Supplementary Materials for:

Glucose Levels Alter the Mga Virulence Regulon in the Group A Streptococcus

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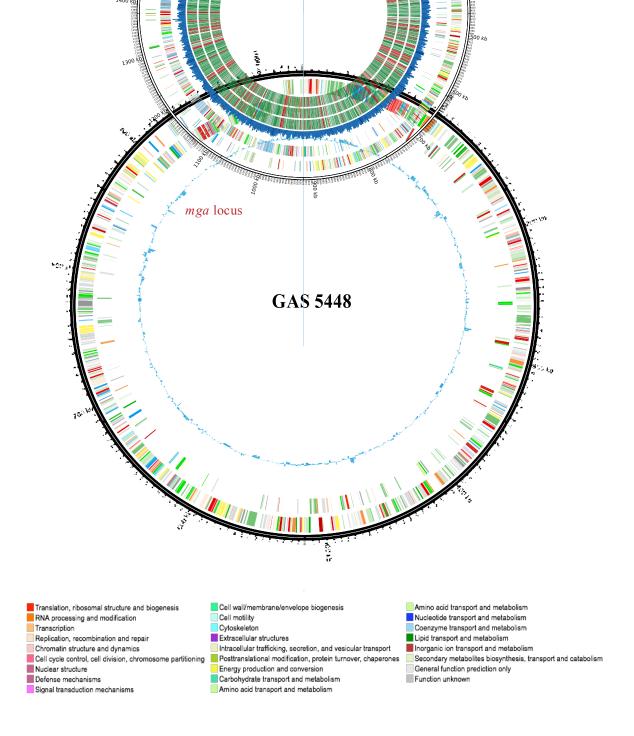


Figure S1. Transcriptomic landscape of M1T1 5448 during growth in C media compared to THY. Circos plot of GAS 5448 differentially expressed (DE) genes during growth in C media (low glucose) versus THY (high glucose) at late logarithmic growth. The outer most ring represents a size ruler and the next two rings represent the GAS open reading frames on the (+) and (-) strand of the genome, respectively, with color representing function COG category as indicted in legend. The inner ring (blue) shows transcript levels of genes, where the height of the bars represent the log₂ fold-change in expression in C media vs. THY (inside is down, outside is up). The position of the *mga* locus is indicated.

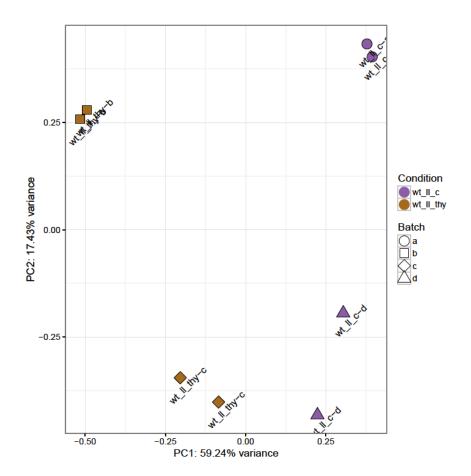
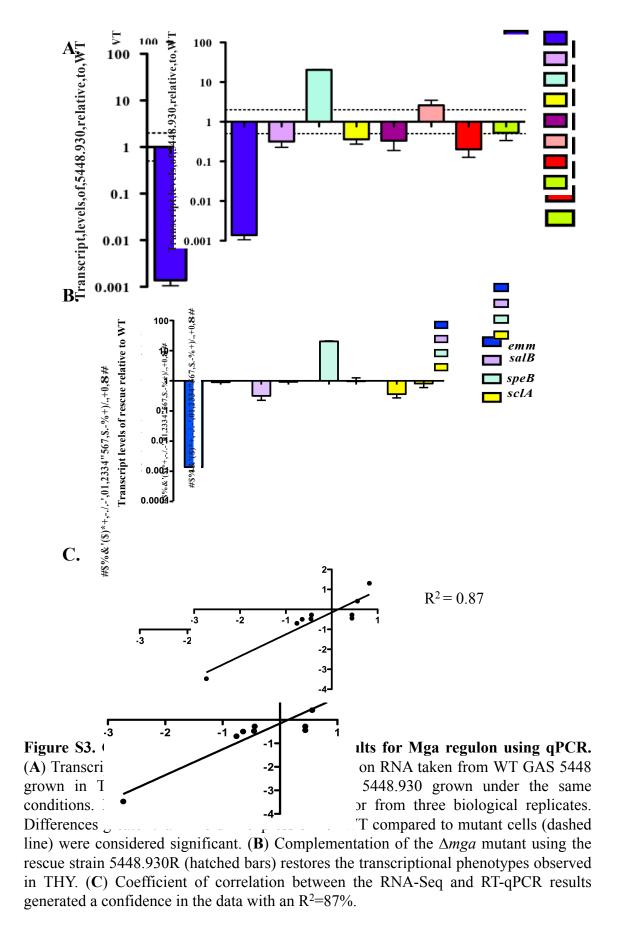


Figure S2. Clustering of WT 5448 samples based on RNA-seq data. A principal component analysis (PCA) plot is shown for the log₂ fold change data of WT 5448 grown in THY (high glucose) or C media (low glucose) at late logarithmic growth phase prior to normalization for batch effects. The PCA plots show the first two principal components on the X and Y axes, respectively, and the percentage of total variance specific to that principle component is indicated. Each biological replicate is represented as a single point with color corresponding to the media condition (THY, brown; C media, purple) and shapes represent different experimental batches.



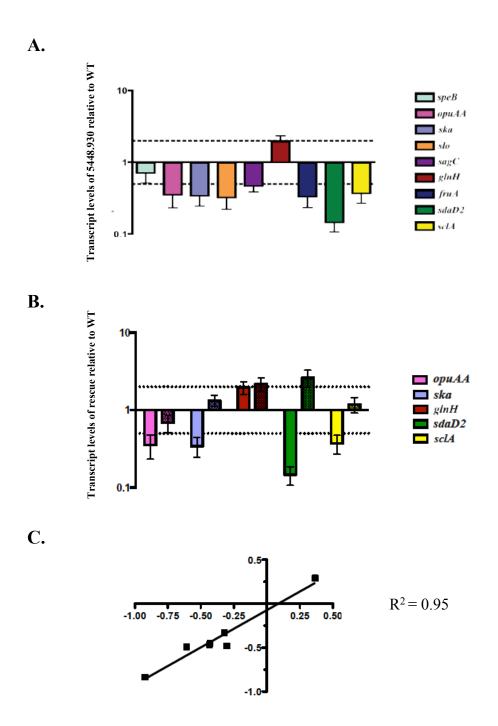
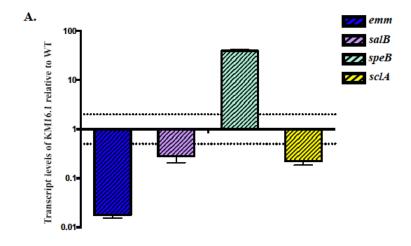


Figure S4. Confirmation of C media RNA-Seq results for Mga regulon using qPCR. (A) Transcript levels were determined using qPCR on RNA taken from WT GAS 5448 grown in C media at late-log compared to Δmga 5448.930 grown under the same conditions. Error bars represent the standard error from three biological replicates. Differences greater than 2-fold in expression for WT compared to mutant cells (dashed line) were considered significant. (B) Complementation of the Δmga mutant using the rescue strain 5448.930R (hashed bars) restores the transcriptional phenotypes observed in C media. (C) Coefficient of correlation between the RNA-Seq and RT-qPCR results. Generated a confidence in the data with an $R_2 = 95\%$.



Figure S5. Sda1 levels are reduced in Δmga M1T1 GAS supernatants grown in C media. Western blot analysis of 44-kDa Sda1 streptodornase (encoded by sdaD2) in culture supernatants grown to late-log phase (Klett ~70) from WT 5448 and two Δmga mutants (5448.930 and KM16.1). Concentrated culture supernatants were isolated as described in Methods and equal amounts were separated by 10% SDS-PAGE, transferred to nitrocellulose, and probed using a polyclonal α-Sda1 antibody.



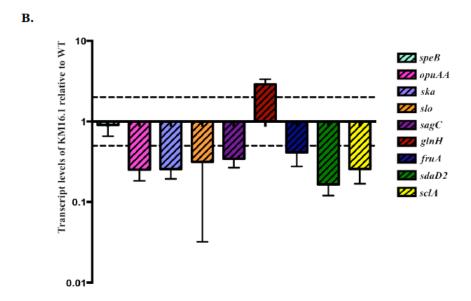


Figure S6. Media-dependent transcriptional regulation by Mga is recapitulated in an independent 5448 Δmga mutant. Transcript levels were determined using qPCR on RNA taken from WT GAS 5448 grown in THY (A) or C media (B) at late-log compared to Δmga KM16.1 grown under the same conditions. Error bars represent the standard error from three biological replicates. Differences greater than 2-fold in expression for WT compared to mutant cells (dashed line) were considered significant.

Supplemental Table S1: Genes in 5448 differentially expressed (DE) in THY compared to C Media

