Perspective: Stabilizing the Microbiome Skin-Gut-Brain Axis with Natural Plant Botanical Ingredients in Cosmetics

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Abstract: The microbiome of the gut and skin have recently been shown to have a strong connection through the host immune system. Various skin and gut inflammatory conditions are interrelated and connected through intricate immune pathways that affect the host barrier functions both in the skin and the gut. Microbiome ‘dysbiosis’ of the skin and gut leads to various alterations in host immune pathways that can alter the barrier and lead to disease. In this perspective article, we discuss the role of plant botanicals in cosmetics and their effect on the skin-gut-brain axis.

Keywords: microbiome; skin-gut-brain axis; plant botanical ingredients

1. Introduction

The surface microbiome opens a new dimension in looking at the very intricate communication between the external environment and the human body [1]. With a unique signature to each host, the microbiome forms a cloud of microorganisms that have specific interactions with each other and with the immune system of the host. The neuroendocrine and immune pathways of the host play an important role in establishing communication between the microbial commensals and the internal host organs in a skin-gut-brain axis [2]. The human body perhaps can be looked at as being 40% human cells to about 60% bacterial microorganisms [3]. This fact leads us to take a more serious interest in how these microorganisms interact with their specific host to impact the regulatory functions of the host [4].

2. Microbiome–Host Interaction through Immune System

Microorganisms are present from the surface to the deeper layers of the skin as well as in the epithelial lining of the gut. The specificity of microorganisms to each host as elucidated through metagenomic sequencing has led to the recognition of the microbiome as a unique signature to each host [4]. Several factors have been implicated in affecting the surface microbial colonies both on the skin and in the gut, from birth to environmental changes, geographic location, and dietary changes as well as a modern lifestyle [5]. A shift towards more immunological health conditions affecting the population, e.g., allergies, autoimmune conditions, atopic skin conditions, etc. can be attributed to these factors, which can influence the host and microbial interaction [5]. It can be inferred that between the bacterial organisms and the host surface cells there exists a symbiotic relationship. A harmonious balance between their communication exists in a physiologic state. A “dysbiotic” state adversely impacts physiological functions such as barrier protection, and may therefore lead to pathologic states. The immune cells in the surface layer are primarily the mediators of this communication in the host through a complex immune signaling pathway. Both the innate and adaptive immune system play a
role in this dialogue, affecting the various host immune organs like the lymph nodes, liver, and spleen as well as creating an interaction between the gut and skin [1,5]. Detailing the interaction of the host immune system and microorganisms is beyond the scope of this publication. However, several recent papers have elaborated the major role of Toll-like receptor (TLR) cells, dendritic cells, natural killer cells (NK cells), and Langerhans cells in microbial interactions and in passing on the effects of these interactions through an immunological cascade in the host [1–5].

3. Elaborating the Skin-Gut-Brain Axis and Various Host Neuroendocrine Pathways

Possible immunologic connections between the skin and the gut can be seen in conditions such as celiac disease, which often has an associated dermatitis, and psoriasis, where gut-associated bacteria can be more frequently found circulating in the bloodstream [6,7]. A study recently published in 2018 by Codoner et al. in Nature Scientific Reports presented data that showed a “psoriatic core intestinal microbiome” exists that clearly differs from the one present in the healthy population. They also reported that these patients had a higher incidence of gut to bloodstream translocations of microbial components with poor immune and barrier functions in the gut. Recent findings have also indicated the effect of gut microbiome on atopic dermatitis [7,8].

Gut and skin both have a neuroendocrine pathway that influences the brain function. For example, dysbiosis of the skin and gut microbes in atopic dermatitis has been connected to mental states such as depression and anxiety [9]. An established gut-brain axis is understood by the communication through multiple parallel pathways including the autonomic nervous system as well as the pituitary-hypothalamus-adrenal axis [9]. In a recent publication, the author emphasized the role of gut bacteria in aiding the delivery of nutrients needed for absorption, for example, complex oligosaccharides that need gut microbes for processing, which then helps with the absorption of minerals like calcium, magnesium, and iron from the gut [10]. These nutrients are responsible for physiological effects such as protecting the gastric mucosa, improving emotional state, supplying energy in the sleeping state, and modulating subsequent behavior, leading to brain activation and behavioral modulation in response to signaling through the gut-brain axis [9,10]. Complex oligosaccharides, essential amino acids, and various minerals are absorbed in the gut with the help of healthy gut flora [10]. Certain factors that were noted to alter gut bacteria, for example in terms of biodiversity, include animal-based foods and products where the animal source was treated with antibiotics and hormones [11]. Plant-based foods have shown a positive influence on moderate acne vulgaris, the treatment of which up until now has been through antibiotics [11].

4. Plant Botanicals that Affect Gut and Skin

Not much data is available in the field of specific botanicals regarding their role on the skin microbiome, however, there is evidence of a plant-based diet being beneficial for gut microbiome [11,12]. Given the important relationship between the skin and gut microbiome and the noted impact of plant-based diets on gut health, we can speculate that plant-based topicals in cosmetics may also act as a healthy source of nutrients for the skin microbiome and perhaps overall health.

