

Supplementary Table 5. Genes predicted under selection in *E. coli* by selective signatures and the MK test

COG	log ₂ v	log ₂ FI	dN/dS	Func	Description
112	2.13 **	2.47 *	0.12	E	serine hydroxymethyltransferase
118	2.02 **	0.82	0.07	E	imidazole glycerol phosphate synthase subunit HisH
150	1.96 **	0.69	0.03	F	phosphoribosylaminoimidazole synthetase
2973	1.86 *	0.41	0.00	K	Trp operon repressor
157	1.86 **	0.68	0.02	H	nicotinate-nucleotide pyrophosphorylase
2980	1.73 **	0.43	0.07	M	minor lipoprotein
704	1.54 *	0.84	3.21	P	negative regulator of PhoR/PhoB two-component regulator
357	1.51 **	0.61	0.02	M	glucose-inhibited division protein B
99	1.48 *	0.74	0.24	J	30S ribosomal protein S13
649	1.48 *	1.87 *	0.02	C	NADH:ubiquinone oxidoreductase, chain C,D
196	1.33 **	0.54	0.03	H	hypothetical protein
414	1.30 *	2.02 *	0.04	H	pantoate--beta-alanine ligase
2109	1.28 *	0.50	0.00	H	cob(I)yrinic acid a,c-diamide adenosyltransferase
3123	1.28 *	0.94	0.02	S	hypothetical protein
1214	1.27 **	0.94	0.05	O	predicted peptidase
3159	1.21 *	0.90	0.09	S	hypothetical protein
850	1.19 **	0.37	0.06	D	septum formation inhibitor
221	1.16	1.03	0.09	C	inorganic pyrophosphatase
2861	1.15 *	0.85	0.63	S	orf, hypothetical protein
1983	1.12 *	0.93	0.06	KT	DNA-binding transcriptional activator
529	1.11	0.55	0.05	P	adenylylsulfate kinase
149	1.07 *	2.08	0.08	G	triosephosphate isomerase
205	1.05 *	0.89	0.03	G	6-phosphofructokinase
3076	1.03	0.30	0.02	S	hypothetical protein
229	0.97 *	0.67	0.02	O	methionine sulfoxide reductase B
540	0.96 *	1.68	0.02	F	aspartate carbamoyltransferase catalytic subunit
1005	0.95	1.82 *	0.16	C	NADH dehydrogenase subunit H
751	0.88 *	2.09 *	0.05	J	glycyl-tRNA synthetase subunit beta
17	0.86	1.00	0.05	J	asparaginyl-tRNA synthetase
481	0.85	0.94	0.02	M	GTP-binding protein LepA
1666	0.84	2.52 *	0.07	S	orf, hypothetical protein
344	0.82 *	1.74	0.08	S	hypothetical protein
682	0.80 *	1.93 *	2.50	M	prolipoprotein diacylglycerol transferase
2377	0.80 **	0.37	0.01	O	anhydro-N-acetylmuramic acid kinase
554	0.74	0.27	0.03	C	glycerol kinase
151	0.73	0.29	4.33	F	phosphoribosylamine--glycine ligase
233	0.73	0.95	0.07	J	ribosome releasing factor
249	0.69 *	1.51 *	0.02	L	DNA mismatch repair protein
66	0.69	0.32	0.08	E	isopropylmalate isomerase small subunit
212	0.68 *	0.61	0.01	H	putative ligase
1207	0.64 *	0.91	0.06	M	bifunctional N-acetylglucosamine-1-phosphate uridyltransferase/glucosamine-1-phosphate acetyltransferase
2908	0.64 *	0.44	0.06	S	UDP-2,3-diacylglicosamine hydrolase
165	0.63	0.37	0.09	E	argininosuccinate lyase
468	0.63	0.67	0.01	L	recombinase A
3004	0.62	0.66	0.00	P	pH-dependent sodium/proton antiporter
2360	0.62 *	0.56	0.04	O	leucyl/phenylalanyl-tRNA--protein transferase
1485	0.60 *	0.37	0.04	R	conserved protein with nucleoside triphosphate hydrolase domain N-(5'-phospho-L-ribosyl-formimino)-5-amino-1-(5'- phosphoribosyl)-4-imidazolecarboxamide isomerase
106	0.58	0.32	0.02	E	
323	0.58 *	0.60	0.02	L	DNA mismatch repair protein
418	0.58 *	0.30	0.02	F	dihydroorotase
629	0.56	1.41	0.06	L	single-strand DNA-binding protein

