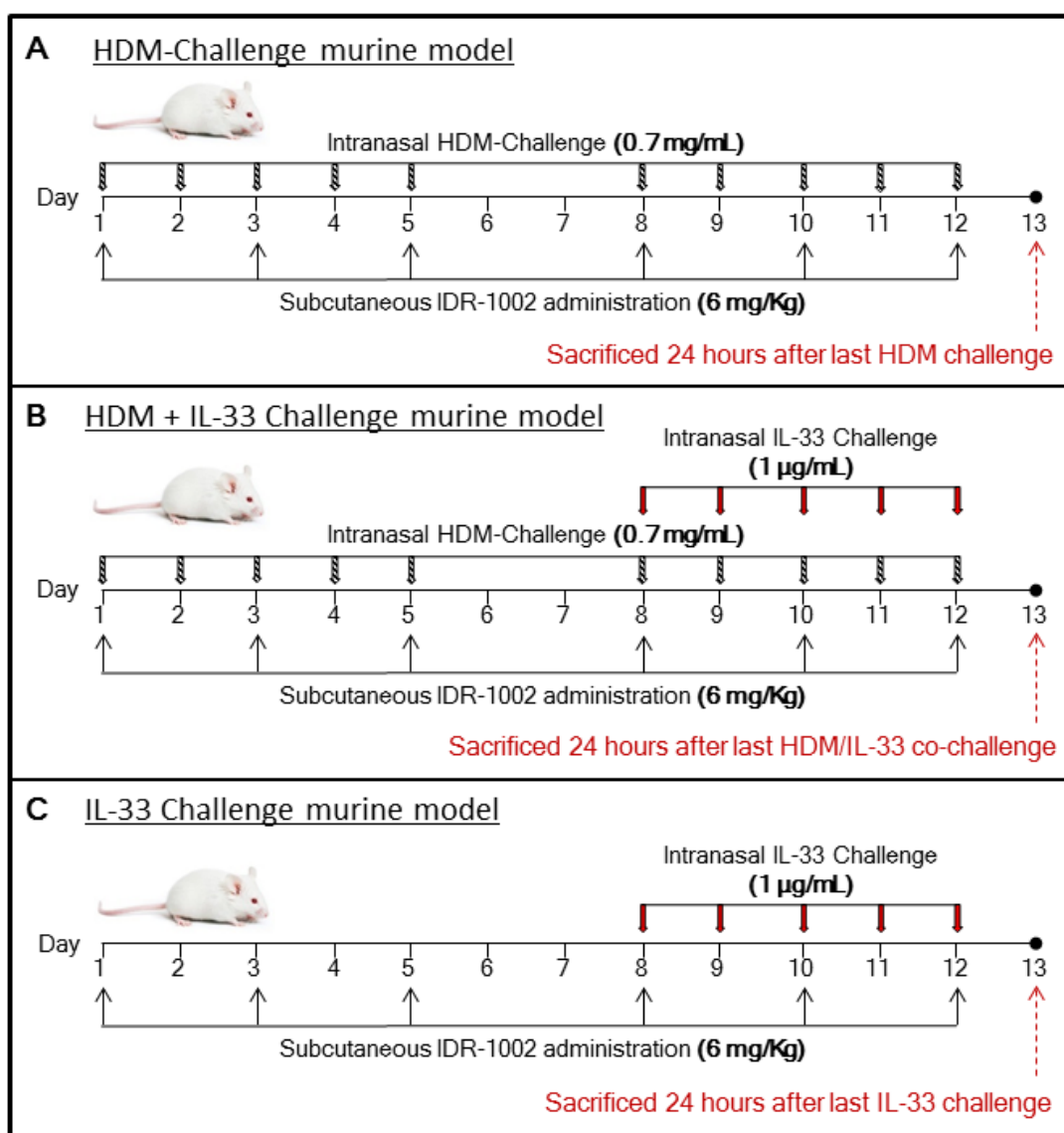
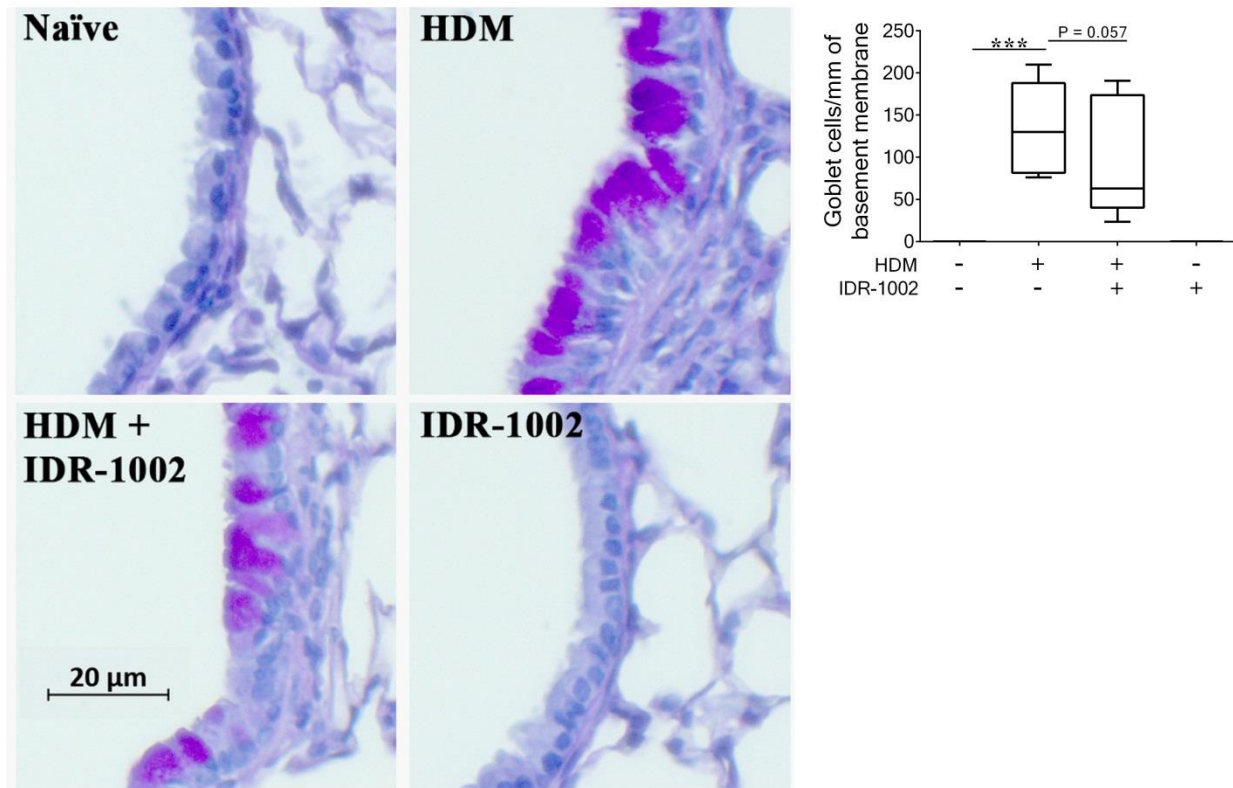


SUPPLEMENTARY INFORMATION

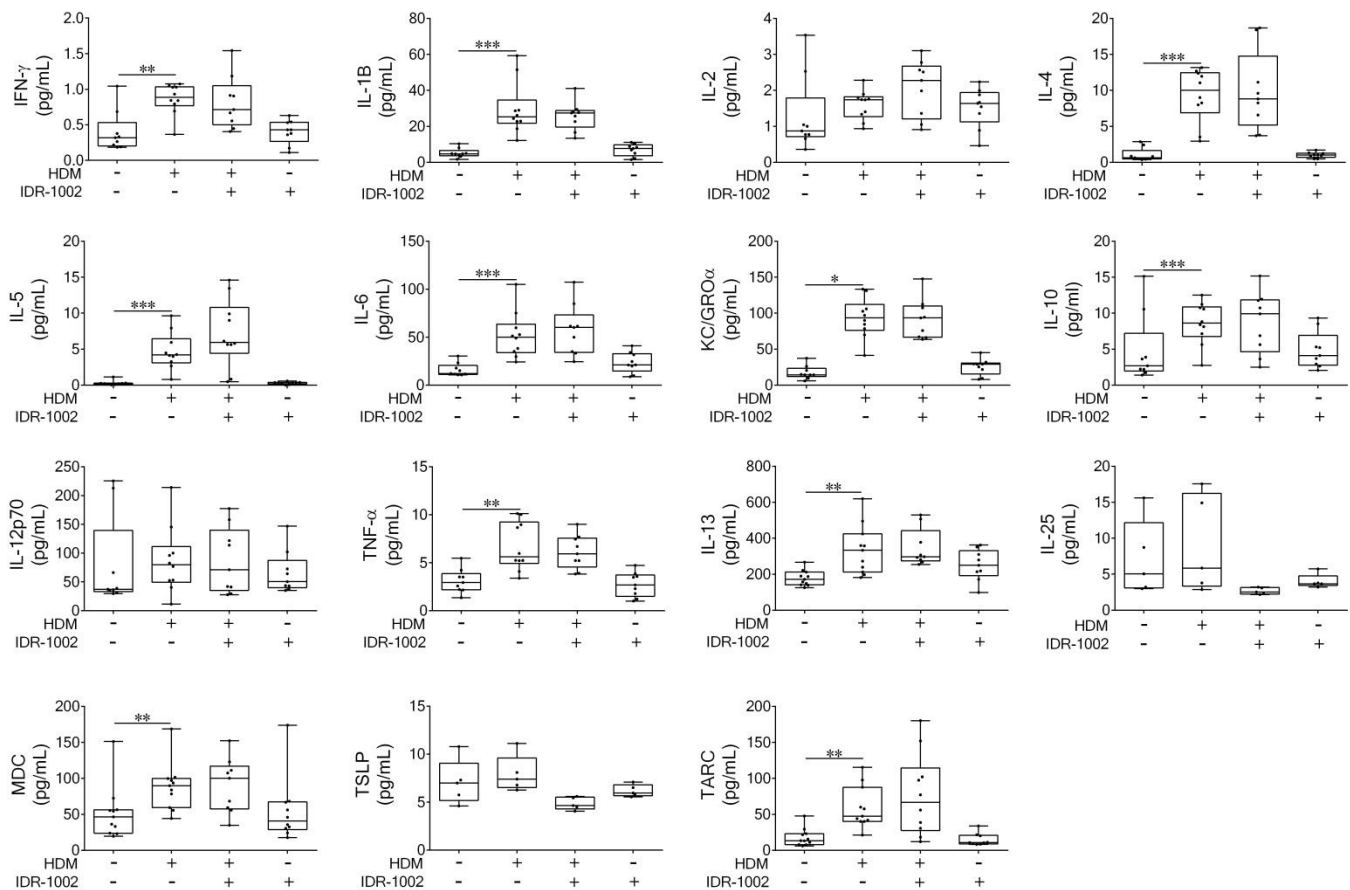
Supplementary Figure 1: Murine models. (A) HDM model; female BALB/c (8-10 wks) mice were challenged with 35 μ l of whole HDM extract (0.7 mg/mL) in saline intranasal (i.n.), for 2 weeks. IDR-1002 was administered subcutaneously (6 mg/kg) 3 times a week. (B) HDM + IL-33 co-challenge model; female BALB/c (8-10 wks) mice were challenged with 35 μ l of whole HDM extract (0.7 mg/mL) in saline intranasal (i.n.), for 2 weeks. 1 μ g of IL-33 was administered i.n. on days 8-12. IDR-1002 was administered subcutaneously 3 times a week at 6 mg/kg. (C) IL-33 model; female BALB/c (8-10 wks) mice received IDR-1002 administered subcutaneously (6 mg/kg) 3 times a week for 2 weeks, recombinant IL-33 (1 μ g per mouse) was administered (i.n.) on days 8-12.



