



US 20120251502A1

(19) **United States**(12) **Patent Application Publication**
Towner et al.(10) **Pub. No.: US 2012/0251502 A1**(43) **Pub. Date: Oct. 4, 2012**(54) **HUMAN EBOLA VIRUS SPECIES AND
COMPOSITIONS AND METHODS THEREOF****Publication Classification**(75) Inventors: **Jonathan S. Towner**, Atlanta, GA
(US); **Stuart T. Nichol**, Atlanta, GA
(US); **James A. Comer**, Atlanta,
GA (US); **Thomas G. Ksiazek**,
Atlanta, GA (US); **Pierre E. Rollin**,
Atlanta, GA (US)(73) Assignee: **The Government of the US as
Represented by the Secretary of
the Dept. of health**, Atlanta, GA
(US)(21) Appl. No.: **13/125,890**(22) PCT Filed: **Oct. 26, 2009**(86) PCT No.: **PCT/US09/62079**§ 371 (c)(1),
(2), (4) Date:**Jun. 21, 2011****Related U.S. Application Data**(62) Division of application No. 61/108,175, filed on Oct.
24, 2008.(51) **Int. Cl.****A61K 35/76** (2006.01)
C07H 21/04 (2006.01)
C12N 7/04 (2006.01)
C07K 14/08 (2006.01)
A61K 38/02 (2006.01)
A61K 31/7088 (2006.01)
C07K 7/06 (2006.01)
C07K 7/08 (2006.01)
C12N 7/00 (2006.01)
C07H 21/02 (2006.01)(52) **U.S. Cl. 424/93.6; 435/235.1; 536/23.72;
435/236; 530/350; 514/1.1; 514/44 R; 530/330;
530/329; 530/328; 530/327; 530/326; 530/325;
530/324**

(57)

ABSTRACT

Compositions and methods including and related to the Ebola Bundibugyo virus (EboBun) are provided. Compositions are provided that are operable as immunogens to elicit and immune response or protection from EboBun challenge in a subject such as a primate. Inventive methods are directed to detection and treatment of EboBun infection.

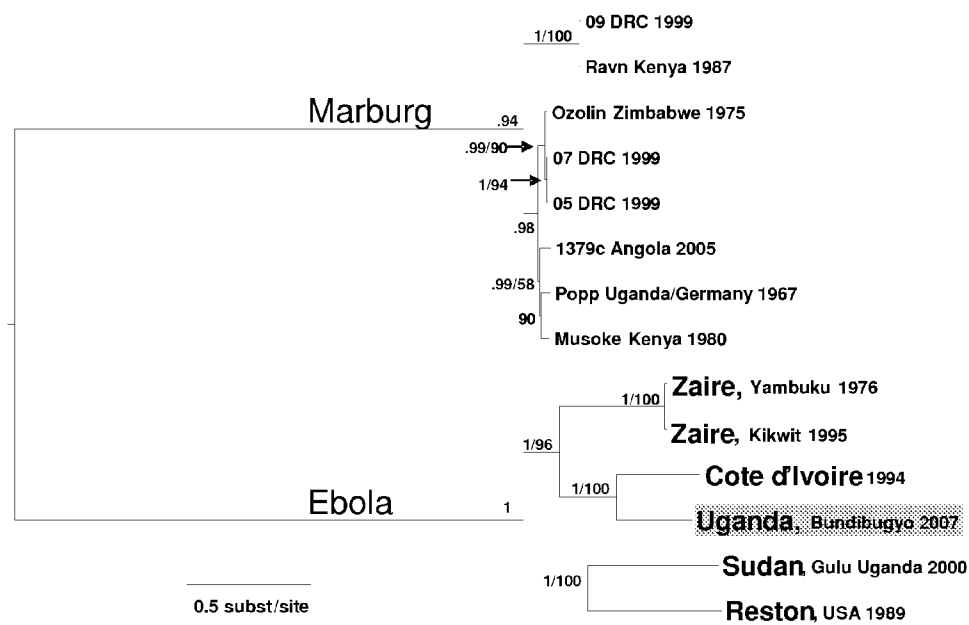


Fig. 1

FIG. 2

[illegible]

FIG. 2

[illegible]

FIG. 2

Ebola Bundibugyo '07	2210	2220	2230	2240	2250	2260	2270	2280	2290	2300
Ebola IC '94	CGACCAACACGATGGTGACAA	TGAAGCAATTCCTCCCTGG	AAATCAGACGACGAGG	GTAGCACTGATCTACT	TGACGAGAAACAAAGCCTGCCACTGC					
Ebola Zaire '76	TGTCACAATGAAGATGACATTC	CCCTCTTGGAAACAGCAAG	AAACACACTGAGACCAAT	TACCAACCAAAAATACCACTGC						
	GGACGATCCGACGAGACGCT	TAGCCCTTCGGCTTGGAT	CAGATGATGAGAGCAGG	ACGGAACCTCCCAACCGCACA	CCCACTGCTGC					
Ebola Bundibugyo '07	2310	2320	2330	2340	2350	2360	2370	2380	2390	2400
Ebola IC '94	ACCTCCCGCTCCCGTCTAC	CGAAGTACTCGTAGAT	TCTCGTAGATCTGCTCCCT	CAGACCAATCCCGACAGT	CCCAATCAACGACCAAT	TGAGCAACAATGC				
Ebola Zaire '76	TCCACGACACTGTTATCGG	AGTAAATTCAGAAAAGG	AGCCCTCCCGCAAGAAAAT	CCCGAGACCAACCAACCA	AGTGAAGTGAAGTGAAGT	TACCAATACC				
	CCCAACCGCTCCCGTATCAG	AGATCCTCTGAAAAAG	AAAGAACTCCCGCAAG	CAGCAGCAACAAGAT	CAGGACCACTCAAGAGG	CCGAGCAACCAAGAC				
Ebola Bundibugyo '07	2410	2420	2430	2440	2450	2460	2470	2480	2490	2500
Ebola IC '94	AGGAACAATGCTAGTCG	GAGCAATCCATTCGAGAA	ATGTAFCACATATCTT	GAAAAACAGAGACGCT	TTTGTGATGCCATCTT	TGATCCCATATGATGA				
Ebola Zaire '76	GACAAATAACCTCACT	CAGACCAATCAGTGG	AGAAATGTATCGACACAT	CTCCAAACACAGAGCA	CCATTTGATGCCATCTT	TATATACATGATGA				
	AGTGACAACACCCAGT	CAGAACACTCTTTT	TGAGAGATGTATCG	CCACATTTCAAGATCA	CAGGGGCCATTTGAT	GCCTGTTTGTATATATCATATGATGA				
Ebola Bundibugyo '07	2510	2520	2530	2540	2550	2560	2570	2580	2590	2600
Ebola IC '94	AAGAAGCCCATCATTT	CAGCACTAGTGTATG	ATGGAAGAGATATACAT	ATCCAGACTCTCTT	GAAATGAGTATCCACCTGGCT	CAGCGAAGGAAGC				
Ebola Zaire '76	CGGAGGAGCCGATCTCT	TTAGCACTAGTGTATG	GAAGATACCTGATCTCT	TTGAGGGAGCATCCAC	CGTGGCTCAGTGA	AAAAAGAGGC				
	AGGATGAGCCCTGTAGT	TTTTCAGTACAGTGT	ACAGTATCCAGCTCC	CTTGAAGAGAAATATCC	CACCATGGCTCACT	GTAAAAAGAGGC				
Ebola Bundibugyo '07	2610	2620	2630	2640	2650	2660	2670	2680	2690	2700
Ebola IC '94	CATGACGAACACATAG	ATTATACCAATGGAT	GGTGCAGAGTTTACT	GGCCTGTGATGAATCAT	ATAGAATAAATTCAT	GGCAATCTCCAGCATCAC				
Ebola Zaire '76	CTTGAATGAGACATAG	GTGTTATCAATGGAT	GTATCAACAAATTTACT	GGCTGTATGAATCAG	AGCAAAATTCATGGCTAT	CTTTCAGACCCAC				
	TATGAATGAAGAGATAG	ATTGTTTACATTTGG	ATGGTTCACAAATTTAT	TGGCGGGTGAATGA	TATCAACAGATAAATTCAT	GGCAATCTCCCAACATCAT				
Ebola Bundibugyo '07	2710	2720	2730	2740	2750	2760	2770	2780	2790	2800
Ebola IC '94	AGGTGATCCGACCTCT	AAACTGAGCT---CC	TATCAAGCTAC	CCATCACTCTGCG	GAATGCCAGAACCTCC	TCCCAACAGTCCCATCGA				
Ebola Zaire '76	AAGTAAATTTCTTCA	TATGACAGATCA---TT	GTGAAGTTATATACC	ACCATCCCTGCA	CAAGACATGA	AAACCACTCAACAGCCCTACACAGG				
	CAGTGAATGACATGGA	CAATGGGATGATTC	AAACCGCAAAATAGCT	ATAATGATGA	GTAGTCAAGGAACG	AAAAAC-----AGGAAGNAATTTT	TGATGTC			
Ebola Bundibugyo '07	2810	2820	2830	2840	2850	2860	2870	2880	2890	2900
Ebola IC '94	GAACCTCCGACCGGT	CACAGCGCAAGACG	CAACCTTAATGAT	GTCTCTGTTCA	CCCAACCGCAACACACT	TTGATCGACTTCCAGAC---AACTGA				
Ebola Zaire '76	ATACCTTGGAGACCA	TACACAGATCAGCAGT	GTGCAACACCCCAT	TCGGAATCCACCA	CCCAACCAACCAACAT---AATCCCAAGCAACCAACCT					
	TAAGGTGTGAATTA	TATACAAATAAAGTGA	TTT---CTTATTT	TGAATTAAGAGT	AGCTATTATTA	TACTAGCCGTTTTC	CAAGTTTCAAGTTTCAATTTGAGTCT			

Ebola Bundibugyo '07	TTATATGTTCTCAAAAATACAAAGTATGAAGATAAGAAAAGCATCTCTTTATCTTTGAGAGGAGCTAAATCTTTTATACTTCACTAATCTTTAAGTAAGTTT
Ebola IC '94	CAAAAGGACTTACAAAACCAAGGGTGATGAAGATTAAAGAAAAGCCCTCTTCAGTTGCAAGGAGCTAAATCTTAAAACTTCACTAGACATAAGGATAAATC
Ebola Zaire '76	TTAAGTAAACC--AAACCAAAAGTATGAAGATAAGAAAACCTTACCTCGGCTGAGAGAGTGTTTTTCATTAACCTTCATCTTGTAAACGTTGAGCCAA
Ebola Bundibugyo '07	GATCACTACCACCATCAGGAGGGCAATTCTACTACTGTGCACCGCCAGAAATACATAGAGGCTGTCTACCCAAATGAGAACGGTTAGTACTAGTATCAACAGT
Ebola IC '94	GATTCCAAATCAGATCAGGAGAAATCATCCTACCCACGACACCTGAATACATGAGGAGCTGTTTACCCAAATGAGAACCAATGAATTTCTGGTGCAGACAAC
Ebola Zaire '76	AATTGTTAAAATAATCAGGGGGTTATATTGCTACTGTCTCCTCCTGAATATATGAGAGGCCATATACCCCTGTGCAGGTCAAAATTCAACAATTTGCTAGAGGT
Ebola Bundibugyo '07	ACTGCCAGTGTGCCGAACTTTCACAGACCGGATGTAAATGATCAGTGTATACACCCCTCCAACTCACTCCGACCAATTCGCTGATGATAAACAATCGATCATCCAA
Ebola IC '94	ACTGCCAGTGGCCCTAAATTACACAACAACACTGGTGTGATGACAAATGATACTCCCTCTTAATTCACTCGACCAAGTTGCCAGATGATAATATGATCATCCGA
Ebola Zaire '76	GGCAACACGAATACAGGCTTCCTCGACACCGGACTCAGTCAATGCGGACACTCCATCAGTCCCAATTCAGGCCCAATTGCCGATGACACCAATCGACCATGCCA
Ebola Bundibugyo '07	GTCATACCAACCAAGTGTTCATCAGCCTTTATACFCGAGGCAATGGTGCATATGATATCGGGCCGAAGTACTTAATGAAGCAAAATCCCTATATGGCT
Ebola IC '94	GCCACAGCCCTAACAGTGTTCCTCTGCAATTATATTTGGAAGCTATGGTGCATGTAAATATCTGGCCCCGAAAGTCTGTGATGAAGCAAAATCCCAATCTGGCT
Ebola Zaire '76	GGCACACACCAGGCAAGTGTCTCATCAGCATTCATCCTTGAAGCTATGGTGAATGTGCATATCGGGCCCCCAAGTCTTAATGAAGCAAAATTTCCAAATTTGGCT
Ebola Bundibugyo '07	CCCTTTGGGTGTGCTGATCAAAAACATATAGTTTTTGACTCAACTACAGCTGCAATATGCTGCAATCTGCATCTGACACCACTCACTACTTTGGCAAAACCTCC
Ebola IC '94	TCCTCTGGGTGTCTCTGACCAGAGACATATAGCTTTTGATTCAACCACTGCTGCCATATATGCTAGCATCATATACCATCACTCACTATTTGGCAAAACCTCA
Ebola Zaire '76	TCCTCTAGGTGTGCTGATCAAAAGACCTACAGCTTTTGACTCAACTACGGCCGCCATCATGCTTCTCATACACTATCACTCCCAATTTCCGGCAAGGCCAAC
Ebola Bundibugyo '07	AAATCCGCTTGTGAGAAATCACTGACTTGTGCTCGGGATCCCGATCACCCGCTTGCGGCTTCTGAAGATCAAGAAATCAAGGCTTTCTTCAAGAGTTTGTGC
Ebola IC '94	AAATCCGCTTGTGAGAAATCAACCGACTTGTGCTCGGCATACCTGATCACCCACTACAGCTCCCTAAGAAATAGGAATATCAAGCTTCTCTCAAGAGTTTGTGC
Ebola Zaire '76	AAATCCACTTGTGAGTCAATCGGCTGGGTCTGTGGAATCCCGGATCATCCCTCAGGCTCTCGGAATTGGAACACAGGCTTTCTCTCCAGGAGTTCTGTTTC
Ebola Bundibugyo '07	TGCTCTCAGTTCAATTGCGCAATTTTCACTTTTGAACCTGACGGCTCTAAAGCTGATCACTCAACCTCTCCCGGACGAACCTTGACCGGATGATATCTCC
Ebola IC '94	TGCTCTCAGTTCAACTGCGCAATTTTCACTTTTGAATCTGACGCTGCAAGCTGATCAACCACTCTCCGACGGGCAACCTTGACCGGATGATATCTCC
Ebola Zaire '76	TTTCGGCAGCTCAACTACCCCAATTTTCACTTTTGAATTTTGACAGCACTCAAACTGATCACCCCACTGCTGCTGTGCAACCTTGCTGCAACCTTGACCGATGACCTCC

Ebela Bundibugyo '07	5110	5120	5130	5140	5150	5160	5170	5180	5190	5200
Ebela IC '94	5210	5220	5230	5240	5250	5260	5270	5280	5290	5300
Ebela Zaire '76	5310	5320	5330	5340	5350	5360	5370	5380	5390	5400
Ebela Bundibugyo '07	5410	5420	5430	5440	5450	5460	5470	5480	5490	5500
Ebela IC '94	5510	5520	5530	5540	5550	5560	5570	5580	5590	5600
Ebela Zaire '76	5610	5620	5630	5640	5650	5660	5670	5680	5690	5700
Ebela Bundibugyo '07	5710	5720	5730	5740	5750	5760	5770	5780	5790	5800
Ebela IC '94	5810	5820	5830	5840	5850	5860	5870	5880	5890	5900
Ebela Zaire '76	5910	5920	5930	5940	5950	5960	5970	5980	5990	6000

[illegible]

Ebola Bundibugyo '07	8210	8220	8230	8240	8250	8260	8270	8280	8290	8300
Ebola IC '94	8310	8320	8330	8340	8350	8360	8370	8380	8390	8400
Ebola Zaire '76	8410	8420	8430	8440	8450	8460	8470	8480	8490	8500
Ebola Bundibugyo '07	8510	8520	8530	8540	8550	8560	8570	8580	8590	8600
Ebola IC '94	8610	8620	8630	8640	8650	8660	8670	8680	8690	8700
Ebola Zaire '76	8710	8720	8730	8740	8750	8760	8770	8780	8790	8800
Ebola Bundibugyo '07	8810	8820	8830	8840	8850	8860	8870	8880	8890	8900
Ebola IC '94	8910	8920	8930	8940	8950	8960	8970	8980	8990	9000
Ebola Zaire '76	9010	9020	9030	9040	9050	9060	9070	9080	9090	9100