Plants are equipped with regenerative and inflammatory pathways of their own involving potent antioxidants, phyto-acids, oleic compounds, and reactive byproducts that have been shown to have positive effects on human tissue [12,13]. Plant stem cells in cosmetics have become popular in topical formulations. Modern extraction methods have been used to obtain potent compounds from plant callus cells that have shown some anti-aging benefits [13]. It is worth considering, however, that processing, just as in food, may alter and perhaps have detrimental effects on the living plant extracts [14–17].

Ayurveda and Siddha medicine have used potent natural herbs to treat many ailments. These ancient medical systems that originated perhaps five thousand years ago in India, believe that a harmonious state of being is achieved when the personal constitution, or Dosha, is balanced with its surroundings [18]. An interrelationship is postulated between personal composition (also known as
the “Dosha”) and the environment. It is interesting to consider the similarities of this concept with that of the microbiome and host [1,18]. Recent research in herbal medicine, although limited, has shown the effects of plant botanical on keratinocyte proliferation, the reduction of inflammation, and the stimulation of wound healing [19,20]. From a modern biomedical perspective, one could consider that the “balancing of the doshas” that Ayurveda [20,21] aims for may be through actions on the skin and gut microbiome that foster symbiosis.

The fragrance of natural essential oils has been used for centuries therapeutically, and recent studies have shown a positive effect of aromatherapy in reducing stress in prepartum anxiety and overall mental state of patients going through cancer therapy [22]. The protein-based biomarker analysis showed differential effects of these 10 EOs (essential oils) in a pre-inflamed human dermal fibroblast system. Bergamot, cilantro, and spikenard EOs significantly inhibited some inflammatory, tissue remodeling, and immunomodulatory biomarkers, suggesting anti-inflammatory, immunomodulatory, and wound healing properties [23]. Most essential oils and potent herbs have multiple modalities of action on human cells and tissues by influencing cytokines, inflammatory markers, and neuroendocrine pathways (Figure 1) [24]. More research is required to understand how these pathways can relate to the microbiome–host interaction.

![Brain-Gut-Skin diagram](image)

**Figure 1.** Skin-gut-brain axis.

A list of a few plant botanicals with strong use in Ayurveda and Siddha medicine that have had recent evidence-based clinical studies showing positive skin effects can be found in Table 2.

**Table 1.** Ayurvedic herbs with recent clinical studies. Ref [25–38].

<table>
<thead>
<tr>
<th>Plant Herb/Extract</th>
<th>Skin Effect</th>
<th>Gut Effect/Systemic Effect. Mechanism of Action</th>
</tr>
</thead>
<tbody>
<tr>
<td>Amla (Indian Goose Berry)</td>
<td>Cooling effect, antioxidant As has high concentrations of Ascorbic acid</td>
<td>Aids in digestion when ingested orally, often used as a post meal digestive drink in Ayurveda to balance the Dosha imbalance Maintenance of telomere length is facilitated by an increase in telomerase activity *</td>
</tr>
<tr>
<td>Phyllanthus emblica</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Rosehip seed</td>
<td>Stimulates dermal fibroblast Improving tone texture and clarity</td>
<td>Inflammation, immune response, and tissue remodeling. Aromatic effects.</td>
</tr>
<tr>
<td>Boscovia in the family Burseraceae</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Aloe vera gel/juice</td>
<td>Wound healing and regeneration</td>
<td>Reduces diabetic neuropathy *</td>
</tr>
</tbody>
</table>

*For more details on each herb and its effects, refer to the cited references.*
Table 2. Ayurvedic herbs with recent clinical studies. Ref [25–38].

<table>
<thead>
<tr>
<th>Plant Herb/Extract</th>
<th>Skin Effect</th>
<th>Gut Effect/Systemic Effect. Mechanism of Action</th>
</tr>
</thead>
<tbody>
<tr>
<td>Rose, genus Rosa</td>
<td>Aromatic, soothing and cooling effect</td>
<td>Immunomodulation, anti-inflammatory actions but needs more detailed testing to confirm.</td>
</tr>
<tr>
<td>Rosewater/rose extract/rose essential oil/tincture</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Neem Azadirachta indica</td>
<td>Antibacterial, anti-inflammatory, antioxidant, anti-diabetic, antihelminthic</td>
<td>Anticancer. Anti-inflammatory</td>
</tr>
<tr>
<td>Tree in the mahogany family Meliaceae.</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Turmeric (Curcuma longa)</td>
<td>Anti-inflammatory, antimicrobial</td>
<td>Inflammatory bowel disease Action via various pathways</td>
</tr>
<tr>
<td>Wound healing</td>
<td>Anticancer?</td>
<td></td>
</tr>
<tr>
<td>Grapeseed</td>
<td>Antioxidant, anti-pigment, anti-inflammatory</td>
<td>Ingested reduced blood pressure and cardiovascular risk</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Anticancer, anti-inflammatory effects on cells.</td>
</tr>
<tr>
<td>Sandalwood genus Santalam</td>
<td>Anti-inflammatory, anti-pigment</td>
<td>Known to act through multiple immune pathways.</td>
</tr>
<tr>
<td>Oil/powder</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

* Currently in trials.

