- 1 Capacitive Imaging for Skin Characterizations and Solvent Penetration
- 2 Measurements
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13 Abstract

- 14 Capacitive contact imaging has shown potential in measuring skin properties
- 15 including hydration, micro relief analysis, as well as solvent penetration
- measurements. Through calibration we can also measure the absolute permittivity
- of the skin, and from absolute permittivity we then work out the absolute solvent
- content in skin. In this paper, we present our latest study of capacitive contact
- imaging for skin characterizations and vivo skin solvent penetration. We will show

- 20 how to use capacitive occlusion measurements to characterize skin damages, and
- 21 how to calculate the absolute water content and solvent content in skin.

Keywords

- 23 Capacitive imaging, skin hydration, solvent penetration, water content, solvent
- 24 content.

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1. Introduction

27 The Epsilon permittivity imaging system which is based on Fujistu fingerprint sensors 28 (Fujistu Ltd), has shown potentials in skin hydration imaging, skin characterization 29 analysis, and skin solvent penetration measurements (Levegue et al., 2003; Batisse 30 et al., 2006; Xiao et al., 2007; Bevilacqua et al., 2008; Singh et al., 2008). It is based 31 on capacitance measurement principles, and the measurement results depend on 32 the sample's dielectric permittivity, because of calibration, we can get the absolute 33 dielectric permittivity so that we can work out the absolute water hydration. Our latest 34 studies showed that Epsilon permittivity imaging system can be used in many 35 solvents, due to their high dielectric permittivity. This makes the technique very useful for in-vivo trans-dermal drug delivery studies (Xiao et al., 2012a; Xiao et al., 36 37 2012b).

2. Materials and Methods

40 2.1 Epsilon permittivity imaging system

41 The Epsilon permittivity imaging system, based on Fujistu fingerprint sensors

42 (Fujistu Ltd), has 256x300 pixels with 50µm spatial resolution, as shown in Fig. 1

43 (Xiao et al., 2012b). Each pixel is equivalent of a capacitive sensor, which measures

44 the dielectric constant or permittivity of the sample. Epsilon has an 8-bit grey-scale

45 capacitance resolution per pixel (0 - 255).

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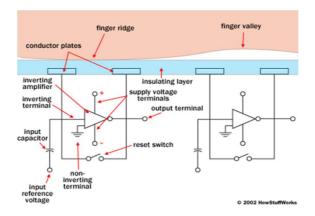
A) Epsilon and the in-vivo stand



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B) Epsilon and the in-vitro stand



52 C) Capacitive fingerprint sensor measurement principle [8]

Fig. 1. Epsilon Permittivity Imaging System: A) Epsilon and the *in-vivo* stand, B)

Epsilon and the *in-vitro* stand, and C) its measurement principle.

2.2 Epsilon calibration

The Epsilon differs from other similar systems in its calibrated, linear response to near-surface dielectric permittivity, see **Fig. 2**. The linear response is important because hydration is linearly related to permittivity. The calibration ensures consistency from instrument to instrument and from time to time. With calibrated Epsilon imaging systems we can measure the absolute dielectric permittivity of the material. A RGB colour scheme is used to represent the dielectric constant values, the brighter the colour, the higher the dielectric constant, and darker the colour the lower the dielectric constant.

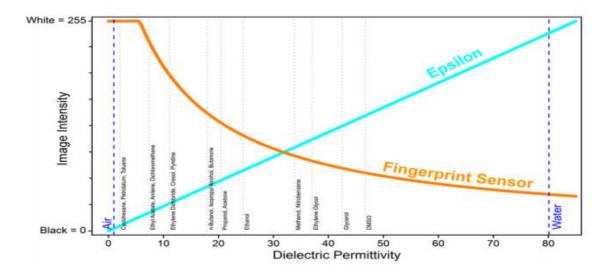


Fig. 2. The Epsilon Calibration curve and linear response to dielectric permittivity

2.3 Skin Hydration and Solvent Concentration Calculations

- 70 If we assume measured skin dielectric constant value (ε_m) is linearly dependent on
- 71 that of dry skin (ε_{dry}) and water (ε_{water}), i.e.

