## Additional file 5 - Comparison with other bacterial promoter prediction approaches

<u>E. coli</u>

Name	Sensitivity	FP/100 nt	Promoters	Regions	Spacing	g Method	Comment	Reference
Genomic Distribution	97.1% 42.4% 28.1%	9.26 1.13 0.59	377	335	16-20	Organism-specialized matrix	See discussion.	This work
	31.0%	1.09				General matrix		
Statistical over- representation generating PSWM	N/D	N/D	N/D	~2500	0-30	Words from 3 to 5 nucleotides were analyzed in the first 300 nt of IRs located upstream of all operons.Statistically significant dimmers with fixed-length spacers were grouped into clusters of related sequences. Putative functions were assigned to these sequence clusters by examining their location in the genome.	No sigma70 consensus sequence was identified. This was attributed to the greater variability of promoters in this organism.	[32]
PSWM	100%	15.10				PSWM search limited on 250 nt of each region, cutoff mean- $3\sigma$ .	Any approach based on PSWMs depends on the availability of an	
PSWM Cover	47.7% 86.0%	0.87 1.88	599	470	13-24	PSWM search limited on 250 nt of each region, cutoff mean- $0.5\sigma$ . PSWM with several constraints such as grouping signals and filtering with the distance from the start codon	extensive experimental dataset to create a representative description of the As the authors noted, a distance filter is "hard to reconcile with a direct computational modeling of RNAP binding and transcription initiation"	[24]
MITRA	N/D	N/D	N/D	589	3-23	The significance of hexanucleotide pairs in the first 310 nt of divergent IRs was estimated by comparaison with those of convergent IRs considering three type of information: the strength score, the dyad score, and the positional score.	No statistically strong signal corresponding to a principal sigma factor- dependent promoter sequence was identified in this organism.	[30]
NNPP2.2	31.0%	0.62	671	510	N/D	Neural network trained with 272 <i>E. coli</i> promoters. Promoter searches were limited to the first 500 nt of every tested regions. Default cutoff was set to 0.8 and a shift of $\pm$ 3 nt was tolerated with respect to the location of the experimentally identified promoter.		[78]
TLS-NNPP2.2	30.1%	0.22	0,1	010	102	Distance filter combined to NNPP2.2. Cutoff set to 0.0269.	See comment on the use of a distance filter in the "Cover" approach.	[39]
НММ	N/D	N/D	N/A	N/A	N/A	Same methodology as for <i>C. jejuni</i> (see below).	A TATA-box of varying intensity was identified but no periodic signal could be seen in sequence logos of predicted promoters aligned by the model.	[27]
SVM	N/A	N/A	450	450	N/D	A kernel with strings of length 5 and 1 mismatch trained with sets of 200 nt each.	The results presentation do not allowed sensitivity and FP rate extraction.	[28]
DNA stability	32.0%	1.20	227	227	N/D	The free energy of a 15 nt moving window is calculated on regions of 1000 nt each. If two positive signals (higher than thresholds) are within 25 nt of each other, they were considered as 1 segment. A TP is a segment that overlap the 200 nt region spanning from -150 to +50 from the characterized TSS.	As the authors noted, this "method tries to find a promoter region" rather than identifying precisely promoter boxes.	[18]

B. subtilis

Name	Sensitivity	FP/100 nt	Promoters	Regions	Spacing	Method	Comment	Reference
Genomic Distribution	100% 84.5% 56.8% 50.0%	8.42 4.29 0.99 0.93	148	142	16-20	Organism-specialized matrix General matrix	See discussion.	This work
Statistical over- representation generating PSWM	82.5%	*0.2%	132	2729	3-30	Methodology based on [32]. Words of 4 and 5 nts. Their PSWM correspondig to consensus promoter sequence (WM1= $N_7$ TTGAN <sub>19</sub> TATAATAN <sub>6</sub> ) predicted 1141 promoter sequences, including 109 of the 132 containng a spacing of 17.	*The FP rate was estimated from an expected normal distribution of the data. No actual count of false predictions. Both sensitivity and FP rate would have to be considered for fair comparison.	[31]
MITRA	N/D	N/D	N/D	552	3-23	The significance of hexanucleotide pairs in the first 310 nt of divergent IRs was estimated by comparaison with those of convergent IRs considering three type of information: the strength score, the dyad score, and the positional score.	The identified consensus sequence was not used to make predictions.	[30]
НММ	70.0%	*0.0	130	N/A	N/D	Trained with 100 nt sequences from approximately -85 to $+15$ relative to the transcription start site.	*The FP rate was estimated from 1000 random sequences of 100 nt, with respect to the GC% of the genome. Similarly, almost no FPs are produced by our algorithm with the second version of shuffled genomes.	[26]
DNA stability	N/A	N/A	89	89	N/D	The free energy of a 15 nt moving window is calculated on regions of 1000 nt each. If two positive signals (higher than thresholds) are within 25 nt of each other, they were considered as 1 segment. A TP is a segment that overlap the 200 nt region spanning from $-150$ to $+50$ from the characterized TSS.	As the authors noted, this "method tries to find a promoter region" rather than identifying precisely promoter boxes.	[18]

