



Figure 2 | Predictive modelling of pharmacological sensitivity using CCLE genomic data. **a, b**, Drug responses for panobinostat (green) and PLX4720 (orange/purple) represented by the high-concentration effect level (A_{\max}) and transitional concentration (EC_{50}) for a sigmoidal fit to the response curve (**b**). **c**, Elastic net regression modelling of genomic features that predict sensitivity to PD-0325901. The bottom curve indicates drug response, measured as the area over the dose-response curve (activity area), for each cell line. The central heat map shows the CCLE features in the model (continuous z-score for expression and copy number, dark red for discrete mutation calls), across all cell lines (x axis). Bar plot (left): weight of the top predictive features for sensitivity (bottom) or insensitivity (top). Parentheses indicate features present in >80% of models after bootstrapping. LOF, loss of function mutation; nnMS, non-neutral missense mutation (Supplementary Methods).

d, Specificity and sensitivity (receiver operating characteristic curves) of cross-validated categorical models predicting the response to a MEK inhibitor, PD-0325901 (activity area). Mean true positive rate and standard deviation ($n = 5$) are shown when models are built using all lines (global categorical model, in blue and orange), or within only melanoma lines (green). **e**, Activity area values for panobinostat between cell lines derived from haematopoietic ($n = 61$) and solid tumours ($n = 387$). The middle bar, median; box, inter-quartile range; bars extend to $1.5\times$ the inter-quartile range. **f**, Distribution of activity area values for AEW541 relative to *IGF1* mRNA expression. Orange dots, multiple myeloma cell lines ($n = 14$); blue dots, cell lines from other tumour types ($n = 434$). Box-and-whisker plots show the activity area or mRNA expression distributions relative to each cell line type (line, median; box, inter-quartile range), with bars extending to $1.5\times$ the inter-quartile range.