

## Residual Pigment Islands Treated With 88% Phenol Peeling in a Woman With Universal Vitiligo<sup>☆</sup>



### Islotes de pigmentación residual en paciente con vitiligo universal, tratados con *peeling* de fenol al 88%

To the Editor

Vitiligo is a depigmenting skin disease with a global prevalence of 1%. While multiple therapeutic options exist, none of them is completely satisfactory, especially in universal vitiligo, where depigmentation of the residual areas of pigment using chemical methods, such as hydroquinone monobenzyl ether (HMBE), may be cosmetically acceptable, as it produces complete, almost irreversible depigmentation. Other depigmenting substances exist, however, such as phenol, have been described previously but with little clinical evidence.

A 67-year-old woman with vitiligo that had appeared in childhood and was universally distributed since 14 years earlier visited our department seeking alternative depigmenting therapeutic options, as she presented areas of residual pigmentation on the face, chest, and shoulders, which had not responded to prior treatment with HMBE (Fig. 1).

An 88% aqueous solution of phenol was applied selectively, starting on the pigmented areas on the right cheek, as they were small areas that minimized systemic absorption, toxicity, and adverse effects. This was achieved by evaluating the response and tolerance of the patients and by applying cold compresses to alleviate pain immediately after frosting was observed. Given the good response, 4 applications were performed on the same section, achieving complete depigmentation, and the same procedure was performed on the other pigmented areas with excellent results and no evidence of repigmentation until 1 year after the procedure; the patient, who complied strictly with photoprotection, was very satisfied (Fig. 2).

The melanotoxic effects of phenol have been previously documented, but its depigmenting mechanism of action appears to be complex. The common characteristic of these products is their chemical structure, which includes a phenol group made up of a benzene ring with a hydroxyl side chain, a structure that is shared with the amino acid tyrosine. Potent phenols act as tyrosine analogs, interfering with the melanogenesis pathway, and initial hypotheses suggested that depigmenting chemicals that entered the melanogenesis pathway generated toxic metabolites that destroy the melanocytes. Tyrosine and other enzymes of melanogenesis bind covalently to phenols, as does tyrosine, generating reactive oxygen species and activating the response of the



Figure 1 Before treatment.



Figure 2 After treatment.

proteins released, autophagy, and exosomes, which supply adjacent immune cells with new antigens, initiate an inflammatory response, and activate autoreactive T cells, thus initiating an autoimmune response that results in their destruction.<sup>1-3</sup>

They also induce glutathione depletion, which may increase the immunogenicity of melanosomal proteins. The pigmented cells exposed to phenol activate specific T cells, also reacting against other melanocytes not directly stressed by exposure. This would explain depigmentation at a distance from these compounds through a mechanism analogous to that which occurs in contact sensitization.<sup>4</sup>

The use of phenol has not been described previously in guidelines on the management of vitiligo, even though its use may be safe and economically viable. In Brazil, however, it has been introduced successfully in clinical practice at a concentration of 88%, as a depigmenting therapy for universal vitiligo; nevertheless, a case reported in Iran obtained no response to single-drug therapy with phenol and treatment was supplemented with cryotherapy.<sup>5,6</sup>

Phenol is an aromatic hydrocarbon derived from coal tar and is used as a chemical peeling agent. Its effect

<sup>☆</sup> Please cite this article as: Alomar A, Marrón Hernández M, Bitencourt F. Islotes de pigmentación residual en paciente con vitiligo universal, tratados con *peeling* de fenol al 88%. *Actas Dermosifiliogr*. 2021;112:284-285.

varies depending on the concentration and the surface area to which it is applied. Concentrations above 80% produce denaturalization and rapid and irreversible coagulation of epidermal proteins, resulting in the formation of a barrier that prevents the chemical from penetrating the deep dermis, whereas when diluted to 50%, it acts as a keratolytic agent and disrupts the sulfur bridges, thus increasing its penetration beyond the dermis and causing greater destruction and systemic absorption.<sup>7</sup>

Complications may include scarring, dyschromia, and eczema herpeticum. High doses are toxic, and it must not, therefore, be applied over large areas, given that it has a marked corrosive action, either due to ingestion, inhalation, or direct contact. Cellular uptake is rapid and passive due to its lipophilic nature and signs of systemic toxicity appear shortly after exposure. Target organs are the liver, kidneys, lungs, and cardiovascular system. When used by qualified experts, however, it does not usually cause complications. Repigmentation of the skin may occur if patients do not protect themselves adequately from the sun.<sup>8</sup>

Although the only treatment currently approved by the US Food and Drugs Administration (FDA) for vitiligo is HMBE, few published cases describing the efficacy of phenol and isolated studies demonstrating its mechanism of action exist. Our patient presented a satisfactory response to the selective application of 88% phenol, with no complications and no relapse; we therefore consider it to be an excellent depigmenting therapeutic option in universal vitiligo with areas of pigmentation that do not respond to HMBE.

