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Amanda L. Lewis

*Washington University School of Medicine in St. Louis*

Grace E. Deitzler

*Washington University School of Medicine in St. Louis*

Maria J. Ruiz

*Washington University School of Medicine in St. Louis*

Cory Weimer

*Washington University School of Medicine in St. Louis*

SoEun Park

*Washington University School of Medicine in St. Louis*

*See next page for additional authors*

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**Authors**

Amanda L. Lewis, Grace E. Deitzler, Maria J. Ruiz, Cory Weimer, SoEun Park, Lloyd S. Robinson, Kymberlie Hallsworth-Pepin, Aye Wollam, Makedonka Mitreva, and Warren G. Lewis

# Genome Sequences of 11 Human Vaginal *Actinobacteria* Strains

Amanda L. Lewis,<sup>a,b,d</sup> Grace E. Deitzler,<sup>a,d</sup> Maria J. Ruiz,<sup>a</sup> Cory Weimer,<sup>a,d</sup> SoEun Park,<sup>a,d</sup> Lloyd S. Robinson,<sup>a,d</sup> Kymerlie Hallsworth-Pepin,<sup>c,e</sup> Aye Wollam,<sup>c,e</sup> Makedonka Mitreva,<sup>c,e</sup> Warren G. Lewis<sup>c,d</sup>

Department of Molecular Microbiology,<sup>a</sup> Department of Obstetrics and Gynecology,<sup>b</sup> Department of Medicine,<sup>c</sup> Center for Women's Infectious Disease Research,<sup>d</sup> and McDonnell Genome Institute,<sup>e</sup> Washington University School of Medicine, St. Louis, Missouri, USA

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**The composition of the vaginal microbiota is an important health determinant. Several members of the phylum *Actinobacteria* have been implicated in bacterial vaginosis, a condition associated with many negative health outcomes. Here, we present 11 strains of vaginal *Actinobacteria* (now available through BEI Resources) along with draft genome sequences.**

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Address correspondence to Amanda L. Lewis, [allewis@wustl.edu](mailto:allewis@wustl.edu), or Warren G. Lewis, [WarrenGLewisPhD@gmail.com](mailto:WarrenGLewisPhD@gmail.com).

Bacterial vaginosis (BV) is a vaginal dysbiosis associated with serious health complications (1–6). It is characterized by the absence of *Lactobacillus* species in the vagina and overgrowth of a polymicrobial community often containing members of the phylum *Actinobacteria*, including *Gardnerella vaginalis*, *Atopobium* spp., and others. In fact, *G. vaginalis* was recently shown to elicit several features of BV in a mouse vaginal infection model (7, 8). Bifidobacteria are also commonly isolated from the vagina, although members of this genus are rarely found in pathological contexts. Here, we isolated 11 vaginal bacteria from the phylum *Actinobacteria*. Vaginal swabs were collected from nonpregnant and pregnant women according to Washington University institutional review board (IRB)-approved protocols (201108155 and 20110382). Organisms isolated from vaginal swabs were cultured anaerobically, and identification was performed by 16S rRNA gene sequencing. Genomic DNA was obtained using the Wizard genomic DNA purification kit (Promega). Methodological details on isolation and clinical information will be described elsewhere.

Genomes were assembled *de novo* using the One Button Velvet assembly pipeline (version 1.1.06) (9) with hash sizes of 31, 33, and 35 after downsizing the sample input data to 100× coverage. An internal core gene screen on the assembly tested for complete-

ness of the genome. After assembly, the minimum length for contigs was set to 200 bp, and an internal core gene screen was performed as defined by the Human Microbiome Project (HMP) (10). Then, adapters were removed, and low-quality regions were trimmed. Finally, a screen for contamination was performed. The process of gene annotation included generating both *ab initio* and evidence-based (BLAST) predictions. Functional predictions of coding sequences were made using GeneMark and Glimmer3 (11, 12). Loci were then defined by clustering predictions with the same reading frame. We evaluated predictions using the nonredundant (NR) and Pfam databases and resolved overlaps between adjacent coding genes. Intergenic regions not spanned by GeneMark and Glimmer3 were subject to a BLAST search against NCBI's NR database and predictions generated based on protein alignments. tRNA genes were determined using tRNAscan-SE (13) and noncoding RNA genes by RNAmmer (14) and Rfam (15). Metabolic pathways and subcellular localization were predicted using KEGG and PSORTb, respectively (16, 17), and functional domains were evaluated using InterProScan (18).

