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(54) Title: METHOD FOR TREATING ECZEMA

(57) Abstract: A method for treating eczema including the steps of applying an isothiocyanate functional surfactant to an area affected by eczema, wherein the isothiocyanate functional surfactant comprises at least one isothiocyanate functional group associated with an aliphatic and/or aromatic carbon atom of the isothiocyanate functional surfactant.



WO 2013/003560 A1

TITLE OF THE INVENTION

METHOD FOR TREATING ECZEMA

CROSS-REFERENCE TO RELATED APPLICATIONS

[0000] This application claims the benefit of United States Provisional Application Serial Number 61/431,977, entitled "METHOD FOR TREATING ECZEMA," filed January 12, 2011 and United States Provisional Application Serial Number 61/502,113, entitled "METHOD FOR TREATING ECZEMA," filed June 28, 2011 – all of which are hereby incorporated herein by reference in their entirety, including all references cited therein.

BACKGROUND OF THE INVENTION

1. Field of the Invention

[0001] The present invention relates in general to a method for treating eczema (*i.e.*, dermatitis) and, more particularly, to a method for treating a plurality of forms of eczema including, but not limited to, atopic eczema.

2. Background Art

[0002] Eczema is a general term for many types of skin inflammation, also known as dermatitis. The most common form of eczema is atopic eczema or dermatitis (*n.b.*, many practitioners use the terms eczema and dermatitis interchangeably). However, there are many other different forms of eczema including, contact eczema, allergic contact eczema, seborrheic eczema, nummular eczema, neurodermatitis, stasis dermatitis, dyshidrotic eczema – among others.

[0003] Eczema occurs in people of all races and can affect people of any age, although the condition is most common in infants, and about 85% of people have an onset prior to five years of age. Typically, eczema will permanently resolve by age three in only about one-half of affected infants. In others, the condition tends to recur throughout life. People with eczema often have a family history of the condition or a family history of other allergic conditions, such as asthma and/or hay fever. Approximately 20% of children and approximately 1%-5% of adults are believed to have eczema. This means that more than approximately 15 million people in the United States alone express symptoms of the disease. While eczema is not known to be contagious, it is believed to be at least partially inherited. As such, it is not uncommon to find members of the same family affected.

[0004] While doctors do not know the exact cause of eczema, a defect of the skin that impairs its function as a barrier, possibly combined with an abnormal function of the immune system, are believed to be important factors. Studies have shown that in people with atopic dermatitis, there are gene defects that lead to abnormalities in certain proteins (*e.g.*, filaggrin) that are important in maintaining the barrier of normal skin. Some forms of eczema can be triggered by substances that come in contact with the skin, such as soaps, cosmetics, clothing, detergents, jewelry, or sweat. Environmental allergens (substances that cause allergic reactions) may also cause outbreaks of eczema. Changes in

temperature or humidity, or even psychological stress, can lead to outbreaks of eczema in some people.

[0005] Eczema most commonly causes dry, reddened skin that itches and/or burns, although the appearance of eczema varies from person to person and varies according to the specific type of eczema. Intense itching is generally the first symptom in most people with eczema. Sometimes, eczema may lead to blisters and oozing lesions, but eczema can also result in dry and scaly skin. Repeated scratching may lead to thickened, crusty skin.

[0006] While any region of the body may be affected by eczema, in children and adults, eczema typically occurs on the face, neck, and the insides of the elbows, knees, and ankles. In infants, eczema typically occurs on the forehead, cheeks, forearms, legs, scalp, and neck.

[0007] Eczema can sometimes occur as a brief reaction that only leads to symptoms for a few hours or days, but in other cases, the symptoms persist over a longer time and are referred to as chronic dermatitis.

[0008] There are many different forms of eczema including atopic eczema or dermatitis, contact eczema, allergic contact eczema, seborrheic eczema, nummular eczema, neurodermatitis, stasis dermatitis, and dyshidrotic eczema.

[0009] Atopic dermatitis is a chronic skin disease characterized by itchy, inflamed skin and is the most common cause of eczema. The condition tends to come and go, depending upon exposures to triggers or causative factors. Factors that may cause atopic dermatitis (allergens) include environmental factors like

molds, pollen, or pollutants; contact irritants like soaps, detergents, nickel (present in jewelry), or perfumes; food allergies; or other allergies. Around two-thirds of those who develop the condition do so prior to one year of age. When the disease starts in infancy, it is sometimes termed infantile eczema.

5 [0010] Contact eczema (*i.e.*, contact dermatitis) is a localized reaction that includes redness, itching, and burning in areas where the skin has come into contact with an allergen (an allergy-causing substance to which an individual is sensitized) or with a general irritant such as an acid, a cleaning agent, or other chemical. Other examples of contact eczema include reactions to laundry
10 detergents, soaps, nickel (present in jewelry), cosmetics, fabrics, clothing, and perfume. Due to the vast number of substances with which individuals have contact, it can be difficult to determine the trigger for contact dermatitis. The condition is sometimes referred to as allergic contact eczema (*i.e.*, allergic contact dermatitis) if the trigger is an allergen, and irritant contact eczema (*i.e.*,
15 irritant contact dermatitis) if the trigger is an irritant. Skin reactions to poison ivy, oak and/or sumac are examples of allergic contact eczema. People who have a history of allergies have an increased risk for developing contact eczema.

[0011] Seborrheic eczema (*i.e.*, seborrheic dermatitis) is a form of skin inflammation of unknown cause. The signs and symptoms of seborrheic eczema
20 include yellowish, oily, scaly patches of skin on the scalp, face, and occasionally other parts of the body. Dandruff and "cradle cap" in infants are examples of seborrheic eczema. It is commonplace for seborrheic dermatitis to inflame the

face at the creases of the cheeks and/or the nasal folds. Seborrheic dermatitis is not necessarily associated with itching. This condition tends to run in families. Emotional stress, oily skin, infrequent shampooing, and weather conditions may all increase a person's risk of developing seborrheic eczema. One type of seborrheic eczema is also common in people with AIDS.

[0012] Nummular eczema (*i.e.*, nummular dermatitis) is characterized by coin-shaped patches of irritated skin – most commonly located on the arms, back, buttocks, and lower legs – that may be crusted, scaling, and extremely itchy. This form of eczema is relatively uncommon and occurs most frequently in elderly men. Nummular eczema is usually a chronic condition. A personal or family history of atopic dermatitis, asthma, or allergies increases the risk of developing the condition.

[0013] Neurodermatitis, also known as lichen simplex chronicus, is a chronic skin inflammation caused by a scratch-itch cycle that begins with a localized itch (*e.g.*, an insect bite) that becomes intensely irritated when scratched. Women are more commonly affected by neurodermatitis than men, and the condition is most frequent in people 20-50 years of age. This form of eczema results in scaly patches of skin on the head, lower legs, wrists, or forearms. Over time, the skin can become thickened and leathery. Stress can exacerbate the symptoms of neurodermatitis.

