

Appendix

Was Kaposi's Sarcoma-Associated Herpesvirus Introduced into China via

Ancient Silk Road: An Evolutionary Perspective?

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Appendix Table1. Viral strains used in this analysis

Name	Continent	Country/area	Subtype	Sample year	Access Number
EU-ITA17-C0-1995	Europe	Italy	C0 ^ζ	1995	AF130286.1
EU-ITA18-A0-1995	Europe	Italy	A0 ^ζ	1995	AF130285.1
EU-ITA19-A0-1995	Europe	Italy	A5	1995	AF130284.1
EU-ITA20-A0-1995	Europe	Italy	A5	1995	AF130283.1
EU-ITA21-A0-1995	Europe	Italy	A5	1995	AF130282.1
EU-ITA22-C0-1993	Europe	Italy	C0	1993	AF130281.1
EU-ITA23-A3-1998	Europe	Italy	A3	1998	AF130280.1
EU-ITA24-A3-1998	Europe	Italy	A3	1998	AF130279.1
EU-ITA25-C0-1997	Europe	Italy	C0	1997	AF130278.1
EU-ITA26-C0-1993	Europe	Italy	C0	1993	AF130277.1
EU-ITA27-C0-1995	Europe	Italy	C0	1995	AF130276.1
EU-ITA28-A2-1994	Europe	Italy	A2	1994	AF130275.1
EU-ITA29-C0-1997	Europe	Italy	C3	1997	AF130274.1
EU-ITA30-C0-1996	Europe	Italy	C1	1996	AF130271.1
EU-ITA31-C0-1995	Europe	Italy	C0	1995	AF130270.1
EU-ITA32-A0-1988	Europe	Italy	A0	1988	AF130269.1
EU-ITA36-A4-2003	Europe	Italy	A4	2003	GU097430.1
EU-ITA37-C0-1991	Europe	Italy	C0	1991	GU097425.1
EU-ITA38-A3-2002	Europe	Italy	A3	2002	GU097421.1
EU-ITA39-A4-1990	Europe	Italy	A4	1990	FJ884615.1
EU-ITA40-C3-1991	Europe	Italy	C3	1991	FJ884614.1
EU-ITA41-A0-1985	Europe	Italy	A3	1985	FJ884613.1
EU-ITA42-A1-1984	Europe	Italy	A1	1984	FJ884612.1

EU-ITA43-A3-1983	Europe	Italy	A3	1983	FJ884611.1
EU-GRE01-C0-1994	Europe	Greece	C0	1994	AF130268.1
EU-GRE02-C0-1994	Europe	Greece	C1	1994	AF130267.1
EU-GRE03-C1-1992	Europe	Greece	C1	1992	GU097423.1
EU-GRE04-A3-1992	Europe	Greece	A3	1992	GU097420.1
EU-GRE05-A3-1992	Europe	Greece	A3	1992	GU097419.1
EU-GRE06-C0-1992	Europe	Greece	C0	1992	GU097417.1
EU-GRE07-A3-1992	Europe	Greece	A3	1992	FJ884610.1
EU-GRE08-A3-1992	Europe	Greece	A3	1992	FJ884608.1
EU-GRE09-C0-1992	Europe	Greece	C0	1992	FJ884609.1
AS-CHN01-C2-2006	Asia	China	C2	2006	FJ853388.1
AS-CHN02-C2-2005	Asia	China	C2	2005	FJ853387.1
AS-CHN03-A1-2004	Asia	China	A1	2004	FJ853386.1
AS-CHN04-C2-2003	Asia	China	C2	2003	FJ853385.1
AS-CHN05-A1-2002	Asia	China	A1	2002	FJ853384.1
AS-CHN06-A1-2001	Asia	China	A1	2001	FJ853383.1
AS-CHN07-C2-2000	Asia	China	C2	2000	FJ853382.1
AS-CHN08-C2-1999	Asia	China	C2	1999	FJ853381.1
AS-CHN09-C2-1998	Asia	China	C2	1998	FJ853380.1
AS-CHN10-C3-1997	Asia	China	C3	1997	FJ853379.1
AS-CHN11-C2-1998	Asia	China	C2	1998	FJ853378.1
AS-CHN12-C2-1999	Asia	China	C2	1999	FJ853377.1
AS-CHN13-C2-2000	Asia	China	C2	2000	FJ853376.1
AS-CHN14-C-2001	Asia	China	C2	2001	FJ853375.1
AS-CHN15-C3-2002	Asia	China	C3	2002	FJ853374.1
AS-CHN16-C2-2003	Asia	China	C2	2003	FJ853373.1
AS-CHN17-C2-2004	Asia	China	C2	2004	FJ853372.1
AS-CHN18-A0-2005	Asia	China	A3	2005	FJ853371.1
AS-CHN19-C2-2006	Asia	China	C2	2006	FJ853370.1
AS-CHN20-C2-2005	Asia	China	C2	2005	FJ853369.1
AS-CHN21-C2-2004	Asia	China	C2	2004	FJ853368.1
AS-CHN22-C2-2003	Asia	China	C2	2003	FJ853367.1