Spy Number	cre (CcpA) ^a	Annotation	Gene	Log ₂ FC	p value
M5005_Spy0011		tRNA(Ile)-lysidine synthetase	tilS	-1.45	2.6E-04
M5005 Spy0017	(R^2)	CHAP-domain-containing cell wall hydrolase; pcsB/sibA	cdhA	2.34	7.0E-07
M5005_Spy0019	cre	DNA repair protein	recO	-1.05	1.1E-03
M5005_Spy0022		phosphoribosylaminoimidazole-succinocarboxamide synthase		2.45	1.1E -0 4
M5005_Spy0023		phosphoribosylformylglycinamidine synthase		1.97	2.7E-04
M5005_Spy0024		amidophosphoribosyltransferase	purF	2.76	1.8E-06
M5005_Spy0025		phosphoribosylformylglycinamidine cyclo-ligase	purM	2.38	1.0E-04
M5005_Spy0026	cre	phosphoribosylaminoimidazolecarboxamide formyltransferase	purN	2.05	1.4E-04
M5005_Spy0027		cyclohydrolase	D	2.00	2.7E-05
M5005_Spy0029 M5005_Spy0030		phosphoribosylamineglycine ligase phosphoribosylaminoimidazole carboxylase catalytic subunit	purD E	2.21 1.63	7.2E-04 7.6E-03
M5005_Spy0030 M5005_Spy0031		phosphoribosylaminoimidazole carboxylase catalytic subunit	purE purK	1.03	5.1E-03
M5005_Spy0031 M5005_Spy0032		hypothetical protein	purk	1.06	9.2E-03
M5005_Spy0032 M5005_Spy0034		transcriptional regulator		1.24	3.0E-03
M5005_Spy0035		holliday junction DNA helicase	ruvB	1.46	1.5E-04
M5005_Spy0039	$cre2 (R^{3,4})$	alcohol dehydrogenase/acetaldehyde dehydrogenase	adh2	-2.07	6.8E-03
M5005_Spy0040	$cre(\mathbf{R}^{2,3})$	alcohol dehydrogenase	adhA	-1.94	2.3E-02
M5005_Spy0041	,	Na+ driven multidrug efflux pump		-1.03	9.0E-03
M5005_Spy0043		SSU ribosomal protein S10P	rpsJ	1.15	2.2E-02
M5005_Spy0044		LSU ribosomal protein L3P	rplC	1.04	1.0E-02
M5005_Spy0052		LSU ribosomal protein L29P	rpmC	1.54	1.0E-02
M5005_Spy0064		protein translocase subunit	secY	1.22	7.7E-03
M5005_Spy0065		adenylate kinase	adk	1.97	4.3E-04
M5005_Spy0066		bacterial protein translation initiation factor 1	infA	1.54	3.7E-04
M5005_Spy0068		SSU ribosomal protein S13P	rpsM	1.25	1.8E-03
M5005_Spy0070		DNA-directed RNA polymerase alpha chain	rpoA	1.49	9.2E-04
M5005_Spy0071		LSU ribosomal protein L17P	rplQ	1.70	2.6E-04
M5005_Spy0073		hypothetical protein		2.57	5.5E-03
M5005_Spy0076		4-diphosphocytidyl-2-C-methyl-D-erythritol kinase	a do D	1.16	1.7E-02
M5005_Spy0077		transcriptional regulator, MarR family high-affinity zinc uptake system ATP-binding protein	adcR adcC	1.09 1.10	8.8E-05 1.1E-03
M5005_Spy0078 M5005_Spy0085		putative DNA binding protein	aacc	2.62	9.1E-06
M5005_Spy0085 M5005_Spy0086	cre	comG operon protein 1	comYA	2.02	5.2E-03
M5005_Spy0088	Cre	comG operon protein 3	comYC	1.76	2.4E-03
M5005 Spy0094	$cre(\mathbb{R}^{2,3})$	acetate kinase	ackA	1.21	2.0E-03
M5005 Spy0095	(R^4)	hypothetical protein		2.59	1.7E-04
M5005 Spy0106		transcriptional regulator; Regulator of protein F	rofA	-1.39	8.0E-04
M5005_Spy0107		fibronectin-binding protein; FCT Pilus Region	сра	2.50	3.8E-07
M5005_Spy0108		signal peptidase I; FCT Pilus Region	sipA1	2.52	2.7E-06
M5005_Spy0109		fibronectin-binding protein; FCT Pilus Region		2.37	1.0E-04
M5005_Spy0110		hypothetical protein	srtC2	2.61	7.6E-06
M5005_Spy0111		hypothetical protein		1.63	1.6E-03
M5005_Spy0113		transposase	_	2.14	4.2E-03
M5005_Spy0114		sortase; FCT Pilus Region	srtB	2.59	3.0E-04
M5005_Spy0115		hypothetical protein		-1.26	3.9E-04
M5005_Spy0134		tellurite resistance protein		-1.33 1.39	1.3E-05
M5005_Spy0137		nucleoside-binding protein hypothetical protein		2.34	1.6E-05 2.5E-03
M5005_Spy0142 M5005_Spy0143	cre	hypothetical protein		3.97	1.9E-03
M5005_Spy0145 M5005_Spy0148	cre	PTS system, 3-keto-L-gulonate specific IIC	ptxC	-2.37	5.9E-03
M5005_Spy0148	(R^4)	PTS system, 3-keto-L-gulonate specific IIB	ptxB	-4.53	2.0E-05
M5005_Spy0150	(R^4)	PTS system, 3-keto-L-gulonate specific IIA	ptxA	-4.61	3.3E-04
M5005_Spy0151	$(R^{2,4})$	3-keto-L-gulonate-6-phosphate decarboxylase	pun	-3.52	4.3E-03
M5005 Spy0152	()	L-xylulose 5-phosphate 3-epimerase		-3.57	7.4E-04
M5005_Spy0153		L-ribulose-5-phosphate 4-epimerase	araD	-5.77	3.5E-06
M5005_Spy0154		hypothetical protein		-3.95	5.7E-05
M5005_Spy0156		metal-dependent hydrolase		-2.14	4.0E-04
M5005_Spy0157	(A^3)	glycine betaine transport ATP-binding protein	opuAA	2.11	1.7E-07
M5005_Spy0158		glycine betaine transport system permease protein	opuABC	2.01	2.5E-06
M5005_Spy0174		hypothetical protein		1.96	1.2E-02
M5005_Spy0175		queuine tRNA-ribosyltransferase	tgt	1.06	2.1E-03
M5005_Spy0177		bioY protein		1.21	2.0E-02
M5005_Spy0185	cre	glucose-6-phosphate isomerase	pgi	1.09	3.8E-02
M5005_Spy0192		UTPglucose-1-phosphate uridylyltransferase	hasC.2	-1.19	1.4E-03

M5005 Spy0194	(R^4)	glycerol-3-phosphate dehydrogenase; GAPDH	gapA	-1.27	2.1E-04
M5005 Spy0195		Heme-sensitive transcriptional repressor, MarR-family	pefR	1.17	1.3E-02
M5005_Spy0204		TCS-1 histidine kinase FasB	fasB	1.10	1.4E-03
M5005_Spy0206		TCS-1 response regulator FasA	fasA	1.13	1.0E-02
M5005_Spy0211		LSU ribosomal protein L34P	rpmH	1.64	1.6E-03
M5005_Spy0213	$(R^{2,3,4})$	ABC Sialic Acid; N-acetylneuraminate-binding protein	•	-1.98	1.6E-03
M5005 Spy0214	$cre(\mathbf{R}^{2,3,4})$	ABC Sialic Acid; N-acetylneuraminate permease protein		-2.11	5.6E-04
M5005 Spy0215	$(R^{2,3,4})$	ABC Sialic Acid; N-acetylneuraminate permease protein		-1.83	3.2E-03
M5005 Spy0216	$(\mathbb{R}^{2,3,4})$	hypothetical membrane spanning protein		-2.87	1.1E-04
M5005_Spy0217	$(R^{2,3,4})$	N-acetylneuraminate lyase	nanH	-1.94	3.3E-03
M5005_Spy0218	$(R^{3,4})$	N-acetylmannosamine kinase		-1.95	2.8E-03
M5005_Spy0221	()	ribonuclease M5		1.31	1.8E-03
M5005_Spy0224		ribulose-phosphate 3-epimerase	rpe	-1.36	1.2E-04
M5005_Spy0230		SSU ribosomal protein S12P	rpsL	1.71	4.2E-04
M5005_Spy0231		SSU ribosomal protein S7P	rpsG	1.32	4.4E-03
M5005_Spy0238		putative undecaprenol kinase	bacA	-1.20	1.7E-03
M5005 Spy0241		hypothetical cytosolic protein	rgpG	1.62	2.5E-02
M5005_Spy0247		D-alanyl-D-alanine carboxypeptidase	180	1.79	1.2E-05
M5005_Spy0277		ABC transporter permease protein		1.32	4.4E-02
M5005_Spy0274	cre	branched-chain amino acid transport system carrier protein	braB	4.77	5.8E-07
M5005_Spy0275	cre	serine/threonine sodium symporter	orab	3.26	2.2E-07
M5005_Spy0275 M5005_Spy0295	Cre	60 kDa inner membrane protein YIDC		-1.20	1.6E-03
M5005_Spy0295 M5005_Spy0296		acylphosphatase		-1.23	2.0E-03
M5005_Spy0270 M5005_Spy0307		DNA integration/recombination/invertion protein		-1.22	1.3E-04
M5005_Spy0307 M5005_Spy0318		pyruvate formate-lyase activating enzyme	pflC	1.35	3.5E-04
M5005_Spy0318 M5005_Spy0321	(R^2)	ferrichrome transport system permease protein	fhuG	1.33	5.2E-04
	(R^2)	ferrichrome-binding protein	fhuD	1.13	7.5E-03
M5005_Spy0323					
M5005_Spy0324	(R^2)	ferrichrome transport ATP-binding protein	fhuA	1.85	1.2E-04
M5005_Spy0329		hypothetical cytosolic protein		-1.11	5.1E-05
M5005_Spy0333		signal peptidase-like protein		-1.23	4.7E-04
M5005_Spy0335		corrin/porphyrin methyltransferase		-0.97	1.8E-04
M5005_Spy0336		hypothetical membrane associated protein		-1.13	1.0E-02
M5005_Spy0338		arsenate reductase family protein	CED	1.24	1.3E-03
M5005_Spy0341	cre	lactocepin; SpyCEP; ScpC; IL-8 protease	spyCEP	-1.78	1.4E-04
M5005_Spy0346	(R^2)	hypothetical protein	15. 1	-1.41	2.8E-02
M5005_Spy0347	(R^2)	ribonucleoside-diphosphate reductase beta chain	nrdF.1	-1.25	2.0E-03
M5005_Spy0348	(R^2)	hypothetical protein	nrdI	-1.49	5.0E-03
M5005_Spy0349	(R^2)	ribonucleoside-diphosphate reductase alpha chain	nrdE.1	-1.56	4.3E-03
M5005_Spy0361	$cre(\mathbf{R}^{2,4})$	phosphoglycerate transporter protein	177	-1.05	3.3E-02
M5005_Spy0374		LSU ribosomal protein L11P	rplK	1.84	4.9E-03
M5005_Spy0375		LSU ribosomal protein L1P	rplA	1.86	3.2E-03
M5005_Spy0385	cre	67 kDa Myosin-crossreactive antigen		-1.23	5.2E-03
M5005_Spy0387		uracil DNA glycosylase superfamily protein		-0.98	1.3E-02
M5005_Spy0398		bacteriocin		1.53	4.0E-02
M5005_Spy0421		lactoylglutathione lyase	gloA	-1.70	1.2E-05
M5005_Spy0422		NAD(P)H-dependent quinone reductase		-1.94	4.0E-06
M5005_Spy0423	cre2	Xaa-Pro dipeptidase	pepQ	-1.61	1.8E-04
M5005_Spy0425		glycosyltransferase		-1.29	2.1E-04
M5005_Spy0426		1,2-diacylglycerol 3-glucosyltransferase		-1.52	2.8E-04
M5005_Spy0457		plasmid stabilization system protein		-1.08	2.5E-02
M5005_Spy0461		hypothetical cytosolic protein		-1.67	1.6E-04
M5005_Spy0462		hypothetical protein		-1.50	6.6E-03
M5005_Spy0463		hypothetical cytosolic protein		-1.13	5.4E-03
M5005_Spy0464		microcin C7 self-immunity protein	mccF	-1.14	1.3E-03
M5005_Spy0465		hypothetical protein		-0.95	1.5E-02
M5005_Spy0466		hypothetical protein		-1.43	2.9E-05
M5005_Spy0467		transposase		-1.09	2.8E-02
M5005_Spy0469		hypothetical protein		-1.74	9.1E-04
M5005_Spy0473	224	multidrug resistance protein B		1.46	1.4E-03
M5005_Spy0474	$(R^{2,3,4})$	transcription antiterminator, BglG family	licT	2.43	7.5E-05
M5005_Spy0475	$cre(\mathbf{R}^{2,3,4})$	PTS system, b-Glucoside -specific IIABC		3.78	2.7E-08
M5005_Spy0476	$(R^{2,3,4})$	6-phospho-beta-glucosidase	bglA	2.94	4.4E-06
M5005_Spy0477		hypothetical membrane spanning protein		-1.67	3.6E-05
M5005_Spy0478		hypothetical membrane spanning protein		-1.65	3.7E-04
M5005_Spy0479		hypothetical membrane spanning protein		-3.11	5.0E-02
M5005_Spy0483		stress-responsive transcriptional regulator		-2.31	4.4E-08
M5005_Spy0485		prolipoprotein diacylglyceryl transferase	lgt	-0.96	5.6E-04
M5005_Spy0486		hypothetical protein	-	-1.21	3.0E-03
M5005_Spy0496		hydrolase, HAD superfamily		1.90	5.5E-06

