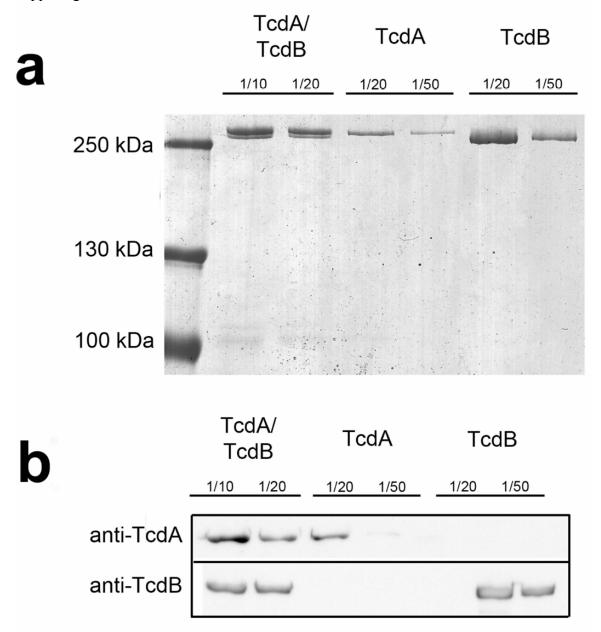
**Supplementary Figure 1.** Purification of TcdA and TcdB from filtered supernatants of *C.difficile* NAP-1/027 cultures. **(a)** Coomassie-stained SDS-PAGE gel with TcdA/TcdB (100kDa filtered input), TcdA- and TcdB-containing fractions. **(b)** Western blotting of the TcdA/TcdB, TcdA- and TcdB-containing fractions with mouse monoclonal anti-TcdA and TcdB antibodies.

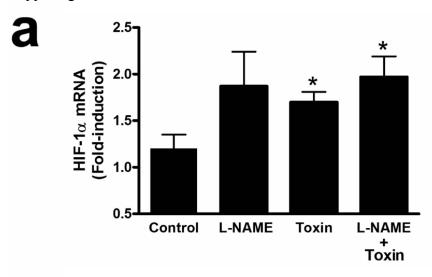
**Supplementary Figure 2.** Inhibition of NO synthases with L-NAME (100μM) has no effect on toxin-induced (100μg/mL) upregulation of HIF-1α gene expression (a) as assessed with qPCR; \* denotes p<0.05 compared to control, n=4. (b) L-NAME did not alter toxin-induced changes in TER. Caco-2 cells were exposed to vehicle (control), *C.difficile* toxin (60μg/mL), L-NAME (100μM) with *C.difficile* toxin (60μg/mL) or L-NAME (100μM) alone. Time (min) indicates the time elapsed following addition of toxin to monolayer, data are expressed as a percentage of the control TER, #p<0.005 compared to control, n=4.

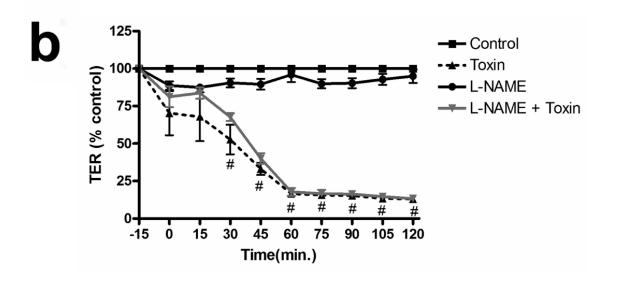
**Supplementary Figure 3.** Treatment of Caco-2 cells with dimethyloxallyl glycine (DMOG, 500  $\mu$ M for 2h) upregulates HIF-1 $\alpha$  protein.

Suppl. Figure 1.



Suppl. Figure 2.





Suppl. Figure 3.