AS-CHN22-C3-2009	Asia	China	C3	2009	KM582707.1
AS-CHN23-A1-2009	Asia	China	A1	2009	KM582706.1
AS-CHN24-A2-2009	Asia	China	A2	2009	KM582705.1
AS-CHN25-A2-2009	Asia	China	A2	2009	KM582704.1
AS-CHN26-C3-2009	Asia	China	C3	2009	KM582703.1
AS-CHN27-A2-2008	Asia	China	A2	2008	KM582702.1
AS-CHN28-A2-2008	Asia	China	A2	2008	KM582701.1
AS-CHN29-A2-2008	Asia	China	A2	2008	KM582700.1
AS-CHN30-A2-2008	Asia	China	A2	2008	KM582699.1
AS-CHN31-A3-2008	Asia	China	A3	2008	KM582698.1
AS-CHN32-A2-2008	Asia	China	A2	2008	KM582696.1
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AS-CHN38-C2-2008	Asia	China	C2	2008	KM582690.1
AS-CHN39-C2-2008	Asia	China	C2	2008	KM582689.1
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AS-CHN41-A4-2007	Asia	China	A4	2007	KM582687.1
AS-CHN42-C3-2007	Asia	China	C3	2007	KM582686.1
AS-CHN43-A3-2007	Asia	China	A3	2007	KM582685.1
AS-CHN44-A2-2007	Asia	China	A2	2007	KM582684.1
AS-CHN45-A1-2007	Asia	China	A1	2007	KM582683.1
AS-CHN46-A3-2007	Asia	China	A3	2007	KM582682.1
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AS-CHN48-A2-2007	Asia	China	A2	2007	KM582680.1
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AS-IRA02-C0-2009	Asia	Iran	C3	2009	HQ395618.1
AS-IRA03-A0-2009	Asia	Iran	A0	2009	HQ395617.1
AS-IRA04-A0-2009	Asia	Iran	A0	2009	HQ395616.1
AS-IRA05-A0-2009	Asia	Iran	A0	2009	HQ395615.1
AS-IRA06-A0-2009	Asia	Iran	A0	2009	HQ395614.1
AS-IRA07-A0-2009	Asia	Iran	A0	2009	HQ395613.1
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AS-IRA09-A0-2009	Asia	Iran	A0	2009	HQ395611.1
AS-IRA20-C0-2001	Asia	Iran	C2	2001	JN242284.1
AS-IRA21-C0-2005	Asia	Iran	C0	2005	JN242283.1
AS-IRA22-C0-2009	Asia	Iran	C2	2009	JN242282.1
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AS-IRA24-C0-2002	Asia	Iran	C0	2002	JN242280.1
AS-IRA25-C0-2000	Asia	Iran	C0	2000	JN242279.1
AS-IRA26-C0-2007	Asia	Iran	C0	2007	JN242278.1
AS-IRA27-C0-2006	Asia	Iran	C1	2006	JN242277.1
AS-IRA28-C0-2007	Asia	Iran	C3	2007	JN242276.1
AS-IRA29-C0-2007	Asia	Iran	C2	2007	JN242275.1
AS-IRA30-A0-2006	Asia	Iran	A0	2006	JN242274.1
AS-IRA31-C0-2008	Asia	Iran	C0	2008	JN242273.1
AS-IRA32-C0-2007	Asia	Iran	C1	2007	JN242272.1
AS-IRA33-C0-2006	Asia	Iran	C2	2006	JN242271.1
AS-IRA34-C0-2007	Asia	Iran	C2	2007	JN242270.1
AS-IRA35-C0-2008	Asia	Iran	C3	2008	JN242269.1
AS-IRA36-C0-2007	Asia	Iran	C0	2007	JN242268.1
AS-IRA37-C0-2008	Asia	Iran	C3	2008	JN242267.1
AS-IRA38-C0-2007	Asia	Iran	C2	2007	JN242266.1
AS-IRA39-C0-2008	Asia	Iran	C3	2008	JN242265.1
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AS-IRA42-C0-2007	Asia	Iran	C0	2007	JN242262.1
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AS-IRA44-C0-2001	Asia	Iran	C2	2001	JN242260.1
AS-IRA45-C0-2006	Asia	Iran	C2	2006	JN242259.1
AS-IRA46-C0-2006	Asia	Iran	C3	2006	JN242258.1
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AS-IRA49-C0-2004	Asia	Iran	C0	2004	JN242255.1
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AS-IRA51-C0-2008	Asia	Iran	C0	2008	JN242253.1
AS-IRA52-C0-2007	Asia	Iran	C3	2007	JN242252.1
AS-IRA52-C0-2009	Asia	Iran	C0	2009	JN242292.1
AS-IRA53-C0-2009	Asia	Iran	C0	2009	JN242291.1
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AS-IRA56-C0-2002	Asia	Iran	C2	2002	JN242288.1
AS-IRA57-C0-2008	Asia	Iran	C3	2008	JN242287.1
AS-IRA58-C0-2009	Asia	Iran	C0	2009	JN242286.1
AS-IRA59-C0-2008	Asia	Iran	C0	2008	JN242285.1
EU-SSI01-N0-1995	Europe	Southern Siberia	A3	1995	GQ861492.1
EU-SSI02-N0-1995	Europe	Southern Siberia	A3	1995	GQ861491.1
EU-SSI03-N0-1995	Europe	Southern Siberia	A3	1995	GQ861490.1
EU-SSI04-N0-1995	Europe	Southern Siberia	A3	1995	GQ861489.1
EU-SSI05-N0-1995	Europe	Southern Siberia	A3	1995	GQ861488.1
EU-SSI06-N0-1995	Europe	Southern Siberia	A3	1995	GQ861487.1
EU-SSI07-N0-1995	Europe	Southern Siberia	C2	1995	GQ861486.1
EU-SSI08-N0-1995	Europe	Southern Siberia	A3	1995	GQ861485.1
EU-SSI09-N0-1995	Europe	Southern Siberia	A3	1995	GQ861484.1
EU-SSI10-N0-1995	Europe	Southern Siberia	A3	1995	GQ861483.1
EU-SSI11-N0-1995	Europe	Southern Siberia	A2	1995	GQ861482.1
EU-SSI12-N0-1995	Europe	Southern Siberia	C2	1995	GQ861481.1
EU-SSI13-N0-1995	Europe	Southern Siberia	A3	1995	GQ861480.1
EU-SSI14-N0-1995	Europe	Southern Siberia	A3	1995	GQ861479.1
EU-SSI15-N0-1995	Europe	Southern Siberia	A3	1995	GQ861478.1

