1	Α
Salicylic acid		0.5	_	0.5	0.5	-	0.5		0.7	_	-	<u>-</u>	-	Α
Triethanolamine	0.1		-	0.1	0.1			-	0.1	-	-	0.9	0.5	С
Sodium	-	-	0.1	0.1	0.1	-	-	0.1	-	0.1	-	_	-	С
Hydroxide														
Polyquaternium	-		-	1.0		-	0.5	-	0.6	-	-	-	0.5	В
10					ļ	<u> </u>	ļ							_
Polymer KG 30 ⁴	_	0.7	-		0.7			-		0.8	-	-	-	В
Fragrance	0.5	0.5	0.5	0.5	0.5	<u> </u> -	0.5	0.5	0.5	0.5	0.5	0.5	0.5	E
Cetyl Alcohol	3	-	<u> </u>		2.6		_	2.2		2.2		2.5		С

Stearyl Alcohol	2	_	_		1.5	_	_	3.2	-	1.1	_	1.6	_	C
Cetearyl Alcohol	-	_	4	_	0.4	-	-	_	3.4	_	-	_	3.4	С
Behenyl Alcohol	2	-	2	-	-	_	-	0.6	_	0.7	_	-	-	С
Natrosol 330 Plus	1	_	-	-	0.4	-	-	0.7	0.6		-	0.5	0.2	С
													5	
Natrosol 250	-	-	1	-	0.3	-	-	-	-	1.2	-	-	0.3	С
HHR														
Jaguar HP 105 ⁵	-	1.2	-	-	-	-	-	_	-	-	0.8	-	-	D
Jaguar C14S ⁵	-		-	1	-	-	-	-	-	-	-	-	-	D
Jaguar C13S ⁵	<u>-</u>	-	-	-	-	1.2		-	-	-	-	-	-	D
Xanthan Gum	-		-	-	-	-	2.0	_	_	-	-	-	-	D
Sodium Chloride	-	-	0.1	-	-	-	0.5	0.2	-	0.3	-	-	-	Α
Water	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	Α

Example	14	15	16	17	18	19	20	21	22	23	24	25	26	
				l	<u></u>		Weigh	t %	I		I.	ļ.,		
Quat. Amm. Agt (see notes below)	8	8	8	8	8	8	8	8	8	8	8	8	8	А
Glycerine	32	32	32	32	32	32	32		-		-	-	-	Α
Urea	-	-	-	-	-	-	-	22	22	22	22	22	22	A
Petrolatum	-	-	10	-	12	15	-	_	1	10	-	12	15	В
Lanolin 1		-	-	7.5	5	•	15	-	-	1	10	10	_	В
Coronet Lanolin 1	-	-	-	-	-	-	5	-		•	-	_	-	В
Super Sterol	-	-	-	-	-	+	5	+	•	-	-	-	-	В
Sefa Cottonate	21	21	12	12	-	-	-	21	21	12	12	-	_	В
Dimethicone 10Cst	-	0.5	0.5	0.5	-	0.5	0.5	0.5	-	-	-	0.5	0.5	В
Dimethicone 1000Cst	-	1.0	1.0	1.0	-	1.0	1.0	1.0	-	-	-	1.0	1.0	В
Cyclomethicone	-	2.5	2.5	2.5	-	2.5	2.5	2.5	-	-	-	2.5	2.5	В
Florasun PEG-10	-	2	-	2	_	3	-	1		-	3	-	-	В

