

Virus Name: La Crosse		Abbreviation: LACV
Status Arbovirus	Select Agent No	SALS Level 2
SALS Basis Results of SALS surveys and information from the Catalogue.		
Other Information		
Antigenic Group Califrona		

SECTION I - Full Virus Name and Prototype Number

Prototype Strain Number / Designation	Accession Number	Original Date Submitted 12/29/1984
Family Bunyaviridae	Genus Bunyavirus	
Information From Wayne H. Thompson	Address Department of Prev. Med., University of Wisconsin, Madison, WI	
Information Footnote Reviewed by editor		

Section II - Original Source

Isolated By (name) W.H. Thompson	Isolated at Institute State Lab of Hygiene, Madison, WI	
Host Genus Man	Species	Host Age/Stage 4 year-old
Sex Female		
<u>Isolated From</u>	<u>Isolation Details</u>	
Organs/Tissues	Brain tissue	
Signs and Symptoms of Illness A fatal case of meningo-encephalitis*	Arthropod	
Time Held Alive before Inoculation		
Collection Method Brain tissues taken at autopsy by B.F.Kalfayan (1)	Collection Date 9/14/1960	
Place Collected (Minimum of City, State, Country) Pathology, La Crosse, Lutheran Hosp., Wisconsin		
Latitude 44° N	Longitude 91° W	
Macrohabitat Child had lived in a forested valley area near the Mississippi river, upper midwestern USA;alt=600ft	Microhabitat	Method of Storage until Inoculated Original tissues frozen in glass jars at -40dC
Footnotes		

Section III - Method of Isolation

Inoculation Date
5/4/1964

Animal (Details will be in Section 6)
nb mice

Route Inoculated ic and ip	Reisolation Yes
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Other Reasons
Sera of the mother, father, and three of the eight other children had antibodies.

Homologous Antibody Formation by Source Animal

Test(s) Used
HI, CF, NT

Footnotes

Section IV - Virus Properties

Physicochemical
RNA

Pieces (number of genome segments)	Infectivity	Sedimentation Coefficients(s) (S)
Percentage wt, of Virion Protein	Lipid	Carbohydrate
Virion Polypeptides: Number	Details	
Non-virion Polypeptides: Number	Details	
Virion Density	Sedimentation Coefficients(s) (S)	
Nucleocapsid Density	Sedimentation Coefficients(s) (S)	

Stability of Infectivity (effects)

pH (infective range)

Lipid Solvent (ether - % used to test) 1:2	After Treatment Titer 2.3 dex	Control Titer 5.8 dex
Lipid Solvent (chloroform)	After Treatment Titer	Control Titer
Lipid Solvent (deoxycholate)	After Treatment Titer	Control Titer
Other (formalin, radiation)		

Virion Morphology

Shape	Dimensions 95 nm	
Mean nm	Range nm	
Measurement Method Electron microscopy (19)	Surface Projections/Envelope Envelope observed	Nucleocapsid Dimensions, Symmetry

Morphogenesis

Site of Constituent Formation in Cell Site of Virion Assembly Site of Virion Accumulation

Inclusion Bodies Other

Hemagglutination

Hemagglutination Antigen Source Erythrocytes (species used)
Yes **SMB ext. by sucrose-acetone** **Goose**

pH Range pH Optimum
6.0-6.2

Temperature Range Temperature Optimum

Remarks

HA activity has not been consistent quality or high enough titer to be used in HI tests. Maximum titer has been 1:40 *
Hospitalized three days after onset of fever, headache, vomiting, convulsion and pleocytosis. ** A suit**

Serologic Methods Recommended

HI, CF, NT, and immunodiffusion tests

Footnotes

HA activity has not been consistent quality or high enough titer to be used in HI tests. Maximum titer has been 1:40 *
Hospitalized three days after onset of fever, headache, vomiting, convulsion and pleocytosis. ** A suit**

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

Results of complement-fixation tests with the human isolate and other members of the CAL group [1].

Antigens	Mouse hyperimmune sera:				
	LAC	BFS283	SSH	TVT	Melao
La Crosse	128/>512	8/>512	8/512	8/512	8/512
BFS-283	32/64	32/128	8/256	8/128	8/256
Snowshoe hare	32/256	8/256	32/256	16/128	8/256
TVT	<8/<8	8/128	<8/<8	64/128	8/64
Melao	32/256	<8/<8	<8/<8	16/128	32/256

Antigen titer/antibody titer

Results of additional comparative CF, NT and immunodiffusion tests have since been reported also indicating serologic relationship of the La Crosse isolate to, but antigenic differences from, other California group viruses. [4], [5], [6]
Results of neutralization tests with antisera to isolates from Wisconsin mosquitoes [7].

Virus

Antisera	La Crosse (199 TCID50)	Snowshoe hare (31 TCID50)	Jamestown Canyon (31 TCID50)	Trivittatus (63 TCID50)
W-1805-64	40 ^a	20	20	10
W-2827-65	120	60	<10	<10
W-2985-65	160	20	80	20
W-401-67	640	40	15	30
W-581-67	80	30	30	15
W-968-67	>1280	30	80	80
W-946-67	160	15	20	15
W-1042-68	320	160	80	20
LAC	640	80	20	10
SSH	40	320	20	<10
JC	<10	<10	640	<10
TVT	<10	<1-	20	160

Antisera: prepared in guinea pigs.