EU-SSI16-N0-1995	Europe	Southern Siberia	A3	1995	GQ861477.1
EU-SSI17-N0-1995	Europe	Southern Siberia	A3	1995	GQ861476.1
EU-SSI18-N0-1995	Europe	Southern Siberia	A3	1995	GQ861475.1

ζ : Sequence subtype was not explicitly determined via phylogenetic analysis.

Appendix Table2: Genetic distance between groups and within groups

	China	Iran	Italy-Greece	Southern Siberia
China	0.094	0.101	0.097	0.094
Iran	-	0.074	0.102	0.125
Italy-Greece	-	-	0.097	0.095
Southern Siberia	-	-	-	0.036

Appendix Table3: Sites found to be under positive selection pressure via three approaches within Chinese sequences.

Codon	SLAC		FEL		REL	
	dN-dS	p value	dN-dS	p value	dN-dS	Bayes Factor
31	3.907	0.066	42.514	0.022	13.172	>10000
33	3.224	0.022	32.170	0.005	13.047	>10000
35	2.738	0.252	28.223	0.089	13.009	>10000
36	3.914	0.120	55.246	0.013	13.164	>10000
39	1.645	0.229	18.231	0.092	9.452	4702.75
40	2.954	0.063	30.498	0.026	13.073	>10000
42	2.769	0.099	32.330	0.013	13.111	>10000
44	2.004	0.181	23.283	0.036	12.100	>10000
74	3.690	0.032	37.842	0.008	13.176	>10000

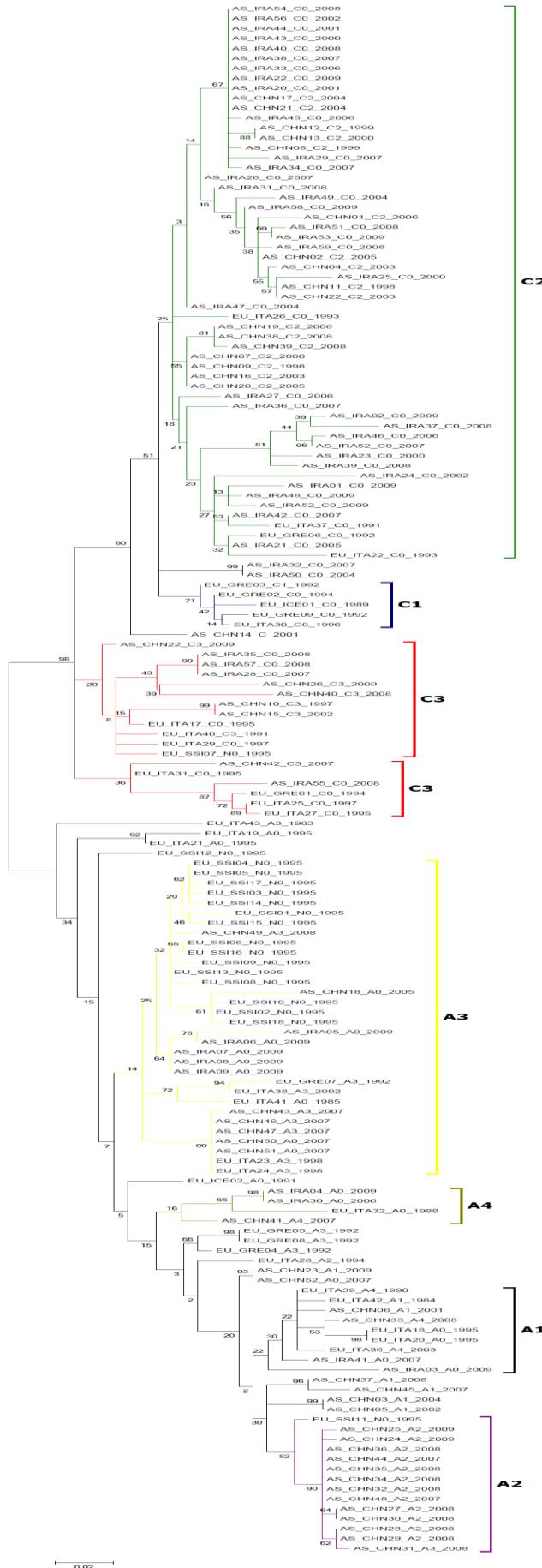
^[1]SLAC P value < 0.5;^[2]FEL P value < 0.25; REL Bayes factor > 50^[3].

