

## LUND UNIVERSITY

### Genome Sequence of Moraxella catarrhalis RH4, an Isolate of Seroresistant Lineage

Zomer, Aldert; de Vries, Stefan P. W.; Riesbeck, Kristian; Meinke, Andreas L.; Hermans, Peter W. M.; Bootsma, Hester J.

Published in: Journal of Bacteriology

DOI: 10.1128/JB.01833-12

2012

Link to publication

Citation for published version (APA): Zomer, A., de Vries, S. P. W., Riesbeck, K., Meinke, A. L., Hermans, P. W. M., & Bootsma, H. J. (2012). Genome Sequence of Moraxella catarrhalis RH4, an Isolate of Seroresistant Lineage. *Journal of Bacteriology*, 194(24), 6969-6969. https://doi.org/10.1128/JB.01833-12

Total number of authors: 6

General rights

Unless other specific re-use rights are stated the following general rights apply: Copyright and moral rights for the publications made accessible in the public portal are retained by the authors and/or other copyright owners and it is a condition of accessing publications that users recognise and abide by the legal requirements associated with these rights.

· Users may download and print one copy of any publication from the public portal for the purpose of private study

or research.
You may not further distribute the material or use it for any profit-making activity or commercial gain

· You may freely distribute the URL identifying the publication in the public portal

Read more about Creative commons licenses: https://creativecommons.org/licenses/

#### Take down policy

If you believe that this document breaches copyright please contact us providing details, and we will remove access to the work immediately and investigate your claim.

LUND UNIVERSITY

**PO Box 117** 221 00 Lund +46 46-222 00 00

# Genome sequence of *Moraxella catarrhalis* RH4, an isolate of seroresistant lineage.

Aldert Zomer<sup>1,2,†</sup>, Stefan P.W. de Vries<sup>1,†</sup>, Kristian Riesbeck<sup>3</sup>, Andreas L. Meinke<sup>4</sup>, Peter W.M Hermans<sup>1</sup>, Hester J. Bootsma<sup>1\*</sup>

<sup>1</sup>Laboratory of Pediatric Infectious Diseases, Radboud University Medical Centre, Nijmegen, The Netherlands, <sup>2</sup>Centre for Molecular and Biomolecular Informatics, Nijmegen Centre for Molecular Life Sciences, Radboud University Medical Centre, Nijmegen, The Netherlands, <sup>3</sup>Medical Microbiology, Department of Laboratory Medicine Malmö, Skåne University Hospital, Lund University, Malmö, Sweden, <sup>4</sup>Intercell AG, , Vienna, Austria

†These authors contributed equally to this work.

\* Corresponding author. Mailing address: Laboratory of Pediatric Infectious Diseases, Radboud University Medical Centre, P.O. Box 9101 (Route 224), 6500 HB Nijmegen, The Netherlands. Phone:
31-24-3666406. Fax: 31-24-3666352. E-mail: <u>H.Bootsma@cukz.umcn.nl</u>

#### ABSTRACT

Here we report the annotated genome sequence of *Moraxella catarrhalis* strain RH4, a seroresistant lineage strain isolated from the blood of an infected patient. This genome sequence will allow us to gain further insight into the genetic diversity of clinical *M*. *catarrhalis* isolates and will facilitate study of *M. catarrhalis* pathogenesis.

#### **GENOME ANNOUNCEMENT**

The Gram-negative diplococcus *Moraxella catarrhalis* is an emerging human-restricted respiratory tract pathogen. It is the third most common cause of childhood otitis media, and is frequently associated with exacerbations of chronic obstructive pulmonary disease (COPD) in adults. We reported the first completely assembled and annotated *M. catarrhalis* genome in 2010 (6) of strain BBH18 (erroneously referred to as RH4 at the time), a sputum isolate from a COPD patient during an exacerbation (8). In 2011, an additional 10 genome sequences of clinical *Moraxella* isolates were published (5) and compared to the BBH18 genome and the partial genome sequence of strain ATCC 43617 (10). This indicated a modest diversity in gene content and chromosomal organization between these isolates. Here we present the annotated genome sequence of the clinically relevant RH4 strain, which was originally isolated from the blood of an infected patient (4).

The draft genome sequence of *M. catarrhalis* RH4 was obtained using Illumina 50 bp pairedend technology (total of 13,826,736 reads, with 700x coverage). Reads were assembled with the Ray assembler software program (3), resulting in a total of 31 contigs (>100 bp in size). Contigs were ordered using the program Projector 2 (9) with the BBH18 sequence as a scaffold and the correct order was verified by gap-spanning PCR's. When possible, gaps were filled or corrected by Sanger sequencing of PCR products, followed by use of GapFiller (2).

2

After manual contig assembly, 9 contigs covering a total of 1,836,691 bp were obtained, which is within the size range of the reported genomes (1.78 to 1.96 Mbp).

