

## Uniformity of rotavirus strain nomenclature proposed by the Rotavirus Classification Working Group (RCWG)

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**Abstract** In April 2008, a nucleotide-sequence-based, complete genome classification system was developed for group A rotaviruses (RVs). This system assigns a specific genotype to each of the 11 genome segments of a particular RV strain according to established nucleotide percent cutoff values. Using this approach, the genome of individual RV strains are given the complete descriptor of Gx-P[x]-Ix-Rx-Cx-Mx-Ax-Nx-Tx-Ex-Hx. The Rotavirus Classification Working Group (RCWG) was formed by scientists in the field to maintain, evaluate and develop

the RV genotype classification system, in particular to aid in the designation of new genotypes. Since its conception, the group has ratified 51 new genotypes: as of April 2011, new genotypes for VP7 (G20-G27), VP4 (P[28]-P[35]), VP6 (I12-I16), VP1 (R5-R9), VP2 (C6-C9), VP3 (M7-M8), NSP1 (A15-A16), NSP2 (N6-N9), NSP3 (T8-T12), NSP4 (E12-E14) and NSP5/6 (H7-H11) have been defined for RV strains recovered from humans, cows, pigs, horses, mice, South American camelids (guanaco), chickens, turkeys, pheasants, bats and a sugar glider. With

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increasing numbers of complete RV genome sequences becoming available, a standardized RV strain nomenclature system is needed, and the RCWG proposes that individual RV strains are named as follows: RV group/species of origin/country of identification/common name/year of identification/G- and P-type. In collaboration with the National Center for Biotechnology Information (NCBI), the RCWG is also working on developing a RV-specific resource for the deposition of nucleotide sequences. This resource will provide useful information regarding RV strains, including, but not limited to, the individual gene genotypes and epidemiological and clinical information. Together, the proposed nomenclature system and the NCBI RV resource will offer highly useful tools for investigators to search for, retrieve, and analyze the ever-growing volume of RV genomic data.

## Introduction

Rotaviruses (RVs) are members of the family *Reoviridae* and cause severe diarrheal illness in the young of various animal species [25]. In humans, RV infections lead to the death of more than 500,000 infants and young children each year, particularly in developing regions of the world [72]. RVs possess a genome consisting of 11 segments of double-stranded (ds) RNA [25]. Most segments encode a single polypeptide, allowing the virus to express six structural viral proteins (VPs) and five non-structural proteins (NSPs) [25]. However, in some group A RV strains, a second open reading frame (ORF) is detected in genome segment 11, leading to the expression of another protein product (NSP6) [35] in addition to NSP5. The viral particle has icosahedral symmetry and is composed of three concentric protein layers [25, 65]. VP7 and VP4 are the

components of the outermost protein layer (outer capsid), and each carries neutralizing epitopes [25]. The middle protein layer (inner capsid) is composed of VP6 and surrounds the inner layer (the core shell), which is composed of VP2 [65]. Packaged within the core shell are the viral RNA-dependent RNA polymerase (VP1) and RNA capping enzyme (VP3), as well as the 11 dsRNA genome segments [25]. The RV NSPs have various functions in the replication and morphogenesis of RV progeny and in evasion of the host immune response [25]. Based on the antigenic properties of VP6, RVs have been subdivided into five serological species (A-E) and two additional tentative species (F and G) according to the International Committee on Taxonomy of Viruses (ICTV) [4, 79]. These “RV species” are commonly referred to as “RV groups”. RVs belonging to species A, B and C (RVA, RVB and RVC, respectively) are known to infect humans and various animals, whereas RVs of species D, E, F and G (RVD, RVE, RVF and RVG, respectively) thus far have only been recovered from animals, mostly birds [4, 61]. Epidemiologically, RVA is the most important for human infection and disease and has been classified further using various approaches. Specifically, RVA strains have been categorized based on (i) the antigenic properties of VP6, VP7 and VP4 (subgroups, G-serotypes and P-serotypes, respectively); (ii) the migration pattern of the RNA genome segments when subjected to polyacrylamide gel electrophoresis (long, short, supershort or atypical electrophoretotypes); (iii) whole-genome RNA hybridization patterns (genogroups); and (iv) nucleotide sequence analysis (genotypes) [25, 57]. Due to the segmented nature of the RV genome, reassortment events can occur after co-infection with RV strains belonging to the same group/species both *in vitro* and *in vivo* [27, 34, 64]. Numerous examples of such gene exchanges are available in the

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literature for RVA strains [50, 61]. Recent data also suggest that reassortment involving the NSP1-encoding genome segment may have occurred between ancestral strains of avian RVA and RVD [92]. However, there is no evidence for reassortment among contemporary RVs that belong to different groups/species. Their inability to undergo segment exchange, even under experimental conditions, indicates that the RV groups can be thought of as unique viral species [4].

In April 2008, a nucleotide-sequence-based, complete genome classification system was developed for RVA strains [54]. This system assigns a specific genotype to each of the 11 RV genome segments according to established nucleotide percent cutoff values. The VP7-VP4-VP6-VP1-VP2-VP3-NSP1-NSP2-NSP3-NSP4-NSP5/6 genes of RV strains are described using the abbreviations Gx-P[x]-Ix-Rx-Cx-Mx-Ax-Nx-Tx-Ex-Hx ( $x =$  Arabic numbers starting from 1), respectively [54]. Full-genome analyses have increased our recognition of the relatedness between animal and human RV strains, and this highlights the relevance of a common nomenclature for both animal and public health. To maintain, evaluate, and develop this system, the Rotavirus Classification Working Group (RCWG) was formed [55], which includes researchers worldwide.

In this article, we report on the current status of genotype diversity for RVA strains that has been identified to date by the RCWG. Moreover, the RCWG is also working with researchers at the National Center for Biotechnology Information (NCBI) to develop a specific database for the deposition of RV sequence information. This RV resource will contain complete genome sequences that are annotated using VIGOR (Viral Genome ORF Reader) [99], as well as multiple types of metadata (epidemiological, clinical, etc.) to provide useful additional information to investigators. The new RV database will be very similar to the “Influenza Virus Resource”, which is widely used to monitor influenza virus strains [10]. As part of the database development, the RCWG herein proposes a standardized nomenclature for RV strains, which is similar to that already established for influenza viruses. Universal adoption of this new system by the RV community will be crucial for the retrieval of relevant information from the RV resource.

### Update from the RCWG

The task of the RCWG is to maintain, evaluate, and develop the RV genotype classification system. When the nucleotide sequence from a RVA strain of a potential new genotype is submitted to the RCWG, a thorough phylogenetic analysis is performed, and this analysis is reviewed by the members of the RCWG. When a consensus is reached,

the submitter receives an e-mail with the novel genotype number(s) of the particular gene(s), which can then be used in publications. To facilitate the utilization of the RV genotype classification system, a regularly updated online automatic web application, RotaC (<http://rotac.regatools.be/>), was developed [47].

Since the establishment of the RCWG, its activities have been reviewed at annual face-to-face meetings, held in conjunction with major scientific virology meetings. The first three RCWG-meetings were held at (i) the 27th annual meeting of the American Society for Virology (ASV) (July 14th, 2008, Ithaca, New York), (ii) the 10th International Symposium on dsRNA Viruses (July 24th, 2009, Hamilton Island, Australia) and (iii) the 29th annual meeting of the ASV (July 19th, 2010, Bozeman, Montana). The fourth and fifth RCWG meetings are planned to take place during the 30th annual meeting of the ASV in Minneapolis, Minnesota (July 16th-20th, 2011) and the 11th International Symposium on dsRNA Viruses in San Juan, Puerto Rico (November 27th-December 1st, 2012), respectively.

