

Supplemental Table S1. Proteins found in the supernatant of *S. aureus* Newman (wt) and the mutant Newman $\Delta secDF$ (*secDF*) and their predicted localization.

NWMN	Protein name	Gene	wt				<i>secDF</i>				σ wt	σ <i>secDF</i>	P	SignalP [1]	Sec SP [2]	No SP(*) [2]	SecretomeP [3]	Localization [4,5]	
			wt I	wt II	wt III	wt IV	<i>secDF</i> I	<i>secDF</i> II	<i>secDF</i> III	<i>secDF</i> IV									
			fold change (log 2)																
0002	DNA polymerase III subunit beta	<i>dnaN</i>	0	0.16			0.01	-0.13			0.08	-0.06						Cytoplasm	
0014	50S ribosomal protein L9	<i>rplI</i>	0	-0.51			-0.61	-0.95			-0.26	-0.78					Yes	Cytoplasm	
0030	Probable tRNA-dihydrouridine synthase	<i>dus</i>	0	-0.12			-0.99	-0.32			-0.06	-0.66						Cytoplasm	
0055	Immunoglobulin G-binding protein A	<i>spa</i>	0	-0.37	0	-0.29	-3.01	-3.10	-2.40	-2.52	-0.16	-2.76	***	Yes	Yes		Yes	Cell wall	Sortase substrate
0113	Putative aldehyde dehydrogenase	<i>aldA</i>	0	0.16	0	-0.24	-0.15	-0.78	-0.46	-0.56	-0.02	-0.49	*					Cytoplasm	
0129	Ornithine aminotransferase 2	<i>argD</i>	0	0.05	0	-0.56	-0.34	-0.44	-0.68	-1.10	-0.13	-0.64						Cytoplasm	
0166	Staphylocoagulase	<i>coa</i>	0	0.14	0	0.08	-4.33	-3.91	-2.61	-3.26	0.06	-3.53	**	Yes	Yes		Yes	Extracellular	
0172	Putative zinc metalloprotease	<i>0172</i>	0	0.10			-0.67	-0.65			0.05	-0.66	*				Yes	Membrane	
0189	2-C-methyl-D-erythritol 4-phosphate cytidylyl-transferase 2	<i>ispD2</i>	0	0.17			-0.06	-0.35			0.08	-0.21						Unknown	
0210	Glycyl-glycine endopeptidase	<i>lytM</i>	0	0.03	0	-0.35	2.00	2.06	1.72	1.34	-0.08	1.78	***	Yes	Yes		Yes	Extracellular	
0249	5'-nucleotidase, lipoprotein e (P4)	<i>0249</i>	0	-0.27	0	0.22	-2.22	-2.16	-1.58	-1.75	-0.01	-1.93	***	Yes	Yes		Yes	Extracellular	
0262	Lipase 2	<i>geh</i>	0	-0.18	0	-0.19	-2.93	-2.92	-1.63	-1.75	-0.09	-2.31	**	Yes	Yes		Yes	Extracellular	
0357	30S ribosomal protein S6	<i>rpsF</i>	0	-0.14	0	-0.24	-0.36	-0.22	0.20	-0.09	-0.09	-0.12						Cytoplasm	
0371	Alkyl hydroperoxide reductase subunit F	<i>ahpF</i>	0	-0.19	0	-0.66	-0.47	-0.70	-0.86	-1.08	-0.21	-0.78	*			x		Cytoplasm	
0372	Alkyl hydroperoxide reductase subunit C	<i>ahpC</i>	0	0.13	0	-0.34	-0.05	-0.37	-0.07	-0.39	-0.05	-0.22				x	Yes	Cytoplasm	
0375	Conserved hypothetical protein	<i>0375</i>	0	-0.32			-1.06	-1.05			-0.16	-1.06					Yes	Cytoplasm	
0377	Conserved hypothetical protein	<i>0377</i>	0	0.00	0	-0.46	-0.59	-0.81	-0.50	-0.52	-0.12	-0.60	*				Yes	Cytoplasm	
0380	Inosine-5'-monophosphate dehydrogenase	<i>guaB</i>	0	-0.06	0	-0.01	0.02	-0.05	-0.40	-0.57	-0.02	-0.25				x		Cytoplasm	
0381	GMP synthase [glutamine-hydrolyzing]	<i>guaA</i>	0	-0.20			-0.31	-0.63			-0.10	-0.47				x		Cytoplasm	
0429	N-acetylmuramoyl-L-alanine amidase	<i>sle1</i>	0	-0.38	0	0.51	-1.45	-0.92	-0.12	-1.05	0.03	-0.89	*	Yes	Yes		Yes	Cell wall, non-cov. LysM domain [6]	
0443	Conserved hypothetical protein	<i>0443</i>	0	-0.07			-0.27	-0.39			-0.03	-0.33						Unknown	
0453	Methionyl-tRNA synthetase	<i>metS</i>	0	-0.33	0	-0.05	-0.87	-1.20	-0.35	-0.74	-0.09	-0.79	*					Cytoplasm	
0463	Ribose-phosphate pyrophosphokinase	<i>prs</i>	0	0.36			0.37	-0.04			0.18	0.17						Cytoplasm	
0464	50S ribosomal protein L25	<i>rplY</i>	0	0.07	0	-0.56	0.37	0.19	-0.21	-0.01	-0.12	0.08						Cytoplasm	
0472	Hypoxanthine-guanine phosphoribosyl-transferase	<i>hpt</i>	0	0.48	0	-0.65	0.11	0.44	2.31	1.36	-0.04	1.06						Cytoplasm	
0475	Cysteine synthase homolog	<i>cysK</i>	0	0.06	0	-0.08	-0.37	-0.45	-0.29	-0.44	-0.01	-0.39	***					Cytoplasm	
0479	Lysyl-tRNA synthetase	<i>lysS</i>	0	-0.25			-0.56	-0.42			-0.12	-0.49						Cytoplasm	
0481	Pyridoxal biosynthesis lyase	<i>pdxS</i>	0	-0.28	0	-1.02	-0.44	-0.39	-0.32	-0.85	-0.33	-0.50						Cytoplasm	
0482	Glutamine amidotransferase subunit	<i>pdxT</i>	0	-0.47			-0.26	-0.18			-0.23	-0.22						Cytoplasm	
