Supplementary Figures

Cisplatin induced neurotoxicity is mediated by Sarm1 and calpain activation

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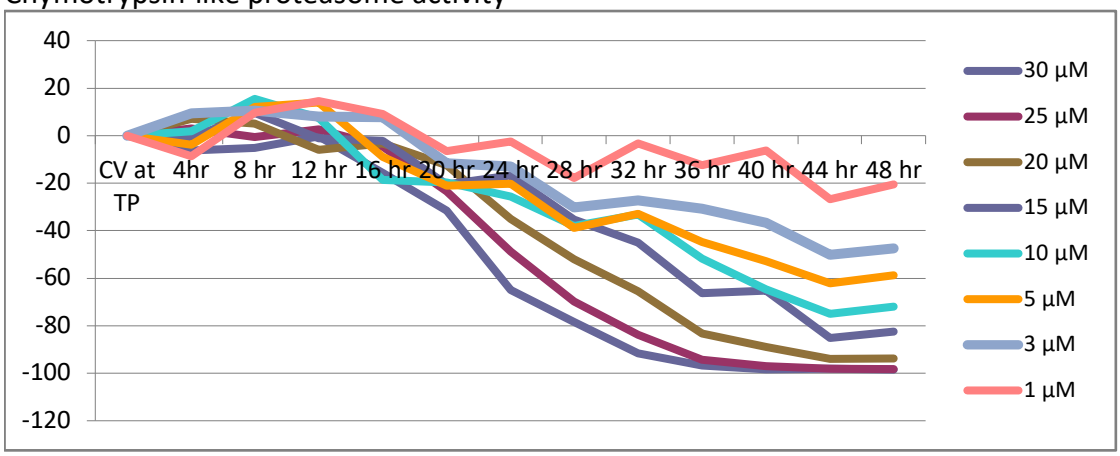
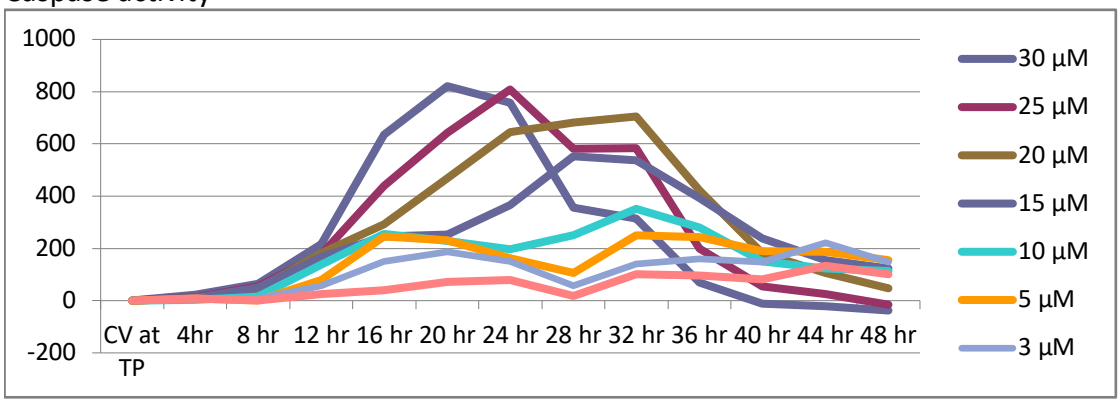
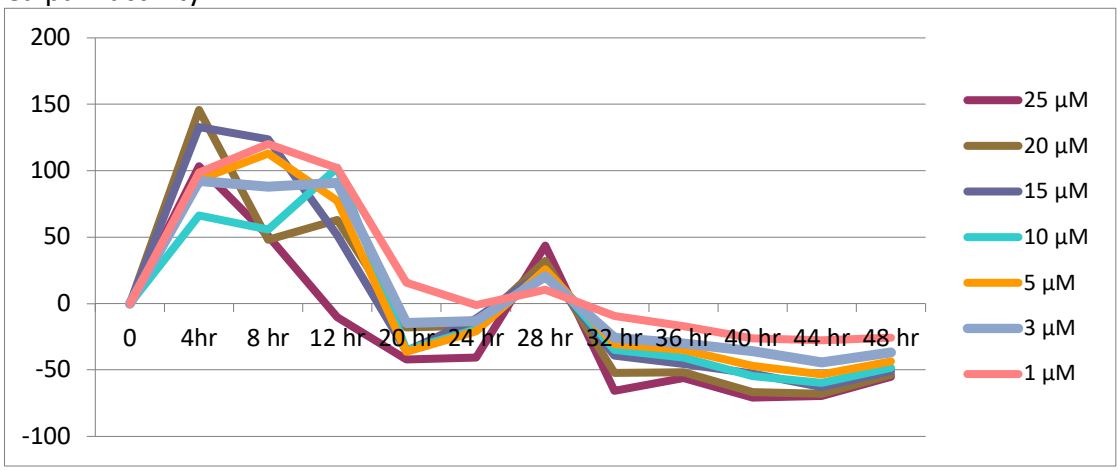
Chymotrypsin-like proteasome activity