**Accession number(s).** These whole-genome shotgun projects have been deposited in GenBank under the accession numbers listed in Table 1. We have also made the strains available to the

**TABLE 1** Identifiers and nucleotide sequences for sequenced strains of vaginal *Actinobacteria*

Genus/species	Strain	BEI catalog no.	Nucleotide accession no.
<i>Actinomyces neuii</i>	MJR8396A	HMS-1266	LRPJ00000000
<i>Alloscardovia omnicoles</i>	CMW7705A	HMS-1282	LRPK00000000
<i>Atopobium vaginae</i>	CMW7778A	HMS-1300	LSOA00000000
<i>Bifidobacterium bifidum</i>	MJR8628B	HMS-1264	LRPO00000000
<i>Bifidobacterium breve</i>	GED8481	HMS-1261	LRPP00000000
<i>Bifidobacterium longum</i>	CMW7750	HMS-1299	LRPQ00000000
<i>Corynebacterium</i> sp.	CMW7794	HMS-1295	LSRB00000000
<i>Gardnerella vaginalis</i>	GED7275B	HMS-1272	LRPZ00000000
<i>Gardnerella vaginalis</i>	GED7760B	HMS-1284	LRQA00000000
<i>Gardnerella vaginalis</i>	CMW7778B	HMS-1298	LSRC00000000
<i>Propionibacterium avidum</i>	MJR7694	HMS-1291	LRVD00000000

research community by depositing them with the Biodefense and Emerging Infections (BEI) Research Resource Repository (see BEI numbers in Table 1).