[0014] Stasis dermatitis is a skin irritation on the lower legs, generally related to the circulatory problem known as venous insufficiency, in which the

function of the valves within the veins has been compromised. Stasis dermatitis occurs almost exclusively in middle-aged and elderly people, with approximately 6%-7% of the population over 50 years of age being affected by the condition. The risk of developing stasis dermatitis increases with advancing age. Symptoms include itching and/or reddish-brown discoloration of the skin on one or both legs. Progression of the condition can lead to the blistering, oozing skin lesions seen with other forms of eczema, and ulcers may develop in affected areas. The chronic circulatory problems lead to an increase in fluid buildup or edema in the legs. Stasis dermatitis has also been referred to as varicose eczema.

[0015] Dyshidrotic eczema (*i.e.*, dyshidrotic dermatitis) is an irritation of the skin on the palms of hands and soles of the feet characterized by clear, deep blisters that itch and burn. The cause of dyshidrotic eczema is unknown. Dyshidrotic eczema is also known as vesicular palmoplantar dermatitis, dyshidrosis, or pompholyx. This form of eczema occurs in up to 20% of people with hand eczema and is more common during the spring and summer months and in warmer climates.

[0016] To the best of Applicant's knowledge, and until now, the traditional goals for the treatment of eczema were merely to minimize itching, inflammation, and/or worsening of the condition. Treatment for eczema typically involved both lifestyle changes and the use of medications.

[0017] In some cases corticosteroid creams have been prescribed to decrease the inflammatory reaction in the skin. However, such creams are

replete with drawbacks. In addition, two topical (cream) medications have been approved by the U.S. FDA for the treatment of eczema: tacrolimus (Protopic) and pimecrolimus (Elidel). These drugs belong to a class of immune suppressant drugs known as calcineurin inhibitors. In 2005, the FDA issued a warning about the use of these drugs, citing studies in animals that showed a possible association between the use of these drugs and the development of certain types of cancer. As such, use of calcineurin inhibitors appears to be problematic from carcinogenic and other perspectives.

[0018] While the above-identified medical treatments do appear to provide at least some relief to those who are afflicted by eczema, such treatment remains non-desirous and/or problematic inasmuch as, among other things, none of the above-identified treatments provide sufficient therapeutic relief from the debilitating effects of eczema without material drawbacks.

[0019] It is therefore an object of the present invention to provide a method for treating eczema which offers timely relief from the symptoms presented when one is afflicted with eczema.

[0020] These and other objects of the present invention will become apparent in light of the present specification, claims, and drawings.

SUMMARY OF THE INVENTION

[0021] In one embodiment, the present invention is directed to a method for treating eczema comprising the step of: applying an isothiocyanate functional surfactant to an area affected by eczema, wherein the isothiocyanate functional surfactant comprises at least one isothiocyanate functional group associated with an aliphatic and/or aromatic carbon atom of the isothiocyanate functional surfactant.

[0022] In another embodiment of the present invention, the method for treating eczema further comprises the step of removing the isothiocyanate functional surfactant from the area affected by eczema.

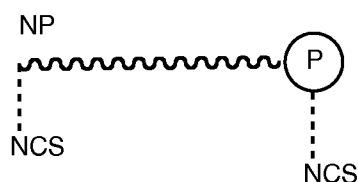
[0023] In yet another exemplary embodiment, the present invention is directed to a method for treating eczema comprising the steps of: (a) applying an isothiocyanate functional surfactant to an area affected by eczema, wherein the isothiocyanate functional surfactant comprises at least one isothiocyanate functional group associated with an aliphatic and/or aromatic carbon atom of the isothiocyanate functional surfactant; (b) removing the isothiocyanate functional surfactant from the area affected by eczema; and (c) repeating the steps of applying and removing the isothiocyanate functional surfactant to/from the affected area.

[0024] The present invention is also directed to a method for treating eczema comprising the step of: washing an area affected by eczema with an isothiocyanate functional surfactant, wherein the isothiocyanate functional

surfactant comprises at least one isothiocyanate functional group associated with an aliphatic and/or aromatic carbon atom of the isothiocyanate functional surfactant.

[0025] The present invention is further directed to a method for treating eczema comprising the step of: applying a lysine derivative to an area affected by eczema, wherein the lysine derivative comprises an α -nitrogen and a ϵ -nitrogen, and wherein an alkyl and/or alkanoyl substituent comprising at least approximately 8 carbon atoms is associated with the α -nitrogen, and further wherein at least one isothiocyanate functional group is associated with the ϵ -nitrogen.

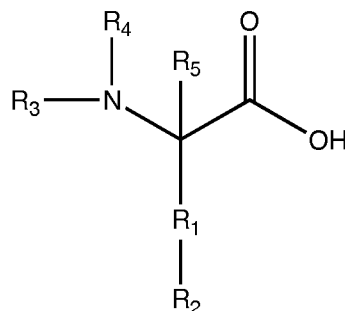
[0026] The present invention is still further directed to a method for treating eczema comprising the step of: applying a surfactant to an area affected by eczema, wherein the protonated form of the surfactant is represented by the following chemical structure:



wherein the surfactant comprises a non-polar moiety (NP) and a polar moiety (P), and wherein at least one isothiocyanate functional group (NCS) is associated with the polar and/or non-polar moiety.

[0027] In another embodiment, the present invention is directed to a method for treating eczema, comprising the step of: applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein

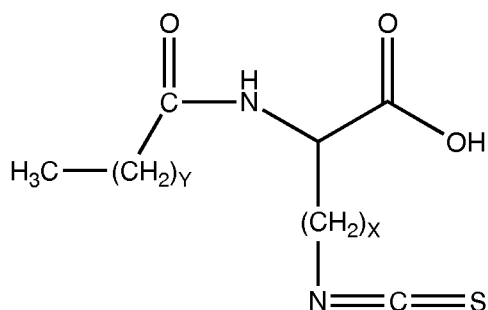
the protonated form of the surfactant is represented by the following chemical structure:



wherein R_1 comprises an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer; wherein R_2 comprises NCS; and wherein $\text{R}_3\text{-R}_5$ are the same or different and comprise H; OH; an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer with the proviso that at least one of $\text{R}_3\text{-R}_5$ comprise an alkyl, cycloalkyl, polycycloalkyl,

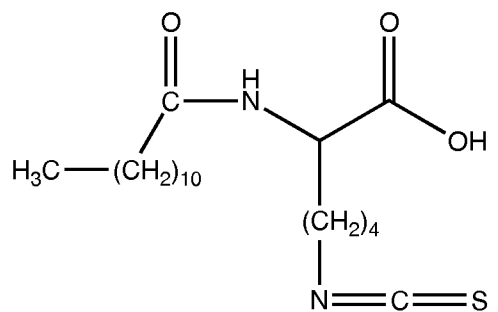
heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 8 to approximately 25 carbon atom(s).

[0028] The present invention is also directed to a method for treating eczema comprising the step of: applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein the protonated form of the surfactant is represented by the following chemical structure:

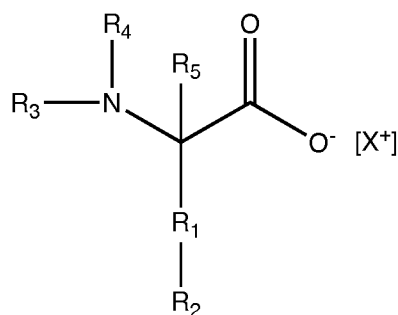


wherein X comprises an integer ranging from approximately 1 to approximately 25, and wherein Y comprises an integer ranging from approximately 6 to approximately 25.