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$$\varepsilon_{\rm m} = \varepsilon_{dry} \times (1 - \frac{H}{100}) + \varepsilon_{water} \times \frac{H}{100}$$
 (1)

- 73 where H is water concentration in skin as percentage of volume ratio, then we can
- work out the water content using following equation [6]

75 .
$$H = \frac{\varepsilon_{\text{m}} - \varepsilon_{dry}}{\varepsilon_{water} - \varepsilon_{dry}} \times 100$$
 (2)

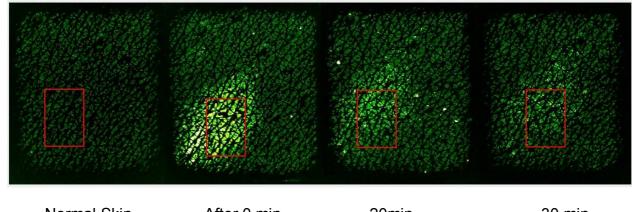
- For the solvent penetration through skin, measured dielectric constant value ($\varepsilon_{\rm m}$) is
- a combination of pure solvent (ε_{sol}) and skin (ε_{skin}), similar to Eq.2, we also can
- calculate the solvent concentration (**C**, volume ratio percentage),

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$$C = \frac{\varepsilon_{\rm m} - \varepsilon_{\rm skin}}{\varepsilon_{\rm sol} - \varepsilon_{\rm skin}} \times 100 \tag{3}$$

80 In the paper, the dielectric constants we used for water is $\boldsymbol{\epsilon}_{water} = 80.1$, for dry skin is $\boldsymbol{\epsilon}_{dry} = 1$, for pure alcohol is $\boldsymbol{\epsilon}_{sol} = 24.3$, for pure ethylene glycol is $\boldsymbol{\epsilon}_{sol} = 55.81$, and for pure glycerol is $\boldsymbol{\epsilon}_{sol} = 38.91$.

2.4 Re-positioning Using Normalized Cross-Correlation Algorithm

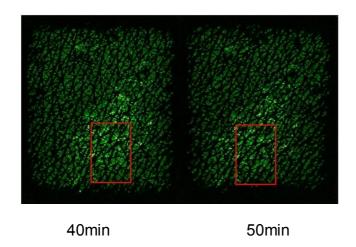
Re-positioning is very important in image processing. In many occasions, we would prefer to compare the results from exactly the same skin site. To select the same skin site manually is cumbersome and inaccurate. We have developed an efficient repositioning technique based on normalized cross-correlation algorithm. As illustrated in **Fig.3.**, users can select the region of interests (RoI) in the first image, the re-positioning technique will automatically find the exactly same position in the subsequent images.



Normal Skin After 0 min 20min 30 min

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96 **Fig.3.** Capacitive contact images of skin before and after Glycerol application. Red 97 squares show the repositioning of the Rol in Normal Skin image (before application) 98 and in the consequent images (after application).

100 If we use matrix **R** to represent the region of interests, and matrix **T** to represent the 101 target images, we first normalized the **R** using following formula,

$$R_N = \frac{R - min}{max - min} \times 255 \tag{4}$$

103 Where **max** and **min** are the maximum value and minimum value of **R**, and R_N is
104 the normalized **R**. Similarly we can also normalize **T** to get T_N . Then we can
105 calculate the cross-correlation of R_N and T_N as,

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$$Corr = \sum_{i} \left(\frac{\sum_{j} (R_N(j) - RM) \times (T_N(j-i) - TM)}{\sqrt{\sum_{j} (R_N(j) - RM)^2} \times \sqrt{\sum_{j} (T_N(j-1) - TM)^2}} \right)$$
 (5)

107 Where **i** and **j** are the matrix indexes, **RM** is the mean of R_N and **TM** is the mean of T_N . From calculation result **Corr**, we can find out at which position that two matrix

are best correlated, and that will be the position of the original region of interests **R** in target image **T**.

2.5 Experimental Procedures

All the measurements were taken under normal ambient laboratory conditions, of 21 ± 1 °C and 45±5 % RH. The volar forearm skin sites from healthy volunteers (aged 20 – 40) were chosen for this study. The volunteers were acclimatized in the laboratory for 20 min prior to the experiments.

