

**Table S2.**

<b>Protein expression quantiles</b>	<b>q1 (least expressed)</b>	<b>q2</b>	<b>q3</b>	<b>q4 (most expressed)</b>
<b>Number of genes with identified rare codon cluster</b>	90	67	99	345

**Table S2. Genes with identified rare codon clusters are not disproportionately sampled from lowly expressed genes.** Could it be that large changes in ribosomal occupancy are not observed after rare clusters (Fig. 2A, Fig. 3A) because the clusters we identify are more likely to come from lowly expressed genes, i.e. genes which do not have high translation levels and for which it may be less likely that ribosomal footprints will be sampled? We used the average footprint count of a gene (total number of footprints within the coding sequence divided by gene length) as a proxy for protein expression levels. If anything, there are more genes with non-optimal codon clusters from genes which have more footprint reads ( $\chi^2$ ,  $P < 2.2e-16$ ) so we do not consider this an issue.