FIG. 2

Ebola Bundibugyo '07	8910	8920	8930	8940	8950	8960	8970	8980	8990	9000
Ebola IC '94	AACTGGCGCTCCCTTGAACACAAATGAACATCACTGCTCTAAAGATACACAGATTAGCAATGCCAATGCGATGATTTCCAAACAAAAGACACGGCCCA	GACATGTGGATCCACGGAAACAACTAAGCATAGTTGCTCCAAAGATTCACTGCTGGCTAATCTCTATGCTGAGGATTTCCAAACAAAAGATGGCGCT	GACTTGTGGATCAGTAGAACAAATTAATAATACTGCAACCAAGGACTCGCGCTTAGCAATCCCAAGGCTGATGATTTCCAGCAAGAGGAAGGTCCA						
Ebola Zaire '76	9010	9020	9030	9040	9050	9060	9070	9080	9090	9100
Ebola Bundibugyo '07	AAAAATTACACTATTGACACTTTTGGAGACTGCGGAGTATTTGGTCAAAAACAAAGATATCAAGGGCAATTGATGACTCAAGACTTAAGAGCATTTACTAAACCCCTTT	AAAGTAACACTGTCGATGCTTATAGACACGACAGAGTATTTGGTCCAAAACAGGACATTAAGAACATTCGATGATTCAGATTAAGAGCTTTATTTGACCCCTTT	AAAATTACCTTTGCTGACATGATCAAGACGCGAGAACACTGGCGGAGACAAAGACATCAGAACCATAGAGGATTCAAAATTAAGAGCATTTGTTGACTCTAT						
Ebola IC '94	GTGCGGTCAATGACGAGGAAATTTCTCAAAATCCACAGCTTAGTCTATTTGTGTGAGAGTCACTCTACGACGAGAAAGGCTTAGGACAGGATCAATTCAGAACTCTGT	GTGCTGTTATGACGCGCAAAFTTCTCAAAATCTCACTTAGCTTGTATGTGAAGCCACTTACGCGGAGAAAGGACTTGGTCAAGACCAATTCAGAGTCAGT	GTGCTGTGATGACGAGGAAATTTCTCAAAATCCACAGCTGAGTCTTTTATGTGAGACACACACTTAAGCGCGAGGGGCTTGGGCAAGATCAGGCAGAACCCGCT						
Ebola Zaire '76	9110	9120	9130	9140	9150	9160	9170	9180	9190	9200
Ebola Bundibugyo '07	TCCTTGAAGTGTATCAGCGCTTACATAGCGACAAAGCGGAAATTTTGAAGCAGCCCTATGGCAACAAATGGGACCGACAGTCCCTTGATCATGTTTATAACA	TCCTGGAGGTATATCAACGCTTACACAGCGATTAAGGTGGAAATTCGAGCAGCACTATGGCAGCAGTGGGATGGGCAATCATGTGATTAATGTTTCATACAA	TCCTCGAAGTATATCAACGATTAACACAGTGTATAAGGAGCGCAGTTTGTGAAGCTGCACATATGGCAACAAATGGGACCGACAAATCCCTTAATTTATGTTTATCACT						
Ebola IC '94	GCATTTCTTAATATTGCTTTTACAATTACCCCTGTGAAGTTTCATCTGTTGTTTATTCAGGATTAAGGCTGCTAGTGCCTCAATCAGAAGATACCGAGACCT	GCATTTTAAATATTGCAATTACCAATTACCATGTGAGAGTTTCATCTGTTGTTTATTCAGGTTTGAGATTCGTGATACCCAGTCGGAAGCCACTGAGGTTG	GCATTTCTGAATATTGCTCTCCAGTTACCGTGTGAAGTTTCTGCTGCTGTTGTTTCAGGTTTAAAGAACATTTGGTTCCTCAATCAGATAATGAGGAAGCTT						
Ebola Zaire '76	9310	9320	9330	9340	9350	9360	9370	9380	9390	9400
Ebola Bundibugyo '07	CAACCTACACCGAGACACGTCATGCTGTCAGAGGAAGGTGGCCGCCCATTAACATCTTC-----CACAGTCGAATCTACCAATAATTTCCC--TATTCAACGC	TAACCCCTCCGAAACCTGCACATGCTCAGAGGAGGAGTTCCTCATTTAGGCCCA-----AATCAAGGGGAGCTAATAAATCCC--TTTTGACAT	CAACCAACCCGGGACATGCTCATGCTGCTGATGAGGTTACCCCTTAATAAGGCTGACTAAACACATATATAACCTTCTTACTTTCATCAATACTCCGTAT						
Ebola IC '94	AGATAAGAAATCAGTACTAAACACACAACTGCAAAAATTAACAA--AACACAGCATATAAGTGAATCTCTGCT--GTGATTAGCAACACG--AATGATCTTCAAT	GCATAACATCACATACAAATTTCAAAGGCATTGGATAAATGAGTATTTCAAGGAGATTAGTTTGGCTCAAAATCAGATCCGAGCAATTAATCATCTAC	ACCTATCATCATATATTATTAACAAGACGATATCCTTTAAACTTAT--TCAGTACTATATCACTCTCGTTCGTTTCAAAATTAATAAGATG---TGCATGATGTC						
Ebola Zaire '76	9510	9520	9530	9540	9550	9560	9570	9580	9590	9600

[illegible]

[illegible]

[illegible]

EBola Bundibugyo '07	TTTCGGTAAAGACGAATATTTAAATGGCGTCCAAATTTGGCTCAGTCCTTAAAGAGGCGTACCAAGACTGCTACACGAATGCTCCCTGTCAGATGCGAATCTTTTGATGACCTTTC
EBola IC '94	TTTGGCAAAACAAATATTTGAATGAGTACAAATTAACCTCAATCTCGAAACTGCTACTAGAAATGCAACCTTGTCAGATGCTATCTTTTGATGATCTTTC
EBola Zaire '76	TTTGGAAAAACAAATATTTGAATGGGGTCCAAATTTGGCTCAGTCCTTAAAGAGGCGTACCAAGATGCGACCAATGTTCTGTATGCCAATTTTGTGATGATCTTTC
EBola Bundibugyo '07	AGGAACTCTGGCTAGTATAGGAACGCAATTTGAGAGATCTATATTCGGAGACTAGACATGTATACCTTTCCGGGTGGTGGCCGCAATTCCTCAATCAATCTTT
EBola IC '94	AAGGACACTAGCTAGCATAGGCACGGCTTTTGAAGATCTATCTCCGAAACTAGCAGCTAGTCCCTGTGTAGTAGTAGCAGCTGCATTCCTCAATCCATCCCTTTT
EBola Zaire '76	AAGGACCCCTGGCTAGTATAGGCACCTGCTTTTGAAGCGATCCCATCTCTGAGACACGACATATCTTTCCCTTGGAGGATTAACCGACGCTTTTCCCATACGCTTTT
EBola Bundibugyo '07	CTTCGGTTAGGATCTCTCCAAATPACCAACCACCTTTGGTTTCAACAAAGGAACGGATCTAGGTCAACATPATCACTAAGCAAAACCGTTTGGATTTTCGGAACTACTCACT
EBola IC '94	TTCCGTAAGAACTCTTACAAATATCATCATCTTGGCTTCAACAAAGGNAACAGACTGGGTCAATTTGTCTAATTAAGCAAGCCATTAGATTTTGGAACTATAACT
EBola Zaire '76	TTCCGTTAGCAATCTTGCAATATCATCATCTCGGGTTCATAAAGTTTTGACCTTGGNCAAGTTPAACACTCGCGCAAACTCCTGGGATTTTCGGAAACAATATCA
EBola Bundibugyo '07	CTTGTCTTAAAGCGGTACCTCAAGTCTCTAGGAGGTTTATTCGTTTTAAACCCAGAGAAATGTTTTTATTCGCAACCTTTGGAGACCCCGTGACCTCCGGCCCTAT
EBola IC '94	TTGGCTTTGGCAGTACACAAAGTCTTGGGTGGCTTATCATCTCTAAATCCAGAAATGTTTTTATAGAAATCTGGGTGATGCTCTGTGTACTTCAGGCGCTGT
EBola Zaire '76	TTGGCACTAGCGGTACCGCAGGTCTTTGGAGGTTATCTCTCTTGAATCTCGAANAATGTTTCTACCGGAATCTAGGAGATCCAGTTCACCTCCAGGCTTAT
EBola Bundibugyo '07	TCCAACTTAGACTTACTCTGCAAAATGATCAACATGGACACCTTATTTCTACCTTTAATGCCAAGAACCCCGGGAACCTGTAGTGCATTAAGCTTTGTACT
EBola IC '94	TTTCAGTCTAAGACATATCTTCAATGATCCACATGGATGATTTGTTTTACCTTTGATCGCAAGAACCCAGGGAACCTGTAGCGCAATAGCTTTGTGT
EBola Zaire '76	TCCAGTTAAABACTTATCTCCGAATGATTTGAGATGGATGATTATTTCTTACCTTTAATTTGGCAAGAACCCCTGGGAACCTGCCACTGCCATTTGCTTGTGCT
EBola Bundibugyo '07	CAACCCACCGGATGATGATTCCTCGGTTCACAGATCTTACATCTTTCTTACGTAGATAGTGGCTAGAACCAATCTACTTTCAGTGCCTGCAAAATAAATTTA
EBola IC '94	AAACCTACTGGTGTAAACGTACCGGGTCCACAGATTTGACATCTCTTCTACGTACAGTGGCGGCAACCAATTACTACTTAAGTGCCTCAAAAATAAATTTA
EBola Zaire '76	AAATCTAGCGGATTAATGTCCTCGGTGCGAAGACTTAACCTTCATTTCTCGCGCCAGATTTGTACGGCAGGACCATCAACCTTAAGTGGCGAATAACAACACTT
EBola Bundibugyo '07	ATAAACACTTTGTTTCACTCTCAGCCGATTTAGAAATGAGATGGTATGTAATGGCTACTTTCTTCAACACCTGTAAATCAGTGGTGTGTGCTGCTGATA
EBola IC '94	ATAAACACTTTGTTTCACTCTTCTGCTGATTTAGNAAGATGAAATGGTTTGGTTTCTTCTTCTACACAGATCTAGTAGGTTTGGCCGCCGATA
EBola Zaire '76	ATTAATCTTATTTATCGGCTCAGCTGACTTGGAAAGCGAAATGGTTTGTAAATGGCTATTATATCACTCAACTCTCTGTTATGAGTCTGTTTGGCGCCGATA
EBola Bundibugyo '07	TATTTCTCTGTACTCCGAGTGGGAAGCGCTTGCGAGTCTCTAGGTTATTTAGAAAGGACTAGACCTTGTGTAGCCTCCAAAGTCACTCAATAACAATGCAGA
EBola IC '94	TATTTTCTCGACTCCOAGTGGGAACGTTTACAGATCTTGTAGGTTTACCTTTGAAGGACATAGACATTTGTATGCTCTTAAATATTAAATTAATAATTAAGTCA
EBola Zaire '76	CTTTTTCAGCGACCGCGAGGGGAAGCTGTGCAAAATTTCTAGGATCTCTAGGAAAGCAACGACCAATTTAGGACTCTCTTAAGATCATCAACAATAATACAGA

[illegible]

FIG. 2

Ebola Bundibugyo '07 Ebola IC '94 Ebola Zaire '76	15610 15620 15630 15640 15650 15660 15670 15680 15690 15700 GAGTTCTCAGGTGGGGGCAATCAGCAAGGCACACAATATCATCTTCCAAATGTAATCAATTTTCGGTTGGCCCTAATTTGATTTACGATTTCCGAACA GAAATTTTCAGGAGGAGCTCAGTCAGCAAGACAGATAGTATATCTCCAGAAATGTTTAAATTTTCCTTTGCTTTGCTTTGATACGATTTAGGAACG GAGTTTTTCAGGAGGTGGCCAGTCTGCACGCCACAGCAATATATATTTTCCAGAAATGTTTAAATTTATGTCAGTTGCAGTTGCATTTGATTTAAATTTAGAAACA
Ebola Bundibugyo '07 Ebola IC '94 Ebola Zaire '76	15710 15720 15730 15740 15750 15760 15770 15780 15790 15800 CCGAAACATCCCTCCATTTCAGCATATCGTGCCCATCTCCATCTTTCACAGTGTTCACACGGGAGCTCCAGCTCAATACCTACACGCTACGCT TGGCTACTCTTCTATACACATCATCGGGCTCACTCTTTCATTTGTGNAAGTGTTCACGGCAGAGGTTCCAGCCCAATATTTAGTTATACATCAACATTT CTGAGGCTACAGATATCCAAATATAATCGTGCTCACTTCATCTTAATAGTGTTCACCCGGGAGTACCAGCTCAGTATTTAAACATACACATCTACATT
Ebola Bundibugyo '07 Ebola IC '94 Ebola Zaire '76	15810 15820 15830 15840 15850 15860 15870 15880 15890 15900 TTCCCTTGGATCTCACAAAGTACCGAGAGATGAGTTAAATTTATGATAACAAATCCGTTTAAAGGTGGACTTAATTTGCAACCTATCCTTTGATATCCACTT GCCATTTGACCTTACACGGTATCGGGATAATGAGTTGATTTAGATGACATCCATCCATTAAGAGTGGTTTAAATTTGCAATCTTTCTTTGATTAATCCGCTT GGATTTAGATTTTAAACAGATACCGAGAAACGAAATGATTTATGACAGTAACTCTCTTAAAGGAGGACCTCAATTTGCAATATCTCATTCGATATATCCATTTT
Ebola Bundibugyo '07 Ebola IC '94 Ebola Zaire '76	15910 15920 15930 15940 15950 15960 15970 15980 15990 16000 TTCAAGGGCCAAAGGCTCAATATCATAGAGGAGGATTTGATAGATTTCTCATCTATCTGGTGGGAACTTGGGAAACCATCATTCAGTCCCATATCT TTCAAGGGCCAGAGCTTAACTAATTTGAAGAGACTTGTAGACTTACTTACTTATTCAGGATGGAGCTAGCTAAACCTGTTATCCCAATCTATATATTT TTCCAAGGTAAACGGCTGAACATATATAGATGATCTTATTCGACTCGCTCACTTATCTGGATGGAGCTGCTGCAAGACCATCATGCAATCAATATATTT
Ebola Bundibugyo '07 Ebola IC '94 Ebola Zaire '76	16010 16020 16030 16040 16050 16060 16070 16080 16090 16100 CAGACAGCAATAACTCATCCACAGACCCCATTTAGCAGTGGAGAAACACGATCATTCACACTCACCTTTTCACATATCTTAAGGTGGGCTCCCTATAG CTGACAGCAACAAATTCATCAACGGATCCAAATCAGTAGTGGGAAACACGATCATTCACCACTCACTCTTCGACATATCTTAAGATGGACTACTATATAG CAGATAGCAACAAATTCATCTACAGACCCCAATTAGCAGTGGAGAAACAGATCATCTACCTACCACTTTCTTAACCTATCCCAAGATAGGACTTCTGTACAG
Ebola Bundibugyo '07 Ebola IC '94 Ebola Zaire '76	16110 16120 16130 16140 16150 16160 16170 16180 16190 16200 TTTCGGCGCCATCGTCAGTTATTACTTAGGGAATACCAATTTAGGACCAAAAGCTAGACCTCAGCTCATTTTATGTTATTTAACAACCTCAAAATCCAT TTTTGGTGACATCATGATTATATCTAGGCAACACCAATTTAGAACCAAAATTCAGCTCTTAACAATTCATATATTAACCTTATATTCAGTACTCAAAATACAT TTTTGGGCGCTTTTAAAGTTATTATCTTGGCAATACAAATCTTCGGACTAAGAAATTAACACTTGCAATTTTTTATATTTACTTTAACTACTCAAAATTCAT
Ebola Bundibugyo '07 Ebola IC '94 Ebola Zaire '76	16210 16220 16230 16240 16250 16260 16270 16280 16290 16300 TTTGCCACATCGCTCGTTGAGGATACTTAAGCCACCTTTAAACAGTGTAGTGTATCAGACTAATGAGTATTCCTCTCATTTTTCATCTCAATCTTACA AATTTACCTCATCGCTCGTTGAGAAATCTTAAACCTACTTTGAAACACACGATGTTATCTCGAGATTAATAGTATTCAGCTCTCACTTCTCAATTTATA AATCTACCAATCGCTCATTTGGGAATACTTAAGCCCAACATTCAAACATGCAAGGTTATGTCACGGTTAATGAGTATTCAGTCTCTCATTTTCTATTTACA
Ebola Bundibugyo '07 Ebola IC '94 Ebola Zaire '76	16310 16320 16330 16340 16350 16360 16370 16380 16390 16400 TCGGGGTACGGCAGGTGATCGAGGGCTTTTCGGATGCTACCACTATTCTCTCGAGTGGCCATTTCTCTCTTCCCTTCAATTTATCAAAAAATGGATCGT TTGAGGAACTCTGCTGTGTGATCGAGGACCTTCGGATGGCGGACAGATTGTTCTTTAGAACTGCCATTAATGCTCTCTCTCAATTCGTTAGAAAGTGGATAGT TAGCGGTGCTGCAGGTGACAGAGGACTCTCAGATGCGGCCAGGTTATTTTGGAGACGCTCCATTTTCATCTTTTCTTACATTTTGTAAAGATGGATAAT

[illegible]

[illegible]

FIG. 2

	17910	17920	17930	17940	17950	17960	17970	17980	17990	18000
Ebola Bundibugyo '07	AAAGGTTCTTTACCATCGATATAAACCTAGTTGATTCACGGAAGGTCCTGCTCGATCCCTTTACCATTTAACACACTTGCACGAGAGATTAGAGAA								
Ebola IC '94	GAGGTTTATACCATAGATACAAATTTAGTCGATTCCTAGAAAGCCCTTTGACTTCCATTTGCCACATCTAGCGACCTGCAGACCCGAGATTAGGGAG								
Ebola Zaire '76	GAAAGTACTATACCAACAGGTATAACCTCGTCGATTCAAAAGAGGTCCTAGTCTCTATCACTCAGCACTTAGCACATCTTTAGACGACAGATTTCGAGAA								
	18010	18020	18030	18040	18050	18060	18070	18080	18090	18100
Ebola Bundibugyo '07	TTAGTGTGTGACTATATATCAGCAACGACAAAGTCGAACCCAAACATACCACTTCATCAAACGACAAAGGCCGGATTACAAAATTAGTCAATGACTACC								
Ebola IC '94	TTGGTTAATGACTATATATCAACAAAGTCGAACCCAAACATATCATTTTCATTAACAAATAAAAAGGTCGTATTACAAAATTGGTAAATGATTACC								
Ebola Zaire '76	TTAACAATGATATATATCAACAGCGCAAAAGTCGGACTCAACCATATCATTTATTCTGTACTGCAAAAGGACGAATCACAACACTAGTCAATGATTATT								
	18110	18120	18130	18140	18150	18160	18170	18180	18190	18200
Ebola Bundibugyo '07	TTAAATTTTATCTCGTAGTCGAAGCACTGAAGCATAAATGTCTTTGGCAGGAAGAACTCAGAACACTCTCTGACTTAATCAATGTGTTGCAATCGATTTTA								
Ebola IC '94	TTAAGTTCTTTCTAATAATACAAAGCCTTAAAGCAAAATGACACATGGCAAGAGGAAGTAAGAGCTCTCCAGATCTAATAGTCTCTGCACTCGATTCTA								
Ebola Zaire '76	TAAATTTCTTTCTTATTGTGCAAGCAATTAAACATAATGGGACATGGCAAGCTGAGTTTAAGAAATTACCAGAGTTGATTAGTGTGCAATAGGTTCTA								
	18210	18220	18230	18240	18250	18260	18270	18280	18290	18300
Ebola Bundibugyo '07	CCATATAAGGAGCTGCTCATGTGAAAGATCGATTTTAACTTAACTTTTACTTAACCCGTAATGCAAGACTCAGAACTCAAGAACTTAATGAGAGAGATTAAAC								
Ebola IC '94	TCATACTCGAAACTGTTTCATGTGAAACCGGTTCTCTAGTACAGACTTTTACTTATCACGGATGCGAGATTCCGAAATCAAACTTAATAGATAGATTGACC								
Ebola Zaire '76	CCATATTAGAGATTGCAATTTGTGAAGAACGTTTCTTCTAGTTCAAACTTATATTACATAGAATGCAAGATTCTGAAGTTAAGCTTATCGAAAGGCTGACA								
	18310	18320	18330	18340	18350	18360	18370	18380	18390	18400
Ebola Bundibugyo '07	GGGTTTCTAGGATTGTATCCTAATGGTATTAAACGCTTAAGATCCCTTAGAGGCAT-CGCAATATAGCTCCAAACATTAAATGATATTGCTGTCAATACA								
Ebola IC '94	GGCCTTCTTAGTCTATGTCCTCAATGGTTTTCGCGTAAGGACTCTTGAGCTACAACTCCACATAGTTATACA-ATGGTACCAGGACACTATATGTAAA								
Ebola Zaire '76	GGGCTTCTGAGTTTATTTCCGGATGGTCTCTACAGGTTTGATTGAATTAC--CGTGATAGTATCCTGATAC---TTGCAAAAGGTTGGTTATTAAACATA								
	18410	18420	18430	18440	18450	18460	18470	18480	18490	18500
Ebola Bundibugyo '07	TCTACCTGACCGAGAGCAAGGTTTA-TTATAAATAAACCTATATACATGACTGCAATGCGTAATTTATACCGAAACACAGTGGGCTGCACATGCAGGTT								
Ebola IC '94	TTGACCCCTAA-GAAGAGTAATTCG-ACACACAGAGTTCTCAAGTGAAACCCCTCATCTCAGATTATCTGTGGTTGCAATTCATAATCCGATT--GTTA								
Ebola Zaire '76	CAGATTATAAATAAACCTCATAAATTGCTCTCATACATCATATTGATCTTAATCTCAATAAACA-CTATTAAATAACGAAGAGTCCCTATATATTATATAC								
	18510	18520	18530	18540	18550	18560	18570	18580	18590	18600
Ebola Bundibugyo '07	CCTGTTGAGCTTTAAAGATCATGC-ATATATAAATGATTTTGTATACATAATCTAGTTAGTACTAATAACAGTACTACATCATATATCTATCAATT								
Ebola IC '94	CCCCGTGAG----TATAACTCCAGATTATATAGAAAATACCT-TTTGTCTGCAAAATTTATCTTAATTCAGATACATACGCTCCCAATCGTATAAAAT								
Ebola Zaire '76	TATATTAGCC-TCTCTCCCTGGGTGATAATCAAAAAATTAC---AATGCAGCATGTGTGCATAT-----TACTCCCGCAATGAATTTAACGCAAC								

HUMAN EBOLA VIRUS SPECIES AND COMPOSITIONS AND METHODS THEREOF

RELATED APPLICATIONS

[0001] This application claims priority benefit of U.S. Provisional Application 61/108,175 filed 24 Oct. 2008; the contents of which are hereby incorporated by reference.