Table 1 contains a list of RV strains possessing new genotypes that have been recognised by the RCWG since the introduction of the classification system in April 2008. The new genotypes were found for RVs identified in a variety of host species including humans, cows, pigs, horses, mice, chickens, turkeys, pheasants, South American camelids, bats and a sugar glider. Several of these new genotypes have been published [1, 19, 23, 30, 44, 58, 62, 85, 87, 91, 95]. Table 2 shows a large selection of RV strains of all recognised genotypes that are not represented in Table 1. Table 3 contains a list with representative RV strains and accession numbers for each of the 166 currently established genotypes divided over the 11 genome segments.

Although the RV classification system has so far been limited to that of RVA strains, the RCWG plans to develop similar systems for RV strains belonging to other RV groups/species once a critical number of complete genome sequences has become available. The complete genomes of four human RVB strains were recently sequenced and analyzed by Yamamoto and colleagues, bringing the total number of genomes to eight (seven human RVB strains and a single murine RVB strain) [70, 103]. So far, all of the fully analyzed human RVB strains were detected in China and South-East Asia (Bangladesh, India and Myanmar). Regarding RVC strains, the entire genome sequences have been determined for only eight strains: the porcine strain Cowden and seven human strains detected in the UK, Japan, China, Bangladesh and India [12, 15, 104]. The first complete genome sequence of a RVD strain (RVD/chicken-wt/DEU/05V0049/2005/GXP[X]) was published recently [92]. Furthermore, the complete genome sequences of two novel adult diarrhea rotavirus (NADRV) strains (RVX/Human-tc/CHN/NADRV-J19/1997/GXP[X] and

**Table 1** RV strains possessing novel genotypes assigned by the RCWG since its formation in April 2008 [54, 55]

Strain name	New genotypes											Ref.
	VP7	VP4	VP6	VP1	VP2	VP3	NSP1	NSP2	NSP3	NSP4	NSP5	
RVA/Human-wt/ECU/Ecu534/2006/G20P[28]	<b>G20</b>	<b>P[28]</b>	<b>I13</b>									[87]
RVA/Cow-wt/JPN/Azuk-1/2006/G21P[29]	<b>G21</b>	<b>P[29]</b>	I2	R2	C2	M2	A13	N2	<b>T9</b>	E2	H3	[1]
RVA/Guanaco-wt/ARG/Chubut/1999/G8P[14]	G8	P[14]	I2	<b>R5</b>	C2	M2	A3	N2	T6	<b>E12</b>	H3	[58]
RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	G19	<b>P[30]</b>	I11	<b>R6</b>	<b>C6</b>	<b>M7</b>	<b>A16</b>	<b>N6</b>	<b>T8</b>	E10	<b>H8</b>	[91]
RVA/Turkey-tc/DEU/03V0002E10/2003/G22P[35]	<b>G22</b>	<b>P[35]</b>	I4								H4	[85], NP
RVA/Chicken-tc/DEU/06V0661/2006/G19P[31]	G19	<b>P[31]</b>	I11								H8	[85]
RVA/Human-wt/NPL/KTM368/2004/G11P[25]	G11	P[25]	<b>I12</b>	R1	C1	M1	A1	N1	T1	E1	H1	[62]
RVA/Human-tc/ITA/260-97/1997/G3P[3]	G3	P[3]	I3	R3	C3	M3	<b>A15</b>	N2	T3	E3	H6	[63]
RVA/Horse-wt/ARG/E30/1993/G3P[12]	G3	P[12]	I6	R2	C2	M3	A10	N2	T3	E2	<b>H7</b>	NP
RVA/Pig-wt/IRL/61/07-ire/2007/G2P[32]	G2	<b>P[32]</b>	I5							E9		[19]
RVA/Pheasant-wt/HUN/Phea14246/2008/G23P[?]	<b>G23</b>											[95]
RVA/Pig-wt/CAN/CE-M-06-0003/2005/G2P[27]	G2	P[27]	<b>I14</b>									[44]
RVA/Cow-tc/JPN/Dai-10/2007/G24P[33]	<b>G24</b>	<b>P[33]</b>	I2	R2	C2	M2	A13	N2	T9	E2	H3	[1]
RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	G16	P[16]	I7	<b>R7</b>	<b>C7</b>	<b>M8</b>	A7	<b>N7</b>	<b>T10</b>	E7	<b>H9</b>	NP
RVA/Bat-wt/KEN/KE4852/07/2007/G25P[6]	<b>G25</b>	P[6]	<b>I15</b>	RY <sup>a</sup>	<b>C8</b>			<b>N8</b>	<b>T11</b>	E2	<b>H10</b>	[23]
RVA/Pig-wt/JPN/FGP51/2009/G4P[34]	G4	<b>P[34]</b>										NP
RVA/Human-tc/KEN/B10/1987/G3P[2]	G3	P[2]	<b>I16</b>	<b>R8</b>	C5	M5	A5	N5	T5	<b>E13</b>	H5	[30]
RVA/Pig-wt/JPN/TJ4-1/2010/G26P[?]	<b>G26</b>											NP
RVA/Horse-tc/GBR/L338/1991/G13P[18]	G13	P[18]	I6	<b>R9</b>	<b>C9</b>	M6	A6	<b>N9</b>	<b>T12</b>	<b>E14</b>	<b>H11</b>	NP
RVA/SugarGlider-wt/JPN/SG33/2010/G27P[X]	<b>G27</b>											NP

The strain name, available genotype constellation and publication are shown. Novel assigned genotypes are in bold

An open space means the genotype is not known because sequencing has not been performed

An "X" in the name of a strain indicates that this information is missing

NP not published

<sup>a</sup> Partial sequences of this gene segment have been determined, but it could not be assigned to any of the established genotypes. It may be the representatives of a new genotype, but the entire ORF needs to be determined, as stated in the guidelines from the RCWG [55], before this can be confirmed

RVX/Human-wt/BAN/NADRV-B219/2002/GXP[X]), causing diarrhea in adults, have been determined [39, 70, 105]. These NADRV strains have not yet been assigned to an RV group/species by the ICTV. Recently, VP6 sequence information was obtained for avian RVF and RVG strains [40]. In the future, analysis of these sequence data should allow the calculation of sequence-based thresholds to define (new) RV groups/species and genotypes within groups/species. A detailed overview of the current classification for non-RVA strains was published recently [61].

### Proposed nomenclature of RV strains

Currently, no guidelines exist for the naming of RV strains. The lack of a standardized nomenclature system has led to discrepancies in the literature where researchers have developed their own naming systems. With increasing numbers of complete RV genome sequences becoming available, and

with the development of specific resources to retrieve and analyze these sequences, a more uniform nomenclature is clearly needed. The RCWG has discussed this issue, and we propose the following nomenclature for individual strains:

RV group/species of origin/country of identification/  
common name/year of identification/G- and P-type

#### A) Guidelines for *wild-type* RV strains

These guidelines apply to any wild-type RV strain or naturally occurring reassortant RV strain recovered from human or animal populations that have been sequenced directly from clinical specimens (such as stool, blood or tissue samples) or environmental samples. Guidelines for nomenclature of i) tissue-culture-adapted RV strains, ii) RV vaccine strains, iii) RV strains that have been generated in the laboratory by reassortment or iv) RV strains engineered in the laboratory by reverse genetics or equivalent mechanisms are provided further below in sections B), C), D), and E).

