

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization

International Bureau



(10) International Publication Number

WO 2014/100851 A1

(43) International Publication Date

3 July 2014 (03.07.2014)

(51) International Patent Classification:
A01N 31/02 (2006.01) *A61K 31/00* (2006.01)
A01N 25/04 (2006.01) *A61L 2/18* (2006.01)

(21) International Application Number:
PCT/AU2013/001489

(22) International Filing Date:
19 December 2013 (19.12.2013)

(25) Filing Language: English

(26) Publication Language: English

(30) Priority Data:
2012905697 24 December 2012 (24.12.2012) AU

(71) Applicant: NOVAPHARM RESEARCH (AUSTRALIA) PTY LTD [AU/AU]; 3-11 Primrose Avenue, Rosebery, New South Wales 2018 (AU).

(72) Inventors: KRITZLER, Steven; 9 Redgum Avenue, Cronulla, New South Wales 2230 (AU). KWON, Hyo Sang; 3-11 Primrose Avenue, Rosebery, New South Wales 2018 (AU).

(74) Agent: SHELSTON IP; Level 21, 60 Margaret Street, Sydney, New South Wales 2000 (AU).

(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IR, IS, JP, KE, KG, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LR, LS, MW, MZ, NA, RW, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MK, MT, NL, NO, PL, PT, RO, RS, SE, SI, SK, SM, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, KM, ML, MR, NE, SN, TD, TG).

Declarations under Rule 4.17:

— *of inventorship (Rule 4.17(iv))*

Published:

— *with international search report (Art. 21(3))*



WO 2014/100851 A1

(54) Title: IMPROVED ANTIMICROBIAL COMPOSITIONS

(57) Abstract: Antiseptic hand rub composition comprising at least 0.2% w/w of isopropyl myristate, free from other derivatives of myristic acid. Phenoxyethanol is preferably present in the composition at a concentration of 1% by wt or less. A glycol may also be present, such as dipropylene glycol if the composition is an ethanol based hand rub gel or propylene glycol if the composition is in the form of an emulsion or dispersion. Additional biocides such as triclosan and chlorhexidine gluconate may be included.

IMPROVED ANTIMICROBIAL COMPOSITION

BACKGROUND

The use of alcohol as an antimicrobial dates to biblical times and earlier. Alcohol-containing 5 antimicrobial compositions have been widely used in hospitals since at least the 1990's. In 1993, Bruch et al (US 5,403,864) stated:-

"Infection control and epidemiology experts have repeatedly emphasized that the single most important element in reducing the spread of infection is hand washing because a common method of transfer among individuals in the health care environment is with the hands. This fact has been painfully 10 demonstrated in the analysis of epidemic spread.

However obvious and simple this may seem, medical care personnel, including physicians and nurses, are reluctant to wash or scrub their hands as frequently as required by their own protocols. It is estimated that the average time of washing between patients is 10 sec or less. The effectiveness of soap-and-water washing is measured in terms of minutes. Most simply do not wash frequently enough...

15 *When a health care worker handles equipment or patients, bacteria which are not a part of the normal skin flora are picked up and adhere loosely to the topmost skin layer, the stratum corneum."*

These statements remain as true in 2012 as when written in 1993. However in the intervening 20 years, new and improved antimicrobial preparations have been developed and the significance of two additional facts has become apparent. Firstly, some microorganisms may reside more deeply in sub 20 corneum strata. Secondly, the major reason for non-compliance with protocols is drying and chapping of the hands and skin irritation caused by repeated use of alcoholic rubs or water based antiseptic washes. Attempts to minimize irritation by inclusion of emollients have not been effective at increasing compliance either (i) because the emollients contributed to a feeling of greasiness after use or (ii) because they reduced the speed with which the hand wash was effective, or (iii) at the concentrations 25 required, the emollients were ineffective at skin irritation reduction or for a combination of these three reasons.

In 1995 Bruch et al disclosed an antimicrobial composition comprising Triclosan, chloroxyphenol and an alcohol but this composition was not effective against subcutaneous organisms and although the inventors claimed that no signs of irritation were exhibited after multiple uses in the laboratory, many 30 cases of skin irritation were exhibited in hospital use.

In 1998 Jampani et al (US 6,022,551) noted a need for an antimicrobial composition that is effective while also being non-irritating to users, and described a composition containing specific thickeners, and phospholipids

The present Inventors have found that subjective feel of the composition also plays an important role in compliance, irrespective of other factors, and it is not sufficient for a composition to be "non-irritating". Thus, staff who may have to apply compositions to their hands as frequently as 100 times a day if they are to fully comply with protocols have been found to have a much higher compliance rate if using 5 preparations which they judge to feel good, than if using preparations which they do not judge to feel good, or which they judge to feel inferior to other preparations they have used which they judge to feel better. One of the factors influencing feel is the tendency to pill exhibited by some alcoholic gel preparations, but other factors include greasiness, and other subjective factors which play a major role in affecting how the composition feels when used and hence compliance rates. . Preparations which are 10 generally judged by staff in use to feel superior to prior art preparations are herein referred to as having "improved feel" One Internationally accepted benchmark for biocidal efficacy is that a specified dose of an antiseptic composition left in contact with the hands for a specified time is required to produce at least the same biocidal efficacy as 6 ml of 60% v/v isopropyl alcohol with 60 secs contact time. (The test method is fully described in European Standard EN 1500:1997, entitled 'Chemical disinfectants and 15 antiseptics Hygienic Handrub – Test method and requirements (phase 2/step2)' against *E. coli* NCTC 10538.herinafter referred to as "EN1500:1977")

Compositions for use as antiseptic hand rubs have contained materials added to "improve skin conditioning" and moisturization e.g. humectants such as glycerine, anti-inflammatories such as isolene, 20 and anchoring agents /conditioners such as phenyldimethicone quaternary compounds. However skin "conditioners" are intended to affect the moisturization, emolliency and condition of the skin, in order to reduce irritation, rather than to affect the feel of the composition on the skin.

Any discussion of the prior art throughout the specification should in no way be considered as an admission that such prior art is widely known or forms part of common general knowledge in the field.

25 **OBJECT OF THE INVENTION**

It is an object of the present invention to overcome or ameliorate at least one of the disadvantages of the prior art, or to provide a useful alternative.

More particularly, it is an object of the present invention to provide an antiseptic handrub or handwash which avoids or ameliorates at least some of the above discussed disadvantages of prior art and which 30 in preferred embodiments produces at least the same biocidal efficacy as is produced by hand rubbing with 6ml of 60% v/v isopropyl alcohol in 60 seconds, but does so with improved skin feel.

Preferred embodiments are suitable for repeated use by health care personnel when moving from patient to patient or procedure to procedure with the same patient and comply with the internationally accepted standard for efficacy.

35 The use of preferred embodiments promotes improved compliance with antiseptic protocols.

BRIEF STATEMENT OF THE INVENTION

According to a first aspect, the invention consists in an antiseptic hand rub composition which when used at a rate of less than 6ml of composition for up to 60 seconds produces a level of biocidal efficacy equal to or greater than that produced by 6ml of 60% v/v aqueous isopropyl alcohol in 60 secs (as measured according to the test method of EN1500:1977), said composition characterised in that it comprises at least 0.2% w/w of isopropyl myristate.

Isopropyl myristate is the isopropyl ester of myristic acid (linear C14 saturated acid), having the following structure:

10



In one preferred embodiment according to the first aspect, handrubbing with only 3ml of an ethanol based antiseptic hand rub gel composition produces in 30 seconds at least the same biocidal efficacy as is produced by hand rubbing with 6ml of 60% v/v isopropyl alcohol in 60 seconds (as measured according to the test method of EN1500:1977). Because preferred embodiments provide the same or greater biocidal efficacy with half the quantity of the preparation and in half the time, this together with the improved feel of compositions according to the invention, greatly enhances compliance with handrubbing protocols. Since ethanol is inferior in biocidal efficacy to isopropanol it is surprising that an ethanol based antiseptic gel is as efficacious.

20 Unless the context clearly requires otherwise, throughout the description and the claims, the words "comprise", "comprising", and the like are to be construed in an inclusive sense as opposed to an exclusive or exhaustive sense; that is to say, in the sense of "including, but not limited to".

In one preferred embodiment of the first aspect of the invention, the antiseptic hand rub composition is an ethanol based antiseptic hand rub composition that may comprise one or more glycols (preferably 25 dipropylene glycol), or phenoxyethanol, or both a glycol and phenoxyethanol.

In one preferred embodiment the antiseptic is an ethanol based hand rub composition in the form of a gel.

In an alternative preferred embodiment of the first aspect of the invention, the antiseptic hand rub composition is in the form of an emulsion or dispersion that may comprise one or more glycols, of which 30 at least one is a low molecular weight glycol (preferably propylene glycol), or phenoxyethanol, or both a glycol and phenoxyethanol.

Preferably the isopropyl myristate acts as an antipilling agent.

For preference, the isopropyl myristate is the only derivative of myristic acid present in the composition, that is, the composition is specifically free from all other esters, salts or derivatives of myristic acid such as ethyl myristate or butyl myristate, or sodium myristate, myristamide, myristyl aldehyde, myristamide DEA etc.