DEPOSIT STATEMENT

[0002] The invention provides the isolated human Ebola (hEbola) viruses denoted as Bundibugyo (EboBun) deposited with the Centers for Disease Control and Prevention ("CDC"; Atlanta, Ga., United States of America) on Nov. 26, 2007 and accorded an accession number 200706291. This deposit was not made to an International Depository Authority (IDA) as established under the Budapest Treaty on the International Recognition of the Deposit of Microorganisms for the Purposes of Patent Procedure, and is a non-Budapest treaty deposit. The deposited organism is not acceptable by American Type Culture Collection (ATCC), Manassas, Va., an International Depository Authority (IDA) as established under the Budapest Treaty on the International Recognition of the Deposit of Microorganisms for the Purposes of Patent Procedure. Samples of the stated Deposit Accession No. 200706291 will be made available to approved facilities for thirty years from the date of deposit, and for the lifetime of the patent issuing from, or claiming priority to this application.

FIELD OF THE INVENTION

[0003] The invention is related to compositions and methods directed to a novel species of human Ebola (hEbola) virus.

BACKGROUND OF THE INVENTION

[0004] The family Filoviridae consists of two genera, Marburgvirus and Ebolavirus, which have likely evolved from a common ancestor¹. The genus Ebolavirus includes four species: Zaire, Sudan, Reston and Côte d'Ivoire (Ivory Coast) ebolaviruses, which have, with the exception of Reston and Côte d'Ivoire ebolaviruses, been associated with large hemorrhagic fever (HF) outbreaks in Africa with high case fatality (53-90%)².

[0005] Viruses of each species have genomes that are at least 30-40% divergent from one another, a level of diversity that presumably reflects differences in the ecological niche they occupy and in their evolutionary history. Identification of the natural reservoir of ebolaviruses remains somewhat elusive, although recent PCR and antibody data suggest that three species of arboreal fruit bats may be carriers of Zaire ebolavirus³. No data has yet been published to suggest reservoirs for the Sudan, Reston and Côte d'Ivoire ebolavirus species. However, a cave-dwelling fruit bat has been recently implicated as a natural host for marburgvirus^{4, 5}, supporting the hypothesis that different bat species may be the reservoir hosts for the various filoviruses.

[0006] Filovirus outbreaks are sporadic, sometimes interspersed by years or even decades of no apparent disease activity. The last new species of ebolavirus was discovered 14 years ago (1994), in Côte d'Ivoire (Ivory Coast), and involved a single non-fatal case, a veterinarian who performed an autopsy on an infected chimpanzee found in the Tai Forest⁶. No further disease reports have been associated with Côte

d'Ivoire ebolavirus, in contrast to Zaire and Sudan ebolaviruses which have each caused multiple large outbreaks over the same time period.

[0007] In late November 2007, HF cases were reported in the townships of Bundibugyo and Kikyo in Bundibugyo District, Western Uganda. The outbreak continued through January 2008, and resulted in approximately 149 cases and 37 deaths⁷. Laboratory investigation of the initial 29 suspect-case blood specimens by classic methods (antigen capture, IgM and IgG ELISA) and a recently developed random-primed pyrosequencing approach identified this to be an Ebola HF outbreak associated with a new discovered ebolavirus species. These specimens were negative when initially tested with highly sensitive real-time RT-PCR assays specific for all known Zaire and Sudan ebolaviruses and Marburg viruses. This new species is referred to herein as "the Bundibugyo species", abbreviated "EboBun".

[0008] Accordingly, compositions and methods directed to the new Ebola virus species are described herein and the most closely related Ebola Ivory Coast species, which compositions and methods are useful for diagnosis and prevention of human Ebola virus infection; including related vaccine development, and prevention of hemorrhagic fever in a human population.

SUMMARY OF THE INVENTION

[0009] The present invention is based upon the isolation and identification of a new human Ebola virus species, EboBun. EboBun was isolated from the patients suffering from hemorrhagic fever in a recent outbreak in Uganda. The isolated virus is a member of the Filoviridae family, a family of negative sense RNA viruses. Accordingly, the invention relates to the isolated EboBun virus that morphologically and phylogenetically relates to known members filoviridae.

[0010] In one aspect, the invention provides the isolated EboBun virus deposited with the Centers for Disease Control and Prevention ("CDC"; Atlanta, Ga., United States of America) on Nov. 26, 2007 and accorded an accession number 200706291, as stated in the paragraph entitled "DEPOSIT STATEMENT" supra.

[0011] In another aspect, the invention provides an isolated hEbola EboBun virus comprising a nucleic acid molecule comprising a nucleotide sequence selected from the group consisting of: a) a nucleotide sequence set forth in SEQ ID NO: 1; b) a nucleotide sequence that hybridizes to the sequence set forth in SEQ ID NO: 1 under stringent conditions; and c) a nucleotide sequence that has at least 70%, 75%, 80%, 85%, 90%, 95%, 96%, 97%, 98%, or 99% identity to the SEQ ID NO: 1. In another aspect, the invention provides the complete genomic sequence of the hEbola virus EboBun.

[0012] In a related aspect, the invention provides nucleic acid molecules isolated from EboBun, or fragments thereof.

[0013] In another aspect, the invention provides proteins or polypeptides that are isolated from the EboBun, including viral proteins isolated from cells infected with the virus but not present in comparable uninfected cells; or fragments thereof. In one embodiment of the present invention, the amino acid sequences of the proteins or polypeptides are set forth in SEQ ID NOS: 2-9 and 59, or fragments thereof.

[0014] In a related aspect, the invention provides an isolated polypeptide encoded by the nucleic acid molecule of the inventive hEbola EboIC (Sequence ID No. 10) virus described above.

[0015] In another aspect, the invention provides an isolated hEbola EboIC virus comprising a nucleic acid molecule comprising a nucleotide sequence selected from the group consisting of: a) a nucleotide sequence set forth in SEQ ID NO: 10; b) a nucleotide sequence that hybridizes to the sequence set forth in SEQ ID NO: 10 under stringent conditions; and c) a nucleotide sequence that has at least 70%, 75%, 80%, 85%, 90%, 95%, 96%, 97%, 98%, or 99% identity to the SEQ ID NO: 10. In another aspect, the invention provides the complete genomic sequence of the hEbola virus EboIC.

[0016] In a related aspect, the invention provides nucleic acid molecules isolated from EboIC, or fragments thereof.

[0017] In another aspect, the invention provides proteins or polypeptides that are isolated from the EboIC, including viral proteins isolated from cells infected with the virus but not present in comparable uninfected cells; or fragments thereof. In one embodiment of the present invention, the amino acid sequences of the proteins or polypeptides are set forth in SEQ ID NOs: 11-19, or fragments thereof.

[0018] In a related aspect, the invention provides an isolated polypeptide encoded by the nucleic acid molecule of the inventive hEbola EboIC virus described above.

[0019] In other aspects, the invention relates to the use of the isolated hEbola virus for diagnostic and therapeutic methods based on EbBun, EboIC, or a combination thereof. In one embodiment, the invention provides a method of detecting in a biological sample an antibody immunospecific for the genus of West African Ebola Species constituting hEbola EbBun and EboIC virus using at least one the inventive isolated hEbola virus described herein, or any of the inventive proteins or polypeptides as described herein. In another specific embodiment, the invention provides a method of screening for an antibody which immunospecifically binds and neutralizes hEbola EbBun. Such an antibody is useful for a passive immunization or immunotherapy of a subject infected with hEbola.

[0020] In another aspect, the invention provides an isolated antibody or an antigen-binding fragment thereof which immunospecifically binds to the hEbola virus of the invention described above.

[0021] In other aspects, the invention provides methods for detecting the presence, activity or expression of the Glade of Bundibungyo-Ivory Coast hEbola virus in a biological material, such as cells, blood, saliva, urine, feces and so forth; and specifically at least one of EbBun or EboIC.

[0022] In a related aspect, the invention provides a method for detecting the presence of the inventive hEbola virus described above in a biological sample, the method includes (a) contacting the sample with an agent that selectively binds to a West African hEbola virus; and (b) detecting whether the compound binds to the West African hEbola virus in the sample.

[0023] In another aspect, the invention provides a method for detecting the presence of the inventive polypeptide described above, in a biological sample, said method includes (a) contacting the biological sample with an agent that selectively binds to the polypeptide; and (b) detecting whether the agent binds to the polypeptide in the sample. In another aspect, the invention provides a method for detecting the presence of a first nucleic acid molecule derived from the inventive hEbola virus described above in a biological sample, the method comprising: (a) contacting the biological

sample with an agent that selectively binds to the polypeptide; and (b) detecting whether the agent binds to the polypeptide in the sample.

[0024] In another aspect, the invention provides a method for propagating the hEbola virus in host cells comprising infecting the host cells with the inventive isolated hEbola virus described above, culturing the host cells to allow the virus to multiply, and harvesting the resulting virions. Also provided by the present invention are host cells infected with the inventive hEbola virus described above.

[0025] In another aspect, the invention provides a method of detecting in a biological sample the presence of an antibody that immunospecifically binds hEbola virus, the method comprising: (a) contacting the biological sample with the inventive host cell host described above; and (b) detecting the antibody bound to the cell.

[0026] In another aspect, the invention provides vaccine preparations, comprising the inventive hEbola virus, including recombinant and chimeric forms of the virus, nucleic acid molecules comprised by the virus, or protein subunits of the virus. The invention also provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of the inventive hEbola virus described above, and a pharmaceutically acceptable carrier. In one embodiment, the invention provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of a protein extract of the inventive hEbola virus described above, or a subunit thereof; and a pharmaceutically acceptable carrier. In another, the invention provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of a nucleic acid molecule comprising the nucleotide sequence of SEQ ID NO: 1 or a complement thereof, and a pharmaceutically acceptable carrier. In another, the invention provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of a nucleic acid molecule comprising any of inventive the nucleotide sequences as described above, or a complement thereof, and a pharmaceutically acceptable carrier.

[0027] In a related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of the inventive hEbola virus described above, and a pharmaceutically acceptable carrier. In another related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of a protein extract of the inventive hEbola virus described above or a subunit thereof, and a pharmaceutically acceptable carrier. In another related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of a nucleic acid molecule comprising the nucleotide sequence of SEQ ID NO: 1 or a complement thereof, and a pharmaceutically acceptable carrier. In another related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of a nucleic acid molecule comprising the inventive nucleotide sequence as described above or a complement thereof, and a pharmaceutically acceptable carrier. In another related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of any of the inventive polypeptides described above.

[0028] In another aspect, the present invention provides pharmaceutical compositions comprising antiviral agents of the present invention and a pharmaceutically acceptable carrier. In a specific embodiment, the antiviral agent of the invention is an antibody that immunospecifically binds hEbola

virus or any hEbola epitope. In another specific embodiment, the antiviral agent is a polypeptide or protein of the present invention or nucleic acid molecule of the invention.

[0029] In a related aspect, the invention provides a pharmaceutical composition comprising a prophylactically or therapeutically effective amount of an anti-hEbola EboBun agent and a pharmaceutically acceptable carrier.

[0030] The invention also provides kits containing compositions and formulations of the present invention. Thus, in another aspect, the invention provides a kit comprising a container containing the inventive immunogenic formulation described above. In another aspect, the invention provides a kit comprising a container containing the inventive vaccine formulation described above. In another, the invention provides a kit comprising a container containing the inventive pharmaceutical composition described above. In another, the invention provides a kit comprising a container containing the inventive vaccine formulation described above. In another, the invention provides a method for identifying a subject infected with the inventive hEbola virus described above, comprising: (a) obtaining total RNA from a biological sample obtained from the subject; (b) reverse transcribing the total RNA to obtain cDNA; and (c) amplifying the cDNA using a set of primers derived from a nucleotide sequence of the inventive hEbola virus described above.

[0031] The invention further relates to the use of the sequence information of the isolated virus for diagnostic and therapeutic methods.

[0032] In another aspect, the present invention provides methods for screening antiviral agents that inhibit the infectivity or replication of hEbola virus or variants thereof.

[0033] The invention further provides methods of preparing recombinant or chimeric forms of hEbola.

BRIEF DESCRIPTION OF THE DRAWINGS

[0034] FIG. 1 represents a Phylogenetic tree comparing full-length genomes of Ebolavirus and Marburg virus by Bayesian analysis;

[0035] FIG. 2 represents an alignment of genomes of novel hEbola EboBun (SEQ ID NO: 1) referred to below as “Ebola Bundibugyo” or “EboBun”, and hEbola Zaire (SEQ ID NO: 20); referred to below as “Ebola Zaire ’76” or “EboZ” and hEbola Ivory Coast (SEQ ID NO: 10) also referred to below as “EboIC”.

DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENTS

[0036] It is to be understood that the present invention is not limited to particular embodiments described, as such may, of course, vary. It is also to be understood that the terminology used herein is for the purpose of describing particular embodiments only, and is not intended to be limiting.

[0037] Due to the sequence divergence of EboBun relative to all previously recognized ebolaviruses, the present invention has utility in design of diagnostic assays to monitor Ebola HF disease in humans and animals, and develop effective antivirals and vaccines.

[0038] The EboBun virus of the present invention is genetically distinct, differing by more than 30% at the genome level from all other known ebolavirus species. The unique nature of this virus created challenges for traditional filovirus molecular based diagnostic assays and genome sequencing approaches. Instead, over 70% of the virus genome was

sequenced using a recently developed random-primed pyrosequencing approach which allowed the rapid development of molecular detection assay which were deployed in the disease outbreak response. This random-primed pyrosequencing draft sequence allowed faster completion of the whole genome sequence using traditional primer walking approach and confirmation that the EboBun virus represented a new ebolavirus species.

Definitions

[0039] The definitions herein provided are operative throughout the entire description of the invention set forth herein, including the Summary of the Invention.

[0040] The term “an antibody or an antibody fragment that immunospecifically binds a polypeptide of the invention” as used herein refers to an antibody or a fragment thereof that immunospecifically binds to the polypeptide encoded by the nucleotide sequence of SEQ ID NO: 1 (EboBun), or a fragment thereof, and does not non-specifically bind to other polypeptides. An antibody or a fragment thereof that immunospecifically binds to the polypeptide of the invention may cross-react with other antigens. Preferably, an antibody or a fragment thereof that immunospecifically binds to a polypeptide of the invention does not cross-react with other antigens. An antibody or a fragment thereof that immunospecifically binds to the polypeptide of the invention can be identified by, for example, immunoassays or other techniques known to those skilled in the art, or otherwise as described herein.

[0041] An “isolated” or “purified” peptide or protein is substantially free of cellular material or other contaminating proteins from the cell or tissue source from which the protein is derived, or substantially free of chemical precursors or other chemicals when chemically synthesized. The language “substantially free of cellular material” includes preparations of a polypeptide/protein in which the polypeptide/protein is separated from cellular components of the cells from which it is isolated or recombinantly produced. Thus, a polypeptide/protein that is substantially free of cellular material includes preparations of the polypeptide/protein having less than about 30%, 20%, 10%, 5%, 2.5%, or 1% (by dry weight) of contaminating protein. When the polypeptide/protein is recombinantly produced, it is also preferably substantially free of culture medium, i.e., culture medium represents less than about 20%, 10%, or 5% of the volume of the protein preparation.

[0042] When polypeptide/protein is produced by chemical synthesis, it is preferably substantially free of chemical precursors or other chemicals, i.e., it is separated from chemical precursors or other chemicals which are involved in the synthesis of the protein. Accordingly, such preparations of the polypeptide/protein have less than about 30%, 20%, 10%, 5% (by dry weight) of chemical precursors or compounds other than polypeptide/protein fragment of interest. In a preferred embodiment of the present invention, polypeptides/proteins are isolated or purified.

[0043] An “isolated” nucleic acid molecule is one which is separated from other nucleic acid molecules which are present in the natural source of the nucleic acid molecule. Moreover, an “isolated” nucleic acid molecule, such as a cDNA molecule, can be substantially free of other cellular material, or culture medium when produced by recombinant techniques, or substantially free of chemical precursors or other chemicals when chemically synthesized. In a preferred embodiment of the invention, nucleic acid molecules encoded

ing polypeptides/proteins of the invention are isolated or purified. The term “isolated” nucleic acid molecule does not include a nucleic acid that is a member of a library that has not been purified away from other library clones containing other nucleic acid molecules.