M5005 G 0502		.1.4.411		1.45	1.25.04
M5005_Spy0503		glutathione peroxidase	F	-1.45	1.2E-04
M5005_Spy0504		oligoendopeptidase F	pepF	-1.70	1.0E-05
M5005_Spy0512	cre	hydrolase, HAD superfamily		-1.02	8.9E-05
M5005_Spy0523		hypothetical protein		-1.63	2.3E-02
M5005_Spy0528		glucose-1-phosphate phosphodismutase		-1.13	1.2E-03
M5005_Spy0539		ATP-binding protein		-1.30	2.2E-05
M5005_Spy0540		transporter		-1.27	4.1E-04
M5005_Spy0541		hypothetical cytosolic protein		-1.50	6.2E-05
M5005 Spy0542		dipeptidase	pepD	-1.77	8.8E-05
M5005_Spy0543	cre2	high-affinity zinc uptake system protein znuA precursor		1.49	3.5E-03
M5005_Spy0545		galactosamine-6-phosphate deaminase (isomerizing)	agaS	-0.93	1.5E-02
M5005_Spy0546		LSU ribosomal protein L31P	rpmE	1.77	5.0E-06
M5005_Spy0549	cre	chorismate mutase	<i>P</i> ·	1.13	1.0E-02
M5005_Spy0551	c, c	LSU ribosomal protein L19P	rplS	2.14	4.6E-05
M5005_Spy0555		hypothetical cytosolic protein	TPIS	-1.40	7.8E-04
M5005_Spy0558				-1.40	4.6E-02
	$(R^{2,3,4})$	transposase	~~~1	-2.37	
M5005_Spy0562	$(R^{2,3,4})$	streptolysin S precursor	sagA		2.1E-04
M5005_Spy0563	$(R^{2,3,4})$	streptolysin S biosynthesis protein	sagB	-1.86	1.9E-04
M5005_Spy0564	$(R^{2,3,7})$	streptolysin S biosynthesis protein	sagC	-1.87	1.0E-04
M5005_Spy0565	$(R^{2,3,4})$	streptolysin S biosynthesis protein	sagD	-1.25	1.9E-03
M5005_Spy0566	$(R^{2,3,4})$	streptolysin S putative self-immunity protein	sagE	-2.32	5.5E-05
M5005_Spy0567	$(R^{2,3,4})$	streptolysin S biosynthesis protein	sagF	-1.82	1.0E-03
M5005_Spy0568	$(R^{2,3,4})$	streptolysin S export ATP-binding protein	sagG	-1.88	7.7E-05
M5005_Spy0569	$(R^{2,3,4})$	streptolysin S export transmembrane protein	sagH	-1.76	1.9E-04
M5005_Spy0570	$(R^{2,3,4})$	streptolysin S export transmembrane protein	sagI	-1.14	2.7E-02
M5005_Spy0575		ATP synthase C chain	atpE	1.60	1.4E-04
M5005_Spy0576	(A^3)	ATP synthase A chain	atpB	1.26	1.7E-03
M5005_Spy0579	(A^3)	ATP synthase alpha chain	atpA	1.43	2.0E-02
M5005_Spy0593	()	neutral zinc metallopeptidase family		-1.35	7.7E-05
M5005 Spy0594		ATP-dependent nuclease subunit B	rexB	-1.18	6.8E-04
M5005_Spy0595		ATP-dependent nuclease subunit A	rexA	-1.26	4.1E-04
M5005_Spy0597		SSU ribosomal protein S21P	I CAII	1.39	1.1E-03
		Group A Carbohydrate; Membrane protein	and I	-1.02	
M5005_Spy0611			gacJ		8.6E-03
M5005_Spy0615		pore forming protein	ebsA	-1.66	3.5E-04
M5005_Spy0616		ferredoxin	1	-1.78	4.7E-02
M5005_Spy0618		cytidylate kinase	cmk	-1.08	4.3E-04
M5005_Spy0619		bacterial protein translation initiation factor 3	infC	2.03	4.4E-04
M5005_Spy0620	cre	LSU ribosomal protein L35P	rpl36	1.93	2.3E-03
M5005_Spy0621		LSU ribosomal protein L20P	rplT	2.34	1.2E-05
M5005_Spy0624		3-dehydroquinate dehydratase	aroD	1.25	4.3E-04
M5005_Spy0625		chorismate synthase	aroF	1.20	5.5E-03
M5005_Spy0628		folylpolyglutamate synthase/dihydrofolate synthase	folC.2	-1.18	1.5E-04
M5005_Spy0633		LSU ribosomal protein L21P	rplU	1.84	3.9E-05
M5005_Spy0634		hypothetical ribosome-associated protein		1.05	1.4E-03
M5005_Spy0635		LSU ribosomal protein L27P	rpmA	2.05	2.7E-05
M5005 Spy0636		transcriptional regulator, LysR family		-1.36	2.0E-05
M5005_Spy0637		lipoprotein signal peptidase	lsp	-1.45	5.0E-05
M5005_Spy0638		ribosomal large subunit pseudouridine synthase D	•	-1.21	1.1E-05
M5005 Spy0639		pyrR bifunctional protein	pyrR	1.57	1.6E-03
M5005 Spy0640		uracil permease	pyrP	1.30	5.0E-03
M5005_Spy0641		aspartate carbamoyltransferase	pyrB	2.18	1.7E-05
M5005_Spy0642		carbamoyl-phosphate synthase small chain	carA	1.73	3.8E-04
M5005_Spy0643		carbamoyl-phosphate synthase large chain	carB	1.62	1.6E-03
M5005_Spy0648		SSU ribosomal protein S16P	rpsP	1.74	4.3E-04
		RNA binding protein	rpsi	1.59	7.4E-05
M5005_Spy0649	(R^2)		hunV	1.11	1.0E-02
M5005_Spy0651	(K)	Heme-utilization cell surface protein	hup Y		
M5005_Spy0652		Heme-oxygenase cytosolic protein	hupZ	1.76	4.9E-04
M5005_Spy0653	cre	cobalt-zinc-cadmium resistance protein	czcD	-3.16	3.4E-05
M5005_Spy0655		16S rRNA processing protein	rimM	1.25	6.3E-04
M5005_Spy0668		IgG-degrading protease	mac	1.82	4.0E-03
M5005_Spy0669		phage protein		1.84	3.4E-02
M5005_Spy0676		hypothetical protein		1.25	1.6E-02
M5005_Spy0686		3-hydroxy-3-methylglutaryl-coenzyme A reductase		-1.29	5.4E-04
M5005_Spy0687	4	hydroxymethylglutaryl-CoA synthase	mvaS.1	-0.95	1.5E-02
M5005_Spy0696	(R^4)	phosphopentomutase	deoB	-1.00	7.2E-03
M5005_Spy0698	(R^4)	purine nucleoside phosphorylase	punA	-1.01	2.4E-02
M5005_Spy0703		orotidine 5'-phosphate decarboxylase	pyrF	1.62	6.4E-03
M5005_Spy0704		orotate phosphoribosyltransferase	pyrE	2.61	1.4E-03
M5005_Spy0706		cystine-binding protein	- ·	1.67	4.8E-05

M5005_Spy0707		cystine transport system permease protein		1.07	8.2E-03
M5005 Spy0717	cre	rhodanese-related sulfurtransferases		1.23	6.7E-04
M5005 Spy0718		hypothetical protein		1.11	7.4E-03
		glutathione S-transferase		2.40	2.2E-06
M5005_Spy0719					
M5005_Spy0721		hypothetical protein		-0.99	3.2E-02
M5005_Spy0732		NIF3-related protein		1.24	2.4E-02
M5005_Spy0735		dTDP-4-dehydrorhamnose 3,5-epimerase	cpsFP	-1.04	8.1E-04
M5005_Spy0737		mutator mutT protein/7,8-dihydro-8-oxoguanine-triphosphatase	1	-1.48	6.3E-05
M5005_Spy0738		hypothetical membrane spanning protein		-1.64	1.7E-06
M5005_Spy0739		tetratricopeptide repeat family protein		-1.19	1.2E-03
M5005_Spy0751		pyruvate dehydrogenase E1 component alpha subunit	acoA	1.95	5.3E-03
M5005_Spy0752		pyruvate dehydrogenase E1 component beta subunit	acoB	1.85	9.3E-04
M5005_Spy0753		dihydrolipoamide acetyltransferase of pyruvate dehydrogenase	acoC	1.02	5.2E-03
			ucoc		
M5005_Spy0774		nucleoside diphosphate kinase		2.02	5.4E-03
M5005_Spy0775	224	nucleoside diphosphate kinase		1.61	6.3E-04
M5005_Spy0780	$cre(\mathbf{R}^{2,3,4})$	PTS system, Mannose/Fructose family IIA component	ptsA	1.51	7.6E-03
M5005 Spy0781	$(R^{2,3,4})$	PTS system, Mannose/Fructose family IIB component	ptsB	1.33	1.3E-03
M5005_Spy0783	$(R^{2,3,4})$	PTS system, Mannose/Fructose family IID component	ptsD	1.12	1.0E-02
	$cre(\mathbb{R}^{3,4})$				
M5005_Spy0790	cre (K)	succinate-semialdehyde dehydrogenase	gabD	-1.23	1.5E-02
M5005_Spy0791	(R^4)	excinuclease ABC subunit C	uvrC	-1.98	5.5E-04
M5005_Spy0792		NAD(P)H-dependent quinone reductase		-2.64	1.2E-04
M5005 Spy0793		Xaa-His dipeptidase		-1.74	1.8E-02
M5005_Spy0794		tRNA (5-carboxymethylaminomethyl-2-thiouridylate) synthase	thdF	2.16	4.5E-06
M5005_Spy0795		LSU ribosomal protein L10P	rplJ	1.52	9.0E-03
M5005_Spy0796		LSU ribosomal protein L12P	rplL	2.69	1.0E-03
M5005_Spy0804		TCS-6 response regulator; Streptin biosynthesis	srtR	-0.94	3.6E-04
M5005 Spy0806		lantibiotic precursor	srtA	-1.24	2.1E-03
M5005_Spy0807		ABC transporter (ATP binding)-lantibiotic associated	srtT	-1.63	6.4E-06
M5005_Spy0808		lantibiotic transport ATP-binding protein	srtF	-0.95	3.5E-03
M5005_Spy0809		lantibiotic transport permease protein	srtE	-1.98	3.8E-05
M5005_Spy0810		lantibiotic transport permease protein	srtG	-1.62	1.1E-04
M5005 Spy0811		transcriptional regulator, Cro/CI family		-1.29	4.6E-02
M5005_Spy0820		tetrahydropholate synth./dihydrofolate synthase	folC.1	-2.04	5.6E-08
	2		-		
M5005_Spy0821	cre2	tetrahydropholate synth., GTP cyclohydrolase I	fol E	-1.20	3.7E-04
M5005_Spy0822		tetrahydropholate synth., dihydropteroate synthase	folP	-1.11	2.1E-02
M5005 Spy0823		tetrahydropholate synth., dihydroneopterin aldolase	folQ	-1.34	3.4E-04
M5005_Spy0824		tetrahydropholate synth., pyrophosphokinase	folK	-1.51	1.2E-04
M5005_Spy0825		UDP-N-acetylenolpyruvoylglucosamine reductase	murB	1.29	2.1E-04
M5005_Spy0826		spermidine/putrescine transport system ATP-binding protein	potA	2.58	4.0E-05
M5005_Spy0827		spermidine/putrescine transport system permease protein	potB	1.36	7.8E-04
M5005 Spy0828		spermidine/putrescine transport system permease protein	potC	1.52	4.0E-03
M5005 Spy0829		spermidine/putrescine-binding protein	P	2.29	3.4E-04
	(n 2.3)				
M5005_Spy0835	$cre(\mathbb{R}^{2,3})$	class B acid phosphatase		-1.26	2.4E-03
M5005_Spy0841		glutamine amidotransferase, class I		-1.06	2.8 E-03
M5005_Spy0843		hypothetical cytosolic protein		-1.46	7.7E-05
M5005 Spy0845		ribose-phosphate pyrophosphokinase		-1.00	1.7E-03
M5005_Spy0852	$(R^{3,4})$	short chain dehydrogenase		2.02	1.6E-06
M5005_Spy0853	(R^3)	short chain dehydrogenase		2.86	4.2E-07
M5005_Spy0858	(A^4)	xanthine phosphoribosyltransferase	xpt	4.52	1.5E-06
M5005_Spy0859	(A^4)	xanthine permease		4.41	1.6E-09
M5005 Spy0873	cre	L-lactate dehydrogenase	ldh	1.60	1.4E-03
M5005_Spy0879	0.0	hypothetical protein		-2.03	3.5E-04
M5005_Spy0880		conserved membrane protein		-5.13	1.1E-06
M5005_Spy0881	cre	hypothetical cytosolic protein		-5.63	7.1E-09
M5005_Spy0883		ribonuclease HII		1.06	3.5 E-05
M5005_Spy0884		hypothetical protein	smf	2.51	1.1E-08
M5005_Spy0890		D-lactate dehydrogenase	ddh	1.57	6.8E-03
			aan		
M5005_Spy0911		hypothetical protein		-1.01	8.4E-03
M5005_Spy0914		phage transcriptional repressor		1.30	5.0E-03
M5005 Spy0943	cre	cytidine deaminase	cdd	1.42	5.4E-04
M5005 Spy0945	cre (R ⁴)	pantothenate kinase	coaA	-1.61	4.4E-04
	cre (IC)	•	соил		
M5005_Spy0965	m2 . 4	ABC transporter permease protein		-1.09	6.2E-04
M5005_Spy0982	$cre(R^2,A^4)$	histidine-binding protein		1.14	1.3E-03
M5005_Spy1017		phage protein		3.15	6.5E-03
M5005_Spy1023		terminase large subunit		1.45	3.6E-02
M5005_Spy1028		phage protein		3.67	1.9E-03
M5005_Spy1046		phage protein		-1.28	5.0E-03
M5005_Spy1062	2.2	maltodextrose utilization protein	malA	-1.96	4.0E-02
M5005_Spy1063	$(R^{2,3})$	ABC Cyclomaltodextrin permease protein	malD	-1.85	3.9E-02

