

| Supplemental Table 1 | | |
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| Constructs | Source | Catalogue number |
| pCK-Drosha-Flag | addgene | Plasmid #164690 |
| pFLAG/HA-DGCR8 | addgene | Plasmid #10921 |
| VP5-FA hDGCR8 (NM_022720) | Gunter Meister | N/A |
| pLVTHM | addgene | Plasmid #12247 |

| Supplemental Table 2 | | |
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| Primers/Oligo | Source | Catalogue number |
| Synthetic RNA Oligo | | |
| hsa-miR-155-5p miRCURY LNA miRNA Mimic UUA AUGCUAAUCGUGAUAGGGGU | Exiqon/Qiagen | Cat. YM00472490-ADB |
| Negative control RNA | Exiqon/Qiagen | Cat. YM00479903-ADA |
| hsa-miR-155-5p miRCURY LNA miRNA Detection probe UUA AUGCUAAUCGUGAUAGGGGU | Exiqon/Qiagen | Cat. YD00619862-BCA |
| hsa-miR-148b-3p miRCURY LNA miRNA Detection probe UCAGUGCAUCACAGAACUUUGU | Exiqon/Qiagen | Cat. YD00613165-BCA |
| hsa-miR-let-7f1 LNA detection probe AACTATACAATCTACTACCTCA | Exiqon/Qiagen | Cat. YD00611752-BCA |
| hsa-miR-let7d LNA detection probe AACTATGCAACCTACTACCTCT | Exiqon/Qiagen | Cat. YD00616777-BCA |
| hsa-miR-30c-1 LNA detection probe GCTGAGAGTGTAGGATGTTTACA | Exiqon/Qiagen | Cat. YD00610995-BEA |
| hsa-miR-30a LNA detection probe CTTCCAGTCGAGGATGTTTACA | Exiqon/Qiagen | Cat. YD00611442 |
| hsa-miR-30d LNA detection probe CTTCCAGTCGGGGATGTTTACA | Exiqon/Qiagen | Cat. YD00613716 |
| Human U6 detection probe CACGAATTTGCGTGTCATCCTT | Sigma | N/A |
| miR-aU14 LNA detection probe GCGACGTCTCGACAGAGCTCTTA | Exiqon/Qiagen | Cat. YCD0075760-BEA |
| miR-aU14 mimic UAAGAGCUCUGUCGAGACGUCGC | IDT DNA Technologies | N/A |

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| miR-aU14 mutant 1 UAAGUGGAUUGUCGAGACGUCGC | IDT DNA Technologies | N/A |
| miR-aU14 mutant 2 UAAUCGAUUGUCGAGACGUCGC | IDT DNA Technologies | N/A |
| miR-155 mimic UUA AUGCUAAUCGUGAUAGGGGUU | IDT DNA Technologies | N/A |
| Anti-Let-7d.1 UACA UCCCUGCCCGUGCUAUGC | IDT DNA Technologies | N/A |
| Anti-Let-7d.2 UACUCCUUGUGGGCUGCUAUGC | IDT DNA Technologies | N/A |
| Anti-Let-7f.1 UACAAUCAGUGGGCUGCUAUGC | IDT DNA Technologies | N/A |
| Anti-Let-7f.2 UACCUCCUGAACUGCUAUGC | IDT DNA Technologies | N/A |
| control RNA for interaction studies UACGUACCACAGUGCUAUGC | IDT DNA Technologies | N/A |
| hsa-miR-30c-5p miRCURY LNA miRNA Mimic UGUAAACA UCCUACACUCUCAGC | Exiqon/Qiagen | YM00471185 |
| hsa-miR-30c-5p miRCURY LNA miRNA Power Inhibitor UGUAAACA UCCUACACUCUCAGC | Exiqon/Qiagen | YI04105292 |
| miRCURY LNA miRNA inhibitor control TAACACGTCTATACGCCCA | Exiqon/Qiagen | YI00199006 |
| Biotinylated pre-miR-30c-1 /5Biosg/ACCAUGCUGUAGUGUGUGUAA ACA UCCUACACUCUCAGCUGUGAGCUCA AGGUGGCUGGGAGAGGGUUGUUUACUCC UUCUGCCAUGGA | IDT DNA Technologies | N/A |
| DNA Oligo / Primers | | |
| miR-aU14 lentivirus (Forward) | Sigma | N/A |