[0044] The term “portion” or “fragment” as used herein includes the specified fragment lengths, and all integers in between, inclusive of the specified end points in a specified range, and inclusive of any length up to the full length of a protein, polypeptide, or nucleic acid.

[0045] The term “having a biological activity of the protein” or “having biological activities of the polypeptides of the invention” refers to the characteristics of the polypeptides or proteins having a common biological activity, similar or identical structural domain, and/or having sufficient amino acid identity to the polypeptide encoded by the nucleotide sequence of SEQ ID NO: 1 (EboBun). Such common biological activities of the polypeptides of the invention include antigenicity and immunogenicity.

[0046] The term “under stringent condition” refers to hybridization and washing conditions under which nucleotide sequences having at least 70%, at least 75%, at least 80%, at least 85%, at least 90%, or at least 95% identity to each other remain hybridized to each other. Such hybridization conditions are described in, for example but not limited to, Current Protocols in Molecular Biology, John Wiley & Sons, NY (1989), 6.3.1-6.3.6; Basic Methods in Molecular Biology, Elsevier Science Publishing Co., Inc., NY (1986), pp. 75-78, and 84-87; and Molecular Cloning, Cold Spring Harbor Laboratory, NY (1982), pp. 387-389, and are well known to those skilled in the art. A preferred, non-limiting example of stringent hybridization conditions is hybridization in 6× sodium chloride/sodium citrate (SSC), 0.5% SDS at about 68° C. followed by one or more washes in 2×SSC, 0.5% SDS at room temperature. Another preferred, non-limiting example of stringent hybridization conditions is hybridization in 6×SSC at about 45° C., followed by one or more washes in 0.2×SSC, 0.1% SDS at about 50-65° C.

[0047] The term “variant” as used herein refers either to a naturally occurring genetic mutant of hEbola EboBun, or hEbola EboIC, or a recombinantly prepared variation of these hEbola species, each of which contain one or more mutations in its genome compared to the hEbola of SEQ ID NO: 1 or 10. The term “variant” may also refer either to a naturally occurring variation of a given peptide or a recombinantly prepared variation of a given peptide or protein in which one or more amino acid residues have been modified by amino acid substitution, addition, or deletion.

[0048] “Homology” refers to sequence similarity or, alternatively, sequence identity, between two or more polynucleotide sequences or two or more polypeptide sequences.

[0049] The terms “percent identity” and “% identity,” as applied to polynucleotide sequences, refer to the percentage of identical nucleotide matches between at least two polynucleotide sequences aligned using a standardized algorithm. Such an algorithm may insert, in a standardized and reproducible way, gaps in the sequences being compared in order to optimize alignment between two sequences, and therefore achieve a more meaningful comparison of the two sequences.

[0050] Percent identity between polynucleotide sequences may be determined using one or more computer algorithms or programs known in the art or described herein. For example, percent identity can be determined using the default parameters of the CLUSTAL V algorithm as incorporated into the

MEGALIGN version 3.12e sequence alignment program. This program is part of the LASERGENE software package, a suite of molecular biological analysis programs (DNASTAR, Madison, Wis.). CLUSTAL V is described in Higgins, D. G. and P. M. Sharp (1989; CABIOS 5:151-153) and in Higgins, D. G. et al. (1992; CABIOS 8:189-191). For pairwise alignments of polynucleotide sequences, the default parameters are set as follows: Ktuple=2, gap penalty=5, window=4, and “diagonals saved”=4. The “weighted” residue weight table is selected as the default.

[0051] Alternatively, a suite of commonly used and freely available sequence comparison algorithms which can be used is provided by the National Center for Biotechnology Information (NCBI) Basic Local Alignment Search Tool (BLAST) (Altschul, S. F. et al. (1990) J. Mol. Biol. 215:403-410), which is available from several sources, including the NCBI, Bethesda, Md., and on the NCBI World Wide Web site available on the Internet. The BLAST software suite includes various sequence analysis programs including “blastn,” that is used to align a known polynucleotide sequence with other polynucleotide sequences from a variety of databases. Also available is a tool called “BLAST 2 Sequences” that is used for direct pairwise comparison of two nucleotide sequences. “BLAST 2 Sequences” can be accessed and used interactively on the Internet via the NCBI World Wide Web site as well. The “BLAST 2 Sequences” tool can be used for both blastn and blastp (discussed below). BLAST programs are commonly used with gap and other parameters set to default settings. For example, to compare two nucleotide sequences, one may use blastn with the “BLAST 2 Sequences” tool Version 2.0.12 (Apr. 21, 2000) set at default parameters. Such default parameters may be, for example: Matrix:BLOSUM62; Reward for match: 1; Penalty for mismatch: -2; Open Gap: 5 and Extension Gap: 2 penalties; Gap×drop-off: 50; Expect: 10; Word Size: 11; Filter: on.

[0052] Percent identity may be measured over the length of an entire defined sequence, for example, as defined by a particular SEQ ID number, or may be measured over a shorter length, for example, over the length of a fragment taken from a larger, defined sequence, for instance, a fragment of at least 20, at least 30, at least 40, at least 50, at least 70, at least 100, or at least 200 contiguous nucleotides. Such lengths are exemplary only, and it is understood that any fragment length supported by the sequences shown herein, in the tables, figures, or sequence listing, may be used to describe a length over which percentage identity may be measured.

[0053] The phrases “percent identity” and “% identity”, as applied to polypeptide sequences, refer to the percentage of identical residue matches between at least two polypeptide sequences aligned using a standardized algorithm. Methods of polypeptide sequence alignment are well known. Some alignment methods take into account conservative amino acid substitutions. Such conservative substitutions, explained in more detail above, generally preserve the charge and hydrophobicity at the site of substitution, thus preserving the structure (and therefore function) of the polypeptide. The phrases “percent similarity” and “% similarity”, as applied to polypeptide sequences, refer to the percentage of residue matches, including identical residue matches and conservative substitutions, between at least two polypeptide sequences aligned using a standardized algorithm. In contrast, conservative substitutions are not included in the calculation of percent identity between polypeptide sequences.

[0054] Percent identity between polypeptide sequences may be determined using the default parameters of the CLUSTAL V algorithm as incorporated into the MEGA-LIGN version 3.12e sequence alignment program (described and referenced above). For pairwise alignments of polypeptide sequences using CLUSTAL V, the default parameters are set as follows: Ktuple=1, gap penalty=3, window=5, and “diagonals saved”=5. The PAM250 matrix is selected as the default residue weight table.

[0055] Alternatively the NCBI BLAST software suite may be used. For example, for a pairwise comparison of two polypeptide sequences, one may use the “BLAST 2 Sequences” tool Version 2.0.12 (Apr. 21, 2000) with blastp set at default parameters. Such default parameters may be, for example: Matrix: BLOSUM62; Open Gap: 11 and Extension Gap: 1 penalties; Gap×drop-off: 50; Expect: 10; Word Size: 3; Filter: on.

[0056] Percent identity may be measured over the length of an entire defined polypeptide sequence, for example, as defined by a particular SEQ ID number, or may be measured over a shorter length, for example, over the length of a fragment taken from a larger, defined polypeptide sequence, for instance, a fragment of at least 15, at least 20, at least 30, at least 40, at least 50, at least 70 or at least 150 contiguous residues. Such lengths are exemplary only, and it is understood that any fragment length supported by the sequences shown herein, in the tables, figures or sequence listing, may be used to describe a length over which percentage identity may be measured.

[0057] The term “agent” encompasses any chemical, biochemical, or biological molecule; such as small molecules, proteins, polypeptides, antibodies, nucleic acid molecules including DNA or RNA, and the like.

Methods and Compositions Related to the Inventive hEbola

[0058] The present invention is based upon the isolation and identification of a new human Ebola virus species, EboBun and the sequencing of the only other known West African Ebola species EboIC. EboBun was isolated from the patients suffering from hemorrhagic fever in a recent outbreak in Uganda. The isolated virus is a member of the Filoviridae family, a family of negative sense RNA viruses. Accordingly, the invention relates to the isolated EboBun or EBOIC virus that morphologically and phylogenetically relates to known members filoviridae.

[0059] In another aspect, the invention provides an isolated hEbola virus including a nucleic acid molecule with a nucleotide sequence that is preferably: a) a nucleotide sequence set forth in SEQ ID NO: 1; b) a nucleotide sequence that hybridizes to the sequence set forth in SEQ ID NO: 1 under stringent conditions; or c) a nucleotide sequence that has at least 70%, 75%, 80%, 85%, 90%, 95%, 96%, 97%, 98%, or 99% identity to the SEQ ID NO: 1. In one embodiment of the present invention, the hEbola virus is killed. In another, the virus is attenuated. In another, the infectivity of the attenuated hEbola virus is reduced. In another, the infectivity is reduced by at least 5-fold, 10-fold, 25-fold, 50-fold, 100-fold, 250-fold, 500-fold, or 10,000-fold. In another, the replication ability of the attenuated hEbola virus is reduced. In another, the replication ability of the attenuated virus is reduced by at least 5-fold, 10-fold, 25-fold, 50-fold, 100-fold, 250-fold, 500-fold, 1,000-fold, or 10,000-fold. In another, the protein synthesis ability of the attenuated virus is reduced. In another, the protein synthesis ability is reduced by at least 5-fold, 10-fold, 25-fold, 50-fold, 100-fold, 250-fold, 500-fold, 1,000-fold, or

10,000-fold. In another, the assembling ability of the attenuated hEbola virus is reduced. In another, the assembling ability of the attenuated virus is reduced by at least 5-fold, 10-fold, 25-fold, 50-fold, 100-fold, 250-fold, 500-fold, 1,000-fold, or 10,000-fold. In another, the cytopathic effect of the attenuated hEbola virus is reduced. In another, the cytopathic effect is reduced by at least 5-fold, 10-fold, 25-fold, 50-fold, 100-fold, 250-fold, 500-fold, 1,000-fold, or 10,000-fold.

[0060] In another aspect, the invention provides the complete genomic sequence of the hEbola virus EboBun or EboIC. In a specific embodiment, the virus includes a nucleotide sequence of SEQ ID NOs: 1 or 10, respectively.

[0061] In a related aspect, the invention provides nucleic acid molecules isolated from EboBun, EboIC, or fragments thereof. In one embodiment of the present invention, the isolated nucleic acid molecule includes the nucleotide sequence of SEQ ID NOs: 1 or 10, or a complement thereof. In another, the nucleic acid molecule includes a nucleotide sequence having at least 4, 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 350, 400, 450, 500, 550, 600, 650, 700, 750, 800, 850, 900, 950, 1000, 1500, 2000, 2500, 3000, 3500, 4000, 4500, 4600, 4700, 4800, or 4900 contiguous nucleotides of the nucleotide sequence of SEQ ID NO: 1, or a complement thereof; with the proviso that the nucleotide sequence is not comprised by the nucleotide sequence set forth in SEQ ID NO: 20 (Ebola Zaire nucleotide sequence); or at least 5000, 5500, 5600, 5700, 5800, 5900, 6000, 6100, 6200, 6300, 6400, 6500, or 6600 contiguous nucleotides of the nucleotide sequence of SEQ ID NOs: 1 or 10, or a complement thereof. In another embodiment, the isolated nucleic acid molecule includes a nucleotide sequence that encodes the EboBun amino acid sequence of SEQ ID NOs: 2-9 or 59, the EboIC amino acid sequence of SEQ ID NOs: 11-19, or a complement of the nucleotide sequence that encodes the EboBun amino acid sequences of SEQ ID NOs: 2-9 or 59 or the EboIC amino acid sequences of SEQ ID NOs: 11-19. In another, the isolated nucleic acid molecule hybridizes under stringent conditions to a nucleic acid molecule having the nucleotide sequence of SEQ ID NOs: 1 or 10 or a complement thereof, wherein the nucleic acid molecule encodes an amino acid sequence which has a biological activity exhibited by a polypeptide encoded by the nucleotide sequence of SEQ ID NOs: 1 or 10. In another, nucleic acid molecule is RNA. In another, nucleic acid molecule is DNA.

[0062] In another aspect, the invention provides proteins or polypeptides that are isolated from the EboBun, including viral proteins isolated from cells infected with the virus but not present in comparable uninfected cells. In one embodiment of the present invention, the amino acid sequences of the proteins or polypeptides are set forth in SEQ ID NOs: 2-9, 59, or 11-19, or fragments thereof. In one embodiment, polypeptides or proteins of the present invention have a biological activity of the protein (including antigenicity and/or immunogenicity) encoded by the sequence of SEQ ID NOs: 1 or 10. In another, the polypeptides or the proteins of the present invention have a biological activity of at least one protein having the amino acid sequence (including antigenicity and/or immunogenicity) set forth in SEQ ID NOS: 2-9, 59, or 11-19, or a fragment thereof.

[0063] In a related aspect, the invention provides an isolated polypeptide encoded by the nucleic acid molecule of the invention described above. In one embodiment of the present invention, the isolated polypeptide includes the amino acid

sequence selected from the group consisting of: a) an amino acid sequence set forth in SEQ ID NO: 2, 3, 4, 5, 6, 7, 8, or 9; 11, 12, 13, 14, 15, 16, 17, 18 or 19; and b) an amino acid sequence that has 70%, 75%, 80%, 85%, 90%, 95%, 96%, 97%, 98%, or 99% homology to the amino acid sequence according to a). In another, the isolated polypeptide comprises the amino acid sequence having at least 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 210, 220, 230, 240 or 250 contiguous amino acid residues of the amino acid sequence of SEQ ID NOs: 5 or 18 (VP24); 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 210, 220, 230, 240, 250, 260, 270, 280 contiguous amino acid residues of the amino acid sequence of SEQ ID NOs: 6 or 17 (VP30); 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 310, or 320 contiguous amino acid residues of the amino acid sequence of SEQ ID NOs: 8 or 13 (VP40); 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 310, 320, 330, or 340 contiguous amino acid residues of the amino acid sequence of SEQ ID NOs: 7 or 12 (VP35); 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 310, 320, 330, 340, 350, 360, or 370 contiguous amino acid residues of the amino acid sequence of SEQ ID NOs: 4 or 15 (SGP); 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 310, 320, 330, 340, 350, 360, or 370 contiguous amino acid residues of the amino acid sequence of SEQ ID NOs: 59 or 16 (SSGP); 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 350, 400, 450, 450, 500, 550, 600, 610, 620, 630, 640, 650, 660, or 670 contiguous amino acid residues of the amino acid sequence of SEQ ID NOs: 9 or 14 (GP); 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 350, 400, 450, 450, 500, 550, 600, 650, 700, 710, 720, or 730 contiguous amino acid residues of the amino acid sequence of SEQ ID NOs: 3 or 11 (NP); or 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 350, 400, 450, 500, 550, 600, 650, 700, 750, 800, 850, 900, 950, 1000, 1050, 1100, 1150, 1200, 1250, 1300, 1350, 1400, 1450, 1500, 1550, 1600, 1650, 1700, 1750, 1800, 1850, 1900, 1950, 2000, 2050, 2100, 2150, 2160, 2170, 2180, 2190, or 2200 contiguous amino acid residues of the amino acid sequence of SEQ ID NOs: 2 or 19 (L).

[0064] In other aspects, the invention relates to the use of an isolated West African hEbola virus for diagnostic and therapeutic methods. In one embodiment, the invention provides a method of detecting in a biological sample an antibody immunospecific for the hEbola virus using the inventive isolated hEbola virus described herein, or any of the inventive proteins or polypeptides as described herein. In another specific embodiment, the invention provides a method of screening for an antibody which immunospecifically binds and neutralizes hEbola EboBun or EboIC or a combination thereof. Such an antibody is useful for a passive immunization or immunotherapy of a subject infected with hEbola.

[0065] In another aspect, the invention provides an isolated antibody or an antigen-binding fragment thereof which immunospecifically binds to a West African genus hEbola virus of the invention described above, and illustratively including EboBun or EboIC. In one embodiment of the present invention, the isolated antibody or an antigen-binding fragment thereof neutralizes a West African genus hEbola virus. In another, the isolated antibody or an antigen-binding fragment thereof immunospecifically binds to the inventive polypeptide described above. The invention further provides antibodies that specifically bind a polypeptide of the inven-

tion encoded by the nucleotide sequence of SEQ ID NOs: 1 (EboBun) or 10 (EboIC), a fragment thereof, or encoded by a nucleic acid comprising a nucleotide sequence that hybridizes under stringent conditions to the nucleotide sequence of SEQ ID NOs: 1 (EboBun) or 10 (EboIC) and/or any hEbola EboBun epitope, having one or more biological activities of a polypeptide of the invention. These polypeptides include those shown in SEQ ID NOs: 2-9, 59, and 11-19. Such antibodies include, but are not limited to, polyclonal, monoclonal, bi-specific, multi-specific, human, humanized, chimeric antibodies, single chain antibodies, Fab fragments, F(ab')₂ fragments, disulfide-linked Fvs, intrabodies and fragments containing either a VL or VH domain or even a complementary determining region (CDR) that specifically binds to a polypeptide of the invention.

[0066] In other aspects, the invention provides methods for detecting the presence, activity or expression of the hEbola virus of the invention in a biological material, such as cells, blood, saliva, urine, and so forth. The increased or decreased activity or expression of the hEbola virus in a sample relative to a control sample can be determined by contacting the biological material with an agent which can detect directly or indirectly the presence, activity or expression of the hEbola virus. In one embodiment of the present invention, the detecting agents are the antibodies or nucleic acid molecules of the present invention. Antibodies of the invention can also be used to treat hemorrhagic fever.

[0067] In a related aspect, the invention provides a method for detecting the presence of the inventive hEbola virus described above in a biological sample, the method comprising: (a) contacting the sample with an agent that selectively binds to the hEbola virus; and (b) detecting whether the compound binds to the hEbola virus in the sample. In one embodiment of the present invention, the biological sample is selected from the group consisting of cells; blood; serum; plasma; feces; rectal, vaginal and conjunctival swabs. In another, the agent that binds to the virus is an antibody. In another, the agent that binds to the virus is a nucleic acid molecule comprising the nucleotide sequence of SEQ ID NO: 1 or a complement thereof. In another, the agent that binds to the virus is a nucleic acid molecule comprising a nucleotide sequence having at least 4, 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 350, 400, 450, 500, 550, 600, 650, 700, 750, 800, 850, 900, 950, 1000, 1500, 2000, 2500, 3000, 3500, 4000, 4500, 4600, 4700, 4800, 4900, 5000, 5500, 5600, 5700, 5800, 5900, 6000, 6100, 6200, 6300, 6400, 6500, or 6600 contiguous nucleotides of the nucleotide sequence of SEQ ID NOs: 1 or 10, or a complement thereof.