The RH4 genome sequence was annotated using the RAST (rapid annotations using subsystems technology) server (1) and manually corrected for errors in open reading frame (ORF) calling. The total genome has a G+C content of 41.6% and is composed of 1,904 genes, including 1,845 protein-encoding genes, 4 rRNA operons, and at least 43 tRNA's. RH4 has a novel sequence variant for the *abcZ* allele and thus a novel multilocus sequence type (MLST), but clearly belongs to the seroresistant lineage (11). The RH4 genome contains the *bro-1*  $\beta$ -lactamase gene and all of the major known *M. catarrhalis* virulence factors, among which UspA1, UspA2H, MID/Hag, and, in contrast to, for instance, BBH18, a complete *mha* locus. Compared to all *Moraxella* genomes published to date, the RH4 genome contains 10 unique genes, 8 of which are located consecutively on a 10.1 kb fragment. In addition to 4 putative restriction-modification protein-encoding genes, this cluster contains a putative CiaB-encoding gene, where the ortholog in *Campylobacter jejuni* is involved in internalization into mammalian cells (7).

#### Nucleotide sequence accession number.

This whole-genome shotgun project has been deposited in DDBJ/EMBL/GenBank under the accession no. AMSO00000000. The version described in this article is the first version, AMSO01000000.1.

#### Acknowledgements

This work was supported by the Vienna Spot of Excellence (VSOE) grant (ID337956), the Anna and Edwin Berger foundation (to K.R.) and the Swedish Medical Research Council (grant number 521-2010-4221; to K.R.).

3

#### REFERENCES

- Aziz, R. K., D. Bartels, A. A. Best, M. DeJongh, T. Disz, R. A. Edwards, K. Formsma, S. Gerdes, E. M. Glass, M. Kubal, F. Meyer, G. J. Olsen, R. Olson, A. L. Osterman, R. A. Overbeek, L. K. McNeil, D. Paarmann, T. Paczian, B. Parrello, G. D. Pusch, C. Reich, R. Stevens, O. Vassieva, V. Vonstein, A. Wilke, and O. Zagnitko. 2008. The RAST Server: rapid annotations using subsystems technology. BMC Genomics 9:75.
- Boetzer, M., and W. Pirovano. 2012. Toward almost closed genomes with GapFiller. Genome Biol. 13:R56.
- 3. **Boisvert, S., F. Laviolette, and J. Corbeil.** 2010. Ray: simultaneous assembly of reads from a mix of high-throughput sequencing technologies. J. Comput. Biol. **17:**1519-1533.
- Christensen, J. J., J. Ursing, and B. Bruun. 1994. Genotypic and phenotypic relatedness of 80 strains of *Branhamella catarrhalis* of worldwide origin. FEMS Microbiol. Lett. 119:155-159.
- Davie, J. J., J. Earl, S. P. de Vries, A. Ahmed, F. Z. Hu, H. J. Bootsma, K. Stol, P. W. Hermans, R. M. Wadowsky, G. D. Ehrlich, J. P. Hays, and A. A. Campagnari. 2011. Comparative analysis and supragenome modeling of twelve *Moraxella catarrhalis* clinical isolates. BMC Genomics 12:70.
- de Vries, S. P., S. A. van Hijum, W. Schueler, K. Riesbeck, J. P. Hays, P. W. Hermans, and H. J. Bootsma. 2010. Genome analysis of *Moraxella catarrhalis* strain RH4, a human respiratory tract pathogen. J. Bacteriol. 192:3574-3583.
- Konkel, M. E., B. J. Kim, V. Rivera-Amill, and S. G. Garvis. 1999. Bacterial secreted proteins are required for the internaliztion of *Campylobacter jejuni* into cultured mammalian cells. Mol. Microbiol. 32:691-701.
- Mollenkvist, A., T. Nordstrom, C. Hallden, J. J. Christensen, A. Forsgren, and K. Riesbeck. 2003. The *Moraxella catarrhalis* immunoglobulin D-binding protein MID has conserved sequences and is regulated by a mechanism corresponding to phase variation. J. Bacteriol. 185:2285-2295.

- van Hijum, S. A., A. L. Zomer, O. P. Kuipers, and J. Kok. 2005. Projector 2: contigR mapping for efficient gap-closure of prokaryotic genome sequence assemblies. Nucleic Acids Res. 33:W560-566.
- Wang, W., L. Reitzer, D. A. Rasko, M. M. Pearson, R. J. Blick, C. Laurence, and E. J. Hansen. 2007. Metabolic analysis of *Moraxella catarrhalis* and the effect of selected in vitro growth conditions on global gene expression. Infect. Immun. 75:4959-4971.
- Wirth, T., G. Morelli, B. Kusecek, A. van Belkum, C. van der Schee, A. Meyer, and M.
   Achtman. 2007. The rise and spread of a new pathogen: seroresistant *Moraxella catarrhalis*.
   Genome Res. 17:1647-1656.