



Not actual patients.

Revealing the damage being done in AD and PN¹⁻³

Learn more about the role of a neuroimmune cytokine in atopic dermatitis (AD) and prurigo nodularis (PN) by clicking below.

[WHAT IS IL-31? >](#)

[THE DAMAGE IN AD >](#)

[THE DAMAGE IN PN >](#)

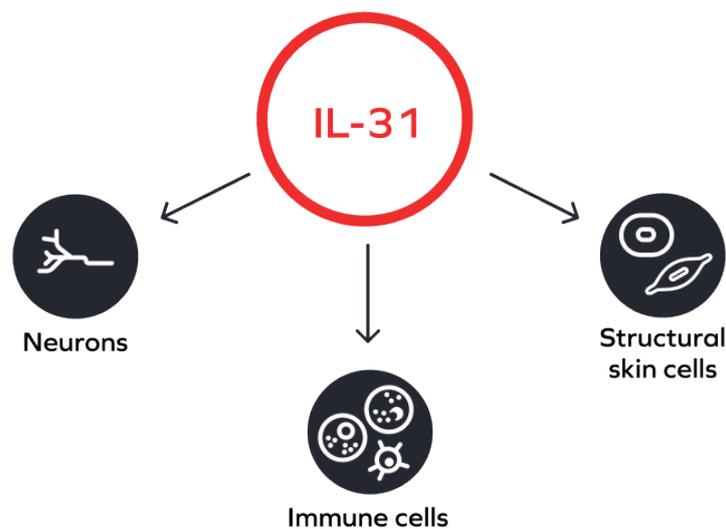
GALDERMA

IL-31 is a central mediator of neuroimmune skin dysfunction

IL-31 plays a role in many inflammatory skin diseases, including AD and PN.¹⁻³ IL-31 bridges the peripheral nervous system and immune system and targets structural skin cells, driving itch, inflammation, and skin abnormalities.¹



The center of neuroimmune skin dysfunction¹



Ready to dive deeper?

Visit [IL31role.com](https://www.il31role.com) to watch videos that give you a closer look at the role of IL-31 in AD and PN

[IL-31 IN AD >](#)

[IL-31 IN PN >](#)

A man with dark hair and a beard is shown from the chest up, wearing a dark blue long-sleeved shirt. He has visible skin conditions, including redness and small bumps on his neck and chest. He is scratching his chest with his right hand. On the right side of his neck, there are bright orange and yellow flames rising from his skin, with some sparks falling. The background is dark and smoky.

In AD,
**there may be more
damage than you think**

Learn more about a central mediator of neuroimmune dysfunction that ignites and fuels a pathologic process characterized by itch, inflammation, and skin barrier dysfunction in patients with AD.^{1,2}

[DISEASE BURDEN >](#)

Not an actual patient.

A decorative graphic at the bottom of the page consisting of several overlapping red circular arcs on a light blue background.

The signs and symptoms of AD are disrupting the lives of patients

AD is a common, chronic, inflammatory skin disease that is characterized by persistent itch, embarrassing eczematous lesions, and frequent skin infections.^{4,5}

Persistent itch is much more than just an irritating sensation—it is the most burdensome symptom for patients with AD and can significantly disrupt their daily activities, sleep, and psychological well-being.^{5,6}

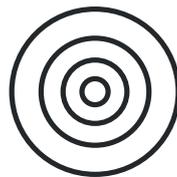
79% of patients with AD and their caregivers reported that itch was among their top 3 most problematic symptoms^{5*}

AD is driven by both inflammatory and neural mechanisms

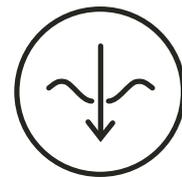
Current understanding of AD pathophysiology reveals that it is a neuroimmune-mediated disease, driven by^{2,7,8}:



Itch



Inflammation



Skin barrier dysfunction

[ROLE OF IL-31 >](#)

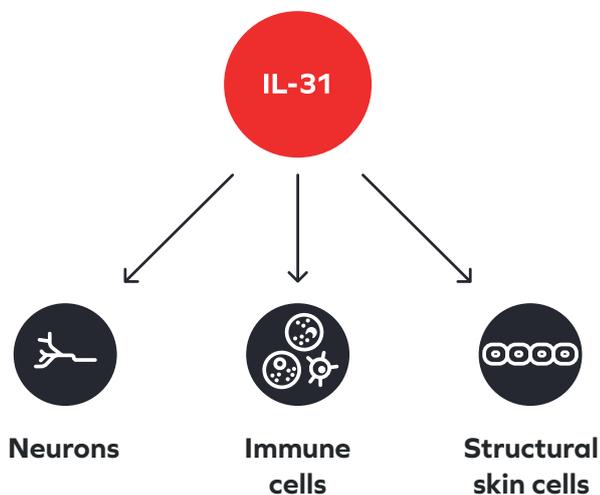
*Results were based on an online 32-item survey of 1508 patients with AD and their caregivers from the United States and 57 other countries. Survey respondents were asked which 3 symptoms have been the most problematic for the patient with AD.⁵