- 5 The isopropyl myristate acts as an anti-pilling agent and improves the feel of the composition on the skin. The present inventors have surprisingly found that inclusion of at least 0.2% w/w isopropyl myristate in alcoholic hand rubs containing glycol and phenoxyethanol (both gels and water based emulsions), significantly improves the feel otherwise produced by the same and similar compositions omitting the isopropyl myristate. Even more surprisingly, the selection of isopropyl myristate (a C14 ester) as an antipilling agent produces benefits including skin feel not obtained with the C12 or C14 isopropyl esters and also not obtained with the C16 or C18 isopropyl esters, or with other myristates.
- 10

- 15 For instance, the same highly desirable feel is not obtained by substituting the C10 isopropyl ester, isopropyl caproate (decanoate) or the C12 isopropyl ester, isopropyl laurate (dodecanoate), nor is it obtained by substituting the C16 ester isopropyl palmitate or the C18 ester isopropyl stearate. Nor by substituting other similar compositions such myristyl myristate for the isopropyl myristate

Preferably the alcoholic antiseptic hand rub composition is substantially free from C10, C12, C16 or C18 isopropyl esters, i.e. free from isopropyl caproate (C10), isopropyl laurate (C12), isopropyl palmitate C16 and isopropyl stearate (C18). By substantially free is meant that the isopropyl myristate is at least 90% pure C14)

- 20 Preferably the isopropyl myristate is present in an amount of 0.1-2.0% w/w. More preferably, it may be present in an amount of 0.2-0.8% w/w, or more preferably still 0.2 - 0.5% w/w or 0.2 – 0.3% w/w.

Preferably if the alcoholic antiseptic hand rub composition is a gel then the isopropyl myristate is present in an amount of around 0.2%.

- 25 In another embodiment the invention provides a method of improving the skin feel of an ethanol based antiseptic hand rub composition which produces a level of biocidal efficacy equal to or greater than that produced by hand rubbing with 6ml of 60% v/v isopropyl alcohol in 60 seconds (as measured in accord with EN1500:1977) applying less than 6ml of such handrub for up to 60 seconds, said method comprising the step of incorporating at least 0.2% w/w of isopropyl myristate in the composition.

- 30 The alcoholic antiseptic hand rub composition may comprise a glycol, or phenoxyethanol or a glycol and up to 1% by wt of phenoxyethanol. In preferred embodiments it may be in the form of a gel or an emulsion or suspension.

Preferably the isopropyl myristate is added as an antipilling agent.

Preferably the isopropyl myristate is selected as the only derivative of myristic acid present in the composition and the method avoids the use of a formulation containing C10, C12, C16 or C18 isopropyl

- 5 -

esters, i.e. the method of improving the skin feel avoids the use of isopropyl caproate (C10), isopropyl laurate (C12), isopropyl palmitate C16 and isopropyl stearate (C18).

In another aspect, the invention provides an antiseptic hand rub composition characterised in that it comprises at least 0.2% w/w of isopropyl myristate. Preferably the composition is free from any of isopropyl caproate, isopropyl laurate, isopropyl palmitate and isopropyl stearate. Preferably, isopropyl myristate is the only derivative of myristic acid present in the composition.

5 Preferably the isopropyl myristate is present in a concentration of from 0.2% to 0.5%

Preferably, the composition comprises phenoxyethanol. The phenoxyethanol is preferably present in a concentration of 1% by wt or less.

10 The composition is preferably in the form of an ethanol based hand rub or in the form of an emulsion or dispersion.

The composition preferably contains one or more glycols.

In one preferred embodiment, the composition is an ethanol based hand rub gel, in which case the glycol is preferably dipropylene glycol. The dipropylene glycol is preferably present in an amount of from 15 0.25 to 4 times the amount by wt of isopropyl myristate. Preferably the dipropylene glycol is present in an amount of up to 1% by wt of the composition.

In one particularly preferred embodiment of the present invention, the invention provides a formulation comprising: ethanol, phenoxyethanol, a glycol and isopropyl myristate; and free from any other myristic acid derivative. The formulation is free from any other myristic acid derivative.

20 The amount of ethanol is preferably 50-80% w/w, more preferably ethanol 60-65 w/w%; the amount of dipropylene glycol is preferably 0.2-0.8% w/w, more preferably 0.4-0.6% w/w; the amount of phenoxyethanol is preferably 0.2-up to 1.0% w/w, more preferably 0.5-0.6% w/w; and the amount of isopropyl myristate is preferably 0.1 to 0.3, more preferably around 0.2% w/w.

25 In an alternative preferred embodiment, the composition is in the form of an emulsion or dispersion in which case at least one of the glycols is a low molecular weight glycol, preferably propylene glycol. The propylene glycol is preferably present in an amount of from 0.25 to 4 times the amount by wt of isopropyl myristate. Preferably the propylene glycol is present in an amount of up to 1% by wt of the composition.

The antiseptic compositions may include one or more additional biocides. Preferred additional biocides are triclosan and chlorhexidine gluconate.

- 6 -

The formulations in examples 1 to 8 were prepared. Each formulation contains from 0.2 to 0.5% w/w of isopropyl myristate. Examples 1 to 5 below exemplify alcohol based handrub gels according to the invention. Examples 6 to 8 below exemplify water based emulsions or dispersions.

5 **Example 1:** (70% v/v Ethanol, CHG 0.5%w/w antiseptic gel):

| | |
|--------------------------------|-----------------|
| Ethanol | 62.00%w/w |
| Chlorhexidine gluconate | 0.50%w/w |
| Hydroxypropyl methyl cellulose | 0.50%w/w |
| Glycerol | 0.50%w/w |
| 10 Quarternium-80 | 0.05%w/w |
| Phenoxyethanol | 1.0%w/w |
| Isopropyl myristate | 0.2%w/w |
| Fragrance | 0.10%w/w |
| Aminomethylpropanol | 0.02%w/w |
| 15 Lactic acid | 0.05%w/w |
| Red No. 33 | q.s as required |
| Deionised water | q.s to 100. |

Example 2: (70%v/v Ethanol, Triclosan 1.0%w/w antiseptic gel)

| | |
|------------------------|-----------------|
| 20 Ethanol | 62.00%w/w |
| Triclosan | 1.00%w/w |
| Carbopol | 0.40%w/w |
| Glycerol | 0.50%w/w |
| Propylene glycol | 0.50%w/w |
| 25 Isopropyl myristate | 0.20%w/w |
| Quarternium-80 | 0.05%w/w |
| Phenoxyethanol | 1.00%w/w |
| Fragrance | q.s as required |
| Aminomethylpropanol | q.s as required |
| 30 FD&C Green no3 | q.s as required |
| Deionised water | q.s to 100. |

Example 3: (70% v/v Ethanol antiseptic gel")

| | |
|---------------------|-----------------|
| Ethanol | 62.00%w/w |
| 35 Carbopol | 0.30%w/w |
| Dipropylene glycol | 0.50%w/w |
| Isopropyl myristate | 0.20%w/w |
| Phenoxyethanol | 0.55%w/w |
| Fragrance | q.s as required |

- 7 -

| | |
|---------------------|-----------------|
| Aminomethylpropanol | q.s as required |
| Dye stuff | q.s as required |

Example 4: (70% v/v Ethanol antiseptic gel")

| | | |
|----|---------------------|-----------------|
| 5 | Ethanol | 62.00%w/w |
| | Carbopol | 0.35%w/w |
| | Glycerol | 0.50%w/w |
| | Dipropylene glycol | 0.50%w/w |
| | Isopropyl myristate | 0.20%w/w |
| 10 | Quarternium 80 | 0.05%w/w |
| | Phenoxyethanol | 0.50%w/w |
| | Fragrance | q.s as required |
| | Aminomethylpropanol | q.s as required |
| | FD&C Blue No. 1 | q.s as required |
| 15 | FD&C Yellow No. 5 | q.s as required |
| | Deionised water | q.s to 100 |

Example 5: (70% v/v Ethanol, CHG 2.0% w/w surgical gel)

| | | |
|----|---------------------------------|-----------------|
| 20 | Ethanol | 62.00%w/w |
| | Chlorhexidine gluconate | 2.00%w/w |
| | Hydroxy propyl methyl cellulose | 0.30%w/w |
| | Glycerol | 0.50%w/w |
| | Phenoxyethanol | 1.00%w/w |
| | Isopropyl myristate | 0.20%w/w |
| 25 | Aminomethyl propanol | q.s as required |
| | Lactic acid | q.s as required |
| | Red No. 33 | q.s as required |
| | Deionised water | q.s to 100. |

30 Examples 6-8 below are water based antiseptic emulsions or dispersions

Example 6: (CHG 1% w/w antiseptic lotion)

| | | |
|----|--------------------------------|----------|
| 35 | Chlorhexidine gluconate | 1.00%w/w |
| | Cetostearyl alcohol | 2.50%w/w |
| | Cetyl palmitate | 0.50%w/w |
| | Ceteareth 20 | 0.60%w/w |
| | POE-40 hydrogenated castor oil | 0.20%w/w |
| | Polyethylen glycol-4000 | 2.70%w/w |
| | Glyceryl stearate | 0.50%w/w |