2														
Methyl Paraben	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	A
Propyl Paraben	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	В
Disodium EDTA	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.0	0.0	0.1	A
Sodium Benzoate	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	В
Niacinamide	5	5	3	0.0	5	5	5	3	5	0.0	0.0	5	5.5	A
Panthenol	3	3	1		3	3	3	1	3			3	3	A
Tocopherol	2	2	0.5		2	2	2	0.5	2			2	2	В
Acetate	-	-	0.0		_	_	_	0.0				-	-	
Retinol Palmitata	0.1	_		_	_	0.05	_	_	_	_	_	_	0.15	В
Parsol 1789 ³	-	1		1	0.5	-	0.1	_	1	<u>-</u>	1	_	1	В
Parsol MCX 3	-	5	_	5	2.5	-	0.5	-	5	_	5	_	5	В
Citric acid	1	0.5	1	0.5	0.5	1	0.5	1	0.3	0.75	1	1	1	A
Salicylic acid		0.5	-	0.5	0.5	_	0.5	-	0.7	-	-	-	-	Α
Triethanolamine	0.1	-	_	0.1	0.1	-	-	-	0.1	-	-	0.9	0.5	С
Sodium	-	-	0.1	0.1	0.1	-	-	0.1	-	0.1	-	-	_	С
Hydroxide														
Polyquaternium	-		-	1.0		-	0.5	-	0.6	-	-	-	0.5	В
10									,					
Polymer KG 30 ⁴		0.7	-		0.7	-		•		0.8	-	-		В
Fragrance	0.5	0.5	0.5	0.5	0.5	-	0.5	0.5	0.5	0.5	0.5	0.5	0.5	Ε
Cetyl Alcohol	3	_	-	-	2.6	-	-	2.2	-	2.2	-	2.5	-	С
Stearyl Alcohol	2	-		-	1.5	-	-	3.2	-	1.1	-	1.6	-	С
Cetearyl Alcohol	-		4	-	0.4	-	-	-	3.4	-	-	-	3.4	С
Behenyl Alcohol	2	-	2	-	-	-	-	0.6	-	0.7	-	-	-	С
Natrosol 330 Plus	1	-	-	-	0.4	-	-	0.7	0.6		-	0.5	0.25	С
Natrosol 250	-	-	1	-	0.3	-	-	-	-	1.2	-	-	0.3	С
HHR														
Jaguar HP 105 ⁵	-	1.2	-	-	-	-	-	-	-		0.8	-	-	D
Jaguar C14S ⁵	_		-	1	-	-	-				-	-	-	D
Jaguar C13S ⁵	-	_	-	-	-	1.2	-	-	-	-	-	-	-	D
Xanthan Gum	<u> </u>		-	-	-	-	2.0	-	-	-	-	-	-	D
Sodium Chloride	-	_	0.1	<u>-</u>	-		0.5	0.2	-	0.3	-	-	-	_A
Water	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	Α

Example	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	
							w	eight	%	1						
Quat. Amm. Agt	3	3	3	3	3	3	3	3	3	3	3	3	3	3	3	А
Glycerine	15	16	22	9	8	9	-	32	32	32	32	32	32	32	32	Α
Propylene Glycol	-	16	-	-	-	5	12	-	-	-			-			A
Butylene Glycol	-	-	8_	8		8	12	-		-	-	_	_			Α
Urea		-		12		2	9		ļ -	-	_	-		_		Α
Petrolatum			10	-	12	15	-	-	4	7.5	-	12	5	-	6	В
Lanolin ¹	-	-	-	7.5	7.5	-	15	<u>-</u>	3.5	<u>-</u>	3			-		В
Coronet Lanolin 1	-	-	-	-	-	-	-		7	-	6.7			6	<u>-</u>	В
Super Sterol Esters ¹	-	-	-	-	-	-	-	-	3.5	-	6.5	-	3	6	3	В
Isohexadecane	-	-	-	-	-	-	-		-	5		3	-	-	3	В
Isononyl	-	-	-	-	-	-	-	-	-	3	_	5	-	-	-	В
Isononanoate																
Sefa Cottonate	21	21	12	12	_	-	-	13	-	_8			8	12	10	В
Sefa Behenate	-	-	-	-	-	-	-	9	-	-	3	-	5		-	В
Dimethicone 10Cst	_	0.5	0.5	0.5	-	0.5	0.5	0.5	-	-	-	0.5	0.5	-	-	В
Dimethicone	-	1.0	1.0	1.0	-	1.0	1.0	1.0	-	-	-	1.0	1.0	-	-	В
Cyclomethicone	-	2.5	2.5	2.5	-	2.5	2.5	2.5	-	_	_	2.5	2.5	_	_	В
Florasun PEG-10 ²		2	-	2	•	3		1	1		3	-	-	2.5	-	В
Methyl Paraben	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	Α
Propyl Paraben	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	В
Disodium EDTA	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	Α
Sodium Benzoate	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	В
Niacinamide	3	5	5			5	5	5	3	3		5	5	5	3	Α
Panthenol	1	3	3_			3	3	3	1	1		3	3	3	1	Α
Tocopherol Acetate	0.5	2	2			2	2	2	0.5	0.5		2	2	2	0.5	В
Retinol Palmitate		-		0.1	-	0.05	_	-	0.1	-	-	-	-		-	В