[0068] In another aspect, the invention provides a method for detecting the presence of the inventive polypeptide described above, in a biological sample, the method comprising: (a) contacting the biological sample with an agent that selectively binds to the polypeptide; and (b) detecting whether the agent binds to the polypeptide in the sample. In one embodiment of the present invention, the biological sample is selected from the group consisting of cells; blood; serum; plasma; feces; rectal, vaginal and conjunctival swabs. In another, the agent that binds to the polypeptide is an antibody or an antigen-binding fragment thereof.

[0069] In another aspect, the invention provides a method for detecting the presence of a first nucleic acid molecule derived from the inventive hEbola virus described above in a biological sample, the method includes (a) contacting the biological sample with an agent that selectively binds to the

nucleic acid; and (b) detecting whether the agent binds to the nucleotide in the sample. In one embodiment of the present invention, the agent that binds to the first nucleic acid molecule is a second nucleic acid molecule comprising the nucleotide sequence of SEQ ID NO: 1 or a complement thereof. In another, the second nucleic acid molecule comprises at least 4, 5, 10, 15, 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200, 250, 300, 350, 400, 450, 500, 550, 600, 650, 700, 750, 800, 850, 900, 950, 1000, 1500, 2000, 2500, 3000, 3500, 4000, 4500, 4600, 4700, 4800, 4900, 5000, 5500, 5600, 5700, 5800, 5900, 6000, 6100, 6200, 6300, 6400, 6500, or 6600 contiguous nucleotides of the nucleotide sequence of SEQ ID NOs: 1 or 10, or a complement thereof.

[0070] In another aspect, the invention provides a method for propagating the hEbola virus in host cells comprising infecting the host cells with an inventive isolated West African hEbola virus described above, culturing the host cells to allow the virus to multiply, and harvesting the resulting virions. Also provided by the present invention are host cells infected with the inventive hEbola virus described above. In one embodiment of the present invention, the host cell is a primate cell.

[0071] In another aspect, the invention provides a method of detecting in a biological sample the presence of an antibody that immunospecifically binds hEbola virus, the method includes: (a) contacting the biological sample with the inventive host cell described above; and (b) detecting the antibody bound to the cell.

[0072] In another aspect, the invention provides vaccine preparations, including the inventive hEbola virus, including recombinant and chimeric forms of the virus, nucleic acid molecules comprised by the virus, or protein subunits of the virus. In one embodiment, the vaccine preparations of the present invention includes live but attenuated hEbola virus with or without pharmaceutically acceptable carriers, including adjuvants. In another, the vaccine preparations of the invention comprise an inactivated or killed hEbola Ebola virus, Ebola virus, or a combination thereof, with or without pharmaceutically acceptable carriers, including adjuvants. Such attenuated or inactivated viruses may be prepared by a series of passages of the virus through the host cells or by preparing recombinant or chimeric forms of virus. Accordingly, the present invention further provides methods of preparing recombinant or chimeric forms of the inventive hEbola viruses described herein.

[0073] In another specific embodiment, the invention provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of the inventive hEbola virus described above, and a pharmaceutically acceptable carrier. In another, the invention provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of a protein extract of the inventive hEbola virus described above, or a subunit thereof; and a pharmaceutically acceptable carrier. In another aspect, the invention provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of a nucleic acid molecule comprising the nucleotide sequence of SEQ ID NOs: 1 or 10, or a complement thereof, and a pharmaceutically acceptable carrier. In another, the invention provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of a nucleic acid molecule comprising any of inventive the nucleotide sequences as described above, or a complement thereof, and a pharmaceutically acceptable carrier. In another aspect, the invention provides a vaccine for-

mulation comprising a therapeutically or prophylactically effective amount of a protein extract of the inventive hEbola virus described above, or a subunit thereof; and a pharmaceutically acceptable carrier. In another aspect, the invention provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of a nucleic acid molecule comprising the nucleotide sequence of SEQ ID NOs: 1 or 10, or a complement thereof, and a pharmaceutically acceptable carrier. In another, the invention provides a vaccine formulation comprising a therapeutically or prophylactically effective amount of a nucleic acid molecule comprising any of inventive the nucleotide sequences as described above, or a complement thereof, and a pharmaceutically acceptable carrier.

[0074] In yet another specific embodiment, the vaccine preparations of the present invention comprise a nucleic acid or fragment of the hEbola virus, e.g., the virus having Accession No. 200706291, or nucleic acid molecules having the sequence of SEQ ID NOs: 1 or 10, or a fragment thereof. In another, the vaccine preparations comprise a polypeptide of the invention encoded by the nucleotide sequence of SEQ ID NOs: 1 or 10 or a fragment thereof. In a specific embodiment, the vaccine preparations comprise polypeptides of the invention as shown in SEQ ID NOs: 2-9, 59, or 11-19, or encoded by the nucleotide sequence of SEQ ID NOs: 1 or 10, or a fragment thereof.

[0075] Furthermore, the present invention provides methods for treating, ameliorating, managing or preventing hemorrhagic fever by administering the vaccine preparations or antibodies of the present invention alone or in combination with adjuvants, or other pharmaceutically acceptable excipients. Furthermore, the present invention provides methods for treating, ameliorating, managing, or preventing hemorrhagic fever by administering the inventive compositions and formulations including the vaccine preparations or antibodies of the present invention alone or in combination with antivirals [e.g., amantadine, rimantadine, gancyclovir, acyclovir, ribavirin, penciclovir, oseltamivir, foscarnet zidovudine (AZT), didanosine (ddI), lamivudine (3TC), zalcitabine (ddC), stavudine (d4T), nevirapine, delavirdine, indinavir, ritonavir, vidarabine, nelfinavir, saquinavir, relenza, tamiflu, pleconaril, interferons, etc.], steroids and corticosteroids such as prednisone, cortisone, fluticasone and glucocorticoid, antibiotics, analgesics, bronchodilators, or other treatments for respiratory and/or viral infections.

[0076] In a related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of the inventive hEbola virus described above, and a pharmaceutically acceptable carrier.

[0077] In another related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of a protein extract of the inventive hEbola virus described above or a subunit thereof, and a pharmaceutically acceptable carrier.

[0078] In another related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of a nucleic acid molecule comprising the nucleotide sequence of SEQ ID NOs: 1, 10, a combination thereof, or a complement thereof, and a pharmaceutically acceptable carrier.

[0079] In another related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of a nucleic acid molecule comprising the

inventive nucleotide sequence as described above or a complement thereof, and a pharmaceutically acceptable carrier.

[0080] In another related aspect, the invention provides an immunogenic formulation comprising an immunogenically effective amount of any of the inventive polypeptides described above.

[0081] In another aspect, the present invention provides pharmaceutical compositions comprising antiviral agents of the present invention and a pharmaceutically acceptable carrier. In a specific embodiment, the antiviral agent of the invention is an antibody that immunospecifically binds hEbola virus or any hEbola epitope. In another specific embodiment, the antiviral agent is a polypeptide or protein of the present invention or nucleic acid molecule of the invention.

[0082] In a related aspect, the invention provides a pharmaceutical composition comprising a prophylactically or therapeutically effective amount of an anti-hEbola EboBun agent and a pharmaceutically acceptable carrier. In one embodiment of the present invention, the anti-hEbola EboBun agent is an antibody or an antigen-binding fragment thereof which immunospecifically binds to the hEbola virus of Deposit Accession No. 200706291, or polypeptides or protein derived therefrom. In another, the anti-hEbola agent is a nucleic acid molecule comprising the nucleotide sequence of SEQ ID NOs: 1, 10, a combination thereof, or a fragment thereof. In another, the anti-hEbola agent is a polypeptide encoded by a nucleic acid molecule comprising the nucleotide sequence of SEQ ID NOs: 1, 10, a combination thereof, or a fragment thereof having a biological activity of the polypeptide.

[0083] The invention also provides kits containing compositions and formulations of the present invention. Thus, in another aspect, the invention provides a kit comprising a container containing the inventive immunogenic formulation described above.

[0084] In another aspect, the invention provides a kit includes a container containing the inventive vaccine formulation described above.

[0085] In another aspect, the invention provides a kit including a container containing the inventive pharmaceutical composition described above.

[0086] In another aspect, the invention provides a kit including a container containing the inventive vaccine formulation described above.

[0087] In another aspect, the invention provides a method for identifying a subject infected with the inventive hEbola virus described above, including: (a) obtaining total RNA from a biological sample obtained from the subject; (b) reverse transcribing the total RNA to obtain cDNA; and (c) amplifying the cDNA using a set of primers derived from a nucleotide sequence of the inventive hEbola virus described above.

[0088] In one embodiment of the present invention, the set of primers are derived from the nucleotide sequence of the genome of the hEbola virus of Deposit Accession No. 200706291. In another, the set of primers are derived from the nucleotide sequence of SEQ ID NOs: 1 or 10 or any of the inventive nucleotide sequences as described above, or a complement thereof.

[0089] The invention further relates to the use of the sequence information of the isolated virus for diagnostic and therapeutic methods. In a specific embodiment, the invention provides nucleic acid molecules which are suitable for use as primers consisting of or including the nucleotide sequence of

SEQ ID NOs: 1 or 10, or a complement thereof, or at least a portion of the nucleotide sequence thereof. In another specific embodiment, the invention provides nucleic acid molecules which are suitable for hybridization to the inventive hEbola nucleic acid; including, but not limited to PCR primers, Reverse Transcriptase primers, probes for Southern analysis or other nucleic acid hybridization analysis for the detection of hEbola nucleic acids, e.g., consisting of or including the nucleotide sequence of SEQ ID NOs: 1, 10 a combination thereof, a complement thereof, or a portion thereof. The invention further encompasses chimeric or recombinant viruses encoded in whole or in part by the nucleotide sequences.

[0090] In another aspect, the present invention provides methods for screening antiviral agents that inhibit the infectivity or replication of hEbola virus or variants thereof.

[0091] The invention further provides methods of preparing recombinant or chimeric forms of hEbola.

[0092] In another aspect, the invention provides vaccine preparations including the hEbola virus, including recombinant and chimeric forms of the virus, or subunits of the virus. The present invention encompasses recombinant or chimeric viruses encoded by viral vectors derived from the genome of the inventive hEbola virus described herein or natural variants thereof. In a specific embodiment, a recombinant virus is one derived from the hEbola virus of Deposit Accession No. 200706291. It is recognized that natural variants of the inventive hEbola viruses described herein comprise one or more mutations, including, but not limited to, point mutations, rearrangements, insertions, deletions etc., to the genomic sequence. It is recognized that the mutations may or may not result in a phenotypic change.

[0093] In another specific embodiment, a chimeric virus of the invention is a recombinant hEbola EboBun or EboIC virus which further comprises a heterologous nucleotide sequence. In accordance with the invention, a chimeric virus may be encoded by a nucleotide sequence in which heterologous nucleotide sequences have been added to the genome or in which endogenous or native nucleotide sequences have been replaced with heterologous nucleotide sequences.

[0094] According to the present invention, the chimeric viruses are encoded by the viral vectors of the invention which further comprise a heterologous nucleotide sequence. In accordance with the present invention a chimeric virus is encoded by a viral vector that may or may not include nucleic acids that are non-native to the viral genome. In accordance with the invention a chimeric virus is encoded by a viral vector to which heterologous nucleotide sequences have been added, inserted or substituted for native or non-native sequences. In accordance with the present invention, the chimeric virus may be encoded by nucleotide sequences derived from different species or variants of hEbola virus. In particular, the chimeric virus is encoded by nucleotide sequences that encode antigenic polypeptides derived from different species or variants of hEbola virus.

[0095] A chimeric virus may be of particular use for the generation of recombinant vaccines protecting against two or more viruses (Tao et al., J. Virol. 72, 2955-2961; Durbin et al., 2000, J. Virol. 74, 6821-6831; Skiadopoulos et al., 1998, J. Virol. 72, 1762-1768 (1998); Teng et al., 2000, J. Virol. 74, 9317-9321). For example, it can be envisaged that a virus vector derived from the hEbola virus expressing one or more proteins of variants of hEbola virus including hEbola EboBun, or vice versa, will protect a subject vaccinated with

such vector against infections by both the native hEbola and the variant. Attenuated and replication-defective viruses may be of use for vaccination purposes with live vaccines as has been suggested for other viruses. (See, for example, PCT WO 02/057302, at pp. 6 and 23; and United States Patent Application Publication 2008/0069838 incorporated by reference herein).

[0096] In accordance with the present invention the heterologous sequence to be incorporated into the viral vectors encoding the recombinant or chimeric viruses of the invention include sequences obtained or derived from different species or variants of hEbola.

[0097] In certain embodiments, the chimeric or recombinant viruses of the invention are encoded by viral vectors derived from viral genomes wherein one or more sequences, intergenic regions, termini sequences, or portions or entire ORF have been substituted with a heterologous or non-native sequence. In certain embodiments of the invention, the chimeric viruses of the invention are encoded by viral vectors derived from viral genomes wherein one or more heterologous sequences have been inserted or added to the vector.

[0098] The selection of the viral vector may depend on the species of the subject that is to be treated or protected from a viral infection. If the subject is human, then an attenuated hEbola virus can be used to provide the antigenic sequences.

[0099] In accordance with the present invention, the viral vectors can be engineered to provide antigenic sequences which confer protection against infection by the inventive hEbola and natural variants thereof. The viral vectors may be engineered to provide one, two, three or more antigenic sequences. In accordance with the present invention the antigenic sequences may be derived from the same virus, from different species or variants of the same type of virus, or from different viruses.

[0100] The expression products and/or recombinant or chimeric virions obtained in accordance with the invention may advantageously be utilized in vaccine formulations. The expression products and chimeric virions of the present invention may be engineered to create vaccines against a broad range of pathogens, including viral and bacterial antigens, tumor antigens, allergen antigens, and auto antigens involved in autoimmune disorders. One way to achieve this goal involves modifying existing hEbola genes to contain foreign sequences in their respective external domains. Where the heterologous sequences are epitopes or antigens of pathogens, these chimeric viruses may be used to induce a protective immune response against the disease agent from which these determinants are derived. In particular, the chimeric virions of the present invention may be engineered to create vaccines for the protection of a subject from infections with hEbola virus and variants thereof.

[0101] Thus, the present invention further relates to the use of viral vectors and recombinant or chimeric viruses to formulate vaccines against a broad range of viruses and/or antigens. The present invention also encompasses recombinant viruses including a viral vector derived from the hEbola or variants thereof which contains sequences which result in a virus having a phenotype more suitable for use in vaccine formulations, e.g., attenuated phenotype or enhanced antigenicity. The mutations and modifications can be in coding regions, in intergenic regions and in the leader and trailer sequences of the virus.

[0102] The invention provides a host cell including a nucleic acid or a vector according to the invention. Plasmid or

viral vectors containing the polymerase components of hEbola virus are generated in prokaryotic cells for the expression of the components in relevant cell types (bacteria, insect cells, eukaryotic cells). Plasmid or viral vectors containing full-length or partial copies of the hEbola genome will be generated in prokaryotic cells for the expression of viral nucleic acids in vitro or in vivo. The latter vectors optionally contain other viral sequences for the generation of chimeric viruses or chimeric virus proteins, optionally lack parts of the viral genome for the generation of replication defective virus, and optionally contain mutations, deletions or insertions for the generation of attenuated viruses. In addition, the present invention provides a host cell infected with hEbola virus of Deposit Accession No. 200706291,

[0103] Infectious copies of West African hEbola (being wild type, attenuated, replication-defective or chimeric) are optionally produced upon co-expression of the polymerase components according to the state-of-the-art technologies described above.

[0104] In addition, eukaryotic cells, transiently or stably expressing one or more full-length or partial hEbola proteins are optionally used. Such cells are preferably made by transfection (proteins or nucleic acid vectors), infection (viral vectors) or transduction (viral vectors) and are useful for complementation of mentioned wild type, attenuated, replication-defective or chimeric viruses.

[0105] The viral vectors and chimeric viruses of the present invention optionally modulate a subject's immune system by stimulating a humoral immune response, a cellular immune response or by stimulating tolerance to an antigen. As used herein, a subject means: humans, primates, horses, cows, sheep, pigs, goats, dogs, cats, avian species and rodents.

Formulation of Vaccines and Antivirals

[0106] In a preferred embodiment, the invention provides a proteinaceous molecule or hEbola virus specific viral protein or functional fragment thereof encoded by a nucleic acid according to the invention. Useful proteinaceous molecules are for example derived from any of the genes or genomic fragments derivable from the virus according to the invention, preferably the GP, L, NP, sGP, VP24, VP30, VP35, and VP 40 proteins described herein. Such molecules, or antigenic fragments thereof, as provided herein, are for example useful in diagnostic methods or kits and in pharmaceutical compositions such as subunit vaccines. Particularly useful are polypeptides encoded by the nucleotide sequence of SEQ ID NOs: 1 or 10; or antigenic fragments thereof for inclusion as antigen or subunit immunogen, but inactivated whole virus can also be used. Particularly useful are also those proteinaceous substances that are encoded by recombinant nucleic acid fragments of the hEbola genome, of course preferred are those that are within the preferred bounds and metes of ORFs, in particular, for eliciting hEbola specific antibody or T cell responses, whether in vivo (e.g. for protective or therapeutic purposes or for providing diagnostic antibodies) or in vitro (e.g. by phage display technology or another technique useful for generating synthetic antibodies).

[0107] It is recognized that numerous variants, analogues, or homologues of EboBun polypeptides are within the scope of the present invention including amino acid substitutions, alterations, modifications, or other amino acid changes that increase, decrease, or do not alter the function or immunogenic propensity of the inventive immunogen or vaccine. Several post-translational modifications are similarly envi-

sioned as within the scope of the present invention illustratively including incorporation of a non-naturally occurring amino acid(s), phosphorylation, glycosylation, sulfation, and addition of pendent groups such as biotinylation, fluorophores, lumiphores, radioactive groups, antigens, or other molecules.

[0108] Methods of expressing and purifying natural or recombinant peptides and proteins are well known in the art. Illustratively, peptides and proteins are recombinantly expressed in eukaryotic cells. Exemplary eukaryotic cells include yeast, HeLa cells, 293 cells, COS cells, Chinese hamster ovary cells (CHO), and many other cell types known in the art. Both eukaryotic and prokaryotic expression systems and cells are available illustratively from Invitrogen Corp., Carlsbad, Calif. It is appreciated that cell-free expression systems are similarly operable.