**Table 2** List of RV strains representing all of the currently established genotypes except for those represented in Table 1

Strain name	Genotypes											Ref.
	VP7	VP4	VP6	VP1	VP2	VP3	NSP1	NSP2	NSP3	NSP4	NSP5	
RVA/Human-tc/USA/Wa/1974/G1P1A[8]	G1	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[54]
RVA/Human-wt/BGD/Dhaka16-03/2003/G1P[8]	G1	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[76]
RVA/Human-wt/USA/LB2719/2006/G1P[8]	G1	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[7]
RVA/Human-wt/IND/06361/2006/G1P[8]	G1	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[2]
RVA/Human-wt/IND/0613158/2006/G1P[8]	G1	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[2]
RVA/Human-wt/IND/061060/2006/G1P[8]	G1	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[2]
RVA/Human-wt/IND/NIV929893/1992/G1P[19]	G1	P[19]	I1							E1		[16]
RVA/Pig-wt/SVN/P21-5/2004/G1P[27]	G1	P[27]								E9		[88]
RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	G2	P[4]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[54]
RVA/Human-wt/CHN/TB-Chen/1996/G2P[4]	G2	P[4]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[13]
RVA/Human-wt/USA/LB2744/2006/G2P[4]	G2	P[4]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[7]
RVA/Human-wt/BGD/MMC6/2005/G2P[4]	G2	P[4]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[33]
RVA/Human-wt/BGD/MMC88/2005/G2P[4]	G2	P[4]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[33]
RVA/Pig-wt/ESP/34461-4/2003/G2P[23]	G2	P[23]	I5							E1	H1	[42]
RVA/Pig-wt/THA/CMP034/2000/G2P[27]	G2	P[27]	I5							E9	H1	[42]
RVA/Simian-tc/ZAF/SA11-H96/1958/G3P5B[2]	G3	P[2]	I2	R2	C5	M5	A5	N5	T5	E2	H5	[86]
RVA/Simian-tc/USA/RRV/1975/G3P[3]	G3	P[3]	I2	R2	C3	M3	A9	N2	T3	E3	H6	[59]
RVA/Human-tc/USA/HCR3A/1984/G3P[3]	G3	P[3]	I3	R3	C2	M3	A9	N2	T3	E3	H6	[93]
RVA/Cat-tc/AUS/Cat97/1984/G3P[3]	G3	P[3]	I3	R3	C2	M3	A9	N2	T3	E3	H6	[93]
RVA/Dog-tc/USA/CU-1/1982/G3P[3]	G3	P[3]	I3	R3	C2	M3	A9	N2	T3	E3	H6	[93]
RVA/Human-wt/THA/CMH222/2001/G3P[3]	G3	P[3]	I8							E3		[41]
RVA/Human-tc/AUS/RV3/1977/G3P2A[6]	G3	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[82]
RVA/Cow-lab/GBR/PP-1/1976/G3P[7]	G3	P[7]					A3			E8		[22]
RVA/Pig-tc/VEN/A131/1988/G3P9[7]	G3	P[7]	I5	R1	C2	M1	A1	N1	T1	E1	H1	[54]
RVA/Human-tc/USA/P/1974/G3P1A[8]	G3	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[54]
RVA/Human-wt/USA/DC5544-Bethesda/1991/G3P[8]	G3	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[66]
RVA/Cat-wt/ITA/BA222/2005/G3P[9]	G3	P[9]	I2	R2	C2	M2	A3	N1	T3	E2	H3	[51]
RVA/Human-wt/ITA/PAH136/1996/G3P[9]	G3	P[9]	I2	R2	C2	M2	A3	N1	T6	E2	H3	[20]
RVA/Human-wt/ITA/PAI58/1996/G3P[9]	G3	P[9]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[20]
RVA/Cat-tc/AUS/Cat2/1984/G3P[9]	G3	P[9]	I3	R3	C2	M3	A3	N1	T6	E3	H3	[93]
RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	G3	P[9]	I3	R3	C3	M3	A3	N3	T3	E3	H3	[54]
RVA/Human-wt/BEL/B4106/2000/G3P[14]	G3	P[14]	I2	R2	C2	M3	A9	N2	T6	E5	H3	[52]
RVA/Rabbit-tc/ITA/30/96/1996/G3P[14]	G3	P[14]	I2	R2	C2	M3	A9	N2	T6	E5	H3	[52]
RVA/Rabbit-wt/ITA/229/01/2001/G3P[22] <sup>a</sup>	G3	P[22]								E5		[48]
RVA/Rhesus-tc/USA/TUCH/2002/G3P[24]	G3	P[24]	I9	R3	C3	M3	A9	N1	T3	E3	H6	[59]
RVA/Human-wt/IND/mani-253/2007/G4P[4]	G4	P[4]	I1	R1	C1	M2	A8	N1	T1	E1	H1	[69]
RVA/Human-wt/IND/mani-362/2007/G4P[6]	G4	P[6]	I1	R1	C1	M1	A8	N1	T1	E1	H1	[69]
RVA/Human-tc/GBR/ST3/1975/G4P2A[6]	G4	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[54]
RVA/Pig-tc/USA/Gottfried/1983/G4P[6]	G4	P[6]	I1	R1	C1	M1	A8	N1	T1	E1	H1	[62]
RVA/Human-tc/CHN/R479/2004/G4P[6]	G4	P[6]	I5	R1	C1	M1	A1	N1	T7	E1	H1	[100]
RVA/Pig-tc/USA/OSU/1977/G5P9[7]	G5	P[7]	I5	R1	C1	M1	A1	N1	T1	E1	H1	[62]
RVA/Human-tc/BRA/IAL28/1992/G5P[8]	G5	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[54]
RVA/Pig-tc/VEN/A34/1985/G5P[23]	G5	P[23]								E1		[45]
RVA/Pig-wt/ITA/134/04-15/2003/G5P[26]	G5	P[26]	I5							E1		[49]
RVA/Cow-tc/VEN/BRV033/1990/G6P6[1]	G6	P[1]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[54]
RVA/Cow-tc/FRA/RF/1982/G6P[1]	G6	P[1]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[18]
RVA/Goat-tc/BGD/GO34/1999/G6P[1]	G6	P[1]	I2	R2	C2	M2	A11	N2	T6	E2	H3	[29]