0487	ATP-dependent Clp protease, ATP-binding subunit ClpC	<i>clpC</i>	0	0.40	0	-0.17	-0.05	0.02	-0.28	-0.44	0.06	-0.19				x		Cytoplasm	
0490	Glutamyl-tRNA synthetase	<i>gltX</i>	0	0.61	0	-0.54	0.01	-0.17	-0.44	-0.91	0.02	-0.38				x		Cytoplasm	
0498	Transcription antitermination protein	<i>nusG</i>	0	0.13			-0.32	-0.39			0.06	-0.35						Cytoplasm	
0499	50S ribosomal protein L11	<i>rplK</i>	0	0.15	0	-0.26	0.21	0.23	0.13	0.31	-0.03	0.22						Cytoplasm	
0500	50S ribosomal protein L1	<i>rplA</i>	0	0.23	0	-0.25	0.12	-0.16	0.06	-0.14	0.00	-0.03					Yes	Cytoplasm	
0501	50S ribosomal protein L10	<i>rplJ</i>	0	0.06	0	-0.48	0.09	-0.04	0.00	0.04	-0.10	0.02						Cytoplasm	

NWMN	Protein name	Gene	fold change (log 2)										P	SignalP [1]	Sec SP [2]	No SP(*) [2]	SecretomeP [3]	Localization [4,5]		
			wt I	wt II	wt III	wt IV	secDF I	secDF II	secDF III	secDF IV	σ wt	σ secDF								
0502	50S ribosomal protein L7/L12	<i>rplL</i>	0	0.06	0	-0.36	0.30	-0.02	-0.07	-0.12	-0.07	0.02							Cytoplasm	
0504	DNA-directed RNA polymerase subunit beta	<i>rpoB</i>	0	0.03			-0.30	-0.29				0.01	-0.30	*					Cytoplasm	
0505	DNA-directed RNA polymerase subunit beta'	<i>rpoC</i>	0	0.16	0	-0.34	-0.41	-0.49	0.07	-0.48	-0.05	-0.33							Cytoplasm	
0508	30S ribosomal protein S7	<i>rpsG</i>	0	-0.38	0	-0.70	0.42	0.36	-0.40	-0.24	-0.27	0.03							Cytoplasm	
0509	Elongation factor G	<i>fus</i>	0	0.06	0	-0.38	0.33	0.26	-0.63	-0.51	-0.08	-0.14				x			Cytoplasm	
0510	Elongation factor Tu	<i>tufA</i>	0	-0.05	0	-0.21	0.77	0.65	0.67	0.72	-0.07	0.71	***				x		Cytoplasm	
0512	Conserved hypothetical protein	<i>0512</i>	0	-0.12			-0.53	-0.89				-0.06	-0.71						Cytoplasm	
0513	Similar to chaperone protein HchA	<i>hchA</i>	0	-0.27	0	-0.44	0.05	-0.01	-0.38	-0.39	-0.18	-0.18					Yes		Cytoplasm	
0516	Probable branched-chain-amino-acid amino-transferase	<i>ilvE</i>	0	0.11	0	-0.32	0.49	0.02	-0.51	-0.65	-0.05	-0.16							Cytoplasm	
0525	Serine-aspartate repeat-containing protein E	<i>sdrE</i>	0	-0.08	0	-0.44	-0.92	-1.00	-0.81	-1.03	-0.13	-0.94	**	Yes	Pot.			Yes	Cell wall	Sortase substrate
0533	3-hexulose-6-phosphate synthase	<i>0533</i>	0	-0.32	0	-0.52	0.05	-0.16	0.37	-0.09	-0.21	0.04							Cytoplasm	
0550	Conserved hypothetical protein	<i>0550</i>	0	-0.03	0	0.56	0.62	-0.56	0.86	0.71	0.13	0.41							Unknown	
0551	Phosphate acetyltransferase	<i>pta</i>	0	0.02	0	-0.39	-0.21	-0.51	-0.53	-0.43	-0.09	-0.42	*				x		Cytoplasm	
0579	Arginyl-tRNA synthetase	<i>argS</i>	0	0.31			-1.65	-0.87			0.16	-1.26							Cytoplasm	
0588	Transcriptional regulator	<i>sarA</i>	0	0.00			-0.03	-0.26			0.00	-0.15							Cytoplasm	
0655	MarR family regulatory protein	<i>mgrA</i>	0	-0.29			0.01	-0.22			-0.14	-0.10						Yes	Cytoplasm	
0687	Glycerol phosphate lipoteichoic acid synthase	<i>ltaS</i>	0	0.04	0	-0.15	-0.70	-0.69	-0.69	-0.75	-0.03	-0.71	***					Yes	Membrane	
0721	Ribosomal protein S30EA	<i>0721</i>	0	-0.35			-0.67	-0.55			-0.18	-0.61					x		Cytoplasm	
0732	Thioredoxin reductase	<i>trxB</i>	0	0.27			-0.06	-0.15			0.14	-0.11					x		Cytoplasm	
0736	ATP-dependent Clp protease, ATP-binding subunit ClpP	<i>clpP</i>	0	-0.33			-0.36	-0.69			-0.16	-0.53					x		Cytoplasm	
0741	Glyceraldehyde-3-phosphate dehydrogenase 1	<i>gapA</i>	0	0.00	0	-0.14	0.78	0.55	0.43	0.97	-0.03	0.68	**				x		Cytoplasm	
0742	Phosphoglycerate kinase	<i>pgk</i>	0	0.05	0	0.04	0.23	0.08	-0.07	-0.39	0.02	-0.04					x		Cytoplasm	
0743	Triosephosphate isomerase	<i>tpi</i>	0	-0.18	0	-0.45	0.31	0.24	-0.36	0.19	-0.16	0.09					x		Cytoplasm	
0744	2,3-bisphosphoglycerate-independent phosphoglycerate mutase	<i>pgm</i>	0	-0.71			-3.04	-3.31			-0.35	-3.17					x		Cytoplasm	
0745	Enolase	<i>eno</i>	0	-0.13	0	-0.14	-0.01	-0.23	-0.38	-0.28	-0.07	-0.23					x		Cytoplasm	MLS
0756	Clumping factor A	<i>clfA</i>	0	0.13	0	-0.23	-1.48	-1.55	-1.34	-1.34	-0.02	-1.42	***	Yes	Yes			Yes	Cell wall	Sortase substrate