[0109] In a preferred embodiment an immunogenic polypeptide is a full length EboBun protein. Preferably, an immunogen is a full length EboBun protein of SEQ ID NOs: 2-9 or 59, or EboC SEQ ID NOs: 11-19, or a fragment thereof as described herein. Preferably, an immunogen is has a minimum of 5 amino acids. As used herein an immunogen is preferably a polypeptide. In the context of an immunogenic polypeptide the terms immunogen, polypeptide, and antigen are used interchangeably.

[0110] Modifications and changes can be made in the structure of the inventive immunogens that are the subject of the application and still obtain a molecule having similar or improved characteristics as the wild-type sequence (e.g., a conservative amino acid substitution). For example, certain amino acids are optionally substituted for other amino acids in a sequence without appreciable loss of immunogenic activity. Because it is the interactive capacity and nature of a polypeptide that defines that polypeptide's biological functional activity, certain amino acid sequence substitutions can be made in a polypeptide sequence and nevertheless obtain a polypeptide with like or improved properties. Optionally, a polypeptide is used that has less or more immunogenic activity compared to the wild-type sequence.

[0111] In making such changes, the hydropathic index of amino acids is preferably considered. The importance of the hydropathic amino acid index in conferring interactive biologic function on a polypeptide is generally understood in the art. It is known that certain amino acids can be substituted for other amino acids having a similar hydropathic index or score and still result in a polypeptide with similar biological activity. Each amino acid has been assigned a hydropathic index on the basis of its hydrophobicity and charge characteristics. Those indices are: isoleucine (+4.5); valine (+4.2); leucine (+3.8); phenylalanine (+2.8); cysteine/cysteine (+2.5); methionine (+1.9); alanine (+1.8); glycine (−0.4); threonine (−0.7); serine (−0.8); tryptophan (−0.9); tyrosine (−1.3); proline (−1.6); histidine (−3.2); glutamate (−3.5); glutamine (−3.5); aspartate (−3.5); asparagine (−3.5); lysine (−3.9); and arginine (−4.5).

[0112] It is believed that the relative hydropathic character of the amino acid determines the secondary structure of the resultant polypeptide, which in turn defines the interaction of the polypeptide with other molecules, such as enzymes, substrates, receptors, antibodies, antigens, and the like. It is known in the art that an amino acid can be substituted by another amino acid having a similar hydropathic index and still obtain a functionally equivalent immunogen. In such changes, the substitution of amino acids whose hydropathic

indices are within ± 2 is preferred, those within ± 1 are particularly preferred, and those within ± 0.5 are even more particularly preferred.

[0113] As outlined above, amino acid substitutions are generally based on the relative similarity of the amino acid side-chain substituents, for example, their hydrophobicity, hydrophilicity, charge, size, and the like. Exemplary substitutions that take various of the foregoing characteristics into consideration are well known to those of skill in the art and include (original residue: exemplary substitution): (Ala: Gly, Ser), (Arg: Lys), (Asn: Gln, His), (Asp: Glu, Cys, Ser), (Gln: Asn), (Glu: Asp), (Gly: Ala), (His: Asn, Gln), (Ile: Leu, Val), (Leu: Ile, Val), (Lys: Arg), (Met: Leu, Tyr), (Ser: Thr), (Thr: Ser), (Tyr: Trp, Phe), and (Val: Ile, Leu). Embodiments of this disclosure thus contemplate functional or biological equivalents of a polypeptide and immunogen as set forth above. In particular, embodiments of the polypeptides and immunogens optionally include variants having about 50%, 60%, 70%, 80%, 90%, and 95% sequence identity to the polypeptide of interest.

[0114] The invention provides vaccine formulations for the prevention and treatment of infections with hEbola virus. In certain embodiments, the vaccine of the invention comprises recombinant and chimeric viruses of the hEbola virus. In certain embodiments, the virus is attenuated.

[0115] In another embodiment of this aspect of the invention, inactivated vaccine formulations are prepared using conventional techniques to "kill" the chimeric viruses. Inactivated vaccines are "dead" in the sense that their infectivity has been destroyed. Ideally, the infectivity of the virus is destroyed without affecting its immunogenicity. In order to prepare inactivated vaccines, the chimeric virus may be grown in cell culture or in the allantois of the chick embryo, purified by zonal ultracentrifugation, inactivated by formaldehyde or β -propiolactone, and pooled. The resulting vaccine is usually inoculated intramuscularly or intranasally.

[0116] Inactivated viruses are optionally formulated with a suitable adjuvant in order to enhance the immunological response. Such adjuvants illustratively include but are not limited to mineral gels, e.g., aluminum hydroxide; surface active substances such as lysolecithin, pluronic polyols, polyanions; peptides; oil emulsions; and potentially useful human adjuvants such as BCG and *Corynebacterium parvum*.

[0117] In another aspect, the present invention also provides DNA vaccine formulations including a nucleic acid or fragment of the inventive hEbola virus, e.g., the virus having Accession No. 200706291, or nucleic acid molecules having the sequence of SEQ ID NOs: 1 or 10, or a fragment thereof. In another specific embodiment, the DNA vaccine formulations of the present invention comprise a nucleic acid or fragment thereof encoding the antibodies which immunospecifically bind hEbola viruses. In DNA vaccine formulations, a vaccine DNA comprises a viral vector, such as that derived from the hEbola virus, bacterial plasmid, or other expression vector, bearing an insert including a nucleic acid molecule of the present invention operably linked to one or more control elements, thereby allowing expression of the vaccinating proteins encoded by the nucleic acid molecule in a vaccinated subject. Such vectors can be prepared by recombinant DNA technology as recombinant or chimeric viral vectors carrying a nucleic acid molecule of the present invention.

[0118] A nucleic acid as used herein refers to single- or double-stranded molecules which are optionally DNA,

including the nucleotide bases A, T, C and G, or RNA, including the bases A, U (substitutes for T), C, and G. The nucleic acid may represent a coding strand or its complement. Nucleic acids are optionally identical in sequence to the sequence which is naturally occurring or include alternative codons which encode the same amino acid as that which is found in the naturally occurring sequence. Furthermore, nucleic acids optionally include codons which represent conservative substitutions of amino acids as are well known in the art.

[0119] As used herein, the term “isolated nucleic acid” means a nucleic acid separated or substantially free from at least some of the other components of the naturally occurring organism, for example, the cell structural components commonly found associated with nucleic acids in a cellular environment and/or other nucleic acids. The isolation of nucleic acids is illustratively accomplished by techniques such as cell lysis followed by phenol plus chloroform extraction, followed by ethanol precipitation of the nucleic acids. The nucleic acids of this invention are illustratively isolated from cells according to methods well known in the art for isolating nucleic acids. Alternatively, the nucleic acids of the present invention are optionally synthesized according to standard protocols well described in the literature for synthesizing nucleic acids. Modifications to the nucleic acids of the invention are also contemplated, provided that the essential structure and function of the peptide or polypeptide encoded by the nucleic acid are maintained.

[0120] The nucleic acid encoding the peptide or polypeptide of this invention is optionally part of a recombinant nucleic acid construct comprising any combination of restriction sites and/or functional elements as are well known in the art which facilitate molecular cloning and other recombinant DNA manipulations. Thus, the present invention further provides a recombinant nucleic acid construct including a nucleic acid encoding a polypeptide of this invention.

[0121] Generally, it may be more convenient to employ as the recombinant polynucleotide a cDNA version of the polynucleotide. It is believed that the use of a cDNA version will provide advantages in that the size of the gene will generally be much smaller and more readily employed to transfect the targeted cell than will a genomic gene, which will typically be up to an order of magnitude larger than the cDNA gene. However, the inventor does not exclude the possibility of employing a genomic version of a particular gene where desired.

[0122] As used herein, the terms “engineered” and “recombinant” cells are synonymous with “host” cells and are intended to refer to a cell into which an exogenous DNA segment or gene, such as a cDNA or gene has been introduced. Therefore, engineered cells are distinguishable from naturally occurring cells which do not contain a recombinantly introduced exogenous DNA segment or gene. A host cell is optionally a naturally occurring cell that is transformed with an exogenous DNA segment or gene or a cell that is not modified. A host cell preferably does not possess a naturally occurring gene encoding RSV G protein. Engineered cells are, thus, cells having a gene or genes introduced through the hand of man. Recombinant cells illustratively include those having an introduced cDNA or genomic DNA, and also include genes positioned adjacent to a promoter not naturally associated with the particular introduced gene.

[0123] To express a recombinant encoded polypeptide in accordance with the present invention one optionally pre-

pares an expression vector that comprises a polynucleotide under the control of one or more promoters. To bring a coding sequence “under the control of” a promoter, one positions the 5' end of the translational initiation site of the reading frame generally between about 1 and 50 nucleotides “downstream” of (i.e., 3' of) the chosen promoter. The “upstream” promoter stimulates transcription of the inserted DNA and promotes expression of the encoded recombinant protein. This is the meaning of “recombinant expression” in the context used here.

[0124] Many standard techniques are available to construct expression vectors containing the appropriate nucleic acids and transcriptional/translational control sequences in order to achieve protein or peptide expression in a variety of host-expression systems. Cell types available for expression include, but are not limited to, bacteria, such as *E. coli* and *B. subtilis* transformed with recombinant phage DNA, plasmid DNA or cosmid DNA expression vectors.

[0125] Certain examples of prokaryotic hosts illustratively include *E. coli* strain RR1, *E. coli* LE392, *E. coli* B, *E. coli* 1776 (ATCC No. 31537) as well as *E. coli* W3110 (F-, lambda-, prototrophic, ATCC No. 27325); bacilli such as *Bacillus subtilis*; and other enterobacteria such as *Salmonella typhimurium*, *Serratia marcescens*, and various *Pseudomonas* species.

[0126] In general, plasmid vectors containing replicon and control sequences that are derived from species compatible with the host cell are used in connection with these hosts. The vector ordinarily carries a replication site, as well as marking sequences that are capable of providing phenotypic selection in transformed cells. For example, *E. coli* is often transformed using pBR322, a plasmid derived from an *E. coli* species. Plasmid pBR322 contains genes for ampicillin and tetracycline resistance and thus provides easy means for identifying transformed cells. The pBR322 plasmid, or other microbial plasmid or phage may also contain, or be modified to contain, promoters that can be used by the microbial organism for expression of its own proteins.

[0127] In addition, phage vectors containing replicon and control sequences that are compatible with the host microorganism are optionally used as transforming vectors in connection with these hosts. For example, the phage lambda is optionally utilized in making a recombinant phage vector that can be used to transform host cells, such as *E. coli* LE392.

[0128] Further useful vectors include pIN vectors and pGEX vectors, for use in generating glutathione S-transferase (GST) soluble fusion proteins for later purification and separation or cleavage. Other suitable fusion proteins are those with β -galactosidase, ubiquitin, or the like.

[0129] Promoters that are most commonly used in recombinant DNA construction include the β -lactamase (penicillinase), lactose and tryptophan (trp) promoter systems. While these are the most commonly used, other microbial promoters have been discovered and utilized, and details concerning their nucleotide sequences have been published, enabling those of skill in the art to ligate them functionally with plasmid vectors.

[0130] For expression in *Saccharomyces*, the plasmid YRp7, for example, is commonly used. This plasmid contains the trp1 gene, which provides a selection marker for a mutant strain of yeast lacking the ability to grow in tryptophan, for example ATCC No. 44076 or PEP4-1. The presence of the trp1 lesion as a characteristic of the yeast host cell genome

then provides an effective environment for detecting transformation by growth in the absence of tryptophan.

[0131] Suitable promoting sequences in yeast vectors illustratively include the promoters for 3-phosphoglycerate kinase or other glycolytic enzymes, such as enolase, glyceraldehyde-3-phosphate dehydrogenase, hexokinase, pyruvate decarboxylase, phosphofructokinase, glucose-6-phosphate isomerase, 3-phosphoglycerate mutase, pyruvate kinase, triosephosphate isomerase, phosphoglucose isomerase, and glucokinase. In constructing suitable expression plasmids, the termination sequences associated with these genes are also preferably ligated into the expression vector 3' of the sequence desired to be expressed to provide polyadenylation of the mRNA and termination.

[0132] Other suitable promoters, which have the additional advantage of transcription controlled by growth conditions, illustratively include the promoter region for alcohol dehydrogenase 2, isocytichrome C, acid phosphatase, degradative enzymes associated with nitrogen metabolism, and the aforementioned glyceraldehyde-3-phosphate dehydrogenase, and enzymes responsible for maltose and galactose utilization.

[0133] In addition to microorganisms, cultures of cells derived from multicellular organisms are also operable as hosts. In principle, any such cell culture is operable, whether from vertebrate or invertebrate culture. In addition to mammalian cells, these include insect cell systems infected with recombinant virus expression vectors (e.g., baculovirus); and plant cell systems infected with recombinant virus expression vectors (e.g., cauliflower mosaic virus, CaMV; tobacco mosaic virus, TMV) or transformed with recombinant plasmid expression vectors (e.g., Ti plasmid) containing one or more coding sequences.

[0134] In a useful insect system, *Autographica californica* nuclear polyhedrosis virus (AcNPV) is used as a vector to express foreign genes. The virus grows in *Spodoptera frugiperda* cells. The isolated nucleic acid coding sequences are cloned into non-essential regions (for example the polyhedron gene) of the virus and placed under control of an AcNPV promoter (for example, the polyhedron promoter). Successful insertion of the coding sequences results in the inactivation of the polyhedron gene and production of non-occluded recombinant virus (i.e., virus lacking the proteinaceous coat coded for by the polyhedron gene). These recombinant viruses are then used to infect *Spodoptera frugiperda* cells in which the inserted gene is expressed (e.g., U.S. Pat. No. 4,215,051).

[0135] Examples of useful mammalian host cell lines include VERO and HeLa cells, Chinese hamster ovary (CHO) cell lines, W138, BHK, COS-7, 293, HepG2, NIH3T3, RIN and MDCK cell lines. In addition, a host cell is preferably chosen that modulates the expression of the inserted sequences, or modifies and processes the gene product in the specific fashion desired. Such modifications (e.g., glycosylation) and processing (e.g., cleavage) of protein products may be important for the function of the encoded protein.

[0136] Different host cells have characteristic and specific mechanisms for the post-translational processing and modification of proteins. Appropriate cell lines or host systems are preferably chosen to ensure the correct modification and processing of the foreign protein expressed. Expression vectors for use in mammalian cells ordinarily include an origin of replication (as necessary), a promoter located in front of the gene to be expressed, along with any necessary ribosome binding sites, RNA splice sites, polyadenylation site, and transcriptional terminator sequences. The origin of replica-

tion is preferably provided either by construction of the vector to include an exogenous origin, such as may be derived from SV40 or other viral (e.g., Polyoma, Adeno, VSV, BPV) source, or may be provided by the host cell chromosomal replication mechanism. If the vector is integrated into the host cell chromosome, the latter is often sufficient.

[0137] The promoters are optionally derived from the genome of mammalian cells (e.g., metallothionein promoter) or from mammalian viruses (e.g., the adenovirus late promoter; the vaccinia virus 7.5K promoter). Further, it is also possible, and may be desirable, to utilize promoter or control sequences normally associated with the desired gene sequence, provided such control sequences are compatible with the host cell systems.

[0138] A number of viral based expression systems are operable herein, for example, commonly used promoters are derived from polyoma, Adenovirus 2, Adenovirus 5, cytomegalovirus and Simian Virus 40 (SV40). The early and late promoters of SV40 virus are useful because both are obtained easily from the virus as a fragment which also contains the SV40 viral origin of replication. Smaller or larger SV40 fragments are also operable, particularly when there is included the approximately 250 bp sequence extending from the HindIII site toward the BglII site located in the viral origin of replication.

[0139] In cases where an adenovirus is used as an expression vector, the coding sequences are preferably ligated to an adenovirus transcription/translation control complex, e.g., the late promoter and tripartite leader sequence. This chimeric gene is then optionally inserted in the adenovirus genome by in vitro or in vivo recombination. Insertion in a non-essential region of the viral genome (e.g., region E1 or E3) will result in a recombinant virus that is viable and capable of expressing proteins in infected hosts.

[0140] Specific initiation signals may also be required for efficient translation of the claimed isolated nucleic acid coding sequences. These signals include the ATG initiation codon and adjacent sequences. Exogenous translational control signals, including the ATG initiation codon, may additionally need to be provided. One of ordinary skill in the art would readily be capable of determining this need and providing the necessary signals. It is well known that the initiation codon must be in-frame (or in-phase) with the reading frame of the desired coding sequence to ensure translation of the entire insert. These exogenous translational control signals and initiation codons are optionally of a variety of origins, both natural and synthetic. The efficiency of expression is optionally enhanced by the inclusion of appropriate transcription enhancer elements or transcription terminators.

[0141] In eukaryotic expression, one will also typically desire to incorporate into the transcriptional unit an appropriate polyadenylation site if one was not contained within the original cloned segment. Typically, the poly A addition site is placed about 30 to 2000 nucleotides "downstream" of the termination site of the protein at a position prior to transcription termination.

[0142] For long-term, high-yield production of recombinant proteins, stable expression is preferred. For example, cell lines that stably express constructs encoding proteins are engineered. Rather than using expression vectors that contain viral origins of replication, host cells are preferably transformed with vectors controlled by appropriate expression control elements (e.g., promoter, enhancer, sequences, transcription terminators, polyadenylation sites, etc.), and a

selectable marker. Following the introduction of foreign DNA, engineered cells may be allowed to grow for 1-2 days in an enriched medium, and then are switched to a selective medium. The selectable marker in the recombinant plasmid confers resistance to the selection and allows cells to stably integrate the plasmid into their chromosomes and grow to form foci, which in turn can be cloned and expanded into cell lines.

[0143] A number of selection systems are illustratively used, including, but not limited to, the herpes simplex virus thymidine kinase, hypoxanthine-guanine phosphoribosyltransferase and adenine phosphoribosyltransferase genes, in tk⁻, hgp⁺ or ap⁺ cells, respectively. Also, antimetabolite resistance is optionally used as the basis of selection for dhfr, which confers resistance to methotrexate; gpt, which confers resistance to mycophenolic acid; neo, which confers resistance to the aminoglycoside G-418; and hyg⁺, which confers resistance to hygromycin. It is appreciated that numerous other selection systems are known in the art that are similarly operable in the present invention.

[0144] The nucleic acids encoding the peptides and polypeptides of this invention are optionally administered as nucleic acid vaccines. For the purposes of vaccine delivery, a nucleic acid encoding a peptide or polypeptide of this invention is preferably in an expression vector that includes viral nucleic acid including, but not limited to, vaccinia virus, adenovirus, retrovirus and/or adeno-associated virus nucleic acid. The nucleic acid or vector of this invention is optionally in a liposome or a delivery vehicle which can be taken up by a cell via receptor-mediated or other type of endocytosis. The nucleic acid vaccines of this invention are preferably in a pharmaceutically acceptable carrier or administered with an adjuvant. The nucleic acids encoding the peptides and polypeptides of this invention can also be administered to cells in vivo or ex vivo.