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| (Phos)CGCGTCCCCTAAGAGCTCTGTCTGA GACGTGCTTCAAGAGAGCGACGTCTCG ACAGAGCTCTTATTTTTGGAAAT | | |
| miR-aU14 lentivirus (Reverse) (Phos)CGATTTCCAAAAATAAGAGCTCTG TCGAGACGTGCTCTCTTGAAGCGACGTC TCGACAGAGCTCTTAGGGGA | Sigma | N/A |
| miR-aU14 mutant lentivirus (Forward) (Phos)CGCGTCCCCTAATATCGCTGTCTGA GACGTGCTTCAAGAGAGCGACGTCTCG ACAGCGATATTATTTTTGGAAAT | Sigma | N/A |
| miR-aU14 mutant lentivirus (Reverse) (Phos)CGATTTCCAAAAATAATATCGCTG TCGAGACGTGCTCTCTTGAAGCGACGTC TCGACAGCGATATTAGGGGA | Sigma | N/A |
| miR-A lentivirus (Forward) (Phos)CGCGTCCCCACTATACGTCCAACA CTGACAGCTGTGAGCTCAAGGTGGGCTG TCAGTGTTGGACGTATAGTTTTTTGGAA AT | Sigma | N/A |
| miR-A lentivirus (Reverse) (Phos)CGATTTCCAAAAAATACTATACGTCC AACTGACAGCCACCTTGAGCTCACAG CTGTCAGTGTTGGACGTATAGTGGGGA | Sigma | N/A |
| Probe for miR-A detection GCTGTCAGTGTTGGACGTATAGT | This paper | N/A |
| miR-B lentivirus (Forward) (Phos)CGCGTCCCCACCATGCTGTAGTGT GTCTATAGTTCCTACACACTCAGCTGTGA GCTCAAGGTGGCTGGGTGAGGGTACTAT AGTCCTTCTGCCATGGTTTTTTGGAAAT | Sigma | N/A |
| miR-B lentivirus (Reverse) | Sigma | N/A |

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| (Phos)CGATTTCCAAAAACCATGGCAGAA GGACTATAGTACCCTCACCCAGCCACCTT GAGCTCACAGCTGAGTGTGTAGGAACTA TAGACACACTACAGCATGGTGGGGA | | |
| miR-30c-1 sponge (Forward, Oligo 1) (Phos)CGCGTCCCCCTGAGGATGTAGGAT GTTTACATGACTGAGGATGTAGGATGTT TACAAGCGGCTGAGGATGTAGGATGTTT ACA CCCGGTGCATGACTAAGCTAGCCTGAGG AT | Sigma | N/A |
| miR-30c-1 sponge (Forward, Oligo 2) GTAGGATGTTTACACGACTGAGGATGTA GGATGTTTACAACGCGCTGAGGATGTAG GATGTTTACATTTTTTGAAAT | Sigma | N/A |
| miR-30c-1 sponge (Reverse, Oligo 1) GCTAGCTTAGTCATGCACCGGGTGTA CATCCTACATCCTCAGCGCCTTGTAACA TCCTACATCCTCAGTCATGTAAACATCCT ACATCCTCAGGGGGA | Sigma | N/A |
| miR-30c-1 sponge (Reverse, Oligo 2) CGATTTCCAAAAATGTAAACATCCTACA TCCTCAGCCGCTTGTAACATCCTACATC CTCAGTCGTGTAAACATCCTACATCCTCA G | Sigma | N/A |
| Probe for miR-B detection GCTGAGTGTGTAGGAAGTATAGA | This paper | N/A |
| TaqMan Actin (Internal control) primers Hs.PT.39a.22214847 | IDT DNA Technology | Cat. 226901553 |
| TaqMan IFNbeta primers Hs.PT.58.39481063.g | IDT DNA Technology | Cat. 226901557 |
| HHV-6A U14 mutant BAC Forward | This paper | N/A |

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| TACAATGATGAAAAGTCTTTTCGACACC GAAGAAGCCATGAGGAGATTAGATCGTG CATTGATGGTAGGGATAACAGGGTAATC GATTT | | |
| HHV-6A U14 mutant BAC Reverse GGCATATATGAACTACCACGAGCATCGA CAAACAGCCCCATCAATGCACGATCTAA TCTCCTCATGGCCAGTGTTACAACCAATT AACC | This paper | N/A |
| HHV-6 P41 PCR forward CCTGTTTTGATGCCAACGCA | Sigma | N/A |
| HHV-6 P41 PCR reverse AAAGCACGTTGTTGACGGTG | Sigma | N/A |
| HHV-6 U90 PCR forward CCAGCTAAAGTTTTCAACTCTCCA | Sigma | N/A |
| HHV-6 U90 PCR reverse GTGATACATCCTTCGATGACCTGA | Sigma | N/A |
| IFNbeta PCR forward GAACTTTGACATCCCTGAGGAGATTAAG CAGC | Sigma | N/A |
| IFNbeta P41 PCR reverse GTTCCTTAGGATTTCCACTCTGACTATGG TCC | Sigma | N/A |
| IFIT1 PCR forward ACAGCAACCATGAGTACAAATGG | Sigma | N/A |
| IFIT1 P41 PCR reverse CATCGTCATCAATGGATAACTCCC | Sigma | N/A |
| 5S PCR forward GTCTACGGCCATACCACCC | Sigma | N/A |
| 5S PCR reverse AAAGCCTACAGCACCCGGT | Sigma | N/A |

