INCI: Polyglyceryl-10 Mono/Dioleate (and) Polyglyceryl-3 Oleate (and) Glycerin (and) Phosphatidylglycerol

December 31, 2013 rev.

DC3897

## **Bio-Active Emulsifier**

Distinctive Emul-Lipid BA is a unique, "bio-mimetic", oil-in-water emulsifier, offering a natural choice for improving product stability and performance while minimizing the potential of bio-incompatibility and irritation. It is derived from plant origin and can be formulated into a wide variety of o/w emulsions. Distinctive Emul-Lipid BA is recommended for use in thin/low-viscosity emulsions where stability may be challenging. In addition, this emulsifier offers unique biological interactions for anti-aging, calming irritated skin, and skin hydration applications enabling the creation of base formulations which re-balance skin's natural regenerative processes, and support delivery of actives ingredients.



Distinctive Emul-Lipid BA contains Phosphatidylglycerol (PG), an important constituent of cell membranes typically found in animal tissue at levels between 1-11% of the total lipid content. Research suggests PG offers a regenerative signaling pathway that prompts skin cells regulate cell proliferation and differentiation. It is this key, bio-identical constituent that helps make Distinctive Emul-Lipid BA highly skin compliant and allows it to replenish naturally occurring components to the skin, rebalancing cellular homeostasis and restoring barrier function to protect against drying and environmental stress.

#### **BENEFITS**

- Hydrating/Moisturizing
- Cosmetic anti-aging benefits
- Unique sensory properties
- Unique "mini-emulsification" properties
- Rebalances cellular homeostasis
- Glycerol chemistry (PEG-free)
- Highly skin compliant
- 100% Plant Origin (Non-GMO)

#### **APPLICATIONS**

- Creams & Lotions
- Cleansers
- Hair care

- Sensitive skin
- Anti-aging
- Color cosmetics

### **TYPICAL PROPERTIES**

Appearance
Color
Odor
Specific Gravity
Loss on Drying

(2 hrs, 2 grams, 105°C)

Liquid
Dark Yellow to Light Brown
Characteristic
0.99 – 1.10
< 3.00





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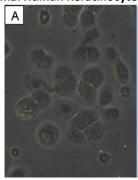
#### IN-VITRO STUDIES: Emul-lipid BA vs. Emul-lipid Control (No PG)

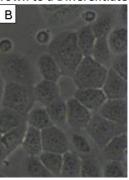
#### Human Gene Expression of Distinctive® Emul-lipid BA

| Human dene Expression of Distinctive Emul-lipid DA |                 |                                |   |  |
|--|-----------------|--------------------------------|---|--|
| TABLE II   | Modulation of w | nt pathway genes by            | 0.05% EMU-BA with phosphalidylalycerol, as compared to EMU-CTR (without phosphalidylalycerol).  |  |
| Position on array                                  | Symbol          | Fold Regulation<br>vs. EMU CTR | Comments  |  |
| C06  | FRAT1           | 3.6                            | Activator of Wnt canonical signaling through inhibition of GSK-3.   |  |
| C10  | FZD2            | 2.0                            | Increased in differentiated tissues (Choi et al., 2008). Accordingly, Frizzled 2 increases the intracellular $Ca^{2+}$ level, consistently with the role of this ion in keratinocyte differentiation (Niu et al., 2012).  |  |
| D03  | FZD8            | 7.3                            | Frizzled 8 decreases with age in progenitor cells. Its upregulation may "rejuvenate" these cells, making them more capable of tissue regeneration (Brunt et al., 2012).   |  |
| D06  | JUN             | 2.4                            | Jun is a target of Wnt canonical pathway. Jun is an early differentiation marker (Blatti & Scott, 1992; Murray et al., 2013) and an effector of TGF-beta – a key effector in skin homeostasis.  |  |
| D07  | KREMEN1         | -2.1                           | Kremen 1 (Krm1) is a negative regulator of the canonical Wnt signaling pathway.   |  |
| E09  | SFRP1           | 2.2                            | SFRP1 Induces differentiation, inhibits proliferation of epithelial cells and negatively regulates Wnt pathway.   |  |
| F10  | WNT10A          | 2.1                            | Induced by TGF-beta. Activator of WNT/β-catenin signaling. WNT10A, in addition to the formation of teeth and hair follicles, is of importance for the formation of nails, regeneration of the epidermis, papillae of the tongue and sweat gland function. Loss of function results in dry skin, abnormal hair patterns and nail malformations (Nawaz et al., 2009). |  |
| G10  | WNT7B           |                                | Wnt7b plays an important role in stem cell homeostasis and in the tissue repair and regeneration (Lin et al., 2010; Kandyba et al., 2013).  |  |

8 out of 84 genes on the Wnt PCR array panel were differentially expressed by Emul-lipid BA. The directionality of the modulation indicates a controlled increase of expression of Wnt genes involved in proliferative/proregenerative progenitor cell homeostasis (FZD8, WNT7b, WNT10a), as well as cell differentiation (FZD2, JUN), consistent with the morphological changes observed microscopically (Fig. 2). This increase may be balanced by the negative regulator SFRP1, itself a powerful pro-differentiation effector. In conclusion, Emul-lipid BA is a bioactive material with progenitor (basal layer stem) cell - normalizing and skin -regenerative benefits, which could result in improved overall skin homeostasis.