[0145] It is contemplated that the isolated nucleic acids of the disclosure are optionally "overexpressed", i.e., expressed in increased levels relative to its natural expression in cells of its indigenous organism, or even relative to the expression of other proteins in the recombinant host cell. Such overexpression is assessed by a variety of methods illustratively including radio-labeling and/or protein purification. However, simple and direct methods are preferred, for example, those involving SDS/PAGE and protein staining or immunoblotting, followed by quantitative analyses, such as densitometric scanning of the resultant gel or blot. A specific increase in the level of the recombinant protein or peptide in comparison to the level in natural in transfected cells is indicative of overexpression, as is a relative abundance of the specific protein in relation to the other proteins produced by the host cell and, e.g., visible on a gel.

[0146] Various heterologous vectors are described for DNA vaccinations against viral infections. For example, the vectors described in the following references, incorporated herein by reference, may be used to express hEbola sequences instead of the sequences of the viruses or other pathogens described; in particular, vectors described for hepatitis B virus (Michel, M. L. et al., 1995, DAN-mediated immunization to the hepatitis B surface antigen in mice: Aspects of the humoral response mimic hepatitis B viral infection in humans, *Proc. Natl. Acad. Sci. USA* 92:5307-5311; Davis, H. L. et al., 1993, DNA-based immunization induces continuous secretion of hepatitis B surface antigen and high levels of circulating antibody, *Human Molec. Genetics* 2:1847-1851),

HIV virus (Wang, B. et al., 1993, Gene inoculation generates immune responses against human immunodeficiency virus type 1, *Proc. Natl. Acad. Sci. USA* 90:4156-4160; Lu, S. et al., 1996, Simian immunodeficiency virus DNA vaccine trial in Macaques, *J. Virol.* 70:3978-3991; Letvin, N. L. et al., 1997, Potent, protective anti-HIV immune responses generated by bimodal HIV envelope DNA plus protein vaccination, *Proc Natl Acad Sci USA*. 94(17):9378-83), and influenza viruses (Robinson, H. L. et al., 1993, Protection against a lethal influenza virus challenge by immunization with a haemagglutinin-expressing plasmid DNA, *Vaccine* 11:957-960; Ulmer, J. B. et al., Heterologous protection against influenza by injection of DNA encoding a viral protein, *Science* 259:1745-1749), as well as bacterial infections, such as tuberculosis (Tascon, R. E. et al., 1996, Vaccination against tuberculosis by DNA injection, *Nature Med.* 2:888-892; Huygen, K. et al., 1996, Immunogenicity and protective efficacy of a tuberculosis DNA vaccine, *Nature Med.*, 2:893-898), and parasitic infection, such as malaria (Sedegah, M., 1994, Protection against malaria by immunization with plasmid DNA encoding circumsporozoite protein, *Proc. Natl. Acad. Sci. USA* 91:9866-9870; Doolan, D. L. et al., 1996, Circumventing genetic restriction of protection against malaria with multi-gene DNA immunization: CD8⁺T cell-interferon .delta., and nitric oxide-dependent immunity, *J. Exper. Med.*, 1183:1739-1746).

[0147] Many methods are optionally used to introduce the vaccine formulations described above. These include, but are not limited to, oral, intradermal, intramuscular, intraperitoneal, intravenous, subcutaneous, and intranasal routes. Alternatively, in a preferred embodiment the chimeric virus vaccine formulation is introduced via the natural route of infection of the pathogen for which the vaccine is designed. The DNA vaccines of the present invention are optionally administered in saline solutions by injections into muscle or skin using a syringe and needle (Wolff J. A. et al., 1990, Direct gene transfer into mouse muscle in vivo, *Science* 247:1465-1468; Raz, E., 1994, Intradermal gene immunization: The possible role of DNA uptake in the induction of cellular immunity to viruses, *c. Natl. Acad. Sci. USA* 91:9519-9523). Another way to administer DNA vaccines operable herein is called the "gene gun" method, whereby microscopic gold beads coated with the DNA molecules of interest is fired into cells (Tang, D. et al., 1992, Genetic immunization is a simple method for eliciting an immune response, *Nature* 356:152-154). For general reviews of the methods for DNA vaccines, see Robinson, H. L., 1999, DNA vaccines: basic mechanism and immune responses (Review), *Int. J. Mol. Med.* 4(5):549-555; Barber, B., 1997, Introduction: Emerging vaccine strategies, *Seminars in Immunology* 9(5):269-270; and Robinson, H. L. et al., 1997, DNA vaccines, *Seminars in Immunology* 9(5):271-283.

Attenuation of hEbola Virus or Variants Thereof

[0148] The hEbola virus or variants thereof of the invention are optionally genetically engineered to exhibit an attenuated phenotype. In particular, the viruses of the invention exhibit an attenuated phenotype in a subject to which the virus is administered as a vaccine. Attenuation can be achieved by any method known to a skilled artisan. Without being bound by theory, the attenuated phenotype of the viruses of the invention is caused, e.g., by using a virus that naturally does not replicate well in an intended host species, for example, by reduced replication of the viral genome, by reduced ability of the virus to infect a host cell, or by reduced ability of the viral

proteins to assemble to an infectious viral particle relative to the wild type species of the virus.

[0149] The attenuated phenotypes of hEbola virus or variants thereof are optionally tested by any method known to the artisan. A candidate virus, for example, is optionally tested for its ability to infect a host or for the rate of replication in a cell culture system. In certain embodiments, growth curves at different temperatures are used to test the attenuated phenotype of the virus. For example, an attenuated virus is able to grow at 35° C., but not at 39° C. or 40° C. In certain embodiments, different cell lines are used to evaluate the attenuated phenotype of the virus. For example, an attenuated virus may only be able to grow in monkey cell lines but not the human cell lines, or the achievable virus titers in different cell lines are different for the attenuated virus. In certain embodiments, viral replication in the respiratory tract of a small animal model, including but not limited to, hamsters, cotton rats, mice and guinea pigs, is used to evaluate the attenuated phenotypes of the virus. In other embodiments, the immune response induced by the virus, including but not limited to, the antibody titers (e.g., assayed by plaque reduction neutralization assay or ELISA) is used to evaluate the attenuated phenotypes of the virus. In a specific embodiment, the plaque reduction neutralization assay or ELISA is carried out at a low dose. In certain embodiments, the ability of the hEbola virus to elicit pathological symptoms in an animal model is tested. A reduced ability of the virus to elicit pathological symptoms in an animal model system is indicative of its attenuated phenotype. In a specific embodiment, the candidate viruses are tested in a monkey model for nasal infection, indicated by mucus production.

[0150] The viruses of the invention are optionally attenuated such that one or more of the functional characteristics of the virus are impaired. In certain embodiments, attenuation is measured in comparison to the wild type species of the virus from which the attenuated virus is derived. In other embodiments, attenuation is determined by comparing the growth of an attenuated virus in different host systems. Thus, for a non-limiting example, hEbola virus or a variant thereof is attenuated when grown in a human host if the growth of the hEbola or variant thereof in the human host is reduced compared to the non-attenuated hEbola or variant thereof.

[0151] In certain embodiments, the attenuated virus of the invention is capable of infecting a host, is capable of replicating in a host such that infectious viral particles are produced. In comparison to the wild type species, however, the attenuated species grows to lower titers or grows more slowly. Any technique known to the skilled artisan can be used to determine the growth curve of the attenuated virus and compare it to the growth curve of the wild type virus.

[0152] In certain embodiments, the attenuated virus of the invention (e.g., a recombinant or chimeric hEbola) cannot replicate in human cells as well as the wild type virus (e.g., wild type hEbola) does. However, the attenuated virus can replicate well in a cell line that lacks interferon functions, such as Vero cells.

[0153] In other embodiments, the attenuated virus of the invention is capable of infecting a host, of replicating in the host, and of causing proteins of the virus of the invention to be inserted into the cytoplasmic membrane, but the attenuated virus does not cause the host to produce new infectious viral particles. In certain embodiments, the attenuated virus infects the host, replicates in the host, and causes viral proteins to be inserted in the cytoplasmic membrane of the host with the

same efficiency as the wild type hEbola. In other embodiments, the ability of the attenuated virus to cause viral proteins to be inserted into the cytoplasmic membrane into the host cell is reduced compared to the wild type virus. In certain embodiments, the ability of the attenuated hEbola virus to replicate in the host is reduced compared to the wild type virus. Any technique known to the skilled artisan can be used to determine whether a virus is capable of infecting a mammalian cell, of replicating within the host, and of causing viral proteins to be inserted into the cytoplasmic membrane of the host.

[0154] In certain embodiments, the attenuated virus of the invention is capable of infecting a host. In contrast to the wild type hEbola, however, the attenuated hEbola cannot be replicated in the host. In a specific embodiment, the attenuated hEbola virus can infect a host and can cause the host to insert viral proteins in its cytoplasmic membranes, but the attenuated virus is incapable of being replicated in the host. Any method known to the skilled artisan can be used to test whether the attenuated hEbola has infected the host and has caused the host to insert viral proteins in its cytoplasmic membranes.

[0155] In certain embodiments, the ability of the attenuated virus to infect a host is reduced compared to the ability of the wild type virus to infect the same host. Any technique known to the skilled artisan can be used to determine whether a virus is capable of infecting a host.

[0156] In certain embodiments, mutations (e.g., missense mutations) are introduced into the genome of the virus, for example, into the sequence of SEQ ID NOs: 1 or 10, or to generate a virus with an attenuated phenotype. Mutations (e.g., missense mutations) can be introduced into the structural genes and/or regulatory genes of the hEbola. Mutations are optionally additions, substitutions, deletions, or combinations thereof. Such variant of hEbola can be screened for a predicted functionality, such as infectivity, replication ability, protein synthesis ability, assembling ability, as well as cytopathic effect in cell cultures. In a specific embodiment, the missense mutation is a cold-sensitive mutation. In another embodiment, the missense mutation is a heat-sensitive mutation. In another embodiment, the missense mutation prevents a normal processing or cleavage of the viral proteins.

[0157] In other embodiments, deletions are introduced into the genome of the hEbola virus, which result in the attenuation of the virus.

[0158] In certain embodiments, attenuation of the virus is achieved by replacing a gene of the wild type virus with a gene of a virus of a different species, of a different subgroup, or of a different variant. In another aspect, attenuation of the virus is achieved by replacing one or more specific domains of a protein of the wild type virus with domains derived from the corresponding protein of a virus of a different species. In certain other embodiments, attenuation of the virus is achieved by deleting one or more specific domains of a protein of the wild type virus.

[0159] When a live attenuated vaccine is used, its safety should also be considered. The vaccine preferably does not cause disease. Any techniques known in the art for improving vaccine safety are operable in the present invention. In addition to attenuation techniques, other techniques are optionally be used. One non-limiting example is to use a soluble heterologous gene that cannot be incorporated into the virion membrane. For example, a single copy of the soluble version

of a viral transmembrane protein lacking the transmembrane and cytosolic domains thereof is used.

[0160] Various assays are optionally used to test the safety of a vaccine. For example, sucrose gradients and neutralization assays are used to test the safety. A sucrose gradient assay is optionally used to determine whether a heterologous protein is inserted in a virion. If the heterologous protein is inserted in the virion, the virion is preferably tested for its ability to cause symptoms in an appropriate animal model since the virus may have acquired new, possibly pathological, properties.

5.4 Adjuvants and Carrier Molecules

[0161] hEbola-associated antigens are administered with one or more adjuvants. In one embodiment, the hEbola-associated antigen is administered together with a mineral salt adjuvants or mineral salt gel adjuvant. Such mineral salt and mineral salt gel adjuvants include, but are not limited to, aluminum hydroxide (ALHYDROGEL, REHYDRAGEL), aluminum phosphate gel, aluminum hydroxyphosphate (ADJU-PHOS), and calcium phosphate.

[0162] In another embodiment, hEbola-associated antigen is administered with an immunostimulatory adjuvant. Such class of adjuvants include, but are not limited to, cytokines (e.g., interleukin-2, interleukin-7, interleukin-12, granulocyte-macrophage colony stimulating factor (GM-CSF), interferon- γ interleukin-1 β (IL-1 β), and IL-1 β peptide or Sclavo Peptide), cytokine-containing liposomes, triterpenoid glycosides or saponins (e.g., QuilA and QS-21, also sold under the trademark STIMULON, ISCOPREP), Muramyl Dipeptide (MDP) derivatives, such as N-acetyl-muramyl-L-threonyl-D-isoglutamine (Threonyl-MDP, sold under the trademark TERMURTIDE), GMDP, N-acetyl-nor-muramyl-L-alanyl-D-isoglutamine, N-acetylmuramyl-L-alanyl-D-isoglutaminyl-L-alanine-2-(1'-2'-dipalmitoyl-s-n-glycero-3-hydroxyphosphoryloxy)-ethylamine, muramyl tripeptide phosphatidylethanolamine (MTP-PE), unmethylated CpG dinucleotides and oligonucleotides, such as bacterial DNA and fragments thereof, LPS, monophosphoryl Lipid A (3D-MLA sold under the trademark MPL), and polyphosphazenes.

[0163] In another embodiment, the adjuvant used is a particular adjuvant, including, but not limited to, emulsions, e.g., Freund's Complete Adjuvant, Freund's Incomplete Adjuvant, squalene or squalane oil-in-water adjuvant formulations, such as SAF and MF59, e.g., prepared with block-copolymers, such as L-121 (polyoxypropylene/polyoxyethylene) sold under the trademark PLURONIC L-121, Liposomes, Virosomes, cochleates, and immune stimulating complex, which is sold under the trademark ISCOM.

[0164] In another embodiment, a microparticulate adjuvant is used. Microparticulate adjuvants include, but are not limited to, biodegradable and biocompatible polyesters, homo- and copolymers of lactic acid (PLA) and glycolic acid (PGA), poly(lactide-co-glycolides) (PLGA) microparticles, polymers that self-associate into particulates (poloxamer particles), soluble polymers (polyphosphazenes), and virus-like particles (VLPs) such as recombinant protein particulates, e.g., hepatitis B surface antigen (HbsAg).

[0165] Yet another class of adjuvants that are optionally used include mucosal adjuvants, including but not limited to heat-labile enterotoxin from *Escherichia coli* (LT), cholera holotoxin (CT) and cholera Toxin B Subunit (CTB) from *Vibrio cholerae*, mutant toxins (e.g., LTK63 and LTR72), microparticles, and polymerized liposomes.

[0166] In other embodiments, any of the above classes of adjuvants are optionally used in combination with each other or with other adjuvants. For example, non-limiting examples of combination adjuvant preparations used to administer the hEbola-associated antigens of the invention include liposomes containing immunostimulatory protein, cytokines, T-cell and/or B-cell peptides, or microbes with or without entrapped IL-2 or microparticles containing enterotoxin. Other adjuvants known in the art are also included within the scope of the invention (see Vaccine Design: The Subunit and Adjuvant Approach, Chap. 7, Michael F. Powell and Mark J. Newman (eds.), Plenum Press, New York, 1995, which is incorporated herein in its entirety).

[0167] The effectiveness of an adjuvant is illustratively determined by measuring the induction of antibodies directed against an immunogenic polypeptide containing a hEbola polypeptide epitope, the antibodies resulting from administration of this polypeptide in vaccines which are also comprised of the various adjuvants.

[0168] The polypeptides are optionally formulated into the vaccine as neutral or salt forms. Pharmaceutically acceptable salts include the acid additional salts (formed with free amino groups of the peptide) and which are formed with inorganic acids, such as, for example, hydrochloric or phosphoric acids, or organic acids such as acetic, oxalic, tartaric, maleic, and the like. Salts formed with free carboxyl groups are optionally derived from inorganic bases, such as, for example, sodium potassium, ammonium, calcium, or ferric hydroxides, and such organic bases as isopropylamine, trimethylamine, 2-ethylamino ethanol, histidine, procaine and the like.

[0169] The vaccines of the invention are preferably multivalent or univalent. Multivalent vaccines are made from recombinant viruses that direct the expression of more than one antigen.

[0170] Many methods are operable herein to introduce the vaccine formulations of the invention; these include but are not limited to oral, intradermal, intramuscular, intraperitoneal, intravenous, subcutaneous, intranasal routes, and via scarification (scratching through the top layers of skin, e.g., using a bifurcated needle).

[0171] The patient to which the vaccine is administered is preferably a mammal, most preferably a human, but is also optionally a non-human animal including but not limited to lower primates, cows, horses, sheep, pigs, fowl (e.g., chickens), goats, cats, dogs, hamsters, mice and rats.

Preparation of Antibodies

[0172] Antibodies that specifically recognize a polypeptide of the invention, such as, but not limited to, polypeptides including the sequence of SEQ ID NOs: 2-9, 59, or 11-19 and other polypeptides as described herein, or hEbola epitope or antigen-binding fragments thereof are used in a preferred embodiment for detecting, screening, and isolating the polypeptide of the invention or fragments thereof, or similar sequences that might encode similar enzymes from the other organisms. For example, in one specific embodiment, an antibody which immunospecifically binds hEbola epitope, or a fragment thereof, is used for various in vitro detection assays, including enzyme-linked immunosorbent assays (ELISA), radioimmunoassays, western blot, etc., for the detection of a polypeptide of the invention or, preferably, hEbola, in samples, for example, a biological material, including cells, cell culture media (e.g., bacterial cell culture media, mammalian cell culture media, insect cell culture media, yeast cell

culture media, etc.), blood, plasma, serum, tissues, sputum, nasopharyngeal aspirates, etc.

[0173] Antibodies specific for a polypeptide of the invention or any epitope of hEbola are optionally generated by any suitable method known in the art. Polyclonal antibodies to an antigen of interest, for example, the hEbola virus from Deposit Accession No. 200706291, or including a nucleotide sequence of SEQ ID NOs: 1 or 10, are optionally produced by various procedures well known in the art. For example, an antigen is optionally administered to various host animals including, but not limited to, rabbits, mice, rats, etc., to induce the production of antisera containing polyclonal antibodies specific for the antigen. Various adjuvants are optionally used to increase the immunological response, depending on the host species, and include but are not limited to, Freund's (complete and incomplete) adjuvant, mineral gels such as aluminum hydroxide, surface active substances such as lysolecithin, pluronic polyols, polyanions, peptides, oil emulsions, keyhole limpet hemocyanins, dinitrophenol, and potentially useful adjuvants for humans such as BCG (*Bacille Calmette-Guerin*) and *Corynebacterium parvum*. Such adjuvants are also well known in the art.