M5005 Spy1064	$(R^{2,3})$	ABC Cyclomaltodextrin permease protein	malC	-1.57	5.1E-02
M5005_Spy1004 M5005_Spy1065	$(R^{2,3})$	alpha-amylase	amyA	-1.70	2.9E-02
M5005_Spy1066	$(R^{2,3})$	neopullulanase/cyclomaltodextrinase/maltogenic alpha-amylase	amyB	-1.65	5.4E-02
M5005_Spy1069	$(R^{2,3})$	esterase	u,2	-0.99	2.8E-03
M5005_Spy1070	()	protein precursor	dltD	-0.92	1.0E-02
M5005_Spy1072		protein DltB	dltB	-1.47	6.7E-05
M5005 Spy1073		D-alanine-activating enzyme	dltA	-1.43	1.0E-06
M5005_Spy1074		hypothetical protein		-1.79	3.1E-04
M5005_Spy1075	2.4	excinuclease ABC subunit B	uvrB	-1.57	6.0E-07
M5005_Spy1079	$(R_{2}^{2,4})$	PTS system, Cellobiose-specific IIC component		-2.09	3.4E-03
M5005_Spy1080	$(R_{2,4}^{2,4})$	hypothetical protein		-1.43	6.7E-03
M5005_Spy1081	$(R^{2,4})$	PTS system, Cellobiose-specific IIA component		-1.76	8.4E-03
M5005_Spy1082	$(R^{2,4})$	PTS system, Cellobiose-specific IIB component		-1.99	4.1E-03
M5005_Spy1083	$cre(\mathbf{R}^{2,4})$	PTS system, Cellobiose-specific IIA component		-1.83	5.8E-04
M5005_Spy1084	$(R^{2,4})$ cre (R^4)	outer surface protein	h~112	-1.25	2.2E-02
M5005_Spy1085 M5005_Spy1093	$cre(\mathbf{R}^{2})$ $cre(\mathbf{R}^{2,3,4})$	beta-glucosidase hypothetical protein	bglA.2	-1.61 -3.31	6.6E-03 2.3E-06
M5005_Spy1095 M5005_Spy1096	cre(K)	thioesterase superfamily protein		1.85	7.6E-03
M5005_Spy1090 M5005_Spy1097		phosphorylase, Pnp/Udp family		-0.97	2.2E-02
M5005_Spy1007 M5005_Spy1100		shikimate kinase	aroK	-1.23	4.6E-03
M5005_Spy1102		ribonuclease BN		-1.46	1.7E-04
M5005 Spy1106	cre	protein G-related alpha 2M-binding protein	grab	1.90	7.1E-05
M5005_Spy1107		UDP-N-acetylglucosamine 1-carboxyvinyl transferase	murZ	1.34	6.5E-03
M5005 Spy1109		internalin protein	inlA	1.27	1.0E-03
M5005_Spy1114		hypothetical membrane spanning protein		-1.93	9.0E-04
M5005_Spy1117		ATP-dependent RNA helicase	deaD2	-1.00	4.3E-03
M5005_Spy1120		phosphoenolpyruvate-protein phosphotransferase	pstI	1.55	6.2E-04
M5005_Spy1122		glutaredoxin	nrdH	-1.87	4.9E-05
M5005_Spy1123		ribonucleoside-diphosphate reductase alpha chain	nrdE.2	-1.62	1.0E-03
M5005_Spy1124		ribonucleoside-diphosphate reductase beta chain	nrdF.2	-1.07	4.8E-03
M5005_Spy1125		chloride channel protein		-1.59 2.11	3.1E-06
M5005_Spy1131 M5005_Spy1132		transcriptional regulator, Cro/CI family alanyl-tRNA synthetase	alaS	2.11	3.7E-05 9.0E-06
M5005_Spy1132 M5005_Spy1137		putative competence protein/transcription factor	шиз	1.62	2.3E-03
M5005_Spy1137 M5005_Spy1139		glucosamine-6-phosphate isomerase	nagB	1.24	2.7E-03
M5005_Spy1139		hypothetical protein	nage	-2.17	9.2E-03
M5005_Spy1143		hypothetical protein		-2.16	1.9E-02
M5005_Spy1144	cre	hypothetical protein		-2.26	1.1E-03
M5005_Spy1149		1-acyl-sn-glycerol-3-phosphate acyltransferase		1.23	3.3E-02
M5005_Spy1152		kup system potassium uptake protein		1.56	3.1E-04
M5005_Spy1167		Pb, Cd, Zn and Hg-transporting ATPase	pmtA	1.43	1.9E-03
M5005_Spy1199		phage protein		2.11	2.8E-02
M5005_Spy1205		phage protein		1.51	1.0E-02
M5005_Spy1227		hypothetical protein	D.1	1.08	2.6E-03
M5005_Spy1229		arginine repressor	argR1	1.72	7.1E-03
M5005_Spy1230 M5005_Spy1231		hemolysin dimethylallyltransferase/geranyltranstransferase	Gora	1.18 1.55	5.0E-02 1.7E-02
M5005_Spy1231 M5005_Spy1232		exodeoxyribonuclease VII small subunit	fps xseB	2.43	1.7E-02 1.1E-03
M5005_Spy1232 M5005_Spy1237	(R^2)	arginine transport ATP-binding protein	artP	2.19	7.8E-05
M5005_Spy1238	(R^2)	arginine transport system permease protein	artQ	1.52	1.3E-03
M5005_Spy1241	(/	mutator protein (7,8-dihydro-8-oxoguanine-triphosphatase)	mutT	-0.95	2.1E-03
M5005_Spy1256	(R^4)	rhodanese-related sulfurtransferases		1.12	8.3E-03
M5005_Spy1257	(R^3)	glucokinase/Xylose repressor	glcK	1.98	2.7E-04
M5005_Spy1258	cre	hypothetical cytosolic protein		2.50	3.0E-06
M5005_Spy1263	224	hypothetical protein		-1.65	3.3E-03
M5005_Spy1270	$(R_{2,3,4}^{2,3,4})$	carbamate kinase	arcC	-3.88	3.1E-05
M5005_Spy1271	$(R^{2,3,4})$	Xaa-His dipeptidase		-4.11	5.7E-09
M5005_Spy1272	$(R^{2,3,4})$	arginine/ornithine antiporter	ъ.	-4.09	2.0E-07
M5005_Spy1273	$(R^{2,3,4})$ $(R^{2,3,4})$	ornithine carbamoyltransferase	arcB	-4.04 2.70	3.8E-06
M5005_Spy1274	$(R^{2,3,7})$ $cre(R^{2,3,4})$	acetyltransferase	ans 1	-3.79	1.9E-07
M5005_Spy1275	cre (K · ·)	arginine deiminase TCS-9 response regulator yesN-like	arcA tcs9R	-3.22 -0.99	2.5E-04 1.6E-02
M5005_Spy1281 M5005_Spy1297		phospho-2-dehydro-3-deoxyheptonate aldolase	icstr	-0.99 1.91	7.1E-04
M5005_Spy1297 M5005_Spy1298		3-dehydroquinate synthase	aroB	2.02	8.4E-03
M5005_Spy1299		hypothetical protein	W. OD	-1.91	4.4E-02
M5005_Spy1299		TCS-10 response regulator; TCS-X; YesN-like	trxR	-1.20	1.4E-02
M5005_Spy1308		ABC unknown sugar-binding protein		-1.79	8.8E-03
M5005_Spy1309		ABC unknown sugar permease protein		-2.78	7.0E-03
M5005_Spy1310	cre2	ABC unknown sugar permease protein		-2.82	6.2E-03

