

# Python Implementation of Codon Adaptation Index

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## Software

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## Summary

Amino acids, the building blocks of proteins, are encoded in DNA by triplets of nucleotides called codons. Notably, the synonymous codons for an amino acid are not used in equal proportions in coding DNA sequences. Rather, they exhibit a bias which varies from organism to organism. The codon adaptation index (CAI) is a measurement of this bias with respect to a set of reference genes (Sharp & Li, 1987). It has been used in the context of heterologous protein expression (Grote et al., 2005), virus attenuation (Eschke, Trimpert, Osterrieder, & Kunec, 2018), and cotranslational protein folding prediction (Rodriguez, Wright, Emrich, & Clark, 2017).

CAI is a Python package for the efficient calculation of this metric, along with the associated relative synonymous codon usage (RSCU) and relative adaptiveness metrics. In addition, CAI includes a command line interface for the calculation of CAI from FASTA files containing DNA sequences. For example, to find the CAI of the sequence within `sequence.fasta` with respect to the sequences within `reference.fasta`, one need only run:

```
$ CAI -s sequence.fasta -r reference.fasta
0.24948128951724224
```

Similarly, using the Python API:

```
>>> from CAI import CAI
>>> from Bio import SeqIO # to parse FASTA files
>>> reference = [seq.seq for seq in SeqIO.parse("reference.fasta", "fasta")]
>>> sequence = SeqIO.read("sequence.fasta", "fasta")
>>> CAI(sequence, reference=reference)
0.24948128951724224
```

In comparison to other Python implementations of the CAI metric (Cock et al., 2009), CAI features a CLI, supports multiple genetic codes, can yield the RSCU of reference sets, and correctly handles the case of missing codons in the reference set. Moreover, on a benchmark to determine the CAI of 100 genes consisting of 3,000 random base pairs each with respect to highly expressed genes in *Escherichia coli*, CAI performed 39.6% faster than Biopython's implementation.

In conclusion, CAI is a significantly faster and more versatile method to determine the CAI, RSCU, and relative adaptiveness of DNA sequences.

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