(): Dosage of virus in test.

^a Reciprocal of highest dilution which neutralized CPE activity of virus in BHK-21 cells.

<10: 50% of cells alive, but some CPE evident at this dilution.

Section VI - Biologic Characteristics

Virus Source (all VERTEBRATE isolates)
Serum (M)

Lab Methods of Virus Recovery (ALL ISOLATIONS)
Newborn mice

Cell system (a)	Virus passage history (b)	Evidence of Infection							Growth Without CPE +/- (g)
		CPE			PLAQUES				
		Day (c)	Extent (d)	Titer TC50/ml (e)	Day (c)	Size (f)	Titer PFU/ml (e)		
Vero (CL)	SM 4				6	1,4 mm	5.5* (25)		
LLC-MK2 (CL)					3	0.5-2.0- 8.0 mm	6.0 (25)		
BHK-21 (CL)		2-3	CPE						

* Expressed in dex

Vertebrate (species and organ) and arthropod	No. isolations/No. tested	No. with antibody/No. tested Test used	Country and region
Tamias striatus (eastern chipmunk; blood)	7		Wisconsin, USA (21)
Sentinel gray squirrels (<i>Sciurus carolinensis</i> ; blood)	6		Wisconsin, USA (22)
Man (brain)	1		Wisconsin, USA (24)

In addition to the original isolate from the brain tissues of the fatal case of 1960, isolates have also been obtained from insects and the blood of sentinel rabbits caged in forested sites.

Of eight isolates of LAC obtained from mosquitoes in Wisconsin during 1964 through 1968, five were from 1,985 *Aedes triseriatus*, and one each from 35,865 *Ae communis*, 36,800 *Ae trivittatus*, and 1,818 *Culex pipiens* (7). Fourteen more isolates were obtained from *Ae triseriatus* collected in hardwood deciduous forest study areas during 1969-1970. Fifty-nine other isolates reported from or obtained by the CDC from other states, are from various species, but again, highest infection rates were in *Ae triseriatus* (8).

Neutralizing antibodies are present in sera of small forest-dwelling mammals in Wisconsin; in 62/117 (53%) of chipmunk, *Tamias striatus*; 68/176 (39%) of tree squirrels, *Sciurus carolinensis* and *niger*; 2/13 (15%) of cottontail, *Sylvilagus floridans*; 2/40 (5%) of flying squirrels, *Glaucomys volans*; and 0/23 *Peromyscus leucopus* (9).

Although antibodies neutralizing CAL group viruses are commonly found in rural outdoor workers (10), antibodies which neutralize LAC virus in higher titers than with other CAL group viruses from the state (JC, TVT, and SSH) are usually related to those with exposure to the hardwood deciduous forests common to southwestern Wisconsin and portions of other midwestern states. Convalescent serums from cases of California encephalitis consistently neutralize LAC virus in highest titer (14).

Section VIII - Susceptibility to Experimental Infection (include viremia)

Experimental host and age	Passage history and strain	Inoculation Route-Dose	Evidence of infection	AST (days)	Titer log10/ml
Mice (nb)	SMB 3	ic 0.02	Encephalitis, death	2	8.5
Mice (nb)		ip 0.02	---	2-3	8.0
Mice (nb)		sc			
Mice (wn)		ic 0.03	Encephalitis, death	5	7.4
Mice (wn)		ip 0.03	---	6	1.2
<p>Chipmunk and tree squirrels: 5/6 <i>Tamias striatus griseus</i> and 4/5 <i>Sciurus carolinensis</i> developed viremias of 1.5 to 4.4 dex 2 to 5 days after sc inoculation of LAC virus, and all developed neutralizing antibodies after 6 to 17 days (14).</p>					
<p>Rabbits develop viremia after inoculation (20).</p>					

Section IX - Experimental Arthropod Infection and Transmission

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
<i>Aedes triseriatus</i>	Transmitted LAC virus to suckling mice when tested 15 to 17 days following feeding on viremic chipmunks and squirrels.								
<i>Aedes triseriatus</i>	Venereal transmission from male to female. Infected females transmitted by bite to mice and eggs (23).								
	Natural and exp. transovarial transmission demonstrated in <i>Aedes triseriatus</i> (26).								

Section X - Histopathology

Character of lesions (specify host)

Histologic changes included neuronal degeneration and inflammatory response with some perivascular edema in cerebral cortex, some cuffing within the cerebrum and meninges.

Inclusion Bodies

Intranuclear

Organs/Tissues Affected

Category of tropism

Section XI - Human Disease

In Nature

Residual

Death

Significant

Reported

Subclinical

Overt Disease

Clinical Manifestations

Fever (S), headache (S), prostration (R), conjunctival inflammation (R), CNS signs (including encephalitis) (S), CNS Pleocytosis (S), lymphadenopathy (R), convulsions.

Number of Cases

Category (i.e. febrile illness, etc.)

Section XII - Geographic Distribution

Known (Virus detected)

Numerous cases in the USA particularly in OH, WI, MN, IA, IN, IL, and 13 other states as well (8,17).

Suspected (Antibody only detected)

Section XIII - References

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