## Supplemental materials

Fig. S1. Surface-enhanced Raman spectrometric analysis. For simplicity, pinocmebrin but not propolis was used in the experiment, where solutions of pinocmebrin $(100 \mu \mathrm{M})$ in PBS with or without $50 \mu \mathrm{M} \mathrm{FeSO}$ (pinocmebrin $+\mathrm{Fe}^{2+}$ or pinocmebrin) were analyzed and $\mathrm{H}_{2} \mathrm{O}_{2}(20 \mu \mathrm{M})$ was used as the control. The arrow shows the peak presumably due to $\mathrm{H}_{2} \mathrm{O}_{2}$. Technical service was provided by TopTekEnwave Optronics Inc., (Nei-Hu,Taipei).

Fig. S2. Propolis-Induced oxidative DNA damage is subject to repair. AGS cells in overnight culture at $50 \%$ confluency were treated with $0.3 \mu \mathrm{~g} / \mathrm{ml}$ propolis for 1 h before being analyzed for oxidative DNA damage at the indicated time points (filled circle) Assays without Endo III and Fpg (E+F) were used as control (open circle), indicating that propolis did not cause strand break.

Fig. S3. Cellular adaptation to propolis-induced oxidative DNA damage. AGS cells in overnight culture at $50 \%$ confluency were treated with or without $0.3 \mu \mathrm{~g} / \mathrm{ml}$ propolis for 8 h and then treated again with propolis at the same dose for 1 h before being harvested for analysis of oxidative DNA damage with the comet assay (columns 3 and 4). Cells without any treatment or cells received treatment with propolis for 8 h alone, columns 1 and 2, respectively, were used as controls. The asterisk (*) indicates statistical significance ( $p<0.01 ; n=3$ ) between the indicated experiments. Error bars for SE.

Fig. S4. Glutathione reductase (GR) activity of cell extracts of CL1-0 cells and the GR RNAi knockdown strain. The knockdown strain was selected from CL1-0 cells transfected with the pLKO.1-shGR vector in puromycin ( $0.8 \mu \mathrm{~g} / \mathrm{ml}$ ) containing medium. GR activity was measured as described previously (Rescigno et al., Biochemistry 1994, 33: 5721-5727). The asterisk (*) indicates statistical significance ( $p<0.01$; $n=3$ ) between the indicated experiments. Error bars for SE.

Fig. S1.


Fig. S2.


Time after removal of propolis (h)

Fig. S3.


Fig. S4.


