

Figure S6. Performance of cancer recurrence and MRD detection using the simulation data.

(a-d) Zoom-in of Figure 3 (a-d). The area under the ROC curve (AUC) of the MRD/recurrence detection on (a) the validation dataset and (c) the independent dataset with different tumor fractions and sequencing coverage. The sensitivity and specificity with different tumor fractions and sequencing coverage on (b) the validation dataset and (d) the independent dataset. (e) AUCs of MRD/recurrence detection with and without error suppression (ES) on the independent dataset at 200x depth with different tumor fractions. (f) The sensitivity and specificity of MRD/recurrence detection with and without error suppression on the independent dataset at 200x depth with different tumor fractions. In (a), (c) and (e), the dots indicated the averaged AUC, and the vertical bars indicated average  $\pm$  SD of the AUC (see Methods). In (b), (d) and (f), the dots show the averaged sensitivity using a cutoff p-value = 0.05 of the background noise distribution; the vertical bars indicated average  $\pm$  SD of the sensitivity; the specificity was shown in the legend in the format of (averaged specificity, (average - SD, average - SD)). The solid lines show the smoothed performance fitted with logit functions.