0758	Extracellular matrix protein	<i>emp/ssp</i>	0	-0.04	0	-0.36	-3.18	-3.05	-2.68	-3.00	-0.10	-2.98	***	Yes	Yes			Yes	Extracellular / CW	Can re-bind to CW, non-cov. [7]
0776	Glycine cleavage system H protein	<i>gcvH</i>	0	-0.20	0	-0.52	0.10	-0.11	-0.25	-0.42	-0.18	-0.17							Cytoplasm	
0783	CsbD-like superfamily protein	<i>0783</i>	0	-0.06	0	-0.29	-0.75	-0.87	-0.14	-0.83	-0.09	-0.65	*					Yes	Unknown	
0800	Hydrolase	<i>nagD</i>	0	0.14	0	-0.08	-0.29	-0.68	-0.33	-0.42	0.02	-0.43	**					Yes	Unknown	
0803	D-alanine-D-alanyl carrier protein ligase	<i>dltA</i>	0	-0.15			0.02	-0.24			-0.08	-0.11							Cytoplasm	
0805	D-alanyl carrier protein	<i>dltC</i>	0	-0.26	0	-0.31	0.11	0.12	0.00	0.48	-0.14	0.18							Cytoplasm	
0824	Putative peptidyl-prolyl cis-trans isomerase	<i>ppi</i>	0	-0.24	0	-0.19	0.06	-0.25	0.01	-0.38	-0.11	-0.14						Yes	Cytoplasm	
0833	Glucose-6-phosphate isomerase	<i>pgi</i>	0	-0.19	0	-0.13	-0.70	-0.51	-0.26	-0.26	-0.08	-0.43	*				x		Cytoplasm	
0836	Signal peptidase IB	<i>spsB</i>	0	-0.51			-0.77	-0.08			-0.26	-0.43						Yes	Membrane	MLS
0839	Fumarylacetylacetate hydrolase family protein	<i>0839</i>	0	-0.02	0	-0.32	0.11	0.15	-0.15	-0.15	-0.09	-0.01						Yes	Cytoplasm	

NWMN	Protein name	Gene	wt I	wt II	wt III	wt IV	fold change (log 2)				σ wt	σ secDF	P	SignalP [1]	Sec SP [2]	No SP(*) [2]	SecretomeP [3]	Localization [4,5]		
							secDF I	secDF II	secDF III	secDF IV										
0845	Chaperone protein	<i>clpB</i>	0	-0.38	0	-0.01	0.20	-0.09	-2.34	-2.73	-0.10	-1.24							Cytoplasm	
0854	3-oxoacyl-[acyl-carrier-protein] synthase 2	<i>fabF</i>	0	0.00	0	-0.63	0.44	0.63	-0.27	-0.36	-0.16	0.11							Cytoplasm	
0885	2', 5' RNA ligase	<i>0885</i>	0	-0.04	0	-0.09	-0.33	-0.33	-0.44	-0.33	-0.03	-0.36	***						Cytoplasm	
0915	1,4-Dihydroxy-2-naphthoyl-CoA synthase	<i>memB</i>	0	0.22	0	-0.51	-0.18	-0.32	-0.27	-0.30	-0.07	-0.27							Cytoplasm	
0917	Cysteine protease	<i>sspB</i>	0	-0.02	0	-0.22	-1.38	-1.33	-0.65	-1.14	-0.06	-1.13	**	Yes	Yes		Yes		Extracellular	
0918	V8 protease, glutamyl endopeptidase precursor	<i>sspA</i>	0	-0.16			-1.19	-1.64			-0.08	-1.41		Yes	Yes		Yes		Extracellular	
0922	Bifunctional autolysin	<i>atl</i>	0	1.21	0	-0.34	1.67	0.91	0.19	0.12	0.22	0.72		Yes	Yes		Yes		Extracellular	
0925	Similar to cell envelope-related transcriptional attenuator	<i>0925</i>	0	-0.25	0	-0.54	-1.27	-1.31	-1.28	-1.90	-0.20	-1.44	***	Yes		x	Yes		Membrane	predicted uncleaved SP
0930	Probable quinol oxidase subunit 2	<i>qoxA</i>	0	-0.11			-0.88	-0.94			-0.06	-0.91	*				Yes		Membrane	
0932	Bifunctional protein folD	<i>folD</i>	0	-0.17			-0.40	-0.18			-0.09	-0.29							Cytoplasm	
0949	Histidine-containing phosphocarrier protein	<i>ptsH</i>	0	-0.20	0	-0.38	0.24	-0.01	-0.22	-0.15	-0.14	-0.04				x			Cytoplasm	
0950	Phosphoenolpyruvate-protein phosphotransferase	<i>ptsI</i>	0	-0.24	0	-0.46	0.04	0.06	-0.42	-0.73	-0.17	-0.26				x			Cytoplasm	
0959	Pyruvate dehydrogenase E1 component subunit alpha	<i>phdA</i>	0	-0.05	0	-0.52	0.05	-0.22	-0.29	-0.63	-0.14	-0.27				x			Cytoplasm	
0960	Pyruvate dehydrogenase E1 component subunit beta	<i>phdB</i>	0	-0.07	0	-0.28	0.20	-0.04	0.00	-0.10	-0.09	0.01				x			Cytoplasm	
0961	Dihydropolyllysine-residue acetyltransferase component of pyruvate dehydrogenase complex	<i>pdhC</i>	0	0.08	0	-0.35	0.02	-0.11	-0.13	-0.37	-0.07	-0.15				x			Cytoplasm	
0962	Dihydropolyl dehydrogenase	<i>pdhD</i>	0	-0.21	0	0.02	-0.39	-0.56	-0.24	-0.31	-0.05	-0.37	*			x			Cytoplasm	
1019	Putative major head protein Orf47	<i>1019</i>	0	0.12			-1.65	-1.53			0.06	-1.59	**						Cytoplasm	
1041	Iron-regulated surface determinant protein A	<i>isdA</i>	0	-0.08	0	-0.26	-0.53	-0.64	-0.31	-0.59	-0.08	-0.51	**	Yes	Yes		Yes		Cell wall	Sortase substrate
