## Supporting information

## Tannic acid – iron complex-based nanoparticles as a novel tool against oxidative stress

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Figure S1. Chemical structure of tannic acid.



**Figure S2.** Normalized absorbance at 570 nm for different concentrations of L-TAFe NPs obtained in the total antioxidant capacity assay. The three chosen concentrations fall within the linearity range.



**Figure S3**. Human primary skin fibroblasts viability after treatment with different concentrations of TBH for 1 h. Analyses were normalized to control non-treated cells (CTRL; \* p < 0.05 with respect to CTRL).



**Figure S4**. *In vivo* biocompatibility results. Planarians survival is 100% after the administration of L-TAFe NPs at both 10 and 100  $\mu$ g/mL after seven days of incubation.



**Figure S5**. Intensity distribution (%) as a function of the hydrodynamic diameter (nm) of L-TAFe NPs in different conditions (H<sub>2</sub>O, PBS, DMEM+FBS 10%, DMEM+FBS+H<sub>2</sub>O<sub>2</sub>) at different time points (0, 1, 3, 6, 10 days).



**Figure S6**. UV/Vis spectra of L-TAFe NPs in water (black) and PBS (blue). In the inset, a picture showing the color of L-TAFe NPs in the two different solutions is reported.



**Figure S7.** (A) Derivative weight (%) of tannic acid, DSPE-PEG, and L-TAFe NPs. (B) Peak deconvolution of the L-TAFe NP thermal event.



**Figure S8**. Human primary skin fibroblasts viability after treatment with 5  $\mu$ M of tannic acid, L-ascorbic acid, or and N-acetyl-L-cysteine for 24 h (WST-1 assay). Analyses were normalized to control non-treated cells (CTRL).



**Figure S9.** Micrographs showing either dead or live planarians (prepared and fixed for imaging), as typically obtained for each of the indicated conditions in terms of TBH concentration, presence/absence of L-TAFe NPs, and time point (in terms of experimental day). In particular, a freshly dead specimen is shown on the left, and four whole planarians are portrayed in the remaining images. All pictures have the same magnification.



**Figure S10**. *In vivo* toxicology results following DPI (2.5  $\mu$ M) administration, used as antioxidant positive control. DPI may exert a modest protective activity at low TBH concentrations (150  $\mu$ M and 200  $\mu$ M) compared to untreated animals, but such effects are not statistically significant (two-tailed unpaired *t*-test,  $\alpha = 0.05$ ). Plots have been obtained through the assignment of a time-consistent mortality score, starting from 8 for the day of TBH administration down to a score of 1 for day 7 (0 was assigned to survived specimens), followed by averaging scores for each experimental class and computing of standard error.