

# Differences of Translation Termination Sites Among the Three Stop Codons

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## 1 Introduction

Most organisms use different mechanisms of translation termination for the three types of stop codons, i.e. UAA, UAG, and UGA [1]. Using the GenBank database, we have conducted comprehensive computer analyses of translation termination sites in order to find the difference of consensus sequence patterns around each type of stop codons. We also systematically analyzed the preference of stop codons (O/E ratio: frequency of UAA, UAG, or UGA used as stop codons divided by frequency of the same triplet over the entire genome). It is known that Release Factor 1 (RF1) is required for translation termination for the stop codon UAG, and Release Factor 2 (RF2) for the stop codon UGA. The stop codon UAA requires either RF1 or RF2, but not necessarily both. For the five species of which genes for RF1 and RF2 are annotated in the GenBank database, we computed Codon Adaptation Index (CAI) values [2] for these genes.

## 2 Results and Discussion

Tables 1 and 2 show the results of our analysis. For the stop codon UAA, patterns AAA and GAA are frequently observed at position  $-3$  (from the stop codon). For the stop codon UGA, on the other hand, pattern GAG is often observed at position  $-12$ . Consensus patterns observed for the stop codon UAG are not as strong as the other two stop codons.

Table 3 shows that the preference of each stop codon is correlated with the CAI value of its corresponding Release Factor gene; if O/E ratio of stop codon is high, the CAI value of the corresponding release factor gene is also high. Greater CAI values usually imply higher expression rates. We therefore conclude that many organisms preferably use the type of stop codon which corresponds to the most abundant Release Factor.

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species	cod.	pat.	pos.	Z val.
<i>A.fulgidus</i>	uga	gag	-12	15.20
<i>A.aeolicus</i>	uga	gag	-12	12.87
<i>B.subtilus</i>	uga	gag	-12	14.99
<i>E.coli</i>	uga	gag	-12	19.71
<i>M.tuberculosis</i>	uga	gag	-12	13.67
<i>P.horikoshii</i>	uga	gag	-12	14.71
<i>T.maritima</i>	uga	gag	-12	15.76
<i>M.genitalium</i>	uag	gaa	-12	7.19
<i>H.pylori</i>	uga	gaa	-9	12.35
<i>A.fulgidus</i>	uga	gtg	-9	18.35
<i>M.jannaschii</i>	uga	gtg	-9	11.38
<i>T.maritima</i>	uga	gtg	-9	18.86
<i>A.fulgidus</i>	uga	cta	-3	11.07
<i>H.influenzae</i>	uga	cta	-3	10.58
<i>E.coli</i>	uaa	gag	-3	17.21
<i>M.jannaschii</i>	uaa	aaa	-3	16.03
<i>E.coli</i>	uaa	aaa	-3	21.65
<i>H.influenza</i>	uaa	aaa	-3	24.17
<i>H.influenzae</i>	uaa	gaa	-3	16.58
<i>B.subtilus</i>	uaa	gaa	-3	17.06
<i>M.jannaschii</i>	uaa	gaa	-3	24.05
<i>E.coli</i>	uag	gag	-3	8.15
<i>M.pneumoniae</i>	uag	aaa	-3	9.05
<i>H.influenza</i>	uag	aaa	-3	8.91

Table 1: Frequent triplets before stop codons

species	cod.	pat.	pos.	Z val.
<i>A.aeolicus</i>	uaa	ttatg	+12	11.24
<i>M.jannaschii</i>	uaa	ttttt	+10	11.97
<i>A.fulgidus</i>	uaa	ttaaa	+7	18.89
<i>M.genitalium</i>	uag	tgcga	+6	20.48
<i>B.subtilus</i>	uaa	aaaaa	+5	23.87
<i>A.aeolicus</i>	uaa	gaggt	+5	14.55
<i>A.fulgidus</i>	uaa	tttta	+5	26.64
<i>P.horikoshii</i>	uaa	ctttt	+4	10.78
<i>T.maritima</i>	uaa	gaggt	+4	13.78
<i>A.aeolicus</i>	uaa	ggagg	+4	21.42
<i>A.fulgidus</i>	uaa	ttttt	+4	38.44
<i>M.jannaschii</i>	uag	ggtga	+4	22.19
<i>A.fulgidus</i>	uga	gttgt	+4	27.10
<i>P.horikoshii</i>	uga	gttat	+4	30.86
<i>B.subtilus</i>	uaa	aaaag	+3	19.44
<i>A.aeolicus</i>	uaa	aataa	+3	10.75
<i>B.subtilus</i>	uaa	aggag	+3	13.56
<i>A.aeolicus</i>	uaa	aggag	+3	12.66
<i>A.fulgidus</i>	uaa	atttt	+3	12.65
<i>H.influenzae</i>	uaa	ggaga	+3	13.63
<i>T.maritima</i>	uaa	ggagg	+3	16.72
<i>B.subtilus</i>	uaa	ggagg	+3	17.44
<i>E.coli</i>	uaa	ggagt	+3	14.47
<i>T.pallidum</i>	uaa	ggggt	+3	11.28
<i>M.jannaschii</i>	uaa	ggtga	+3	13.70
<i>M.jannaschii</i>	uag	gtgag	+3	16.56
<i>T.maritima</i>	uag	gaggt	+3	14.70
<i>B.subtilus</i>	uag	gaggt	+3	19.01
<i>M.pneumoniae</i>	uag	ttatg	+3	21.46
<i>T.maritima</i>	uga	ggggg	+3	31.68
<i>A.fulgidus</i>	uga	ggtgg	+3	29.10
<i>P.horikoshii</i>	uga	ggtga	+3	46.02

Table 2: Frequent pattern consist of five bases observed after UAA

Species	O/E-UAA (RF1 or RF2)	O/E-UGA (RF2)	O/E-UAG (RF1)	RF1	RF2
<i>A.aeolicus</i>	1.10	1.22	0.51	0.719	0.745
<i>B.burgdorferi</i>	1.10	0.78	0.97	0.707	0.625
<i>E.coli</i>	1.66	0.63	0.50	0.757	0.761
<i>H.pylori</i>	1.17	0.97	0.70	0.713	0.744
<i>T.pallidum</i>	0.90	0.74	1.68	0.742	0.711

Table 3: O/E ratio of stop codon and CAI value of the corresponding Release Factor

## References

- [1] Brown, C.M., Stockwell, P.A., Trotman, C.N. A., and Tate, W.P., The signal for the termination of protein synthesis in prokaryotes, *Nucleic Acids Res.*, 18:2079–2086, 1990.
- [2] Sharp, P.M. and Li, W.-H., The codon adaptation index – a measure of directional synonymous codon usage bias, and its potential applications, *Nucleic Acids Res.*, 15:1281–1295, 1987.