Appendix Table4: Sites found to be under positive selection pressure via three approaches within all sequences.

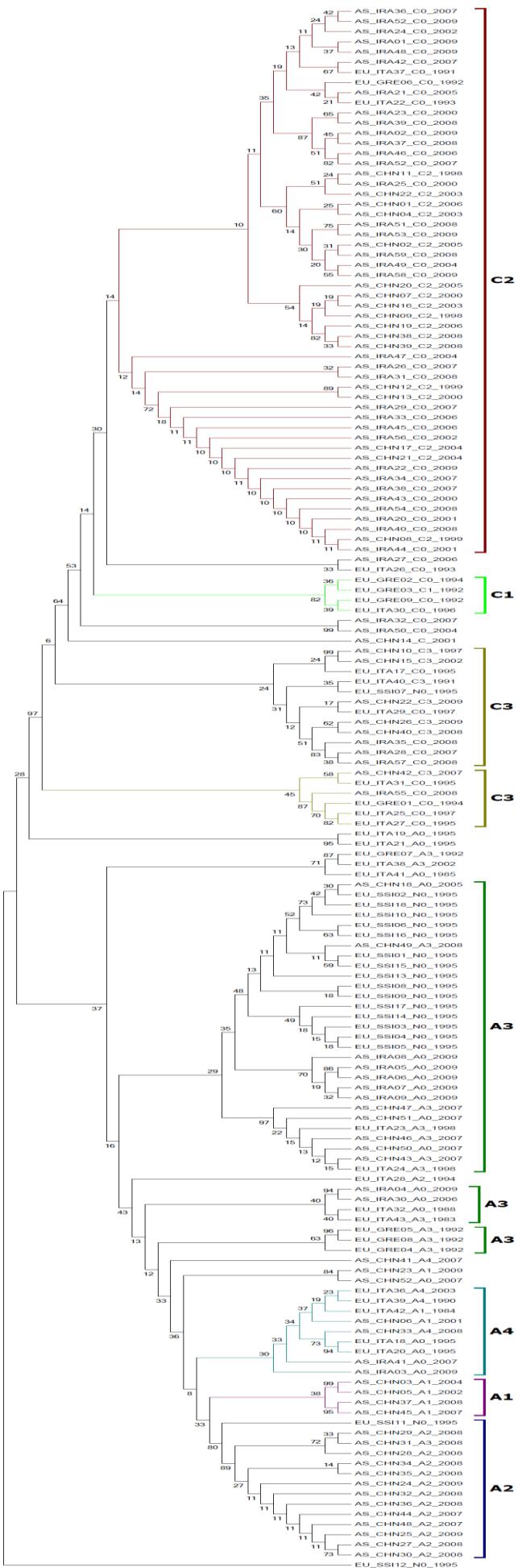
Codon	SLAC		FEL		REL	
	dN-dS	p value	dN-dS	p value	dN-dS	Bayes Factor
20	3.589	0.002	10.562	0.000	3.657	360.439
22	2.486	0.001	7.776	0.000	12.221	2154.34
24	3.084	0.019	9.892	0.005	16.327	475.574
25	4.336	0.000	13.680	0.000	21.175	>10000
26	0.995	0.091	3.096	0.023	5.047	>10000
28	1.443	0.036	4.568	0.012	6.449	>10000
29	2.135	0.015	6.026	0.022	13.432	>10000
30	2.842	0.004	9.568	0.000	12.438	>10000
31	3.100	0.007	11.823	0.000	5.887	2702.75
54	2.640	0.002	7.807	0.000	12.988	>10000
63	2.162	0.014	6.263	0.009	12.133	112.556

^[1]SLAC P value < 0.5;^[2]FEL P value < 0.25; REL Bayes factor > 50^[3].