M5005_Spy1311	cre	glucokinase		-1.95	2.4E-03
M5005_Spy1312		hypothetical protein		-3.23	3.4E-03
M5005_Spy1319		tRNA (uracil-5-)-methyltransferase		1.24	2.0E-02
M5005_Spy1323		transposase		-1.12	2.0E-02
	(A^3)		angM		
M5005_Spy1329	(A)	cysteine synthase	cysM	3.54	1.0E-05
M5005_Spy1334		transporter	yvqF	-1.13	1.0E-02
M5005_Spy1340		DNA-directed RNA polymerase omega chain		2.04	1.9E-05
M5005_Spy1341		guanylate kinase	gmk	1.23	4.2E-03
M5005 Spy1342		hydrolase, HAD superfamily	8	1.30	4.2E-04
			D		
M5005_Spy1344		acetyl-CoA acetyltransferase	atoB	-1.16	5.2E-04
M5005_Spy1346		acetate CoA-transferase beta subunit	ato A	-1.44	6.9E-04
M5005_Spy1347		D-beta-hydroxybutyrate dehydrogenase		-1.76	1.0E-03
M5005_Spy1348		D-beta-hydroxybutyrate permease		-2.18	5.5E-08
M5005_Spy1354		recombination protein	recU	-1.14	3.5E-03
M5005_Spy1356	cre	aminopeptidase C	pepC	-1.25	1.8E-02
M5005_Spy1359		amino acid permease		1.36	1.8E-04
M5005 Spy1362		transporter		1.78	4.0E-06
M5005_Spy1363		amino acid ABC transporter permease protein		1.17	1.6E-03
M5005_Spy1374		hypothetical protein		1.17	1.4E-02
	(D 2.3.4)				
M5005_Spy1376	$cre(\mathbf{R}^{2,3,4})$	transaldolase		-2.82	1.0E-04
M5005_Spy1377	$(R^{2,3,4})$	trans-acting positive regulator		-2.26	3.4E-03
M5005_Spy1378	$cre\left(\mathbb{R}^{2,3,4}\right)$	NADH peroxidase		-1.90	1.7E-02
M5005_Spy1379	$(R^{2,3,4})$	glycerol uptake facilitator protein	glpF	-4.84	1.0E-05
M5005 Spy1380	$(R^{2,4})$	alpha-glycerophosphate oxidase	glpO	-4.44	2.1E-07
	(D 2.3.4)				
M5005_Spy1381	$cre(\mathbf{R}^{2,3,4})$	glycerol kinase	glpK	-4.13	5.4E-08
M5005_Spy1391		degV family protein		-1.76	7.6E-06
M5005_Spy1393		hydrolase, HAD superfamily		-1.17	5.1E-05
M5005_Spy1395	$(R^{2,4})$	tagatose-bisphosphate aldolase; SpeB regulator	lacD.1	-2.93	2.7E-04
	(R^2)	tagatose-6-phosphate kinase	lacC.1	-2.29	
M5005_Spy1396	(R)				6.0E-03
M5005_Spy1397	$(R_{2}^{2,4})$	galactose-6-phosphate isomerase lacB subunit	lacB.1	-2.54	8.1E-04
M5005_Spy1398	$(\mathbb{R}^{2,4})$	galactose-6-phosphate isomerase lacA subunit	lacA.1	-2.63	4.2E-03
M5005_Spy1399	$(R^{2,4})$	PTS system, Galactose-specific IIC component		-2.16	2.1E-02
M5005_Spy1408	()	ribosome-binding factor A	rbfA	1.44	2.4E-03
M5005_Spy1409		bacterial protein translation initiation factor 2	-	1.69	2.0E-04
			infB		
M5005_Spy1410		LSU ribosomal protein L7AE		1.47	2.8E-04
M5005_Spy1412		N utilization substance protein A	nusA	1.09	4.2E-03
M5005_Spy1420		phage protein		1.52	3.1E-03
M5005_Spy1428		phage protein		1.79	3.1E-03
M5005_Spy1432		phage protein		1.08	3.2E-02
M5005_Spy1441		phage terminase small subunit		1.52	1.3E-02
M5005_Spy1442		phage transcriptional activator		2.16	2.7E-04
M5005 Spy1444		adenine-specific methyltransferase		2.09	5.8E-03
M5005_Spy1445		phage protein		2.05	5.4E-03
M5005_Spy1447		phage-related DNA helicase			
	(p .2)			1.20	2.0E-02
M5005_Spy1448	(R^2)	hypothetical protein		1.52	4.4E-02
M5005 Spy1449		DNA primase		1.45	2.0E-02
M5005 Spy1450		phage-encoded DNA polymerase		1.45	7.7E-03
M5005_Spy1454		phage protein		1.20	1.0E-02
M5005_Spy1455		phage protein		1.60	1.5E-02
M5005_Spy1459		phage protein		2.58	1.3E-03
M5005 Spy1466		phage protein		1.29	3.2E-02
M5005_Spy1472		bis(5'-nucleosyl)-tetraphosphatase (asymmetrical)	hit	-1.32	2.2E-03
M5005 Spy1476		ATP/GTP hydrolase		1.16	2.2E-04
	(D.4)				
M5005_Spy1477	$(R^4)_{3}$	guanine-hypoxanthine permease		2.27	1.6E-05
M5005_Spy1479	$cre(R^3)$	PTS system, Mannose-specific IIAB; Primary Glucose PTS	manL	2.99	6.8E-05
M5005_Spy1480	$cre\left(\mathbb{R}^{3,4}\right)$	PTS system, Mannose-specific IIC; Primary Glucose PTS	manM	3.36	1.5E-05
M5005 Spy1481	$(R^{3,4})$	PTS system, Mannose-specific IID; Primary Glucose PTS	manN	2.78	4.1E-05
M5005_Spy1482	(11)	hypothetical cytosolic protein	manO	-0.97	3.1E-02
M5005_Spy1493		acyl carrier protein	acpP	1.39	2.0E-02
M5005_Spy1497		chaperone protein	dnaJ	1.27	3.3E-03
M5005_Spy1504		hypothetical membrane spanning protein		1.14	2.4E-02
M5005_Spy1508		glutamyl-tRNA(Gln) amidotransferase subunit C	gatC	1.04	4.3E-04
M5005_Spy1500		pyrazinamidase/nicotinamidase	8	1.01	3.4E-02
		* -			
M5005_Spy1518		transporter	=:	1.52	4.0E-03
M5005_Spy1519		ATP-dependent DNA helicase	recG	-1.14	5.2E-04
M5005_Spy1520		hypothetical protein		-4.09	7.4E-04
M5005_Spy1523		ABC transporter, ATP-binding protein	siaF	-1.09	6.9E-04
M5005_Spy1523 M5005_Spy1533		holo-[acyl-carrier protein] synthase	alr	1.13	8.5E-03
	(D2)				
M5005_Spy1538	(R^2)	mannose-6-phosphate isomerase	pmi	-1.39	8.1E-04

	2				
M5005_Spy1539	(R^2)	fructokinase	scrK	1.59	4.6E-04
M5005_Spy1553		SSU ribosomal protein S18P	rpsR	1.93	1.2E-03
M5005_Spy1559		thioredoxin	trx	-1.26	2.8E-03
M5005_Spy1569	cre	formate acetyltransferase	pfl	2.57	4.0E-07
M5005_Spy1574		universal stress protein family	PJ.	8.91	1.6E-10
M5005_Spy1571	$cre(A^4)$	quinolone resistance protein	norA	9.81	1.4E-12
	cre(A)		norA		
M5005_Spy1581		transcriptional regulator, MerR family	1 0	1.32	8.1E-03
M5005_Spy1582		DNA polymerase III, epsilon chain	dnaQ	1.26	6.5E-03
M5005_Spy1598	cre	hypothetical protein		-1.38	2.1E-04
M5005_Spy1606		LSU ribosomal protein L28P	rpmB	2.04	1.7E-05
M5005 Spy1610	cre2	CTP synthase	pyrG	2.63	7.9E-06
M5005_Spy1611		DNA-directed RNA polymerase delta chain	rpoE	1.33	1.0E-04
M5005_Spy1627		ABC transporter permease protein	salY	-1.16	5.1E-04
M5005_Spy1644	(R^2)	DNA integration/recombination/invertion protein	5411	2.93	3.1E-02
	(14)	SSU ribosomal protein S9P	una I	1.92	3.6E-04
M5005_Spy1646			rpsI		
M5005_Spy1647	- 231	LSU ribosomal protein L13P	rplM	1.51	1.3E-03
M5005_Spy1661	$cre(\mathbf{R}^{2,3,4})$	transaldolase		-1.72	2.3E-03
M5005_Spy1662	$(R^{2,4})$	putative transport protein		-1.70	1.1E-03
M5005_Spy1664	$cre\left(\mathbb{R}^{2,4}\right)$	PTS system, Mannitol-specific IIA component		-1.17	3.2E-02
M5005_Spy1666		SSU ribosomal protein S15P		1.83	4.3E-04
M5005_Spy1667	(R^2)	hypothetical protein		1.58	1.2E-03
M5005 Spy1671	(11)	transcriptional regulator, MarR family		-1.00	8.1E-03
M5005_Spy1675		phosphatidate cytidylyltransferase	cdsA	-1.21	5.1E-04
	(\mathbf{p}^2)		CUSA		
M5005_Spy1691	(R^2)	exodeoxyribonuclease III		-1.63	2.1E-04
M5005_Spy1697		4-amino-4-deoxychorismate lyase		-1.09	3.0E-04
M5005_Spy1698		anthranilate synthase component II	trpG	-1.45	1.2E-04
M5005_Spy1699		ATPase, AAA family		-1.70	9.1E-06
M5005_Spy1710		streptococcal histidine triad protein		3.01	4.2E-08
M5005 Spy1711		laminin binding lipoprotein; AdcR- repressed	lsp	2.25	2.5E-05
M5005_Spy1714	(R^2)	Fn-binding surface protein; Mga-regulated	fba	2.51	1.9E-03
M5005_Spy1715	(R^2)	C5A peptidase; Mga-regulated	scpA	2.20	2.6E-03
	(14)		sepn		
M5005_Spy1716		transposase		-1.66	8.7E-05
M5005_Spy1717		transposase		-2.47	1.0E-03
M5005_Spy1718		secreted inhibitor of complement; Mga-regulated	sic1.0	2.32	2.3E-03
M5005_Spy1721	cre	hypothetical protein; sORF upstream of <i>mga</i>		-2.86	8.2E-03
M5005 Spy1730	(A^3)	hypothetical protein		-1.45	1.5E-04
M5005_Spy1736		hypothetical protein		-2.04	2.0E-02
M5005_Spy1737		Stand-alone transcriptional regulator Rgg, RopB	rgg	-1.43	1.0E-02
M5005_Spy1738	$cre(R^3)$	Secreted DNase-Streptodornase-chromosomal	spd	1.45	8.4E-04
	(R^3)	glycerol dehydrogenase	gldA	-1.50	1.0E - 04
M5005_Spy1741	(K)		_		
M5005_Spy1742		transaldolase	mipB	-1.35	8.9E-04
M5005_Spy1743	. 24	formate acetyltransferase	pflD	-1.35	2.0E-05
M5005_Spy1745	$(\mathbb{R}^{2,4})$	PTS system, Cellobiose-specific IIB component	celB	-1.64	3.2E-02
M5005_Spy1747		sorbitol operon regulator		1.07	2.7E-02
M5005_Spy1765		cold shock protein	csp	1.67	7.8E-03
M5005_Spy1768	(R^4)	peroxiredoxin reductase (NAD(P)H)	ahpC	-1.07	6.0E-03
M5005 Spy1769	(R^4)	peroxiredoxin reductase (NAD(P)H)	ahpF	-1.24	5.7E-03
M5005_Spy1770	$(R^{2,3,4})$	imidazolonepropionase	hutI	-5.24	1.0E-03
M5005_Spy1770					
M5005_Spy1771	$cre(\mathbf{R}^{2,4})$	urocanate hydratase	hutU	-5.28	1.4E-02
M5005_Spy1773	$(R^{2,4})$	formiminotetrahydrofolate cyclodeaminase	<i>n</i> -	-7.25	3.5E-03
M5005_Spy1774	$(R^{2,4})$	formatetetrahydrofolate ligase	fhs.2	-7.37	2.1E-03
M5005_Spy1775	$(R^{2,4})$	hypothetical cytosolic protein		-6.67	8.6 E-03
M5005_Spy1776	$(R^{2,4})$	amino acid permease		-7.13	3.5E-03
M5005_Spy1777	$(\mathbb{R}^{2,4})$	histidine ammonia-lyase	hutH	-6.67	2.4E-03
M5005 Spy1778	(R^2)	formiminoglutamase	hutG	-1.82	6.1E-03
M5005_Spy1780	(11)	SSU ribosomal protein S2P	rpsB	1.79	3.3E-04
M5005_Spy1780 M5005_Spy1781		protein translation elongation factor Ts	трзы	1.72	5.7E-04
M5005_Spy1782		neutral endopeptidase	pepO	-1.38	2.1E-04
M5005_Spy1783		trehalose-6-phosphate hydrolase	treC	-1.55	3.4E-04
M5005_Spy1784		PTS system, Trehalose-specific IIBC component	treB	-0.95	3.6E-02
M5005_Spy1786		transcriptional regulator, MarR family		2.52	3.3E-04
M5005_Spy1787		glyoxalase family protein		-1.05	7.8E-03
M5005_Spy1789		anaerobic ribonucleoside-triphosphate reductase	nrdG	1.44	2.4E-03
M5005 Spy1790		acetyltransferase		2.23	1.4E-02
M5005_Spy1790 M5005_Spy1792		hypothetical protein		1.02	2.5E-03
M5005_Spy1797		hypothetical cytosolic protein	13	-1.38	1.2E-04
M5005_Spy1798		suppressor of clpP & X SpxA homolog, allele 2	spxA2	-2.61	2.3E-04
M5005_Spy1802		holliday junction DNA helicase	ruvA	-1.19	6.1E-04
M5005_Spy1803		multidrug resistance protein B	lmrP	-1.12	1.7E-03