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| Pri-miR30c_Forward_KpnI GAATTTCGGTACCTGTTGTCCCAGGCAGG GGCACC | Sigma | N/A |
| Pri-miR30c_Reverse_XhoI GAATTCCTCGAGAGGGGGCCTGATGGAC AGCTCTCCGC | Sigma | N/A |
| Pre-miR30c_Forward T7 IVT GCTTATCGAAATTAATACGACTCACTAT AGGGACCATGCTGTAGTGTGTGTAAAC | Sigma | N/A |
| Pri-miR30c_Reverse T7 IVT TCCATGGCAGAAGGAGTAAACAACCC | Sigma | N/A |
| Pri-miR30a_Forward_KpnI GAATTCGGTACCGTACTTTACAGAATCG TTGCC | Sigma | N/A |
| Pri-miR30a_Reverse_XhoI GAATTCCTCGAGCCATTTTAATTCAGCTT TG | Sigma | N/A |
| Pre-miR30a_Forward T7 IVT GCTTATCGAAATTAATACGACTCACTAT AGG GCGACTGTAAACATCCTCGACTG | Sigma | N/A |
| Pri-miR30a_Reverse T7 IVT GCAGCTGCAAACATCCGACTG | Sigma | N/A |
| Pri-miR30d_Forward_KpnI GAATTCGGTACCTCACAGCTATTGTTCA GCAC | Sigma | N/A |
| Pri-miR30d_Reverse_XhoI GAATTCCTCGAGTAAAAGAATGCAGCTA AAGATTTCTG | Sigma | N/A |
| Pre-miR30d_Forward T7 IVT GCTTATCGAAATTAATACGACTCACTAT AGG GTTGTGTGTAAACATCCCCGACTG | Sigma | N/A |
| Pri-miR30d_Reverse T7 IVT | Sigma | N/A |

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| GTAGCAGCAAACATCTGACTG | | |
| Pri-miR148b_Forward_KpnI GAATTCGGTACCGAGGACAGCATTCTTG TTTTCTTG | Sigma | N/A |
| Pri-miR148b_Reverse_XhoI GAATTCCTCGAGCCTTGTAACACATTAA ATGCTGAC | Sigma | N/A |
| Pre-miR148b_Forward T7 IVT GCTTATCGAAATTAATACGACTCACTAT AGGGCAAGCACGATTAGCATTTGAGG | Sigma | N/A |
| Pri-miR148b_Reverse T7 IVT TAGAAAGCTTTCGAGACAAAGTTC | Sigma | N/A |
| TaqMan Non-coding RNA Assay Assay ID: Hs04231430_sH (miR30a) | Thermo Scientific | 4426961 |
| TaqMan Non-coding RNA Assay Assay ID: Hs04231453_s1 (miR30d) | Thermo Scientific | 4426961 |
| TaqMan Gene Expression Assay Assay ID: Hs06628134_s1 (miR30c1) | Thermo Scientific | 4351372 |
| TaqMan Advanced miRNA Assay Assay ID: 479448_mir (hsa-miR-30a-5p) | Thermo Scientific | A25576 |
| TaqMan Advanced miRNA Assay Assay ID: 478008_mir (hsa-miR-30c-5p) | Thermo Scientific | A25576 |
| TaqMan Advanced miRNA Assay Assay ID: 478606_mir (hsa-miR-30d-5p) | Thermo Scientific | A25576 |
| TaqMan miRNA Assay Assay ID: 001973 (U6 snRNA) | Thermo Scientific | 4427975 |
| TaqMan Gene Expression Assay Assay ID: Hs00360261_m1 (NFYC) | Thermo Scientific | 4448892 |
| TaqMan Gene Expression Assay (FAM) Assay ID: Hs02621180_s1 (IFNbeta) | Thermo Scientific | 4331182 |

| Supplemental Table 3 | | | |
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| Antibodies | Source | Catalogue Number | Dilutions used |
| Drp1 (clone C-5) | Santa Cruz | Cat. SC-271583 | 1:100 ^a 1:1000 ^b |
| Drp1 (clone 6Z-82) | Santa Cruz | Cat. SC-101270 | 1:100 ^a 1:1000 ^b |
| Drp1 | GeneTex | Cat. GTX31901 | 1:100 |
| p53 (clone DO-1) | Santa Cruz | Cat. SC-126 | 1:100 |
| Beta Actin (clone 4C2) | Sigma | Cat. MABT825 | 1:100 |
| GAPDH | Santa Cruz | Cat. SC-47724 | 1:100 |
| HHV-6 P41 (clone 9A5D12) | Santa Cruz | Cat. SC-65447 | 1:100 ^a 1:1000 ^b |
| HHV-6 gp82/105 Clone 2-D6 late antigen | Balachandran et al., 1989 Available at HHV-6 Foundation, USA repository. (https://hhv-6foundation.org/repository) | N/A | 1:100 |
| Anti-Rabbit Cy-5 secondary antibody | Dianova | Cat. 111-175-144 | 1:100 ^b |
| Anti-Mouse Cy-5 secondary antibody | Dianova | Cat. 115-175-146 | 1:100 ^b |
| Donkey anti-mouse ATTO 643, custom labeled with ATTO 643-NHS (ATTO-TEC GmbH, # AD 643) | Jackson ImmunoResearch | Cat. 715-005-150 | 1:500 ^b |

^a dilution for immunofluorescence

^b dilution for immunoblotting