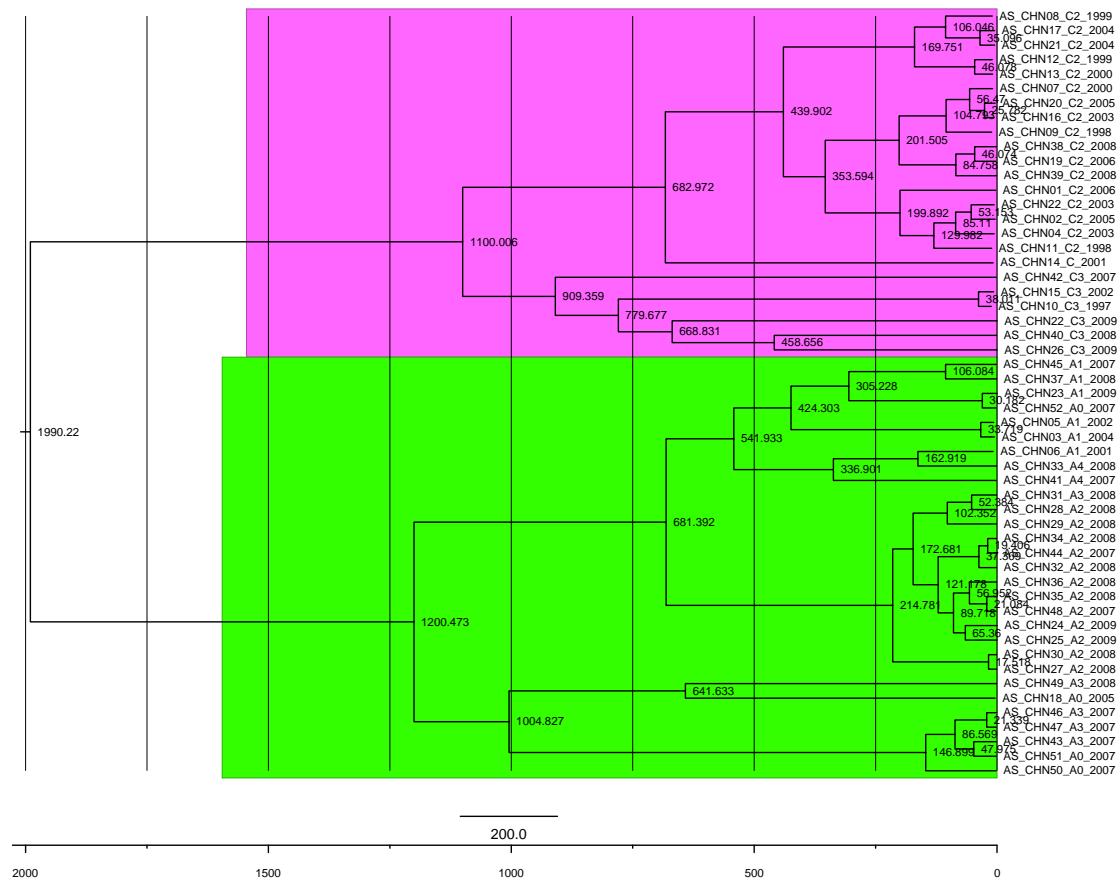
Appendix Figure1: The Maximum Likelihood tree of KSHV sequences with following parameters: 1,000 bootstrap replicates, randomly broken site, and general time-reversible distance measure with a gamma distribution shape parameter of 0.5.



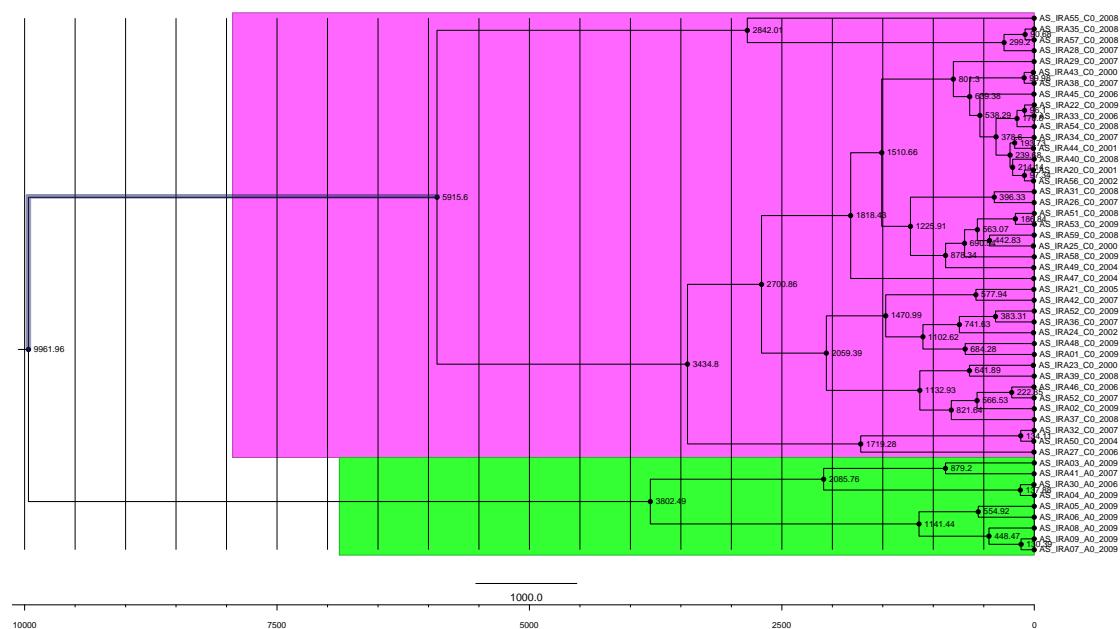
Appendix Figure2: The Maximum Parsimony tree of KSHV sequences with following parameters: 1,000 bootstrap replicates, randomly broken site, and general time-reversible distance measure with a gamma distribution shape parameter of 0.5.