- 8 -

| | | |
|----|--|-------------|
| | PEG-100 stearate | 0.50%w/w |
| | Sorbitan monostearate | 0.70%w/w |
| | Paraffin oil | 4.00%w/w |
| | Isopropyl myristate | 0.50%w/w |
| 5 | Phenoxyethanol | 1.00%w/w |
| | Ethanol | 5.00%w/w |
| | Propylene glycol | 0.50%w/w |
| | PEG-150/ stearyl alcohol/ SMDI copolymer | 4.00%w/w |
| | Barium sulphate | 2.00%w/w |
| 10 | Quarternium 80 | 0.20%w/w |
| | Fragrance | 0.05%w/w |
| | Deionised water | q.s to 100. |

Example 7 (CHG 2% w/w surgical lotion)

| | | |
|----|--|-------------|
| 15 | Chlorhexidine gluconate | 2.00%w/w |
| | Cetosterayl alcohol | 1.80%w/w |
| | Cetyl palmitate | 0.50%w/w |
| | Ceteareth 20 | 0.60%w/w |
| | POE-40 hydrogenated castor oil | 0.20%w/w |
| 20 | Polyethylen glycol-4000 | 1.80%w/w |
| | Glyceryl stearate | 0.40%w/w |
| | PEG-100 stearate | 0.40%w/w |
| | Sorbitan monostearate | 0.40%w/w |
| | Paraffin oil | 3.0%w/w |
| 25 | Isopropyl myristate | 1.00%w/w |
| | Phenoxyethanol | 1.00%w/w |
| | Ethanol | 10.00%w/w |
| | Propylene glycol | 0.50%w/w |
| | PEG-150/stearyl alcohol/SMDI copolymer | 2.00%w/w |
| 30 | Barium sulphate | 2.00%w/w |
| | Quarternium 80 | 0.20%w/w |
| | Deionised water | q.s to 100. |

Example 8: (Triclosan 0.5% w/w antiseptic lotion)

| | | |
|----|--------------------------------|----------|
| 35 | Triclosan | 0.50%w/w |
| | Cetosterayl alcohol | 2.50%w/w |
| | Cetyl palmitate | 0.50%w/w |
| | Ceteareth 20 | 0.60%w/w |
| | POE-40 hydrogenated castor oil | 0.20%w/w |

- 9 -

| | | |
|----|--|-------------|
| | Polyethylen glycol-4000 | 2.70%w/w |
| | Glyceryl stearate | 0.50%w/w |
| | PEG-100 stearate | 0.50%w/w |
| | Sorbitan monostearate | 0.70%w/w |
| 5 | Paraffin oil | 4.00%w/w |
| | Isopropyl myristate | 0.50%w/w |
| | Phenoxyethanol | 1.00%w/w |
| | Chlorhexidine gluconate | 0.20%w/w |
| | Ethanol | 5.00%w/w |
| 10 | Propylene glycol | 0.50%w/w |
| | Povidone | 0.20%w/w |
| | PEG-150/ stearyl alcohol/ SMDI copolymer | 2.00%w/w |
| | Barium sulphate | 2.00%w/w |
| | Quarternium 80 | 0.20%w/w |
| 15 | Fragrance | 0.05%w/w |
| | Deionised water | q.s to 100. |

METHOD- part 1 Tests A, B, C-Relating to ethanol based antiseptic hand rub gels

20 Comparative tests as to skin feel were conducted as follows:

Various alcoholic gel antiseptic formulations based on Example 3 were prepared with the isopropyl myristate removed from the formulation which was otherwise left intact and various isopropyl esters, myristyl myristate, glycerol laurate – all of which are considered to be emollients. These and a market leading alcoholic gel antiseptic were compared by an experienced panel for their skin feel after repeated 25 usage.

A. The ethanol antiseptic gel formula described in Example 3 above was used to prepare skin feel test formulations as follows:

1. Omitting the isopropyl myristate without replacement.
2. Omitting the isopropyl myristate; replacing the isopropyl myristate in Example 3 with the C10 Isopropyl ester.
3. Omitting the isopropyl myristate; replacing the isopropyl myristate in Example 3 with the C12 Isopropyl ester.
4. Incorporating isopropyl myristate.
5. Omitting the isopropyl myristate; replacing the isopropyl myristate in Example 3 with the C16 Isopropyl ester.
6. Omitting the isopropyl myristate; replacing the isopropyl myristate in Example 3 with the C18 Isopropyl ester.
7. Omitting the isopropyl myristate; replacing the isopropyl myristate in Example 3 with myristyl myristate.

- 10 -

8. Omitting the isopropyl myristate; replacing the isopropyl myristate in Example 3 with glyceryl laurate.
9. Alcoholic hand rub "Angel"® from Johnson and Johnson.

B. Various Levels of isopropyl myristate. The ethanol antiseptic gel formula described in Example 3 above was used to prepare skin feel test formulations as follows:

1. The formulation without isopropyl myristate.
2. The formulation with 0.1%w/w isopropyl myristate.
3. The formulation with 0.2%w/w isopropyl myristate.
4. The formulation with 0.35%w/w isopropyl myristate.
5. The formulation with 0.5%w/w isopropyl myristate.
6. The formulation with 0.8%w/w isopropyl myristate.

C. Various Levels of glycol, in this case dipropylene glycol (DPG) were added to Example 3 which otherwise was unchanged and these assessed for skin feel. The levels of dipropylene glycol(DPG) added to Example 3 were:

1. The formulation with 0.1%w/w Dipropylene glycol
2. The formulation with 0.5%w/w Dipropylene glycol
3. The formulation with 0.8%w/w Dipropylene glycol
4. The formulation with 1.0%w/w Dipropylene glycol
5. The formulation with 1.3%w/w Dipropylene glycol

The hands were first washed with antiseptic handwash following standard hospital handwash procedure and dried with paper towels and then allowed to dry for 5 minutes. 3 ml of the respective test product was then applied to both hands and rubbed until dry. After a further minute skin feel was noted. After 5 minutes from the first application, a second application of the test product was applied to both hands and the test procedure was repeated. This sequence was repeated until 4 further 3ml applications had been made with 5 minute intervals between each. The skin feel and pilling were both noted 1 minute after the hands were rubbed dry after the 5th application.

The tests were conducted with a panel of 10 experienced staff on 3 occasions, each at a different site. The results were as follows: "IPM" refers to isopropyl myristate. "DPG" refers to dipropylene glycol.

The respondents were able to choose from a limited number of descriptive results with a sufficient range of choices to encompass the full spectrum of possible results. The respondents were asked to classify the feel as "light" or "heavy" and to classify the feel as one or more of "dry" or "smooth" or "waxy", "greasy" or "oily". In the results columns the number of respondents providing the predominant skin feel result ("res") were noted.

METHOD-part 2 Tests D, E, F relating to antiseptic handrub emulsions/suspensions

Various aqueous emulsion (lotion) antiseptic formulations were prepared with the isopropyl myristate removed from the formulation which was otherwise left intact and various Isopropyl esters, myristyl myristate, glycerol laurate – all of which are considered to be emollients. These were compared by an experienced panel for their skin feel after repeated usage.

D. The aqueous antiseptic emulsion formula described in Example 8 above was used to prepare skin feel test formulations as follows:

1. Omitting the isopropyl myristate without replacement.
2. Omitting the isopropyl myristate; replacing the Isopropyl myristate in Example 8 with the C10 isopropyl ester.
3. Omitting the isopropyl myristate; replacing the Isopropyl myristate in Example 8 with the C12 isopropyl ester.
4. Incorporating isopropyl myristate.
5. Omitting the isopropyl myristate; replacing the Isopropyl myristate in Example 8 with the C16 Isopropyl ester.
6. Omitting the isopropyl myristate; replacing the Isopropyl myristate in Example 8 with the C18 Isopropyl ester.
7. Omitting the isopropyl myristate; replacing the Isopropyl myristate in Example 8 with myristyl myristate.
8. Omitting the isopropyl myristate; replacing the Isopropyl myristate in Example 8 with glyceryl laurate.

E. Various Levels of Isopropyl Myristate:

The aqueous antiseptic emulsion formula described in Example 8 above was used to prepare skin feel test formulations as follows:

1. The formulation without isopropyl myristate.
2. The formulation with 0.2%w/w isopropyl myristate.
3. The formulation with 0.5%w/w isopropyl myristate.
4. The formulation with 0.8%w/w isopropyl myristate.
5. The formulation with 1.0%w/w isopropyl myristate.
6. The formulation with 1.3%w/w isopropyl myristate.

F. Various Levels of Glycol, in this case propylene glycol (PG), were added to Example 8 which otherwise was unchanged and these assessed for skin feel:

1. The formulation with 0.2%w/w propylene glycol.
2. The formulation with 0.5%w/w propylene glycol.
3. The formulation with 0.8%w/w propylene glycol.

