## **Supporting Figures:**

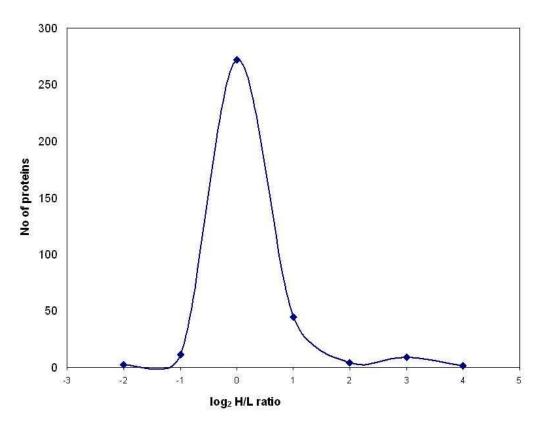


Figure S1. Lognormal distribution of the quantified proteins based on the variability at the heart proteome after 8 Gy. Fold changes (H/L ratio) of all identified proteins were log2 transformed and binned into indicated intervals. Normalization factor:  $1.004 \pm 0.173$ 

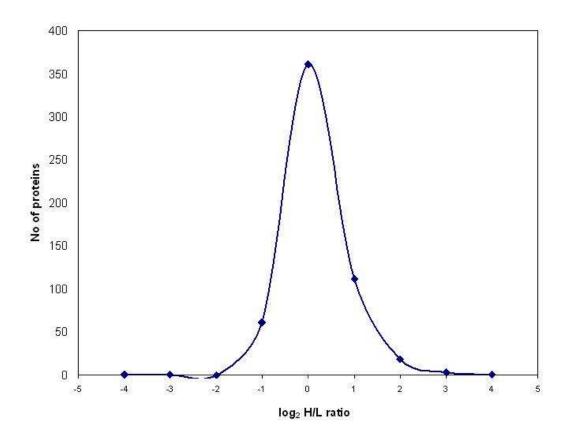
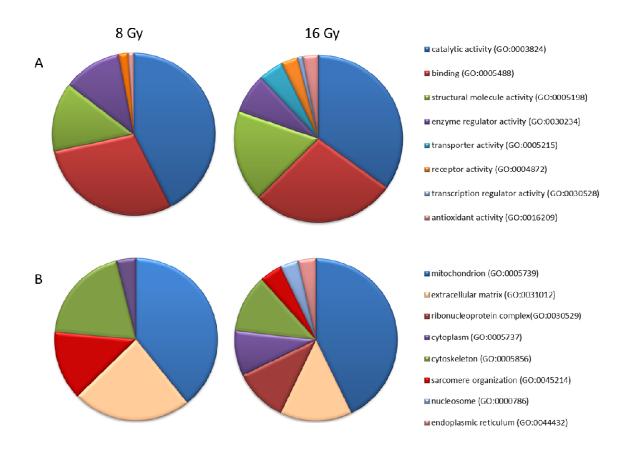


Figure S2. Lognormal distribution of the quantified proteins based on the variability at the heart proteome after 16 Gy. Fold changes (H/L ratio) of all identified proteins were log2 transformed and binned into indicated intervals. Normalization factor:  $0.82 \pm 0.078$ 



**Figure S3.** Gene ontology (GO) analysis of significantly differentially regulated proteins. The most radiation-responsive GO categories "Molecular function" (A), and "Cellular compartment" (B) using the PANTHER (<a href="http://www.pantherdb.org">http://www.pantherdb.org</a>) and DAVID bioinformatics tools (<a href="http://www.david.niaid.nih.gov">http://www.david.niaid.nih.gov</a>) are shown. Catalytic activity and binding were the most altered cellular functions by ionizing radiation. The mitochondrial proteins were the most affected protein class.

