PROKARYOTES





Complete Genome Sequence of Streptomyces lavendulae subsp. lavendulae CCM 3239 (Formerly "Streptomyces aureofaciens CCM 3239"), a Producer of the Angucycline-Type **Antibiotic Auricin**

Tobias Busche,^a Renata Novakova,^b Arwa Al'Dilaimi,^a Dagmar Homerova,^b Lubomira Feckova,^b Bronislava Rezuchova,^b Erik Mingyar, b Dominika Csolleiova, b Carmen Bekeova, b Anika Winkler, a Beatrica Sevcikova, b Jörn Kalinowski, a Jan Kormanec, b Christian Rückerta

^aCenter for Biotechnology (CeBiTec), Universität Bielefeld, Bielefeld, Germany bInstitute of Molecular Biology, Slovak Academy of Sciences, Bratislava, Slovakia

ABSTRACT Streptomyces lavendulae subsp. lavendulae CCM 3239 produces the angucycline antibiotic auricin and was thought to be the type strain of Streptomyces aureofaciens. We report the complete genome sequence of this strain, which consists of a linear chromosome and the linear plasmid pSA3239, and demonstrate it to be S. lavendulae subsp. lavendulae.

he Gram-positive soil bacteria of the genus Streptomyces undergo exceptional morphological differentiation, accompanied by "physiological differentiation," characterized by production of biologically active secondary metabolites like antibiotics (1). The strain "Streptomyces aureofaciens CCM 3239" (where the quotes indicate that the strain was originally misidentified as S. aureofaciens), received from the Czech Collection of Microorganisms (CCM), has been investigated for more than 25 years by our group with regard to morphological differentiation and the role of sigma factors therein (2-9).

We also identified the type II polyketide gene cluster aur1, located on the linear plasmid pSA3239. This gene cluster encodes production of the unique angucycline antibiotic auricin, which is produced during a narrow interval of several hours after entry into stationary phase, after which it is degraded due to its instability at high pH values reached later in the stationary phase. This unusual pattern of auricin production arises from its strict complex regulation, involving both feed-forward and feedback control by auricin intermediates via several transcriptional regulators (9-12). Thus, the genome sequence of "S. aureofaciens CCM 3239" should aid in the understanding of differentiation and global regulatory mechanisms for auricin biosynthesis in this strain.

Purified genomic DNA (gDNA) of "S. aureofaciens CCM 3239" was isolated with standard methods (13). Sequencing, assembly, and finishing were done as described previously (14) using a TrueSeq DNA PCR-free library instead of a Nextera WGS library. Assembly of 1.5 Gbp raw data resulted in 4 scaffolds containing 181 contigs. Gene prediction and annotation were performed using Prokka (15). The linear chromosome has a size of 8,691,831 bp (72.63% G+C content), while the linear plasmid pSA3239 has a size of 241,081 bp (72.12% G+C). Automated annotation of the complete genome sequence revealed the presence of 8,101 genes, including 159 RNA-encoding genes. Analysis of the genome sequence with the tool antiSMASH (16) revealed the presence of 26 respectively 5 putative secondary metabolite gene clusters located on the chromosome respectively pSA3239.

Received 26 January 2018 Accepted 2 February 2018 Published 1 March 2018

Citation Busche T, Novakova R, Al'Dilaimi A, Homerova D, Feckova L, Rezuchova B, Mingyar E, Csolleiova D, Bekeova C, Winkler A, Sevcikova B, Kalinowski J, Kormanec J, Rückert C. 2018. Complete genome sequence of Streptomyces lavendulae subsp. lavendulae CCM 3239 (formerly "Streptomyces aureofaciens CCM 3239"), a producer of the angucycline-type antibiotic auricin. Genome Announc 6:e00103-18. https:// doi.org/10.1128/genomeA.00103-18.

Copyright © 2018 Busche et al. This is an open-access article distributed under the terms of the Creative Commons Attribution 4.0 International license.

Address correspondence to Christian Rückert christian.rueckert@cebitec.uni-bielefeld.de.

genæmeAnnouncements™ genomea.asm.org 1

genameAnnouncements^T Busche et al.

Intriguingly, comparison of the 16S rRNA of "S. aureofaciens CCM 3239" with that of the type strain S. aureofaciens NBRC 12594 (17) revealed a low similarity (93% identity). Analysis of the 16S rRNA sequence in the ribosomal database project (18) using type strains revealed the highest similarity (99.3%) to Streptomyces lavendulae IFO 12789. Moreover, the genome sequence of "S. aureofaciens CCM 3239" was 100% identical with contigs of the draft sequence of Streptomyces lavendulae subsp. lavendulae NRRL B-2774 (GenBank accession number NZ_JOEW0000000), including the 16S rRNA genes. In addition, the genome sequence contains a gene cluster for the synthesis of the antibiotic streptothricin, which is identical on the nucleotide level to the gene cluster from Streptomyces lavendulae subsp. lavendulae NBRC 12789 (19). The lavender spore color and the production of soluble brown pigments provide additional phenotypic evidence that strain CCM 3239 is wrongly filed by CCM and is actually S. lavendulae subsp. lavendulae CCM 3239.