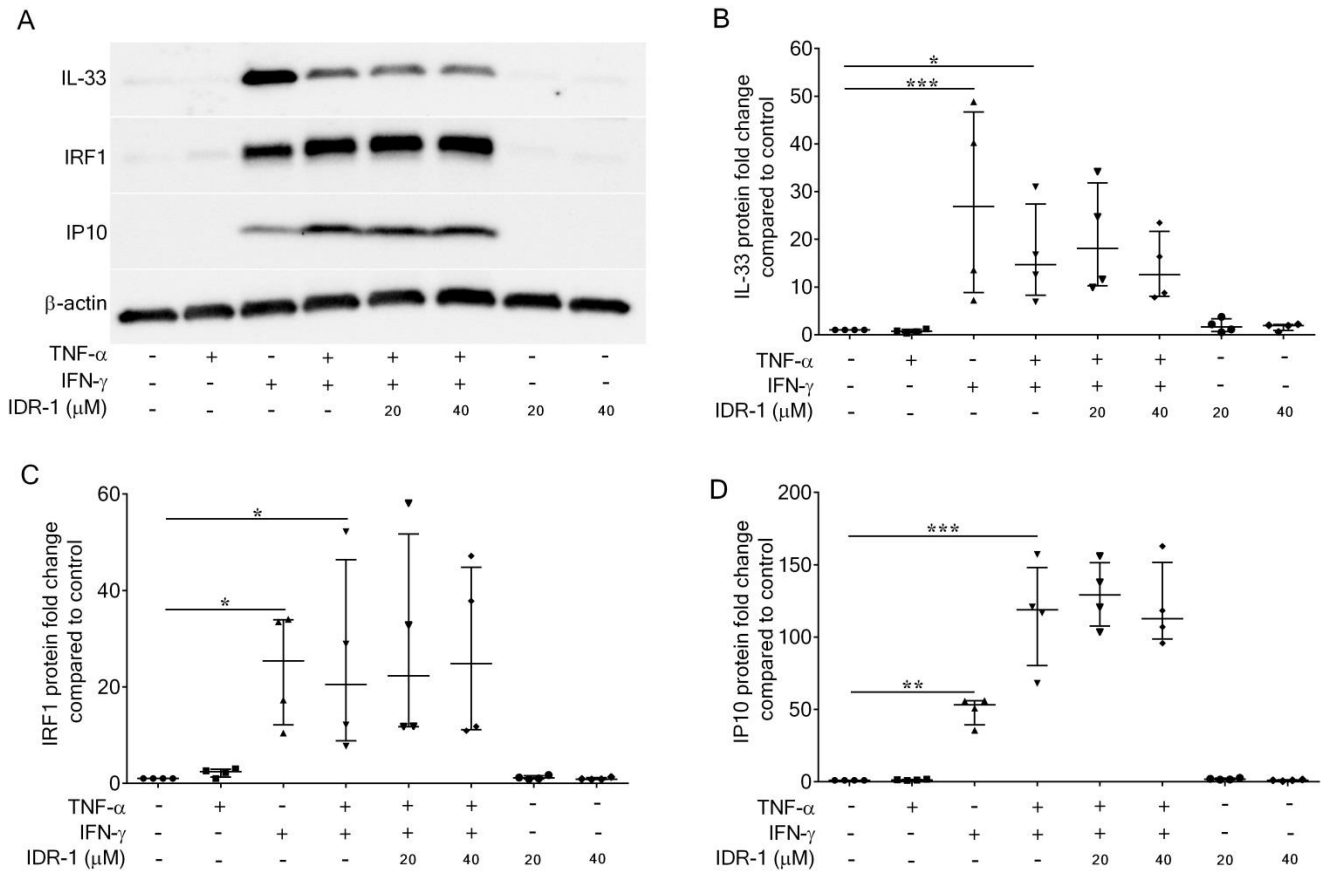
Supplementary Figure 2: Subcutaneous administration of IDR-1002 markedly decreases goblet cell hyperplasia. Female BALB/c (8-10 wks) mice (n=2-4 mice per group) were challenged with 35 μ l of whole HDM extract (0.7 mg/mL) in saline intranasal (i.n.), for 2 weeks. IDR-1002 was administered subcutaneously at a dose of 6 mg/Kg per mouse three times per week. Lung sections (6 μ m) were stained with PAS staining. **(A)** Representative images of PAS staining of the lung tissue. **(B)** Number of goblet cells per mm of basement membrane length. Each dot represents individual airways. Statistical significance was determined by one-way ANOVA with Tukey's multiple comparisons test



Supplementary Figure 3. Cytokine expression profile in lung homogenates of HDM-challenged mice, in the presence and absence of IDR-1002. Female BALB/c (8-10 wks) mice (n=9-10 mice per group) were challenged with 35 μ L of whole HDM extract (0.7 mg/mL) in saline intranasal (i.n.), for 2 weeks. IDR-1002 was administered subcutaneously at a dose of 6 mg/Kg per mouse three times per week. Lung tissue homogenates were monitored for a panel of cytokines, 24 hr after the last HDM challenge. Production of IFN- γ , IL-1 β , IL-10, IL-12 p70, IL-2, IL-4, IL-5, IL-6, KC, TNF- α , IL-33, TSLP, IL-25, MDC and TARC were monitored by the multiplex Meso Scale Discovery (MSD) platform or ELISA. Bar's shows median and interquartile range, whiskers show min and max points. One-way ANOVA with Tukey's multiple comparisons test was used to assess statistical significance (* $p \leq 0.05$, ** $p \leq 0.01$, *** $p \leq 0.001$).



