Supplemental Figure 1: Controls for mBD3 and CRAMP Immunostaining:

Immunostaining was performed for mBD3 (A-C) and CRAMP (D-F) and in CRAMP k.o. mice (B, D). Panels C and F were immunostained with secondary antibody only. Note marked decrease in CRAMP staining in CRAMP k.o. mice (E); in contrast, the epidermis of some CRAMP k.o. mice demonstrates increased mBD3 protein (B). Mag bars = 50 μm. G: The full-length mBD3 (3.5-4 kDa) is detectible on a 10% tricine gel between 110 and 140 kDa. The western immunoblotting shows the full-length peptide (left) and mBD3 protein in normal (non-stressed) mouse epidermis (right).
Supplemental Figure 2: **Exogenous GC Down-Regulate CRAMP (but not mBD3)**

**mRNA Expression In Vitro:** Second-passage, cultured human keratinocytes were treated with a single dose of dexamethasone (10 ng/ml ethanol) or same volume of vehicle alone (controls) for 18 hrs, followed by RNA extraction and quantitation by rt-PCR, as in Methods.

Supplementary Figure 3: **Normal Permeability Barrier Recovery in Adrenalectomized Mice:**

Adrenalectomized and sham-operated mice (n=5 each) were
tape-stripped until TEWL levels $\geq 5x$ normal, and recovery rates were compared 2 and 4 hours later. The differences in recovery rates in the two groups were not significant.

Supplementary Figure 4: Exogenous Physiologic Lipids Restore Lamellar Bodies in Psychologically Stressed (PS) Epidermis: A: PS epidermis under basal conditions (PS + Ba) displays few lamellar bodies (LB) in cytosol of outer stratum granulosum (SG) cells. Note proximity of secreted LB contents at SG-stratum corneum (SC) interface (open arrows). B: PS animals treated with vehicle alone (PS + Veh). C: PS animals treated with physiologic lipids (PS + L). Few LB are present in SG cytosol of vehicle-treated, PS mice (arrowheads), and secreted LB contents at SG-SC interface remain diminished (B, open arrows). In contrast, PS and lipid-treated animals reveal numerous LB in SG cytosol (C, arrowheads), and abundant, newly-secreted LB contents at SG-SC
interface (C, solid arrows). A, ruthenium tetroxide post-fixation; B & C, osmium tetroxide post-fixation. Mag bars = 0.5 _m.

Supplementary Figure 5: Topical Physiologic Lipids Do Not Normalize

**Immunostaining for CRAMP after PS/GC:** Hairless mice (n= 4 or 5 in each cohort) received either equimolar mixture of ceramides, cholesterol and free fatty acids (1:1:1 molar ratio; 2% final concentration) in propylene glycol:ethanol (7:3 vols) vehicle (60 _l vols to 3cm²), or vehicle alone, while being co-treated with either PS (E, F) or topical clobetasol (GC) (B, C), as above. Frozen sections were immunostained for CRAMP and mBD3, as above. Mag bars = 50 _m.