Frequency A (% of total cLP M ₀)	100- 80- 60- 40- 20- 0-	CTI		□ Residen ■ Infiltratin * DSS	t g
В	CTL	DSS		Top 10 DAVID GO Cate	gories
			#	DAVID GO Category	p-value
			1	Immune response	1.20E-16
			2	Response to wounding	3.50E-16
			3	Inflammatory response	1.30E-13
			4	Detense response	7.00E-13
				IL DOMOTAVIS	

6

7

8

9

10

Taxis

Cell migration

Hemopoiesis

Regulation of cell activation

Regulation of leukocyte activation

1

	Top 10 genes						
#	Gene Symbol	Fold Change					
> 1	II1f9	5.418079					
2	Pglyrp1	3.213495					
3	Cfb	3.16281					
4	Gbp2	2.925675					
5	ll1rap	2.582913					
6	Cxcl9	2.564114					
7	Rab27a	2.525242					
8	ll18rap	2.514167					
9	Cd300lf	2.395304					
10	Olr1	2.343935					

Ц

4.80E-08

3.00E-07

8.90E-07

2.40E-06

2.90E-06



Supplemental Figure 1. Inflammatory M ϕ accumulate in colons of DSS-treated mice and express IL-36 γ . (A) Frequency of resident (Cx3cr1^{hi}Ly6C⁻) and infiltrating (Cx3cr1^{lo/int}Ly6C⁺) cLP in the colon of WT mice treated for 5 d with 3% DSS. **P* < 0.05. Data are representative of three independent experiments with three to five mice per group. (B) Microarray expression analyses of cLP M ϕ from CTL or 5 d DSS-treated WT mice. Hierarchical clustering (heatmap) and top 10 GO categorization are shown along with the top 10 genes within the Immune response category. (C) IL-36 γ mRNA expression in bone marrow-derived macrophages treated for 12 h with LPS or CpG. Data are representative of two independent experiments. **P* < 0.05.



Supplemental Figure 2. The IL-36/IL-36R axis regulates neutrophil accumulation in colonic wounds and CXCL1 and CXCL2 expression by IECs. (A) Immunofluoresence on frozen sections of resealing colonic wounds in *II1rl2*^{+/+} and *II1rl2*^{-/-} mice on 2 d showing F-actin (red) and Ly6G (green), original magnification ×40. (B) Quantification of Ly6G+ cells per high power field (HPF). (C) ELISA of CXCL1 and CXCL2 in supernatants of colonic explant cultures stimulated for 24 h with IL-36 γ (100ng/mL). (D) ELISA of CXCL1 and CXCL2 on supernatants of Mode-K IEC cells stimulated as in (C). Data are representative of two independent experiments. **P* < 0.05.