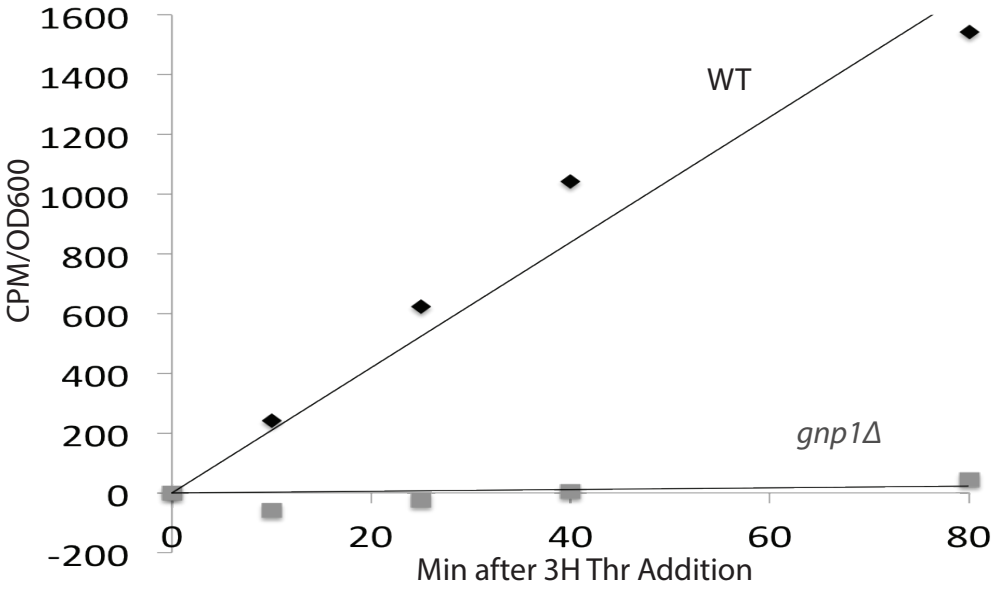**Figure 3-figure supplement 1**