	,			,				,							, <u> </u>	
Parsol 1789 3		1	-	1	0.5	_	1	-	1		1		0.1	1	_	В
Parsol MCX 3	-	5	-	5	2.5		5	-	5		5		0.5	5		В
Citric acid	1	0.5	1	0.5	0.5	1	0.5	1	0.3	0.75	1	1	1	1	<u>-</u>	Α
Salicylic acid	-	0.5	-	0.5	0.5	-	0.5		0.7	-		-		-	0.5	Α
Triethanolamine	0.1		-	0.1	0.1	<u>-</u>	-	-	0.1	-	-	0.9	0.5		-	С
Sodium Hydroxide	-	-	0.1	0.1	0.1	-	-	0.1	-	0.1	-	-		-	0.6	С
Polyquaternium 10	-		-	1.0	-	<u> </u>	0.5		0.6	-	-		0.5	1.0	-	В
Polymer KG 30 ⁴	_	0.7	-	-	0.7	_		-	-	0.8	-	-	-		0.8	В
Fragrance	0.5	0.5	0.5	0.5	0.5	-	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	Ε
Cetyl Alcohol	3		-	-	2.6	-	-	2.2		2.2	_	2.5	_	-	3.2	С
Stearyl Alcohol	2	-		-	1.5		_	3.2	_	1.1		1.6	_	_	2.6	С
Cetearyl Alcohol		-	4_	_	0.4	-	-	-	3.4	-	-	<u>.</u>	3.4	-		С
Behenyl Alcohol	2		2	-	-		-	0.6	-	0.7	-	-	-	-	0.2	С
Natrosol 330 Plus	1			-	0.4	-	-	0.7	0.6			0.5	0.25	-	0.6	С
Natrosol 250 HHR	-	-	1_	-	0.3	-	-	-	-	1.2	-	-	0.3	-		С
Jaguar HP 105 ⁵	-	1.2	-	-	-	-	-	_	-	-	0.8	-		_	_	D
Jaguar C14S ⁵	_	-	-	1	-	-		-	-	-	-	-	-	-	-	D
Jaguar C13S ⁵	_	-	_	-		1.2	-	-	-	-	-	-	-	0.8	-	D
Xanthan Gum	-		-	-			3.0	-	-	-	-	-		-	-	D
Sodium Chloride		-	0.1	-	-	-	0.5	0.2	-	0.3	-	-	-	0.2	0.1	Α
Water	qs	qs	qs	qs	qs	gs	qs	qs	qs	qs	qs	qs	qs	qs	qs	Α

Example	42	43	44	45	46	47	48	49	50	51	52	53	54	55	56	
			<u> </u>				l	Weig	jht %				1			
Quat. Amm. Agent (see notes below)	8	8	8	8	8	8	8	8	8	8	8	8	8	8	8	А
Glycerine	15	16	22	9	8	9	-	32	32	32	32	32	32	32	32	А
Propylene Glycol	-	16	-		-	5	12	-	-	-	-	-		-		Α
Butylene Glycol	-		8	8_	-	8	12		-	-	-	-	<u>-</u>	-		Α
Urea	_	_	_	12	-	2	9	_	-	-	-	-	_	-		Α
Petrolatum	-	-	10	_	12	15		_	4	7.5	-	12	5		6	В
Lanolin ¹		_	-	7.5	7.5	-	15	-	3.5	-	3	_	-	-	_	В
Coronet Lanolin 1	_	-	-	-	-	-	_	-	7	_	6.7		-	6	_	В