1057	Thioredoxin	<i>trxA</i>	0	-0.11	0	-0.35	0.79	0.56	0.18	-0.01	-0.12	0.38							Cytoplasm	
1067	FPRL1 inhibitory protein / FLIPr	<i>flr</i>	0	-0.16	0	-0.31	-4.07	-3.87	-3.04	-3.25	-0.12	-3.56	***	Yes	Yes		Yes		Extracellular	
1069	Fibrinogen-binding protein	<i>efb/fib</i>	0	0.27	0	0.09	-1.49	-1.11	-1.38	-1.34	0.09	-1.33	***	Yes	Yes		Yes		Extracellular / CW	Can re-bind to CW, non-cov. [7]
1073	Alpha-hemolysin	<i>hla</i>	0	0.02	0	-0.46	-1.16	-1.44	-0.67	-0.97	-0.11	-1.06	**	Yes	Yes		Yes		Extracellular	
1086	Uncharacterized N-acetyltransferase	<i>1086</i>	0	0.07			-0.68	-0.54			0.04	-0.61	*						Unknown	
1096	Cell division protein	<i>ftsZ</i>	0	-0.19	0	-0.60	-0.29	-0.16	-0.47	-0.24	-0.20	-0.29							Cytoplasm	
1103	Isoleucyl-tRNA synthetase OS	<i>ileS</i>	0	0.08	0	-0.19	-0.45	-0.44	-0.12	-0.51	-0.03	-0.38	*			x			Cytoplasm	
1125	Peptide deformylase	<i>def</i>	0	-0.60			-0.34	-0.79			-0.30	-0.57							Cytoplasm	
1139	Phosphate acyltransferase	<i>plsX</i>	0	1.27			-0.86	0.15			0.63	-0.36							Cytoplasm	
1140	Malonyl CoA-acyl carrier protein transacylase	<i>fabD</i>	0	-0.63			-0.15	-0.77			-0.31	-0.46							Cytoplasm	
1141	3-oxoacyl-[acyl-carrier-protein] reductase	<i>fabG</i>	0	-0.40	0	0.12	-0.65	-0.83	-0.47	-0.32	-0.07	-0.57	*						Cytoplasm	
1148	30S ribosomal protein S16	<i>rpsP</i>	0	-0.01	0	-0.39	0.13	-0.14	-0.42	-0.44	-0.10	-0.22					Yes		Cytoplasm	
1155	Succinyl-CoA ligase [ADP-forming] subunit β	<i>sucC</i>	0	0.02	0	-0.38	-0.40	-0.59	-0.56	-0.85	-0.09	-0.60	**						Cytoplasm	
1156	Succinyl-CoA ligase [ADP-forming] subunit α	<i>sucD</i>	0	0.27	0	-0.12	-0.35	-0.49	0.12	-0.48	0.04	-0.30				x			Cytoplasm	
1165	GTP-sensing transcriptional pleiotropic repressor	<i>codY</i>	0	-0.13	0	0.19	0.08	-0.11	0.53	0.86	0.02	0.34							Cytoplasm	

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							<i>secDF I</i>	<i>secDF II</i>	<i>secDF III</i>	<i>secDF IV</i>	σ wt	σ <i>secDF</i>							
1166	30S ribosomal protein S2	<i>rpsB</i>	0	-0.47	0	-0.22	-0.41	-0.40	-0.48	-0.62	-0.17	-0.48						Cytoplasm	
1167	Elongation factor Ts	<i>tsf</i>	0	-0.12	0	-0.45	0.08	-0.07	-0.06	-0.13	-0.14	-0.05			x			Cytoplasm	
1169	Ribosome-recycling factor	<i>frr</i>	0	0.06	0	-0.44	0.02	-0.22	0.28	0.04	-0.09	0.03						Cytoplasm	
1178	Translation initiation factor IF-2	<i>infB</i>	0	-0.23			-0.07	0.11			-0.11	0.02						Cytoplasm	
1181	Polyribonucleotide nucleotidyltransferase	<i>ribF</i>	0	-0.44			-1.25	-1.02			-0.22	-1.13						Cytoplasm	
1182	30S ribosomal protein S15	<i>rpsO</i>	0	-0.28			-0.39	-0.50			-0.14	-0.44						Cytoplasm	
1208	Glycerol kinase	<i>glpK</i>	0	0.19			-0.58	-0.55			0.09	-0.57						Cytoplasm	
1217	Glutamine synthetase	<i>glnA</i>	0	-0.03	0	0.08	0.32	0.16	0.59	0.45	0.01	0.38	*		x			Cytoplasm	
1236	Thermonuclease	<i>nuc</i>	0	0.00	0	-0.21	-0.57	-0.63	-0.33	-0.66	-0.05	-0.55	**				Yes	Extracellular	
1246	Catalase	<i>katA</i>	0	-0.04	0	-0.42	-0.57	-0.75	-0.79	-0.81	-0.11	-0.73	**		x		Yes	Cytoplasm	
1254	Transketolase	<i>tkt</i>	0	-0.12	0	-0.13	-0.13	-0.17	-0.07	-0.26	-0.06	-0.16			x		Yes	Unknown	
1263	Aconitate hydratase	<i>acnA</i>	0	-0.21	0	-0.11	-0.25	-0.46	-0.39	-0.19	-0.08	-0.32	*				Yes	Cytoplasm	
1274	Regulatory protein	<i>msrR</i>	0	-0.02	0	-0.19	-1.46	-1.00	-1.04	-1.14	-0.05	-1.16	***				Yes	Membrane	
1275	4-oxalocrotonate tautomerase	<i>1275</i>	0	0.12			0.01	-0.07			0.06	-0.03						Cytoplasm	
1313	Cold shock protein	<i>cspA</i>	0	-0.20	0	-0.44	0.44	0.35	0.62	0.12	-0.16	0.38	*		x		Yes	Cytoplasm	
1332	Probable CtpA-like serine protease	<i>1332</i>	0	-0.48			-1.22	-0.89			-0.24	-1.05					Yes	Membrane	
1333	Phosphotransferase system glucose-specific IIA component	<i>crr</i>	0	-0.46			-0.01	-0.32			-0.23	-0.17						Cytoplasm	
1339	Conserved hypothetical protein	<i>1339</i>	0	0.32			-0.36	-0.46			0.16	-0.41						Cytoplasm	
