

**Supplementary Table I:** Co-regulated adjacent TU pairs in *E. coli*

<i>Regulating TFs<sup>a</sup></i>	<i>TU1</i>	<i>TU2</i>	<i>Orientation<sup>b</sup></i>	<i>Mode of control on TU1<sup>c</sup></i>	<i>Mode of control on TU2<sup>c</sup></i>	<i>r<sub>s</sub><sup>d</sup></i>
CRP / caiF	caiTABCDE	fixABCX	Div	P / P	P / P	0.88
CRP / araC <sup>e</sup>	araBAD	araC	Div	P / D	P / N	0.88
arcA / betI <sup>e</sup>	betIBA	betT	Div	N / N	N / N	ND <sup>f</sup>
cynR <sup>e</sup>	cynR	cynTSX	Div	N	P	0.67
Fur	fepB	entCEBA-cstA	Div	N	N	0.75
CRP / nagC <sup>e</sup>	nagBACD	nagE	Div	P / N	P / N	0.8
CRP / arcA	gltA	sdhCDAB-b0725-sucABCD	Div	P / N	P / N	0.86
birA	bioA	bioBFCD	Div	N	N	0.76
csgD <sup>e</sup> / ompR	csgDEFG	csgBA	Div	P / P	P / P	0.84
FlhDC / fliA	flgAMN	flgBCDEFGHIJK	Div	P / P	P / P	0.85
CRP / mall <sup>e</sup>	mall	malXY	Div	N / N	D / N	ND <sup>f</sup>
FliA	fliC	fliDST	Div	P	P	0.69
FlhDC / fliA	fliE	fliFGHIJK	Div	P / P	P / P	0.89
CRP / glpR	glpTQ	glpACB	Div	P / N	P / N	0.8
dsdC <sup>e</sup>	dsdC	dsdXA	Div	N	P	0.51
fliA / ihfAB	hycABCDEFGH	hypABCDE	Div	P / P	P / P	0.77
CRP / fucR <sup>e</sup>	fucAO	fucPIKUR	Div	P / P	P / P	0.67
lysR <sup>e</sup>	lysA	lysR	Div	P	N	0.46
exuR	uxaC	exuT	Div	N	N	0.48
CRP <sup>e</sup>	yhfA	CRP	Div	D	D	ND <sup>f</sup>
asnC <sup>e</sup>	asnC	asnA	Div	N	P	0.38
ilvY <sup>e</sup>	illvY	ilvC	Div	N	P	-0.08
argR	argE	argCBH	Div	N	N	0.83
CRP / malT	malEFG	malK-lamB-malM	Div	P / P	P / P	0.64
lexA	uvrA	Ssb	Div	N	N	0.38
soxR <sup>e</sup>	soxS	soxR	Div	P	N	0.57
CRP/melR <sup>e</sup>	melR	melAB	Div	P/N	P/P	0.91
cpxR <sup>e</sup>	cpxR	cpxP	Div	P	P	0.04
pspF <sup>e</sup>	pspF	pspABCDE	Div	N	P	0.38
rhaS <sup>e</sup>	rhaBAD	rhaSR	Div	P	N	0.77
torR <sup>e</sup>	torR	torCAD	Div	N	P	ND <sup>f</sup>
xylR <sup>e</sup>	xylAB	xylFGHR	Div	P	P	0.29
Rob	ybaO	mdlA	Uni	P	P	0.43
ihfAB	glnHPQ-ybiO	dps	Uni	P	P	0.04
appY / arcA	hyaABCDEF	appCBA	Uni	P/P	P/P	ND <sup>f</sup>
Fnr/ ihfAB / narL	narK	narGHJI	Uni	P/P/P	P/P/P	0.89
arcA	fumC	fumA	Uni	N	N	ND <sup>f</sup>
fliA	tar	motAB-cheAW	Uni	P	P	0.75
FlhDC / fliA	fliFGHIJK	fliLMNOPQR	Uni	P/P	P/P	0.92

CRP / galS <sup>e</sup>	mglBAC	galS	Uni	P/N	P/N	0.53
CRP	caiF	caiTABCD	Con	P	P	0.25
CRP	nupG	speC	Con	P	N	0.57
CRP	glpD	glgCAP	Con	P	P	0.77
rhaS <sup>e</sup>	rhaSR	rhaT	Con	N	P	0.63

<sup>a</sup>If more than one TF regulates the same adjacent TU pair, all common TFs are indicated, separated by ‘/’

<sup>b</sup>Orientation of TU1 and TU2 relative to each other. Div=divergent, Con=convergent and Uni=unidirectional

<sup>c</sup>P=positive regulation, N=negative regulation, D=dual regulation. If more than one TF regulates the same TU pair the type of regulation is indicated for each of the TFs separated by ‘/’

<sup>d</sup>Spearman correlation between the expression profiles of TU1 and TU2

<sup>e</sup>Co-regulation in *cis*

<sup>f</sup>Expression data missing