**TABLE S7.** Summary of the sensitivity of various restriction endonucleases to DNA modifications introduced by the JCM7686\_0815 protein (m<sup>4</sup>C MTase).

REase	Recognition site	Sensitivity to m⁴C	Number of sites in pET30_JCM7686_0815	Cleavage
Cfrl	YGGCCR	U	5	P, L
HaellI	GGC <u>C</u>	S	26	P*
Hin6l	GCGC	U	53	Υ
Hpall	<u>CC</u> GG	S	34	Υ
Mspl	<b>C</b> cGG	S	34	Υ

<sup>\*</sup> the larger fused DNA fragments were observed only in case of incomplete cleavage of BsuRI sites that overlapped CfrI target sequences (YGGCCR).

Y – complete cleavage; P – partial digestion; L – partial plasmid cleavage; S – sensitive to  $m^4C$ ; U – unknown sensitivity to  $m^4C$ ;  $\underline{\mathbf{C}}$  – the enzyme will not cleave if the  $\underline{\mathbf{marked}}$  cytosine is methylated; c – the enzyme is not sensitive to methylation of the marked cytosine; C – there are no data on the sensitivity of the enzyme to methylation of the unmarked cytosine.