(19) Europäisches Patentamt
European Patent Office
Office européen des brevets

(11) EP 1 272 154 B1

(12) **EUROPEAN PATENT SPECIFICATION**

(45) Date of publication and mention of the grant of the patent:
22.09.2004 Bulletin 2004/39

(21) Application number: **00921641.7**

(22) Date of filing: **04.04.2000**

(51) Int Cl.[7]: **A61K 7/48**

(86) International application number:
PCT/US2000/008871

(87) International publication number:
WO 2001/074327 (11.10.2001 Gazette 2001/41)

(54) COMPOSITION FOR IMPROVING SKIN LIPID BARRIER FUNCTION

ZUBEREITUNG ZUR VERBESSERUNG DER EPITHELIALEN LIPIDBARRIEREFUNKTION

COMPOSITION DESTINEE A AMELIORER LA BARRIERE LIPIDIQUE DE LA PEAU

(84) Designated Contracting States:
AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

(43) Date of publication of application:
08.01.2003 Bulletin 2003/02

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EP 1 272 154 B1

Printed by Jouve, 75001 PARIS (FR)

Description

Field of the Invention

[0001] The invention relates to cosmetic and pharmaceutical compositions. More specifically, the invention relates to topical compositions that are useful in enhancing the function of the skin's natural lipid barrier.

Background of the Invention

[0002] Skin is typically characterized as consisting of three distinct layers, namely the stratum corneum, the epidermis and the dermis. The stratum corneum, the outermost layer, is made up of keratinized cells, surrounded by intercellular space filled with lipids. The stratum corneum provides a substantial physical barrier to penetration of most substances to the lower layers of the skin. In addition to preventing transport of substances to the other skin layers, however, this barrier also aids in prevention of water loss from the skin. Both functions are primarily attributable to the presence of the lipids in the stratum corneum.

[0003] There are two sources of the skin surface lipids making up this important barrier: sebaceous glands and the epidermis. The lipids are a diverse group of compounds, comprising triglycerides, diglycerides, ceramides, free fatty acids, wax esters, cholesterol and cholesterol esters, and squalene. The quantity and composition of the skin surface lipids differ from place to place on the body, and may to some extent be related to the number of sebaceous glands in a given area of the skin. The condition of the skin surface lipids may also be affected by an essential fatty acid deficiency. Additionally, the lipid barrier is easily diminished by exposure to harsh detergents or soaps. It is apparent, then, that the quality of the skin lipid barrier can vary widely, depending on a number of different factors, and therefore, may not always be adequate to perform its protective function optimally.

[0004] As an attempt to compensate for what may be a less than adequate lipid barrier, cosmetic compositions frequently incorporate components which compensate for water loss. Examples of such materials are hygroscopic humectants, e.g., urea or propylene glycol, which hold water on the skin; or emollients, e.g., oleyl alcohol or caprylic/capric triglycerides. Certain cosmetic components may be occlusive skin conditioners, which are used to provide an "artificial" barrier; such compounds are frequently lipids which remain on the skin surface, and include various hydrogenated oils, waxes and butters. Although many of these products provide an effective means of stemming water loss from the skin, they do have to be reapplied frequently to maintain the effect, and do not generally constitute a natural-occurring component of the stratum corneum, potentially giving rise to an unnatural, greasy feel to the skin. In addition, various pharmaceutical or cosmetic active agents are also frequently used to treat the symptoms of dry skin-associated conditions; however, in many cases, particularly with pharmaceutical agents, the treatments themselves may cause undesirable side effects in the individual being treated, while ultimately resulting in no actual repair of the lipid barrier.

[0005] The most desirable situation, from a functional point of view, is to find a way to enhance the skin's own ability to maintain and/or repair the strength of its barrier, so that the protective barrier formed is completely natural. It has now been discovered that a combination of specific skin active agents results in an unexpected increase in the strength of the natural barrier, by stimulating the production and maintenance of the barrier's naturally occurring components. There is thus provided a new type of cosmetic or pharmaceutical composition which functions by enhancing the skin's own functions, resulting in a more natural means of preventing dry skin and other undesirable results of a deficient lipid barrier.