- 12 -

4. The formulation with 1.0%w/w propylene glycol.
5. The formulation with 1.3%w/w propylene glycol.

The hands were first washed with antiseptic handwash following standard hospital handwash procedure and dried with paper towels and then allowed to dry for 5 minutes. 4 ml of the respective test product

5 was then applied to both hands and rubbed until dry. After a further minute skin feel was noted. After 5 minutes from the first application, a second application of the test product was applied to both hands and the test procedure was repeated. This sequence was repeated until 2 further 4ml applications had been made with 5 minute intervals between each. The skin feel was both noted 1 minute after the hands were rubbed dry after the 3rd application.

10 The tests were conducted with a panel of 10 experienced staff on 3 occasions, each at a different site. The results were as follows: "IPM" refers to isopropyl myristate "PG" refers to propylene glycol.

In the results columns the number of respondents providing the predominant skin feel result were noted.

The respondents were able to choose from a limited number of descriptive results with a sufficient range of choices to encompass the full spectrum of possible results.

15 **RESULTS- part 1 Tests A, B, C-Relating to ethanol based antiseptic hand rub gels**

Tests to date have shown that the biocidal efficacy of compositions according to the invention was not adversely affected by inclusion of isopropyl myristate (IPM) in amounts of up to at least 1 % w/w.

20 For example, an alcoholic hand rub according to example 3 of the invention complies with the requirements of the European standard test with a log reduction of 3.29 after 30 seconds rubbing with 3 mL of product compared to the reference product, 6 mL (60%v/v Propan-2-ol) for 60 seconds contact time with a log reduction of 3.33. (a difference in log reduction of no statistical significance despite the difference in quantity and time)

Table -1 Results from Test A, Site 1

25 **Effect of various isopropyl esters on Skin Feel in comparison with market leading product**

| | After 1 st Application | Res. | After 5 th Application | Res. | Pilling | Res. | Rank |
|--------------------------|-----------------------------------|------|-----------------------------------|------|---------------|------|------|
| Example 3 without IPM | Light, Dry | 7/10 | Light, Very Dry | 8/10 | Pilling a lot | 9/10 | 5 |
| Example 3 with IPM (C14) | Light, smooth | 6/10 | Soft, Smooth | 8/10 | No Pilling | 8/10 | 1 |
| Example 3 with C10 | Light, Dry | 8/10 | Light, Very Dry | 7/10 | Pilling a bit | 8/10 | 2 |
| Example 3 with C12 | Light, Dry | 7/10 | Light, Dry | 6/10 | Pilling a bit | 7/10 | 2 |
| Example 3 with C16 | Heavy, Waxy | 8/10 | Heavy, Waxy | 8/10 | No Pilling | 7/10 | 3 |

- 13 -

| | | | | | | | |
|-----------------------------------|---------------------|------|--------------------|------|---------------|------|---|
| Example 3 with C18 | Heavy, Waxy | 7/10 | Heavy, Oily & Waxy | 9/10 | No Pilling | 8/10 | 4 |
| Example 3 with Myristyl Myristate | Heavy, Waxy | 7/10 | Heavy, Waxy | 6/10 | No Pilling | 7/10 | 6 |
| Example 3 with Glyceryl Laurate | Heavy, Greasy | 7/10 | Heavy, Greasy | 8/10 | No Pilling | 9/10 | 6 |
| Example with "Angel"® | Light, smooth, Oily | 8/10 | Greasy, Very Oily | 7/10 | Pilling a lot | 9/10 | 8 |

From the results of above Table 1 it has been concluded that:

1. After repeated cycles of use the composition incorporating isopropyl myristate produced skin feel preferred to that produced by any of the other Isopropyl ester containing compositions tested.
 2. After repeated cycles of use the composition incorporating isopropyl myristate produced the preferred combination of skin feel in combination with minimal pilling than the other compositions tested.
- 5

Table 2 Test B – Site 1

Effect of IPM concentration on Skin Feel

| | After 1 st Application | No. | After 5 th Application | No. | Pilling | No. | Rank |
|--------------------------|-----------------------------------|------|-----------------------------------|------|---------------|------|------|
| Example 3 without IPM | Light, Dry | 7/10 | Light, Very Dry | 8/10 | Pilling a lot | 9/10 | 6 |
| Example 3 with 0.1% IPM | Light, Very Dry | 8/10 | Light, Dry | 8/10 | Pilling a bit | 9/10 | 4 |
| Example 3 with 0.2% IPM | Light, Smooth | 6/10 | Soft, Smooth | 8/10 | No Pilling | 8/10 | 1 |
| Example 3 with 0.35% IPM | Light, Smooth | 9/10 | Soft, Smooth, Oily | 6/10 | No Pilling | 9/10 | 2 |
| Example 3 with 0.5% IPM | Soft, Smooth | 6/10 | Soft, Oily | 8/10 | No Pilling | 8/10 | 2 |
| Example 3 with 0.8% IPM | Soft, Smooth | 8/10 | Heavy, Oily | 9/10 | No Pilling | 9/10 | 4 |

From the results of above Table 2 it has been concluded that:

- 10 Of the various levels of isopropyl myristate added to the composition, after repeated cycles of use, the test subjects preferred the compositions incorporating isopropyl myristate at levels of 0.2%, 0.35% and 0.5%. Of these the level of 0.2% was somewhat preferred over the levels of 0.35% and 0.5%.

- 14 -

Table 3 Test C – Site 1**Effect of Dipropylene glycol concentration on Skin Feel**

| | After 1 st Application | No. | After 5 th Application | No. | Pilling | No. | Rank |
|--------------------------------------|--------------------------------------|------|--------------------------------------|------|---------------|------|------|
| Example 3 with 0.2% DPG and 0.2% IPM | Light, Very Dry | 7/10 | Light, Dry | 8/10 | Pilling a bit | 8/10 | 5 |
| Example 3 with 0.5% DPG and 0.2% IPM | Light, Smooth | 6/10 | Soft, Smooth | 8/10 | No Pilling | 8/10 | 1 |
| Example 3 with 0.8% DPG and 0.2% IPM | Light, Smooth | 6/10 | Soft, Smooth | 8/10 | No Pilling | 8/10 | 1 |
| Example 3 with 1% DPG and 0.2% IPM | Light, Smooth | 7/10 | Soft, Tacky | 7/10 | No Pilling | 9/10 | 3 |
| Example 3 with 1.3% DPG and 0.2% IPM | Light, Tacky | 6/10 | Heavy, Tacky | 8/10 | No Pilling | 9/10 | 4 |

From the results of above Table 3 it has been concluded that:

Of the various levels of dipropylene glycol added to the composition, after repeated cycles of use, the 5 test subjects preferred the compositions incorporating dipropylene glycol at levels of 0.5%, 0.8% and 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

Table 4 Results from Test A, Site 2**Effect of various isopropyl esters on Skin Feel in comparison with market leading product**

| | After 1 st Application | No. | After 5 th Application | No. | Pilling | No. | Rank |
|-----------------------------------|--------------------------------------|------|--------------------------------------|------|---------------|------|------|
| Example 3 without IPM | Light, Dry | 9/10 | Light, Very Dry | 8/10 | Pilling a lot | 6/10 | 6 |
| Example 3 with IPM (C14) | Light, smooth | 8/10 | Soft, Smooth | 9/10 | No Pilling | 8/10 | 1 |
| Example 3 with C10 | Light, Dry | 7/10 | Light, Dry | 7/10 | Pilling a bit | 6/10 | 2 |
| Example 3 with C12 | Light, Dry | 8/10 | Light, Dry | 6/10 | Pilling a bit | 7/10 | 3 |
| Example 3 with C16 | Heavy, Oily | 9/10 | Heavy, Oily | 9/10 | No Pilling | 8/10 | 3 |
| Example 3 with C18 | Heavy, Waxy | 8/10 | Heavy, Oily & Waxy | 8/10 | No Pilling | 7/10 | 5 |
| Example 3 with Myristyl Myristate | Heavy, Waxy | 7/10 | Heavy, Waxy | 7/10 | No Pilling | 8/10 | 7 |

- 15 -

| | | | | | | | |
|---------------------------------|---------------------|------|-------------------|------|---------------|------|---|
| Example 3 with Glyceryl Laurate | Heavy, Greasy | 6/10 | Heavy, Greasy | 8/10 | No Pilling | 8/10 | 8 |
| Example with "Angel"® | Light, smooth, Oily | 9/10 | Greasy, Very Oily | 8/10 | Pilling a lot | 9/10 | 8 |

From the results of above Table 4 it has been concluded that:

1. After repeated cycles of use the composition incorporating isopropyl myristate produced skin feel preferred to that produced by any of the other Isopropyl ester containing compositions tested.
2. After repeated cycles of use the composition incorporating isopropyl myristate produced the preferred combination of skin feel in combination with minimal pilling as compared to the other compositions tested.