Table 2 continued

Strain name	Genotypes											Ref.
	VP7	VP4	VP6	VP1	VP2	VP3	NSP1	NSP2	NSP3	NSP4	NSP5	
RVA/Cow-tc/USA/WC3/1981/G6P[5]	G6	P[5]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[54]
RVA/Cow-tc/GBR/UK/1973/G6P7[5]	G6	P[5]	I2	R2	C2	M2	A3	N2	T7	E2	H3	[21]
RVA/Human-wt/BEL/B1711/2002/G6P[6]	G6	P[6]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[56]
RVA/Human-wt/IND/HP140/1987/G6P[13]	G6	P[13]	I2	R1	C1	M1				E1	H1	[97]
RVA/Human-wt/BEL/B10925-97/1997/G6P[14]	G6	P[14]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[58]
RVA/Human-wt/HUN/Hun5/1997/G6P[14]	G6	P[14]	I2	R2	C2	M2	A11	N2	T6	E2	H3	[58]
RVA/Human-wt/HUN/BP1879/2003/G6P[14]	G6	P[14]	I2	R2	C2	M2	A11	N2	T6	E2	H3	[6]
RVA/Turkey-tc/IRL/Ty-3/1979/G7P[17]	G7	P[17]	I4							E11		[68]
RVA/Guanaco-wt/ARG/Rio_Negro/1998/G8P[1]	G8	P[1]	I2	R5	C2	M2	A13	N2	T6	E12	H3	[58]
RVA/Macaque-tc/USA/PTRV/1990/G8P[1]	G8	P[1]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[59]
RVA/Human-wt/NIC/NIC522/2008/G8P[1]	G8	P[1]	I2	R2	C2	M2	A13	N2	T6 <sup>c</sup>	E2	H3	[5]
RVA/Human-tc/KEN/B12/1987/G8P[1]	G8	P[1]	I2	R2	C2	M	A3	N2	T6	E2	H3	[32]
RVA/Cow-tc/THA/A5-13/1988/G8P[1]	G8	P[1]					A14					[71]
RVA/Human-wt/DEU/GER1H-09/2009/G8P[4]	G8	P[4]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[73]
RVA/Human-wt/COD/DRC86/2003/G8P[6]	G8	P[6]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[53]
RVA/Human-wt/COD/DRC88/2003/G8P[8]	G8	P[8]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[53]
RVA/Human-tc/IDN/69M/1980/G8P4[10]	G8	P[10]	I2	R2	C2	M2	A2	N2	T2	E2	H2	[54]
RVA/Human-wt/HUN/BP1062/2004/G8P[14]	G8	P[14]	I2	R2	C2	M2	A11	N2	T6	E2	H3	[9]
RVA/Sheep-tc/ESP/OVR762/2002/G8P[14]	G8	P[14]	I2	R2	C2	M2	A11	N2	T6	E2	H3	[58]
RVA/Human-wt/BEL/B3458/2003/G9P[8]	G9	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[54]
RVA/Human-tc/USA/WI61/1983/G9P1A[8]	G9	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[54]
RVA/Human-wt/USA/OM46/1998/G9P[8]	G9	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[67]
RVA/Human-wt/USA/OM473/2000/G9P[8]	G9	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[67]
RVA/Human-tc/IND/I116E/1985/G9P[11]	G9	P[11]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[82]
RVA/Human-wt/IND/RMC321/1990/G9P[19]	G9	P[19]	I5	R1	C1	M1	A1	N1	T1	E1	H1	[96]
RVA/Human-wt/IND/mani-97/2006/G9P[19]	G9	P[19]	I5	R1	C1	M1	A8	N1	T1	E1	H1	[69]
RVA/Human-wt/IND/mani-265/2007/G10P[6]	G10	P[6]	I2	R2	C2	M2	A3	N2	T2	E2	H2	[69]
RVA/Human-wt/NGA/6717ARN/2002/G10P[8]	G10	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[24]
RVA/Human-wt/CIV/6755ARN/2002/G10P[8]	G10	P[8]	I2	R1	C1	M1	A1	N1	T1	E1	H1	[24]
RVA/Human-wt/IND/N155/2003/G10P[11]	G10	P[11]	I2	R2	C2	M2	A1	N1	T1	E2	H3	[77]
RVA/Cow-tc/CHN/DQ-75/2008/G10P[11]	G10	P[11]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[102]
RVA/Sheep-tc/CHN/Lamb-NT/XXXX/G10P[15]	G10	P[15]	I10	R2	C2	M2	A11	N2	T6	E2	H3	[14]
RVA/Human-wt/ECU/EC2184/2005/G11P[6]	G11	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[8]
RVA/Pig-tc/MEX/YM/1983/G11P9[7]	G11	P[7]	I5	R1	C1	M1	A8	N1	T1	E1	H1	[62]
RVA/Pig-tc/VEN/A253/1988/G11P9[7]	G11	P[7]	I5	R1	C2	M1	A1	N1	T1	E1	H1	[54]
RVA/Human-wt/BGD/Matlab36-02/2002/G11P[8]	G11	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[62]
RVA/Human-wt/BGD/Dhaka6/2001/G11P[25]	G11	P[25]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[62]
RVA/Human-wt/NPL/KTM368/2004/G11P[25]	G11	P[25]	I12	R1	C1	M1	A1	N1	T1	E1	H1	[62]
RVA/Human-tc/PHL/L26/1987/G12P[4]	G12	P[4]	I2	R2	C2	M1	A2	N1	T2	E2	H1	[75]
RVA/Human-wt/BGD/Dhaka12-03/2003/G12P[6]	G12	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[75]
RVA/Human-wt/BGD/Matlab13/2003/G12P[6]	G12	P[6]	I1	R1	C1	M1	A1	N1	T2	E1	H1	[75]
RVA/Human-wt/BGD/RV161/2000/G12P[6]	G12	P[6]	I2	R2	C2	M2	A2	N2	T2	E1	H2	[75]
RVA/Human-wt/BGD/RV176-00/2000/G12P[6]	G12	P[6]	I2	R2	C2	M2	A2	N2	T2	E6	H2	[75]
RVA/Human-tc/KOR/CAU195/200X/G12P[6]	G12	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[89]
RVA/Human-tc/KOR/CAU214/200X/G12P[6]	G12	P[6]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[89]
RVA/Pig-wt/IND/RU172/2002/G12P[7]	G12	P[7]	I5	R1	C1	M1	A1	N1	T1	E1	H1	[31]
RVA/Human-wt/BEL/B4633/2003/G12P[8]	G12	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[75]

**Table 2** continued

Strain name	Genotypes											Ref.
	VP7	VP4	VP6	VP1	VP2	VP3	NSP1	NSP2	NSP3	NSP4	NSP5	
RVA/Human-wt/BGD/Dhaka25-02/2002/G12P[8]	G12	P[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	[75]
RVA/Human-tc/THA/T152/1998/G12P[9]	G12	P[9]	I3	R3	C3	M3	A12	N3	T3	E3	H6	[75]
RVA/Horse-wt/ARG/E403/2006/G14P[12]	G14	P[12]	I2	R2	C2	M3	A10	N2	T3	E2	H7	NP
RVA/Cow-wt/ARG/B383/1998/G15P[11]	G15	P[11]	I2	R5	C2	M2	A13	N2	T6	E12	H3	[58]
RVA/Cow-tc/IND/Hg18/XXXX/G15P[21]	G15	P[21]								E2		[80]
RVA/Mouse-tc/BRA/EHP/1981/G16P[20]	G16	P[20]					A7			E7		[26]
RVA/Turkey-tc/IRL/Ty-1/1979/G17P[17]	G17	P[17]	I4					N4		E4		[68]
RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	G18	P[17]	I4	R4	C4	M4	A4	N4	T4	E4	H4	[38]
RVB/Rat-tc/USA/IDIR/1984/G1P[X] <sup>b</sup>	G1 <sup>b</sup>											[98]
RVB/Human-wt/CHN/WH-1/2002/G2P[X] <sup>b</sup>	G2 <sup>b</sup>											[106]
RVB/Human-wt/BGD/Bang117/2003/G2P[X] <sup>b</sup>	G2 <sup>b</sup>											[103]
RVB/Cow-wt/IND/DB176/2001/G3P[X] <sup>b</sup>	G3 <sup>b</sup>											[11]
RVC/Pig-tc/USA/Cowden/1982/G1P[1] <sup>b</sup>	G1 <sup>b</sup>	P[1] <sup>b</sup>										[74]
RVC/Cow-tc/JPN/Shintoku/1991/G2P[3] <sup>b</sup>	G2 <sup>b</sup>	P[3] <sup>b</sup>										[94]
RVC/Human-tc/GBR/Bristol/1998/G4P[2] <sup>b</sup>	G4 <sup>b</sup>	P[2] <sup>b</sup>										[37]
RVC/Human-wt/BGD/BS347/2005/G4P[2] <sup>b</sup>	G4 <sup>b</sup>	P[2] <sup>b</sup>										[104]
RVD/chicken-wt/DEU/05V0049/2005/GXP[X]												[92]

An open space means the (1) genotype not known because sequencing has not been performed or (2) genotypes have not yet been established for non-group-A RVs. An “X” in the name of a strain indicates that this information is missing. Accession numbers for representative strains of all the different genotypes can be found in Table 3

NP not published

<sup>a</sup> Only the partial VP8\* coding region of the VP4 sequence of P[22] strains is currently available

<sup>b</sup> Genotypes are provisional. Guidelines about the classification of RV strains belonging to RVB and RVC will be determined in the near future by the RCWG

<sup>c</sup> Due to a typographic error, the NSP3 genotype of strain NIC522 was previously misidentified as T2 [5]

- *RV group/species*: RVA, RVB, RVC, RVD, RVE, RVF or RVG. RVX should be used in cases where a strain has not yet been assigned to an established or new RV group/species.
- *Species of origin*: The “species of origin” field contains two components. The first component is human, pig, cow, dog, cat, rhesus, simian, horse, rabbit, mouse, goat, sheep, guanaco, bat, turkey, chicken, pheasant, sugar glider or other, as appropriate. In cases in which the sample was obtained from the environment (i.e., sewage, soil, river, lake, ocean or similar), “Env” should be used. If the species of origin is unknown, “X” should be used. The second component is “-wt” (for wild-type) to distinguish from tissue-culture-adapted (-tc) and laboratory-generated or -engineered (-lab) RV strains. If a tissue-culture-adapted or laboratory-generated strain is introduced into the population (e.g., a vaccine strain) and later recovered from a human or animal, the species from which the sample was recovered should be used, followed by “-wt”. The researcher should be able to give a name to a strain

without the need to perform any analyses. If subsequent analyses reveal that a strain is wholly or partially derived from a vaccine strain, this information can be added in the metadata of the record in GenBank or any other database.

