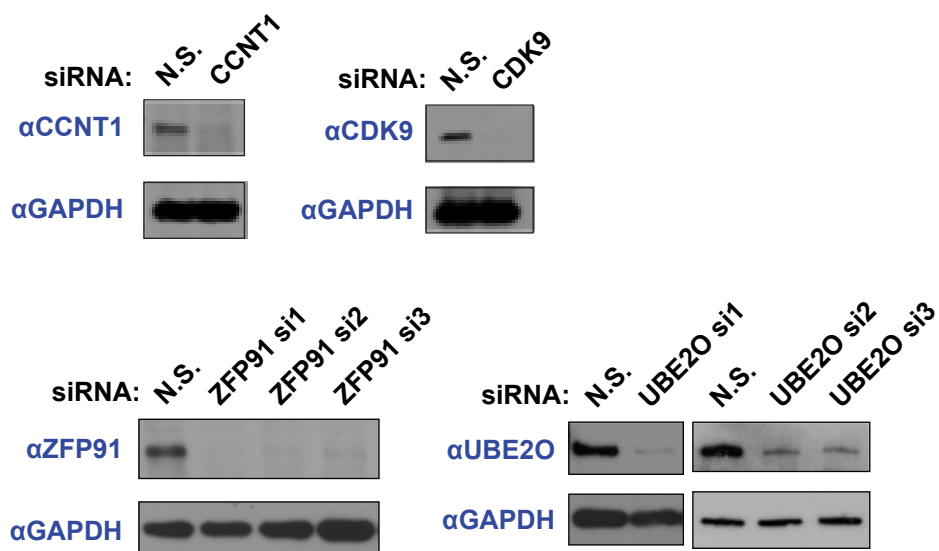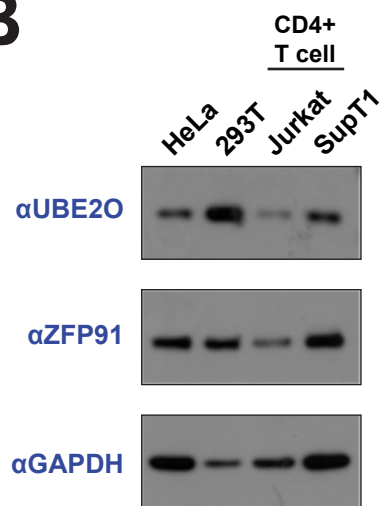Figure 1-figure supplement 1