Supplementary materials

Figures:

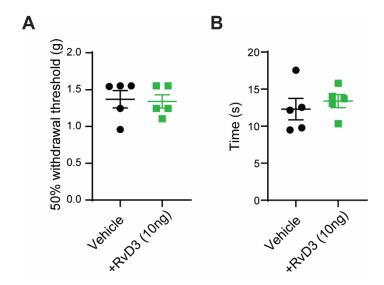


Figure S1: RvD3 does not alter mechanical and thermal sensitivity. (A) Mechanical sensitivity assessed by von Frey's test and (B) heat sensitivity assessed using the Hargreaves' in mice intradermally injected with RvD3 (10 ng) and a vehicle control (n = 5 mice/group). Statistical analysis: (A and B) two-tailed unpaired Student's t-test was used; data are depicted as $mean \pm SEM$.

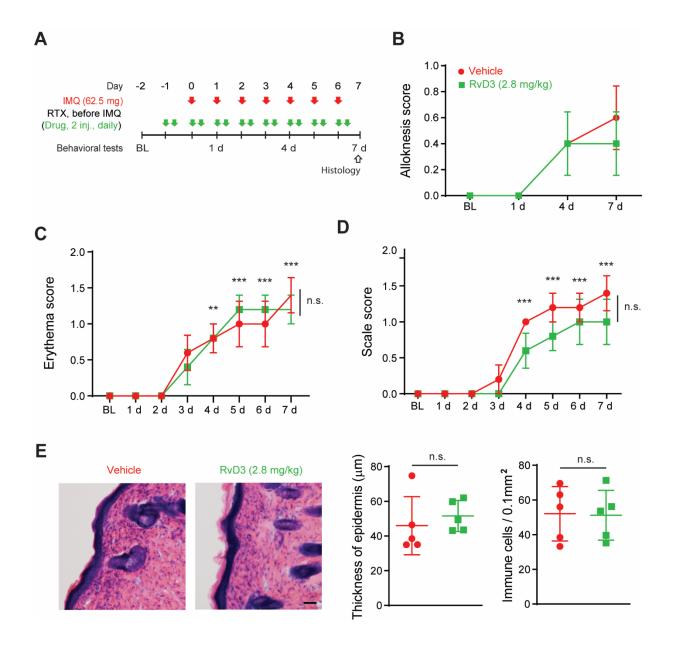


Figure S2: Time course of resiniferatoxin (RTX) and RvD3 treatments on psoriasiform itch and skin inflammation. (A) Experimental schematic indicating the daily topical applications of imiquimod (IMQ) to the nape of mice pre-treated with RTX, as well as the times of the vehicle control or RvD3 treatments and the behavioral tests. BL = Baseline (B) Time course of alloknesis scores, as well as (C and D) cutaneous erythema and scaliness in psoriatic mice (daily topical application of imiquimod cream), which were pre-treated with RTX and injected intraperitoneally with RvD3 (2 daily doses of 2.8 mg/kg, i.e. 2 inj.) or vehicle control (n = 5

mice/group). To note, spontaneous itch, erythema and scaliness significantly develop overtime in mice treated with RTX, but RvD3 treatments failed to show any additional effects compared to mice injected with a vehicle control. (E) Histopathology of skin tissues (Scale bar, $50\mu m$), epidermis thickness and immune cell infiltration at 7 days after the first IMQ application in mice treated with RTX and injected intraperitoneally with RvD3 or vehicle control (n = 5 mice/group). Statistical analysis: (A-D) two-way ANOVA followed by Bonferroni post-hoc test; (E) two-tailed unpaired Student's t-test; data are depicted as mean \pm SEM.; and **p < 0.01, ***p< 0.001. n.s. = non-significant.

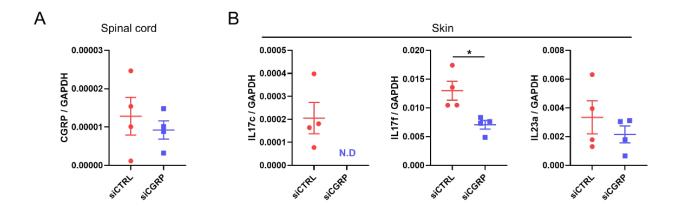


Figure S3: CGRP knockdown in DRG tissue reduces cutaneous levels of interleukin-17. (A) Spinal CGRP mRNA expression levels are unchanged, and (B) cutaneous interleukin (IL)-17c, IL-17f, and IL-23a mRNA expression levels at 7 days after the first imiquimod application in mice treated with intrathecal deliveries of siRNA (3 μ g in 10 μ L on day 0 and 5) against CGRP (siCGRP) or a control (siCTRL). Statistical analysis: (A and B) two-tailed unpaired Student's t-test was used; data are depicted as mean \pm SEM.; and *p < 0.05.

Table:

Table S1. Primer sequences for qPCR

Target gene (product length)	Primer sequences	Accession number
IL17c (283bp)	ATGCTTGTGTCGTGGATG GTGCCTGGAATGTCTGTC	NM_145834.3
IL17f (170bp)	TGCTACTGTTGATGTTGGGAC AATGCCCTGGTTTTGGTTGAA	NM_145856.2
IL22 (124bp)	ATGAGTTTTTCCCTTATGGGGAC GCTGGAAGTTGGACACCTCAA	NM_016971.2
IL23a (119bp)	TGGATACGGGGCACATTATTTTT CAGCAGCTCTCTCGGAATCTC	NM_031252.2
CGRP (102bp)	GAGGGCTCTAGCTTGGACAG AAGGTGTGAAACTTGTTGAGGT	NM_007587.2
NPPB (100bp)	GAGGTCACTCCTATCCTCTGG GCCATTTCCTCCGACTTTTCTC	NM_008726.5
SST (160bp)	ACCGGGAAACAGGAACTGG TTGCTGGGTTCGAGTTGGC	NM_009215.1
GRP (166bp)	CTGTTGGCTCTGGTCCTCTG CATACAGGGACGGGGATTCA	NM_175012.4
TAC1 (86bp)	ATTCCTTTGTTGGACTAATGGGC ACGTCTTCTTTCGTAGTTCTGC	NM_001311060.1