1365	Asparaginyl-tRNA synthetase	<i>asnS</i>	0	-0.10			-0.55	-0.59			-0.05	-0.57	*					Cytoplasm	
1378	Nucleoside diphosphate kinase	<i>ndk</i>	0	-0.05			-0.53	-0.61			-0.02	-0.57	*					Cytoplasm	
1382	DNA-binding protein HU	<i>hup</i>	0	0.10	0	-0.38	0.48	0.24	0.48	0.39	-0.07	0.40	*					Cytoplasm	
1385	30S ribosomal protein S1	<i>rpsA</i>	0	-0.09	0	-0.37	-0.42	-0.55	-0.15	-0.33	-0.11	-0.36			x			Cytoplasm	
1386	Cytidylate kinase	<i>cmk</i>	0	0.34			0.86	0.17			0.17	0.52						Cytoplasm	
1389	Elastin-binding protein	<i>ebpS</i>			0	-0.50			-1.13	-1.23	-0.25	-1.18			x		Yes	Membrane	MLS
1417	6-phosphogluconate dehydrogenase, decarboxylating	<i>gnd</i>	0	0.00			0.24	-0.10			0.00	0.07			x			Cytoplasm	
1456	Superoxide dismutase [Mn/Fe]	<i>sodA</i>	0	0.26	0	-0.48	-0.26	-0.24	-0.21	-0.43	-0.06	-0.28			x		Yes	Extracellular	No SP
1468	Glycyl-tRNA synthetase	<i>glyS</i>	0	0.07	0	-0.07	0.30	0.06	-0.14	-0.08	0.00	0.04						Cytoplasm	
1483	Chaperone protein	<i>dnaK</i>	0	0.02	0	-0.40	0.12	0.02	0.14	0.07	-0.10	0.09			x		Yes	Cytoplasm	
1484	Hsp-70 cofactor	<i>grpE</i>	0	0.08	0	-0.43	-0.51	-0.57	-0.09	-0.35	-0.09	-0.38			x		Yes	Cytoplasm	
1488	30S ribosomal protein S20	<i>rpsT</i>	0	-0.37			-0.59	-0.95			-0.19	-0.77					Yes	Cytoplasm	
1516	Conserved hypothetical protein	<i>1516</i>	0	0.00			0.14	-0.02			0.00	0.06						Cytoplasm	
1547	50S ribosomal protein L27	<i>rpmA</i>	0	0.14	0	-0.53	-0.02	-0.10	-0.42	-0.47	-0.10	-0.25					Yes	Cytoplasm	
1549	50S ribosomal protein L21	<i>rplU</i>	0	0.14	0	-0.97	-0.77	-0.36	-0.55	-0.92	-0.21	-0.65						Cytoplasm	
1558	Valyl-tRNA synthetase	<i>valS</i>	0	0.35			0.19	-0.13			0.18	0.03						Cytoplasm	
1569	Trigger factor	<i>tig</i>	0	-0.13	0	-0.21	-0.03	-0.23	-0.40	-0.40	-0.09	-0.26			x			Cytoplasm	
1574	Translation initiation factor IF-3	<i>infC</i>	0	-0.23			-1.13	-0.50			-0.12	-0.81						Cytoplasm	
1576	Threonyl-tRNA synthetase	<i>thrS</i>	0	0.19			0.03	-0.48			0.09	-0.23						Cytoplasm	
1587	Isocitrate dehydrogenase [NADP]	<i>citC</i>	0	-0.11	0	-0.35	-0.51	-0.71	-0.49	-1.00	-0.11	-0.68	**					Cytoplasm	

NWMN	Protein name	Gene	wt I	wt II	wt III	wt IV	fold change (log 2)				σ wt	σ secDF	P	SignalP [1]	Sec SP [2]	No SP(*) [2]	SecretomeP [3]	Localization [4,5]		
							secDF I	secDF II	secDF III	secDF IV										
1592	Pyruvate kinase	<i>pyk</i>	0	-0.32			-0.72	-0.60											Cytoplasm	
1593	6-phosphofructokinase	<i>pfk</i>	0	-0.14	0	-0.35	-0.56	-0.23	-0.40	-0.24	-0.12	-0.36							Cytoplasm	
1601	Metal-dependent hydrolase	<i>1601</i>	0	0.93			0.81	-0.03			0.46	0.39							Cytoplasm	
1604	Universal stress protein family protein	<i>1604</i>	0	-0.07	0	-0.40	-0.19	-0.25	-0.43	-0.31	-0.12	-0.29				x			Cytoplasm	
1609	Probable thiol peroxidase	<i>thil</i>	0	-0.04	0	-0.09	-0.25	-0.23	0.24	-0.32	-0.03	-0.14							Cytoplasm	
1613	30S ribosomal protein S4	<i>rpsd</i>	0	0.13			0.05	-0.09			0.07	-0.02							Cytoplasm	
1625	Formate--tetrahydrofolate ligase	<i>fhs</i>	0	0.25	0	-0.39	-0.21	-0.25	-0.18	-0.45	-0.04	-0.27				x			Cytoplasm	
1643	D-alanine aminotransferase	<i>dat</i>	0	-0.28			0.29	0.44			-0.14	0.36							Cytoplasm	
1681	Phosphoenolpyruvate carboxykinase [ATP]	<i>pckA</i>	0	-0.11			-0.75	-0.85			-0.05	-0.80	**					Yes	Cytoplasm	
1726	Signal transduction protein TRAP (RNAIII-activating protein TRAP)	<i>traP</i>	0	-0.08			-0.18	-0.56			-0.04	-0.37				x		Yes	Cytoplasm	
1733	Foldase protein	<i>prsA</i>	0	0.01	0	-0.31	-0.49	-0.49	-0.66	-0.82	-0.08	-0.62	**	Yes				Yes	Lipoprotein	
1737	Conserved hypothetical protein	<i>1737</i>	0	0.23	0	-0.40	0.83	0.71	-0.14	-0.03	-0.04	0.34							Cytoplasm	
1767	ThiJ/Pfpl family protein	<i>1767</i>	0	-0.19	0	-0.58	-0.36	-0.68	-0.56	-0.71	-0.19	-0.58							Cytoplasm	
1831	Ferritin	<i>ftnA</i>	0	0.31	0	-0.25	-0.20	-0.49	-0.36	-0.22	0.01	-0.31							Cytoplasm	