Supplementary Figure 4: Peptide IDR-1 does not alter IL-33 production in human primary bronchial epithelial cells (PBEs). Human Primary Bronchial Epithelial cells obtained from 4 donors were stimulated with TNF α (20 ng/mL) and IFN γ (30 ng/mL), in the presence and absence of IDR-1 (20 and 40 μ M). IL-33, IRF1 and IP10 abundance was monitored in cytoplasmic fractions of the cell lysates by western blots, 24 hr post-stimulation. Protein abundance was quantified by densitometry. (A) A representative immunoblot for all proteins, and densitometry analysis (n=4) for (B) IL-33, (C) IRF1 and (D) IP-10 are shown. Protein fold change shown in the graphs represents relative band intensity compared to that in unstimulated cells normalized to 1, after normalization with β -actin for protein input. Each dot represents an individual donor, and bars show the median and interquartile range. RM one-way ANOVA with Fisher's LSD test was used for statistical analyses (* $p \leq 0.05$, ** $p \leq 0.01$).



Supplementary Table 1: Demographics of donors of human PBECs used in this study.

| Donor # | Gender | Age | Height | Weight | BMI | Current smoker | Ex-Smoker | Pack years | Oral Steroids | Inhaled Steroids |
|---------|--------|-----|--------|--------|-------|----------------|-----------|------------|---------------|------------------|
| BR384 | Female | 55 | 168 | 90 | 31.9 | No | Yes | unknown | Yes | No |
| BR390 | Female | 68 | 162 | 61 | 23.24 | No | Yes | 55 | No | No |
| BR421 | Male | 56 | 181 | 83 | 25.3 | No | Yes | unknown | unknown | unknown |
| BR448 | Male | 74 | 160 | 68 | 27 | No | No | 0 | No | No |