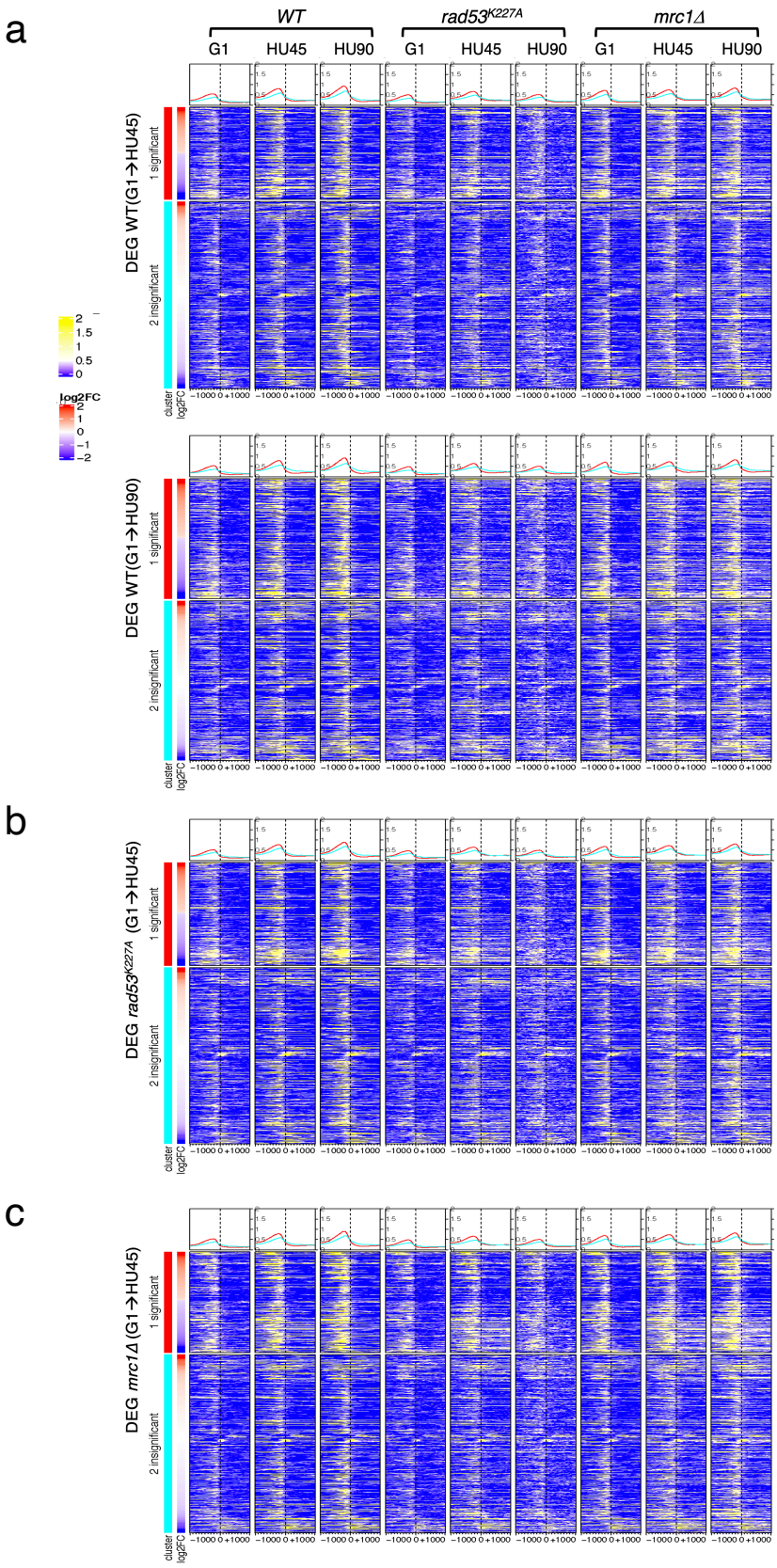Figure 4 – figure supplement 1