1837	Aspartyl/glutamyl-tRNA(Asn/Gln) amidotransferase subunit B	<i>gatB</i>	0	-0.37	0	-0.74	-0.23	-0.56	-0.05	-0.24	-0.28	-0.27							Cytoplasm	
1838	Glutamyl-tRNA(Gln) amidotransferase subunit A	<i>gatA</i>	0	0.16			0.62	0.34			0.08	0.48						Yes	Cytoplasm	
1839	Aspartyl/glutamyl-tRNA(Asn/Gln) amidotransferase subunit C	<i>gatC</i>	0	0.23	0	0.03	-0.40	-0.51	-0.55	-0.62	0.06	-0.52	***					Yes	Cytoplasm	
1847	Staphopain thiol protease	<i>scpA</i>	0	-0.23			-2.14	-2.22			-0.12	-2.18	*	Yes	Yes			Yes	Extracellular	
1857	Probable manganese-dependent inorganic pyrophosphatase	<i>ppaC</i>	0	-0.07	0	-0.03	-1.00	-0.38	-2.13	-2.27	-0.03	-1.45							Cytoplasm	
1859	Conserved hypothetical protein	<i>1859</i>	0	-0.77			-0.91	-1.13			-0.38	-1.02				x			Cytoplasm	
1872	65 kDa membrane protein	<i>eap/map</i>	0	-0.42	0	-0.27	-1.79	-1.61	-2.06	-2.21	-0.17	-1.92	***	Yes	Yes			Yes	Extracellular / CW	Can re-bind to CW, non-cov. [8]
1876	Staphylococcal complement inhibitor	<i>scn</i>	0	0.20	0	-0.05	-1.65	-1.66	-3.18	-3.46	0.04	-2.49	*	Yes	Yes			Yes	Extracellular	
1877	Chemotaxis inhibitory protein	<i>chp</i>	0	0.47	0	0.63	-2.52	-2.29	-1.25	-1.62	0.28	-1.92	**	Yes	Yes			Yes	Extracellular	
1880	Staphylokinase	<i>sak</i>	0	0.08	0	-0.48	-0.87	-0.87	-0.79	-0.82	-0.10	-0.84	**	Yes	Yes			Yes	Extracellular	
1883	Enterotoxin type A	<i>sea</i>	0	0.05	0	-0.08	-1.95	-1.89	-1.29	-1.40	-0.01	-1.63	**	Yes				Yes	Extracellular	
1927	Leukocidin/hemolysin toxin family F subunit	<i>lukF</i>	0	0.12	0	-0.21	-0.42	-0.54	-1.09	-0.87	-0.02	-0.73	*	Yes	Yes			Yes	Extracellular	
1928	Leukocidin/hemolysin toxin family S subunit	<i>lukS</i>	0	-0.03	0	-0.31	-1.42	-1.70	-1.16	-1.14	-0.09	-1.36	***	Yes	Yes			Yes	Extracellular	
1937	Chaperonin 60 kDa subunit	<i>groEL</i>	0	0.03	0	-0.61	-0.28	-0.51	-0.41	-0.47	-0.15	-0.42				x			Cytoplasm	
1938	10 kDa chaperonin	<i>groES</i>			0	-0.79			-0.41	-0.69	-0.40	-0.55				x			Cytoplasm	
1972	Anti-sigma-B factor antagonist	<i>rsbV</i>	0	-0.15	0	-0.11	-0.79	-0.87	-0.29	-0.09	-0.07	-0.51							Cytoplasm	
1999	Probable transglycosylase	<i>sceD</i>	0	0.09	0	-0.49	1.81	1.81	1.60	1.38	-0.10	1.65	***	Yes	Yes			Yes	Extracellular	Can ionically bind to cell wall [9]
2004	UDP-N-acetylglucosamine 1-carboxyvinyltransferase 1	<i>murA1</i>	0	1.01			-1.90	-0.17			0.51	-1.03							Cytoplasm	
2007	ATP synthase subunit beta	<i>atpD</i>	0	-0.18			-0.47	-0.74			-0.09	-0.60							Cytoplasm	

NWMN	Protein name	Gene	fold change (log 2)										P	SignalP [1]	Sec SP [2]	No SP(*) [2]	SecretomeP [3]	Localization [4,5]			
			wt I	wt II	wt III	wt IV	secDF I	secDF II	secDF III	secDF IV	ø wt	ø secDF									
2009	ATP synthase subunit alpha	<i>atpA</i>	0	-0.08			0.08	-0.30					-0.04	-0.11						Cytoplasm	
2016	Uracil phosphoribosyltransferase	<i>upp</i>	0	-0.41			-0.09	-0.14					-0.21	-0.11						Cytoplasm	
2024	50S ribosomal protein L31 type B	<i>rpmE</i>	0	-0.08	0	-0.36	-0.25	-0.48	-0.25	-0.58			-0.11	-0.39				Yes		Cytoplasm	
2026	Putative aldehyde dehydrogenase	<i>2026</i>	0	-0.21			-0.29	-0.55					-0.11	-0.42						Cytoplasm	
2029	Fructose-bisphosphate aldolase	<i>fbaA</i>	0	-0.23	0	-0.24	0.10	-0.06	-0.27	-0.18			-0.12	-0.10			x			Cytoplasm	
2031	CTP synthase	<i>pyrG</i>	0	0.12			-0.28	0.52					0.06	0.12						Cytoplasm	
2032	Probable DNA-directed RNA polymerase subunit delta	<i>rpoE</i>	0	-0.11			-0.06	-0.33					-0.06	-0.20				Yes		Cytoplasm	
2041	Deoxyribose-phosphate aldolase 1	<i>deoC</i>	0	-0.43			-0.63	-0.47					-0.21	-0.55						Cytoplasm	
2056	Glucosamine--fructose-6-phosphate aminotransferase [isomerizing]	<i>glmS</i>	0	0.04			0.51	-0.30					0.02	0.11						Cytoplasm	
2062	Phosphoglucosamine mutase	<i>glmM</i>	0	0.10			0.07	-0.11					0.05	-0.02						Cytoplasm	
2074	Conserved hypothetical protein	<i>2074</i>	0	0.13			-0.33	-0.36					0.06	-0.34			x	Yes		Unknown	
