




Genome Sequence of Human Papillomavirus 23 Strain HPV-23/Lancaster/2015

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ABSTRACT The genome of human papillomavirus type 23 (HPV-23; family *Papillomaviridae*, genus *Betapapillomavirus*, species *Betapapillomavirus 2*, type 23) was assembled by deep sequencing from nasopharyngeal swabs. The assembled genome is 2.7% divergent over its full length from the single complete genome of HPV-23 in GenBank (accession no. U31781). We named the strain HPV-23/Lancaster/2015.

The family *Papillomaviridae* consists of more than 120 viral types divided into 49 genera (1). Within the genus *Betapapillomavirus*, type 23 of species *Betapapillomavirus 2* (HPV-23) is represented in GenBank by a single complete genome (2) composed of a circular double-stranded DNA of 7.3 kb (accession no. U31781), as well as two shorter fragments (3, 4). HPV-23 is a clinically significant papillomavirus, having been implicated in epidermodysplasia verruciformis (2), skin cancer (5), ocular syringoma (6), nongenital seborrhoeic keratosis (7), breast cancer (8), and toenail onycholysis (9).

Volunteers were recruited from a general practice surgery and a general hospital in Lancaster, UK (54.05°N, 2.80°W). Nasopharyngeal swabs were taken between 16 December 2014 and 25 February 2015. Ethical approval was granted by the UK National Research Ethics Service, reference 14/LO/1634, NIHR Clinical Research Network (UKCRN) portfolio, ID 17799. All methods were carried out in accordance with the relevant guidelines and regulations.

Pooled RNA from 51 swabs was deep sequenced using an Illumina Nextera XT library and HiSeq 2500 system (SRA GenBank accession no. SRP092324). An HPV-23 genome was assembled using Bowtie 1.1.1 (10) and BWA 0.7.12-r1039 (11), with U31781 as the template. The assembled genome is 7,317 bases in length, differing from U31781 by 188 substitutions (2.7%). A 6-base deletion in the new genome starts at position 5418, within the L2 gene. A further single-base deletion occurs at position 7193, in the 3' direction from the L1 gene. The predicted protein sequences are derived by reference to U31781 and differ at 62 amino acid residues (2.4%), without nonsense substitutions. de Villiers et al. (12) recommend that a nucleotide divergence of 15% be used as the threshold for designation of a new type of human papillomavirus. The new strain is therefore well within the range of diversity expected within type 23 and has been designated HPV-23/Lancaster/2015. It is only the second full-length genome of HPV-23 to be described.

Forslund et al. (13) found HPV types of the *Betapapillomavirus 2* species in 9% of nasal swabs in a study of 312 Danish health care staff, but they detected HPV-23 in only

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2 individuals (<0.7%). Of 9 deep-sequenced nonoverlapping subsets of individuals from our 51 swabs, we detected HPV-23 in all but one. Our frequency is therefore at least 8/51 (>15%) and possibly much higher. The significance of this clinically important papillomavirus at the prevalence in our sample remains a matter for speculation (BAM files, alignments, and phylogenetic trees are available from <https://doi.org/10.17635/lancaster/researchdata/134>).

Accession number(s). The genome sequence of HPV-23/Lancaster/2015 has been deposited in GenBank under the accession number [KY652675](https://www.ncbi.nlm.nih.gov/nuccore/KY652675).

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REFERENCES

- Harari A, Chen Z, Burk RD. 2014. Human papillomavirus genomics: past, present and future. *Curr Probl Dermatol* 45:1–18. <https://doi.org/10.1159/000355952>.
- Kremsdorf D, Favre M, Jablonska S, Obalek S, Rueda LA, Lutzner MA, Blanchet-Bardon C, Van Voorst Vader PC, Orth G. 1984. Molecular cloning and characterization of the genomes of nine newly recognized human papillomavirus types associated with epidermodysplasia verruciformis. *J Virol* 52:1013–1018.
- Chan SY, Delius H, Halpern AL, Bernard HU. 1995. Analysis of genomic sequences of 95 papillomavirus types: uniting typing, phylogeny, and taxonomy. *J Virol* 69:3074–3083.
- Berkhout RJ, Tieben LM, Smits HL, Bavinck JN, Vermeer BJ, ter Schegget J. 1995. Nested PCR approach for detection and typing of epidermodysplasia verruciformis-associated human papillomavirus types in cutaneous cancers from renal transplant recipients. *J Clin Microbiol* 33: 690–695.
- de Villiers EM. 1998. Human papillomavirus infections in skin cancers. *Biomed Pharmacother* 52:26–33.
- Assadoullina A, Bialasiewicz AA, de Villiers EM, Richard G. 2000. Detection of HPV-20, HPV-23, and HPV-DL332 in a solitary eyelid syringoma. *Am J Ophthalmol* 129:99–101. [https://doi.org/10.1016/S0002-9394\(99\)00292-5](https://doi.org/10.1016/S0002-9394(99)00292-5).
- Li YH, Chen G, Dong XP, Chen HD. 2004. Detection of epidermodysplasia verruciformis-associated human papillomavirus DNA in nongenital seborrhoeic keratosis. *Br J Dermatol* 151:1060–1065. <https://doi.org/10.1111/j.1365-2133.2004.06244.x>.
- Sigaroodi A, Nadji SA, Naghshvar F, Nategh R, Emami H, Velayati AA. 2012. Human papillomavirus is associated with breast cancer in the north part of Iran. *ScientificWorldJournal* 2012:837191. <https://doi.org/10.1100/2012/837191>.
- Umanoff N, Werner B, Rady PL, Tyring S, Carlson JA. 2015. Persistent toenail onycholysis associated with beta-papillomavirus infection of the nail bed. *Am J Dermatopathol* 37:329–333. <https://doi.org/10.1097/DAD.000000000000110>.
- Langmead B, Trapnell C, Pop M, Salzberg SL. 2009. Ultrafast and memory-efficient alignment of short DNA sequences to the human genome. *Genome Biol* 10:R25. <https://doi.org/10.1186/gb-2009-10-3-r25>.
- Li H, Durbin R. 2010. Fast and accurate long-read alignment with Burrows-Wheeler transform. *Bioinformatics* 26:589–595. <https://doi.org/10.1093/bioinformatics/btp698>.
- de Villiers EM, Fauquet C, Broker TR, Bernard HU, zur Hausen H. 2004. Classification of papillomaviruses. *Virology* 324:17–27. <https://doi.org/10.1016/j.virol.2004.03.033>.
- Forslund O, Johansson H, Madsen KG, Kofoed K. 2013. The nasal mucosa contains a large spectrum of human papillomavirus types from the *Betapapillomavirus* and *Gammmapapillomavirus* genera. *J Infect Dis* 208: 1335–1341. <https://doi.org/10.1093/infdis/jit326>.