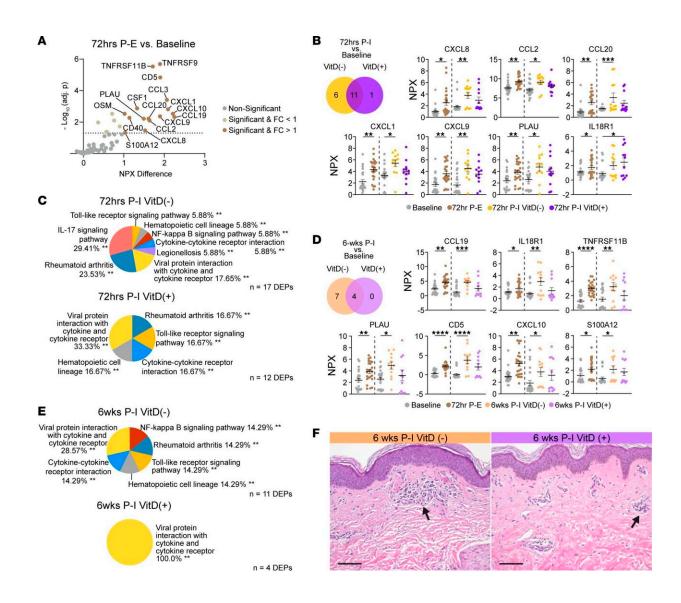


Tackling skin inflammation with vitamin D

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Vitamin D mitigates acute inflammation with durable effects and suppresses markers of NM injury involved in IL-17 signaling pathways. (A) Volcano plot of mean normalized protein expression (NPX) changes for differentially expressed proteins (DEPs) between samples taken 72hr P-E (n = 24) and baseline (n = 24) determined by paired t tests. (B) Venn diagram shows the number of DEPs



found in A that are upregulated 72hrs P-I in VitD(-) (n = 12) and VitD(+) (n = 12) relative to baseline (n = 24). Scatter plots show mean ± SEM of NPX values for labeled groups. (C) Pie charts show the percent composition of significantly enriched KEGG pathways using DEP lists from B as input from comparisons between 72hrs P-I VitD(+) and VitD(-) relative to baseline. (D) Venn diagram shows the number of DEPs found in A that are upregulated 6wks P-I in VitD(-) (n = 12) and VitD(+) (n = 11) relative to baseline. Scatter plots show mean ± SEM of NPX values for labeled groups. (E) Pie charts show the percent composition of significantly enriched KEGG pathways using total DEP lists as input from comparisons between 6wks P-I VitD(+) and VitD(-) relative to baseline. (F) Representative H&E staining from biopsies taken 6wks P-I. Scale bar: 100 μm. Arrows denote areas of inflammation in the dermis. Credit: *JCI Insight* (2023). DOI: 10.1172/jci.insight.163789. https://insight.jci.org/articles/view/163789

High-dose oral vitamin D reduced skin inflammation and increased immunoprotection in patients with chemical-induced skin rashes, according to findings from a recent Northwestern Medicine clinical trial published in *JCI Insight*.

The investigators, led by Kurt Lu, MD, the Eugene and Gloria Bauer Professor of Dermatology, also discovered in these <u>patients</u> a new molecular signature that may help physicians identify which patients have an increased risk of developing severe allergic reactions to treatments including topical chemotherapy.

"Oral <u>vitamin</u> D is a safe and readily available over-the-counter nutritional supplement with anti-inflammatory and immunoprotective properties, while boasting no adverse effects," Lu said. "This study also provides a new avenue of exploration for druggable targets to both reduce consequences of injury and enhance wound healing," Lu said.



In the randomized, double-blinded clinical trial, investigators analyzed skin tissue samples from 28 healthy participants whose inner arm skin was exposed to a test amount of topical nitrogen mustard, an FDA-approved medication with known chemical irritant properties, to produce an experimental rash. Participants then received either a high-dose oral vitamin D intervention or placebo.

Overall, skin tissue in participants that received vitamin D supplements demonstrated less irritation and inflammation than the <u>placebo group</u>.

Subsequent multi-omic analyses of skin tissue and <u>blood samples</u> from the vitamin D group also revealed suppression of the pro-inflammatory IL-17 signaling pathway.

This establishes IL-17 signaling as a promising biomarker, which is important because severe reactions to topical chemotherapy and other medications can either delay or prevent patients from receiving adequate treatment, according to Lu.

"Our molecular signature may help identify patients at-risk for developing a severe allergic response, allowing clinicians to make early adjustments to the treatment regimen or implement supportive therapies. We suggest that giving patients vitamin D when they begin <u>medical</u> <u>treatment</u> may suppress these severe reactions," Lu said.

Moving forward, Lu said his team wants to further investigate the antiinflammatory potential of oral vitamin D supplements in treating rashes caused by other types of treatments and medications, building off their previous findings <u>published in JAMA Dermatology</u> that found that vitamin D can help treat severe rashes caused by chemotherapy, also known as toxic erythema of chemotherapy.

"We would like to conduct further molecular analyses to fully elucidate



the anti-inflammatory mechanisms of Vitamin D and we will look to increase vitamin D efficacy through targeted delivery using biomimetic nanoparticles," Lu said.

More information: Madison K. Ernst et al, Vitamin D3 and deconvoluting a rash, *JCI Insight* (2023). <u>DOI:</u> 10.1172/jci.insight.163789. insight.jci.org/articles/view/163789

Provided by Northwestern University

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