For skin characterization, three types of skin damages were studied: intensive washes, tape stripping and sodium lauryl sulphate (SLS) irritation. For intensive washes, room temperature running water and washing-up liquid were used to rub the skin site gently for 3 min with a finger. After wash, the skin site was carefully patted dry with a tissue before the measurements. Tape stripping was performed 20 times per site by the use of standard stripping tape. SLS irritation was achieved by applying 2% SLS water solution (v/v, volume ratio) on skin. Capacitive contact imaging measurements were performed both before and after. The measurements were done by using capacitive imaging sensor to occlude the skin test sites for a period of one minute, during which skin capacitance images were recorded

continuously at the rate of one image per second. The average water contents of the images were then calculated at different times during occlusion using **Eq.2**.

For skin solvent penetration measurements, three solvents were studies, pure ethanol, pure ethylene glycol, and pure glycerol. A small quantity of solvent, held in a plastic well, was applied on skin site for 5 minutes, the skin site was then carefully wiped clean to make sure no solvent residue left on skin surface. The capacitive imaging measurements were performed both before and after the application. Again, the measurements were done by using capacitive imaging sensor to occlude the skin test sites for a period of one minute, during which skin capacitance images were recorded continuously. The solvent concentrations in skin were calculated by using Eq.4, and the re-positioning technique was used to make sure the calculations are done on exactly the same skin site.

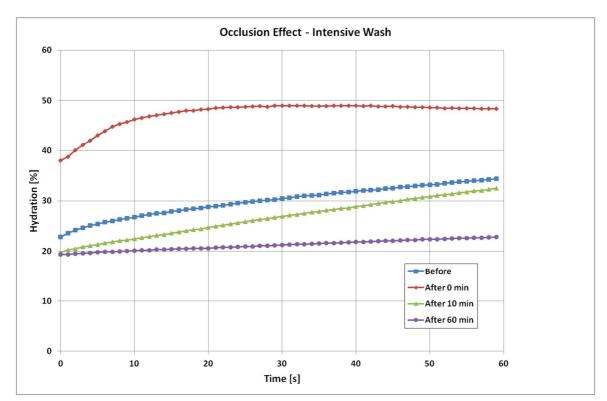
2. Results and Discussions

2.1 Skin Damage Characterizations

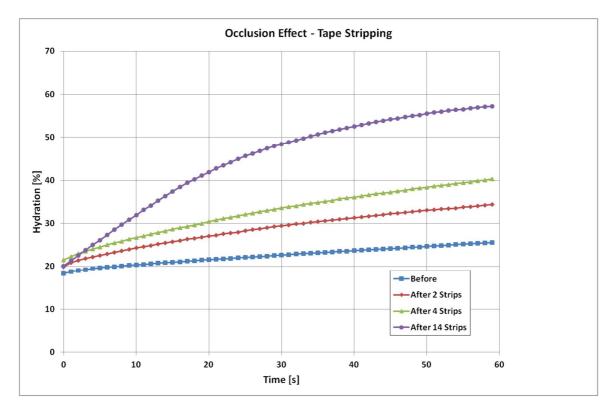
Fig. 4 shows the time dependent skin hydration occlusion curses before and after the intensive wash, tape stripping and SLS irritation. The results show that the levels and the shapes of occlusion curves changed after the skin damage, from the levels

and the shapes we can understand how badly the skin is damaged, and how long does it take to recover. It also is interesting to point out that for different types of skin damages, the time dependent occlusion curves are also different. This suggests that it is possible to use these time dependent occlusion curves to characterize skin, i.e. not only to show how badly the skin is damaged, but also to show what types of damages.

(A)



(B)



(C)