2968	0.52	1.34	0.02	S	hypothetical protein
519	-0.51	*	-1.10	0.01	F bifunctional GMP synthase/glutamine amidotransferase protein
2917	-0.52		-1.13	0.01	D intracellular septation protein A
2925	-0.52	*	-1.45	*	0.01 L exonuclease I
262	-0.57	*	-0.91	0.77	H dihydrofolate reductase
1188	-0.59		-1.46	2.06	J ribosome-associated heat shock protein Hsp15
275	-0.65	*	-0.98	0.01	M S-adenosyl-methyltransferase
1314	-0.71	*	-0.87	0.09	U protein-export membrane protein
4121	-0.73	*	-0.76	*	0.08 S putative peptidase
324	-0.74		-1.90	*	0.05 J tRNA delta(2)-isopentenylpyrophosphate transferase
2976	-0.76	*	-1.58	0.04	S hypothetical protein
93	-0.81		-2.32	0.15	J 50S ribosomal protein L14
1212	-0.83	*	-1.61	*	0.00 M 3-deoxy-manno-octulosonate cytidyllyltransferase
777	-0.99	*	-2.35	*	0.05 I acetyl-CoA carboxylase subunit beta
774	-1.04	*	-1.26	0.04	M UDP-3-O-[3-hydroxymyristoyl] N-acetylglucosamine deacetylase
5633	-1.05	*	-2.65	*	0.08 R hypothetical protein
2190	-1.11	*	-0.99	0.03	G glucose-specific PTS system enzyme IIA component
4657	-1.13	*	-2.93	*	0.03 C Na(+)-translocating NADH-quinone reductase subunit E
360	-1.15	*	-3.00	*	0.06 J 30S ribosomal protein S6
563	-1.25	*	-0.85	0.10	F adenylate kinase
466	-1.28	*	-3.55	*	0.01 O DNA-binding ATP-dependent protease La
3151	-1.37	*	-2.46	0.02	S predicted dehydrogenase
4785	-1.37	**	-0.99	0.08	R hypothetical protein
101	-1.41	*	-1.14	1.40	J tRNA pseudouridine synthase A
821	-1.50	*	-0.81	0.01	I 4-hydroxy-3-methylbut-2-en-1-yl diphosphate synthase
2983	-1.52	*	-1.84	*	0.00 S orf, hypothetical protein
509	-1.58	*	-2.62	*	0.01 E glycine cleavage system protein H
287	-1.58	*	-0.88	1.62	E fused chorismate mutase T/prephenate dehydrogenase
788	-1.63	*	-1.34	1.28	F formyltetrahydrofolate deformylase
634	-1.69	*	-1.68	0.02	F hypoxanthine phosphoribosyltransferase
283	-1.69	*	-1.75	0.01	F cytidylate kinase
131	-1.75	*	-1.71	*	0.02 E imidazoleglycerolphosphate dehydratase and histidinol-phosphate phosphatase
452	-2.01	*	-1.05	0.03	H flavoprotein affecting synthesis of DNA and pantothenate metabolism
41	-2.43	*	-1.23	0.04	F phosphoribosylaminoimidazole carboxylase catalytic subunit
3317	-2.51	*	-1.52	*	0.03 M lipoprotein-34
575	-2.60	*	-1.77	*	1.38 I CDP-diglyceride synthetase
779	-2.62	*	-1.15	0.03	S orf, hypothetical protein
539	-2.71	*	-1.32	0.02	J 30S ribosomal protein S1
3117	-2.77	*	-1.72	*	0.00 S hypothetical protein
103	-2.78	*	-1.42	0.06	J 30S ribosomal protein S9
55	-2.86	*	-1.16	0.01	C F0F1 ATP synthase subunit beta
1385	-2.89	*	-1.68	*	1.34 S orf, hypothetical protein

Legend. Genes are listed only when the MK test and selective signatures agree, using the same criteria as Figure 7 in the main text ($\log_2\text{FI} > 0.26$ and $\log_2v > 0.5$, or $\log_2\text{FI} < -0.74$ and $\log_2v < -0.5$). For v, * denotes Z score > 1 , ** Z > 2 . For FI, * denotes $\chi^2 > 3.84$ ($P < 0.05$ with 1 d.f.).

COG functional category key:

C	Energy Production	K	Transcription
D	Cell Division	L	DNA replication, repair & modification
E	Amino Acid Metabolism	M	Cell Envelope
F	Nucleic Acid Metabolism	N	Motility & Secretion
G	Carbohydrate Metabolism	O	Protein Modification & degradation
H	Coenzyme Metabolism	P	Ion Transport & Metabolism
I	Lipid Metabolism	Q	Secondary Metabolism
J	Ribosome & Translation	R	General Function
S	Unknown	T	Signal Transduction