### **Epidermal Human Keratinocytes Grown to a Differentiated State**





Epidermal human keratinocytes grown in the presence of (A) Control and (B) Emullipid BA . Note the organized tight junctions between cells grown in the presence of Emul-lipid BA suggestive of a differentiated state, while cells in (A) are more scattered and isolated from each other, possibly geared towards further migration and/or proliferation (original mag. X100).

### **Modulating Hydration Related and Inflammatory Genes**

| TABLE II Gene expression in EMU-BA relative to EMU-CTR |      |       |             |
|--|------|-------|-------------|
| 10 LMO-CTK   | AQP3 | COX1  | COX2 (PGS2) |
| Fold regulation  | 1.68 | -1.07 | -2.0        |

While the constitutively-expressed COX1 was not affected by Emul-lipid BA, the inducible proinflammatory COX2 was inhibited by Emul-lipid BA, while AQP3 was upregulated, as compared to the phosphatidylglycerol-free placebo Control



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### IN-VITRO STUDIES: Emul-lipid BA vs. Polysorbate 80

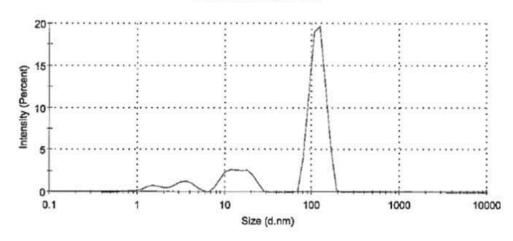
### Collagen I Stimulation & Mitochondrial Metabolism in Human Dermal Fibroblasts

| TABLE II<br>Test Material | Type I Collagen<br>(% Control) | p value | Mitochondrial<br>Metabolism<br>(% Control) | p value |
|---------------------------|--------------------------------|---------|--|---------|
| H2O                       | 100                            | 1       | 100  | 1.000   |
| Emu-BA 0.5% (5mg/ml)      | 60                             | 0.000   | 84   | 0.028   |
| Emu-BA 0.1% (1mg/ml)      | 51                             | 0.000   | 81   | 0.007   |
| Emu-BA 0.02% (200µg/ml)   | 102                            | 0.712   | 107  | 0.202   |
| PS80 0.5% (5mg/ml)        | 3                              | 0.000   | N/A  | 0.000   |
| PS80 0.1% (1mg/ml)        | 3                              | 0.000   | 12   | 0.000   |
| PS80 0.02% (200µg/ml)     | 4                              | 0.000   | 42   | 0.000   |
| MAP                       | 156                            | 0.000   | 111  | 0.069   |

Emul-lipid BA is a non-disruptive emulsifier

## **IN-VITRO STUDIES: Evaluation of Droplet Size in Emulsion**

#### Size Distribution by Intensity



Emul-lipid BA produces stable mini-emulsions



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#### **FORMULATION GUIDELINES**

Recommended use level

2-6% to oil phase Approx. HLB = 9 Emulsifies a wide range of oils. Emulsifies high oil phases >30% Compatible with organic UV filers Formulate between pH 4.5 - 6.5

| THIN SERUM Formula: RON19-44 |                                    |             |   |  |  |  |
|------------------------------|------------------------------------|-------------|---|--|--|--|
| PHASE                        | INGREDIENT                         | % BY WEIGHT | PROCEDURES  |  |  |  |
|                              |                                    |             |   |  |  |  |
| Α                            | Water                              | qs          | <ul> <li>To the main vessel, add water and begin mixing with a propeller</li> </ul> |  |  |  |
| Α                            | Cetyl Hydroxyethylcellulose        | 0.15        | mixer.  |  |  |  |
| Α                            | Glycerin                           | 3.00        | <ul> <li>Add Cetyl Hydroxyethylcellulose and heat to 70°C-75°C.</li> </ul>          |  |  |  |
| Α                            | Xanthan Gum                        | 0.20        | <ul> <li>Premix Glycerin, Butylene Glycol and Xanthan Gum and add to</li> </ul>     |  |  |  |
| Α                            | Butylene Glycol                    | 3.00        | batch.  |  |  |  |
| В                            | Distinctive® Emul-Lipid BA (RON)   | 4.00        | • In a side container, combine Phase B ingredients and heat to 75-                  |  |  |  |
| В                            | Cetyl Alcohol                      | 2.00        | 80°C.   |  |  |  |
| В                            | Dimethicone                        | 2.00        | <ul> <li>Add Phase B to Phase A, mix 15 minutes, until uniform.</li> </ul>          |  |  |  |
| В                            | Butyrospermum Parkii (Shea) Butter | 3.00        | <ul> <li>Begin cooling batch. At 40°C, add Phase C.</li> </ul>                      |  |  |  |
| В                            | Caprylic/Capric Triglyceride       | 3.00        | <ul> <li>Mix until uniform.</li> </ul>  |  |  |  |
| В                            | Vegelight 1214LC (RON)             | 5.00        | <ul> <li>Cool to Room Temperature.</li> </ul>                                       |  |  |  |
| В                            | Dicaprylyl Ether                   | 5.00        |   |  |  |  |
| В                            | Distinctive® Emul-Lipid ST (RON)   | 1.00        | Viscosity: 3600 cps   |  |  |  |
| С                            | Diocide                            | 1.00        |   |  |  |  |
|                              |                                    | 100.00      |   |  |  |  |
|                              |                                    |             |   |  |  |  |