M5005_Spy1806		hypothetical cytosolic protein		1.45	3.8E-03
M5005_Spy1810		hypothetical membrane spanning protein		1.47	6.0E-03
M5005_Spy1811		hypothetical membrane spanning protein		2.32	1.0E-02
M5005_Spy1812		hypothetical membrane spanning protein		1.18	1.1E-02
M5005_Spy1815		LSU ribosomal protein L32P	rpmF	2.49	1.2E-03
M5005_Spy1821		hypothetical protein		1.20	3.4E-03
M5005_Spy1823		integral membrane protein		-1.65	1.0E-05
M5005_Spy1830		transcriptional regulator, TetR family		1.12	7.0E-03
M5005_Spy1831		SSU ribosomal protein S4P	rpsD	1.43	2.5E-03
M5005_Spy1834		hypothetical protein		1.49	1.2E-03
M5005_Spy1840	cre	tRNA (5-methylaminomethyl-2-thiouridylate)-methyltransferase	trmU	1.32	2.4E-02
M5005_Spy1841	(A^3)	L-serine dehydratase	sdhB	3.26	1.1E-03
M5005_Spy1842		L-serine dehydratase	sdhA	3.50	5.8E-04
M5005_Spy1843	(R^2)	transglycosylase SLT domain family protein		2.76	1.2E-04
		CDP-diacylglycerolglycerol-3-phosphate 3-			
M5005_Spy1847		phosphatidyltransferase		-0.96	1.4E-02
M5005_Spy1849		zinc protease		-0.93	9.2E-03
M5005_Spy1851		hyaluronan synthase	hasA	1.30	7.9E-04
M5005_Spy1853	cre	UTPglucose-1-phosphate uridylyltransferase	hasC	1.24	2.2E-03
M5005_Spy1862	cre	ABC transporter, permease protein		-1.42	4.6E-05

^acre: presence of *cre* and *cre2* sites based on Debroy et al. ¹; (ccpA): identifed as CcpA regulated in M1T1 transcriptome study of CcpA regulon; activated (A) or repressed (R) based on Shelburne *et al.* ², Kinkel *et al.* ³, and Debroy *et al.* ¹.

Supplemental Table S2: RNA-Seq results of the mga mutant in THY (WT/ Δmga)

Spy Number	Annotation	Gene	Log ₂ -FC	p value
M5005_Spy0012	hypoxanthine-guanine phosphoribosyltransferase		-1.73	4.0E-05
M5005_Spy0013	cell division protein	ftsH	-1.02	1.3E-03
M5005_Spy0022	phosphoribosylaminoimidazole-succinocarboxamide synthase		-1.89	1.1E-05
M5005_Spy0040	alcohol dehydrogenase	adhA	1.21	5.1E-02
M5005_Spy0041	Na+ driven multidrug efflux pump		-1.16	1.5E-04
M5005_Spy0080	bis(5'-nucleosyl)-tetraphosphatase (asymmetrical)		-1.09	4.9E-03
M5005_Spy0113	transposase		1.47	6.6E-03
M5005_Spy0115	hypothetical protein		1.21	2.2E-05
M5005_Spy0117	transcriptional regulator, LysR family		2.06	1.6E-04
M5005_Spy0118	transcriptional regulator, LysR family		1.32	2.7E-02
M5005_Spy0143	hypothetical protein		2.54	5.2E-03
M5005_Spy0156	metal-dependent hydrolase		1.16	9.1E-03
M5005_Spy0189	hypothetical protein		1.17	4.6E-02
M5005_Spy0212	N-acetylmannosamine-6-phosphate 2-epimerase		1.39	2.0E-03
M5005_Spy0213	ABC Sialic Acid; N-acetylneuraminate-binding protein		1.44	2.9E-03
M5005_Spy0260	putative lipase		-1.04	1.5E-03
M5005_Spy0261	GTP-binding protein		-0.98	1.8E-02
M5005_Spy0297	transposase		2.55	1.5E-02
M5005_Spy0313	riboflavin transporter		-1.20	1.4E-04
M5005_Spy0318	pyruvate formate-lyase activating enzyme	pflC	1.52	8.2E-07
M5005_Spy0319	manganese-dependent inorganic pyrophosphatase	ppaC	0.90	4.8E-05
M5005_Spy0340	L-lactate oxidase	lctO	2.47	2.6E-04
M5005_Spy0341	lactocepin; SpyCEP; IL-8 protease	spyCEP	1.69	4.9E-06
M5005_Spy0344	permease		-1.10	5.8E-04
M5005_Spy0361	phosphoglycerate transporter protein		1.01	7.8E-03
M5005_Spy0383	D-alanyl-D-alanine carboxypeptidase		-1.04	2.7E-03
M5005_Spy0432	acetyl-CoA acetyltransferase		-1.22	4.4E-04
M5005_Spy0456	plasmid stabilization system antitoxin protein		-1.13	1.1E-02
M5005_Spy0469	hypothetical protein		-1.04	4.0E-03
M5005_Spy0471	hydrolase, HAD superfamily		-1.09	3.4E-03
M5005_Spy0494	hypothetical protein		-1.19	4.3E-03
M5005_Spy0518	oligohyaluronate lyase	-	1.58	2.8E-03
M5005_Spy0519	PTS system, N-acetylgalactosamine-specific IID	agaD	1.02	3.6E-02
M5005_Spy0520	PTS system, N-acetylgalactosamine-specific IIC	agaS	1.89	4.9E-05
M5005_Spy0522	unsaturated glucuronyl hydrolase		1.45	8.6E-04
M5005_Spy0549	chorismate mutase		-1.15	3.8E-04
M5005_Spy0585	epuA protein	epuA	-1.20	6.4E-03
M5005_Spy0660	fructose repressor	fruR	1.06	1.8E-02
M5005_Spy0667	exotoxin type C precursor		2.07	7.1E-03
M5005_Spy0668	IgG-degrading protease	mac	1.62	6.1E-04
M5005_Spy0714	hypothetical cytosolic protein		-1.24	2.5E-04
M5005_Spy0742	hypothetical protein		-1.97	2.0E-05
M5005_Spy0743	ABC transporter substrate-binding protein hypothetical protein		-1.59	1.8E-05
M5005_Spy0744	••		-1.55 1.26	2.6E-04
M5005_Spy0745	ABC transporter permease protein ABC transporter ATP-binding protein		-1.26	3.8E-05
M5005_Spy0746		nta A	-1.47 1.50	5.9E-07
M5005_Spy0780	PTS system, Mannose/Fructose family IIA component	ptsA		6.8E-04
M5005_Spy0781	PTS system, Mannose/Fructose family IIB component PTS system, Mannose/Fructose family IIC component	ptsB ptsC	1.30 1.03	7.2E-05 1.4E-03
M5005_Spy0782		ptsC	0.89	
M5005_Spy0786	iron(III)-binding protein		0.89 -1.04	2.0E-03
M5005_Spy0793	Xaa-His dipeptidase			4.8E-02
M5005_Spy0797	hypothetical protein IEN recognes hinding factor 1		-2.85 3.53	1.4E-02 2.2E.04
M5005_Spy0798	IFN-response binding factor 1	$f_{0}1V$	-3.53 1.00	2.2E-04
M5005_Spy0824	2-amino-4-hydroxy-6-hydroxymethyldihydropteridine pyrophosphokinase	folK	-1.00 1.20	3.8E-04
M5005_Spy0834	Zn-dependent alcohol dehydrogenases and related dehydrogenases		1.20	2.6E-0