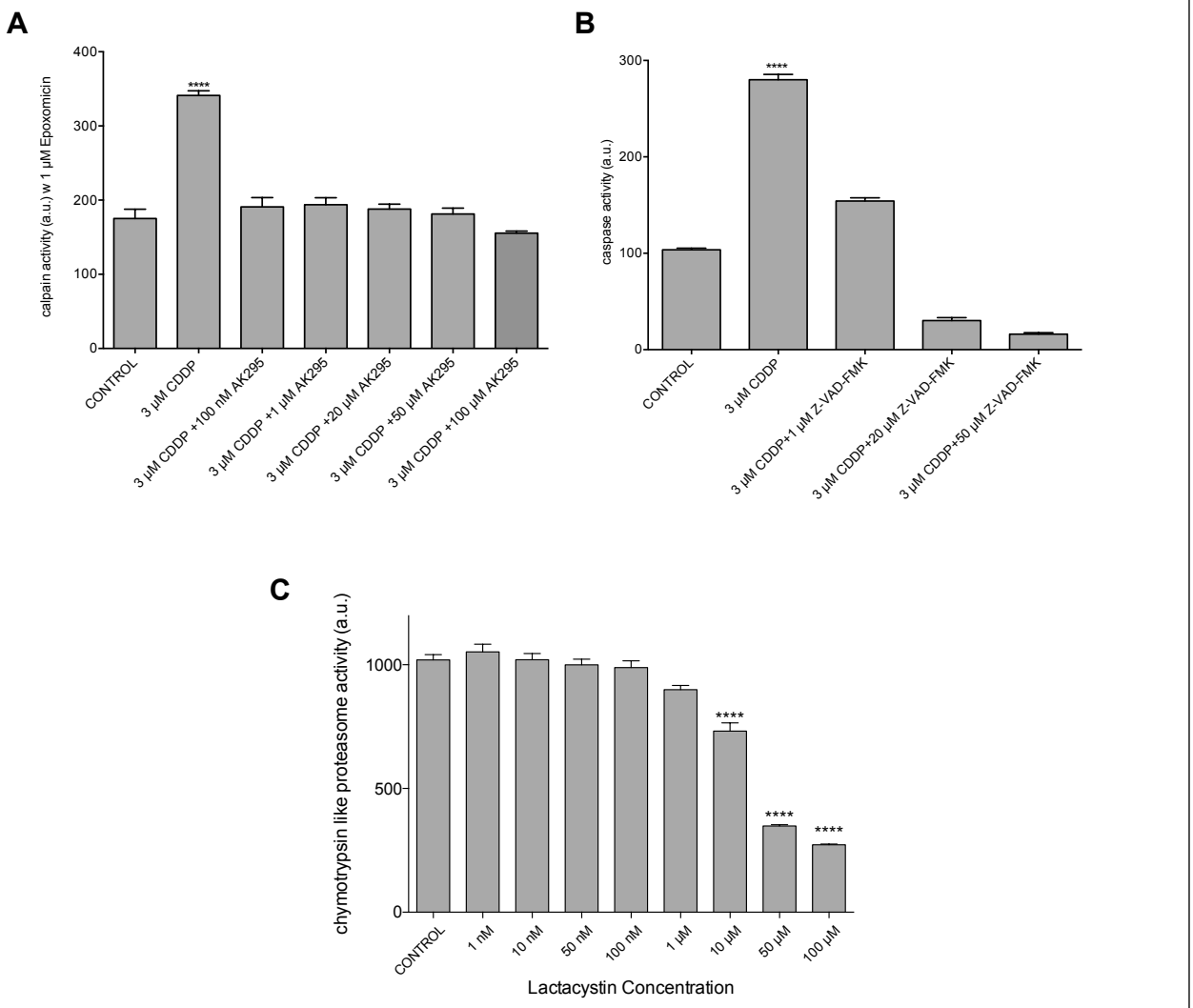
Caspase activity



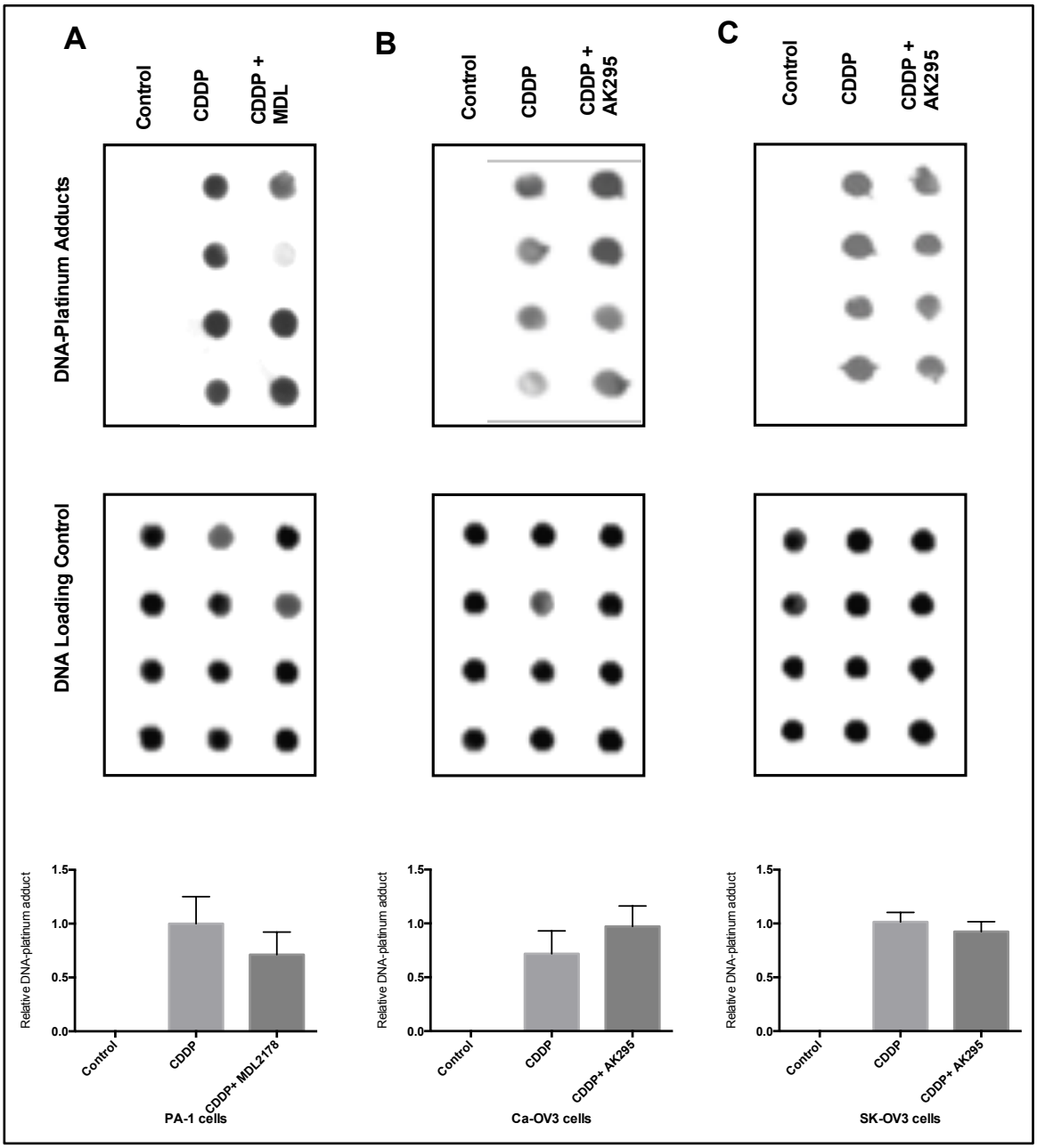
Calpain activity



Supplementary Figure 1: CDDP inhibits the UPS and activates Caspase 3/7 and Calpains in a dose dependent manner. (n=4 at each time point)



Supplementary Figure 2: The specificity of the Calpain-Glo and Caspase-Glo assays were evaluated with corresponding inhibitors (A and B) and inhibition of UPS by lactacystin was confirmed (C). Calpain activity was inhibited by AK295, a specific and highly potent inhibitor of calpain enzymatic activity (A). Similarly, caspase activity was inhibited by Z-VAD-fmk, a pan-caspase inhibitor (B). Lactacystin inhibited the UPS with a profile similar to that of CDDP.



Supplementary Figure 3: Calpain inhibitors MDL2178 and AK295 do not affect DNA-Platinum adduct formation in three different cancer cell lines.