Table 5 Test B, Site 2

Effect of IPM concentration on Skin Feel

| | After 1 st Application | No. | After 5 th Application | No. | Pilling | No. | Rank |
|--------------------------|-----------------------------------|------|-----------------------------------|------|---------------|------|------|
| Example 3 without IPM | Light, Dry | 9/10 | Light, Very Dry | 8/10 | Pilling a lot | 6/10 | 6 |
| Example 3 with 0.1% IPM | Light, Very Dry | 8/10 | Light, Dry | 8/10 | Pilling a bit | 8/10 | 4 |
| Example 3 with 0.2% IPM | Light, Smooth | 8/10 | Soft, Smooth | 9/10 | No Pilling | 8/10 | 1 |
| Example 3 with 0.35% IPM | Light, Smooth | 8/10 | Soft, Smooth, Oily | 6/10 | Pilling a bit | 7/10 | 2 |
| Example 3 with 0.5% IPM | Soft, Smooth | 7/10 | Soft, Oily | 8/10 | No Pilling | 6/10 | 2 |
| Example 3 with 0.8% IPM | Soft, Smooth | 7/10 | Heavy, Oily | 7/10 | No Pilling | 7/10 | 4 |

From the results of above Table 5 it has been concluded that:

- 10 Of the various levels of isopropyl myristate added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating isopropyl myristate at levels of 0.2%, 0.35% and 0.5%. Of these the level of 0.2% was somewhat preferred over the levels of 0.35% and 0.5%.

Table 6 Test C, Site 2

Effect of Dipropylene glycol concentration on Skin Feel

- 16 -

| | After 1 st Application | No. | After 5 th Application | No. | Pilling | No. | Rank |
|--------------------------------------|-----------------------------------|------|-----------------------------------|------|---------------|------|------|
| Example 3 with 0.2% DPG and 0.2% IPM | Light, Very Dry | 8/10 | Light, Dry | 8/10 | Pilling a bit | 7/10 | 5 |
| Example 3 with 0.5% DPG and 0.2% IPM | Light, Smooth | 8/10 | Soft, Smooth | 9/10 | No Pilling | 8/10 | 1 |
| Example 3 with 0.8% DPG and 0.2% IPM | Light, Smooth | 6/10 | Soft, Smooth | 7/10 | No Pilling | 9/10 | 2 |
| Example 3 with 1% DPG and 0.2% IPM | Light, Smooth | 8/10 | Soft, Tacky | 7/10 | No Pilling | 7/10 | 2 |
| Example with 1.3% DPG and 0.2% IPM | Light, Tacky | 7/10 | Heavy, Tacky | 8/10 | No Pilling | 9/10 | 4 |

From the results of above Table 6 it has been concluded that:

Of the various levels of dipropylene glycol added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating dipropylene glycol at levels of 0.5%, 0.8% and 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

5 **Table 7 Results from Test A, Site 3**

Effect of various isopropyl esters on Skin Feel in comparison with market leading product

| | After 1 st Application | No. | After 5 th Application | No. | Pilling | No. | Rank |
|-----------------------------------|-----------------------------------|------|-----------------------------------|------|---------------|------|------|
| Example 3 without IPM | Light, Dry | 9/10 | Light, Very Dry | 8/10 | Pilling a lot | 9/10 | 5 |
| Example 3 with IPM (C14) | Light, smooth | 8/10 | Soft, Smooth | 6/10 | No Pilling | 8/10 | 1 |
| Example 3 with C10 | Light, Dry | 7/10 | Light, Very Dry | 6/10 | Pilling a bit | 8/10 | 2 |
| Example 3 with C12 | Light, Dry | 8/10 | Light, Smooth | 6/10 | Pilling a bit | 7/10 | 2 |
| Example 3 with C16 | Light, Smooth | 9/10 | Heavy, Oily | 7/10 | No Pilling | 9/10 | 2 |
| Example 3 with C18 | Heavy, Waxy | 6/10 | Heavy, Oily | 8/10 | No Pilling | 9/10 | 5 |
| Example 3 with Myristyl Myristate | Heavy, Waxy | 8/10 | Heavy, Waxy | 9/10 | No Pilling | 8/10 | 7 |
| Example 3 with Glyceryl Laurate | Heavy, Greasy | 7/10 | Heavy, Greasy | 6/10 | No Pilling | 8/10 | 7 |

| | | | | | | | |
|-----------------------|---------------------|------|-------------------|------|---------------|------|---|
| Example with "Angel"® | Light, smooth, Oily | 9/10 | Greasy, Very Oily | 8/10 | Pilling a lot | 9/10 | 8 |
|-----------------------|---------------------|------|-------------------|------|---------------|------|---|

From the results of above Table 7 it has been concluded that:

1. After repeated cycles of use the composition incorporating isopropyl myristate produced skin feel preferred to that produced by any of the other Isopropyl ester containing compositions tested.
 2. After repeated cycles of use the composition incorporating isopropyl myristate produced the preferred combination of skin feel in combination with minimal pilling as compared to the other compositions tested.
- 5

Table 8 Test B, Site 3

Effect of IPM concentration on Skin Feel

| | After 1 st Application | No. | After 5 th Application | No. | Pilling | No. | Rank |
|--------------------------|-----------------------------------|------|-----------------------------------|------|---------------|------|------|
| Example 3 without IPM | Light, Dry | 9/10 | Light, Very Dry | 8/10 | Pilling a lot | 9/10 | 6 |
| Example 3 with 0.1% IPM | Light, Very Dry | 8/10 | Light, Dry | 8/10 | Pilling a bit | 8/10 | 4 |
| Example 3 with 0.2% IPM | Light, smooth | 8/10 | Soft, Smooth | 6/10 | No Pilling | 8/10 | 1 |
| Example 3 with 0.35% IPM | Light, Smooth | 8/10 | Soft, Smooth, Oily | 6/10 | Pilling a bit | 7/10 | 2 |
| Example 3 with 0.5% IPM | Soft, Smooth | 8/10 | Soft, Oily | 9/10 | No Pilling | 8/10 | 2 |
| Example 3 with 0.8% IPM | Soft, Smooth | 7/10 | Heavy, Oily | 9/10 | No Pilling | 7/10 | 4 |

From the results of above Table 8 it has been concluded that:

- 10
- Of the various levels of isopropyl myristate added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating isopropyl myristate at levels of 0.2%, 0.35% and 0.5%. Of these the level of 0.2% was somewhat preferred over the levels of 0.35% and 0.5%.

Table 9 Test C, Site 3

Effect of Dipropylene glycol concentration on Skin Feel

| | After 1 st | No. | After 5 th | No. | Pilling | No. | Rank |
|--|-----------------------|-----|-----------------------|-----|---------|-----|------|
| | | | | | | | |

| | Application | | Application | | | | |
|--------------------------------------|-----------------|------|--------------|------|---------------|------|---|
| Example 3 with 0.2% DPG and 0.2% IPM | Light, Very Dry | 9/10 | Light, Dry | 8/10 | Pilling a bit | 7/10 | 5 |
| Example 3 with 0.5% DPG and 0.2% IPM | Light, smooth | 8/10 | Soft, Smooth | 6/10 | No Pilling | 8/10 | 1 |
| Example 3 with 0.8% DPG and 0.2% IPM | Light, Smooth | 7/10 | Soft, Smooth | 7/10 | No Pilling | 7/10 | 2 |
| Example 3 with 1% DPG and 0.2% IPM | Light, Smooth | 8/10 | Soft, Tacky | 9/10 | No Pilling | 7/10 | 2 |
| Example 3 with 1.3% DPG and 0.2% IPM | Light, Tacky | 7/10 | Heavy, Tacky | 9/10 | No Pilling | 9/10 | 4 |

From the results of above Table 9 it has been concluded that:

Of the various levels of dipropylene glycol added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating dipropylene glycol at levels of 0.5%, 0.8% and 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

5 Consolidated Conclusions across the Tests A,B,C and Sites 1-3

It is notable that for Tests A, B and C the preferred compositions chosen by the panellists at each of the sites were the same. This consistency of results is indicative of the experience of the panellists and clearly discernible differences in the results.

Test A

10 The majority of panellists across the three sites agreed on the composition with the preferred emollient with respect to skin feel and pilling. Their clear choice was the formulation with isopropyl myristate.

Test B

15 The majority of panellists agreed that the feel of the formula with IPM was significantly better than that of any other tested formulation. This was the case not only after the first application but more so after repeated applications. Similar results were obtained with the other formulations exemplified and at a range of isopropyl myristate concentrations up to 0.5% w/w and the most preferred level of isopropyl myristate was 0.2%.

20 As will be understood by those skilled in the art the antiseptic compositions can be formulated using other components and in other concentrations without departing from the inventive concept herein disclosed of incorporating isopropyl myristate to improve the feel of the product and enhance its propensity to be used.

Test C

As for Test A the majority of panellists agreed on a composition with a preferred level of dipropylene glycol with respect to skin feel and pilling. The results were consistent across the cycles of use. The preferred composition incorporated dipropylene glycol at a level of 0.5%.