|                   |   | LIPID BA SUNSCREEN  |  |
|-------------------|---|---|--|
| PHASE             | INGREDIENT  | % BY WEIGHT   | PROCEDURES   |
| A<br>A<br>A       | Water<br>Cetyl Hydroxyethylcellulose<br>Glycerin<br>Xanthan Gum   | q.s.<br>0.15<br>3.00<br>0.20  | propeller mixer.  Add Cetyl Hydroxyethylcellulose and heat to 70°C-75°C.   |
| A A B B B B B B C | Disodium EDTA Citric Acid (25% solution)  Distinctive® Emul-Lipid BA (RON) C12-15 Alkyl Benzoate Ethylhexyl Methoxycinnamate Butyl Methoxydibenzoylmethane Homosalate Ethylhexyl Salicylate Vegelight 1214LC (RON) Glyceryl Cocoate Behenyl Behenate Cetyl Alcohol Tocopheryl Acetate | 0.10<br>0.04<br><b>6.00</b><br>4.00<br>7.50<br>2.00<br>10.00<br>3.00<br><b>3.00</b><br>0.70<br>0.50<br>0.50 | <ul> <li>Premix Glycerin, Butylene Glycol and Xanthan Gum and add to batch. Add Disodium EDTA.</li> <li>In a side container, combine Phase B ingredients and heat to 75-80°C.</li> <li>Add Phase B to Phase A, mix 5 - 10 minutes, until uniform. Homogenize batch for 5 minutes at 75°C. Switch to propeller mixer and begin cooling batch.</li> <li>At 40°C, add Phase C. Mix until uniform.</li> <li>At 40°C, add Phase D. Mix until uniform.</li> <li>Cool to Room Temperature.</li> </ul> |
| D                 | Phenoxyethanol (and) Ethylhexylglycerin   | 1.00<br>100.00  | 1  |



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| SQUALANE BARRIER MILK Formula: RON19-63 |                                  |             |       |  |  |  |
|---|----------------------------------|-------------|-------|--|--|--|
| HASE                                    | INGREDIENT                       | % BY WEIGHT | PROCE | DURES  |  |  |
|   |                                  |             |       |  |  |  |
| Α                                       | Water                            | q.s.        | •     | To the main vessel, add water and begin mixing with  |  |  |
| Α                                       | Cetyl Hydroxyethylcellulose      | 0.15        |       | a propeller mixer                                    |  |  |
|   |                                  |             | •     | Add Cetyl Hydroxyethylcellulose and heat to 70-      |  |  |
| В                                       | Glycerin                         | 1.00        |       | 75°C.  |  |  |
| В                                       | Xanthan Gum                      | 0.10        | •     | Premix Phase B ingredients and add to batch at 75°C. |  |  |
| С                                       | Butylene Glycol                  | 5.00        | •     | Add Phase C ingredients and mix until uniform.       |  |  |
| С                                       | Lexgard Natural                  | 1.20        | •     | Add Phase D and mix until uniform.                   |  |  |
|   | -                                |             | •     | In a side container, combine Phase E ingredients and |  |  |
| D                                       | Sodium Benzoate                  | 0.20        |       | heat to 70-75°C.                                     |  |  |
| D                                       | Citric Acid 25% solution         | 0.15        | •     | Add Phase E to main vessel mix 5 - 10 minutes, until |  |  |
|   |                                  |             |       | uniform.   |  |  |
| Е                                       | Distinctive® Emul-Lipid BA (RON) | 6.00        | •     | Homogenize batch for 5 minutes at 70°C.              |  |  |
| Е                                       | C14-C22 Alkane                   | 8.00        | •     | Switch to propeller mixer and begin cooling batch.   |  |  |
| Е                                       | Dicaprylyl Ether                 | 8.00        | •     | Cool to Room Temperature.                            |  |  |
| Е                                       | Squalane                         | 5.00        |       |  |  |  |
| E                                       | Farnesol                         | 3.00        |       |  |  |  |
| E                                       | Distinctive® Emul-Lipid ST (RON) | 0.70        |       |  |  |  |
|   | • • •                            | 100.00      |       |  |  |  |

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