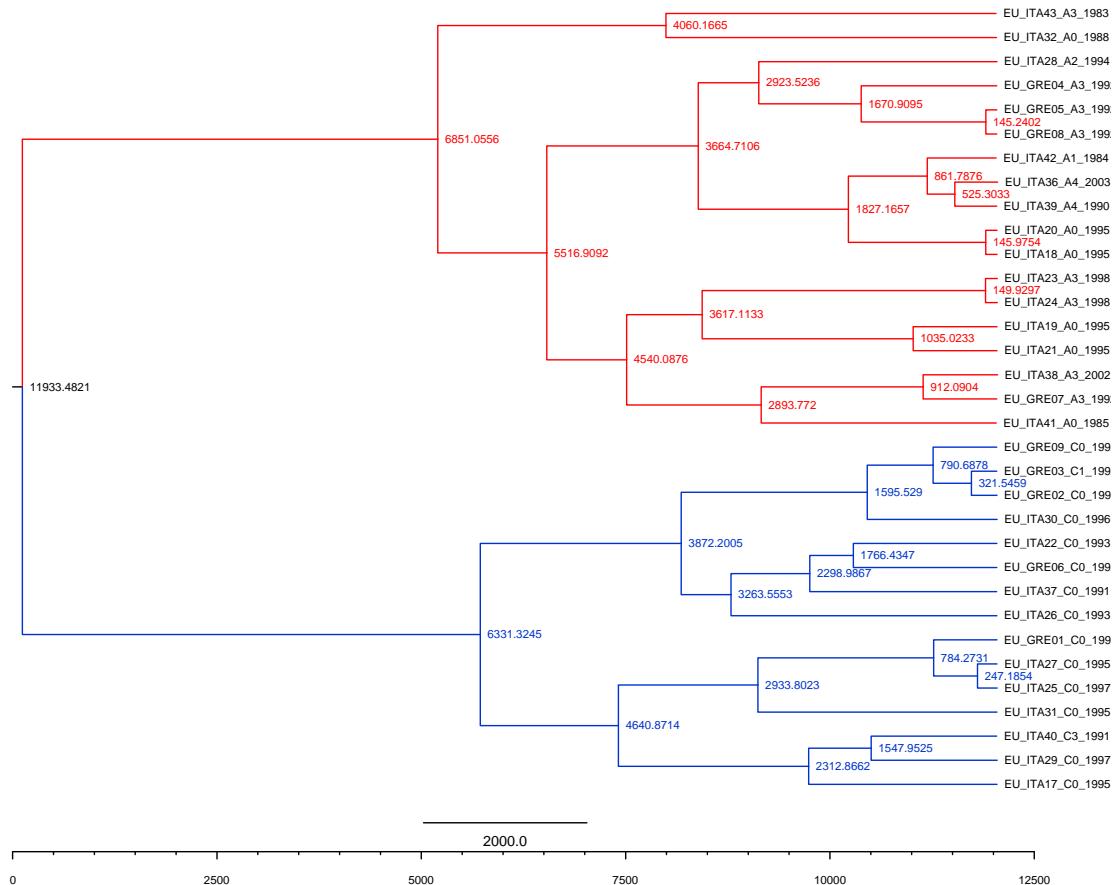
Appendix Figure3: The MCC tree of KSHV sequences from Xinjiang, China.



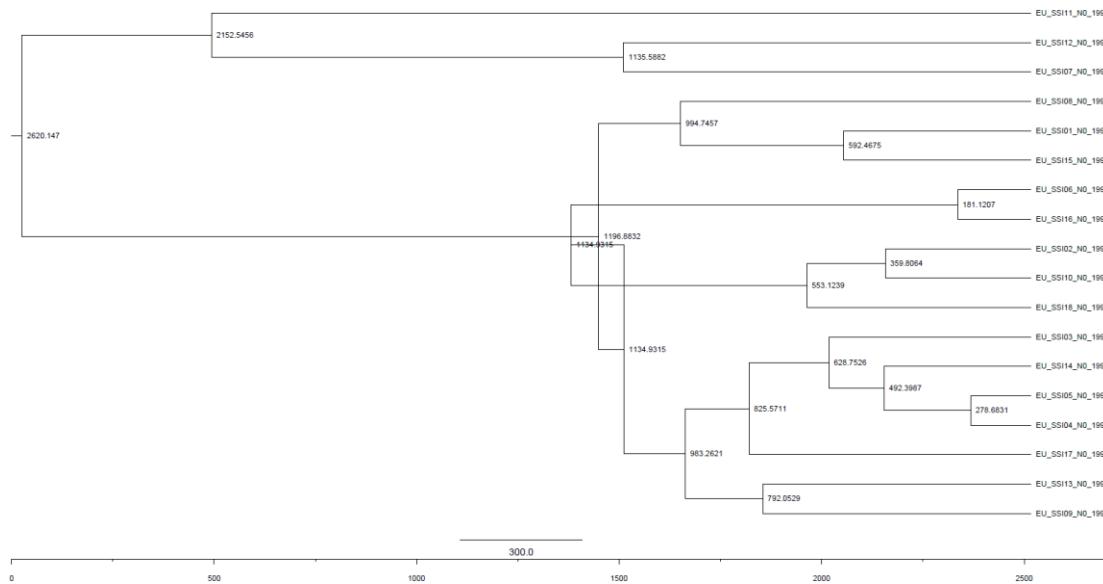
Appendix Figure4: The MCC tree of KSHV sequences from Iran.