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## REFERENCES

1. Wiesenfeld HC, Hillier SL, Krohn MA, Landers DV, Sweet RL. 2003. Bacterial vaginosis is a strong predictor of *Neisseria gonorrhoeae* and *Chlamydia trachomatis* infection. *Clin Infect Dis* 36:663–668. <http://dx.doi.org/10.1086/367658>.
2. Marrazzo JM, Martin DH, Watts DH, Schulte J, Sobel JD, Hillier SL, Deal C, Fredricks DN. 2010. Bacterial vaginosis: identifying research gaps proceedings of a workshop sponsored by DHHS/NIH/NIAID. *Sex Transm Dis* 37:732–744. <http://dx.doi.org/10.1097/OLQ.0b013e3181fbbc95>.
3. Hillier SL, Nugent RP, Eschenbach DA, Krohn MA, Gibbs RS, Martin DH, Cotch MF, Edelman R, Pastorek JG, 2nd, Rao AV, McNellis D, Regan JA, Carey C, Klebanoff MA. 1995. Association between bacterial vaginosis and preterm delivery of a low-birth-weight infant. The Vaginal Infections and Prematurity Study Group. *N Engl J Med* 333:1737–1742. <http://dx.doi.org/10.1056/NEJM199512283332604>.
4. Bradshaw CS, Morton AN, Hocking J, Garland SM, Morris MB, Moss LM, Horvath LB, Kuzevska I, Fairley CK. 2006. High recurrence rates of bacterial vaginosis over the course of 12 months after oral metronidazole therapy and factors associated with recurrence. *J Infect Dis* 193:1478–1486. <http://dx.doi.org/10.1086/503780>.
5. Wilson J. 2004. Managing recurrent bacterial vaginosis. *Sex Transm Infect* 80:8–11. <http://dx.doi.org/10.1136/sti.2002.002733>.
6. Allsworth JE, Peipert JF. 2007. Prevalence of bacterial vaginosis: 2001–2004 National Health and Nutrition Examination Survey data. *Obstet Gynecol* 109:114–120. <http://dx.doi.org/10.1097/01.AOG.0000247627.84791.91>.
7. Gilbert NM, O'Brien VP, Hultgren S, Macones G, Lewis WG, Lewis AL. 2013. Urinary tract infection as a preventable cause of pregnancy complications: opportunities, challenges, and a global call to action. *Global Adv Health Med* 2:59–69. <http://dx.doi.org/10.7453/gahmj.2013.061>.
8. Lewis WG, Robinson LS, Gilbert NM, Perry JC, Lewis AL. 2013. Degradation, foraging, and depletion of mucus sialoglycans by the vagina-adapted actinobacterium *Gardnerella vaginalis*. *J Biol Chem* 288:12067–12079. <http://dx.doi.org/10.1074/jbc.M113.453654>.
9. Zerbino DR, Birney E. 2008. Velvet: algorithms for *de novo* short read assembly using de Bruijn graphs. *Genome Res* 18:821–829. <http://dx.doi.org/10.1101/gr.074492.107>.
10. Nelson KE, Weinstock GM, Highlander SK, Worley KC, Creasy HH, Wortman JR, Rusch DB, Mitreva M, Sodergren E, Chinwalla AT, Feldgarden M, Gevers D, Haas BJ, Madupu R, Ward DV, Birren BW, Gibbs RA, Methe B, Petrosino JF, Strausberg RL, Sutton GG, White OR, Wilson RK, Durkin S, Giglio MG, Gujja S, Howarth C, Kodira CD, Kyrpidis N, Mehta T, Muzny DM, Pearson M, Pepin K, Pati A, Qin X, Yandava C, Zeng Q, Zhang L, Berlin AM, Chen L, Hepburn TA, Johnson J, McCorrison J, Miller J, Minx P, Nusbaum C, Russ C, Sykes SM, Tomlinson CM, Young S, et al. 2010. A catalog of reference genomes from the human microbiome. *Science* 328:994–999. <http://dx.doi.org/10.1126/science.1183605>.
11. Borodovsky M, Mills R, Besemer J, Lomsadze A. 2003. Prokaryotic gene prediction using GeneMark and GeneMark.Hmm. *Curr Protoc Bioinformatics* Chapter 4:Unit 4.5. <http://dx.doi.org/10.1002/0471250953.bi0405s01>.
12. Delcher AL, Harmon D, Kasif S, White O, Salzberg SL. 1999. Improved microbial gene identification with GLIMMER. *Nucleic Acids Res* 27:4636–4641. <http://dx.doi.org/10.1093/nar/27.23.4636>.
13. Lowe TM, Eddy SR. 1997. tRNAscan-SE: a program for improved detection of transfer RNA genes in genomic sequence. *Nucleic Acids Res* 25:955–964. <http://dx.doi.org/10.1093/nar/25.5.0955>.
14. Lagesen K, Hallin P, Rødland EA, Staerfeldt H-H, Rognes T, Ussery DW. 2007. RNAmmer: consistent and rapid annotation of ribosomal RNA genes. *Nucleic Acids Res* 35:3100–3108. <http://dx.doi.org/10.1093/nar/gkm160>.
15. Griffiths-Jones S, Moxon S, Marshall M, Khanna A, Eddy SR, Bateman A. 2005. Rfam: annotating non-coding RNAs in complete genomes. *Nucleic Acids Res* 33:D121–D124. <http://dx.doi.org/10.1093/nar/gki081>.
16. Kanehisa M, Goto S, Kawashima S, Okuno Y, Hattori M. 2004. The KEGG resource for deciphering the genome. *Nucleic Acids Res* 32:D277–D280. <http://dx.doi.org/10.1093/nar/gkh063>.
17. Yu NY, Wagner JR, Laird MR, Melli G, Rey S, Lo R, Dao P, Sahinalp SC, Ester M, Foster LJ, Brinkman FS. 2010. PSORTb 3.0: improved protein subcellular localization prediction with refined localization sub-categories and predictive capabilities for all prokaryotes. *Bioinformatics* 26:1608–1615. <http://dx.doi.org/10.1093/bioinformatics/btq249>.
18. Quevillon E, Silventoinen V, Pillai S, Harte N, Mulder N, Apweiler R, Lopez R. 2005. InterProScan: protein domains identifier. *Nucleic Acids Res* 33:W116–W120. <http://dx.doi.org/10.1093/nar/gki442>.