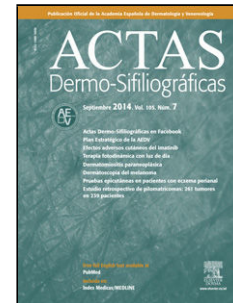


Journal Pre-proof

Phenotypes in Symptomatic Dermographism

M. Prados-Castaño M. Reguero Capilla E. Menéndez -Rivero L. Ruiz Del Barrio



PII: S0001-7310(25)00842-7

DOI: <https://doi.org/doi:10.1016/j.ad.2025.104566>

Reference: AD 104566

To appear in: *Actas dermosifiliograficas*

Received Date: 2 March 2025

Accepted Date: 23 May 2025

Please cite this article as: Prados-Castaño M, Reguero Capilla M, Menéndez -Rivero E, Ruiz Del Barrio L, Phenotypes in Symptomatic Dermographism, *Actas dermosifiliograficas* (2025), doi: <https://doi.org/10.1016/j.ad.2025.104566>

This is a PDF of an article that has undergone enhancements after acceptance, such as the addition of a cover page and metadata, and formatting for readability. This version will undergo additional copyediting, typesetting and review before it is published in its final form. As such, this version is no longer the Accepted Manuscript, but it is not yet the definitive Version of Record; we are providing this early version to give early visibility of the article. Please note that Elsevier's sharing policy for the Published Journal Article applies to this version, see: <https://www.elsevier.com/about/policies-and-standards/sharing#4-published-journal-article>. Please also note that, during the production process, errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

© 2025 Published by Elsevier España, S.L.U. on behalf of AEDV.

Sección. Research Letters

Phenotypes in Symptomatic Dermographism

Fenotipos en el dermatografismo sintomático

M. Prados-Castaño^{a,*}, M. Reguero Capilla^a, E. Menéndez -Rivero^a, and L. Ruiz Del Barrio^a.

^aUnidad de Alergología, hospital universitario Virgen del Rocío, Seville, Spain.

*M. Prados Castaño, M. Reguero Capilla, E. Menéndez Rivero, and L. Ruiz del Barrio

*Corresponding author

*Manuel Prados Castaño

E-mail address: pradosmanuel.cas@gmail.com

To the Editor,

Symptomatic dermatographism (SD), also known as factitious urticaria, is the most common form of inducible physical urticaria. It is characterized by pruritus and/or burning accompanied by a wheal that appears after rubbing, pressure, or scratching of the skin. Lesions resolve within less than 30 minutes to 2 hours after cessation of the triggering stimulus¹. Mild forms (physiologic dermatographism) affect approximately 2–5% of the population, may occur at any age, and have a mean disease duration of 3.6 to 6.9 years². Most studies, however, have evaluated disease progression in patients who were still symptomatic, rather than in those who had achieved remission. Therefore, the true mean duration is longer and, in some cases, spans decades³. SD may occur in isolation or in association with other forms of urticaria and can significantly affect patients' quality of life⁴.

Our aim was to determine whether there are distinct phenotypes that may improve our understanding of this disease and help guide future therapeutic strategies.

We conducted an observational study involving patients older than 14 years within our health care area whose clinical presentation and reason for consultation was SD, not associated with other forms of urticaria, and with a > 6-month history of the disease.

Diagnosis was confirmed by applying pressure to the back or volar aspect of the forearm with a blunt object or using the Fric-Test instrument⁴.

The study included a complete blood count, ESR, biochemical profile, serum tryptase, CRP, TSH, and antithyroid antibodies. Furthermore, we performed skin prick testing with our allergen panel (mites *D. pteronyssinus* and *Lepidoglyphus destructor*, grass pollens, olive, cypress, plane tree, mugwort, salsola, parietaria, molds *Alternaria* and *Aspergillus*, *Anisakis simplex*, latex, peach LTP, and dog, cat, and horse epithelia). Histamine served as the positive control and saline as the negative control. The test was considered positive when a wheal > 3 mm appeared to any tested allergen with a negative control. Patients refrained from antihistamines for 1 week and corticosteroids for 10 days prior to testing.

Specific IgE (ImmunoCAP®) was measured for *Dermatophagoides pteronyssinus* and *Lepidoglyphus destructor*. Sensitization to the remaining allergens was assessed via prick testing and clinical history.

We included a total of 145 patients—57 men and 88 women—with a mean age of 38 years (range, 14–86). No notable abnormalities were found in the blood count, biochemistry, CRP, or TSH. Serum tryptase was normal except in 1 patient (14.5 µg/L). Only 9 patients had elevated antithyroid antibodies.

The results of prick testing, specific IgE, and the rest of the study are shown in Figure 1.