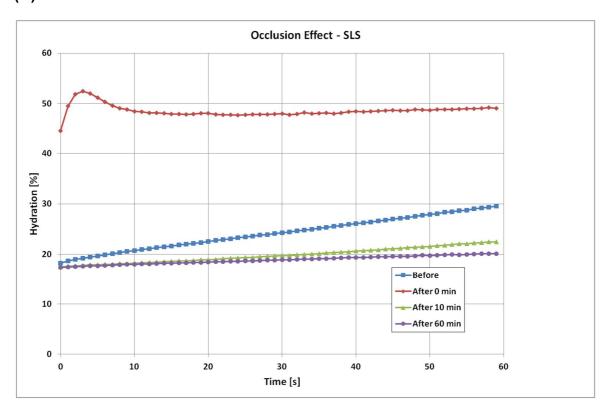
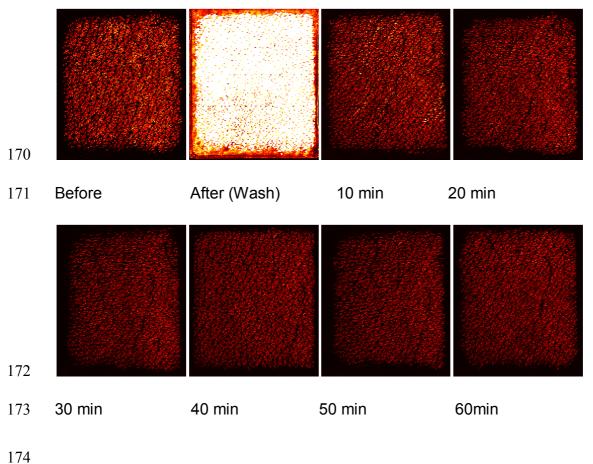


Fig.4. The time dependent skin hydration occlusion curses before and after the intensive wash (A), tape stripping (B) and SLS irritation (C).

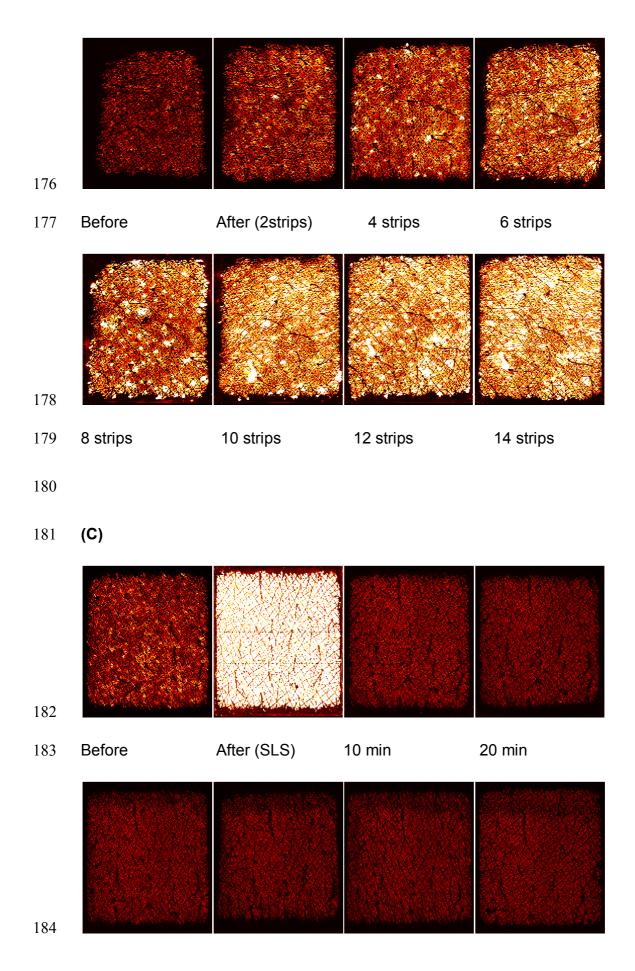
Fig. 5 shows corresponding capacitive contact images before and after intensive washes, tape stripping and SLS irritation measurements. The skin images are generally getting brighter after damage, which indicates higher water content in skin. In both intensive washes and SLS irritation, the darker recovery skin images indicate there is a drying effect after the damage.

(A)



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(B)



185 30 min 40 min 50 min 60min

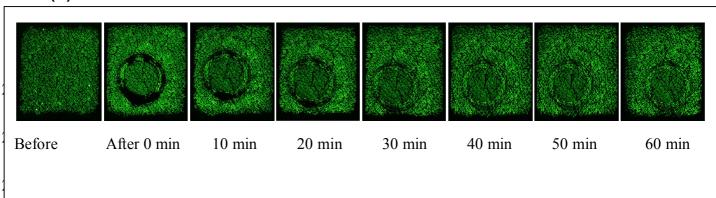
Fig. 5. Typical capacitive contact images of before and after intensive washes (A),

tape stripping (B) and SLS irritation (C).