Summary of the Invention

[0006] The invention relates to cosmetic and pharmaceutical compositions comprising lipid barrier-enhancing effective amounts of at least one protease inhibitor and at least one cellular differentiation enhancer. The composition can also comprise, in a preferred embodiment, effective amounts of a sterol sulfate, an at least one naturally occurring skin lipid component. The composition of the invention can be used in a method for strengthening the natural lipid barrier of the skin, as well as other methods of skin treatment that are made possible by the strengthening of the barrier.

Detailed Description of the Invention

[0007] The combined actives of the composition of the invention have been found to be highly effective in stimulating the repair of a damaged lipid barrier, and therefore, are shown to be useful as well in maintenance of a normal and healthy lipid barrier. For the purposes of the present invention and claims, these abilities will be referred to as strengthening the lipid barrier. The compositions of the invention can strengthen the lipid barrier at least about 40%, relative to a placebo control, as measured by a reduction of transepidermal water loss (TEWL) after a barrier challenge, a standard measurement of barrier function.

Preferably, the compositions are capable of reducing TEWL at least about 50%, more preferably at least about 60%, and most preferably at least about 70%. The invention incorporates as essential elements a protease inhibitor and a cellular differentiation enhancer. Protease inhibitors are compounds, usually naturally occurring, which inhibit the action of proteases in the skin. Skin proteases also occur naturally, and, among other effects, are involved in the breaking down of the collagen and elastin that is required to maintain the healthy

appearance of skin. In the present case, the protease inhibitors used in the invention are thought to act by preventing the breakage of the desmosome bond between corneocytes at the skin surface, thereby keeping the outer layer of skin cells intact, in essence delaying desquamation. This skin layer provides a barrier to water loss, and the enhanced retention of the barrier by the delay of desquamation provides reinforces this barrier to the loss of water further. A variety of protease inhibitors are known. Examples of useful protease inhibitors include, but are not limited to, triterpenoid-containing extracts and refined compounds, for example, white birch bark extract, silver birch bark extract, *Boswellia* extract, bearberry extract, *Centella asiatica* extract, or *Pygeum* (Prunus) *africanum* extract and individual protease inhibitor compounds that may be present in these extracts, including betulinol(betulin), betulinic acid, boswellic acid, ursolic acid, oleanolic acid, oleanol, asiaticoside, asiatic acid, and madagassic acid; phenolic-containing extracts, such as green tea extracts and apple extracts, and compounds contained therein, such as EGCG, ECG, catechins, phenylpropanoids, and phloretin; protein-based extracts, such as soy protein, or egg protease inhibitors, or cholesterol sulfate and phytosterol sulfates. Preferred protease inhibitors are triterpenoids, particularly boswellic acid, betulinol, and betulinic acid, or extracts containing same in substantial quantities.

[0008] It will be recognized from the foregoing that either an extract or the individual protease inhibitor can be used, and that the individual protease inhibitors can also be found in other types of extracts. The amount of active material used will vary depending upon the whether an extract or isolated compound is used, the concentration of active material in a given extract, and the known potency of the active material. However, the concentration of active protease inhibitor in the final product should generally be between 0,001 to 10%, preferably 0,05 to 5%, more preferably 0,1 to 1%, by weight of the total composition.