2086	Alkaline shock protein 23	<i>asp23</i>	0	0.06	0	-0.18	-0.55	-0.97	-0.65	-0.50			-0.03	-0.67	**		x	Yes		Unknown	
2119	30S ribosomal protein S9	<i>rpsI</i>			0	-0.66			-0.30	-0.95			-0.33	-0.63						Cytoplasm	
2120	50S ribosomal protein L13	<i>rplM</i>	0	-0.06	0	-0.51	-0.33	-0.41	-0.49	-0.56			-0.14	-0.45			x	Yes		Cytoplasm	
2125	50S ribosomal protein L17	<i>rplQ</i>	0	0.02	0	-0.24	0.04	-0.06	-0.19	-0.34			-0.06	-0.14						Cytoplasm	
2126	DNA-directed RNA polymerase subunit alpha	<i>rpoA</i>	0	-0.02			-0.14	-0.06					-0.01	-0.10						Cytoplasm	
2127	30S ribosomal protein S11	<i>rpsK</i>	0	0.11			-0.23	-0.29					0.06	-0.26				Yes		Cytoplasm	
2128	30S ribosomal protein S13	<i>rpsM</i>			0	-0.06			0.36	0.54			-0.03	0.45						Cytoplasm	
2130	Translation initiation factor IF-1	<i>infA</i>	0	-0.64	0	-0.10	0.52	0.35	0.88	0.88			-0.18	0.66	**					Cytoplasm	
2131	Adenylate kinase	<i>adk</i>	0	-0.02	0	-0.31	0.24	0.05	-0.06	-0.33			-0.08	-0.03						Cytoplasm	
2133	50S ribosomal protein L15	<i>rplO</i>	0	-0.01	0	-0.40	-0.29	-0.37	-1.00	-0.81			-0.10	-0.62	*					Cytoplasm	
2134	50S ribosomal protein L30	<i>rpmD</i>	0	0.02			-0.02	-0.13					0.01	-0.08				Yes		Cytoplasm	
2135	30S ribosomal protein S5	<i>rpsE</i>	0	0.02	0	-0.77	-0.07	-0.23	0.27	0.01			-0.19	-0.01						Cytoplasm	
2137	50S ribosomal protein L6	<i>rplF</i>	0	0.00	0	-0.29	0.19	0.13	0.09	0.17			-0.07	0.15				Yes		Cytoplasm	
2138	30S ribosomal protein S8	<i>rpsH</i>	0	0.22	0	-0.62	-0.03	0.09	-0.26	-0.20			-0.10	-0.10						Cytoplasm	
2140	50S ribosomal protein L5	<i>rplE</i>	0	0.05	0	-0.53	-0.04	-0.05	-0.42	-0.05			-0.12	-0.14						Cytoplasm	
2141	50S ribosomal protein L24	<i>rplX</i>	0	0.20			0.27	0.09					0.10	0.18				Yes		Cytoplasm	
2143	30S ribosomal protein S17	<i>rpsQ</i>	0	0.07			0.05	0.14					0.04	0.09				Yes		Cytoplasm	
2144	50S ribosomal protein L29	<i>rpmC</i>	0	0.05	0	-0.44	-0.25	-0.24	0.07	-0.08			-0.10	-0.13						Cytoplasm	
2145	50S ribosomal protein L16	<i>rplP</i>	0	-0.51			-0.98	-0.79					-0.25	-0.88						Cytoplasm	
2146	30S ribosomal protein S3	<i>rpsC</i>	0	0.08			0.05	-0.15					0.04	-0.05						Cytoplasm	
2147	50S ribosomal protein L22	<i>rplV</i>	0	0.13	0	-0.65	-0.73	-0.40	-0.28	-0.42			-0.13	-0.46						Cytoplasm	
2148	30S ribosomal protein S19	<i>rpsS</i>	0	0.13	0	-0.28	-0.46	-0.48	-0.26	-0.53			-0.04	-0.43	*					Cytoplasm	
2149	50S ribosomal protein L2	<i>rplB</i>	0	-0.19	0	-0.21	0.05	-0.05	-0.22	-0.31			-0.10	-0.13						Cytoplasm	
2150	50S ribosomal protein L23	<i>rplW</i>	0	0.21			0.66	0.42					0.11	0.54						Cytoplasm	
2151	50S ribosomal protein L4	<i>rplD</i>	0	-0.93			-0.60	-1.20					-0.47	-0.90						Cytoplasm	
2199	Staphylococcal secretory antigen	<i>ssaA</i>	0	0.05	0	-0.06	-1.95	-1.80	-1.75	-1.55			0.00	-1.76	***	Yes	Yes		Yes	Extracellular	Can ionically bind to cell wall [9]

NWMN	Protein name	Gene	wt I	wt II	wt III	wt IV	fold change (log 2)						P	SignalP [1]	Sec SP [2]	No SP(*) [2]	SecretomeP [3]	Localization [4,5]
							<i>secDF I</i>	<i>secDF II</i>	<i>secDF III</i>	<i>secDF IV</i>	σ wt	σ <i>secDF</i>						
2205	Putative 2-hydroxyacid dehydrogenase	2205	0	-0.50	0	-0.60	-0.12	-0.43	-0.25	-0.66	-0.27	-0.36						Cytoplasm
2270	Conserved hypothetical protein	2270	0	-0.24	0	0.04	-0.47	-1.03	-0.65	-1.06	-0.05	-0.80	**	Yes			Yes	Lipoprotein MLS
2317	Immunoglobulin-binding protein	<i>sbi</i>	0	-0.03	0	-0.16	-1.33	-1.23	-1.20	-1.22	-0.05	-1.25	***	Yes	Yes		Yes	Extracellular
2318	Gamma-hemolysin component A	<i>hlgA</i>	0	-0.03	0	-0.33	-0.09	-0.23	-0.67	-0.87	-0.09	-0.46		Yes	Yes		Yes	Extracellular
2319	Gamma-hemolysin component C	<i>hlgC</i>	0	0.08	0	-0.26	-0.32	-0.88	-1.00	-0.99	-0.04	-0.79	*	Yes	Yes		Yes	Extracellular
2320	Gamma-hemolysin component B	<i>hlgB</i>	0	-0.20	0	-0.23	-0.30	-0.58	-0.50	-0.78	-0.11	-0.54	*	Yes	Yes		Yes	Extracellular