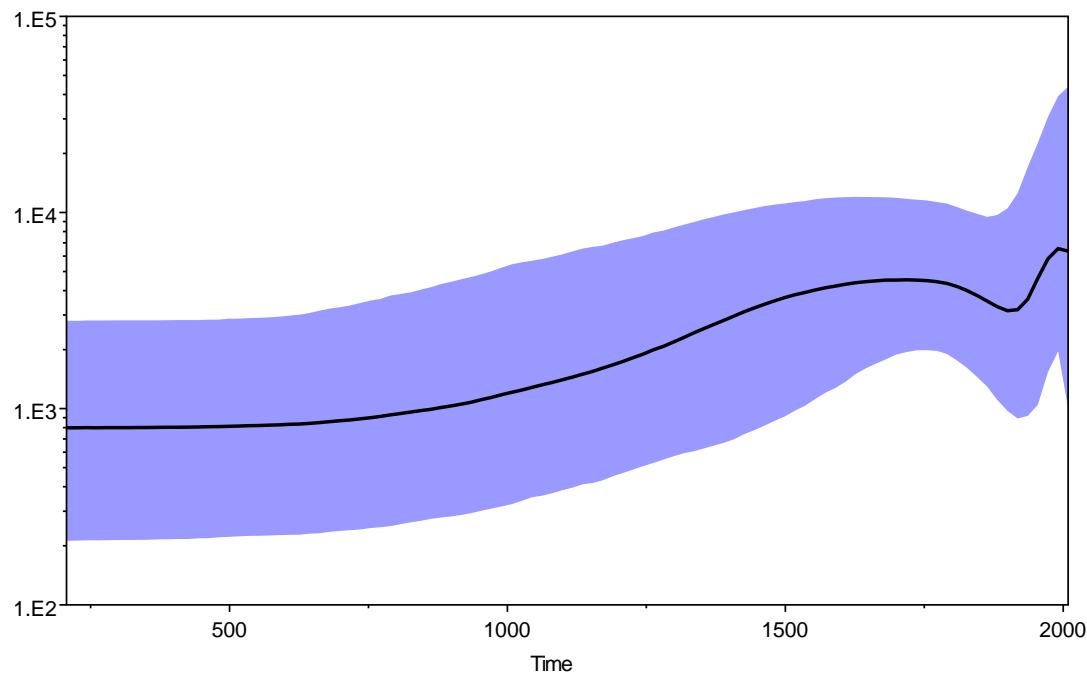
Appendix Figure5: The MCC tree of KSHV sequences from Italy & Greece.



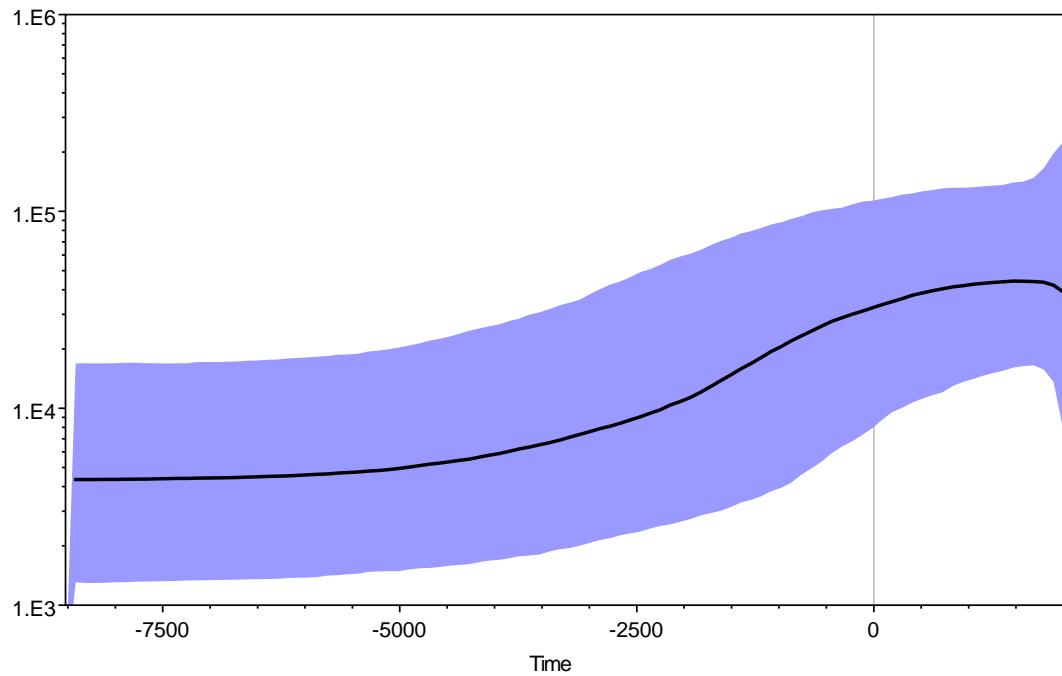
Appendix Figure6: The MCC tree of KSHV sequences from Southern Siberia.