- *Country of identification*: A unique 3-letter abbreviation code is used for each country as listed on the website <https://www.cia.gov/library/publications/the-world-factbook/appendix/appendix-d.html>. If the country of identification is unknown, “XXX” should be used. If a tissue-culture-adapted or laboratory-generated RV strain is introduced into the population and later recovered from a human or animal, the country from which the sample was recovered should be used (further details, e.g., of city/site of identification, can be added in the common name or the metadata; see below).
- *Common name*: This is a short name given by investigators, preferentially not using forward slashes (“/”) or backslashes (“\”) to avoid confusion and computer-processing problems. If appropriate, the common name can contain geographically/clinically

**Table 3** Reference strains and accession numbers representing the 166 currently established genotypes

Geno-type	Reference Strain	Accession number	Geno-type	Reference Strain	Accession number
<b>VP7</b>					
G1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	K02033	P[17]	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	AB009632
G2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	AB118023	P[18]	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712558
G3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	D86271	P[19]	RVA/Human-wt/IND/RMC321/1990/G9P[19]	AF523677
G4	RVA/Human-tc/GBR/ST3/1975/G4P2A[6]	X13603	P[20]	RVA/Mouse-tc/XXX/EHP/1981/G16P[20]	U08424
G5	RVA/Pig-tc/USA/OSU/1977/G5P9[7]	X04613	P[21]	RVA/Cow-tc/IND/Hg18/1995/G15P[21]	AF237665
G6	RVA/Cow-tc/FRA/RF/1982/G6P[1]	X65940	P[22]	RVA/Rabbit-wt/ITA/160-01/2001/G3P[22]	AF526374
G7	RVA/Turkey-tc/IRL/Ty-3/1979/G7P[17]	AB080737	P[23]	RVA/Pig-tc/VEN/A34/1985/G5P[23]	AY174094
G8	RVA/Human-tc/IND/69M/1980/G8P4[10]	EF672560	P[24]	RVA/Rhesus-tc/USA/TUCH/2002/G3P[24]	AY596189
G9	RVA/Human-tc/USA/WI61/1983/G9P1A[8]	AB180969	P[25]	RVA/Human-wt/BGD/Dhaka6/2001/G11P[25]	GU199520
G10	RVA/Human-tc/GBR/A64/1987/G10P11[14]	A01321	P[26]	RVA/Pig-wt/ITA/134-04-15/2003/G5P[26]	DQ061053
G11	RVA/Pig-tc/MEX/YM/1983/G11P9[7]	M23194	P[27]	RVA/Pig-wt/THA/CMF034/2000/G2P[27]	DQ534016
G12	RVA/Human-tc/PHL/L26/1987/G12P[4]	M58290	P[28]	RVA/Human-wt/ECU/Ecu534/2006/G20P[28]	EU805773
G13	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712560	P[29]	RVA/Cow-wt/JPN/Azuk-1/2006/G21P[29]	AB454420
G14	RVA/Horse-wt/ARG/E403/2006/G14P[12]	JF712582	P[30]	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	EU486956
G15	RVA/Cow-wt/ARG/B383/1998/G15P[11]	FJ347116	P[31]	RVA/Chicken-tc/DEU/06V0661/2006/G19P[31]	EU486962
G16	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479955	P[32]	RVA/Pig-wt/IRL/61-07-ire/2007/G2P[32]	FJ492835
G17	RVA/Turkey-tc/IRL/Ty-1/1979/G17P[17]	S58166	P[33]	RVA/Cow-tc/JPN/Dai-10/2007/G24P[33]	AB513836
G18	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	D82979	P[34]	RVA/Pig-wt/JPN/FGP51/2009/G4P[34]	AB571047
G19	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	FJ169861	P[35]	RVA/Turkey-tc/DEU/03V0002E10/2003/G22P[35]	EU486958
G20	RVA/Human-wt/ECU/Ecu534/2006/G20P[28]	EU805773	<b>VP6</b>		
G21	RVA/Cow-wt/JPN/Azuk-1/2006/G21P[29]	AB454421	I1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	K02086
G22	RVA/Turkey-tc/DEU/03V0002E10/2003/G22P[35]	EU486973	I2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	DQ870507
G23	RVA/Pheasant-wt/HUN/Phea14246/2008/G23P[?]	FN393054	I3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	DQ490538
G24	RVA/Cow-tc/JPN/Dai-10/2007/G24P[33]	AB513837	I4	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	D16329
G25	RVA/Bat-wt/KEN/KE4852/07/2007/G25P[6]	GU983676	I5	RVA/Pig-tc/MEX/YM/1983/G11P9[7]	X69487
G26	RVA/Pig-wt/JPN/TJ4-1/2010/G26P[?]	AB605258	I6	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712559
G27	RVA/SugarGlider-wt/JPN/SG33/2010/G27P[X]	AB621363	I7	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479952
<b>VP4</b>					
P[1]	RVA/Cow-tc/FRA/RF/1982/G6P[1]	U65924	I8	RVA/Human-wt/THA/CMH222/2001/G3P[3]	ABC41660
P[2]	RVA/Simian-tc/ZAF/SA11-H96/1958/G3P5B[2]	DQ841262	I9	RVA/Rhesus-tc/USA/TUCH/2002/G3P[24]	AY594670
P[3]	RVA/Dog-tc/AUS/K9/1981/G3P[3]	D14725	I10	RVA/Sheep-tc/CHN/Lamb-NT/XXXX/G10P[15]	FJ032028
P[4]	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	AJ540227	I11	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	DQ096805
P[5]	RVA/Cow-tc/GBR/UK/1973/G6P7[5]	M22306	I12	RVA/Human-wt/NPL/KTM368/2004/G11P[25]	GU199496
P[6]	RVA/Human-tc/GBR/ST3/1975/G4P2A[6]	L33895	I13	RVA/Human-wt/ECU/Ecu534/2006/G20P[28]	EU805774
P[7]	RVA/Pig-tc/USA/OSU/1976/G5P9[7]	M33516	I14	RVA/Pig-wt/CAN/CE-M-06-0003/2006/G2P[27]	GU183245
P[8]	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	L34161	I15	RVA/Bat-wt/KEN/KE4852/07/2007/G25P[6]	GU983675
P[9]	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	D10970	I16	RVA/Human-tc/KEN/B10/1987/G3P[2]	HM627557
P[10]	RVA/Human-tc/IND/69M/1980/G8P4[10]	M60600	<b>VP1</b>		
P[11]	RVA/Human-tc/IND/116E/1985/G9P[11]	L07934	R1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	DQ490539
P[12]	RVA/Horse-wt/ARG/E30/1993/G3P[12]	JF712575	R2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	DQ870505
P[13]	RVA/Human-wt/IND/HP140/1987/G6P[13]	DQ003291	R3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	DQ490533
P[14]	RVA/Human-tc/GBR/A64/1987/G10P11[14]	EF672563	R4	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	AB009629
P[15]	RVA/Sheep-tc/CHN/Lamb-NT/XXXX/G10P[15]	FJ031027	R5	RVA/Guanaco-wt/ARG/Chubut/1999/G8P[14]	FJ347100
P[16]	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479950	R6	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	FJ169853
			R7	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479947
			R8	RVA/Human-tc/KEN/B10/1987/G3P[2]	HM627553

important information or specific (engineered) mutations, as long as the number of characters is less than or equal to 15.

