# Moisturizing Kinetics of Maxrich, a Novel Moisturizing Formula Comprising of Xylitol and Xylitol-based Sugars such as Xylityglucoside and Anhydroxylitol

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#### **Abstract**

**Introduction:** Polyols such as glycerol and xylitol are commonly implicated as moisturizers of the skin and other epithelial tissues. Xylitol was found to augment collagen formation, markedly suppress sodium lauryl sulfate-induced transepidermal water loss (TEWL), exert anti-irritant actions, and increase filaggrin expression and skin hydration.

**Materials and Methods:** This was open-label, comparative, single-arm, pilot study involving 15 healthy volunteers who were prone to develop dry skin. Two test preparations were used in this study, one preparation was with ordinary cream base (control), while the other preparation was novel moisturizing formulation (MaxRich®) containing a complex of xylitol, xylitylglucoside, and anhydroxylitol. Primary objective of the study was to evaluate the hydration of skin, after application, of Maxrich® versus control over the duration of 24 h, while the secondary objective was to evaluate TEWL over the duration of 24 hours. MoistureMeter-SC (MMSC) and VapoMeter were used to measure skin hydration and TEWL, respectively.

**Results:** MaxRich® was associated with a significant increase in skin hydration from baseline after 4 hours as shown by increase in mean MMSC reading (22.86 vs. 9.36; P < 0.05). The skin hydration was maintained above baseline throughout 24 hours which was statistically significant (P < 0.05) as shown by higher mean MMSC reading at 8, 12, and 24 h. Compared to control, MaxRich® was associated with a significant increase in skin hydration (10.12 vs. 22.86, P < 0.05). Similarly, MaxRich® was associated with a significant decrease in transepidermal water loss compared to baseline after 4 h as shown by decrease in mean VapoMeter readings (9.22 vs. 5.99, P < 0.05). TEWL was maintained below baseline value as shown by lower mean VapoMeter readings at 8, 12, and 24 h. Compared to control, MaxRich® was associated with a significant decrease in TEWL (6.93 vs. 5.99, P < 0.05).

**Conclusion:** Results of our study indicate that MaxRich®, a novel moisturizer containing xylitol, xylitylglucoside, and anhydroxylitol, is associated with intense, fast, and long-lasting moisturizing effect.

Key words: MaxRich, Skin hydration, Transepidermal water loss, Xylitol

# INTRODUCTION

It is well known that water content of the stratum corneum is very important factor in the appearance and

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Month of Submission: 10-2018 Month of Peer Review: 11-2018 Month of Acceptance: 12-2018 Month of Publishing: 12-2018 function of the skin.<sup>[1]</sup> Natural moisturizing factor (NMF) along with intercellular lipids is responsible for the water holding capacity and barrier function, respectively, thus is fundamental element in maintaining skin hydration.<sup>[1,2]</sup> Impairment in epidermal barrier, decrease in synthesis of intercellular lipids, and NMF with resultant increase in transepidermal water loss lead to decrease in water content of stratum corneum, leading to dry skin.<sup>[1-4]</sup>

Dry skin or xeroderma is a very frequent condition which occurs at any age. The incidence of dryness related dermatoses is on the rise due to urbanization, adulteration,

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pollution, increased life span, and abandonment of traditional oil massages and baths.<sup>[5]</sup>

Moisturizing products constitute one of the largest and most important skin care product categories.<sup>[5]</sup> The function of moisturizers is to maintain the stratum corneum hydrated. There is plethora of moisturizers available in the market with different constituents and different claims. Despite their widespread use and applications, the scientific literature on moisturizers lacks strong evidence.<sup>[5-7]</sup> Early studies conducted on individuals with both healthy and diseased skin showed that some moisturizers tend to weaken the skin barrier function, whereas others may strengthen it, and these discrepant results were assumed to be caused by the varied compositions of moisturizers.<sup>[8]</sup>

Polyols such as glycerol and xylitol are commonly implicated as moisturizers of the skin and other epithelial tissues. [9] While the moisturizing effects of glycerol are very well documented, very limited data are available on the cutaneous effects of a closely related polyol, namely xylitol and xylitol-based sugars. [9] Despite their similar chemical structures, these polyols induce different gene expression changes in the keratinocytes. [10] In few *in vitro* and *in vivo* studies, xylitol was found to augment collagen formation, markedly suppress sodium lauryl sulfate-induced transepidermal water loss (TEWL), exert anti-irritant actions, and increase filaggrin expression and skin hydration. [9-14]

We conducted this pilot study with the purpose to evaluate *in vivo* the stratum corneum hydration and TEWL by the electrical measurements of skin, after treatment with novel moisturizing formula, MaxRich® comprising xylitol and xylitol-based sugars such as xylitylglucoside and anhydroxylitol.

# **MATERIALS AND METHODS**

Primary objective of the study was to evaluate the hydration of skin, after application, of Maxrich® versus control over the duration of 24 h, while the secondary objective was to evaluate TEWL over the duration of 24 h. MoistureMeter-SC (MMSC) and VapoMeter were used to measure skin hydration and TEWL, respectively.

Two test preparations were used in this study, one preparation was with ordinary cream base (control), while the other preparation was novel moisturizing formulation (MaxRich®) containing a complex of xylitol, xylitylglucoside, and anhydroxylitol. The study was conducted according to the ICH-GCP guidelines.

This study involved 15 healthy human volunteers who were prone to dry skin: Seven women and eight men aged 20–30 years. Inclusion criteria were low hydration values (≤10) on the volar forearms, measured with MoistureMeter-SC. Exclusion criteria were - major skin diseases, pregnancy or breastfeeding, systemic corticosteroid or cytostatic therapy within 30 days, any use of local drugs within 30 days that might influence the skin texture, and any skin condition on that could interfere with a clear-cut assessment of the skin. The volunteers were tested for any allergy to tested products, and 3 days before the study, they were advised not to use any moisturizer.

The volar forearms were selected as the test site for conducting this study. Forearms of each volunteer were cleaned, and the volunteers were acclimatized under controlled conditions for the duration of 1 h at the beginning of the study. The humidity was maintained between 40% and 60% and temperature at 20–22°C for the entire study duration. Two 3 cm² × 3 cm² areas were marked on volar forearm of each participant. Area 1 received control product while Maxrich® was applied on area 2. Approximately 30 mg of study product was applied on the test sites by massaging for 30 s using a fingerstall. Two treatments were applied on test sites for 4 times over the total duration of 24 h at 4, 8, 12, and 24 h.

The MoistureMeter SC measures the hydration of stratum corneum indirectly by measuring the electrical capacitance. The skin is electrically a layered structure. The electrical properties of these layers are related to their water content. The probe, the skin surface, and the deeper skin layers form a structure, similar to an electrical capacitor. The measured capacitance is proportional to the water content of the surface layer of the skin. The higher the reading, the higher the moisture content.

VapoMeter measures the transepidermal water loss indirectly by measuring the relative humidity (RH) and temperature. The VapoMeter is equipped with a closed cylindrical chamber that contains sensors for RH and temperature. There is a linear increase of RH% in the chamber shortly after placing the device in contact with the skin. The TEWL is calculated from the increase in RH%. Values of ambient RH (%) and temperature (°C) are recorded before skin contact. The chamber is passively ventilated between measurements. Reduction in the VapoMeter readings indicates improvement in the skin barrier properties.

Skin hydration and TEWL measurements were carried at baseline and were repeated at 4 h, 8 h, 12 h, and 24 h after product application. Short-term testing, in which the effects are usually determined within a few hours after the initial

application of a defined dose of Maxrich® is extremely useful for the evaluation of the efficacy of product. The interpretation of the results was done by unpaired t-test using Microsoft Excel version 2016. P < 0.05 was considered as significant statistically difference.

# **RESULTS**

# Skin Hydration Measurements Using MoistureMeter SC

The mean MoistureMeter-SC readings are shown in Table 1. At baseline, there was no statistical difference between MMSC readings for MaxRich® and control product.

After application of MaxRich®, there was significant increase in skin hydration from baseline after 4 h as shown by increase in mean MMSC reading (22.86 vs. 9.36; P < 0.05). The skin hydration was maintained above baseline throughout 24 h which was statistically significant (P < 0.05) as shown by higher mean MMSC reading at 8, 12, and 24 h. When compared to control product, there was statistically significant increase in skin hydration with MaxRich® at 4 h as shown by significantly higher mean MMSC readings (10.12 vs. 22.86, P < 0.05). Similar trend was seen at 8 h, 12 h, and 24 h [Figure 1].

#### **TEWL Measurements Using VapoMeter**

The mean VapoMeter readings are shown in Table 2. At baseline, there was no statistical difference between VapoMeter for MaxRich® and control product.

Similar to skin hydration, there was a significant decrease in TEWL compared to baseline seen 4 hours after application of MaxRich® as shown by decrease in mean VapoMeter readings (9.22 vs. 5.99, P < 0.05). TEWL was maintained below baseline value as shown by lower mean VapoMeter readings at 8, 12, and 24 h. When compared to control product, MaxRich® was associated with statistically significant decrease in transepidermal water loss after 4 h

Table 1: Mean of MMSC readings at different time points

	0 h	4 h	8 h	12 h	24 h
Control	8.96	10.12	10.37	9.83	9.17
Maxrich®	9.36	22.86	20.15	18.96	12.13

MMSC: MoistureMeter-SC

Table 2: Mean of VapoMeter readings at different time points

	0 h	4 h	8 h	12 h	24 h
Control	9.75	7.43	7.21	6.94	7.11
Maxrich®	9.22	5.99	6.44	6.32	7.47

as shown by lower mean VapoMeter readings (6.93 vs. 5.99, P < 0.05). Similar trend of lower mean VapoMeter readings was seen with MaxRich® compared to control at 8, 12, and 24 h; however, the difference was not statistically significant [Figure 2].