Although the mean age and female predominance align with previously reported data, the low prevalence of antithyroid antibodies^{5–6} in our cohort may be explained by ethnic differences.

Atopy is more frequent in patients with chronic urticaria⁵ and SD⁶. In our population, it accounted for > 32% (Figure 2), slightly lower than the ~40% reported in former studies⁷. The main sensitization in both chronic urticaria and SD involves mites, and only sensitization to *D. pteronyssinus* has been studied in these conditions^{7–9}. In contrast, sensitization to *L. destructor*—which in our study showed levels comparable to *D. pteronyssinus* in prick testing and slightly lower in specific IgE—has not previously been reported in SD. This finding confirms relevant mite sensitization in these patients, who also report other sensitizations⁸. Mite sensitization has been attributed to increased exposure⁶; however, this does not align with our population, which is predominantly exposed and allergic to pollens¹⁰. Thus, atopy and sensitization to *Dermatophagoides* and *L. destructor* appear characteristic of SD, though their pathogenic role remains unclear.

A small percentage of patients exhibited sensitization to allergens other than mites as the only finding, reinforcing the observation that atopy is more frequent among these patients. Conversely, nearly one third of the cohort had completely negative testing (Figure 2), suggesting the existence of a distinct phenotype, although no explanation or references supporting this finding exist in the literature.

As summarized in Figure 2, four phenotypes were identified in our cohort: 1) sensitized to mites, 2) sensitized to non-mite allergens, 3) sensitized to both groups, and 4) those with negative results.

In conclusion, SD in our population is characterized by sensitization to allergens—primarily mites—or by entirely negative testing.

Additional studies, however, are needed to confirm these findings and evaluate immunotherapy or other treatments as potential therapeutic options.

FUNDING

None declared.

CONFLICTS OF INTEREST

None declared.

REFERENCES

1. Magerl M, Altrichter S, Borzova E, Gimenez-Arnau A, Grattan CE, Lawlor F, et al. The definition, diagnostic testing, and management of chronic inducible urticarias. The EAACI/GA(2) LEN/EDF/UNEV consensus recommendations 2016 update and revision. *Allergy* 2016;71:780-802. <https://dx.doi.org/10.1111/all.12884>. Epub 2016 Apr 6.
2. Schoepke N, Mlynek A, Weller K, Church MK, Maurer M. Symptomatic dermographism: an inadequately described disease. *J Eur Acad Dermatol Venereol* 2015;29:708-12. <https://dx.doi.org/10.1111/jdv.12661>. Epub 2014 Aug 29.
3. Kulthanan K, Ungprasert P, Tuchinda P, Chularojanamontri L, Rujitharanawong Ch, Kiratiwongwan R, Jantanapornchai N, Hawro T, Maurer M. Symptomatic dermographism: a systematic review of treatment options. *J Allergy Clin Immunol Pract* 2020; 8: 3141-61. <https://doi.org/10.1016/j.jaip.2020.05.016>.
4. Maurer M, Fluhr JW, Khan DA. How to approach chronic inducible urticaria. *J Allergy Clin Immunol Pract* 2018;6:1119-30. <https://doi.org/10.1016/j.jaip.2018.03.007>.
5. Chen Q, Yang X, Ni B, Song Z. Atopy in chronic urticaria: an important yet overlooked issue. *Front Immunol*. 2024 Feb 6;15:1279976. <https://dx.doi.org/10.3389/fimmu.2024.1279976>.
6. Cakmak ME, Yegit OO, Öztop. A case control study comparing the general characteristics of patients with symptomatic dermographism and chronic spontaneous urticaria: is atopic a risk factor for symptomatic dermographism? *Int Arch Allergy Immunol* 2024; 185:247252. <https://dx.doi.org/10.1159/000535290>.
7. Rujitharanawong C, Tuchinda P, Chularojanamontri L, Nanchaipruek Y, Jantanapornchai N, Thamlikitkul V, Kulthanan K. Natural history and clinical course of patients with dermographism in a tropical country: a questionnaire-based survey. *Asia Pac Allergy*. 2022 Oct 27;12(4):1-11. <https://dx.doi.org/10.5415/apallergy.2022.12.e39>. eCollection 2022 Oct.
8. Kulthanan K, Wachirakaphan C. Prevalence and clinical characteristics of chronic urticaria and positive skin prick testing to mites. *Acta Derm Venereol* 2008; 88: 584–588. <https://dx.doi.org/10.2340/00015555-0546>.
9. Caliskaner Z, Ozturk Z, Turan M, Karaayvaz M. Skin test positivity to aeroallergens in the patients with chronic urticaria without allergic respiratory disease. *J Invest Allergol Clin Immunol* 2004; Vol. 14(1): 50-54.
10. <https://www.seaic.org/inicio/alergológica> 2015. [Accessed 10 September 2024].