[0009] The second component is a cellular differentiation enhancer. Such compounds act in the present invention to increase the ability of the relevant epidermal cells to synthesize the lipids that constitute the primary component of the barrier. The epidermal cells that produce lipids do not do so during their entire life cycle, however, but do so only at the point of differentiation. If this differentiation is delayed to the point at which the epidermal cells reach their terminal point in migration to the skin surface, the production of lipids is concurrently delayed, and therefore, their effect, if any, is diminished. The differentiation enhancers used in the invention stimulate the earlier production of lipids from epidermal cells, thereby increasing the length of time over which the lipids are being produced, and presumably concurrently increasing the lipid content of the barrier. Several types of cellular differentiation inhibitors are known, and these include, but are not limited to sclareolide, forskolin, 7-dehydrocholesterol, and Vitamin D3 analogs. The differentiation enhancer also will be used in amount consistent with its known activity, 0,001 to 10%, preferably 0,0025 to 5%, more preferably 0,05 to 1%, by weight of the total composition.

[0010] The combination of the protease inhibitor and the differentiation enhancer is shown to have a strong effect on barrier repair when compared with control vehicles and with other compounds commonly used in skin enhancement. Specifically, on skin that had been challenged with tape stripping, it was found that combination of these two components exhibits about a 50% increase in barrier repair, as measurable by transepidermal water loss, in comparison with the placebo, after a period of three days. Thus, this combination on its own is shown to be able to increase barrier function substantially.

[0011] Although the noted combination is highly effective on its own, it has been further shown that the combination achieves even further enhancement of barrier function by its combination with certain other skin enhancement compounds. More specifically, the combination with one or more of certain specific skin enhancers. In a preferred embodiment, the barrier repair combination is further combined with cholesterol sulfate, which in the present specification and claims is intended to refer to the corresponding plant-derived material, phytosterol sulfate. Cholesterol sulfate, as described in Applicants' copending application serial no. 09/246,607, also has an effect on the skin, by increasing the cohesion of the stratum corneum. Thus, its combination with the protease inhibitor/differentiation enhancer even further strengthens the ability of the composition maintain the integrity of the stratum corneum. The amount of cholesterol sulfate employed is preferably about 0.05 to about 10%, preferably from about 0.5 to about 5%, most preferably about 1 to about 3%, by weight of the total composition.

[0012] In all formulations in which cholesterol sulfate is employed, and in a particularly preferred embodiment, it is preferred that the composition also contain other components of the naturally occurring lipid barrier. In a particularly preferred embodiment, the cholesterol sulfate is combined with at least one of each of fatty acids, ceramides, and a sterol, preferably cholesterol or phytosterol. Fatty acids may be up to 24 carbon atoms in length. Examples of preferred fatty acids include butyric acid, caproic acid, octanoic acid, decanoic acid, dodecanoic acid, tetradecanoic acid, palmitic acid, stearic acid, linoleic acid and oleic acid. Particularly preferred are fatty acids with a C_{12} to C_{20} chain length.

[0013] The ceramides to be employed in the compositions of the invention are sphingolipids, having a sphingosine or related molecule backbone with fatty acids or ω-esterified fatty acids linked to an amino group on the sphingosine, and in some cases, with saccharide moieties linked to the terminal hydroxyl of the sphingosine. In particular, the compositions may contain ω-esterified ceramides or acylceramides, cerebrosides, ω-esterified

cerebrosides, or acylglycosyl sphingolipids. Particularly preferred types of ceramides for the present compositions are ceramide III and cerebrosides.

[0014] In those compositions in which cholesterol sulfate is combined with these lipids, the lipid components each can be used in an amount of from about 0.05 to 10%, preferably 0.5 to about 5%, most preferably about 1 to about 3%, all by weight of the total composition. In a particularly preferred embodiment, the cholesterol sulfate and the lipid components are present in substantially equal amounts in the composition. It will be understood from the foregoing that the lipid component need not be pure lipid, but rather may be natural extracts containing one or more desirable lipids, and used in amounts consistent with attaining the concentrations recommended above.

