## SUPPLEMENTARY TABLE S1

## Effect of Topical Application of RTA 408 after Fractionated Radiation Exposure on Skin <br> Histological Grading

| Group ${ }^{\text {a }}$ | Epidermal <br> Thickening | Follicular Hypertrophy | Follicular Atrophy | Thickened Collagen | Inflammation | Ulceration | Necrosis | Parakeratosis | Sum |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Day 16 |  |  |  |  |  |  |  |  |  |
| Vehicle | $2.4 \pm 0.3$ | $2.1 \pm 0.2$ | $1.4 \pm 0.8$ | $2.0 \pm 0.2$ | $1.8 \pm 0.5$ | $1.4 \pm 0.9$ | $0.1 \pm 0.1$ | $2.6 \pm 0.4$ | $13.8 \pm 1.8$ |
| $\begin{gathered} \hline \text { RTA } 408 \\ (0.01 \%) \\ \hline \end{gathered}$ | $3.5 \pm 0.3$ | $2.7 \pm 0.6$ | $2.0 \pm 1.0$ | $3.0 \pm 0.3$ | $1.8 \pm 0.2$ | $2.2 \pm 0.7$ | $1.2 \pm 0.2$ | $3.2 \pm 0.2$ | $19.5 \pm 1.0$ |
| $\begin{gathered} \text { RTA } 408 \\ (0.1 \%) \\ \hline \end{gathered}$ | $3.1 \pm 0.4$ | $2.6 \pm 0.4$ | $0.8 \pm 0.3$ | $2.6 \pm 0.6$ | $2.0 \pm 0.4$ | $0.3 \pm 0.1$ | $0.1 \pm 0.1$ | $1.8 \pm 0.7$ | $13.3 \pm 2.1$ |
| $\begin{gathered} \text { RTA } 408 \\ (1.0 \%) \\ \hline \end{gathered}$ | $3.0 \pm 0.2$ | $2.1 \pm 0.1$ | $0.6 \pm 0.3$ | $2.1 \pm 0.4$ | $1.6 \pm 0.3$ | $0.0 \pm 0.0$ | $0.0 \pm 0.0$ | $1.5 \pm 0.4$ | $11.0 \pm 1.4$ |
| Day 30 |  |  |  |  |  |  |  |  |  |
| Vehicle | $2.2 \pm 0.2$ | $0.4 \pm 0.2$ | $1.4 \pm 0.2$ | $1.8 \pm 0.3$ | $1.7 \pm 0.2$ | $0.1 \pm 0.1$ | $0.0 \pm 0.0$ | $1.4 \pm 0.2$ | $9.0 \pm 0.8$ |
| $\begin{gathered} \hline \text { RTA } 408 \\ (0.01 \%) \\ \hline \end{gathered}$ | $2.3 \pm 0.2$ | $0.8 \pm 0.1$ | $1.7 \pm 0.4$ | $1.4 \pm 0.3$ | $1.6 \pm 0.2$ | $0.4 \pm 0.3$ | $0.1 \pm 0.1$ | $1.9 \pm 0.4$ | $10.2 \pm 1.2$ |
| $\begin{gathered} \text { RTA } 408 \\ (0.1 \%) \\ \hline \end{gathered}$ | $1.8 \pm 0.3$ | $0.5 \pm 0.1$ | $0.4 \pm 0.2 \dagger$ | $0.9 \pm 0.3 *$ | $1.0 \pm 0.2^{* *}$ | $0.0 \pm 0.0$ | $0.0 \pm 0.0$ | $0.9 \pm 0.3$ | $5.5 \pm 1.1 *$ |
| $\begin{gathered} \text { RTA } 408 \\ (1.0 \%) \\ \hline \end{gathered}$ | $1.8 \pm 0.3$ | $0.4 \pm 0.1$ | $0.9 \pm 0.2$ | $0.6 \pm 0.2 * *$ | $0.9 \pm 0.1^{* *}$ | $0.0 \pm 0.0$ | $0.0 \pm 0.0$ | $0.9 \pm 0.2$ | $5.7 \pm 0.9 *$ |
| Day 40 |  |  |  |  |  |  |  |  |  |
| Vehicle | $2.2 \pm 0.1$ | $0.8 \pm 0.3$ | $1.3 \pm 0.2$ | $1.4 \pm 0.3$ | $1.8 \pm 0.3$ | $0.5 \pm 0.2$ | $0.4 \pm 0.1$ | $1.2 \pm 0.2$ | $9.6 \pm 1.0$ |
| $\begin{gathered} \text { RTA } 408 \\ (0.01 \%) \\ \hline \end{gathered}$ | $2.3 \pm 0.5$ | $0.5 \pm 0.2$ | $0.6 \pm 0.1$ | $2.0 \pm 0.4$ | $1.5 \pm 0.4$ | $0.1 \pm 0.1 *$ | $0.1 \pm 0.1$ | $1.0 \pm 0.5$ | $8.1 \pm 1.9$ |
| $\begin{gathered} \text { RTA } 408 \\ (0.1 \%) \\ \hline \end{gathered}$ | $1.5 \pm 0.3 \dagger$ | $0.2 \pm 0.1$ | $0.4 \pm 0.2 *$ | $1.3 \pm 0.1$ | $1.2 \pm 0.2$ | $0.1 \pm 0.1 *$ | $0.1 \pm 0.1^{*}$ | $0.6 \pm 0.3$ | $5.4 \pm 0.9 *$ |
| $\begin{gathered} \text { RTA } 408 \\ (1.0 \%) \\ \hline \end{gathered}$ | $0.7 \pm 0.2 * *$ | $0.1 \pm 0.1$ | $0.9 \pm 0.2$ | $0.2 \pm 0.2 * *$ | $1.0 \pm 0.3$ | $0.0 \pm 0.0^{* *}$ | $0.0 \pm 0.0^{* *}$ | $0.2 \pm 0.1 \dagger$ | $3.1 \pm 0.8^{* *}$ |

${ }^{\text {a }}$ Mice were irradiated with $10 \mathrm{~Gy} /$ day of radiation on days $0-2$ and 5-7 and topically administered vehicle (sesame oil) or RTA $408(0.01 \%, 0.1 \%$, or $1.0 \%$ ) once daily on Days -5 to up to 40 . On days of irradiation, vehicle or RTA 408 was applied after exposure to radiation. H\&E stained slides of skin were evaluated by a board (ACVP)-certified pathologist. Skin was scored subjectively on a scale for epidermal thickening, follicular hypertrophy, follicular atrophy, thickened collagen, inflammation, ulcer, necrosis, keratin cysts, and parakeratosis. Scores were: $1=$ minimally detectible, $2=$ mild, $3=$ moderate, $4=$ marked and $5=$ severe. Summed scores for each animal were also calculated on each day. Data are presented as mean $\pm$ S.E.M. Statistical notations vs. vehicle control group are shown ( ${ }^{* *} \mathrm{p}<0.01,{ }^{*} \mathrm{p}<0.05$, and $\dagger \mathrm{p}<0.1$ ).

Supplementary Fig. S1

A


B