- *Year of identification*: This is given using the “yyyy” format. If the year of identification is unknown, “XXXX” should be used. If a tissue-culture-adapted or laboratory-generated strain is introduced into a population and later recovered from a human or animal, the year in which the sample was recovered should be used.
- *G-type*: This is given using the form G<sub>x</sub>, where x is the established G genotype/serotype number. If the G-type is unknown, GX should be used.
- *P-type*: This is given using the form P<sub>y</sub>[z], where y is the established P serotype number and z [in square brackets] is the established P genotype number. If the P-serotype is unknown, P[z] should be used, and if the P-genotype is unknown, P<sub>y</sub>[X] or P[X] should be used.



Table 3 continued

Geno-type	Reference Strain	Accession number	Geno-type	Reference Strain	Accession number
<b>VP1 continued</b>			N6	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	FJ169860
R9	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712555	N7	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479954
<b>VP2</b>			N8	RVA/Bat-wt/KEN/KE4852/07/2007/G25P[6]	GU983677
C1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	X14942	N9	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712562
C2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	DQ870506	<b>NSP3</b>		
C3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	DQ490536	T1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	X81434
C4	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	AB009630	T2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	EF136660
C5	RVA/Simian-tc/ZAF/SA11-H96/1958/G3P5B[2]	DQ838635	T3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	DQ490535
C6	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	FJ169854	T4	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	AB009626
C7	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479948	T5	RVA/Simian-tc/ZAF/SA11-H96/1958/G3P5B[2]	DQ838610
C8	RVA/Bat-wt/KEN/KE4852/07/2007/G25P[6]	GU983673	T6	RVA/Cow-tc/USA/WC3/1981/G6P[5]	EF990701
C9	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712556	T7	RVA/Cow-tc/GBR/UK/1973/G6P7[5]	K02170
<b>VP3</b>			T8	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	FJ169859
M1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	AY267335	T9	RVA/Cow-wt/JPN/Azuk-1/2006/G21P[29]	AB513838
M2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	AY277914	T10	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479953
M3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	DQ490537	T11	RVA/Bat-wt/KEN/KE4852/07/2007/G25P[6]	GU983678
M4	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	AB009631	T12	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712563
M5	RVA/Simian-tc/ZAF/SA11-H96/1958/G3P5B[2]	DQ838645	<b>NSP4</b>		
M6	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712557	E1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	K02032
M7	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	FJ169855	E2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	AF174305
M8	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479949	E3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	D89873
<b>NSP1</b>			E4	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	AB009627
A1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	L18943	E5	RVA/Human-wt/BEL/B4106/2000/G3P[14]	AY740732
A2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	L18945	E6	RVA/Human-wt/BGD/N26/2000/G12P[6]	DQ146691
A3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	D45244	E7	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479956
A4	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	AB009633	E8	RVA/Cow-lab/GBR/PP-1/1976/G3P[7]	AF427521
A5	RVA/Simian-tc/ZAF/SA11-H96/1958/G3P5B[2]	DQ838599	E9	RVA/Pig-wt/THA/CMP034/2000/G2P[27]	AF427521
A6	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712561	E10	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	FJ169862
A7	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479951	E11	RVA/Turkey-tc/IRL/Ty-3/1979/G7P[17]	AB065286
A8	RVA/Pig-tc/MEX/YM/1983/G11P9[7]	BAA20545	E12	RVA/Guanaco-wt/ARG/Chubut/1999/G8P[14]	FJ347109
A9	RVA/Human-wt/BEL/B4106/2000/G3P[14]	AY740735	E13	RVA/Human-tc/KEN/B10/1987/G3P[2]	HM627562
A10	RVA/Horse-wt/ARG/E30/1993/G3P[12]	JF712572	E14	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712564
A11	RVA/Human-wt/HUN/Hun5/1997/G6P[14]	EF554110	<b>NSP5</b>		
A12	RVA/Human-tc/THA/T152/1998/G12P[9]	AB097459	H1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	AF306494
A13	RVA/Cow-wt/ARG/B383/1998/G15P[11]	FJ347117	H2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	M33608
A14	RVA/Cow-tc/THA/A5-13/XXXX/G8P[1]	D38148	H3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	AB008656
A15	RVA/Human-tc/ITA/PA260-97/1997/G3P[3]	HQ661118	H4	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	AB009628
A16	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	FJ169857	H5	RVA/Simian-tc/ZAF/SA11-H96/1958/G3P5B[2]	DQ838630
<b>NSP2</b>			H6	RVA/Human-tc/THA/T152/1998/G12P[9]	DQ146706
N1	RVA/Human-tc/USA/Wa/1974/G1P1A[8]	L04534	H7	RVA/Horse-wt/ARG/E30/1993/G3P[12]	JF712575
N2	RVA/Human-tc/USA/DS-1/1976/G2P1B[4]	L04529	H8	RVA/Chicken-tc/DEU/02V0002G3/2002/G19P[30]	FJ169863
N3	RVA/Human-tc/JPN/AU-1/1982/G3P3[9]	DQ490534	H9	RVA/Mouse-tc/USA/ETD_822/XXXX/G16P[16]	GQ479957
N4	RVA/Pigeon-tc/JPN/PO-13/1983/G18P[17]	AB009625	H10	RVA/Bat-wt/KEN/KE4852/07/2007/G25P[6]	GU983680
N5	RVA/Simian-tc/ZAF/SA11-H96/1958/G3P5B[2]	DQ838615	H11	RVA/Horse-tc/GBR/L338/1991/G13P[18]	JF712565

An “X” in the name of a strain indicates that this information is missing

Examples of strain names using this nomenclature system can be found in Tables 1 and 2.

#### B) Guidelines for *tissue culture-adapted* RV strains or RV strain passed *in vivo* in their homologous host species

These guidelines apply to any non-vaccine RV strain that has been adapted in tissue culture or passed *in vivo* in a homologous animal model without the intention to introduce specific changes in the genome sequence.

Specific details about the number of passages and the cell lines or animal used should be provided in the metadata. Guidelines for nomenclature of i) RV vaccine strains, ii) RV strains that have been generated in the laboratory by reassortment and iii) RV strains engineered in the laboratory by reverse genetics or equivalent mechanisms are provided below in sections C), D) and E).

- *RV group/species*: See section A) above.
- *Species of origin*: The “species of origin” field contains two components. The first component is human, pig,

cow, dog, cat, rhesus, simian, horse, rabbit, mouse, goat, sheep, bat, guanaco, turkey, chicken, pheasant, sugar glider or other, as appropriate. If the sample was obtained from the environment (i.e., sewage, soil, river, lake, ocean or similar), “Env” should be used. If the species of origin is unknown, “X” should be used. The second component is “-tc” (for tissue cultured) or “-hhp” (homologous host passaged) to distinguish from wild-type (-wt) and laboratory-generated or engineered (-lab) RV strains. If a tissue-culture-adapted or *in vivo*-passaged RV strain is introduced into the population (e.g., a vaccine strain) and later recovered from a human or animal, the species from which the sample was recovered should be used followed by “-wt”.

- *Country of identification*: A unique 3-letter abbreviation code is used for each country as listed on the website <https://www.cia.gov/library/publications/the-world-factbook/appendix/appendix-d.html>. If the country of identification is unknown, “XXX” should be used. If a tissue-culture-adapted RV strain is introduced into the population and later recovered from a human or animal, the country from which the sample was recovered should be used (further details, e.g., of city/site of identification, can be added to the common name or the metadata, see below).
- *Common name*: See section A) above
- *Year of identification*: This is given using the “yyyy” format. If the year of identification is unknown, “XXXX” should be used. If a tissue-cultured strain is introduced into a population and later recovered from a human or animal, the year in which the sample was recovered should be used.
- *G-type*: See section A) above.
- *P-type*: See section A) above.

Examples of strain names using this nomenclature system can be found in Tables 1 and 2.

#### C) Guidelines for RV strains *generated in a laboratory* for which a host species can be assigned unambiguously

These guidelines apply to any RV strain in which small deliberate changes have been introduced (examples: chemically mutagenised RV strains, moderately modified RV strains using reverse genetics techniques, etc.) or RV strains that were passaged in a heterologous animal model and for which there is no ambiguity about the original host species. Specific details about the introduced changes and the methods used should be provided in the metadata. Guidelines for nomenclature for i) RV vaccine strains, ii) RV strains that have been generated in the laboratory by reassortment, iii) RV strains with a designed synthetic sequence generated using reverse genetics or iv) RV strains

in which large modifications have been applied using reverse genetics or equivalent mechanisms are provided below in sections D) and E).