[0015] The effect of the present compositions in effecting barrier repair or maintaining the integrity of the skin's outer layer can be applied to a number of different uses. For example, the compositions can be used to treat any condition in which a deficient or faulty barrier is a factor. In this regard, the compositions can be used to improve the long term moisture retention of the skin, or in prevention or treatment of dry skin conditions generally, or specific dry skin conditions, such as result from regular exposure to detergents, soaps and hot water; seasonal exposure to harsh weather conditions, e.g., cold, wind and/or sun; occupational exposure to harsh chemicals or other drying or damaging agents; or pathological conditions such as eczematous dermatides, psoriasis, ichthyoses, xerosis and the like. It is also well-known that dry skin is commonly associated with aging (both intrinsic and photoaging), and the compositions can be used in prevention of further damage to aging skin, or treatment and/or reversal of already present damage, including the appearance of fine lines and wrinkles, which are frequently associated with dry skin and the thinning of the stratum corneum that occurs with age. The compositions can also be used in the treatment of a defective skin barrier, such as occurs on the soles of the feet, and palms of the hands, where the stratum corneum is very thick, but the lipid barrier is poor. In addition, defective skin barriers frequently occur in association with burns, wounds, blisters, stasis ulcers and bedsores; such injuries can be expected to benefit from application of the compositions.

[0016] A further use of the compositions of the invention is in reduction of the skin's response to irritants and sensitizers. A significant percentage of the population considers itself to have sensitive skin, in that they perceive a frequent, stinging or painful response to various elements to which the skin may be exposed, be it through makeup or skin care products, environmental stimuli such as smoke or pollution, or occupational exposure to chemicals. In addition, even normal skin can have a reaction to exposure to known irritants, such as acids. As it is well known that the stratum corneum and lipids constitute the first line of defense against irritants, by providing a physical barrier to permeability of such materials to the lower skin layers, the application of the compositions of the invention, by increasing the integrity of the barrier, can reduce the reactivity of the skin of both normal and sensitive individuals to irritants and sensitizers. In one embodiment, for example, the compositions can be used to reduce the reaction of the skin to the irritation caused by therapeutic acids such as alpha and beta hydroxy acids, retinoic acid , or to reduce the irritation caused by insect bites or stings, or alleviate the irritation experienced with contact dermatitis.

[0017] The increased cohesion of the stratum corneum brought about by the compositions of the invention also provides other benefits. The stratum corneum represents an important physical barrier between the environment and the deeper skin layers as well as the internal organs. The presence of this thicker layer thus will provide a greater level of protection than is possible with weaker barrier. Perhaps the most important aspect of this effect is the enhanced self-protection from UV rays. The thicker stratum corneum means an increase in the Minimal Erythemal Dose of UV which will result in sunburn or more serious skin damage. In connection with this aspect of the invention, the components of the invention may be beneficially combined with one or more sunscreens for an enhanced UV protective composition which provides both short- and long-term protection. Thus, the invention provides sunscreen compositions comprising effective amounts of the components of the composition of the invention, and one or more sunscreens. Examples of useful sunscreens include, but are not limited to, inorganic sunscreens such as titanium dioxide, zinc oxide, and iron oxide; and organic sunscreens, such as camphor derivatives, cinnamates, salicylates, benzophenones, triazines, PABA derivatives, diphenylacrylate derivatives, and dibenzoylmethane derivatives. In such sunscreen compositions, the components of the invention are present in the amounts described above, and the respective sunscreens are present in the amounts normally used for UV protection.

[0018] An additional use of the compositions of the invention is in the enhancement and prolongation of self-tanning products. One of the recognized limitations of self-tanners, which are normally based on dihydroxyacetone(DHA) as the active component, is that the tan on the skin lasts only as long as the skin cells receiving the DHA remain in place. In the normal course of events, then, a self-applied tan usually lasts no more than 5 days, i.e., for as long as it takes for the stratum corneum layer to which the DHA was applied to fully turn over. When the compositions of the invention are combined with DHA, or any other self-tanning agent, in a typical self-tanning formulation, however, the rate of turnover of the stratum corneum to which the composition is applied is slowed down, thereby permitting a longer rate of retention of the "tanned" cells, and thus prolonging the length of time the tan remains visible on the skin. Thus, the invention provides a self-tanning composition