Super Sterol Esters	-	-	-	-	-	-	-	-	3.5	-	6.5	-	3	6	3	В
Isohexadecane	_	_	-		-	-		_	-	5	_	3		-	3	В
isononyi	-		-	-	-	-	-	-	-	3	-	5	-	-	-	В
Isononanoate									,							
Sefa Cottonate	21	21	12	12	-	-	-	13	-	8		-	8	12	10	В
Sefa Behenate		-			-	-	-	9	-		3	-	5_	-		В
Dimethicone 10Cst		0.5	0.5	0.5	-	0.5	0.5	0.5	-			0.5	0.5	_	-	В
Dimethicone	-	1.0	1.0	1.0	-	1.0	1.0	1.0	-	-	-	1.0	1.0	-	-	В
1000Cst																
Cyclomethicone	-	2.5	2.5	2.5		2.5	2.5	2.5	-	-	-	2.5	2.5	-	-	В
Florasun PEG-10 ²	-	2	-	2	-	3	-	1	-	-	3	-	-	2.5		В
Methyl Paraben	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	Α.
Propyl Paraben	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	В
Disodium EDTA	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	0.1	Α
Sodium Benzoate	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	0.3	В
Niacinamide	3	5	5			5	5	5	3	3		5	5	5	3	A
Panthenol	1	3	3			3	3	3	11	1		3	3_	3	1	Α
Tocopherol Acetate	0.5	2	2			2	2	2	0.5	0.5		2	2	2	0.5	В
Retinol Palmitate		-	-		-	-	-	-	0.05	_		0.1	_	-	-	В
Parsol 1789 ³	-	1	-	1	0.5	_	1	-	1	-	1		0.1	1	-	В
Parsol MCX ³	-	5	-	5	2.5	-	5	-	5	-	5	-	0.5	5	-	В
Citric acid	1	0.5	1	0.5	0.5	1	0.5	1	0.3	0.75	1	1	1	1	-	Α
Salicylic acid		0.5	-	0.5	0.5	-	0.5		0.7	-	-	-	-	-	0.5	Α
Triethanolamine	0.1	<u>.</u>	_	0.1	0.1	-	-		0.1	-	-	0.9	0.5		-	С
Sodium Hydroxide		-	0.1	0.1	0.1			0.1		0.1	-	-	-	-	0.6	С
Polyquaternium 10			-	1.0	-		0.5		0.6	-	-	-	0.5	1.0	-	В
Polymer KG 30 ⁴	-	0.7	-	-	0.7			-	•	0.8	-		-	-	0.8	В
Fragrance	0.5	0.5	0.5	0.5	0.5		0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	0.5	E
Cetyl Alcohol	3				2.6	-	-	2.2	•	2.2	-	2.5	-	-	3.2	С
Stearyl Alcohol	2	-	-		1.5		_	3.2		1.1	-	1.6		-	2.6	С
Cetearyl Alcohol	-	-	4	-	0.4	-		_	3.4	_		-	3.4	-	-	С
Behenyl Alcohol	2		2	-		-		0.6	-	0.7	-	-	-	-	0.2	С
Natrosol 330 Plus	1				0.4	-		0.7	0.6		-	0.5	0.2	•	0.6	С

													5			
Natrosol 250 HHR		_	1		0.3	-	-	-	-	1.2	_	-	0.3	-		С
Jaguar HP 105 ⁵		1.2	•		-		-	-	_	-	0.8	-	-	_	_	D
Jaguar C14S ⁵	_	-		1	-	-	<u>.</u>	-	_	-	_	,	-			D
Jaguar C13S ⁵				-	-	1.2		-			_	-	-	0.8		D
Xanthan Gum		-	-	-	-	-	3.0	-	ı	-	-	-	-	-		D
Sodium Chloride	-	-	0.1	-		-	0.5	0.2	•	0.3		-	-	0.2	0.1	Α
Water	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	qs	А

