



Whole-Genome Sequences of the Archetypal K1 *Escherichia coli* Neonatal Isolate RS218 and Contemporary Neonatal Bacteremia Clinical Isolates SCB11, SCB12, and SCB15

Michael W. Day,^a Lydgia A. Jackson,^a Darrin R. Akins,^a David W. Dyer,^a Susana Chavez-Bueno^b

Departments of Microbiology and Immunology^a and Pediatrics,^b University of Oklahoma Health Sciences Center, Oklahoma City, Oklahoma, USA

Neonatal bacteremia *Escherichia coli* strains commonly belong to the K1 capsular type. Their ability to cause invasive neonatal disease appears to be determined by other virulence factors that have yet to be identified. We report here the genome sequences of four *E. coli* neonatal bacteremia isolates, including that of the archetypal strain RS218.

Received 31 December 2014 Accepted 15 January 2015 Published 26 February 2015

Citation Day MW, Jackson LA, Akins DR, Dyer DW, Chavez-Bueno S. 2015. Whole-genome sequences of the archetypal K1 *Escherichia coli* neonatal isolate RS218 and contemporary neonatal bacteremia clinical isolates SCB11, SCB12, and SCB15. Genome Announc 3(1):e01598-14. doi:10.1128/genomeA.01598-14. Copyright © 2015 Day et al. This is an open-access article distributed under the terms of the Creative Commons Attribution 3.0 Unported license. Address correspondence to Susana Chavez-Bueno, susana-chavez-bueno@ouhsc.edu.

scherichia coli is the most common cause of Gram-negative bacteremia in newborns and young infants and is associated with a mortality rate approaching 40% (1, 2). High-degree E. coli bacteremia can also result in meningitis, which commonly leads to long-term or permanent neurological sequelae (3). The majority of neonatal E. coli isolates causing bacteremia and meningitis express the K1 capsular antigen, which confers serum resistance and protects against phagocytic killing (4-6). RS218 is a wellcharacterized K1-positive neonatal E. coli meningitic isolate recovered in 1974 from the cerebrospinal fluid of a newborn (7, 8). Several RS218 virulence factors, such as OmpA, Ibe, and CNF1, are integral to bacterial passage across the blood-brain barrier (9). The identification and characterization of additional bacterial factors in the pathogenesis of RS218 and other invasive neonatal E. coli strains remain important areas of research. Here, we present the whole-genome sequences of RS218 and three contemporary E. coli blood culture isolates, SCB11, SCB12, and SCB15, which were identified in bacteremic newborns hospitalized at our institution in 2007 (10).

We performed whole-genome sequencing of the four isolates on an Illumina MiSeq using a 250-bp paired-end library. Assembly was performed *de novo* with the A5 assembly pipeline. The sequencing characteristics for each strain are summarized in Table 1. The annotation of the genomes was performed using the NCBI Prokaryotic Genomes Annotation Pipeline. The *adk*, *fumC*, *gyrB*, *icd*, *mdh*, *purA*, and *recA* genes in RS218 were consistent with its multilocus sequence type 95 (ST95). An examination of this

same gene set in SCB11, SCB12, and SCB15 indicated they corresponded to ST141, ST95, and ST501, respectively (11). RS218, SCB11, and SCB12 belong to phylogroup B2, whereas SCB15 is in phylogroup D. All four isolates carry the *kpsM* II group 2 capsule genes and express the K1 capsular antigen, as assessed by agglutination testing. Other known extraintestinal pathogenic E. coli (Ex-PEC) virulence genes, including *cnf1*, *fyuA*, *hek*, *hlyC*, *ibeA*, *iroN*, papGII, and sfa (12), were observed in the RS218, SCB11, and SCB12 genomes, but only *fyuA* and *sfa* were identified in SCB15. As expected, the recently identified plasmid pRS218 was present in strain RS218 (13). SCB12 was found to contain 90% of the pRS218 published sequence. SCB11 and SCB15, in contrast, do not contain pRS218. Future comparisons between these contemporary clinical isolates and the archetypal RS218 strain will yield valuable insight into the molecular pathways exploited by different E. coli isolates causing neonatal septicemia and meningitis.

Nucleotide sequence accession numbers. The nucleotide sequences have been deposited at GenBank under the accession numbers listed in Table 1.

ACKNOWLEDGMENT

This work was supported by the Oklahoma INBRE program, award 8P20GM103447, from the NIH/NIGMS.