- *RV group/species*: See section A) above.
- *Species of origin*: The “species of origin” field contains two components. The first component is human, pig, cow, dog, cat, rhesus, simian, horse, rabbit, mouse, goat, sheep, bat, guanaco, turkey, chicken, pheasant, sugar glider or other, as appropriate. If the sample was obtained from the environment (i.e., sewage, soil, river, lake, ocean or similar), “Env” should be used. If the species of origin is unknown, “X” should be used. The second component is “-lab” (lab engineered) to distinguish from wild-type (-wt) and tissue-culture (-tc)-adapted RV strains. If a laboratory-generated strain is introduced into the population (e.g., a vaccine strain) and later recovered from a human or animal, the species from which the sample was recovered should be used followed by “-wt”.
- *Country of identification*: A unique 3-letter abbreviation code is used for each country as listed on the website <https://www.cia.gov/library/publications/the-world-factbook/appendix/appendix-d.html>. If the country of identification is unknown, “XXX” should be used. If a laboratory-generated strain is introduced into the population and later recovered from a human or animal, the country from which the sample was recovered should be used (further details, e.g., of city/site of identification, can be added to the common name or the metadata, see below).
- *Common name*: See section A) above.
- *Year of identification*: This is given using the “yyyy” format. If the year of identification is unknown, “XXXX” should be used. If a laboratory-engineered strain is introduced into a population and later recovered from a human or animal, the year in which the sample was recovered should be used.
- *G-type*: See section A) above.
- *P-type*: See section A) above.

Examples of strain names using this nomenclature system can be found in Table 2.

#### D) Guidelines for RV strains *generated in a laboratory* for which a host species cannot be assigned unambiguously

These guidelines for nomenclature apply to RV strains that have been generated in the laboratory by reassortment (except for vaccine strains), strains with a designed synthetic sequence generated using reverse genetics, strains in which large modifications have been applied using reverse genetics or RV strains generated by combinations of the above procedures. Specific details about the changes

**Table 4** Hypothetical examples of nomenclature for RV vaccine strains and laboratory-generated/engineered strains

Strain name	Genotypes											Ref.
	VP7	VP4	VP6	VP1	VP2	VP3	NSP1	NSP2	NSP3	NSP4	NSP5	
RVA/Env/CHE/River-Swiss/1998/G3P[X] <sup>a</sup>	G1											[3]
RVA/Simian-lab/USA/SA11-tsE-1400/1982/G3P[2] <sup>b</sup>	G3	P[2]	I2	R2	C5	M5	A5	N2	T5	E2	H5	[78]
RVA/Cow-lab/GBR/PP-1/1976/G3P[7] <sup>c</sup>	G3	P[7]					A3			E8		[22]
RVA/Labstr/USA/RRV-E4/1996/G3P[20] <sup>d</sup>	G3	P[20]	I2	R2	C3	M3	A9	N2	T3	E3	H6	[46]
RVA/Labstr/USA/SA11-huN2/2010/G3P[2] <sup>e</sup>	G3	P[2]	I2	R2	C5	M5	A5	N2	T5	E2	H5	[90]
RVA/Vaccine/USA/RotaTeq-WI79-9/1992/G1P7[5] <sup>f</sup>	G1	P7[5]	I2	R2	C2	M1	A3	N2	T6	E2	H3	[60]
RVA/Vaccine/USA/RotaTeq-SC2-9/1992/G2P7[5] <sup>f</sup>	G2	P7[5]	I2	R2	C2	M1	A3	N2	T6	E2	H3	[60]
RVA/Vaccine/USA/RotaTeq-WI78-8/1992/G3P7[5] <sup>f</sup>	G3	P7[5]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[60]
RVA/Vaccine/USA/RotaTeq-BrB-9/1996/G4P7[5] <sup>f</sup>	G4	P7[5]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[60]
RVA/Vaccine/USA/RotaTeq-WI79-4/1992/G6P1A[8] <sup>f</sup>	G6	P1A[8]	I2	R2	C2	M2	A3	N2	T6	E2	H3	[60]
RVA/Vaccine/USA/Rotarix-RIX4414/1988/G1P1A[8] <sup>g</sup>	G1	P1A[8]	I1	R1	C1	M1	A1	N1	T1	E1	H1	NP

An open space means that the genotype has not been established/determined

<sup>a</sup> Hypothetical name of an RV strain identified in a river in Switzerland [3]

<sup>b</sup> Strain SA11-tsE is a temperature-sensitive RV strain (SA11) generated using chemical mutagenesis [78]

<sup>c</sup> Strain PP-1 is a bovine RV strain that has been passaged several times in the heterologous pig model [22]

<sup>d</sup> Hypothetical name of a RV strain generated using *in vitro* reassortment in cell culture, possessing an RRV-like gene background with the VP4 gene (genotype P[20]) of the murine RV strain EHP [46]

<sup>e</sup> Hypothetical name of a RV strain generated using reverse genetic techniques, possessing an SA11-like gene background with the NSP2 gene (genotype N2) of the human RV strain DS-1 [90]

<sup>f</sup> Hypothetical names of the 5 reassortant RV strains present in the RotaTeq vaccine [60]

<sup>g</sup> Hypothetical name of the RV strain present in the Rotarix vaccine

introduced and the methods used should be provided in the metadata.

- *RV group/species*: See section A) above.
- *Species of origin*: “LabStr”. More specific details about the procedure used to create the virus should be provided in the metadata.
- *Country of identification*: The country in which the RV strain was generated should be provided according to the unique 3-letter abbreviation code for each country: <https://www.cia.gov/library/publications/the-world-factbook/appendix/appendix-d.html>.
- *Common name*: This is a short name given by investigators, preferentially without the use of forward slashes (“/”) or backslashes (“\”) to avoid confusion. If appropriate, the common name can contain specific information about the laboratory strain, such as the procedure used to generate it, as long as the number of characters is less than or equal to 15.
- *Year of identification*: The year in which the RV strain was generated should be provided using the “yyyy” format.
- *G-type*: See section A) above.
- *P-type*: See section A) above.

Examples using this nomenclature system for naming laboratory strains can be found in Table 4.

## E) Guidelines for RV vaccine strains

Due to the increasing use of live-attenuated RV vaccine, it will be important to distinguish vaccine strains from naturally-occurring wild-type strains. Therefore, a specific notation for vaccine strains was developed and should be used if the strain is the sole component or part of a multi-component live vaccine. Only the sequences obtained from virus production lots or batches are officially recognized as vaccine strains and should have “vaccine” in their strain name. Other “pre-vaccine” strains or sequences of strains that were used in laboratories and were not generated by sequencing the actual vaccine production lots of a given manufacturer should use the normal guidelines to name their strains. Additional information such as the “pre-vaccine” status of a strain could be added in the “common name” section or in the metadata of the record.

- *RV group/species*: See section A) above.
- *Species of origin*: “Vaccine”. Further specific details about the generation of the vaccine strain should be provided in the metadata.
- *Country of identification*: The country in which the vaccine strain was recovered (in the case of a vaccine containing a tissue-culture-adapted strain) or developed (in the case of a vaccine developed using reassortment or *in vitro* engineering) should be provided according to

the unique 3-letter abbreviation code for each country: <https://www.cia.gov/library/publications/the-world-factbook/appendix/appendix-d.html>.

- *Common name*: This is a short name given by investigators, preferentially without the use of forward slashes (“/”) or backslashes (“\”) to avoid confusion. It is recommended that the commercial name of the vaccine is included in the common name. The number of characters should be less than or equal to 15.
- *Year of identification*: The year in which the RV strain was recovered (in the case of a vaccine containing a tissue-culture-adapted strain) or developed (in the case of a vaccine developed using reassortment or *in vitro* engineering) should be provided using the “yyyy” format.
- *G-type*: See section A) above.
- *P-type*: See section A) above.