Accession number(s). The complete genome sequence was deposited at DDBJ/ ENA/GenBank under the accessions numbers CP024985 (chromosome) and CP024986 (plasmid).

ACKNOWLEDGMENTS

This work was supported by the Slovak Research and Development Agency under contract APVV-15-0410 and by VEGA grant 2/0002/16 from the Slovak Academy of Sciences. The research leading to these results has received funding from the European Commission's Seventh Framework Programme (FP7/2007-2013) under grant agreement STREPSYNTH (project 613877). This work was cofunded by the Slovak Research and Development Agency under contract DO7RP-0037-12. We acknowledge the financial support of the German Research Foundation (DFG) and the Open Access Publication Fund of Bielefeld University for the article processing charge.

REFERENCES

- 1. Chater KF. 2016. Recent advances in understanding Streptomyces. F1000Res 5:2795. https://doi.org/10.12688/f1000research.9534.1.
- 2. Kormanec J, Farkasovský M, Potúcková L. 1992. Four genes in Streptomyces aureofaciens containing a domain characteristic of principal sigma factors. Gene 122:63-70. https://doi.org/10.1016/0378-1119(92)90032-K.
- 3. Kormanec J, Farkasovský M. 1993. Differential expression of principal sigma factor homologues of Streptomyces aureofaciens correlates with the developmental stage. Nucleic Acids Res 21:3647-3652. https://doi .org/10.1093/nar/21.16.3647.
- 4. Kormanec J, Homerová D, Potúčková L, Nováková R, Rezuchová B. 1996. Differential expression of two sporulation specific σ factors of *Strepto*myces aureofaciens correlates with the developmental stage. Gene 181: 19-27. https://doi.org/10.1016/S0378-1119(96)00395-2.
- 5. Rezuchová B, Barák I, Kormanec J. 1997. Disruption of a sigma factor gene, sigF, affects an intermediate stage of spore pigment production in Streptomyces aureofaciens. FEMS Microbiol Lett 153:371. https://doi.org/ 10.1111/j.1574-6968.1997.tb12598.x.
- 6. Kormanec J, Sevcíková B, Sprusanský O, Benada O, Kofronová O, Nováková R, Rezuchová B, Potúcková L, Homérová D. 1998. The Streptomyces aureofaciens homologue of the whiB gene is essential for sporulation; its expression correlates with the developmental stage. Folia Microbiol (Praha) 43:605-612. https://doi.org/10.1007/BF02816376.
- 7. Nováková R, Sevcíková B, Kormanec J. 1998. A method for the identification

- of promoters recognized by RNA polymerase containing a particular sigma factor: cloning of a developmentally regulated promoter and corresponding gene directed by the Streptomyces aureofaciens sigma factor RpoZ. Gene 208:43-50. https://doi.org/10.1016/S0378-1119(97)00645-8.
- 8. Novakova R, Bistakova J, Kormanec J. 2004. Characterization of the polyketide spore pigment cluster whiESa in Streptomyces aureofaciens CCM3239. Arch Microbiol 182:388-395. https://doi.org/10.1007/s00203 -004-0720-2
- 9. Kormanec J, Novakova R, Mingyar E, Feckova L. 2014. Intriguing properties of the angucycline antibiotic auricin and complex regulation of its biosynthesis. Appl Microbiol Biotechnol 98:45-60. https://doi.org/10 .1007/s00253-013-5373-0.
- 10. Gordon D, Green P. 2013. Consed: a graphical editor for next-generation sequencing. Bioinformatics 29:2936-2937. https://doi.org/10.1093/ bioinformatics/btt515.
- 11. Seemann T. 2014. Prokka: rapid prokaryotic genome annotation. Bioinformatics 30:2068-2069, https://doi.org/10.1093/bioinformatics/btu153.
- 12. Blin K, Wolf T, Chevrette MG, Lu X, Schwalen CJ, Kautsar SA, Suarez Duran HG, de Los Santos ELC, Kim HU, Nave M, Dickschat JS, Mitchell DA, Shelest E, Breitling R, Takano E, Lee SY, Weber T, Medema MH. 2017. antiSMASH 4.0-improvements in chemistry prediction and gene cluster boundary identification. Nucleic Acids Res 45:W36. https://doi.org/10 .1093/nar/gkx319.

Volume 6 Issue 9 e00103-18