# **Template for Taxonomic Proposal to the ICTV Executive Committee Creating Species in an existing genus**

Code <sup>†</sup> 2005.119V.04	To designate the following as species in the genus:	
	_	Hantavirus
	belonging to the family° :	Bunyaviridae
	<i>Saaremaa virus</i> Saaremaa virus – 160V	(SAAV-160) AJ410618
<sup>†</sup> Assigned by ICTV officers		

° leave blank if inappropriate or in the case of an unassigned genus

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## **Old Taxonomic Order**

Order		
Family	Bunyaviridae	
Genus	Hantavirus	
Type Species		ntaan virus
Species in the Genus		22 altogether
Tentative Species in the Genus		none identified
Unassigned Species in the family		none identified
New Taxe	onomic Order	

#### Order

Family	Bunyaviridae	
Genus	Hantavirus	
Type Species	Hantaan virus	
Species in the Genus	23 altogether (SAAV included)	
Tentative Species in the Genu	s none identified	
Unassigned Species in the fan	nily none identified	
<b>ICTV-EC</b> comments and	d response of the SG	

# Species demarcation criteria in the genus

Hantavirus species (from the VIIth report of ICTV):

- are found in a unique ecological niche, i.e. in a different primary rodent reservoir species or subspecies;
- (2) exhibit at least a 7% difference in an identity on comparison of the complete GPC and N protein sequences;
- (3) show at least a 4-fold difference in two-way cross-neutralization tests; and
- (4) do not naturally form reassortants with other hantavirus species

# Argumentation to justify the designation of new species in the genus

When **Saaremaa virus** (SAAV) was identified and subsequently isolated from the striped field mouse (*Apodemus agrarius*) trapped on Saaremaa island, Estonia, it was regarded as an *A. agrarius* -carried variant of *Dobrava virus* DOBV [1,2]. Later it was observed that SAAV and DOBV are sympatric: in Slovenia [3] and Slovakia [4] these hantaviruses were found to be steadily maintained in their respective hosts, field and yellow-necked mice, that inhabited the same localities. The findings were interpreted as evidence of a reproductive isolation of SAAV from DOBV [5]. Serological study revealed 4-fold or higher cross-neutralization titers to DOBV or SAAV in the majority of human sera from Estonia and the Balkans, thus proving that they represent distinct hantavirus serotypes [6].

To summarize, SAAV fulfills the species demarcation criteria 1, 3 and 4 and comes close to the ARBITRARY SELECTED cutoff level of 7% in the aa identity of the GPC (6.1%) [5]. Notably, the two hantaviruses seem to possess different pathogenicity for humans and laboratory mice. In humans, DOBV causes severe HFRS with a case fatality rate of 8-12% [7, 8] while SAAV has not so far been associated with fatal HFRS (and in some areas the seroprevalence is higher that 3% [9]). In suckling mice, DOBV infection is lethal while SAAV infection is not [10].

In our opinion, we are dealing here with a case of host switching, which occurred in the evolution of these hantaviruses [11]. Phylogenetic analysis revealed a discrepancy in the relationships of DOBV, SAAV and Hantaan virus and their respective rodent hosts. This discrepancy is consistent with the transmission of (pre)DOBV/SAAV between *A. flavicollis* and *A. agrarius*, which resulted in the establishing of SAAV *via* its ecological and reproductive isolation from DOBV. Crucially, the estimated time-point of the host switching, 2.7-4.0 MYA, was closer to the present than the expected time of split between the two *Apodemus* species (<6.5MYA).

Saaremaa virus

## References

- 1. **Plyusnin et al.** Dobrava hantavirus in Estonia: does the virus exist throughout Europe? Lancet **349**:1369-1370, 1997.
- 2. Nemirov et al. Isolation and characterization of Dobrava hantavirus carried by the striped field mouse (*Apodemus agrarius*) in Estonia. J. Gen. Virol., 80:371-379, 1999.
- 3. **Avsic-Zupanc et al.** Genetic analysis of wild-type Dobrava hantavirus in Slovenia: coexistence of two distinct lineages within the same natural foci. J Gen Virol 81:1747-1755, 2000.
- 4. **Sibold et al.** Dobrava hantavirus causes hemorrhagic fever with renal syndrome in central Europe and is carried by two different *Apodemus* mice species. J Med Virol 63:158-167, 2001
- 5. **Plyusnin A.** 2002. Genetics of hantaviruses: implications to taxonomy (review). Arch Virol, 147:665-682.
- 6. **Brus-Sjölander et al.** Serological divergence of Dobrava and Saaremaa hantaviruses: evidence for two distinct serotypes. J. Epidemiol. Infect. **128**:99-103, 2002.
- 7. **Papa et al.** Retrospective serological and genetic study of the distribution of hantaviruses in Greece. J Med Virol 55:321-327, 1998.
- 8. **Avzic-Zupanc et al.** Hemorrhagic fever with renal syndrome in the Dolenjska region of Slovenia a ten-year survey. Clin Infect Dis 28:860-865, 1999.
- 9. Golovljova et al. Hantaviruses in Estonia. J. Med. Virol. 68:589-598, 2002.
- 10. **Klingström J,** Hantaviruses animal models, immunology and pathogenesis. PhD Thesis, Karolinska Institute, 2004.
- Nemirov et al. Phylogenetic evidence for host switching in the evolution of hantaviruses carried by *Apodemus* mice. Virus Res. 90: 207-215, 2002 (Corrigendum: Virus Res. 92:125-126, 2003).

# Annexes: