SUPPLEMENTARY FIGURES AND TABLES



Supplementary Figure S1: Colony numbers of the soft agar colony formation assays in Figure 1H. Cell clusters of three or more cells are counted. Of note, in contrast to miR-K12-1 expressing Rat-1 cells which formed big colonies (Figure 1H), vector-expressing Rat-1 cells only formed small cell clusters of no more than 4 cells.



Supplementary Figure S2: IκBα expression or IL-6 knockdown had no effect on the expression of transfected miR-K12-1 in Hela cells. Real-time RT-PCR analysis was performed to examine the expression level of miR-K12-1 in Hela cells transfected with miR-K12-1 together with an empty vector, IκBα, or IL-6 shRNA.



Supplementary Figure S3: SOCS3 inhibits STAT3 but not NF-κB activation induced by miR-K12-1. Luciferase assay was performed to measure STAT3 or NF-κB transcriptional activity in Hela cells transfected with an empty vector, miR-K12-1, or miR-K12-1 plus SOCS3, together with STAT3 or NF-κB luciferase reporter, respectively.



Supplementary Figure S4: Expression of I κ B α , or knockdown of IL-6 or STAT3, had no effect on the expression of miR-K12-1 in Rat-1 cells. Real-time RT-PCR analysis was performed to examine the expression level of miR-K12-1 in Rat-1 cells stably expressing an empty vector, miR-K12-1, or miR-K12-1 together with I κ B α , IL-6 shRNA, or STAT3 shRNA.



Supplementary Figure S5: Colony numbers of the soft agar colony formation assays in Figure 3C. Cell clusters of three or more cells are counted. Of note, in contrast to Rat-1 cells stably expressing miR-K12-1, which formed big colonies (Figure 3C), Rat-1 cells stably expressing an empty vector, or miR-K12-1 together with IkBa, IL-6 shRNA, or STAT3 shRNA only formed small cell clusters of no more than 4 cells.



Supplementary Figure S6: Knockdown of RelA inhibits nuclear expression of STAT3 in BCBL-1 cells. Immunofluorescence assay was performed to examine the expression of STAT3 and RelA in BCBL-1 cells stably expressing RelA shRNA or scrambled control shRNA, together with GFP. Scale bar: 5 µm.



Supplementary Figure S7: Colony numbers of the soft agar colony formation assays in Figure 4E.



Supplementary Figure S8: Knockdown of RelA, II-6 or STAT3 blocks BCBL-1 cell growth in cell culture.



Supplementary Figure S9: Modeling of miR-K12-1/IκBα/NF-κB/IL-6/STAT3 signaling in KSHV tumorigenesis. NF-κB and STAT3 function as dimers but are indicated only by single protein.

Gene	Accession number	Target sequence
Human RelA	NM_021975.3	GGACATATGAGACCTTCAAGA
Human IL-6	NM_000600.3	AGACATGTAACAAGAGTAA
Human STAT3	NM_139276.2	GCACAATCTACGAAGAATCAA
Rat IL-6	NM_012589.2	GCTTCCAAACTGGATATAACC
Rat STAT3	NM_012747.2	GCTGCACCTGATCACCTTTGA

Supplementary Table S1: target sites of shRNAs

Gene	Accession	Primer pairs		
	number	Forward (5' to 3')	Reverse (5' to 3')	
GAPDH	NM_002046.3	CCGAGCCACATCGCTCAGACAC	GTGACCAGGCGCCCAATACGAC	
ΙκΒα	NM_020529.2	GGAGTACGAGCAGATGGTCAA	CCTTCACCTGGCGGATCAC	
IL-6	NM_000600.3	TAGTGAGGAACAAGCCAGAGC	TGGGTCAGGGGTGGTTATTG	
Survivin	NM_001168.2	TGACGACCCCATAGAGGAACA	CGCACTTTCTCCGCAGTTTC	
Bcl-2	NM_000633.2	ATGTGTGTGGAGAGCGTCAACC	TGAGCAGAGTCTTCAGAGACAGCC	
Bcl-xL	NM_138578.1	GAATGACCACCTAGAGCCTTGG	TGTTCCCATAGAGTTCCACAAAAG	
Cyclin D1	NM_053056.2	CCGTCCATGCGGAAGATC	ATGGCCAGCGGGAAGAC	
STAT3	NM_139276.2	CTAGATCGGCTAGAAAACTGGATAACGTC	ATGGGGTCCCCTTTGTAGGAAACT	
RelA	NM_021975.3	CCAGACCAACAACAACCCCT	GATCTTGAGCTCGGCAGTGT	
U6 snRNA	X59362.1	CTCGCTTCGGCAGCACA	AACGCTTCACGAATTTGCGT	
miR-K12-1 reverse- transcription		GTCGTATCCAGTGCAGGGTCCGAGGTA	TTCGCACTGGATACGACGCTTAC	
miR-K12-1		CGCGCATTACAGGAAACTGGG	GTGCAGGGTCCGAGGT	

Supplementary Table S2: primer pairs used for real-time PCR