# **DISCUSSION**

Moisturizing products constitute one of the largest and most important skin care product categories. <sup>[5]</sup> The function of moisturizers is to maintain the stratum corneum hydrated. Despite their widespread use and applications, the scientific literature on moisturizers lacks strong evidence. <sup>[5-7]</sup> Various studies with moisturizers showed that not every moisturizer is beneficial to skin and some moisturizers may weaken the skin barrier function which is assumed to be caused by varied composition of moisturizers. <sup>[15-18]</sup>

Polyols like glycerol are most commonly used ingredient in moisturizing products. A structurally similar polyol, xylitol, is also used as humectant and moisturizer. In few *in vitro* and *in vivo* studies, xylitol was found to augment collagen formation, markedly suppress sodium lauryl sulfate-induced TEWL, exert anti-irritant actions, and increase filaggrin expression and skin hydration.<sup>[9-14]</sup>

Szél et al. in their experiment have shown the skin hydrating and anti-irritant effects of xylitol in murine model of

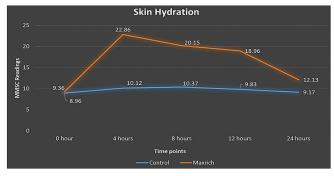


Figure 1: Mean of MoistureMeter-SC readings at different time points

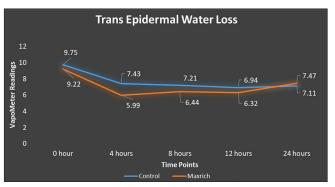


Figure 2: Mean of VapoMeter readings at different time points

irritant contact dermatitis.<sup>[19]</sup> Katsuyama *et al.* in their study showed the beneficial effect of xylitol-containing cream on atopic dry skin.<sup>[20]</sup> Erős *et al.*, their study revealed that the combined application of glycerol and xylitol significantly elevated the hydration of the SC in healthy probands within 2 h.<sup>[21]</sup> Leite e Silva *et al.* in their clinical trial concluded that moisturizer containing xylitol, xylitylglucoside, and anhydroxylitol produced most intense hydration level of the skin surface compared with the gel containing herbal extract and gel containing NMF components.<sup>[1]</sup> Results of these studies clearly demonstrate the skin hydrating benefits of xylitol.

One of the possible explanations of this skin hydrating effect of xylitol is water-binding capacity originating in their chemical structure. However, this is not the only possible explanation as xylitol and xylitol-based sugars reported to exert profound effects on the expression of certain genes and signaling pathways in keratinocytes. Páyer *et al.* in their experiment reported that both glycerol and xylitol upregulated the expression of proteins associated with barrier repair and skin hydration such as filaggrin, loricrin, involucrin, and occludin, but xylitol exerted somewhat more profound effects. [9] Authors concluded that the cutaneous barrier-repairing, differentiation-promoting, and anti-inflammatory effects of xylitol might be superior to those of glycerol. [9]

Szabó-Papp *et al.* in their study reported that xylitol was associated with a marked upregulation of filaggrin, a key molecule of epithelial regeneration and barrier repair. [10] Authors concluded that "since filaggrin levels were shown to be suppressed in dry skin, our intriguing data suggest that xylitol treatment may induce accelerated regeneration and premature differentiation of the keratinocytes resulting in a faster barrier recovery."[10]

Zappelli *et al.* in their review article commented that xylitylglucoside, a sugar derivative of two plant sugars, xylitol and glucose, stimulated the synthesis of proteins, enzymes (different types of keratin, loricrin, and transglutaminase), and ceramides essential to the function of the skin barrier. Authors also commented that these xylitylglucosides are associated with the increased synthesis of dermal macromolecules such as hyaluronic acid and chondroitin sulfate in fibroblast culture. [22] Korponyai *et al.* also reported similar results in their long-term clinical trial with xylitol, authors reported that xylitol leads to an elevated protein expression of filaggrin, a source of NMF.[14]

The results of the above studies clearly demonstrate that besides skin hydration xylitol effectively improves the barrier function of the skin. In the present study, there was a significant increase in skin hydration compared to baseline as well as control within 4 h after application of MaxRich®, a novel moisturizing formula containing xylitol-based sugars. This is in accordance with previously discussed studies. Similarly, there was a significant decrease in transepidermal water loss associated with MaxRich® compared to both baseline and control product 4 h after application. Decrease in transepidermal water loss demonstrates that MaxRich® is associated with epidermal barrier repair. These results highlight the intense and fast acting moisturizing effect of MaxRich®.

Another important finding seen in this study is that the skin hydration was maintained above baseline value while transepidermal water loss was maintained below baseline throughout 24-h period after single application of with MaxRich®. This point toward the long-lasting effect of MaxRich®.

This being a pilot study, small sample size, and very short evaluation period are the main limitations of the study. However, these limitations do not undermine the results of the study. Long-term clinical trial with larger sample size is warranted in future.

# **CONCLUSION**

Results of our study indicate that MaxRich®, a novel moisturizer containing xylitol, xylitylglucoside, and anhydroxylitol, is associated with intense, fast, and long-lasting moisturizing effect.

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