5. Conclusions: Future Considerations for Correlating Plant Botanicals to Surface Microbiome and Their Role in the Concept of “Vibrational Cosmetics” Innovation

The fast-paced lifestyle of the Western world has increased demand for alternative and holistic practices by the consumer with a new demand and trend for natural and organic skin care. Yoga, meditation, vegan, and Ayurvedic diets have become increasingly popular for the “modern-day yogi.” These customers also seek “natural” and “cruelty-free” ingredients in their skin care [39,40]. The global market value for natural cosmetics experienced a positive increase from almost seven billion dollars in 2007 to roughly 15 billion dollars for the year 2017, showing the growing importance of the natural and organic cosmetics market [41].

With increased recognition in the presence and functionality of the microbiome, it is clear that the human body is not what we perceived. The evidence points to the fact that our microbiome proportionally occupies a more predominant structural and functional role in the human body than previously thought. When designing a cosmetic product for the future, we need to make a shift towards understanding and developing formulations that can influence the host–microbiome balance and create a more natural and symbiotic state. For centuries, Eastern medicine has used natural herbs with minimal processing in treatment and rejuvenation with the aim of balance between the patient and their environment. The creation of modern cosmetic formulations influenced by these ancient systems of medicine could be a consideration.

An Ayurvedic herbal preparation that has recently been studied and published is Triphala [42]. It can be ingested as a tea or powder for digestive improvement and gut and liver health. Scientific papers have shown that Triphala has an apoptotic effect on colon cancer stem cells as well as a positive protective effect on dermal fibroblasts [43,44]. Triphala has three constituents, Terminalia bella, Terminalia chebula, and Emblica officinalis. The role of Triphala and its extract has been emphasized in stimulating neutrophil function. Under stress conditions such as noise, Triphala significantly prevents the elevation of IL-4 levels as well as corrects decreased IL-2 and IFN-γ levels. Under the condition of inflammatory stress, its immunosuppressive activity is attributed to its inhibitory action on the complement system, humoral immunity, cell mediated immunity, and mitogen-induced T-lymphocyte proliferation [45,46].

More information on the effects of herbal preparations on the skin and gut microbiome is needed. New research holds great promise in revealing the mechanisms linking dietary patterns with gut microbiota profiles, obesity, inflammation, and disease states. Evidence points to a distinct gut
microbiome profile in vegan and vegetarian diets that is associated with reduced inflammatory states [47].

As getting closer to nature helps improve mood and stabilize the mind, similarly, a change of dietary habits to more natural organic foods will help to shift to a healthier microbiome of the gut [48,49]. The author would like to extrapolate perhaps the important role of having cosmetic ingredients that are plant derived and minimally processed to be more suited for the skin microbiome. One can consider that with less processing, the inherent potent properties could stay intact.

Currently in a high-quality extract these actives are standardized, thus enabling a uniform high-quality extract that will always have the same concentration of actives. In unprocessed plant raw material, the concentration of actives will vary, their availability may be limited, the concentration will most likely be lower when compared to an extract and there is also the risk or degradation of actives in the unprocessed cells due to the presence of degrading enzymes in the unprocessed plant raw material. It will be worth considering that just as the gut microbiome is better suited to natural unprocessed food, one can hypothesize that the skin microbiome is capable of adjusting to the fluctuation of actives in unprocessed ingredients in a similar manner that could lead to a symbiotic state.

Still, further research and publications are required in co-relating the transdermal delivery of botanicals and their effect on the microbiome of the deeper layers of the skin [50]. A simultaneous analysis of the gut and skin microbiota with the use of topically applied botanicals and ingested as tea/herbs would help in correlating any effective action. The natural aroma of the plant botanicals needs to be individually assessed for its effects on cortisol levels and mood stabilizing effects by affecting surface microbiome when applied topically.

We can be more hopeful for a new direction in innovation for cosmetics perhaps personalized to the individual’s microbiome signature. The balancing concept of the Dosha as per the science of Ayurveda can be applied to cosmetics to achieve a balance/symbiosis of the surface microbiome using natural plant botanicals. Evidence points to the fact that the skin functions as a third brain, the second being the gut [51]. Coining the term “vibrational cosmetics” for this new dimension and approach to cosmetics innovation is plausible. Based on this new concept of vibrational cosmetics, one can envision the future of skin care as not only making one look good, but also feel good. The underpinnings of the timeless idea that healthy eating is the key to glowing skin may now be closer to being scientifically understood through the skin-gut axis.

Conflicts of Interest: The author declares no conflict of interest.

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