comprising a protease inhibitor, a cell-differentiation enhancer, and an effective amount of a self-tanning agent, optionally containing cholesterol sulfate and the lipid component. In a preferred embodiment, the self-tanner is DHA, which is usually applied in an amount of from about 2.5 to about 10% by weight of the formulation. The self-tanner may also be imidazole, preferably in combination with DHA, in an amount of about 1-10%, preferably about 1.5-7.5%.

[0019] The compositions of the invention are employed in a manner appropriate to the intended final use of the product. For example, in the treatment of occasional dry skin due to exposure to weather or other temporary conditions, or in the treatment of occasional skin irritation, the compositions can be used on an as-needed basis until the condition is relieved. When being used to treat a more permanent condition, for example, a condition associated with a defective or deficient lipid barrier, particularly sensitive skin, dry skin associated with any type of aging, or the wrinkling or fine lines associated with a thinning of the stratum corneum with aging, the composition is preferably applied chronically, to prevent recurrence of the condition. For this purpose, it is suggested as an example that topical application of the composition, in an amount of from about 0.1 mg/cm^2 to 2 mg/cm^2 of skin, be performed from about once per week to about 4 or 5 times daily, preferably from about 3 times a week to about 3 times daily, most preferably about once or twice per day. By "chronic" application, it is meant herein that the period of topical application may be over the lifetime of the user, preferably for a period of at least about one month, more preferably from about three months to about twenty years, more preferably from about six months to about ten years, more preferably still from about one year to about five years, thereby resulting in the treatment or prevention of the condition in question.

[0020] When the composition is used in conjunction with a sunscreen, it is applied in the same amounts as specified above, on an as-needed basis, to mitigate the effects of exposure to the sun. When used in combination with a self-tanner, the composition is also applied in similar amounts, on the portion of the skin to be tanned, with repetition, again, on an as-needed basis.

[0021] The invention is further illustrated by the following nonlimiting examples:

EXAMPLES

Example 1.

[0022] A study is conducted to determine the efficacy of certain compositions in enhancing repair of the lipid barrier. Female volunteers with normal skin, who are in good general health, free of any dermatological disorders, participate in the study. Their skin barrier is challenged by tape stripping according to the procedure outlined below. After their evaluation, the subjects receive a treatment every day on the right side of their face. They are also given the product containing the actives to use once every night for three nights, on the right side of the face only, the left side being the untreated control side. A separate group is given a placebo, without actives, as treatment product.

[0023] To challenge the barrier, the subjects are acclimated in an environmental room at 40% relative humidity and 70°C for 15-20 minutes. A 5 cm by 1cm area is marked on the lower right cheek near the jaw line and initial water evaporation measurements are taken in three separate spots approximately 1 cm apart in a row. Five cm of cello-tape is placed on the skin in the outlined area, starting from the top of the cheek and after one firm stroke in each direction it is removed by gently pulling in a downward direction parallel to the skin. The procedure is repeated and water evaporation is measured after every five strips until the barrier is disrupted as indicated by a minimum of 18g/sq.m hr on one of the three spots. Both sides of the face are stripped in the same way. The subjects returned for TEWL evaluation 1,2 and 3 days after tape stripping of the skin to monitor the repair.

[0024] Barrier repair is evaluated by first challenging the skin as described above. One side is product-treated 2 times a day, and the other is the untreated control. Repair is measured in the increase in the recovery of the skin on the stripped and treated site compared to the stripped untreated site. From this, total repair is calculated over three days by calculating the change in the area parameter. The smaller the area, the faster the repair.