[0029] In a preferred embodiment, the present invention is directed to a method for treating eczema comprising the step of: applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein the protonated form of the surfactant is represented by the following chemical structure:



[0030] In another embodiment, the present invention is directed to a method for treating eczema, comprising the step of: applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein the protonated form of the surfactant is represented by the following chemical structure:



wherein R₁ comprises an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the

carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer; wherein R₂ comprises NCS; wherein R₃-R₅ are the same or different and comprise H; OH; an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer with the proviso that at least one of R₃-R₅ comprise an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 8 to approximately 25 carbon atom(s), wherein X comprises a counter cation such as, but not limited to, alkali metals, alkaline earth metals, transition metals, s-block metals, d-block metals, p-block metals, NZ₄⁺, wherein Z comprises, H, R₆, and/or OR₆, and wherein R₆ comprises an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional

groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer.

[0031] In yet another preferred embodiment, the present invention is directed to a method for treating eczema as disclosed *supra*, further comprising the step of applying an additional surfactant, wherein the additional surfactant is selected from at least one of the group comprising a non-ionic surfactant, an anionic surfactant, a cationic surfactant, a zwitterionic surfactant, and combinations thereof.

DETAILED DESCRIPTION OF THE INVENTION

[0032] While this invention is susceptible of embodiment in many different forms, there is shown in the drawings and described herein in detail several specific embodiments with the understanding that the present disclosure is to be considered as an exemplification of the principles of the invention and is not intended to limit the invention to the embodiments illustrated.

[0033] In accordance with the present invention, surprisingly effective methods for treating eczema are provided herein. In particular, methods for treating a plurality of types of eczema including atopic eczema, contact eczema, allergic contact eczema, seborrheic eczema, nummular eczema, neurodermatitis, stasis dermatitis, and dyshidrotic eczema are disclosed.

[0034] In one embodiment, the present invention is directed to a method for treating eczema comprising the steps of applying one or more isothiocyanate functional surfactants to an area affected by eczema. Preferably, the isothiocyanate functional surfactant comprises one or more isothiocyanate functional groups associated with an aliphatic and/or aromatic carbon atom of the isothiocyanate functional surfactant. It will be understood that an area affected by eczema may comprise areas proximate and/or contiguous to areas where a manifestation of physical symptoms are present. Physical symptoms include, for example, discomfort, itching, burning, erythema, blistering, epidermal necrosis, desquamation, discoloration, and/or hyperpigmentation – just to name a few. It will be further understood that isothiocyanate functional surfactants, regardless of

their ordinary meaning, are defined herein as a surfactant having an isothiocyanate functional group associated therewith. It will be yet further understood that the term associated as used herein in chemical context, regardless of its ordinary meaning, is defined herein as attached, a covalent bond, a polar covalent bond, an ionic bond, a hydrogen bond, van der Waals forces, electrostatic interaction, directly and/or indirectly linked, etcetera.

[0035] The term surfactant derives from contraction of the terms *surface-active-agent* and is defined herein as a molecule and/or group of molecules which are able to modify the interfacial properties of the liquids (aqueous and non-aqueous) in which they are present. The surfactant properties of these molecules reside in their amphiphilic character which stems from the fact that each surfactant molecule has both a hydrophilic moiety and a hydrophobic (or lipophilic) moiety, and that the extent of each of these moieties is balanced so that at concentrations at or below the critical micelle concentration (*i.e.*, CMC) they generally concentrate at the air-liquid interface and materially decrease the interfacial tension. For example, sodium salts of saturated carboxylic acids are extremely soluble in water up to C8 length and are thus not true surfactants. They become less soluble in water from C9 up to C18 length, the domain of effective surfactants for this class of compounds. The carboxylic acids (fatty acids) can be either saturated or unsaturated starting from C16 chain lengths.

[0036] Without being bound by any one particular theory, it is believed that the isothiocyanate functional surfactants disclosed herein facilitate treatment of

numerous forms of eczema by boosting the body's immune system. It is also believed that the isothiocyanate functional surfactants disclosed herein facilitate elevating phase II enzymes (*e.g.*, HAD(P)H quinine oxidoreductase) which are believed to, among other things regulate inflammatory responses within the body, as well as detoxify carcinogens and/or activated carcinogens.

[0037] In accordance with the present invention, the isothiocyanate functional surfactants may be used as a topical leave-on product in which one or more surfactants remain on the skin and are not immediately and/or ever rinsed off away from the skin. Alternatively, the isothiocyanate functional surfactants of the present invention may be used as a topical wash in an apply-and-rinse fashion. For either case, it is preferred that the isothiocyanate functional surfactants be generally mild to human skin (*e.g.*, non-irritating or low-irritating). In particular, anionic N-alkanoyl surfactants derived from amino acids are especially preferred because, while not completely predictable, they have a tendency to be mild. The methods of preparation detailed in this invention employ, but are not limited to, amino acids that possess at least two amine functionalities, at least one of which is converted to an N-alkanoyl functionality, and at least one of which is converted into isothiocyanate functionality. The amino acids include, but are not limited to, the α -amino acids lysine, ornithine, 2,4-diaminobutanoic acid, 2,3-diaminopropionic acid, 2,7-diaminoheptanoic acid, and 2,8-diaminooctanoic acid. Additionally, amino acids other than α -amino acids may be employed, such as β -amino acids, etcetera. It will be understood

that amino acid derived surfactants are preferred due to their mild nature, but any one of a number of other surfactants are likewise contemplated for use in accordance with the present invention.

[0038] Methods for preparing isothiocyanate functional surfactants and/or their precursors can involve, but are not limited to, conversion of an amine functionality to an isothiocyanate functionality. The methods of conversion of amine functionalities to isothiocyanate functionalities include, but are not limited to: (1) reaction with carbon disulfide to yield an intermediate dithiocarbamate, followed by reaction with ethylchloroformate or its functional equivalent such as bis(trichloromethyl)-carbonate, trichloromethyl chloroformate, or phosgene; (2) reaction with thiophosgene; (3) reaction with 1,1'-thiocarbonyldiimidazole; (4) reaction with phenylthiochloroformate; (5) reaction with ammonium or alkali metal thiocyanate to prepare an intermediate thiourea followed by cleaving to the isothiocyanate via heating; and (6) reaction with an isothiocyanato acyl halide [SCN-(CH₂)_n-CO-Cl]. The resulting isothiocyanate functional surfactant, depending on the method of preparation, can be isolated as a pure material or as a mixture with other surfactants. The resulting isothiocyanate functional surfactant, depending on the method of preparation, can be isolated and used directly in nonionic form, anionic form, cationic form, zwitterionic (amphoteric) form, and/or in a neutral surfactant-precursor form in combination with a base such as sodium hydroxide or triethanol amine if the neutral surfactant-precursor form possesses a protonated carboxylic acid group such that reaction

(deprotonation) with the base converts the neutral surfactant-precursor form to an anionic surfactant, or in neutral surfactant-precursor form in combination with an acid if the neutral surfactant-precursor form possess amine functionality such that reaction (protonation) with the acid converts the neutral surfactant-precursor form to a cationic surfactant.