5 **RESULTS- part 2 Tests D, E, F Relating to antiseptic hand rub emulsions and dispersions**

The aqueous hand rub (lotion) according to the invention complies with the requirements of the European standard test with a log reduction of 3.77 after 60 seconds rubbing with 4 mL of product compared to the reference product, 6 mL (60%v/v propan-2-ol) for 60 seconds contact time with a log reduction of 4.04. (a difference in log reduction of no statistical significance despite the difference in

10 quantity and time)

Table 10. Results from Test D, Site 1

SKIN FEEL Comparison with various Isopropyl esters

| | After 1 st Application | Res. | After 3 rd Application | Res. | Rank |
|--------------------------------------|--------------------------------------|------|--------------------------------------|------|------|
| Example 8 without IPM | Light, Dry | 7/10 | Light, Dry | 7/10 | 3 |
| Example 8 with IPM (C14) | Light, Smooth | 6/10 | Soft, Smooth | 7/10 | 1 |
| Example 8 with C10 | Light, Dry | 8/10 | Light, Dry | 8/10 | 3 |
| Example 8 with C12 | Light, Dry | 7/10 | Light, Smooth | 8/10 | 2 |
| Example 8 with C16 | Heavy, Smooth | 9/10 | Heavy, Waxy | 8/10 | 5 |
| Example 8 with C18 | Heavy, Waxy | 7/10 | Heavy, Oily & Waxy | 7/10 | 5 |
| Example 8 with Myristyl Myristate | Heavy, Waxy | 7/10 | Heavy, Waxy | 8/10 | 7 |
| Example 8 with Glyceryl Laureate | Heavy, Greasy | 7/10 | Heavy, Greasy | 7/10 | 8 |

From the above Table 10 results it has been concluded that:

After repeated cycles of use the composition incorporating isopropyl myristate produced skin feel

15 preferred to that produced by any of the other isopropyl ester containing compositions tested.

Table 11. Results from Test D, Site 2

SKIN FEEL Comparison with various Isopropyl esters

- 20 -

| | After 1 st Application | No. | After 3 rd Application | No. | Rank |
|--------------------------------------|--------------------------------------|------|--------------------------------------|------|------|
| Example 8 without IPM | Light, Dry | 6/10 | Light, Dry | 8/10 | 4 |
| Example 8 with IPM (C14) | Light, Smooth | 8/10 | Soft, Smooth | 7/10 | 1 |
| Example 8 with C10 | Light, Dry | 7/10 | Light, Dry | 7/10 | 2 |
| Example 8 with C12 | Light, Dry | 7/10 | Light, Smooth | 8/10 | 2 |
| Example 8 with C16 | Heavy, Smooth | 8/10 | Heavy, Waxy | 6/10 | 5 |
| Example 8 with C18 | Heavy, Waxy | 7/10 | Heavy, Oily & Waxy | 8/10 | 6 |
| Example 8 with Myristyl Myristate | Heavy, Waxy | 7/10 | Heavy, Waxy | 9/10 | 7 |
| Example 8 with Glyceryl Laurate | Heavy, Greasy | 7/10 | Heavy, Greasy | 7/10 | 7 |

From the above Table 11 results it has been concluded that:

After repeated cycles of use the composition incorporating isopropyl myristate produced skin feel preferred to that produced by any of the other isopropyl ester containing compositions tested.

Table 12. Results from Test D, Site 3

5 SKIN FEEL Comparison with various Isopropyl esters

| | After 1 st Application | No. | After 3 rd Application | No. | Rank |
|--------------------------|--------------------------------------|------|--------------------------------------|------|------|
| Example 8 without IPM | Light, Dry | 9/10 | Light, Dry | 8/10 | 2 |
| Example 8 with IPM (C14) | Light, Smooth | 8/10 | Soft, Smooth | 8/10 | 1 |
| Example 8 with C10 | Light, Dry | 6/10 | Light, Dry | 7/10 | 2 |
| Example 8 with C12 | Light, Dry | 7/10 | Light, Smooth | 7/10 | 4 |
| Example 8 with C16 | Heavy, Smooth | 8/10 | Heavy, Waxy | 7/10 | 5 |
| Example 8 with C18 | Heavy, Waxy | 7/10 | Heavy, Oily & Waxy | 7/10 | 5 |

- 21 -

| | | | | | |
|-----------------------------------|---------------|------|---------------|------|---|
| Example 8 with Myristyl Myristate | Heavy, Waxy | 7/10 | Heavy, Waxy | 8/10 | 7 |
| Example 8 with Glyceryl Laurate | Heavy, Greasy | 7/10 | Heavy, Greasy | 7/10 | 7 |

From the above Table 12 results it has been concluded that:

After repeated cycles of use the composition incorporating isopropyl myristate produced skin feel preferred to that produced by any of the other isopropyl ester containing compositions tested.

Table 13. Test E – Site 1

5 Effect on feel of varying IPM concentration

| | After 1 st Application | No. | After 3 rd Application | No. | Rank |
|-------------------------|-----------------------------------|------|-----------------------------------|------|------|
| Example 8 without IPM | Light, Dry | 6/10 | Light, Dry | 8/10 | 6 |
| Example 8 with 0.2% IPM | Light, Dry | 7/10 | Light, Dry/Smooth | 8/10 | 2 |
| Example 8 with 0.5% IPM | Light, Smooth | 6/10 | Soft, Smooth | 7/10 | 1 |
| Example 8 with 0.8% IPM | Soft, Smooth | 7/10 | Soft, Smooth, Oily | 6/10 | 2 |
| Example 8 with 1.0% IPM | Soft, Smooth | 6/10 | Heavy, Oily | 8/10 | 2 |
| Example 8 with 1.3% IPM | Heavy, Smooth | 8/10 | Heavy, Oily | 7/10 | 5 |

From the above Table 13 results it has been concluded that:

Of the various levels of isopropyl myristate added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating isopropyl myristate at levels of 0.5%, 0.8% and 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

10 Table 14. Test E, Site 2

Effect on feel of varying IPM concentration

| Effect on feel of varying IPM concentration | After 1 st Application | No. | After 3 rd Application | No. | Rank |
|---|-----------------------------------|------|-----------------------------------|------|------|
| Example 8 without IPM | Light, Dry | 6/10 | Light, Dry | 8/10 | 6 |

- 22 -

| | | | | | |
|-------------------------|---------------|------|--------------------|------|---|
| Example 8 with 0.2% IPM | Light, Dry | 7/10 | Light, Dry/Smooth | 7/10 | 5 |
| Example 8 with 0.5% IPM | Light, Smooth | 6/10 | Soft, Smooth | 7/10 | 1 |
| Example 8 with 0.8% IPM | Soft, Smooth | 9/10 | Soft, Smooth, Oily | 7/10 | 1 |
| Example 8 with 1.0% IPM | Soft, Smooth | 8/10 | Heavy, Oily | 8/10 | 3 |
| Example 8 with 1.3% IPM | Heavy, Smooth | 7/10 | Heavy, Oily | 9/10 | 4 |

From the above Table 14 results it has been concluded that:

Of the various levels of isopropyl myristate added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating isopropyl myristate at levels of 0.5%, 0.8% and 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

5 **Table 15. Test E, Site 3**

Effect on feel of varying IPM concentration

| | After 1 st Application | No. | After 3 rd Application | No. | Rank |
|-------------------------|-----------------------------------|------|-----------------------------------|------|------|
| Example 8 without IPM | Light, Dry | 9/10 | Light, Dry | 8/10 | 5 |
| Example 8 with 0.2% IPM | Light, Dry | 8/10 | Light, Dry/Smooth | 6/10 | 4 |
| Example 8 with 0.5% IPM | Light, Smooth | 8/10 | Soft, Smooth | 8/10 | 1 |
| Example 8 with 0.8% IPM | Soft, Smooth | 7/10 | Soft, Smooth, Oily | 7/10 | 2 |
| Example 8 with 1.0% IPM | Soft, Smooth | 6/10 | Heavy, Oily | 7/10 | 2 |
| Example 8 with 1.3% IPM | Heavy, Smooth | 7/10 | Heavy, Oily | 8/10 | 6 |

From the above Table 15 results it has been concluded that:

Of the various levels of isopropyl myristate added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating isopropyl myristate at levels of 0.5%, 0.8% and 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

10 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

Table 16. Test F – Site 1**Effect on Feel of variation in level of Propylene Glycol**

| | After 1 st Application | No. | After 3 rd Application | No. | Rank |
|-------------------------------------|--------------------------------------|------|--------------------------------------|------|------|
| Example 8 with 0.2% PG and 0.5% IPM | Light, Dry | 7/10 | Light, Dry/Smooth | 7/10 | 4 |
| Example 8 with 0.5% PG and 0.5% IPM | Light, Smooth | 6/10 | Soft, Smooth | 7/10 | 1 |
| Example 8 with 0.8% PG and 0.5% IPM | Soft, Smooth | 6/10 | Soft, Smooth, a bit tacky | 7/10 | 1 |
| Example 8 with 1% PG and 0.5% IPM | Soft, Smooth | 8/10 | Soft, Tacky | 7/10 | 3 |
| Example 8 with 1.3% PG and 0.5% IPM | Soft, Tacky | 6/10 | Heavy, Tacky | 7/10 | 4 |