[0025] Treatment products are (1) a composition containing 0.1% sclareolide and 0.2% white birch extract; (2) a composition containing 0.1% sclareolide and 0.2% boswellic acid; (3) a composition containing .2% each of phytocohesine(phytosterol sulfate), cennamides (wheat-derived ceramides), boswellic acid, cholesterol, and linoleic acid, and .1% sclareolide. The results obtained indicate for composition (1), barrier repair is 50% over placebo; for composition (2), repair is 59% over placebo, and composition (3) is shows barrier repair at 78% over the placebo, each composition therefore showing substantial efficacy in barrier repair.

Claims

1. A composition for topical application to the skin comprising effective amounts of at least one protease inhibitor and at least one cell differentiation enhancer.

2. The composition of claim 1 in which the protease inhibitor is selected from the group consisting of triterpenoid-containing extracts, and active components thereof, phenolic-containing extracts and active components thereof, and protein-based extract

and active components thereof.

3. The composition of claim 1 in which the protease inhibitor is selected from the group consisting of white birch extract, silver birch extract, Boswellia extract, bearberry extract, Centella asiatica extract, Pygeum (Prunus) africanum extract, betulinol, betulinic acid, boswellic acid, ursolic acid, oleanolic acid, oleanol, asiaticoside, asiatic acid, madagassic acid, green tea extract, apple extracts, EGCG, ECG, catechins, phenylpropanoids, phloretin, soy protein, egg protease inhibitors, cholesterol sulfate, phytosterol sulfate, and combinations thereof.

4. The composition of claim 3 in which the protease inhibitor is selected from the group consisting of white birch extract, betulinol, betulinic acid, Boswellia extract, boswellic acid, and combinations thereof.

5. The composition of any of claims 1 to 4 in which the cell differentiation enhancer is selected from the group consisting of forskolin, sclareolide, 7-dehydrocholesterol, and Vitamin D3 analogs.

6. The composition of claim 5 in which the enhancer is sclareolide.

7. The composition of any of claims 1 to 6 comprising 0,001 to 10% of a protease inhibitor and 0,001-10% cell differentiation enhancer.

8. The composition of claim 7 comprising 0,1 to 1% of a protease inhibitor and 0,05-1% cell differentiation enhancer.

9. The composition of any of claims 1 to 8 which also contains cholesterol sulfate.

10. The composition of any of claims 1 to 9 which also contains at least one fatty acid, at least one ceramide, and at least one sterol.

11. The composition of claim 10 in which the fatty acid is a C12-C20 fatty acid.

12. The composition of claim 10 or 11 containing 0,05 to 10 % of each of cholesterol sulfate or phytosterol sulfate, fatty acid, ceramide, and sterol.

13. The composition of any of claims 1 to 12 which further comprises an effective amount of at least one self-tanning agent.

14. The composition of claim 13 in which the self-tanning agent is DHA.

15. The composition of any of claims 1 to 14 which further comprises at least one sunscreen.

16. A cosmetic method for treatment or prevention of dry skin which comprises applying to the skin a composition according to any of claims 1 to 15.

17. A cosmetic method for reducing or preventing skin's response to irritants or sensitizers comprising applying to the skin a composition of any of claims 1 to 15.

18. A cosmetic method for protecting skin against effects of exposure to UV radiation comprising applying to the skin a composition of any of claims 1 to 15

19. A method for tanning the skin without exposure the sun comprising applying to the skin a composition according to claim 13 or 14.

Patentansprüche

1. Eine Zusammensetzung für die örtliche Anwendung auf der Haut, die wirksame Mengen wenigstens eines Proteaseinhibitors und wenigstens eines Zelldifferenzierungsverstärkers umfasst.

2. Die Zusammensetzung nach Anspruch 1, bei der der Proteaseinhibitor aus der Gruppe, die aus triterpenoidhaltigen Extrakten und aktiven Komponenten davon, phenolhaltigen Extrakten und aktiven Komponenten davon und einem Extrakt auf Proteinbasis und aktiven Komponenten davon besteht, ausgewählt wird.