[0039] In accordance with the present invention the step of applying comprises, but is not limited to, spraying, dripping, dabbing, rubbing, blotting, dipping, and any combination thereof.

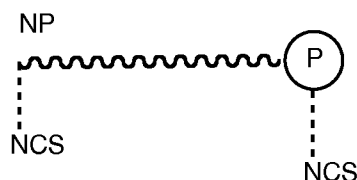
[0040] In a preferred embodiment of the present invention, the isothiocyanate functional surfactant is removed from the affected area after a period of time. Such a period comprises, but is not limited to, seconds (*e.g.*, 1 second, 2 seconds, 5 seconds, 10 seconds, 15 seconds, 20 seconds, 30 seconds, 45 seconds, and 60 seconds), minutes (*e.g.*, 1 minute, 2 minutes, 5 minutes, 10 minutes, 15 minutes, 20 minutes, 30 minutes, 45 minutes, and 60 minutes), hours (*e.g.*, 1 hour, 2 hours, 4 hours, 5 hours, 8 hours, 10 hours, 15 hours, 24 hours, 36 hours, 48 hours, and 60 hours), days (*e.g.*, 1 day, 2 days, 3 days, 4 days, 5 days, 6 days, 7 days, 10 days, 14 days, 21 days, 30 days), etcetera. It will be understood that the step of removing preferably occurs via rinsing, wiping, and/or extracting – just to name a few.

[0041] Depending upon the subject and/or the severity of the eczema, multiple applications may be necessary. As such, the steps of applying and/or

removing the isothiocyanate functional surfactant may be repeated one or a plurality of times.

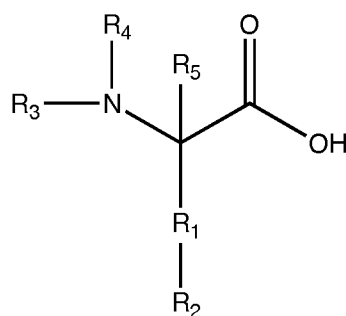
[0042] The present invention is also directed to a method for treating eczema comprising the steps of applying a lysine derivative to an area affected by eczema, wherein the lysine derivative comprises an α -nitrogen and a ϵ -nitrogen. Preferably, an alkyl substituent comprising at least approximately 8 carbon atoms is associated with the α -nitrogen. Preferably, at least one isothiocyanate functional group is associated with the ϵ -nitrogen.

[0043] The present invention is further directed to a method for treating eczema comprising the steps of: applying a surfactant to an area affected by eczema, wherein the surfactant is represented by the following chemical structure:



and wherein the surfactant comprises a non-polar moiety (NP) and a polar moiety (P), and wherein at least one isothiocyanate functional group (NCS) is associated with the polar and/or non-polar moiety.

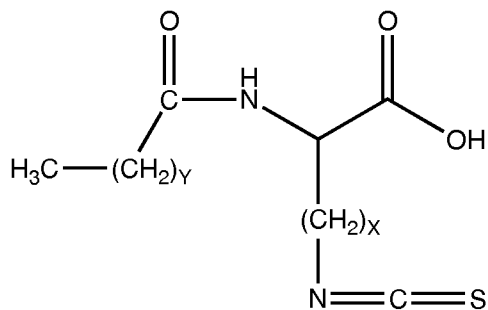
[0044] The present invention is yet further directed to a method for treating eczema, comprising the step of: applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein the protonated form of the surfactant is represented by the following chemical structure:



wherein R_1 comprises an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer; wherein R_2 comprises NCS; and wherein $\text{R}_3\text{-R}_5$ are the same or different and comprise H; OH; an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer with the proviso that at least one of $\text{R}_3\text{-R}_5$ comprise an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl

and/or cyano group containing approximately 8 to approximately 25 carbon atom(s).

[0045] In this embodiment, the surfactant is preferably represented by the following chemical structure:

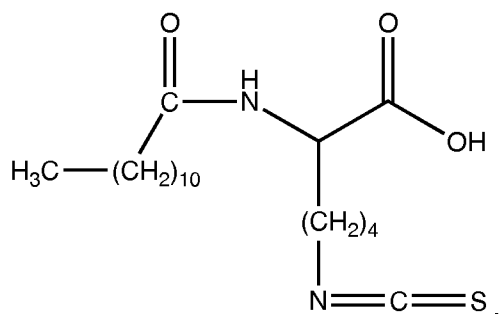


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wherein X comprises an integer ranging from approximately 1 to approximately 25, and wherein Y comprises an integer ranging from approximately 6 to approximately 25.

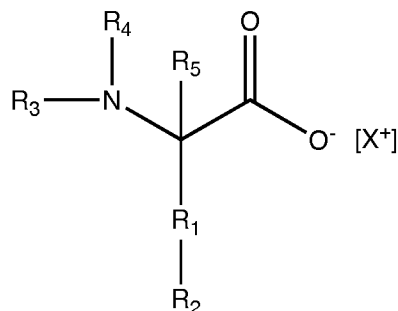
[0046] More preferably, the surfactant is represented by the following chemical structure:

10



[0047] In another embodiment, the present invention is directed to a method for treating eczema comprising the step of: applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein

the protonated form of the surfactant is represented by the following chemical structure:



wherein R₁ comprises an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer; wherein R₂ comprises NCS; wherein R₃-R₅ are the same or different and comprise H; OH; an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer with the proviso that at least one of R₃-R₅ comprise an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl,

aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 8 to approximately 25 carbon atom(s), wherein X comprises a counter cation such as, but not limited to, alkali metals, alkaline earth metals, transition metals, s-block metals, d-block metals, p-block metals, NZ_4^+ , wherein Z comprises, H, R_6 , and/or OR_6 , and wherein R_6 comprises an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer.

[0048] In accordance with the present invention, the isothiocyanate functional surfactant may also be associated with one or more additional surfactants, wherein the additional surfactants are selected from at least one of the group comprising a non-ionic surfactant, an anionic surfactant, a cationic surfactant, a zwitterionic surfactant, and combinations thereof.

[0049] Non-limiting examples of preferred anionic surfactants include taurates; isethionates; alkyl and alkyl ether sulfates; succinamates; alkyl sulfonates, alkylaryl sulfonates; olefin sulfonates; alkoxy alkane sulfonates; sodium and potassium salts of fatty acids derived from natural plant or animal sources or synthetically prepared; sodium, potassium, ammonium, and alkylated

ammonium salts of alkylated and acylated amino acids and peptides; alkylated sulfoacetates; alkylated sulfosuccinates; acylglyceride sulfonates, alkoxyether sulfonates; phosphoric acid esters; phospholipids; and combinations thereof. Specific anionic surfactants contemplated for use include, but are by no means limited to, ammonium cocoyl isethionate, sodium cocoyl isethionate, sodium lauroyl isethionate, sodium stearyl isethionate, sodium lauroyl sarcosinate, sodium cocoyl sarcosinate, sodium lauryl sarcosinate, disodium laureth sulfosuccinate, sodium lauryl sulfoacetate, sodium cocoyl glutamate, TEA-cocoyl glutamate, TEA cocoyl alaninate, sodium cocoyl taurate, potassium cetyl phosphate.