- 1; Available from Croda
- 2: Available from Floratech, AZ, USA
- 3: Available from Hoffman La Roche, NJ, USA
- 4; Available from Amerchol, NJ, USA
- 5; Available from Rhodia, NJ, USA
- In examples 1, 2, 3, 4, 5, 6, 7, 8, 9, 10, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 26, 27, 28, 31, 33, 34, 35, 36, 37,38, 41, 45, 47, 48, 49, 50, 51, 52, 53, 54, 55, 56 the quaternary ammonium compound used is N,N-di(canolyl-oxyethyl)-N methyl,N-(2-hydroxyethyl)Ammonium Methyl Sulfate supplied by Goldschmidt, trade name Rewoquat V3620.
- In Example 11,25,29 the quaternary ammonium compound used is N,N-di(canolyl-oxy-ethyl)-N methyl,N-(2-hydroxyethyl)Ammonium Methyl Sulfate supplied by Goldschmidt, trade name Rewoguat WE18
- In Example 12,30,32 the quaternary ammonium compound used is N,N-di(canolyl-oxy-ethyl)-N methyl,N-(2-hydroxyethyl)Ammonium Chloride supplied by Goldschmidt, developmental material (WE25)
- In Example13,39,40,42 the quaternary ammonium compound used is Methylbis(hydrogenated tallowamidoethyl)(2-hydroxyethyl)ammonium methyl sulfate supplied by Goldschmidt, trade name Varisoft 110.
- In Example 14,43,44 the quaternary ammonium compound used is Methylbis(tallowamidoethyl)(2-hydroxyethyl)ammonium methyl sulfate supplied by Goldschmidt, trade name Varisoft 222.

Process

For those examples including component C:

1. Premix 1: Combine components of group A together at a temperature greater than the transition temperature of the chosen quaternary ammonium compound, keeping back a predetermined part of the quaternary ammonium compound and water. Vigorously agitate this premix.

- 2. Premix 2: Combine components of groups B and C with the remaining parts of the quaternary ammonium compounds and water not previously used in premix 1. Heat above the melting point of the quat and the oils.
- 3. Combine Premix 1 & 2 and allow to cool until 40°C, stir in the perfume.

For those examples including component D:

- 1. Premix 1: Combine components of group A together at a temperature greater than the transition temperature of the chosen quaternary ammonium compound, keeping back a predetermined part of the water. Vigorously agitate this premix.
- 2. Premix 2: Combine with agitation the components of group D with the water not previously used in premix 1.
- **3.** Combine Premix 1 & 2 and components of groups B & E. Vigorously agitate.

The compositions of the above examples provide good skin care benefits, such as good moisturisation, good hydration, good skin feel, good skin softness and/or good skin smoothness, with low levels of negatives such as greasiness, stickiness or tack.

Claims

- 1. A cosmetic composition comprising:
 - (a) at least one readily biodegradable quaternary ammonium agent; and
 - (b) at least 5%, by weight, of humectant.
- 2. A composition according to Claim 1 wherein the composition comprises 0.01%, more preferably at least 1%, by weight, of readily biodegradable quaternary ammonium agent.
- 3. A composition according to Claim 1 or 2 wherein the composition comprises at least 7.5%, more preferably at least 10%, by weight of humectant.
- 4. A composition according to any of the preceding claims wherein the humectant is selected from glycerine, urea, butylene glycol, polyethylene glycol and derivatives thereof, and mixtures thereof.
- 5. A composition according to any of the preceding claims wherein the composition comprises vesicles wherein said vesicles comprise the quaternary ammonium agent and humectant.
- 6. A composition according to any of the preceding claims wherein the quaternary ammonium agent are selected from those that are able to form vesicles in polar solvents, as detected by microscopic analysis.
- 7. A composition according to any of the preceding claims wherein the quaternary ammonium agent is selected from N,N-di(canolyl-oxy-ethyl)-N,N-dimethyl ammonium chloride, N,N- di(canolyl-oxy-ethyl)-N-methyl,N-(2-hydroxyethyl) ammonium methyl sulfate, N,N-di(canolyl-oxy-ethyl)-N-methyl, N-(2-hydroxyethyl) ammonium chloride and mixtures thereof.
- 8. A composition according to any of the preceding claims wherein composition additionally comprises an emollient.

9. A composition according to any of the preceding claims wherein said composition comprises less than 5%, preferably less than 3%, preferably 0%, by weight, of anionic surfactant.