2399	Fibronectin-binding protein A	<i>fnbA</i>	0	-0.08	0	-0.07	-2.01	-1.92	-2.78	-2.78	-0.04	-2.37	**	Yes	Yes		Yes	Cell wall Sortase substrate ¹
2448	ATP-dependent Clp protease, ATP-binding subunit	<i>clpL</i>	0	0.11			-0.71	-0.88			0.05	-0.80	*				Yes	Cytoplasm MLS
2454	1-pyrroline-5-carboxylate dehydrogenase	<i>rocA</i>	0	-0.21	0	-0.58	-0.67	-0.81	-0.76	-0.84	-0.20	-0.77	*			x		Cytoplasm
2467	O-acetyltransferase	<i>oatA</i>	0	0.35	0	-0.23	-1.07	-1.00	-1.13	-1.32	0.03	-1.13	***				Yes	Membrane
2469	Probable transglycosylase	<i>isaA</i>	0	-0.11	0	-0.04	-1.83	-1.87	-1.43	-1.35	-0.04	-1.62	**	Yes	Yes		Yes	Extracellular Can ionically bind to cell wall [9]
2480	Alpha/beta hydrolase fold family protein	2480	0	-0.04			-0.52	-0.22			-0.02	-0.37					Yes	Cytoplasm
2503	Fructose-bisphosphate aldolase class 1	<i>fda</i>	0	-0.03	0	-0.39	0.37	0.49	0.24	0.10	-0.10	0.30	*			x		Cytoplasm
2504	Probable malate:quinone oxidoreductase 2	<i>mqa2</i>	0	0.31			-0.52	-0.54			0.15	-0.53						Cytoplasm MLS
2529	Clumping factor B	<i>clfB</i>	0	-0.17	0	-0.30	-2.58	-2.64	-1.55	-1.80	-0.12	-2.15	**	yes	Yes		yes	Cell wall Sortase substrate
2537	Immunodominant staphylococcal antigen B	<i>isaB</i>	0	-0.05	0	-0.07	-0.60	-0.56	-1.09	-1.40	-0.03	-0.91	*	yes	Yes		yes	Extracellular
2569	Lipase 1	<i>lip</i>	0	0.27	0	-0.15	-0.68	-0.80	-0.54	-0.69	0.03	-0.67	**	yes	Yes		yes	Extracellular
2624	Delta-hemolysin	<i>hld</i>	0	-0.78	0	-1.09	-2.17	-2.12	-1.95	-2.12	-0.47	-2.09	**					Extracellular No SP

(*) Extracellular, without known signal peptide; MLS, multiple localization sites; ¹, FnbPA is truncated due to a point mutation leading to a stop codon. Thus it lacks the sortase motif LPXTG and is entirely secreted [10]; *, $P < 0.05$; **, $P < 0.01$; *** $P < 0.001$.

- Petersen TN, Brunak S, von Heijne G, Nielsen H (2011) SignalP 4.0: discriminating signal peptides from transmembrane regions. *Nat Methods* 8: 785-786.
- Sibbald MJJB, Ziebandt AK, Engelmann S, Hecker M, de Jong A, et al. (2006) Mapping the pathways to staphylococcal pathogenesis by comparative secretomics. *Microbiol Mol Biol Rev* 70: 755-788.
- Bendtsen J, Kiemer L, Fausboll A, Brunak S (2005) Non-classical protein secretion in bacteria. *BMC Microbiol* 5: 58.
- Yu NY, Wagner JR, Laird MR, Melli G, Rey S, et al. (2010) PSORTb 3.0: improved protein subcellular localization prediction with refined localization subcategories and predictive capabilities for all prokaryotes. *Bioinformatics* 26: 1608-1615.
- Krogh A, Larsson Br, von Heijne G, Sonnhammer ELL (2001) Predicting transmembrane protein topology with a hidden markov model: application to complete genomes. *J Mol Biol* 305: 567-580.
- Buist G, Steen A, Kok J, Kuipers OP (2008) LysM, a widely distributed protein motif for binding to (peptido)glycans. *Mol Microbiol* 68: 838-847.
- Chavakis T, Wiechmann K, Preissner KT, Herrmann M (2005) *Staphylococcus aureus* interactions with the endothelium: the role of bacterial "secretable expanded repertoire adhesive molecules" (SERAM) in disturbing host defense systems. *Thromb Haemost* 94: 278-285.
- Palma M, Hagggar A, Flock J-I (1999) Adherence of *Staphylococcus aureus* is enhanced by an endogenous secreted protein with broad binding activity. *J Bacteriol* 181: 2840-2845.
- Stapleton MR, Horsburgh MJ, Hayhurst EJ, Wright L, Jonsson I-M, et al. (2007) Characterization of IsaA and SceD, two putative lytic transglycosylases of *Staphylococcus aureus*. *J Bacteriol* 189: 7316-7325.
- Grundmeier M, Hussain M, Becker P, Heilmann C, Peters G, et al. (2004) Truncation of fibronectin-binding proteins in *Staphylococcus aureus* strain Newman leads to deficient adherence and host cell invasion due to loss of the cell wall anchor function. *Infect Immun* 72: 7155-7163.