[0050] Non-limiting examples of preferred cationic surfactants include alkylated quaternary ammonium salts R_4NX ; alkylated amino-amides ($RCONH-(CH_2)_nNR_3X$); alkylimidazolines; alkoxylated amines; and combinations thereof. Specific examples of anionic surfactants contemplated for use include, but are by no means limited to, cetyl ammonium chloride, cetyl ammonium bromide, lauryl ammonium chloride, lauryl ammonium bromide, stearyl ammonium chloride, stearyl ammonium bromide, cetyl dimethyl ammonium chloride, cetyl dimethyl ammonium bromide, lauryl dimethyl ammonium chloride, lauryl dimethyl ammonium bromide, stearyl dimethyl ammonium chloride, stearyl dimethyl ammonium bromide, cetyl trimethyl ammonium chloride, cetyl trimethyl ammonium bromide, lauryl trimethyl ammonium chloride, lauryl trimethyl ammonium bromide, stearyl trimethyl ammonium chloride, stearyl trimethyl

ammonium bromide, lauryl dimethyl ammonium chloride, stearyl dimethyl cetyl
ditallow dimethyl ammonium chloride, dicetyl ammonium chloride, dilauryl
ammonium chloride, dilauryl ammonium bromide, distearyl ammonium chloride,
distearyl ammonium bromide, dicetyl methyl ammonium chloride, dicetyl methyl
5 ammonium bromide, dilauryl methyl ammonium chloride, distearyl methyl
ammonium chloride, distearyl methyl ammonium bromide, ditallow dimethyl
ammonium chloride, ditallow dimethyl ammonium sulfate, di(hydrogenated tallow)
dimethyl ammonium chloride, di(hydrogenated tallow) dimethyl ammonium
acetate, ditallow dipropyl ammonium phosphate, ditallow dimethyl ammonium
10 nitrate, di(coconutalkyl)dimethyl ammonium chloride, di(coconutalkyl)dimethyl
ammonium bromide, tallow ammonium chloride, coconut ammonium chloride,
stearamidopropyl PG-imonium chloride phosphate, stearamidopropyl
ethyldimonium ethosulfate, stearimidopropyldimethyl (myristyl acetate)
ammonium chloride, stearamidopropyl dimethyl cetearyl ammonium tosylate,
15 stearamidopropyl dimethyl ammonium chloride, stearamidopropyl dimethyl
ammonium lactate, ditallowyl oxyethyl dimethyl ammonium chloride,
behenamidopropyl PG dimonium chloride, dilauryl dimethyl ammonium chloride,
distearyl dimethyl ammonium chloride, dimyristyl dimethyl ammonium chloride,
dipalmityl dimethyl ammonium chloride, distearyl dimethyl ammonium chloride,
20 stearamidopropyl PG-dimonium chloride phosphate, stearamidopropyl
ethyldiammonium ethosulfate, stearamidopropyl dimethyl (myristyl acetate)
ammonium chloride, stearimidopropyl diemthyl cetaryl ammonium tosylate,

stearamido propyl dimethyl ammonium chloride, stearamidopropyl dimethyl ammonium lactate.

[0051] Non-limiting examples of preferred non-ionic surfactants include alcohols, alkanolamides, amine oxides, esters (including glycerides, ethoxylated glycerides, polyglyceryl esters, sorbitan esters, carbohydrate esters, ethoxylated carboxylic acids, phosphoric acid triesters), ethers (including ethoxylated alcohols, alkyl glucosides, ethoxylated polypropylene oxide ethers, alkylated polyethylene oxides, alkylated polypropylene oxides, alkylated PEG/PPO copolymers), silicone copolyols. Specific examples of non-ionic surfactants contemplated for use include, but are by no means limited to, cetearyl alcohol, cetareth-20, nonoxynol-9, C12-15 pareth-9, POE(4) lauryl ether, cocamide DEA, glycol distearate, glyceryl stearate, PEG-100 stearate, sorbitan stearate, PEG-8 laurate, polyglyceryl-10 trilaurate, lauryl glucoside, octylphenoxy-polyethoxyethanol, PEG-4 laurate, polyglyceryl diisostearate, polysorbate-60, PEG-200 isostearyl palmitate, sorbitan monooleate, polysorbate-80.

[0052] Non-limiting examples of preferred zwitterionic or amphoteric surfactants include betaines; sultaines; hydroxysultaines, amido betaines, amidosulfo betaines; and combinations thereof. Specific examples of amphoteric surfactants contemplated for use include, but are by no means limited to, cocoamidopropyl sultaine, cocoamidopropyl hydroxyl sultaine, cocoamidopropylbetaine, coco dimethyl carboxymethyl betaine, lauryl dimethyl carboxymethyl betaine, lauryl dimethyl alphacarboxyethyl betaine, cetyl dimethyl

carboxymethyl betaine, cetyl dimethyl betaine, lauryl (2-bishydroxy) carboxymethyl betaine, stearyl bis-(2-hydroxyethyl) carboxymethyl betaine, oelyl dimethyl gamma-carboxypropyl betaine, lauryl bis-(2-hydroxypropyl)alpha carboxymethyl betaine, coco dimethyl sulfopropyl betaine, stearyl dimethyl sulfopropyl betaine, lauryl dimethyl sulfoethyl betaine, lauryl bis(2-hydroxyethyl) sulfopropyl betaine, oleyl betaine, cocamidopropyl betaine.

[0053] The invention is further described by the following examples.

EXAMPLE 1

Preparation of a mixture of N_α-lauroyl-N_ε-isothiocyanato-L-Lysine with N_α,N_ε-bis-lauroyl-L-lysine

[0054] A 1 liter beaker equipped with an overhead mechanical stainless steel paddle stirrer was charged with 100 mL of 1 M NaOH (0.100 mol). Stirring was initiated and the beaker cooled to - 5 °C to -10 °C using a salt/ice bath. Next, 23.4 g (0.100 mol) of N_ε-benzylidene-L-lysine (prepared via the method of Bezas, B and Zervas, L., JACS, 83, 1961, 719-722) was added. Immediately afterward and while keeping the solution cold, 140 mL (0.140 mol) of precooled (in a salt/ice bath) 1 M NaOH and 26.1 mL of lauroyl chloride was added in two equal portions over a period of 6 minutes. The mixture was stirred for 10 more minutes at -5 to -10 °C, then the ice bath was removed and the reaction mixture allowed to stir for another 1 hour while warming to room temperature. Next, the reaction mixture was cooled using a salt/ice bath and then sufficient concentrated HCl was added to adjust the pH to 7.5-7.8. With