IL-31 is at the center of AD pathogenesis

IL-31, a neuroimmune cytokine expressed in AD skin, is a central mediator that ignites and fuels the 3 key pathophysiologic drivers of the disease: itch, inflammation, and skin barrier dysfunction.^{1,2,9}

There are many cytokines involved in the pathogenesis of AD, including IL-4, IL-5, IL-13, and IL-31.^{10,11}

The effects of IL-31 in AD



IL-31 bridges the peripheral nervous system and immune system¹

- Stimulating peripheral sensory neurons to cause itch and dermal nerve elongation and branching
- Activating immune cells to release proinflammatory cytokines

Overexpressed IL-31 also targets structural skin cells¹

- Disrupting keratinocyte differentiation and decreasing the expression of key proteins, leading to skin barrier dysfunction

There are still unmet needs in AD

AD is a chronic and extremely heterogeneous disease. Despite currently available treatment options, some patients with AD are still experiencing adverse events and a lack of disease control.^{12,13}

[IL-31 IN PN >](#)

[SUMMARY >](#)



In PN,
**there may be more
damage than you think**

Learn more about a central mediator of neuroimmune dysfunction that ignites and fuels a pathologic process characterized by itch, inflammation, and the formation of fibrotic tissue in the skin of patients with PN.^{1,3}

[WHAT IS PN? >](#)

Not an actual patient.

PN is a distinct, damaging skin disease

PN is a distinct, underrecognized, and chronic inflammatory skin disease that is characterized by disfiguring, often excoriated nodules and intractable itch.^{3,14,15} Although PN is sometimes associated with AD, most patients with PN do not have a history of AD.¹⁶⁻¹⁸

PN affects an estimated 72 per 100,000 adults aged 18 to 64 years in the United States, primarily middle-aged women and disproportionately people of African descent.^{3,16}

Common diseases associated with PN

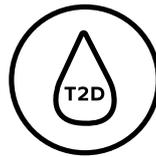
PN is associated with a variety of diseases, including³:



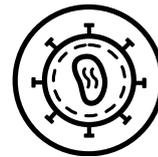
Hypertension



Chronic kidney disease



Type 2 diabetes



HIV infection



Obesity

[DISEASE BURDEN >](#)

The burden of PN is oppressive

PN severely impacts many aspects of patients' lives—it leaves some suffering with intractable itch, disfiguring nodules, debilitating sleep disturbance, and impacts on their psychological well-being.¹⁹

Studies have shown that patients with PN experience a higher burden of disease than patients with several other chronic pruritic skin diseases. This may be due to the itch of PN being more severe than that of the other skin diseases.²⁰

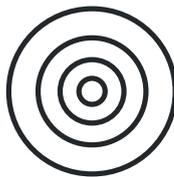
49% of adults with PN reported that itch was the most burdensome symptom of their skin disease^{17*}

What's driving PN?

A growing understanding of PN pathophysiology reveals that it is driven by neuroimmune dysfunction, which manifests as³:



Itch



Inflammation



**Fibrotic tissue
in the skin**

ROLE OF IL-31 >

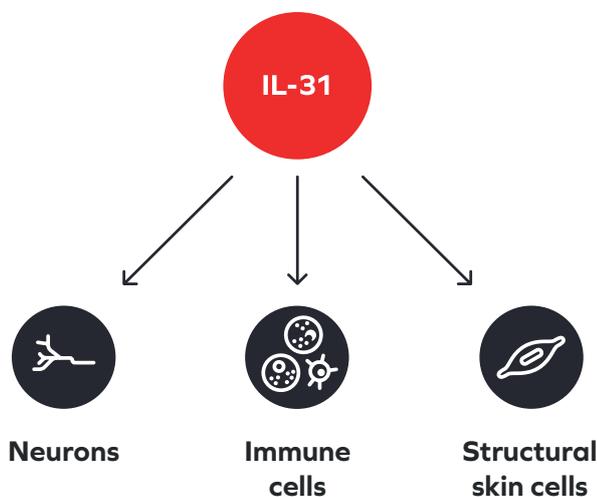
*Results were based on a multicenter, cross-sectional European study of 509 adults with PN. This prospective, questionnaire-based study assessed the clinical profile of PN, as well as its associated burdens.¹⁷