10. Use of a composition according to any of the preceding claims for the treatment of skin.

INTERNATIONAL SEARCH REPORT

donal Application No PCT/US 00/17645

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 A61K7/50 A61K7/00

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) A61K IPC 7

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 practical, search terms used)

EPO-Internal, CHEM ABS Data, WPI Data, PAJ

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 96 03970 A (WITCO CORP) 15 February 1996 (1996-02-15) page 10, line 15 - line 31 page 11, line 1 - line 17 page 27 -page 28; claims	1,2,4,8,
X	WO 92 09264 A (RICHARDSON VICKS INC) 11 June 1992 (1992-06-11) the whole document	1-4,8,10
X	WO 90 03161 A (INOLEX CHEMICAL COMPANY) 5 April 1990 (1990-04-05) examples claims	1,2,4,8,
	-/	
- II		

X Further documents are listed in the continuation of box C.	Patent family members are listed in 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 document 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	Date of mailing of the international search report
19 October 2000	07/11/2000
Name and mailing address of the ISA	Authorized officer
European Patent Ojffice, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijewijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Pelli Wablat, B

1

INTERNATIONAL SEARCH REPORT

Inti ional Application No PCT/US 00/17645

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT					
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.			
P,X	WO 00 00173 A (PROCTER & GAMBLE (US)) 6 January 2000 (2000-01-06) page 5, last paragraph page 6 -page 9 page 20, paragraph 3 claims	1,2,4,8, 10			
A	WO 98 46199 A (CAPSULIS (FR)) 22 October 1998 (1998-10-22) page 9, line 11 - line 55 page 10, line 15 example 9 claims 1-11	1-10			
A	US 5 145 604 A (NEUMILLER PHILLIP J) 8 September 1992 (1992-09-08) column 12; example 14 	1-10			

1

INTERNATIONAL SEARCH REPORT

Information on patent family members

Into Ional Application No
PCT/US 00/17645

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 9603970	A	15-02-1996	US AU AU BR CA CN EP JP US	5552137 A 689940 B 3214495 A 9508862 A 2196803 A 1158560 A 0767653 A 10506882 T 5610187 A	03-09-1996 09-04-1998 04-03-1996 16-09-1997 15-02-1996 03-09-1997 16-04-1997 07-07-1998 11-03-1997
WO 9209264	A	11-06-1992	US US AU AU BR CA CN DE EP ES FI HK IE JP MX NO PT NZ	5073371 A 5073372 A 666470 B 9101091 A 9107097 A 2095607 A,C 1063038 A 69111466 D 69111466 T 0559786 A 2074868 T 932454 A 1007505 A 66747 B 6503086 T 9102332 A 931885 A 99617 A,B 240795 A	17-12-1991 17-12-1991 15-02-1996 25-06-1992 09-11-1993 31-05-1992 29-07-1992 24-08-1995 04-04-1996 15-09-1993 16-09-1995 28-05-1993 16-04-1999 07-02-1996 07-04-1994 01-06-1992 29-07-1993 30-10-1994
WO 9003161	A	05-04-1990	US AT AU CA DE DE EP	4978526 A 124861 T 4210789 A 1327014 A 68923454 D 68923454 T 0435900 A	18-12-1990 15-07-1995 18-04-1990 15-02-1994 17-08-1995 21-03-1996 10-07-1991
WO 0000173	Α	06-01-2000	AU	4962999 A	17-01-2000
WO 9846199	A	22-10-1998	FR AU EP	2761912 A 7340798 A 0975307 A	16-10-1998 11-11-1998 02-02-2000
US 5145604	A	08-09-1992	US AT AU CA DE EP ES JP MX NZ PT WO	5091111 A 138969 T 641289 B 8645991 A 2111122 A,C 69120087 D 69120087 T 0553121 A 2088018 T 6501201 T 9101174 A 239857 A 99004 A,B 9205229 A	25-02-1992 15-06-1996 16-09-1993 15-04-1992 02-04-1992 11-07-1996 23-01-1997 04-08-1993 01-08-1996 10-02-1994 04-05-1992 28-04-1993 31-07-1992 02-04-1992