

Virus Name: Bluetongue		Abbreviation: BLUV
Status Arbovirus	Select Agent No	SALS Level 2
SALS Basis Results of SALS surveys and information from the Catalogue.		
Other Information USDA Permit Required, USDA Restricted, USDA High Consequence Agent		
Antigenic Group Bluetongue		

SECTION I - Full Virus Name and Prototype Number

Prototype Strain Number / Designation	Accession Number	Original Date Submitted 2/10/1985
Family Reoviridae	Genus Orbivirus	
Information From Arthropod-borne Animal Dis. Res. Lab.	Address Arthropod-borne Animal Disease Lab. USDA, ARS, Denver, Colorado 80225, USA	
Information Footnote Revised		

Section II - Original Source

Isolated By (name) Spreull (26)	Isolated at Institute at Republic of South Africa	
Host Genus Ovine	Species	Host Age/Stage
Sex Not Answered		
<u>Isolated From</u>	<u>Isolation Details</u>	
Whole Blood		
Serum/Plasma		
Organs/Tissues	spleen pulp (26)	
Signs and Symptoms of Illness Fever, facial edema, catarrhal discharge, lesions of mouth, feet, death often due to pneumonia (noncontagious disease)	Arthropod	
Time Held Alive before Inoculation		
Collection Method Blood of sick sheep	Collection Date 6/6/1901	
Place Collected (Minimum of City, State, Country) South Africa (Cape Colony)		
Latitude	Longitude	
Macrohabitat	Microhabitat	Method of Storage until Inoculated Defibrinated blood + glycerin + phenol (virulent to 174 days)
Footnotes		

Section III - Method of Isolation

Inoculation Date		
Animal (Details will be in Section 6) Ovine** ** or goat		
Route Inoculated	Reisolation Yes	
Other Reasons South Africa; repeated isolation; vaccine production (28); plurality of strains (23)		
Homologous Antibody Formation by <u>Source Animal</u>		
Test(s) Used		
Footnotes		

Section IV - Virus Properties

Physicochemical RNA, Double Strand
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Pieces (number of genome segments) 10(31)	Infectivity	Sedimentation Coefficients (s) (S)
Percentage wt, of Virion Protein	Lipid	Carbohydrate
Virion Polypeptides: Number 7 (14)	Details The core has 2 major and 3 minor polypeptides and is surrounded by outer capsid layer with 2 major polypeptides, No. 2 and 5; No. 2 appears to be main determinate of serotype specificity. (14).	
Non-virion Polypeptides: Number 2(12);3(25)	Details Polypeptide 5A associated with a tubular-like	
Virion Density 1.36 gm/cm³ (20) 1.38gm/cm³ in cesium chloride	Sedimentation Coefficients(s) 550 (20)(S)	
Nucleocapsid Density 1.38 (20) 1.42(32) in cesium chloride	Sedimentation Coefficients(s) 470 (20)(S)	

Stability of Infectivity (effects)

pH (infective range)

Lipid Solvent (ether - % used to test)	After Treatment Titer	Control Titer
Lipid Solvent (chloroform)	After Treatment Titer	Control Titer
Lipid Solvent (deoxycholate)	After Treatment Titer	Control Titer

Other (formalin, radiation)

Resis. to ether, chloroform, deoxycholate; resis. depends on purification (3,10,31)

Virion Morphology

Shape Icosahedral, orbivirus (2)	Dimensions 50-70 nm	
Mean nm	Range nm	
Measurement Method Electron microscopy	Surface Projections/Envelope No envelope present (2); occ. pseudoenvelopes (3)	Nucleocapsid Dimensions, Symmetry Capsid diam.: D= 3 nm; L=69 nm (20); 32 capsomeres

Morphogenesis

Site of Constituent Formation in Cell Cytoplasm	Site of Virion Assembly Cytoplasm	Site of Virion Accumulation Cytoplasm
Inclusion Bodies Cytoplasmic (4)	Other	