From the above Table 16 results it has been concluded that:

- Of the various levels of propylene glycol added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating propylene glycol at levels of 0.5%, 0.8% and 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

Table 17. Test F, Site 2**Effect on feel of variation in level of Propylene Glycol**

| | After 1 st Application | No. | After 3 rd Application | No. | Rank |
|-------------------------------------|--------------------------------------|------|--------------------------------------|------|------|
| Example 8 with 0.2% PG and 0.5% IPM | Light, Dry | 7/10 | Light, Dry/Smooth | 8/10 | 5 |
| Example 8 with 0.5% PG and 0.5% IPM | Light, Smooth | 6/10 | Soft, Smooth | 7/10 | 1 |
| Example 8 with 0.8% PG and 0.5% IPM | Soft, Smooth | 6/10 | Soft, Smooth, a bit tacky | 8/10 | 2 |
| Example 8 with 1% PG and 0.5% IPM | Soft, Smooth | 7/10 | Soft, Tacky | 7/10 | 2 |
| Example 8 with 1.3% PG and 0.5% IPM | Soft, Tacky | 6/10 | Heavy, Tacky | 8/10 | 4 |

From the above Table 17 results it has been concluded that:

- 24 -

Of the various levels of propylene glycol added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating propylene glycol at levels of 0.5%, 0.8% and 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

Table 18. Test F, Site 3

5 Effect on feel of variation in level of Propylene Glycol

| | After 1 st Application | No. | After 3 rd Application | No. |
|-------------------------------------|--------------------------------------|------|--------------------------------------|------|
| Example 8 with 0.2% PG and 0.5% IPM | Light, Dry | 7/10 | Light, Dry/Smooth | 7/10 |
| Example 8 with 0.5% PG and 0.5% IPM | Light, Smooth | 8/10 | Soft, Smooth | 8/10 |
| Example 8 with 0.8% PG and 0.5% IPM | Soft, Smooth | 8/10 | Soft, Smooth, a bit tacky | 7/10 |
| Example 8 with 1% PG and 0.5% IPM | Soft, Smooth | 8/10 | Soft, Tacky | 7/10 |
| Example 8 with 1.3% PG and 0.5% IPM | Soft, Tacky | 6/10 | Heavy, Tacky | 7/10 |

From the above Table 18 results it has been concluded that:

Of the various levels of propylene glycol added to the composition, after repeated cycles of use the test subjects preferred the compositions incorporating propylene glycol at levels of 0.5%, 0.8% and 1.0%. Of these the level of 0.5% was somewhat preferred over the levels of 0.8% and 1.0%.

10 Consolidated Conclusions relating to emulsions across all Tests D, E, F and Sites

It is notable that for Tests D, E and F the preferred compositions chosen by the panellists at each of the sites were the same. This consistency of results is indicative of the experience of the panellists and clearly discernible differences in the results.

Test D

15 The majority of panellists across the three sites agreed on the composition with the preferred emollient with respect to skin feel. Their clear choice was the formulation with isopropyl myristate.

Test E

20 The majority of panellists agreed that the feel of the formula with IPM was significantly better than that of any other tested formulation. This was the case not only after the first application but more so after repeated applications. Similar results were obtained with the other formulations exemplified and at a

- 25 -

range of isopropyl myristate concentrations up to 1.0% w/w and the most preferred level of isopropyl myristate was 0.5%.

In Tables 10 to 18 illustrate used Example 8, which contains triclosan as the biocide. Substitution of chlorhexidine gluconate for triclosan unsurprisingly gave substantially equivalent results with respect to

5 skin feel.

As will be understood by those skilled in the art the antiseptic compositions can be formulated using other components and in other concentrations without departing from the inventive concept herein disclosed of incorporating isopropyl myristate to improve the feel of the product and enhance its propensity to be used.

10 **Test F**

As for Test D, the majority of panellists agreed on a composition with a preferred level of propylene glycol with respect to skin feel. The results were consistent across the cycles of use. The preferred composition incorporated propylene glycol at a level of 0.5%.

CLAIMS

1. An antiseptic hand rub composition which when used at a rate of less than 6ml of composition for up to 60 seconds produces a level of biocidal efficacy equal to or greater than that produced by 6ml of 60% v/v aqueous isopropyl alcohol in 60 secs (as measured according to the test method of EN1500:1977), said composition characterised in that it comprises at least 0.2% w/w of isopropyl myristate.
2. An antiseptic hand rub composition according to claim 1 in the form of an ethanol based antiseptic hand rub composition which when used at a rate of less than 6ml of composition for up to 60 seconds produces a level of biocidal efficacy equal to or greater than that produced by 6ml of 60% v/v aqueous isopropyl alcohol in 60 secs (as measured according to the test method of EN1500:1977), said composition characterised in that it comprises at least 0.2% w/w of isopropyl myristate.
3. An ethanol based antiseptic hand rub composition according to claim 2 free from isopropyl caproate, isopropyl laurate, isopropyl palmitate and isopropyl stearate.
4. An ethanol based antiseptic hand rub composition according to any one of claims 2-3 wherein isopropyl myristate is the only derivative of myristic acid present in the composition.
5. An ethanol based antiseptic hand rub composition according to any one of claims 2-4 wherein the isopropyl myristate is present in a concentration of from 0.2% to 0.5%
6. An ethanol based antiseptic hand rub composition according to any one of claims 2-5 comprising a glycol.
7. An ethanol based antiseptic hand rub composition according to claim 6 wherein the glycol is or includes dipropylene glycol in a concentration of from 0.25 to 4 times the concentration by wt of isopropyl myristate.
8. An ethanol based antiseptic hand rub composition according to any one of claims 2-7 comprising phenoxyethanol.
9. An ethanol based antiseptic hand rub composition according to any one of claims 2-8 comprising phenoxyethanol at a concentration of 1% or less by wt.

- 27 -

10. An ethanol based antiseptic hand rub composition according to any one of claims 2-9 comprising a glycol and phenoxyethanol in an amount of up to 1% by wt of the composition.
11. An ethanol based antiseptic hand rub composition according to any one of claims 2-10 comprising dipropylene glycol and phenoxyethanol.
- 5 12. An ethanol based antiseptic hand rub composition according to any one of claims 2-11 in the form of a gel.
- 10 13. An alcoholic antiseptic hand rub composition according to any one of the preceding claims which produces substantially the same or better level of biocidal efficacy by hand rubbing for 30 seconds using 3ml of the composition as is obtained using 6ml of 60% v/v isopropyl alcohol in 60 seconds (as measured according to the test method of EN1500:1977).
14. An alcoholic antiseptic hand rub composition according to any one of the preceding claims wherein the isopropyl myristate acts as an antipilling agent.
- 15 15. An antiseptic hand rub composition according to claim 1 in the form of an emulsion or dispersion based antiseptic hand rub composition which when used at a rate of less than 6ml of composition for up to 60 seconds produces a level of biocidal efficacy equal to or greater than that produced by 6ml of 60% v/v aqueous isopropyl alcohol in 60 secs (as measured according to the test method of EN1500:1977), said composition characterised in that it comprises at least 0.2% w/w of isopropyl myristate.
16. An emulsion or dispersion based antiseptic hand rub composition according to claim 15 free from isopropyl caproate, isopropyl laurate, isopropyl palmitate and isopropyl stearate.
- 20 17. An emulsion or dispersion based antiseptic hand rub composition according to claim 15 or 16 wherein isopropyl myristate is the only derivative of myristic acid present in the composition.
18. An emulsion or dispersion based antiseptic hand rub composition according to any one of claims 15 to 17 wherein the isopropyl myristate is present in a concentration of from 0.2% to 1.0%
- 25 19. An emulsion or dispersion based antiseptic hand rub composition according to any one of claims 15 to 18 comprising a one or more glycols.

- 28 -

20. An emulsion or dispersion based antiseptic hand rub composition according to claim 19 wherein at least one of the glycols is a low molecular weight glycol.

21. An emulsion or dispersion based antiseptic hand rub composition according to claim 20 wherein the low molecular weight glycol is propylene glycol.

5 22. An emulsion or dispersion based antiseptic hand rub composition according to any one of claims 15 to 20 comprising phenoxyethanol.

23. An emulsion or dispersion based antiseptic hand rub composition according to any one of claims 15 to 22 comprising phenoxyethanol at a concentration of 1% by wt or less.

10 24. An emulsion or dispersion based antiseptic hand rub composition according to any one of claims 15 to 23 comprising glycol and phenoxyethanol in an amount of up to 1% by wt of the composition.

25. An emulsion or dispersion based antiseptic hand rub composition according to any one of claims 15 to 23 comprising propyleneglycol and phenoxyethanol.

15 26. A formulation comprising: ethanol, phenoxyethanol, a glycol and isopropyl myristate; and free from any other myristic acid derivative.

27. A formulation according to claim 26 comprising: ethanol, phenoxyethanol, dipropyleneglycol and isopropyl myristate; and free from any other myristic acid derivative.

28. A formulation according to claim 26 or 27 comprising: ethanol 50-70% w/w, dipropylene glycol 0.2-0.8% w/w; phenoxyethanol 0.2-0.8% w/w; and isopropyl myristate 0.1 to 0.3 and free from any other myristic acid derivative.

29. A formulation according to any one of claims 27 to 28 comprising comprising: ethanol 60-65% w/w, dipropylene glycol 0.4-0.6% w/w; phenoxyethanol 0.5-0.6% w/w; and isopropyl myristate 0.1 to 0.3% w/w and free from any other myristic acid derivative.

25

30. A method of improving the skin feel of an antiseptic hand rub composition which produces a level of biocidal efficacy equal to or greater than that produced by hand rubbing with 6ml of 60% v/v

isopropyl alcohol in 60 seconds, said method comprising the step of adding to the hand rub at least 0.2% w/w of isopropyl myristate.

31. A method according to claim 30 wherein the hand rub comprises one or more glycols.

32. A method according to claim 30 or 31 wherein the hand rub comprises phenoxyethanol.

5 33. A method according to any one of claims 30 to 32 wherein the hand rub comprises a glycol and phenoxyethanol.

34. A method according to claim 33 wherein the glycol is or includes dipropylene glycol in a concentration of from 0.25 to 4 times the concentration by wt of isopropyl myristate.

35. A method according to any one of claims 30 to 33 wherein the hand rub is in the form of a gel.

10 36. A method according to any one of claims 30 to 33 wherein the hand rub is in the form of an emulsion.

37. A method according to any one of claims 30 to 35 wherein the isopropyl myristate acts as an antipilling agent.

15 38. A method according to any one of claims 30 to 36 wherein the isopropyl myristate is the only derivative of myristic acid present in the composition.

39. A method according to any one of claims 30 to 37 wherein the isopropyl myristate is free from isopropyl caproate, isopropyl laurate, isopropyl palmitate and isopropyl stearate.

40. A method according to any one of claims 30-39 wherein isopropyl myristate is added in an amount of 0.2-2.0% w/w.

20 41. A method according to claim 40 wherein isopropyl myristate is added in an amount of 0.2-1.0% w/w.

42. A method according to claim 41 wherein isopropyl myristate is added in an amount of 0.2 - 0.5% w/w.

INTERNATIONAL SEARCH REPORT

International application No.

PCT/AU2013/001489

A. CLASSIFICATION OF SUBJECT MATTER

A01N 31/02 (2006.01) A01N 25/04 (2006.01) A61K 31/00 (2006.01) A61L 2/18 (2006.01)

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)

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

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

EPOQUE: "isopropyl myristate", "ethanol", "antiseptic", "antibacterial", "antimicrobial", "biocidal", "gel", "emulsion", "phenoxyethanol", "glycol", "dipropylene glycol", "anti-pilling", like terms, and classification marks.

STN (CAPlus/MedLine): "en1500", "isopropyl myristate", "hand"

GOOGLE PATENTS: "en1500", "isopropyl myristate"

GOOGLE: "isopropyl myristate", "pilling", "emollient", "en 1500 1977"

C. DOCUMENTS CONSIDERED TO BE RELEVANT

| Category* | Citation of document, with indication, where appropriate, of the relevant passages | Relevant to claim No. |
|-----------|--|-----------------------|
| | Documents are listed in the continuation of Box C | |

Further documents are listed in the continuation of Box C See patent family annex

| | | | |
|----------|---|-----|--|
| * "A" | Special categories of cited documents: document defining the general state of the art which is not considered to be of particular relevance | "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 |
| "E" | earlier application or patent but published on or after the international filing date | "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 |
| "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) | "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 |
| "O" | document referring to an oral disclosure, use, exhibition or other means | "&" | document member of the same patent family |
| "P" | document published prior to the international filing date but later than the priority date claimed | | |

Date of the actual completion of the international search
12 March 2014

Date of mailing of the international search report
12 March 2014

Name and mailing address of the ISA/AU

AUSTRALIAN PATENT OFFICE
PO BOX 200, WODEN ACT 2606, AUSTRALIA
Email address: pct@ipaaustralia.gov.au
Facsimile No.: +61 2 6283 7999

Authorised officer

Austin Smith
AUSTRALIAN PATENT OFFICE
(ISO 9001 Quality Certified Service)
Telephone No. +61262832381

| INTERNATIONAL SEARCH REPORT C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT | | International application No. PCT/AU2013/001489 |
|---|--|---|
| Category* | Citation of document, with indication, where appropriate, of the relevant passages | Relevant to claim No. |
| X | WO 2010/147868 A2 (GOJO INDUSTRIES, INC.) 23 December 2010 [0061-0063], [0069], [0076-0077], [0108-0121], [0125] | 1-42 |
| X Y | US 2011/0117048 A1 (KRITZLER) 19 May 2011 [0030-0034], Example 1 as above | 26 1-42 |
| Y | EP 1683416 A1 (AIR LIQUIDE SANTÉ INTERNATIONAL [FR] et al) 26 July 2006 [0004], [0015], [0026], [0029-0038], Example A | 1-42 |
| Y | US 2006/0193745 A1 (ARNDT et al) 31 August 2006 [0042-0047], [0057], [0063-0066], [0071-0073], [0080], [0083] | 1-42 |
| A | GARRUTO et al, "Advice to the Lab Lorn - June 2003 - Free Radical Technology" [retrieved on 28 February 2014]. Retrieved from Internet <URL: http://radicaltechnology.com/articles/advice-lmn2003.pdf > published on 18 October 2012 as per Wayback Machine. | |
| A | FRAISE et al, "Principles and Practice of Disinfection, Preservation & Sterilization," Blackwell Publishing Ltd (2004) 4th edition, ISBN 1-4051-0199-7, pages 545-546. | |
| | | |

| INTERNATIONAL SEARCH REPORT Information on patent family members | | International application No. PCT/AU2013/001489 | |
|---|-------------------------|--|--|
| This Annex lists known patent family members relating to the patent documents cited in the above-mentioned international search report. The Australian Patent Office is in no way liable for these particulars which are merely given for the purpose of information. | | | |
| Patent Document/s Cited in Search Report | | Patent Family Member/s | |
| Publication Number | Publication Date | Publication Number | Publication Date |
| WO 2010/147868 A2 | 23 Dec 2010 | AU 2010260293 A1 BR PI1002135 A2 CA 2707374 A1 CA 2764191 A1 CN 101947194 A CN 102803460 A EP 2272339 A2 EP 2443223 A2 JP 2011148759 A JP 2012530139 A KR 20100134532 A KR 20120047212 A MX 2011013317 A TW 201111495 A TW 201113091 A US 2010317743 A1 US 2012129950 A1 WO 2010147868 A2 | 12 Jan 2012 23 Oct 2012 15 Dec 2010 23 Dec 2010 19 Jan 2011 28 Nov 2012 12 Jan 2011 25 Apr 2012 04 Aug 2011 29 Nov 2012 23 Dec 2010 11 May 2012 30 Jan 2012 01 Apr 2011 16 Apr 2011 16 Dec 2010 24 May 2012 23 Dec 2010 |
| US 2011/0117048 A1 | 19 May 2011 | AU 2006246966 B2 BR PI0612931 A2 CA 2608344 A1 CN 101175404 A CN 101175404 B EP 1887864 A1 JP 2008540581 A JP 2013173758 A KR 20080019610 A MX 2007014372 A TW I417046 B US 2008175811 A1 US 2011117048 A1 WO 2006122345 A1 ZA 200710387 A | 19 Jan 2012 07 Dec 2010 23 Nov 2006 07 May 2008 27 Mar 2013 20 Feb 2008 20 Nov 2008 05 Sep 2013 04 Mar 2008 07 Feb 2008 01 Dec 2013 24 Jul 2008 19 May 2011 23 Nov 2006 31 Dec 2008 |
| EP 1683416 A1 | 26 Jul 2006 | EP 1683416 A1 EP 2314162 A1 | 26 Jul 2006 27 Apr 2011 |

Due to data integration issues this family listing may not include 10 digit Australian applications filed since May 2001.
Form PCT/ISA/210 (Family Annex)(July 2009)

| INTERNATIONAL SEARCH REPORT Information on patent family members | | International application No. PCT/AU2013/001489 | |
|---|------------------|---|------------------|
| This Annex lists known patent family members relating to the patent documents cited in the above-mentioned international search report. The Australian Patent Office is in no way liable for these particulars which are merely given for the purpose of information. | | | |
| Patent Document/s Cited in Search Report | | Patent Family Member/s | |
| Publication Number | Publication Date | Publication Number | Publication Date |
| US 2006/0193745 A1 | 31 Aug 2006 | EP 1685854 A1 | 02 Aug 2006 |
| | | EP 1685854 B1 | 27 Oct 2010 |
| | | US 2006193745 A1 | 31 Aug 2006 |
| | | US 2009252775 A1 | 08 Oct 2009 |
| End of Annex | | | |
| Due to data integration issues this family listing may not include 10 digit Australian applications filed since May 2001. Form PCT/ISA/210 (Family Annex)(July 2009) | | | |