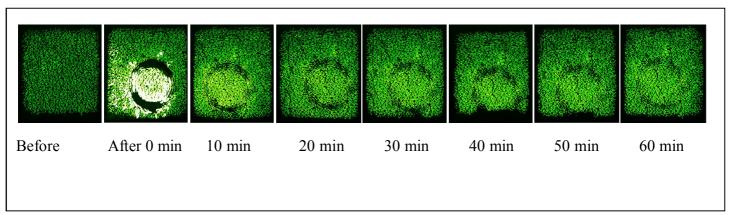
2.2 Skin Solvent Penetrations

Fig. 6 shows the capacitive images before and after the three solvent applications. The image results show that after 5 minutes application, Ethylene Glycol penetrates most into the skin, while Ethanol penetrates the least. This is understandable, as Ethanol is very volatile, and evaporated quickly after the application. Ethylene Glycol also disappears quicker from the skin surface. As Ethylene Glycol is non-volatile, we assume it goes into deeper skin. Glycerol, however, tends to stuck within skin for a long period of time.

(A) Ethanol

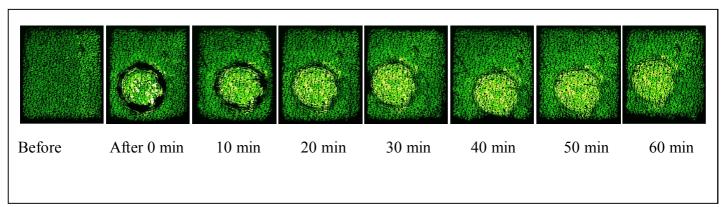


205 (B) Ethylene Glycol



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211 (C) Glycerol



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- Fig. 6. Capacitive contact images of skin before and after solvent applications, (A)
- 218 Ethanol, (B) Ethylene Glycol and (C) Glycerol.

- 220 Fig. 6 shows the corresponding calculated solvent concentration as volume ratio
- 221 percentage before after solvent applications using Eq.4. The calculated results

agree well with the image results. The slight negative undershoots in Ethanol results
 are likely due to the skin drying because of Ethanol application.

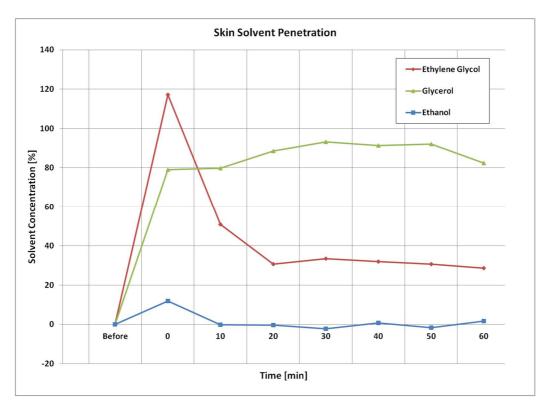


Fig. 6. The calculated solvent concentration as volume ratio percentage before after solvent applications, (A) Ethanol, (B) Ethylene Glycol and (C) Glycerol.

The table below shows the dielectric permittivity of alcohol, ethylene glycol, glycerol, and the concentration of these 3 solvents in the skin from base site to 60 minutes (base site to 66, every 10 minutes).

Figure 6 shows a 5min Glycerol application measurement. Similarly, we can use $C=(\epsilon-\epsilon skin)/(\epsilon Glycerol-\epsilon skin)$ to work out the Glycerol concentration. By using re-positioning technique (Rol + cross correlation), we can select exactly the same skin region we want to analyse in each image, see following red squares. The results show that Glycerol in skin has reached to ~38% (v/v) after just 5min Glycerol application. Then as Glycerol gradually diffuse into to skin, the concentration reduced to 16% (20 min), 9% (30min), 10% (40min) and 6% (50 min).



Glycerol

4 Conclusions

The result shows that, because of volatilizing, the permittivity of alcohol decreased, after t0 (t1-t6), and the capacitive images become darker than base site, the permittivity of both ethylene glycol and glycerol much more increased than base site and diffusion under the surface, the capacitive images of these 2 solvents become brighter than base site, but we can notice the different between these 2 solvents, ethylene glycol is absorbed faster than glycerol.

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