Hemagglutination

Hemagglutination Yes	Antigen Source BHK 21 cell culture; fluoro carbon ext + cent. thru sucrose	Erythrocytes (species used) Several (a)
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Yes	DMR-21 cell culture, micro-carbon extra + cent. and sucrose	Several (a)
	gradients (11,30)	
pH Range 6.0-9.0	pH Optimum 6.0-9.0	
Temperature Range 4dC, 22dC, 37dC	Temperature Optimum 4dC, 22dC, 37dC	
Remarks	(a) Sheep, chicken, goose, guinea pig, mouse erythrocytes	
Serologic Methods Recommended	CF, modified direct CF, agar gel precipitin, NT C	
Footnotes	(a) Sheep, chicken, goose, guinea pig, mouse erythrocytes	

Section V - Antigenic Relationship and Lack of Relationship to Other Viruses

Bluetongue virus is the type species for the Orbivirus genus [21]. The orbiviruses are distinguished from the reoviruses by acid lability, slight solvent sensitivity, serology (CF and HI), and capsid structure [2], [21]. In addition, there is no nucleic acid sequence homology between Reovirus members and members of other genera [21]. At present, there are 12 serological subgroups of genus Orbivirus. The BLU subgroup consists of the 23 serotypes of bluetongue virus [34].

Isolation of field virus has been by inoculation into sheep, embryonated chicken egg, or less so, cell culture and suckling mice; the iv inoculation of embryonated chicken eggs has proven the most sensitive isolation system [3], [10]. The agar gel immunodiffusion test has been the most widely used group specific test for bluetongue virus antibody and has detected all serotypes of BLU [35], [36]. The modified direct CF test (MDCF) and IFA tests also have been used as group specific for bluetongue [35], [36].

A plaque-reduction neutralization test differentiates between BLU and EHD virus antibodies; this test is currently used to identify the 23 serotypes of BLU [34]. Various serum neutralization tests have previously been used to differentiate BLU serotypes [3], [10].

Antigenic relationships have been demonstrated between the bluetongue serogroup and the EHD, Eubenberg and Palyam serogroups [22], [36]. Immunological cross-reaction documented between virus-specified tubules of BLU and EHD viruses [13].

Two-way cross-relationships between Ibaraki virus and EHD virus, serotypes 1 and 2, demonstrated by agar gel precipitin and indirect fluorescent antibody tests [5]. By neutralization tests, Ibaraki virus was more closely related to EHD virus, serotype 2 (Alberta strain). Antigenic relationship not observed between Ibaraki virus and four serotypes of BLU then found in USA.

Section VI - Biologic Characteristics

Virus Source (all VERTEBRATE isolates)
 Blood (LV), CNS (LV), spleen (LV), skin lesions (LV), lymph node (LV); fetal: calf spleen; sheep CNS, liver, lung; lab. mice CNS. Bovine and ovine placenta

Lab Methods of Virus Recovery (ALL ISOLATIONS)
 Newborn mice, chick embryos and Vero cell cultures, BHK-21 cell cultures

Cell system (a)	Virus passage history (b)	Evidence of Infection						
		CPE			PLAQUES			Growth Without CPE +/- (g)
		Day (c)	Extent (d)	Titer TCD50/ml (e)	Day (c)	Size (f)	Titer PFU/ml (e)	

Hamster kidney (BHK-21) cell cultures are commonly used. Virus multiplication is noted and is accompanied by the production of CPE (34). Other cell lines used are Vero and L mouse fibroblasts. Bluetongue virus multiplies without CPE in *Aedes albopictus* and *Ae pseudoscutellaris* cell cultures at 27C and 37C. Maximum titers are detected on days 3 and 6 respectively. Viral multiplication was not demonstrated in *Ae aegypti* cell cultures (15).