REFERENCES

1. Biondi E, Evans R, Mischler M, Bendel-Stenzel M, Horstmann S, Lee V, Aldag J, Gigliotti F. 2013. Epidemiology of bacteremia in febrile infants in

TABLE 1 Genome sequencing statistics and accession numbers for neonatal invasive E. coli clinical isolates

Strain name	Total contig length (bp)	No. of contigs	G+C content (%)	N ₅₀ (bp)	GenBank accession no.	Accession no. version
RS218	5,173,885	68	50.6	277,884	JWZW0000000	JWZW01000000
SCB11	5,105,498	74	50.4	274,108	JSYT0000000	JSYT01000000
SCB12	5,478,295	96	50.5	144,019	JMQO0000000	JMQO01000000
SCB15	4,920,323	141	50.5	174,737	JSYU00000000	JSYU01000000

the United States. Pediatrics 132:990–996. http://dx.doi.org/10.1542/ peds.2013-1759.

- Simonsen KA, Anderson-Berry AL, Delair SF, Davies HD. 2014. Earlyonset neonatal sepsis. Clin Microbiol Rev 27:21–47. http://dx.doi.org/ 10.1128/CMR.00031-13.
- Lin MC, Chi H, Chiu NC, Huang FY, Ho CS. 2012. Factors for poor prognosis of neonatal bacterial meningitis in a medical center in northern Taiwan. J Microbiol Immunol Infect 45:442–447. http://dx.doi.org/ 10.1016/j.jmii.2011.12.034.
- Gaschignard J, Levy C, Romain O, Cohen R, Bingen E, Aujard Y, Boileau P. 2011. Neonatal bacterial meningitis: 444 cases in 7 years. Pediatr Infect Dis J 30:212-217. http://dx.doi.org/10.1097/ INF.0b013e3181fab1e7.
- Mahjoub-Messai F, Bidet P, Caro V, Diancourt L, Biran V, Aujard Y, Bingen E, Bonacorsi S. 2011. *Escherichia coli* isolates causing bacteremia via gut translocation and urinary tract infection in young infants exhibit different virulence genotypes. J Infect Dis 203:1844–1849. http:// dx.doi.org/10.1093/infdis/jir189.
- Cross AS, Gemski P, Sadoff JC, Orskov F, Orskov I. 1984. The importance of the K1 capsule in invasive infections caused by *Escherichia coli*. J Infect Dis 149:184–193. http://dx.doi.org/10.1093/infdis/149.2.184.
- 7. Silver RP, Aaronson W, Sutton A, Schneerson R. 1980. Comparative analysis of plasmids and some metabolic characteristics of *Escherichia*

coli K1 from diseased and healthy individuals. Infect Immun 29: 200–206.

- Achtman M, Mercer A, Kusecek B, Pohl A, Heuzenroeder M, Aaronson W, Sutton A, Silver RP. 1983. Six widespread bacterial clones among *Escherichia coli* K1 isolates. Infect Immun 39:315–335.
- Kim KS. 2012. Current concepts on the pathogenesis of *Escherichia coli* meningitis: implications for therapy and prevention. Curr Opin Infect Dis 25:273–278. http://dx.doi.org/10.1097/QCO.0b013e3283521eb0.
- Shakir SM, Goldbeck JM, Robison D, Eckerd AM, Chavez-Bueno S. 2014. Genotypic and phenotypic characterization of invasive neonatal *Escherichia coli* clinical isolates. Am J Perinatol 31:975–982. http:// dx.doi.org/10.1055/s-0034-1370341.
- Wirth T, Falush D, Lan R, Colles F, Mensa P, Wieler LH, Karch H, Reeves PR, Maiden MC, Ochman H, Achtman M. 2006. Sex and virulence in *Escherichia coli*: an evolutionary perspective. Mol Microbiol 60: 1136–1151. http://dx.doi.org/10.1111/j.1365-2958.2006.05172.x.
- Pitout JD. 2012. Extraintestinal pathogenic *Escherichia coli*: a combination of virulence with antibiotic resistance. Front Microbiol 3:9. http:// dx.doi.org/10.3389/fmicb.2012.00009.
- Wijetunge DS, Karunathilake KH, Chaudhari A, Katani R, Dudley EG, Kapur V, DebRoy C, Kariyawasam S. 2014. Complete nucleotide sequence of pRS218, a large virulence plasmid that augments pathogenic potential of meningitis-associated *Escherichia coli* strain RS218. BMC Microbiol 14:203. http://dx.doi.org/10.1186/s12866-014-0203-9.