<u>H. pylori</u>								
Name	Sensitivity	FP/100 nt	Promoters	Regions	Spacing	Method	Comment	Reference
	100%	6.98						
Genomic	70.6%	3.11	17	16	10.22	Organism-specialized matrix	San diamagin	This most
Distribution	47.1%	0.53	17	10	19-23		See discussion.	THIS WORK
	35.3%	0.70				General matrix		

						These authors used three datasets: a) both strands from all non-convergent IRs; b)	The most statistically significant motif was used to identified 56 putative	
Statistical over-	N/D	N/D	N/D	756, 340	8-10 to	divergent IRs only; c) IRs located upstream of ribosomal genes. They were used to	sigma 80 promoter sequences. However, none of these predictions	[20]
representation	N/D	N/D	IN/D	and 30	22-24	identify hexanucleotide pairs present with at most one mismatch in at least 10% of the	correspond to the 17 characterized promoters described in the literature	[29]
						sequences of the sets.	and used in our work.	
						The significance of hexanucleotide pairs in the first 310 nt of divergent IRs was		
MITRA	N/D	N/D	N/D	169	3-23	estimated by comparaison with those of convergent IRs considering three type of	The identified consensus sequence was not used to make predictions.	[30]
						information: the strength score, the dyad score, and the positional score.		

<u>C. jejuni</u>

Name	Sensitivity	FP/100 nt	Promoters	Regions	Spacing	Method	Comment	Reference
Genomic Distribution	100% 71.4% 42.9%	7.90 3.13 0.84	14	14	16-20	Organism-specialized matrix	See discussion.	This work
	35.7%	0.77				General matrix		
						Trained with 175 divergent IRs of 121 nt each. The authors proposed a consensus	*The FP rate was estimated on sequences generated randomly. Some of	
HMM	70.3%	*0.0	27	22	N/A	sequence composed of an AT-rich periodic signal upstream of a classical -10 box	the 27 promoters used for sensitivity evaluation have not been detected in	[27]
						(TATAAT sequence). No -35 box was formally identified.	the genome.	
						The significance of hexanucleotide pairs in the first 310 nt of divergent IRs was	No statistically strong signal company ding to a minainal signa factor	
MITRA	N/D	N/D	N/D	168	3-23	estimated by comparaison with those of convergent IRs considering three type of	dependent promoter sequence was identified in this organism	[30]
						information: the strength score, the dyad score, and the positional score.	dependent promoter sequence was identified in this organism.	

M. pneumoniae

Name	Sensitivity	FP/100 nt	Promoters	Regions	Spacing	g Method	Comment	Reference
Genomic	100% 76.7%	8.89 3.20	30	27	15-19	Organism-specialized matrix	See discussion.	This work
Distribution	43.3% 30.0%	1.08 1.07				General matrix		
MITRA	N/D	N/D	N/D	52	3-23	The significance of hexanucleotide pairs in the first 310 nt of divergent IRs was estimated by comparaison with those of convergent IRs considering three type of information: the strength score, the dyad score, and the positional score.	No statistically strong signal corresponding to a principal sigma factor- dependent promoter sequence was identified in this organism.	[30]

<u>S. aureus</u>								
Name	Sensitivity	FP/100 nt	Promoters	Regions	Spacing	y Method	Comment	Reference
	87.5%	5.85						
Genomic	62.5%	2.63	0	5	16 20	Organism-specialized matrix	See discussion	This work
Distribution	37.5%	0.59	8	5	10-20	0	See discussion.	THIS WOLK
	37.5%	37.5% 1.13 General matrix						
							A TATA-box of varying intensity but no periodic signal could be seen in	
HMM	N/D	N/D	N/A	N/A	N/A	Same methodology as for C. jejuni (see above).	sequence logos of predicted promoters aligned by the model (data not	[27]
							shown).	

N/D = Non-determined

N/A = Non-available

## **References**

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