IL-31 is at the center of PN pathogenesis

IL-31, a neuroimmune cytokine expressed in PN nodules, is a central mediator that ignites and fuels the 3 key pathophysiologic drivers of the disease: itch, inflammation, and the formation of fibrotic tissue.^{1,3}

There are many cytokines involved in the pathogenesis of PN, including IL-4, IL-13, IL-22, and IL-31.^{1,3,21,22}

The role of IL-31 in PN



IL-31 bridges the peripheral nervous system and immune system¹

- Stimulating peripheral sensory neurons to cause itch^{1,3,23}
- Activating immune cells to release proinflammatory cytokines^{1,3,23}

Overexpressed IL-31 also targets structural skin cells, including dermal fibroblasts^{1,3,24}

- Leading to inhibited cell proliferation and fibrotic remodeling

The unmet need in PN

It is difficult to treat itch in PN.²⁵ Since there are currently no FDA- or EMA-approved therapies for PN, there is a need to relieve itch in affected patients.^{3,25,26} Many adults with PN identified the improvement of itch, nodules, and sleep as their top 3 most important treatment goals.^{27*}

[SUMMARY >](#)

*Results were based on a multicenter, cross-sectional European study of 509 patients with PN. This prospective, questionnaire-based study assessed patient perception of therapeutic goals, as well as previously used therapies, overall satisfaction with therapy, the efficacy of available therapeutic regimens, and out-of-pocket costs.²⁷

Consider the damage being done in AD and PN



Interested in learning more?

Visit [IL31role.com](https://www.il31role.com) to sign up for more information about IL-31, AD, and PN

The images in this material are not of actual patients with AD or PN. They were created based on informed Galderma insights. AD and PN can manifest in individuals differently.

References:

1. Nemmer JM, Kuchner M, Datsi A, et al. Interleukin-31 signaling bridges the gap between immune cells, the nervous system and epithelial tissues. *Front Med (Lausanne)*. 2021;8:639097. doi:10.3389/fmed.2021.639097
2. Datsi A, Steinhoff M, Ahmad F, Alam M, Buddenkotte J. Interleukin-31: the "itchy" cytokine in inflammation and therapy. *Allergy*. 2021;76(10):2982-2997. doi:10.1111/all.14791
3. Williams KA, Roh YS, Brown I, et al. Pathophysiology, diagnosis, and pharmacological treatment of prurigo nodularis. *Expert Rev Clin Pharmacol*. 2021;14(1):67-77. doi:10.1080/17512433.2021.1852080
4. Langan SM, Irvine AD, Weidinger S. Atopic dermatitis. *Lancet*. 2020;396(10247):345-360. doi:10.1016/S0140-6736(20)31286-1
5. McCleary KK. More than skin deep: understanding the lived experience of eczema. March 18, 2020. Accessed December 15, 2021. <http://www.morethanskindeep-eczema.org/report.html>
6. Legat FJ. Itch in atopic dermatitis—what is new? *Front Med (Lausanne)*. 2021;8:644760. doi:10.3389/fmed.2021.644760
7. Dubin C, Del Duca E, Guttman-Yassky E. The IL-4, IL-13 and IL-31 pathways in atopic dermatitis. *Expert Rev Clin Immunol*. 2021;17(8):835-852. doi:10.1080/1744666X.2021.1940962
8. Yosipovitch G, Berger T, Fassett MS. Neuroimmune interactions in chronic itch of atopic dermatitis. *J Eur Acad Dermatol Venereol*. 2020;34(2):239-250. doi:10.1111/jdv.15973
9. Neis MM, Peters B, Dreuw A, et al. Enhanced expression levels of IL-31 correlate with IL-4 and IL-13 in atopic and allergic contact dermatitis. *J Allergy Clin Immunol*. 2006;118(4):930-937. doi:10.1016/j.jaci.2006.07.015
10. Kim J, Kim BE, Leung DYM. Pathophysiology of atopic dermatitis: clinical implications. *Allergy Asthma Proc*. 2019;40(2):84-92. doi:10.2500/aap.2019.40.4202
11. Renert-Yuval Y, Guttman-Yassky E. New treatments for atopic dermatitis targeting beyond IL-4/IL-13 cytokines. *Ann Allergy Asthma Immunol*. 2020;124(1):28-35. doi:10.1016/j.anaai.2019.10.005
12. Pescitelli L, Rosi E, Ricceri F, Pimpinelli N, Prignano F. Novel therapeutic approaches and targets for the treatment of atopic dermatitis. *Curr Pharm Biotechnol*. 2021;22(1):73-84. doi:10.2174/1389201021666200611112755
13. Cork MJ, Danby SG, Ogg GS. Atopic dermatitis epidemiology and unmet need in the United Kingdom. *J Dermatolog Treat*. 2020;31(8):801-809. doi:10.1080/09546634.2019.1655137
14. Elmariah S, Kim B, Berger T, et al. Practical approaches for diagnosis and management of prurigo nodularis: United States expert panel consensus. *J Am Acad Dermatol*. 2021;84(3):747-760. doi:10.1016/j.jaad.2020.07.025
15. Whang KA, Mahadevan V, Bakhshi PR, et al. Prevalence of prurigo nodularis in the United States. *J Allergy Clin Immunol Pract*. 2020;8(9):3240-3241. doi:10.1016/j.jaip.2020.05.051
16. Huang AH, Canner JK, Khanna R, Kang S, Kwatra SG. Real-world prevalence of prurigo nodularis and burden of associated diseases. *J Invest Dermatol*. 2020;140(2):480-483.e4. doi:10.1016/j.jid.2019.07.697
17. Pereira MP, Hoffmann V, Weisshaar E, et al. Chronic nodular prurigo: clinical profile and burden. A European cross-sectional study. *J Eur Acad Dermatol Venereol*. 2020;34(10):2373-2383. doi:10.1111/jdv.16309
18. Kwon CD, Khanna R, Williams KA, Kwatra MM, Kwatra SG. Diagnostic workup and evaluation of patients with prurigo nodularis. *Medicines (Basel)*. 2019;6(4):97. doi:10.3390/medicines6040097
19. Janmohamed SR, Gwillim EC, Yousaf M, Patel KR, Silverberg JI. The impact of prurigo nodularis on quality of life: a systematic review and meta-analysis. *Arch Dermatol Res*. 2021;313(8):669-677. doi:10.1007/s00403-020-02148-0
20. Steinke S, Zeidler C, Riepe C, et al. Humanistic burden of chronic pruritus in patients with inflammatory dermatoses: results of the European Academy of Dermatology and Venereology Network on Assessment of Severity and Burden of Pruritus (PruNet) cross-sectional trial. *J Am Acad Dermatol*. 2018;79(3):457-463.e5. doi:10.1016/j.jaad.2018.04.044
21. Mullins TB, Sharma P, Riley CA, Sonthalia S. Prurigo nodularis. In: StatPearls [Internet]. StatPearls Publishing; 2021. Accessed December 15, 2021. <https://www.ncbi.nlm.nih.gov/books/NBK459204/>
22. Belzberg M, Alphonse MP, Brown I, et al. Prurigo nodularis is characterized by systemic and cutaneous T helper 22 immune polarization. *J Invest Dermatol*. 2021;141(9):2208-2218.e14. doi:10.1016/j.jid.2021.02.749
23. Zeidler C, Ständer S. The pathogenesis of prurigo nodularis—'super-itch' in exploration. *Eur J Pain*. 2016;20(1):37-40. doi:10.1002/ejp.767
24. Singh B, Jegga AG, Shanmukhappa KS, et al. IL-31-driven skin remodeling involves epidermal cell proliferation and thickening that lead to impaired skin-barrier function. *PLoS One*. 2016;11(8):1-15. doi:10.1371/journal.pone.0161877
25. Pereira MP, Basta S, Moore J, Ständer S. Prurigo nodularis: a physician survey to evaluate current perceptions of its classification, clinical experience and unmet need. *J Eur Acad Dermatol Venereol*. 2018;32(12):2224-2229. doi:10.1111/jdv.15107
26. Ständer S, Pereira MP, Berger T, et al. IFSI-guideline on chronic prurigo including nodularis. *Itch*. 2020;5(4):e42. doi:10.1097/itx.000000000000042
27. Pereira MP, Zeidler C, Wallengren J, et al. Chronic nodular prurigo: a European cross-sectional study of patient perspectives on therapeutic goals and satisfaction. *Acta Derm Venereol*. 2021;101(2):adv00403. doi:10.2340/00015555-3726

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