the pH at 7.8-7.8 and with continued cooling and stirring, 4.6 mL (60% of stoichiometric, 0.068 mol) of thiophosgene was added drop-wise via an additional funnel over the period of 1 hour. During this time, sufficient 1 M NaOH was added to maintain a pH range between 7.5-7.8. After the thiophosgene addition was complete, additional 1 M NaOH was added as necessary until the pH stabilized in 7.5-7.8 range. Next, sufficient 30% NaOH was added to adjust the pH to approximately 8.5. Next, 12 mL (0.051 mol) of lauroyl chloride was rapidly added, followed by sufficient 1 M NaOH to keep the pH in the range of 8.00 - 8.50. Next, sufficient concentrated HCl was added to adjust the pH to 1.5. The reaction mixture was filtered via vacuum filtration, and the precipitate washed with dilute HCl (pH = 2). The product, a white moist solid, was dried in vacuo while heating to 60 °C. 45.19 g of white solid product was recovered, a mixture of predominantly N_α-lauroyl-N_ε-isothiocyanato-L-lysine and N_α,N_ε-bis-lauroyl-L-lysine (determined via LC-MS analysis). Both compounds in this mixture can be simultaneously converted into anionic (carboxylate) surfactants via reaction with aqueous NaOH to yield a clear aqueous solution of the surfactants.

EXAMPLE II

Preparation of Pure N_α-lauroyl-N_ε-isothiocyanato-L-Lysine

Step 1: Preparation of N_α-lauroyl-N_ε-carbobenzoxy-L-Lysine

[0055] 60.0 g of N_ε-cbz-L-Lysine (cbz is carbobenzoxy) purchased from
5 Atomole Scientific Company, LTD was added to a three-liter beaker along with
1200 mL of RO water and the mixture was stirred. Next, 39 mL of 30% aqueous
NaOH was added, resulting in dissolution of the N_ε-cbz-L-Lysine. The resulting
solution was cooled in an ice bath and then 52.5 mL of lauroyl chloride was
added. The ice bath was removed 30 minutes later, and stirring continued for an
10 additional six hours, at which time 18 mL of concentrated hydrochloric acid was
added. The reaction mixture was then filtered via vacuum filtration, the white solid
product washed with 1 M aqueous HCl, and then the solid product was dried in
vacuo while heated to approximately 85 °C. 96.5 g of dry white solid product was
obtained. The product is further purified by dissolving it in methanol, filtering off
15 any insoluble precipitate, and removing the methanol in vacuo to recover a white
solid product (mp 99.5 - 103.0 °C)

Step 2: Preparation of N_α-lauroyl-N_ε-ammonium chloride-L-Lysine

[0056] 10.0 g of N_α-lauroyl-N_ε-carbobenzoxy-L-Lysine was weighed into a
one liter Erlenmeyer flask equipped with a magnetic stir bar. 150 mL of
20 concentrated hydrochloric acid was added and the solution was stirred and
heated in an oil bath to 104 °C, then allowed to cool with the oil bath back to
room temperature. The solution was then cooled to 9 °C for approximately four

hours, during which time a large mass of white precipitate formed. The reaction mixture was filtered in vacuo and rinsed with a small amount of cold 1 M HCl. The white solid reaction product was then dried in vacuo while being heated to 78 °C, yielding 7.89 g of white solid product (mp 191 - 193 °C).

5 Step 3: Preparation of N_α-lauroyl-N_ε-isothiocyanato-L-Lysine

[0057] 0.46 mL of thiophosgene was added to 30 mL of dichloromethane in a 125 mL Erlenmeyer flask equipped with a magnetic stir bar. To this solution was drop wise added over 15 minutes a solution consisting of 2.00 g N_α-lauroyl-N_ε-ammonium chloride-L-Lysine, 10 mL RO water, and 2.7 mL 20% aqueous NaOH. Stirring was continued for an additional 30 minutes, after which sufficient concentrated hydrochloric acid was added to lower the pH to 1 as indicated by testing with pHydrion paper. The reaction solution was then transferred into a separatory funnel and the bottom turbid dichloromethane layer was isolated and dried with anhydrous magnesium sulfate and gravity filtered. To the filtrate was added 50 mL of hexanes. The solution was then concentrated via removal of 34 mL of solvent via trap-to-trap distillation and then placed in a -19 °C freezer. A mass of white precipitate formed after a few hours and was isolated via vacuum filtration and then dried in vacuo for 2 hours. 1.130 g of a slightly off white solid powder product was obtained [mp 37.0 - 39.0 °C; IR (cm⁻¹), 3301sb, 2923s, 2852s, 2184m, 2099s, 1721s, 1650s, 1531s, 1456m, 1416w, 1347m, 1216m, 1136w]. Analysis (Midwest Microlab, LLC): Calculated: C, 61.58%; H 9.25%; N,

7.56%; O, 12.95%; S, 8.65%. Actual: C, 61.64%; H, 9.21%; N, 7.58%; O, 13.01%; S, 8.55%.

Step 4: Isolation of Sodium N_α-lauroyl-N_ε-isothiocyanato-L-Lysinate via lyophilization.

5 [0058] 0.147 g of N_α-lauroyl-N_ε-isothiocyanato-L-Lysine was combined and stirred with 2 g of RO water and 0.39 mL of 1.00 M NaOH in a 50 mL single neck round bottom flask and filtered into a 250 mL single neck round bottom flask to yield a clear pale amber solution. The flask was then immersed while rotating into a dry ice/acetone bath to yield a solid coating on the walls of the flask, 10 whereupon the flask was evacuated (0.10 mm Hg) and removed from the ice bath. Evacuation for one hour yielded a dry white solid powder of the water soluble surfactant Sodium N_α-lauroyl-N_ε-isothiocyanato-L-Lysinate. [mp 47 – 55 °C to small droplets of clear colorless viscous liquid; IR (mineral oil mull, cm⁻¹), 3300m amide N-H str; 2188s, 2107s N=C str; 1627s, amide C=O str; 1593s 15 carboxylate C=O str]

EXAMPLE III

Preparation of a Two-Part Formulation for the Treatment of Eczema

[0059] A two-part formulation for topical application to the skin was prepared as follows:

5 [0060] Part I: A 25% by mass mixture of N_α-lauroyl-N_ε-isothiocyanato-L-Lysine in Dow Corning DC344 fluid (a mixture of octamethyl-cyclotetrasiloxane and decamethyl-cyclopentasiloxane) was prepared in a mortar and pestle to produce a paste that was loaded into a 5 ml plastic disposable syringe. A syringe
10 needle was not employed. Rather, the dispensing end of the syringe was capped except for when dispensing without a syringe needle into the palm of a hand occurred.

[0061] Part II: Part II consisted of Cetaphil Moisturizing Lotion to which additional triethanol amine (TEA) was added such that the concentration of the additional triethanol amine was 0.006 g triethanol amine per gram of lotion,
15 raising the pH of the Cetaphil Lotion from 7.74 to 8.77.