3. Die Zusammensetzung nach Anspruch 1, bei der der Proteaseinhibitor aus der Gruppe, die aus Weißbirkenextrakt, Silberbirkenextrakt, Boswellia-Extrakt, Bärentraubenextrakt, Centella asiatica-Extrakt, Pygeum (Prunus) africanum-Extrakt, Betulinol, Betulinsäure, Boswelsäure, Ursolsäure, Oleanolsäure, Oleanol, Asiaticosid, Asiasäure, Madagaskarsäure, Grüner Tee-Extrakt, Apfelextrakt, EGCG, ECG, Catechinen, Phenylpropanoiden, Phloretin, Sojaprotein, Eiproteaseinhibitoren, Cholesterolsulfat, Phytosterolsulfat und Kombinationen davon besteht, ausgewählt wird.

4. Die Zusammensetzung nach Anspruch 3, bei der der Proteaseinhibitor aus der Gruppe, die aus Weißbirkenextrakt, Betulinol, Betulinsäure, Boswellia-Extrakt, Boswelsäure und Kombination davon besteht, ausgewählt wird.

5. Die Zusammensetzung nach einem der Ansprüche 1 bis 4, bei der der Zelldifferenzierungsverstärker aus der Gruppe, die aus Forskolin, Sclareolid, 7-Dehydrocholesterol und Vitamin D3-Analoga be-

steht, ausgewählt wird.

6. Die Zusammensetzung nach Anspruch 5, bei der der Verstärker Sclareolid ist.

7. Die Zusammensetzung nach einem der Ansprüche 1 bis 6, die 0.001-10% eines Proteaseinhibitors und 0.001-10% eines Zelldifferenzierungsverstärkers umfasst.

8. Die Zusammensetzung nach Anspruch 7, die 0.1-1% eines Proteaseinhibitors und 0.05-1% eines Zelldifferenzierungsverstärkers umfasst.

9. Die Zusammensetzung nach einem der Ansprüche 1 bis 8, die ebenfalls Cholesterolsulfat enthält.

10. Die Zusammensetzung nach einem der Ansprüche 1 bis 9, die ebenfalls wenigstens eine Fettsäure, wenigstens ein Ceramid und wenigstens ein Sterol enthält.

11. Die Zusammensetzung nach Anspruch 10, bei der die Fettsäure eine C_{12}-C_{20} Fettsäure ist.

12. Die Zusammensetzung nach Anspruch 10 oder 11, die 0.05-10% eines Cholesterolsulfats oder Phytosterolsulfats, einer Fettsäure, eines Ceramids und eines Sterols enthält.

13. Die Zusammensetzung nach einem der Ansprüche 1 bis 12, die weiter eine wirksame Menge wenigstens eines Selbstbräunungsmittels umfasst.

14. Die Zusammensetzung nach Anspruch 13, bei der das Selbstbräunungsmittel DHA ist.

15. Die Zusammensetzung nach einem der Ansprüche 1 bis 14, die weiter wenigstens eine Sonnencreme umfasst.

16. Ein kosmetisches Verfahren zur Behandlung oder Prävention von trockener Haut, das das Auftragen einer Zusammensetzung gemäß einem der Ansprüche 1 bis 15 auf die Haut umfasst.

17. Ein kosmetisches Verfahren zur Verringerung oder Prävention einer Hautreaktion auf Reizmittel oder Sensibilisierungsmittel, das das Auftragen einer Zusammensetzung nach einem der Ansprüche 1 bis 15 auf die Haut umfasst.

18. Ein kosmetisches Verfahren zum Schutz der Haut gegen die Einwirkung von UV-Strahlung, das das Auftragen einer Zusammensetzung nach einem der Ansprüche 1 bis 15 auf die Haut umfasst.

19. Ein Verfahren zum Bräunen der Haut ohne Einwirkung der Sonne, das das Auftragen einer Zusammensetzung gemäß Anspruch 13 oder 14 auf die Haut umfasst.

