Skin Penetration of Ubiquinone (Co-Q10) In Nanoemulsion Delivery System Using Virgin Coconut Oil (VCO)

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ABSTRACT
Ubiquinone (co-Q10) is a topical anti-aging with low solubility in water, so its penetration into the skin is also low. To increase its penetration ubiquinone (co-Q10) was loaded in the nanoemulsion delivery system using virgin coconut oil (VCO) as oil phase. Ubiquinone (co-Q10) in nanoemulsion formula consists of: ubiquinone (co-Q10), Tween 80, Span 80, ethanol 96%, VCO and acetate buffer solution pH 4.2 ± 0.5 with concentration: 1%; 18.66%; 1.92%; 3.42%; up to 100%, respectively. The characteristics of ubiquinone (co-Q10) nanoemulsion were observed in terms of droplet morphology by transmission electron microscope (TEM) and droplet size by particle analyzer, the pH and viscosity value. The ubiquinone (co-Q10) penetration test in the nanoemulsion system was compared with the emulsion on the rat skin. Result of this research showed the droplet morphology of ubiquinone (co-Q10) in nanoemulsion was spherical. The droplet size was 93.2 ± 2.78 nm. The pH value was 4.12 ± 0.02, and the viscosity was 8.5 cP. Penetration of ubiquinone (co-Q10) in nanoemulsion was deeper (269 ± 48.12 and 993 ± 36.49 μm) than in emulsion (26.33 ± 6.43 and 110.33 ± 9.124 μm), after 2 and 6 hours sample application.

Keywords: Nanoemulsion, Skin Penetration, Ubiquinone (co-Q10), Virgin Coconut Oil (VCO).

INTRODUCTION
Nanoemulsion was known as a system to increase the solubility, and penetration of lipophyllic substance1,2. In this study nanoemulsion delivery system was used as vehicle of ubiquinone (co-Q10) which is known as anti-aging. Ubiquinone (co-Q10) can prevent collagen and elastine damage and help to avoid wrinkle of the skin3. Nanoemulsion is an emulsion of oil in water (O/W) or water in oil (W/O) with the droplet size between 50 - 500 nm. Smaller droplet size results in larger surface area it can increase the penetration of active ingredients4. For O/W nanoemulsion virgin coconut oil (VCO) used as oil phase in nanoemulsion because it can produced smallest droplet size than corn oil and soybean oil5.

MATERIALS AND METHODS
Research Material
Ubiquinone (Kangcare), virgin coconut oil (VCO), Tween 80 (Sigma Aldrich), Span 80 (Sigma Aldrich), ethanol 96% (E.Merck), acetic acid p.a (E.Merck), sodium acetate (E.Merck) p.a.
Research Instrument
Stirrer plate (Dragon Lab MS-Pro), ultrasonic (Branson 3510), Thermo shaker (Wine shake), pH meter (Eutech Instruments pH 700), Viscometer cone and plate, Transmission Electron Microscope (TEM) JEOL JEM 1400, Delsa™ Nano Submicron Particle Size and Zeta Potential Dynamic Light Scattering, Microscope Flourecents and Brightfield Olympus FSX100.

Experimental Animals
Animals used were male Wistar rats with the inclusion criteria: 8-10 weeks of age, 100-250 grams of weight, healthy condition, with no defect or injury.

Ubiquinone (co-Q10) Nanoemulsion and Emulsion Preparation
The nanoemulsion formula in this study refers to the modified formula from Erawati et al, 20146. In this experiment the concentration of ubiquinone (co-Q10), Tween 80, Span 80, ethanol 96%, VCO and acetate buffer solution pH 4.2 ± 0.5 were: 1%; 18.66%; 1.92%; 3.42%; up to 100%, respectively. The emulsion formula consists of: 1% ubiquinone (co-Q10) was dissolved in 35% VCO and then added with 22% of surfactant mixture (Tween 80 and Span 80), it was stirred until the mixture became homogeneous. Subsequently, a solution of acetate buffer pH 4.2 ± 0.5 up to 100%.

RESULTS AND DISCUSSION
The characteristics of ubiquinone (co-Q10) nanoemulsion (QN) and ubiquinone (co-Q10) emulsion (QE) included pH, droplets size and viscosity were presented in Table 1. Ubiquinone (Co-Q10) nanoemulsion and ubiquinone (co-Q10) emulsion both have same pH value were 4.12 ± 0.02. The droplets size of QN (93.2 ± 2.78 nm) was smaller than QE (10.55 ± 0.60 μm). The viscosity of QN (8.5 cP) lower than QE (14.2 cPas). The morphology of QN droplet...
by Transmission Electron Microscope JEOL JEM 1400 was presented in figure 1. Histological preparation of skin penetration of ubiquinone (co-Q10) in emulsion at 2 hours (A1), at 6 hours (A2) after sample application and in nanoemulsion at 2 hours (B1), at 6 hours (B2) after sample application was determined by Microscope Olympus FX-100 42x zoom, presented in Figure 2. The result of in vivo skin penetration tests at two
Table 1: Characteristics of ubiquinone (co-Q10) nanoemulsion (QN) and emulsion (QE).

<table>
<thead>
<tr>
<th>Characteristic</th>
<th>QN</th>
<th>QE</th>
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<tbody>
<tr>
<td>pH</td>
<td>4.12 ± 0.02</td>
<td>4.12 ± 0.02</td>
</tr>
<tr>
<td>Droplet size</td>
<td>93.2 ± 2.78 nm</td>
<td>10.55 ± 0.60 μm</td>
</tr>
<tr>
<td>Viscosity</td>
<td>8.5 cPas</td>
<td>14.2 cPas</td>
</tr>
</tbody>
</table>

was known that penetration depth of ubiquinone (co-Q10) in nanoemulsion were 269 ± 48.12 and 993 ± 36.49 μm, respectively. And than in emulsion were 26.33 ± 6.43 and 110.33 ± 9.124 μm, respectively. It shows that penetration of ubiquinone (co-Q10) in nanoemulsion delivery system into the rat skin faster and deeper then in emulsion. Ubiquinone (co-Q10) penetration rate in nanoemulsion is faster than in the emulsion caused by three factors; the first is the lower nanoemulsion viscosity than the emulsion, the second is the nanoemulsion droplets size smaller than the emulsion, so the movement of the active ingredient is easier in nanoemulsion. The third is the nanoemulsion system which increases the release rate so that it’s much more active ingredients are ready to be penetrated into the skin.6,7

CONCLUSION
Penetration of ubiquinone (co-Q10) in nanoemulsion was deeper (269 ± 48.12 dan 993 ± 36.49 μm) than in emulsion (26.33 ± 6.43 and 110.33 ± 9.124 μm), after 2 and 6 hours sample application on Wistar rats skin.

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