[0062] Preferred Instructions for Application of Formulation to the Skin: A 0.2 mL portion of the N_α-lauroyl-N_ε-isothiocyanato-L-Lysine/DC344 mixture is dispensed from the syringe into the palm of a hand (approximately 0.13 g of the mixture). Next, two full squirts of the Cetaphil/TEA lotion is dispensed on top of
20 the N_α-lauroyl-N_ε-isothiocyanato-L-Lysine/DC344 mixture (approximately 2.8 g of the lotion). Next, using the index finger of the other hand, the components are mixed thoroughly for approximately 30 seconds, during which time the water

insoluble N_{α} -lauroyl- N_{ϵ} -isothiocyanato-L-Lysine surfactant-precursor is deprotonated to yield the water-soluble anionic (carboxylate) surfactant and yield a homogenous smooth white lotion (this reduces the pH to 7.4). This mixture is then applied to the afflicted areas by gently rubbing it on as one would apply any moisturizing lotion. Treatment is recommended two to three times per day until the symptoms of the eczema subside.

EXAMPLE IV

Preparation of a One-Part Formulation for the Treatment of Eczema

[0063] A one-part formulation for topical application to the skin was prepared as follows:

[0064] First, 0.00025% (by wt.; 5.0 micromolar) of Sodium N_{α} -lauroyl- N_{ϵ} -isothiocyanate-L-Lysinate, the sodium salt of the material provided in step three of Example II, was mixed with 2% Lauryl PEG-10 Methyl Ether Dimethicone (commercially available from Clear Chemical Corporation, Holland, Michigan) which was QS to achieve 100% with 2,6,10,15,19,23-Hexamethyltetracosane (commercially available from Sigma-Aldrich). It will be understood that the concentration of Sodium N_{α} -lauroyl- N_{ϵ} -isothiocyanate-L-Lysinate may range from approximately 0.000001% to approximately 50%. Non-limiting examples of additional concentrations include 0.0005%, 0.005%, 0.005%, 0.005%, 0.05%, 0.5%, 5% – just to name a few. It will be further understood that the concentration of Lauryl PEG-10 Methyl Ether Dimethicone may range from approximately 0.00001% to approximately 50%.

[0065] Preferred Instructions for Application of the One-Part Formulation to the Skin: A 0.1-1.0 mL portion of the one-part formulation is dispensed from a container into the palm of a hand for subsequent administration to an affected area and/or is dispensed directly onto an affected area by gently rubbing it on as one would apply a moisturizing lotion. Treatment is recommended one to four times per day until the symptoms of the eczema subside.

EXAMPLE V

Preparation of a One-Part Formulation for the Treatment of Eczema

[0066] A one-part oil-based formulation for topical application to the skin was prepared as follows:

[0067] Lyophilized Sodium N_{α} -lauroyl- N_{ϵ} -isothiocyanato-L-Lysinate (0.15 g) is dissolved in 29.85 g of refined jojoba oil while stirring and warming to 50 °C to give a clear colorless solution that is 0.50% by mass Sodium N_{α} -lauroyl- N_{ϵ} -isothiocyanato-L-Lysinate. Next, 0.10 g of this solution was combined with 69.90 g of refined jojoba oil, 20.0 g of heavy mineral oil, and 10.0 g of squalane to yield an oil-based formulation that is 0.00050% by mass Sodium N_{α} -lauroyl- N_{ϵ} -isothiocyanato-L-Lysinate. The oils employed are provided for the purposes of illustration, and are not to be construed as limiting the invention in any way. As such, the oils may be liquid, solid, or gel, and may be synthetic or of natural origin and include but are not limited to waxes, esters, lipids, fats, glycerides, cyclic silicones, linear silicones, crosslinked silicones, alkylsilicones, silicone copolyols,

alkylated silicone copolyols, and/or hydrocarbons, and/or ethoxylated versions of all of these.

[0068] The foregoing description merely explains and illustrates the invention and the invention is not limited thereto except insofar as the appended
5 claims are so limited, as those skilled in the art who have the disclosure before them will be able to make modifications without departing from the scope of the invention.

**WHAT IS CLAIMED AND DESIRED TO BE SECURED
BY LETTERS PATENT OF THE UNITED STATES IS:**

1. A method for treating eczema, comprising the step(s) of:

5 - applying an isothiocyanate functional surfactant to an area affected by eczema, wherein the isothiocyanate functional surfactant comprises at least one isothiocyanate functional group associated with an aliphatic and/or aromatic carbon atom of the isothiocyanate functional surfactant.

10 2. The method for treating eczema according to claim 1, further comprising the step of removing the isothiocyanate functional surfactant from the area affected by eczema.

3. A method for treating eczema, comprising the steps of:

15 - applying an isothiocyanate functional surfactant to an area affected by eczema, wherein the isothiocyanate functional surfactant comprises at least one isothiocyanate functional group associated with an aliphatic and/or aromatic carbon atom of the isothiocyanate functional surfactant;

 - removing the isothiocyanate functional surfactant from the area affected
20 by eczema; and

 - repeating the steps of applying and removing the isothiocyanate functional surfactant to/from the affected area.

4. A method for treating eczema, comprising the steps of:

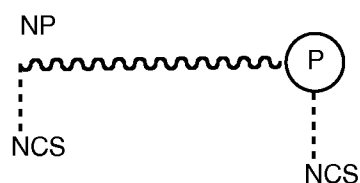
- washing an area affected by eczema with an isothiocyanate functional surfactant, wherein the isothiocyanate functional surfactant comprises at least one isothiocyanate functional group associated with an aliphatic and/or aromatic carbon atom of the isothiocyanate functional surfactant.

5. A method for treating eczema, comprising the step(s) of:

- applying a lysine derivative to an area affected by eczema, wherein the lysine derivative comprises an α -nitrogen and a ϵ -nitrogen, and wherein an alkyl and/or alkanoyl substituent comprising at least approximately 8 carbon atoms is associated with the α -nitrogen, and further wherein at least one isothiocyanate functional group is associated with the ϵ -nitrogen.

6. A method for treating eczema, comprising the step(s) of:

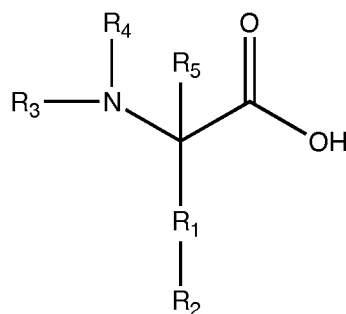
- applying a surfactant to an area affected by eczema, wherein the protonated form of the surfactant is represented by the following chemical structure:



wherein the protonated form of the surfactant comprises a non-polar moiety (NP) and a polar moiety (P), and wherein at least one isothiocyanate functional group (NCS) is associated with the polar and/or non-polar moiety.

5 7. A method for treating eczema, comprising the step(s) of:

- applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein the protonated form of the surfactant is represented by the following chemical structure:

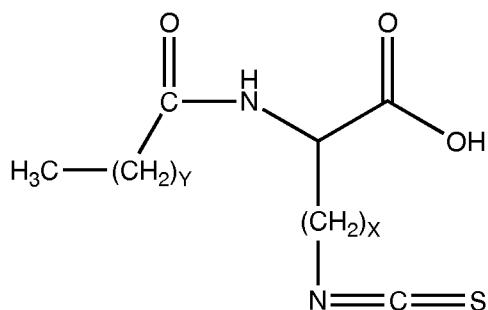


10 wherein R₁ comprises an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols,
 15 esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer; wherein R₂ comprises NCS; and wherein R₃-R₅ are the same or different and comprise H; OH; an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing

approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer with the proviso that at least one of R₃-R₅ comprise an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 8 to approximately 25 carbon atom(s).

