Group of Subsystems: Enterobactin metabolism Matthew Cohoon, Department of Computer Science, University of Chicago, Chicago, IL

Subsystem: Enterobactin biosynthesis

Siderophore enterobactin is a triscatechol derivative of a cyclic triserine lactone that enables bacteria to recover iron from their environment. Believed earlier to be unique to Gram-negative bacteria, enterobactin has been isolated recently from two Gram-positive Streptomy ces species. It is a cyclic compound of three 2,3-dihydroxybenzoylserine units joined in a cyclic structure by lactone linkages (see molecular diagram on slide two). Enterobactin is synthesized under low iron concentration conditions and excreted into the environment where it binds Fe(III) with high affinity and specificity. The ferrisiderophore complexes are taken up into the cell by specific ATP-binding cassette transporter complex (see subsystem "ABC transporter ferric enterobactin (TC 3.A.1.14.)" for details). The biosynthesis of enterobactin starts with conversion of chorismic acid sequentially to isochorismate; 2,3-dihydro-2,3-dihydroxybenzoate; and then into 2,3-dihydroxybenzoic acid (DHB) catalyzed by EntC, EntB, and EntA. The amide linkage of DHB and L-serine is catalyzed by EntD, EntE, EntF, and EntB. Intermolecular cyclization of three molecules of DHB-Ser finally yields enterobactin. The functional role "M enaquinone-specific isochorismate synthase (EC 5.4.4.2)" has been included in this subsystem for clarity. M enF, similar to, although distinct from EntC, is an isochorismate synthase encoded by the *menF* gene.

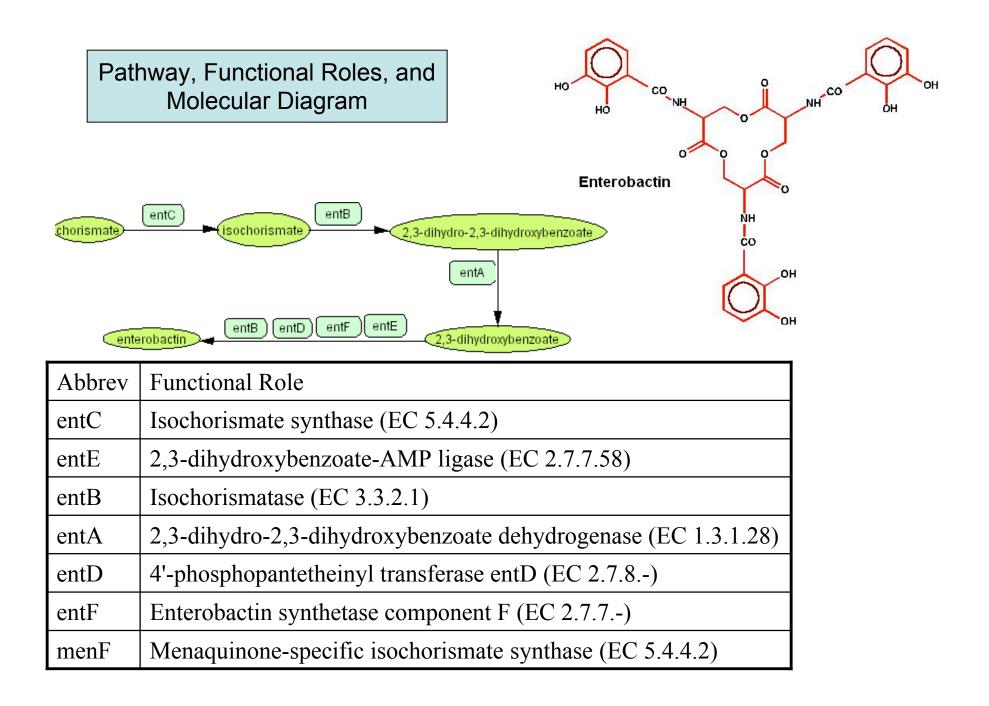
Slide 3 shows a snapshot of the subsystem spreadsheet and graphic representation of conserved chromosomal clusters of *ent* genes. It also illustrates application of SEED tools for detection of un- and mis- called ORFs by comparing gene clusters across genomes.

References:

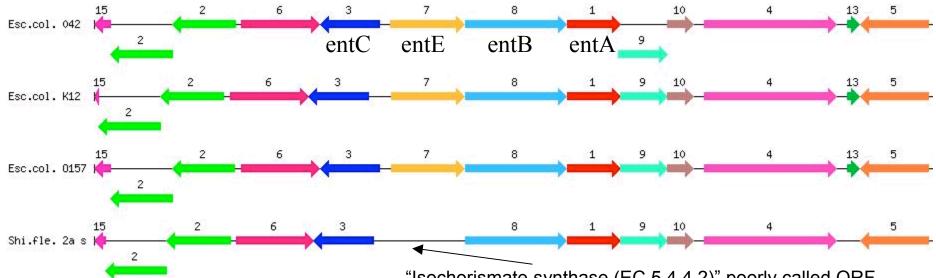
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Chromosomal and spreadsheet views of clustering



"Isochorismate synthase (EC 5.4.4.2)" poorly called ORF

Organism	entC	entE	entB	entA	entD	entF	menF
Escherichia coli 042 [B]	350	351	352	353	337	342	3088
Escherichia coli K12 [B]	593	594	595	596	582	586	2239
Escherichia coli O157:H7 EDL933 [B]	628	629	630	631	617	621	3151
Shigella flexneri 2a str. 301 [B]		489	490	491	478	481	2191

Subsystem: ABC transporter ferric enterobactin (TC 3.A.1.14.2)

This Subsystem is a companion subsystem created to help classify and categorize subunits of the specific ferric enterobactin ATP-binding cassette transporter. The TC number has been ascribed in accordance with the Transport Classification Database. A condensed spread sheet illustrating clustering on the chromosome of the genes encoding ferric enterobactin transporter subunits is shown on the next slide.

Abbrev	Functional Role
fepC	Ferric enterobactin transport ATP-binding protein fepC (TC 3.A.1.14.2)
fepG	Ferric enterobactin transport system permease protein fepG (TC 3.A.1.14.2)
fepD	Ferric enterobactin transport system permease protein fepD (TC 3.A.1.14.2)
fepB	Ferrienterobactin-binding periplasmic protein fepB (TC 3.A.1.14.2)

Organism	fepC	fepG	fepD	fepB
Trichodesmium erythraeum IMS101 [B]		2164	2163	5936
Chromobacterium violaceum ATCC 12472 [B]		2235	2236	2239
Escherichia coli 042 [B]		346	347	349
Escherichia coli CFT073 [B]		651	652	654
Escherichia coli E2348 [B]		3029	3028	3026
Escherichia coli K12 [B]		589	590	592
Escherichia coli O157:H7 [B]		702	703	705
Photorhabdus asymbiotica [B]	1561	1563	1564	1565
Photorhabdus luminescens subsp. laumondii TTO1 [B]	4411	4410	4409	4408
Salmonella enterica serovar Choleraesuis SC-B67 [B]		622	623	625
Salmonella enterica subsp. enterica serovar Gallinarum [B]		857	856	854
Salmonella enterica subsp. enterica serovar Typhi str. CT18 [B]		557	558	560
Salmonella typhimurium LT2 [B]		571	572	574
Shigella dysenteriae M131649 [B]		5060	5059	5057
Shigella flexneri 2a str. 2457T [B]		476	477	479
Shigella flexneri 2a str. 301 [B]	484	485	486	488
Shigella sonnei 53G [B]	597	596	595	593
Yersinia enterocolitica 8081 [B]	3556	3555	3554	3552
Yersinia pseudotuberculosis [B]		2420	312	71
Pseudomonas aeruginosa PAO1 [B]		4160	4159	4158
Pseudomonas aeruginosa UCBPP-PA14 [B]		1762	1761	1760
Salmonella bongori 12149 [V]	560	561	562	564