M5005_Spy0853	short chain dehydrogenase		0.96	7.1E-03
M5005_Spy0858	xanthine phosphoribosyltransferase	xpt	-2.16	7.4E-05
M5005_Spy0859	xanthine permease		-1.81	6.1E-06
M5005_Spy0914	phage transcriptional repressor		-1.32	1.4E-04
M5005_Spy0920	UDP-N-acetylmuramoylpentapeptide-lysine N(6)-alanyltransferase		-1.70	8.6E-05
M5005_Spy0921	ABC transporter ATP-binding protein		-1.54	8.4E-05
M5005_Spy0932	luciferase-like monooxygenase		0.94	1.6E-03
M5005_Spy0933	probable NADH-dependent flavin oxidoreductase		1.00	6.8E-04
M5005_Spy0934	lipoate-protein ligase A		1.07	5.1E-04
M5005_Spy0978	hypothetical protein		-1.17	6.3E-04
M5005_Spy0985	phnA protein		-1.39	6.1E-07
M5005_Spy0986	glucosaminefructose-6-phosphate aminotransferase (isomerizing)	glmS	-1.02	5.0E-04
M5005_Spy1059	ABC Maltodextrin permease protein	malF	0.94	2.8E-04
M5005_Spy1062	maltodextrose utilization protein	malA	1.49	4.1E-02
M5005_Spy1063	ABC Cyclomaltodextrin permease protein	malD	1.54	2.7E-02
M5005_Spy1064	ABC Cyclomaltodextrin permease protein	malC	1.38	2.3E-02
M5005_Spy1065	alpha-amylase	amyA	1.46	1.1E-02
M5005_Spy1066	neopullulanase/cyclomaltodextrinase/maltogenic alpha-amylase	amyB	1.74	7.3E-03
M5005_Spy1067	ABC Cyclomaltodextrin -binding protein	malX	2.52	1.9E-05
M5005_Spy1076	Putative Bicarbonate transporter	glnH	-1.31	2.3E-03
M5005_Spy1077	Putative Bicarbonate transporter	glnQ.2	-1.04	1.3E-02
M5005_Spy1082	PTS system, Cellobiose-specific IIB component	8m Q.2	1.16	3.0E-02
M5005_Spy1082	PTS system, Cellobiose-specific IIA component		1.00	1.1E-02
M5005_Spy1084	outer surface protein		1.02	1.8E-02
M5005_Spy1004	beta-glucosidase	bglA.2	1.46	2.3E-03
M5005_Spy1083	hypothetical protein	UgIA.2	1.15	1.5E-02
M5005_Spy1055	oxalate/formate antiporter		-1.16	3.9E-06
	COME operon protein 1		-1.32	2.0E-02
M5005_Spy1148	dihydroorotate dehydrogenase	nurD.	-1.32 -1.19	4.5E-03
M5005_Spy1165	phage protein	pyrD	-1.19 -1.11	
M5005_Spy1194			1.14	3.5E-03
M5005_Spy1235	phosphoglucomutase			9.2E-05
M5005_Spy1261	radical SAM family enzyme		-1.25	2.9E-03
M5005_Spy1274	acetyltransferase		0.90	3.7E-02
M5005_Spy1275	arginine deiminase	arcA	1.75	4.2E-03
M5005_Spy1284	cytochrome c-type biogenesis protein	ccdA	1.31	2.9E-02
M5005_Spy1299	hypothetical protein	1 1	-2.03	3.1E-03
M5005_Spy1314	hyaluronoglucosaminidase	hyl	0.85	1.0E-02
M5005_Spy1315	transcriptional regulator, GntR family		1.18	3.0E-04
M5005_Spy1316	hypothetical protein		1.35	1.9E-03
M5005_Spy1317	alpha-mannosidase		1.16	5.5E-03
M5005_Spy1340	DNA-directed RNA polymerase omega chain		-1.50	1.7E-05
M5005_Spy1347	D-beta-hydroxybutyrate dehydrogenase		-0.97	9.1E-03
M5005_Spy1376	transaldolase		1.00	5.2E-02
M5005_Spy1377	trans-acting positive regulator		1.09	5.7E-02
M5005_Spy1387	aldo/keto reductase family		1.35	2.4E-05
M5005_Spy1397	galactose-6-phosphate isomerase lacB subunit	lacB.1	1.10	5.0E-02
M5005_Spy1477	guanine-hypoxanthine permease		-1.29	2.1E-04
M5005_Spy1488	biotin carboxyl carrier protein of acetyl-CoA	accB	-1.20	1.6E-02
M5005_Spy1489	3-oxoacyl-[acyl-carrier-protein] synthase	fabF	-1.11	3.0E-02
M5005_Spy1491	malonyl-CoA-[acyl-carrier-protein] transacylase	fabD	-1.21	4.9E-03
M5005_Spy1492	enoyl-[acyl-carrier protein] reductase (NADH)	fabK	-1.13	1.8E-02
M5005_Spy1569	formate acetyltransferase	pfl	1.06	8.6E-04
M5005_Spy1572	hypothetical membrane spanning protein		-1.36	1.4E-04
M5005_Spy1587	uridine phosphorylase	udp	1.12	3.3E-05
M5005_Spy1589	transcriptional regulator, Cathelicidin resistance	crgR	-1.13	7.0E-03
M5005_Spy1618	competence-specific sigma factor	comX.2	-2.41	2.1E-02
M5005_Spy1628	ABC transporter ATP-binding protein		1.40	3.7E-03
M5005_Spy1629	lantibiotic transport ATP-binding protein	salX	1.53	4.3E-03
M5005_Spy1630	serine (threonine) dehydratase	salB	2.13	6.5E-06
M5005_Spy1631	lantibiotic salivaricin A	salA	1.81	3.4E-02

M5005 Spy1632	6-phospho-beta-galactosidase	lacG	2.06	2.7E-03
M5005_Spy1632 M5005_Spy1633	PTS system, Lactose-specific IIBC component	lacE	2.12	4.3E-03
_ 1,	PTS system, Lactose-specific IIA component	lacE	2.12	4.3E-03 1.4E-02
M5005_Spy1634			2.06	
M5005_Spy1635	tagatose-bisphosphate aldolase	lacD.2		5.4E-03
M5005_Spy1636	tagatose-6-phosphate kinase	lacC.2	2.52	1.5E-03
M5005_Spy1637	galactose-6-phosphate isomerase lacB subunit	lacB.2	2.24	4.3E-03
M5005_Spy1638	galactose-6-phosphate isomerase lacA subunit	lacA.2	1.99	2.1E-02
M5005_Spy1644	DNA integration/recombination/invertion protein		-3.39	6.7E-04
M5005_Spy1649	hypothetical membrane spanning protein		-1.00	2.9E-03
M5005_Spy1682	multiple sugar transport ATP-binding protein	msmK	1.01	1.2E-03
M5005_Spy1702	mitogenic exotoxin Z	smeZ	-1.15	1.8E-04
M5005_Spy1705	dipeptide transport system permease protein	dppB	-1.09	6.9E-03
M5005_Spy1709	hypothetical protein		-1.47	7.9E-04
M5005_Spy1714	Fn-binding surface protein; Mga-regulated	fba	3.20	4.5E-06
M5005_Spy1715	C5A peptidase; Mga-regulated	scpA	3.43	2.1E-07
M5005_Spy1717	transposase		-2.57	8.1E-06
M5005_Spy1718	secreted inhibitor of complement; Mga-regulated	sic1.0	4.63	6.2E-09
M5005_Spy1719	M protein; Mga-regulated	emm1	9.08	8.6E-16
M5005_Spy1720	Multi-virulence gene regulator Mga	mga	4.60	2.8E-15
M5005_Spy1721	hypothetical protein		-2.78	6.4E-04
M5005 Spy1733	hypothetical protein		-2.54	4.2E-03
M5005 Spy1734	streptopain SpeB inhibitor	speI	-2.94	3.1E-03
M5005 Spy1735	streptococcal pyrogenic exotoxin B	speB	-2.73	2.1E-04
M5005 Spy1738	Secreted DNase-Streptodornase-chromosomal	spd	0.98	2.2E-03
M5005 Spy1750	hypothetical protein	•	-1.14	2.5E-04
M5005 Spy1758	probable dipeptidase B		1.09	5.3E-04
M5005 Spy1759	transcriptional regulator, MutR family		-1.10	7.0E-03
M5005 Spy1762	10 kDa chaperonin	groES	-1.04	7.1E-03
M5005 Spy1779	transcriptional regulator, LuxR family	8	1.55	1.5E-03
M5005 Spy1798	suppressor of clpP & X SpxA homolog, allele 2	spxA2	-1.86	3.0E-04
M5005_Spy1809	bacteriocin	uviB	-1.13	1.5E-03
M5005_Spy1805	hypothetical membrane spanning protein	u, ib	-1.01	3.0E-03
M5005_Spy1812 M5005_Spy1828	phage infection protein		-1.71	6.7E-07
1415005_Spy 1626	phage infection protein		-1./1	0.715-07

Supplemental Table S3: RNA-Seq results of the mga mutant in C Media (WT/ Δmga)

Spy Number	Annotation	Gene	Log ₂ -FC	p value
M5005_Spy0009	hypothetical protein		1.61	4.9E-02
M5005_Spy0030	phosphoribosylaminoimidazole carboxylase catalytic subunit	purE	1.09	2.9E-02
M5005_Spy0036	protein tyrosine phosphatase		-0.93	4.0E-03
M5005_Spy0091	comG operon protein 6	comYD	1.45	1.4E-02
M5005_Spy0106	transcriptional regulator; Regulator of protein F	rofA	-1.09	1.8E-04
M5005_Spy0113	transposase		1.32	1.8E-02
M5005_Spy0129	V-type ATP synthase subunit C	ntpC	-1.78	6.7E-03
M5005_Spy0130	V-type sodium ATP synthase subunit F	ntpF	-1.44	8.8E-03
M5005_Spy0131	V-type sodium ATP synthase subunit A	ntpA	-1.30	1.4E-02
M5005_Spy0132	V-type sodium ATP synthase subunit B	ntpB	-1.50	1.2E-02
M5005_Spy0133	V-type sodium ATP synthase subunit D	ntpD	-1.57	8.5E-03
M5005_Spy0136	adenylosuccinate synthetase	purA	-1.11	2.9E-05
M5005_Spy0139	NAD glycohydrolase	nga/spn	1.04	4.6E-02
M5005_Spy0140	NAD glycohydrolase inhibitor	ifs	1.14	4.9E-02
M5005_Spy0141	streptolysin O	slo	1.12	2.9E-02
M5005_Spy0143	hypothetical protein		3.35	2.1E-03
M5005_Spy0157	glycine betaine transport ATP-binding protein	opuAA	1.23	4.8E-06
M5005_Spy0167	transposase		1.58	2.9E-02
M5005_Spy0183	hypothetical membrane associated protein		0.91	5.4E-03
M5005_Spy0202	hypothetical protein		-1.34	6.9E-05
M5005_Spy0247	D-alanyl-D-alanine carboxypeptidase		0.99	1.3E-04
M5005_Spy0257	transposase		-1.55	2.2E-02
M5005_Spy0279	hypothetical protein	lemA	-0.91	6.3E-04
M5005_Spy0296	acylphosphatase		-1.04	2.9E-04
M5005_Spy0385	67 kDa Myosin-crossreactive antigen		-1.32	6.1E-05
M5005_Spy0403	hypothetical protein		0.92	4.6E-02
M5005_Spy0422	NAD(P)H-dependent quinone reductase		-1.11	4.8E-05
M5005_Spy0423	Xaa-Pro dipeptidase	pepQ	-1.23	4.7E-05
M5005_Spy0477	hypothetical membrane spanning protein	scfB	-0.97	2.8E-04
M5005_Spy0478	hypothetical membrane spanning protein	scfA	-1.08	6.0E-04
M5005_Spy0486	hypothetical protein		-1.08	2.6E-04
M5005_Spy0522	unsaturated glucuronyl hydrolase		0.98	5.0E-03
M5005_Spy0527	4-hydroxy-2-oxoglutarate aldolase/	kgdA	1.03	5.4E-04
M5005_Spy0555	hypothetical cytosolic protein		-1.09	2.7E-04
M5005_Spy0560	transcriptional regulator		-0.94	1.4E-02
M5005_Spy0562	streptolysin S precursor	sagA	1.33	7.4E-04
M5005_Spy0564	streptolysin S biosynthesis protein	sagC	0.99	6.8E-04
M5005_Spy0565	streptolysin S biosynthesis protein	sagD	1.09	1.5E-04
M5005_Spy0567	streptolysin S biosynthesis protein	sagF	1.46	1.1E-04
M5005_Spy0568	streptolysin S export ATP-binding protein	sagG	1.30	2.5E-05
M5005_Spy0569	streptolysin S export transmembrane protein	sagH	1.09	3.4E-04
M5005_Spy0570	streptolysin S export transmembrane protein	sagI	1.42	1.3E-04
M5005_Spy0571	endonuclease/exonuclease/phosphatase family protein		1.21	1.1E - 06
M5005_Spy0598	large-conductance mechanosensitive channel	mscL	-0.88	2.6E-02
M5005_Spy0615	pore forming protein	ebsA	-1.03	9.6E-04
M5005_Spy0616	ferredoxin		-2.15	1.1E-03
M5005_Spy0641	aspartate carbamoyltransferase	pyrB	0.90	5.7E-03
M5005_Spy0651	Heme-utilization cell surface protein	hupY	2.28	4.6E-09
M5005_Spy0652	Heme-oxygenase cytosolic protein	hupZ	2.16	5.2E-08
M5005_Spy0668	IgG-degrading protease	mac	1.37	1.2E-02
M5005_Spy0679	GTP pyrophosphokinase		-1.01	3.0E-04
M5005_Spy0703	orotidine 5'-phosphate decarboxylase	pyrF	1.15	9.2E-03
M5005_Spy0708	uracil-DNA glycosylase	ung	-0.86	4.1E-02
M5005_Spy0715	SSU ribosomal protein S1P		1.28	9.4E-04
M5005 Spy0720	putative exfoliative toxin		0.86	1.1E-02