Figure 1. Figure illustrates the values obtained from the skin prick test, specific IgE, and the remainder of the diagnostic work-up.

Figure 2. Identified phenotypes and the percentage of patients in each group. A. Negative study. B. Atopic patients. C. Patients with sensitizations other than mites. D. Mite-sensitized patients.

TRADUCCIÓN DE FIGURAS

Figura 1

Prick Test

IgE específica

Otras sensibilizaciones

Estudio negativo

D. *Pteronyssinus*

L. Destructor

+ Ambos

Prick test

Specific IgE

Other sensitizations

Negative study

D. pteronyssinus

L. destructor

+ Both

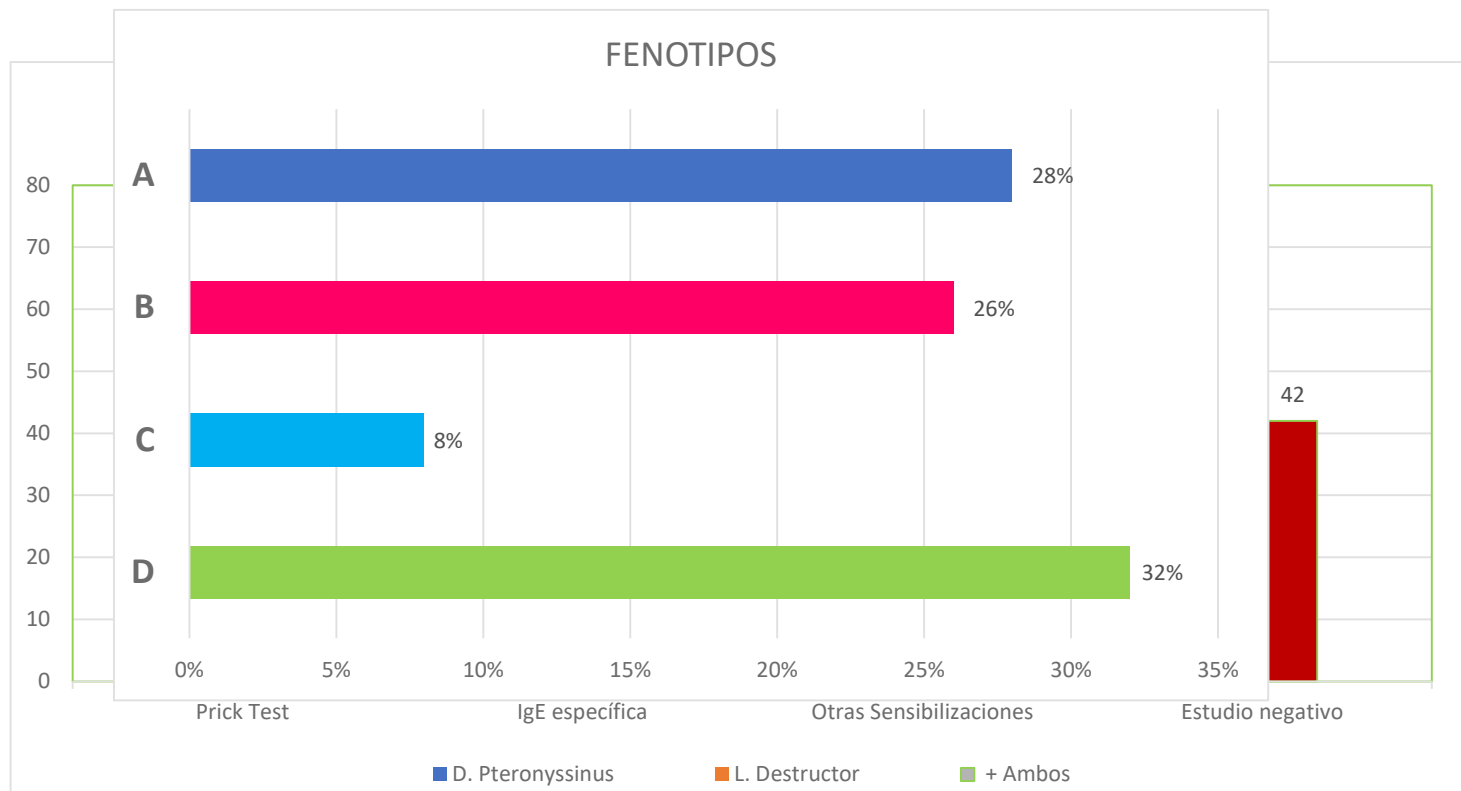


Figura 2

Fenotipos

Phenotypes