Examples using this nomenclature system for naming vaccine strains can be found in Table 4.

### RV resource under development at NCBI

Due to the rapid increase in the number of complete RV genome sequences available, the RCWG is working with NCBI to develop an advanced RV resource, which will include a value-added database and a suite of tools for the analysis of RV sequences. When new sequence records are submitted to GenBank, their contents will be added to the RV database through an automated process. A web-based interface will allow researchers to search the RV database using a variety of genetic, epidemiological and clinical criteria, retrieve relevant sequences and analyze them. With this in mind, the RCWG proposes that the following list of biological descriptors (if available and relevant) should be included with sequence records submitted to GenBank. This list of metadata was agreed upon by members of the RCWG but is by no means exclusive, as additional metadata can also be added.

#### Strain-specific features

- a) Wild-type primary strain
- b) Cell-culture-adapted strain
- c) *In vitro*-generated reassortant strain
- d) Strain engineered by reverse genetics procedures
- e) *In vitro*-generated reassortant RV strain or RV strain engineered by reverse genetics that was recovered in nature from any host (e.g., a vaccine strain recovered in a human infant)
- f) *In vivo*-generated strain resulting from reassortment of a laboratory-generated strain and a wild-type

strain (e.g., a strain recovered from a vaccinated infant that is the result of a reassortment event of the vaccine strain and a wild-type strain).

- In cases b, c and d, more details should be provided to account for any engineered changes. It is important to note that natural reassortant RVs should be categorized as “wild-type strains”.
- Genotypes of the remaining 9 RNA segments according to RCWG guidelines [55].
- Species/group (RVA, RVB, RVC, RVD, RVE, RVF, RVG or RVX)
- For RVA: subgroup (SG, I, II, I+II, non-I/non-II) specificity
- Electropherotype: long, short, super-short or atypical
- Banding pattern (example: 4:2:3:2 for RVA, 4:2:2:3 for RVB, 4:3:2:2 for RVC, etc.)
- Region of identification: country, state, province, city/village
- Latitude / longitude (LAT/LON) coordinates: a Google Earth-like application might be incorporated where the location can be pinpointed on a map, and the coordinates will be added automatically.
- Collection date: time of sample collection (year, month, and day).

#### Host-specific features

- Host species: both the “common English name” and the scientific Latin name
- Host date of birth
- Host age at infection (or date of sample collection)
- Host gender
- Vaccination status of host
- Name of administered vaccine, number of administered doses and vaccination dates
- For animal RV vaccines: adjuvant used (oil or aqueous) and RV genotypes and/or other microbes included in the vaccine.

#### Disease-related and clinical features

- Symptomatic or asymptomatic infection
- Clinical symptoms of host: diarrhea, vomiting, fever, other
- Clinical severity: mild, moderate, severe, using established clinical gastroenteritis severity scoring systems (Vesikari, Clark, other) [17, 28, 83, 84] or World Health Organization (WHO) reference schemes
- Clinical outcome: dehydration (using WHO or Gorelick reference schemes) [36, 101], emergency room or its equivalent (outpatient), hospitalization and/or death

- Sample type: stool, serum, nasal swabs or other systemic tissues (lung, brain, etc.)
- Other co-infecting agents detected
- For RVs detected in domestic animals (calves, foals, piglets, etc.): indicate if the animal received electrolyte solutions, antibiotic, anti-diarrheic or other nonspecific treatment before and/or after sample collection
- Sample derived from a single case of diarrhea or an outbreak. In the case of an outbreak, what were the morbidity and mortality rates?
- For animals indicate type of exploitation (dairy or beef herds; extensive or intensive production; thoroughbred or standard-bred horses, etc.).

#### Analysis-specific information

- Was sequencing performed on stool, rectal swabs, intestinal contents, other types of samples from the patient, environmental samples or cell-culture-adapted virus?
- Cell type (e.g., Vero) and number of times the virus was passaged in cell culture before sequencing
- Sequencing method: Sanger sequencing, pyrosequencing, other
- Termini: were the 5'- and 3'-terminal sequences primer-derived or sequenced de novo?

#### Conclusions

The quickly evolving sequencing capabilities of research laboratories and commercial organizations around the world have resulted in a rapid growth of sequence data for RVs, and dealing with these data has become a major challenge. In an attempt to introduce a systematic method of naming RV strains, the RCWG proposes the following nomenclature: RV group/species of origin/country of identification/common name/year of identification/G- and P-type. In this nomenclature, specific guidelines were developed for wild-type RV strains, tissue-culture-adapted strains, RV vaccine strains and RV strains that have been generated or engineered in a laboratory using reassortment or reverse genetics procedures. The current proposal only leaves room for 15 explanatory characters in the “common name” section of the complete strain name. For all of the other sections (RV group, species of isolation, country of isolation, year of isolation and G/P-genotype), the guidelines are very rigorous. The reason for allowing the freedom of choosing 15 characters in the “common name” section is to provide authors an opportunity to highlight specific or important information. Examples of information that could be added include: i) the city or state of strain

identification, ii) clinical outcomes or viral phenotypic characteristics and iii) mutations or engineered changes in the viral genome. However, we think that it is impossible to include all relevant information about a strain in the actual name. Instead, such information should be added in the metadata linked to each record in GenBank or any other database.

The RCWG is fully aware that it is impossible to define rules for each real-life or hypothetical situation, especially when a strain has undergone several types of manipulations (e.g., a combination of cell culture adaptation, passaging in a homologous host animal and the introduction of reverse-genetic engineered mutations). Therefore, we encourage researchers to choose a strain name that appears to be most appropriate or to contact the RCWG for guidance. While it is not necessary to include in the actual strain name, detailed information about manipulations should be added to the metadata of a record.

This standardized naming procedure is reminiscent of the nomenclature guidelines for influenza viruses and will be very useful for future analysis. However, this system will only succeed if the entire research community supports and adheres to the proposed guidelines. Regarding the use of this new nomenclature in scientific articles, the RCWG suggests that authors use the full strain name the first time it is mentioned in the text and then subsequently use the “common name” (i.e., a suitable abbreviation). In addition, authors should consider using the full strain name in phylogenetic trees, tables and figures if it enhances the interpretation or presentation of the data. However, depending on the individual characteristics of a manuscript, authors have the flexibility to use alternative strategies.

Towards utilizing the large amount of RV nucleotide sequence data, an RV-specific database is being developed by the NCBI, which is expected to be launched in the second half of 2011. This resource will be similar to the NCBI Virus Variation resources developed for influenza and dengue viruses [10, 81] and the resources developed at the Los Alamos National Laboratory for hepatitis C virus (HCV) [43] and for human immunodeficiency virus (HIV) (<http://www.hiv.lanl.gov/content/>). RV nucleotide sequence data, as well as metadata describing the biological and epidemiological context of associated sequences, will be stored in specially designed relational databases. A new, user-friendly web interface will allow investigators to construct and explore resource queries based on a variety of criteria. Additionally, novel web-based tools and displays will allow users to compare retrieved nucleotide and protein sequences on the basis of chronology, geography and biological significance. In order to fully exploit the possibilities of such a resource for the study of viral epidemiology, seasonality, geographical spread, links between specific strains and clinical manifestations, possible

vaccine escape-mutants, etc., it is imperative that nucleotide sequence submissions include relevant metadata. In addition, the RCWG strongly encourages researchers to retrospectively update the names of their RV sequences in GenBank and add relevant metadata to the files. This process is straightforward and will greatly enhance the scientific potential of individual sequence records. From Table 1, it is obvious that close monitoring, updating and genotype assignment activities of the RCWG as practiced during the past three years are absolutely necessary to sustain and develop the new RV genotype-based classification and nomenclature system. With the increasing number of sequence data becoming available for non-RVA strains, the activities of the RCWG will probably be expanded in the near future.

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