Revendications

1. Une composition pour application topique à la peau comprenant des quantités efficaces d'au moins un inhibiteur de protéase et au moins un activateur de différenciation cellulaire.

2. La composition selon la revendication 1, dans laquelle l'inhibiteur de protéase est choisi dans le groupe consistant en extraits contenant des triterpénoïdes et leurs composants actifs, des extraits contenant des substances phénoliques et leurs composants actifs, et des extraits à base de protéine et leurs composants actifs.

3. La composition selon la revendication 1, dans laquelle l'inhibiteur de protéase est choisi dans le groupe consistant en extrait de bouleau blanc, extrait de bouleau argenté, extrait de Boswellia, extrait de raisin d'ours, extrait de Centella asiatica, extrait de Pygeum (Prunus) africanum, betulinol, acide bétulinique, acide boswellique, acide ursolique, acide oléanolique, oléanol, asiaticoside, acide asiatique, acide madagassique, extrait de thé vert, extraits de pomme, EGCG, ECG, catéchines, phénylpropanoïdes, phlorétine, protéine de soja, inhibiteurs de protéase d'oeuf, sulfate de cholestérol, sulfate de phytostérol, et leurs combinaisons.

4. La composition selon la revendication 3, dans laquelle l'inhibiteur de protéase est choisi dans le groupe consistant en extrait de bouleau blanc, bétulinol, acide bétulinique, extrait de Boswellia, acide boswellique et leurs combinaisons.

5. La composition selon l'une quelconque des revendications 1 à 4 dans laquelle l'activateur de différenciation cellulaire est choisi dans le groupe consistant en forskoline, sclaréolide, 7-déhydrocholestérol et analogues de vitamine D3.

6. La composition selon la revendication 5, dans laquelle l'activateur est le sclaréolide.

7. La composition selon l'une quelconque des revendications 1 à 6, comprenant 0,001 à 10 % d'un inhibiteur de protéase et 0,001 à 10 % d'activateur de différenciation cellulaire.

8. La composition selon la revendication 7 comprenant 0,1 à 1 % d'un inhibiteur de protéase et 0,05 à 1 % d'activateur de différenciation cellulaire.

9. La composition selon l'une quelconque des revendications 1 à 8 qui contient en outre du sulfate de cholestérol.

10. La composition selon l'une quelconque des revendications 1 à 9 qui contient en outre au moins un acide gras, au moins une céramide et au moins un stérol.

11. La composition selon la revendication 10, dans laquelle l'acide gras est un acide gras en C12-C20.

12. La composition selon la revendication 10 ou 11 contenant 0,05 à 10 % de chacun des produits suivants : sulfate de cholestérol ou sulfate de phytostérol, acide gras, céramide et stérol.

13. La composition selon l'une quelconque des revendications 1 à 12 qui comprend en outre une quantité efficace d'au moins un agent auto-bronzant.

14. La composition selon la revendication 13 dans laquelle l'agent auto-bronzant est le DHA.

15. La composition selon l'une quelconque des revendications 1 à 14 qui comprent en outre au moins un écran solaire.

16. Une méthode cosmétique pour le traitement ou la prévention de la peau sèche qui comprend l'application sur la peau d'une composition selon l'une quelconque des revendications 1 à 15.

17. Une méthode cosmétique pour diminuer ou prévenir la réponse cutanée à des agents irritants ou sensibilisants comprenant l'application à la peau d'une composition selon l'une quelconque des revendications 1 à 15.

18. Une méthode cosmétique de protection de la peau vis-à-vis des effets de l'exposition au rayonnement UV comprenant l'application à la peau d'une composition selon l'une quelconque des revendications 1 à 15.

19. Une méthode de bronzage de la peau sans exposition au soleil comprenant l'application sur la peau d'une composition selon la revendication 13 ou 14.