M5005_Spy0753	dihydrolipoamide acetyltransferase pyruvate dehydrogenase complex	acoC	-0.85	8.2E-04
M5005_Spy0754	hypothetical protein		-1.50	5.8E-02
M5005_Spy0755	dihydrolipoamide dehydrogenase	acoL	-1.16	2.9E-05
M5005_Spy0794	tRNA (5-carboxymethylaminomethyl-2-thiouridylate) synthase	thdF	1.10	2.7E-04
M5005_Spy0820	tetrahydropholate synth. /dihydrofolate synthase	folC.1	-0.98	2.0E-05
M5005 Spy0821	tetrahydropholate synth., GTP cyclohydrolase I	folE	-0.95	6.7E-05
M5005 Spy0822	tetrahydropholate synth., dihydropteroate synthase	folP	-0.92	5.7E-03
M5005 Spy0824	tetrahydropholate synth., pyrophosphokinase	folK	-0.87	1.4E-3
M5005 Spy0879	hypothetical protein		-1.52	7.9E-05
M5005_Spy0880	conserved membrane protein		-2.45	2.5E-05
M5005 Spy0881	hypothetical cytosolic protein		-1.93	5.2E-05
M5005_Spy0890	D-lactate dehydrogenase	ddh	1.38	5.2E-03
M5005_Spy0090	2-(5"-triphosphoribosyl)-3'-dephosphocoenzyme-A synthase	ddii	1.05	3.0E-02
M5005_Spy0056 M5005_Spy1005	phage protein		-1.47	2.1E-02
M5005_Spy1005 M5005_Spy1035	phage protein		1.44	6.0E-02
	Putative Bicarbonate transporter	glnH	-0.85	3.2E-02
M5005_Spy1076	-	-		
M5005_Spy1077	Putative Bicarbonate transporter	glnQ.2	-1.04	1.0E-02
M5005_Spy1093	hypothetical protein	1.4	-1.16	2.0E-04
M5005_Spy1145	superoxide dismutase	sodA	-0.93	5.7E-03
M5005_Spy1169	phage-encoded streptodornase; secreted DNase	spd3	2.35	1.0E-06
M5005_Spy1170	hypothetical membrane associated protein		2.39	6.1E-07
M5005_Spy1199	phage protein		-1.87	9.6E-03
M5005_Spy1211	phage protein		1.19	5.0E-02
M5005_Spy1213	phage protein		1.28	7.5E-03
M5005_Spy1227	hypothetical protein		1.06	6.2E-05
M5005_Spy1228	DNA repair protein	recN	0.99	6.7E-03
M5005_Spy1229	arginine repressor	argR1	0.99	3.2E-02
M5005_Spy1230	hemolysin		0.92	3.2E-02
M5005_Spy1237	arginine transport ATP-binding protein	artP	1.36	7.8E-04
M5005_Spy1238	arginine transport system permease protein	artQ	1.40	2.1E-04
M5005_Spy1239	hypothetical cytosolic protein		1.14	1.5E-02
M5005_Spy1285	CRISPR2-associated protein Cas2	cas2	1.11	4.9E-04
M5005_Spy1287	CRISPR2-associated protein Cas4	cas4	1.08	2.6E-04
M5005 Spy1289	CRISPR2-associated protein Cds1	cds1	1.06	4.0E-07
M5005_Spy1297	phospho-2-dehydro-3-deoxyheptonate aldolase		1.18	3.0E-03
M5005_Spy1320	regulatory protein	recX	-1.55	1.4E-05
M5005_Spy1323	transposase		0.96	9.6E-03
M5005_Spy1325	ribosome-associated factor Y		-1.16	5.8E-04
M5005_Spy1369	hypothetical protein		1.18	2.4E-02
M5005 Spy1372	ABC transporter permease protein	proB	1.11	2.2E-04
M5005_Spy1382	hypothetical membrane associated protein	ргов	-1.40	1.4E-03
M5005_Spy1400	PTS system, Galactose-specific IIB component		-2.47	2.4E-03
M5005_Spy1400	PTS system, Galactose-specific IIA component		-2.66	2.9E-04
M5005_Spy1407	streptococcal secreted esterase Sse	SSA	1.81	6.5E-04
M5005_Spy1407 M5005_Spy1415	-	sse sdaD2	2.22	8.7E-06
	phage-encoded streptodornase; Secreted DNase	SuaD2		
M5005_Spy1440	terminase large subunit		1.23	1.2E-02
M5005_Spy1441	phage terminase small subunit		1.12	3.4E-02
M5005_Spy1514	universal stress protein family		-1.22	1.2E-03
M5005_Spy1525	conserved hypothetical protein	siaD	1.09	2.2E-03
M5005_Spy1526	ABC transporter, ATP-binding protein, putative	siaC	1.33	7.5E-07
M5005_Spy1527	ABC transporter, ATP-binding protein, putative	siaB	1.39	1.2E-03
M5005_Spy1528	iron compound ABC transporter, ATP-binding protein	siaA	1.44	9.5E-04
M5005_Spy1529	iron compound ABC transporter, permease protein	shp	1.22	6.6E-05
M5005_Spy1530	iron compound ABC transporter, substrate-binding protein	shr	1.27	4.4E-07
M5005_Spy1559	thioredoxin	trx	-0.96	9.7E-04
M5005_Spy1590	SSU ribosomal protein S14P	rpsN2	-1.22	7.5E-03
M5005_Spy1622	type I restriction-modification system specificity subunit	hsdS	1.20	9.4E-07
M5005_Spy1714	Fn-binding surface protein; Mga-regulated	fba	4.79	8.7E-08
M5005_Spy1715	C5A peptidase precursor; Mga-regulated	scpA	5.04	2.9E-10
_ 10		1		

M5005_Spy1716	transposase		-1.07	1.3E-04
M5005_Spy1718	secreted inhibitor of complement; Mga-regulated	sic1.0	5.53	2.1E-09
M5005_Spy1719	M protein; Mga-regulated	emm1	10.06	8.1E-17
M5005_Spy1720	Multi-virulence gene regulator Mga	mga		4.2E-17
M5005_Spy1721	hypothetical protein		-5.00	2.4E-08
M5005_Spy1727	ABC transporter ATP-binding protein		0.99	1.0E-02
M5005_Spy1731	gene regulated by Mga; Mga-regulated	grm	3.85	7.7E-05
M5005_Spy1733	hypothetical protein		1.45	5.2E-02
M5005_Spy1736	hypothetical protein		-1.31	3.0E-02
M5005_Spy1738	Secreted DNase-Streptodornase-chromosomal	spd	1.03	3.2E-04
M5005_Spy1739	hypothetical protein		-3.02	3.2E-05
M5005_Spy1754	translation initiation inhibitor		-0.98	4.6E-02
M5005_Spy1768	peroxiredoxin reductase (NAD(P)H)	ahpC	-0.92	1.1E-03
M5005_Spy1774	formatetetrahydrofolate ligase	fhs.2	1.84	2.5E-02
M5005_Spy1775	hypothetical cytosolic protein		1.59	5.0E-02
M5005_Spy1776	amino acid permease		1.83	2.8E-02
M5005_Spy1777	histidine ammonia-lyase	hutH	2.16	9.1E-03
M5005_Spy1778	formiminoglutamase	hutG	1.01	9.6E-03
M5005_Spy1798	suppressor of clpP & X SpxA homolog, allele 2	spxA2	-1.07	1.7E-02
M5005_Spy1823	integral membrane protein		-0.88	3.5E-04
M5005_Spy1843	transglycosylase SLT domain family protein		1.33	4.5E-03

Supplemental Table S4: PCR primers used in this study

Target	Primer Name	Sequence (5'-3') ^{† ^}	Reference
qPCR			
gyrA	gyrA M1 RT-L	CGACTTGTCTGAACGCCAAAGT	4
0,0	gyrA M1 RT-R	ATCACGTTCCAAACCAGTCAAAC	4
emm	emm1 RT-L	ACTCCAGCTGTTGCCATAACAG	5
omm.	emm1 RT-R	GAGACAGTTACCATCAACAGGTGAA	5
sclA	sclA M1 RT-L	TGCTGACAAAGAAGCTAACCAAAC	5
5611	sclA M1 RT-L	CGTCTGTGGTTGTTGGCTACAG	5
fruA	fruA M1 RT-L	ACCGGGCTTAGTAGCTGGT	6
,	fruA M1 RT-R	ACTTCTCCTCCTGCTGCAA	6
sdaD2	sdaD2 M1T1 RT-L	AGCAAACCAGAGCCAAGTG	This study
54422	sdaD2 M1T1 RT-R	CACATTTTTGGGGCTATGC	This study
speB	speB M1 RT-L	GGTAAAGTAGGCGGACATGCC	7
5702	speB M1 RT-R	CACCCCAACCCCAGTTAACA	7
salB	salB M1T1 RT-L	TGGTGTTTTAGCAGCATTTT	This study
	salB M1T1 RT-R	AATCTCCGCCTGAAACAAT	This study
mac	mac M1T1 RT-L	ACGGCTACCGCCTTAGTCT	This study
	mac M1T1 RT-R	ACGGCGTCAAAAATACCAC	This study
opuAA	opuAA M1 RT-L	TGATTTGCAAGACAGCATGA	7
•	opuAA M1 RT-R	CATCAAAGCAATCCGATCAC	7
ska	ska M1T1 RT-L	TCGAAACGGCAAGGTCTAC	This study
	ska M1T1 RT-R	GCGCACATGTCCCTTTAAC	This study
sagC	sagC M1 RT-L	TGCTGATGCTTGAGGATGAC	3
	sagC M1 RT-R	GCGCCATAAAACGGAAATTA	3
glnH	glnH M1 RT-L	TGAGGGCATTTCTACTGGTG	This study
	glnH M1 RT-R	GCAGCTAAGCCATTGTTGAA	This study
slo	slo M1 RT-L	TTGTTGAGGATAATGTAAGAATGTTTAG	3
	slo M1 RT-R	TCCTGGCTTGCAACTGATTG	3
PCR			
mga	Mga InIn F	ccc GGATCC CAGTGGAGAGAACTAAAATT	This study
mga	Mga Inin I Mga Inin R	ccc GGATCC TAATTCTTCCAAAGAGTTGA	This study This study
M13	1201	AACAGCTATGACCATGATTACG	Genewiz
1.110	1211	GTTGTAAAACGACGCCAGT	Genewiz
aad9	SPR1	cccCTCGAGAATAAAGCCCGCTTCGGCG	8
aaas	SPR2	cccACTAGTAAGGACTTAAAAGAGCGTATTG	8
	5. NE	TOTAL TARINGGIOT TARINGGIO GOTTI I G	

[†] Bold type denotes restriction site

 $^{^{\}updayscript{\wedge}}$ Lowercase letters denote nucleotides not complementary to target DNA

Supplemental References

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