## Funding

This study has not received funding of any kind.

## Conflicts of Interest

The authors declare that they have no conflicts of interest.

## References

1. Taieb A, Alomar A, Böhm M, Dell'Anna ML, De Pase A, Eleftheriadou V, et al. Guidelines for the management of vitiligo: the European Dermatology Forum consensus. *Br J Dermatol*. 2013;168:5–19.
2. Harris J. Chemical-induced vitiligo. *Dermatol Clin*. 2018;35:151–61.
3. Grimes PE, Nashawati R. Depigmentation therapies for vitiligo. *Dermatol Clin*. 2017;35:219–27.
4. Kammeyer A, Willemsen KJ, Ouwerkerk W, Bakker WJ, Ratsma D, Pronk SD, et al. Mechanism of action of 4-substituted phenols to induce vitiligo and anti-melanoma immunity. *Pigment Cell Melanoma Res*. 2019:0–2 [Epub ahead of print].
5. Zanini M, Machado Filho CDAS. Depigmentation therapy for generalized vitiligo with topical 88% phenol solution. *An Bras Dermatol*. 2005;80:415–6.
6. Kavuoosi H. Case Report Induction of depigmentation in a universal vitiligo patient with combination of cryotherapy and phenol. *J Pakistan Assoc Dermatologists*. 2009;19:112–4.
7. O'Connor AA, Lowe PM, Shumack S, Lim AC. Chemical peels: a review of current practice. *Australas J Dermatol*. 2018;59:171–81.
8. Gupta D, Kumari R, Thappa DM. Depigmentation therapies in vitiligo. *Indian J Dermatology, Venereol Leprol*. 2012;78:49.

A. Alomar<sup>a,\*</sup>, M. Marrón Hernández<sup>b</sup>, F. Bittencourt<sup>c</sup>

<sup>a</sup> *Universidad Autónoma de Barcelona, Clínica Dermatológica Moragas, Barcelona, Spain*

<sup>b</sup> *Universidad Central de Venezuela, Cursante del Máster de Dermatología Avanzada de la Universidad Autónoma de Barcelona, Barcelona, Spain*

<sup>c</sup> *Faculdade de Medicina do ABC, Cursante del Máster de Dermatología Avanzada de la Universidad Autónoma de Barcelona, Barcelona, Spain*

\* Corresponding author.

E-mail address: [Agustin.alomar@quironsalud.es](mailto:Agustin.alomar@quironsalud.es) (A. Alomar).

30 July 2019 13 January 2020

<https://doi.org/10.1016/j.adengl.2021.01.018>

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## Allergic Contact Dermatitis Due to Slime<sup>☆</sup>



### Dermatitis alérgica de contacto por *slime*

To the Editor

Slime is a viscoelastic substance used as a toy that has become very popular with children in recent years. Making slime is a common experiment in school camps, play centers, and even at home. It can be made using any of the infinite number of recipes available on internet by mixing everyday

products such as detergent, white glue, shaving foam, and contact lens solution.

A 10-year-old girl with no personal history of atopy consulted after experiencing 2 outbreaks of pruritic erythematous-vesicular papules during the previous year. The papules, which were on the palms and interdigital folds, became scaly (Figs. 1 and 2). The patient was treated with oral and topical corticosteroids, and her lesions resolved completely. Patch testing with the standard series of the Spanish Contact Dermatitis and Skin Allergy Research Group (GEIDAC) was positive at 48 and 96 hours for methylchloroisothiazolinone/methylisothiazolinone (MCI/MI) with the True Test kit and for MI 0.2% in water.

Once the results of the test were known, the patient's history was revisited. She reported that the lesions had appeared when making and using slime without gloves (Fig. 3). She prepared the mixture with liquid deter-

<sup>☆</sup> Please cite this article as: Córdoba S, Blanco-Calvo M, Huerta-Vena A, Borbujo J. Dermatitis alérgica de contacto por slime. *Actas Dermosifiliogr*. 2021;112:285–287.