Vertebrate (species and organ) and arthropod	No. isolations/No. tested	No. with antibody/No. tested Test used	Country and region
<p>Bluetongue is an acute, subacute, and possible chronic virus disease of wild and domestic ruminants (3,10). Sheep often have high morbidity with variable mortality, cattle morbidity is usually low and goat morbidity lower. Although wild ruminants may be severely affected (27), the incrimination of particular species is usually the result of serologic surveys that have shown that a large number of wild ruminants are susceptible (29,33) including epizootics at zoos (9).</p> <p>Culicoides were first incriminated as vectors in South Africa (6). Virus isolations worldwide have incriminated several species from different Culicoides subgenera but many previous isolations are inconclusively labelled because of taxonomic problems within species complexes. These problems and the likely vector species for each geographic area, primarily belonging to the subgenus <i>Avaritia</i>, are discussed (37); the false assessment of a high vector potential for <i>C. brevitarsis</i> in Australia was prevented through carefully related taxonomic, ecologic, and vector competence studies with field populations of the species of the subgenus <i>Avaritia</i> (38). The primary vector of BLU in the USA is <i>C. variipennis</i>, (<i>Monoculicoides</i>), which is probably a species complex and is the only BLU vector species that has been colonized (17). It has been suggested for South Africa that <i>Culicoides</i> may serve as overwintering BLU hosts in endemic areas with mild winters (24); but pathogenesis studies with <i>C. variipennis</i> reconfirmed the unlikelyhood of transovarian transmission of bluetongue virus (39).</p> <p><i>Amblyomma variegatum</i></p>	3		Guinea (41)

Experimental host and age	Passage history and strain	Inoculation Route-Dose	Evidence of infection	AST (days)	Titer log ₁₀ /ml
Mice (nb)		ic			
Mice (nb)		ip			
Mice (nb)		sc			
Mice (wn)		ic			
Mice (wn)		ip			

Experimental hosts are sheep, cattle, goats, some wild ruminants, suckling mice, and hamsters, embryonating chicken eggs, and certain cell lines. CF antigen is usually prepared from baby mice, or cell cultures.

Experimental results depend on the strain of BLU used (3,10).

Several findings promoted experimental BLU research (3,10): the showing of the plurality of strains (23), the successful use of the propagation of BLU in embryonated chicken eggs (1), the development of cell culture as a neutralization assay system (8), and the demonstration for wild and colonized populations of *C. variipennis* that this species was a biological vector of bluetongue virus (7). Subsequent research in the USA showed the viremic curves of BLU in sheep and cattle (18). A genetic selection program for oral susceptibility to BLU developed two highly susceptible and a resistant line from the baseline colony of *C. variipennis* (Sonora strain) that was only 30% susceptible (16).

Arthropod species & virus source(a)	Method of Infection log10/ml (b)		Incubation period (c)		Transmission by bite (d)		Assay of arthropod, log10/ml (e)		
	Feeding	Injected	Days	°C	Host	Ratio	Whole	Organ	System

A carrier state of BLU was identified in cattle (19); this state explained an overwintering mechanism for BLU, partially in relation to bites of the vector that increased circulating virus.

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Section X - Histopathology

Character of lesions (specify host)

Natural infection often inflammation: sheep-lesions buccal, nasal and intestinal mucosae, and striated musculature; fetal lambs-brain lesions; cattle-necroses of oral mucosa, teats and udder (3,10)

Inclusion Bodies

Intranuclear

Lower Vertabrates

Organs/Tissues Affected

Skeletal muscles (LV); sheep-death often pneumonia; death of fetus and congenital abnormalities common in lambs and sheep, cattle, goats, wild ruminants; embryonated chicken eggs;suckling mice, hamsters and guinea pigs.

Category of tropism

Striated muscle, mucosa of head and digestive epithelium (3,10)

Section XI - Human Disease

In Nature	Residual	Death
Subclinical	Overt Disease Reported	
Clinical Manifestations		
Number of Cases	Category (i.e. febrile illness, etc.)	

Section XII - Geographic Distribution

Known (Virus detected)

Most of Africa, Near and Far East, and also from USA, Mexico and most parts of Latin America (40).

Suspected (Antibody only detected)

Section XIII - References

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Remarks

First immunization carried out with bluetongue virus in 1901-2 (26).