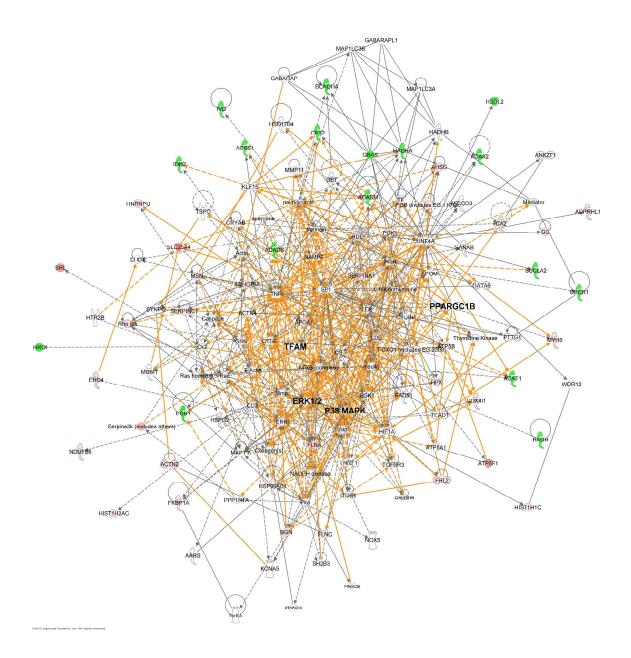


Figure S4. Graphical representation of the merged of the most significant protein interaction network after 8 Gy using Ingenuity Pathway Analysis of differentially regulated proteins. All significantly deregulated proteins identified by ICPL were imported into the Ingenuity Pathway Analysis. The up-regulated proteins are marked in red and the down-regulated in green. The nodes represent proteins that are connected with one or several arrows; the solid arrows represent direct interactions and the dotted arrows indirect interactions; direct and predicted protein interactions were searched for (<a href="http://www.INGENUITY.com">http://www.INGENUITY.com</a>).

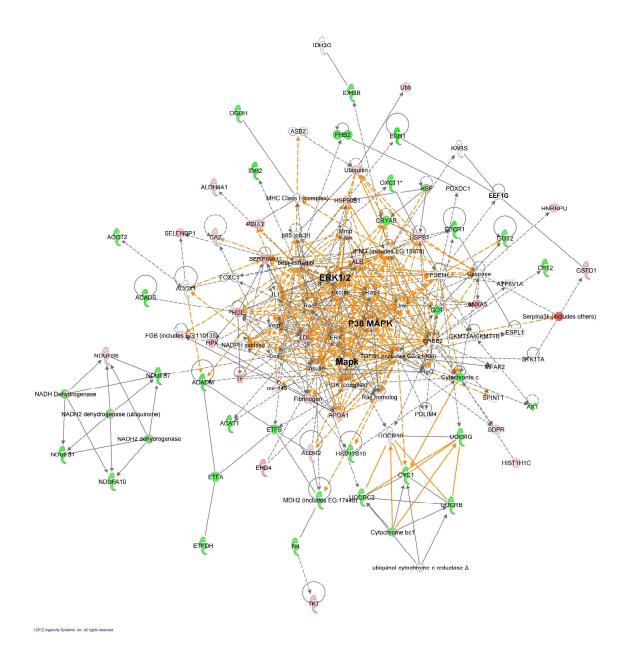


Figure S5. Graphical representation of the merged of the most significant protein interaction network after 16 Gy using Ingenuity Pathway Analysis of differentially regulated proteins. All significantly deregulated proteins identified by ICPL were imported into the Ingenuity Pathway Analysis. The up-regulated proteins are marked in red and the down-regulated in green. The nodes represent proteins that are connected with one or several arrows; the solid arrows represent direct interactions and the dotted arrows indirect interactions; direct and predicted protein interactions searched were for (http://www.INGENUITY.com).

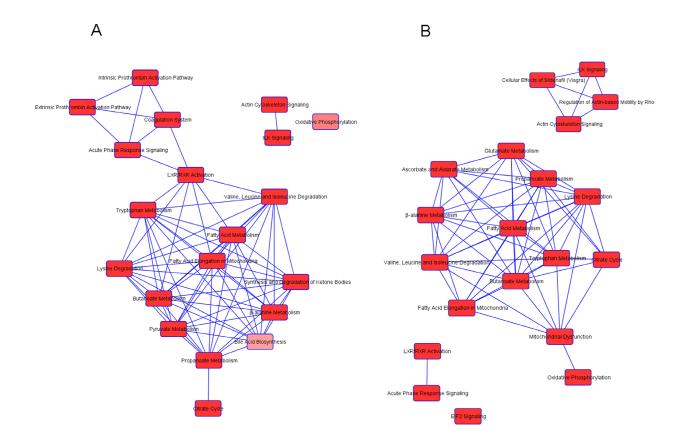
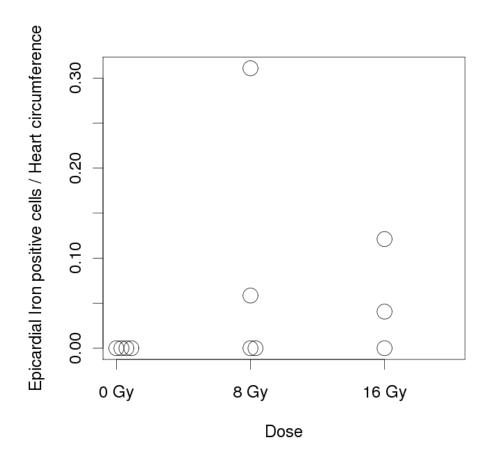


Figure S6. Graphical representation of 20 top canonical pathway networks representing significantly deregulated proteins after 8 Gy (A) and 16 Gy (B). Compiling of top 20 significant canonical pathways of the significantly deregulated proteins showed overlapping and interconnected pathways mainly involved in metabolic activity and cytoskeletal organization.



**Figure S7. Number of epicardial iron positive cells in control and irradiated hearts.** The number of iron positive cells were counted and normalized to heart area in irradiated and control mice. (Kruskal-Wallis rank sum test).