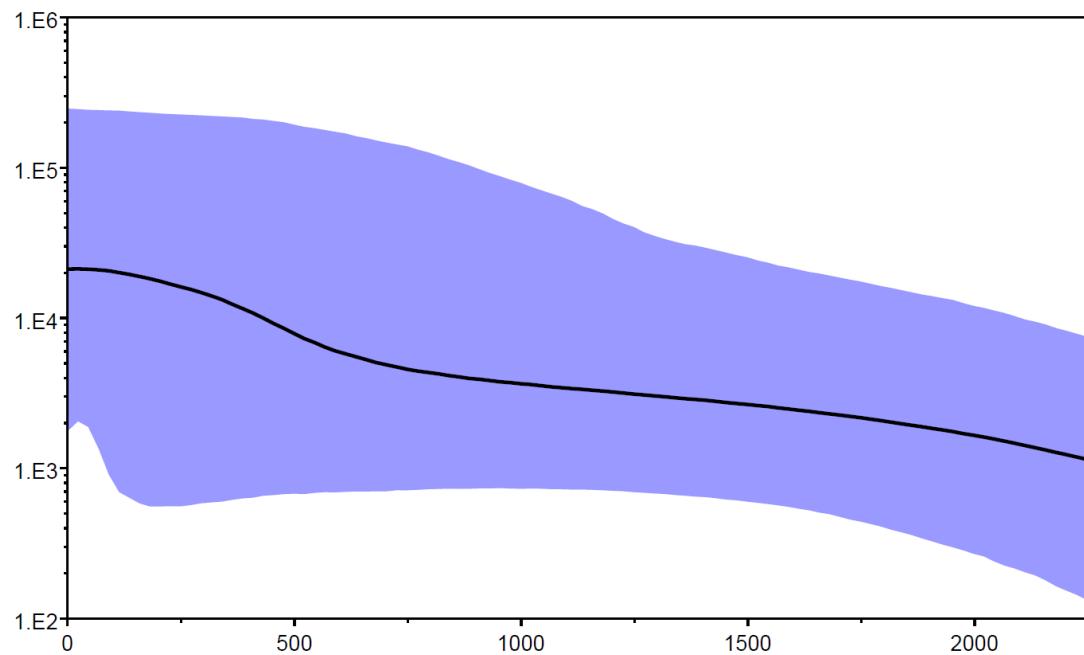
Appendix Figure7: The Bayesian skyline plot of KSHV sequences from China.



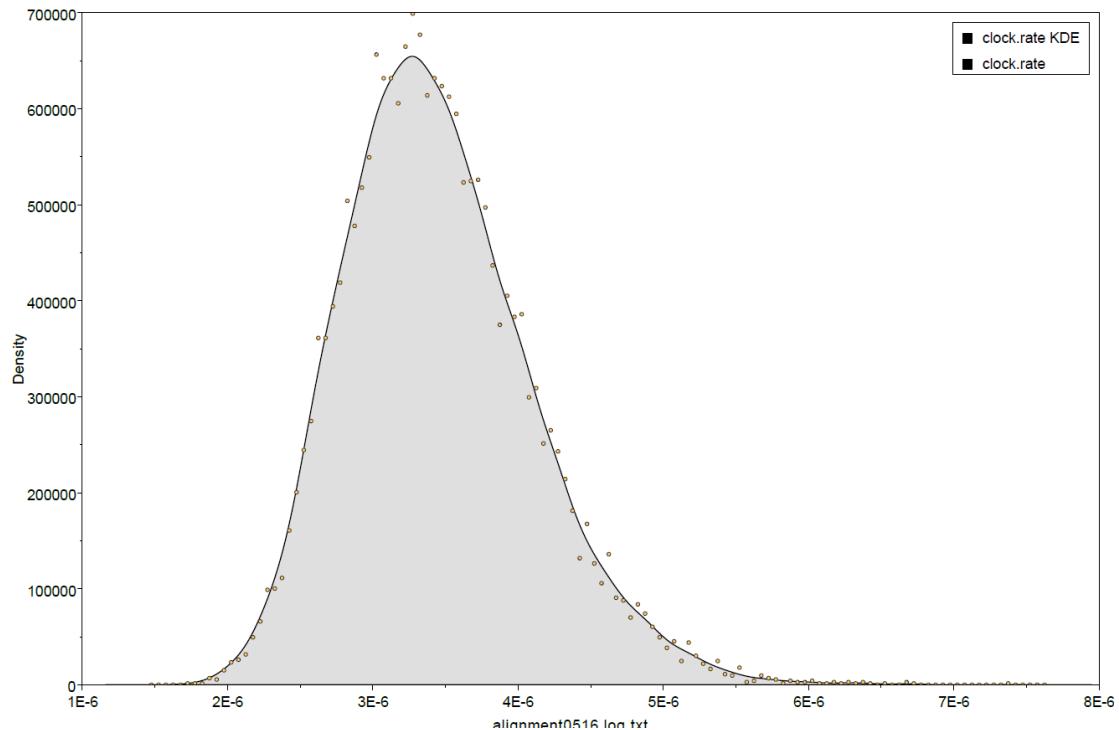
Appendix Figure8: The Bayesian skyline plot of KSHV sequences from Iran, Italy & Greece.



Appendix Figure9: The Bayesian skyline plot of KSHV sequences from Southern Siberia.



Appendix Figure10: The substitution rate of KSHV ORFK1 obtained from Bayesian evolutionary analysis.



Appendix Figure11: The tree height of KSHV ORFK1 obtained from Bayesian evolutionary analysis.