8. A method for treating eczema, comprising the step(s) of:

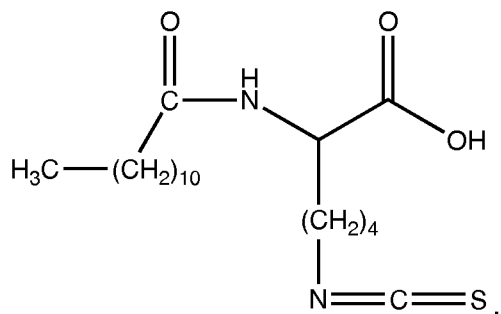
- applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein the protonated form of the surfactant is represented by the following chemical structure:



wherein X comprises an integer ranging from approximately 1 to approximately 25, and wherein Y comprises an integer ranging from approximately 6 to approximately 25.

9. A method for treating eczema, comprising the step(s) of:

- applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein the protonated form of the surfactant is represented by the following chemical structure:

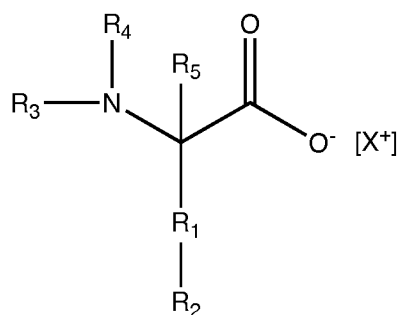


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10. A method for treating eczema, comprising the step(s) of:

- applying a surfactant or a pharmaceutically acceptable salt thereof to an area affected by eczema, wherein the surfactant is represented by the following chemical structure:

10



wherein R₁ comprises an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S

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containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer; wherein R_2 comprises NCS; wherein R_3 - R_5 are the same or different and comprise H; OH; an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer with the proviso that at least one of R_3 - R_5 comprise an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 8 to approximately 25 carbon atom(s), wherein X comprises a counter cation such as, but not limited to, alkali metals, alkaline earth metals, transition metals, s-block metals, d-block metals, p-block metals, NZ_4^+ , wherein Z comprises, H, R_6 , and/or OR_6 , and wherein R_6 comprises an alkyl, cycloalkyl, polycycloalkyl, heterocycloalkyl, aryl, alkaryl, aralkyl, alkoxy, alkanoyl, aroyl, alkenyl, alkynyl and/or cyano group containing approximately 1 to approximately 25 carbon atom(s), wherein the carbon atom(s) may be a linking group to, or part of, a halogen, a N, O, and/or S containing moiety, and/or one or more functional groups comprising alcohols, esters, ammonium salts, phosphonium salts, and

combinations thereof; a linkage to a dimer; a linkage to an oligomer; and/or a linkage to a polymer.

11. The method for treating eczema, according to any one of claims 1 to 10,
5 further comprising the step of applying an additional surfactant, wherein the additional surfactant is selected from at least one of the group comprising a non-ionic surfactant, an anionic surfactant, a cationic surfactant, a zwitterionic surfactant, and combinations thereof.

10 12. The method for treating eczema, according to any one of claims 1 to 11, wherein the eczema is selected from one of the groups comprising atopic eczema, contact eczema, allergic contact eczema, seborrheic eczema, nummular eczema, neurodermatitis, stasis dermatitis, and/or dyshidrotic eczema.

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INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 12/44593

A. CLASSIFICATION OF SUBJECT MATTER

IPC(8) - A01N 47/00; A61K 31/21; A61K 31/26 (2012.01)

USPC - 514/514

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)
USPC- 514/514Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched
USPC- 424/78.02, 78.03, 607; 514/24, 357, 482 (see search terms below)

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

PubWEST, Google Scholar/Patents, Freepatentsonline.com

Search Terms: Isothiocyanate based eczema treatment, Non-corticosteroid based treatment of dermatitis, Method of treating eczema with surfactants, Treating Dermatitis with Surfactants, Treating Dermatitis with lysine isothiocyanate surfactants, Eczema and surfactant.

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	US 7,615,657 B1 (BATHURST et al.) 10 November 2009 (10.11.2009) col 5, ln 55-67; col 6, ln 1-7; col 10, ln 37-46	1-4 and 6
A	US 2009/0005438 A1 (CHENG et al.) 01 January 2009 (01.01.2009) para [0001], [0008]-[0010], [0012], [0023], [0162]-[0169], [0187]	1-4 and 6
A	US 2009/0186853 A1 (YU et al.) 23 July 2009 (23.07.2009) para [0002], [0018]-[0021]	5 and 7-11
A	US 2003/0185864 A1 (KOBAYASHI et al.) 02 October 2003 (02.10.2003) para [0007], [0065]	5 and 7-11
A	US 6,524,594 B1 (SANTORA et al.) 25 February 2003 (25.02.2003) col 1, ln 8-10; col 2, ln 1-5; col 3, ln 4-21; col 6, ln 8-10; col 8, ln 45-49	11
A	US 2004/0037909 A1 (KIM et al.) 26 February 2004 (26.02.2004) para [0027]-[0039], [0093], [0112]	5 and 7-11
A	US 3,969,087 A (SAITO et al.) 13 July 1976 (13.07.1976) col 1, ln 44-57; col 3, ln 25-66; col 13, ln 40-55	5 and 7-11
A	Product Safety Sheet. Allyl Isothiocyanate, Santa Cruz Biotechnology, Inc., 2010, pp 1-14; Section 11, Toxicological Information, pg 11	1-11
X/E	US 2012/0184621 A1 (SILVER) 19 July 2012 (19.07.2012) para [0014]-[0024] http://datasheets.scbl.com/sc-252361.pdf	1-11

☐ Further documents are listed in the continuation of Box C.

* Special categories of cited documents:

"A" document defining the general state of the art which is not considered to be of particular relevance

"E" earlier application or patent but published on or after the international filing date

"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)

"O" document referring to an oral disclosure, use, exhibition or other means

"P" document published prior to the international filing date but later than the priority date claimed

"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art

"&" document member of the same patent family

Date of the actual completion of the international search

23 August 2012 (23.08.2012)

Date of mailing of the international search report

07 SEP 2012

Name and mailing address of the ISA/US

Mail Stop PCT, Attn: ISA/US, Commissioner for Patents
P.O. Box 1450, Alexandria, Virginia 22313-1450

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PCT OSP: 571-272-7774

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 12/44593

Box No. II Observations where certain claims were found unsearchable (Continuation of item 2 of first sheet)

This international search report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☐ Claims Nos.:
because they relate to subject matter not required to be searched by this Authority, namely:

2. ☐ Claims Nos.:
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:

3. ☒ Claims Nos.: 12
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box No. III Observations where unity of invention is lacking (Continuation of item 3 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. ☐ As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying additional fees, this Authority did not invite payment of additional fees.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:

4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee.
- ☐ The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation.
- ☐ No protest accompanied the payment of additional search fees.