[0174] Monoclonal antibodies are optionally prepared using a wide variety of techniques known in the art including the use of hybridoma, recombinant, and phage display technologies, or a combination thereof. In one example, monoclonal antibodies are produced using hybridoma techniques including those known in the art and taught, for example, in Harlow et al., *Antibodies: A Laboratory Manual* (Cold Spring Harbor Laboratory Press, 2nd ed. 1988); Hammerling, et al., in: *Monoclonal Antibodies and T-Cell Hybridomas*, pp. 563-681 (Elsevier, N.Y., 1981) (both of which are incorporated by reference in their entireties). The term "monoclonal antibody" as used herein is not limited to antibodies produced through hybridoma technology. The term "monoclonal antibody" refers to an antibody that is derived from a single clone, including any eukaryotic, prokaryotic, or phage clone, and not the method by which it is produced.

[0175] Methods for producing and screening for specific antibodies using hybridoma technology are routine and well known in the art. In a non-limiting example, mice are immunized with an antigen of interest or a cell expressing such an antigen. Once an immune response is detected, e.g., antibodies specific for the antigen are detected in the mouse serum, the mouse spleen is harvested and splenocytes isolated. The splenocytes are then fused by well known techniques to any suitable myeloma cells. Hybridomas are selected and cloned by limiting dilution. The hybridoma clones are then assayed by methods known in the art for cells that secrete antibodies capable of binding the antigen. Ascites fluid, which generally contains high levels of antibodies, is optionally generated by inoculating mice intraperitoneally with positive hybridoma clones.

[0176] Antibody fragments which recognize specific epitopes are optionally generated by known techniques. For example, Fab and F(ab')₂ fragments are illustratively produced by proteolytic cleavage of immunoglobulin molecules, using enzymes such as papain (to produce Fab fragments) or pepsin (to produce F(ab')₂ fragments). F(ab')₂ fragments preferably contain the complete light chain, and the variable region, the CH1 region and the hinge region of the heavy chain.

[0177] The antibodies of the invention or fragments thereof are optionally produced by any method known in the art for

the synthesis of antibodies, in particular, by chemical synthesis or preferably, by recombinant expression techniques.

[0178] The nucleotide sequence encoding an antibody is obtained from any information available to those skilled in the art (i.e., from Genbank, the literature, or by routine cloning and sequence analysis). If a clone containing a nucleic acid encoding a particular antibody or an epitope-binding fragment thereof is not available, but the sequence of the antibody molecule or epitope-binding fragment thereof is known, a nucleic acid encoding the immunoglobulin may be chemically synthesized or obtained from a suitable source (e.g., an antibody cDNA library, or a cDNA library generated from, or nucleic acid, preferably poly A+RNA, isolated from any tissue or cells expressing the antibody, such as hybridoma cells selected to express an antibody) by PCR amplification using synthetic primers hybridizable to the 3' and 5' ends of the sequence or by cloning using an oligonucleotide probe specific for the particular gene sequence to identify, e.g., a cDNA clone from a cDNA library that encodes the antibody. Amplified nucleic acids generated by PCR are optionally then cloned into replicable cloning vectors using any method known in the art.

[0179] Once the nucleotide sequence of the antibody is determined, the nucleotide sequence of the antibody is optionally manipulated using methods well known in the art for the manipulation of nucleotide sequences, e.g., recombinant DNA techniques, site directed mutagenesis, PCR, etc. (see, for example, the techniques described in Sambrook et al., supra; and Ausubel et al., eds., 1998, *Current Protocols in Molecular Biology*, John Wiley & Sons, NY, which are both incorporated by reference herein in their entireties), to generate antibodies having a different amino acid sequence by, for example, introducing amino acid substitutions, deletions, and/or insertions into the epitope-binding domain regions of the antibodies or any portion of antibodies which may enhance or reduce biological activities of the antibodies.

[0180] Recombinant expression of an antibody requires construction of an expression vector containing a nucleotide sequence that encodes the antibody. Once a nucleotide sequence encoding an antibody molecule or a heavy or light chain of an antibody, or portion thereof has been obtained, the vector for the production of the antibody molecule is optionally produced by recombinant DNA technology using techniques known in the art as discussed in the previous sections. Methods which are known to those skilled in the art are optionally used to construct expression vectors containing antibody coding sequences and appropriate transcriptional and translational control signals. These methods include, for example, in vitro recombinant DNA techniques, synthetic techniques, and in vivo genetic recombination. The nucleotide sequence encoding the heavy-chain variable region, light-chain variable region, both the heavy-chain and light-chain variable regions, an epitope-binding fragment of the heavy- and/or light-chain variable region, or one or more complementarity determining regions (CDRs) of an antibody are optionally cloned into such a vector for expression. Thus, prepared expression vector is optionally then introduced into appropriate host cells for the expression of the antibody. Accordingly, the invention includes host cells containing a polynucleotide encoding an antibody specific for the polypeptides of the invention or fragments thereof.

[0181] The host cell is optionally co-transfected with two expression vectors of the invention, the first vector encoding a heavy chain derived polypeptide and the second vector

encoding a light chain derived polypeptide. The two vectors illustratively contain identical selectable markers which enable equal expression of heavy and light chain polypeptides or different selectable markers to ensure maintenance of both plasmids. Alternatively, a single vector is optionally used which encodes, and is capable of expressing, both heavy and light chain polypeptides. In such situations, the light chain should be placed before the heavy chain to avoid an excess of toxic free heavy chain (Proudfoot, *Nature*, 322:52, 1986; and Kohler, *Proc. Natl. Acad. Sci. USA*, 77:2 197, 1980). The coding sequences for the heavy and light chains optionally include cDNA or genomic DNA.

[0182] In another embodiment, antibodies are generated using various phage display methods known in the art. In phage display methods, functional antibody domains are displayed on the surface of phage particles which carry the polynucleotide sequences encoding them. In a particular embodiment, such phage is utilized to display antigen binding domains, such as Fab and Fv or disulfide-bond stabilized Fv, expressed from a repertoire or combinatorial antibody library (e.g., human or murine). Phage expressing an antigen binding domain that binds the antigen of interest is optionally selected or identified with antigen, e.g., using labeled antigen or antigen bound or captured to a solid surface or bead. Phages used in these methods are typically filamentous phage, including fd and M13. The antigen binding domains are expressed as a recombinantly fused protein to either the phage gene III or gene VIII protein. Examples of phage display methods that can be used to make the immunoglobulins, or fragments thereof, of the present invention include those disclosed in Brinkman et al., *J. Immunol. Methods*, 182:41-50, 1995; Ames et al., *J. Immunol. Methods*, 184:177-186, 1995; Kettleborough et al., *Eur. J. Immunol.*, 24:952-958, 1994; Persic et al., *Gene*, 187:9-18, 1997; Burton et al., *Advances in Immunology*, 57:191-280, 1994; PCT application No. PCT/GB91/01134; PCT publications WO 90/02809; WO 91/10737; WO 92/01047; WO 92/18619; WO 93/11236; WO 95/15982; WO 95/20401; and U.S. Pat. Nos. 5,698,426; 5,223,409; 5,403,484; 5,580,717; 5,427,908; 5,750,753; 5,821,047; 5,571,698; 5,427,908; 5,516,637; 5,780,225; 5,658,727; 5,733,743 and 5,969,108; each of which is incorporated herein by reference in its entirety.

[0183] As described in the above references, after phage selection, the antibody coding regions from the phage is optionally isolated and used to generate whole antibodies, including human antibodies, or any other desired fragments, and expressed in any desired host, including mammalian cells, insect cells, plant cells, yeast, and bacteria, e.g., as described in detail below. For example, techniques to recombinantly produce Fab, Fab' and F(ab')₂ fragments are optionally employed using methods known in the art such as those disclosed in PCT publication WO 92/22324; Mullinax et al., *BioTechniques*, 12(6):864-869, 1992; and Sawai et al., *AJR*, 34:26-34, 1995; and Better et al., *Science*, 240:1041-1043, 1988 (each of which is incorporated by reference in its entirety). Examples of techniques operable to produce single-chain Fvs and antibodies include those described in U.S. Pat. Nos. 4,946,778 and 5,258,498; Huston et al., *Methods in Enzymology*, 203:46-88, 1991; Shu et al., *PNAS*, 90:7995-7999, 1993; and Skerra et al., *Science*, 240:1038-1040, 1988.

[0184] Once an antibody molecule of the invention has been produced by any methods described above, or otherwise known in the art, it is then optionally purified by any method known in the art for purification of an immunoglobulin mol-

ecule, for example, by chromatography (e.g., ion exchange, affinity, particularly by affinity for the specific antigen after Protein A or Protein G purification, and sizing column chromatography), centrifugation, differential solubility, or by any other standard technique(s) for the purification of proteins. Further, the antibodies of the present invention or fragments thereof are optionally fused to heterologous polypeptide sequences described herein or otherwise known in the art to facilitate purification. Illustrative examples include 6×His tag, FLAG tag, biotin, avidin, or other system.

[0185] For some uses, including in vivo use of antibodies in humans and in vitro detection assays, it is preferable to use chimeric, humanized, or human antibodies. A chimeric antibody is a molecule in which different portions of the antibody are derived from different animal species, such as antibodies having a variable region derived from a murine monoclonal antibody and a constant region derived from a human immunoglobulin. Methods for producing chimeric antibodies are known in the art. See e.g., Morrison, *Science*, 229:1202, 1985; Oi et al., *BioTechniques*, 4:214 1986; Gillies et al., *J. Immunol. Methods*, 125:191-202, 1989; U.S. Pat. Nos. 5,807,715; 4,816,567; and 4,816,397, which are incorporated herein by reference in their entireties. Humanized antibodies are antibody molecules from non-human species that bind the desired antigen having one or more complementarity determining regions (CDRs) from the non-human species and framework regions from a human immunoglobulin molecule. Often, framework residues in the human framework regions will be substituted with the corresponding residue from the CDR donor antibody to alter, preferably improve, antigen binding. These framework substitutions are identified by methods well known in the art, e.g., by modeling of the interactions of the CDR and framework residues to identify framework residues important for antigen binding and sequence comparison to identify unusual framework residues at particular positions. See, e.g., Queen et al., U.S. Pat. No. 5,585,089; Riechmann et al., *Nature*, 332:323, 1988, which are incorporated herein by reference in their entireties. Antibodies are humanized using a variety of techniques known in the art including, for example, CDR-grafting (EP 239,400; PCT publication WO 91/09967; U.S. Pat. Nos. 5,225,539; 5,530,101 and 5,585,089), veneering or resurfacing (EP 592,106; EP 519,596; Padlan, *Molecular Immunology*, 28(4/5): 489-498, 1991; Studnicka et al., *Protein Engineering*, 7(6): 805-814, 1994; Roguska et al., *Proc Natl. Acad. Sci. USA*, 91:969-973, 1994), and chain shuffling (U.S. Pat. No. 5,565,332), all of which are hereby incorporated by reference in their entireties.

[0186] Completely human antibodies are particularly desirable for therapeutic treatment of human patients. Human antibodies are made by a variety of methods known in the art illustratively including phage display methods described above using antibody libraries derived from human immunoglobulin sequences. See U.S. Pat. Nos. 4,444,887 and 4,716,111; and PCT publications WO 98/46645; WO 98/50433; WO 98/24893; WO 98/16654; WO 96/34096; WO 96/33735; and WO 91/10741, each of which is incorporated herein by reference in its entirety.

[0187] Human antibodies are also illustratively produced using transgenic mice which are incapable of expressing functional endogenous immunoglobulins, but which can express human immunoglobulin genes. For an overview of this technology for producing human antibodies, see Lonberg and Huszar, *Int. Rev. Immunol.*, 13:65-93, 1995. For a

detailed discussion of this technology for producing human antibodies and human monoclonal antibodies and protocols for producing such antibodies, see, e.g., PCT publications WO 98/24893; WO 92/01047; WO 96/34096; WO 96/33735; European Patent No. 0 598 877; U.S. Pat. Nos. 5,413,923; 5,625,126; 5,633,425; 5,569,825; 5,661,016; 5,545,806; 5,814,318; 5,885,793; 5,916,771; and 5,939,598, which are incorporated by reference herein in their entireties. In addition, companies such as Abgenix, Inc. (Fremont, Calif.), Medarex (NJ) and Genpharm (San Jose, Calif.) can be engaged to provide human antibodies directed against a selected antigen using technology similar to that described above.

[0188] Completely human antibodies which recognize a selected epitope are optionally generated using a technique referred to as “guided selection.” In this approach a selected non-human monoclonal antibody, e.g., a mouse antibody, is used to guide the selection of a completely human antibody recognizing the same epitope. (Jespers et al., *Bio/technology*, 12:899-903, 1988).

[0189] Antibodies fused or conjugated to heterologous polypeptides are optionally used in *in vitro* immunoassays and in purification methods (e.g., affinity chromatography) known in the art. See e.g., PCT publication No. WO 93/21232; EP 439,095; Naramura et al., *Immunol. Lett.*, 39:91-99, 1994; U.S. Pat. No. 5,474,981; Gillies et al., *PNAS*, 89:1428-1432, 1992; and Fell et al., *J. Immunol.*, 146:2446-2452, 1991, which are incorporated herein by reference in their entireties.

[0190] Antibodies may also be illustratively attached to solid supports, which are particularly useful for immunoassays or purification of the polypeptides of the invention or fragments, derivatives, analogs, or variants thereof, or similar molecules having the similar enzymatic activities as the polypeptide of the invention. Such solid supports include, but are not limited to, glass, cellulose, polyacrylamide, nylon, polystyrene, polyvinyl chloride or polypropylene.

Pharmaceutical Compositions and Kits

[0191] The present invention encompasses pharmaceutical compositions including antiviral agents of the present invention. In a specific embodiment, the antiviral agent is preferably an antibody which immunospecifically binds and neutralizes the hEbola virus or variants thereof, or any proteins derived therefrom. In another specific embodiment, the antiviral agent is a polypeptide or nucleic acid molecule of the invention. The pharmaceutical compositions have utility as an antiviral prophylactic agent are illustratively administered to a subject where the subject has been exposed or is expected to be exposed to a virus.

[0192] Various delivery systems are known and operable to administer the pharmaceutical composition of the invention, illustratively, encapsulation in liposomes, microparticles, microcapsules, recombinant cells capable of expressing the mutant viruses, and receptor mediated endocytosis (see, e.g., Wu and Wu, 1987, *J. Biol. Chem.* 262:4429-4432). Methods of introduction include but are not limited to intradermal, intramuscular, intraperitoneal, intravenous, subcutaneous, intranasal, epidural, and oral routes. The compounds may be administered by any convenient route, for example by infusion or bolus injection, by absorption through epithelial or mucocutaneous linings (e.g., oral mucosa, rectal and intestinal mucosa, etc.) and optionally administered together with other biologically active agents. Administration is systemic

or local. In a preferred embodiment, it is desirable to introduce the pharmaceutical compositions of the invention into the lungs by any suitable route. Pulmonary administration can also be employed, e.g., by use of an inhaler or nebulizer, and formulation with an aerosolizing agent.

[0193] In a specific embodiment, it is desirable to administer the pharmaceutical compositions of the invention locally to the area in need of treatment. This administration may be achieved by, for example, and not by way of limitation, local infusion during surgery, topical application, e.g., in conjunction with a wound dressing after surgery, by injection, by means of a catheter, by means of a suppository, by means of nasal spray, or by means of an implant, the implant being of a porous, non-porous, or gelatinous material, including membranes, such as sialastic membranes, or fibers. In one embodiment, administration can be by direct injection at the site (or former site) infected tissues.

[0194] In another embodiment, the pharmaceutical composition is delivered in a vesicle, in particular a liposome (see Langer, 1990, *Science* 249:1527-1533; Treat et al., in *Liposomes in the Therapy of Infectious Disease and Cancer*, Lopez Berestein and Fidler (eds.), Liss, New York, pp. 353-365 (1989); Lopez-Berestein, *ibid.*, pp. 317-327; see generally *ibid.*).

[0195] In yet another embodiment, the pharmaceutical composition is delivered in a controlled release system. In one embodiment, a pump is used (see Langer, *supra*; Sefton, 1987, *CRC Crit. Ref. Biomed. Eng.* 14:201; Buchwald et al., 1980, *Surgery* 88:507; and Saudek et al., 1989, *N. Engl. J. Med.* 321:574). In another embodiment, polymeric materials are used (see *Medical Applications of Controlled Release*, Langer and Wise (eds.), CRC Pres., Boca Raton, Fla. (1974); *Controlled Drug Bioavailability, Drug Product Design and Performance*, Smolen and Ball (eds.), Wiley, New York (1984); Ranger and Peppas, *J. Macromol. Sci. Rev. Macromol. Chem.* 23:61 (1983); see also Levy et al., 1985, *Science* 228:190; During et al., 1989, *Ann. Neurol.* 25:351; Howard et al., 1989, *J. Neurosurg.* 71:105). In yet another embodiment, a controlled release system is placed in proximity of the composition's target, i.e., the lung, thus, requiring only a fraction of the systemic dose (see, e.g., Goodson, in *Medical Applications of Controlled Release*, *supra*, vol. 2, pp. 115-138 (1984)).

[0196] Other controlled release systems are discussed in the review by Langer (*Science* 249:1527-1533 (1990)) the contents of which are incorporated herein by reference.

[0197] The pharmaceutical compositions of the present invention illustratively include a therapeutically effective amount of a live attenuated, inactivated or killed West African hEbola virus, or recombinant or chimeric hEbola virus, and a pharmaceutically acceptable carrier. In a specific embodiment, the term “pharmaceutically acceptable” means approved by a regulatory agency of the Federal or a state government or listed in the U.S. Pharmacopeia or other generally recognized pharmacopeia for use in animals, and more particularly in humans. The term “carrier” refers to a diluent, adjuvant, excipient, or vehicle with which the pharmaceutical composition is administered. Such pharmaceutical carriers are illustratively sterile liquids, such as water and oils, including those of petroleum, animal, vegetable or synthetic origin, such as peanut oil, soybean oil, mineral oil, sesame oil and the like. Water is a preferred carrier when the pharmaceutical composition is administered intravenously. Saline solutions and aqueous dextrose and glycerol solutions are optionally

employed as liquid carriers, particularly for injectable solutions. Suitable pharmaceutical excipients include starch, glucose, lactose, sucrose, gelatin, malt, rice, flour, chalk, silica gel, sodium stearate, glycerol monostearate, talc, sodium chloride, dried skim milk, glycerol, propylene, glycol, water, ethanol and the like. The composition, if desired, also contains wetting or emulsifying agents, or pH buffering agents. These compositions optionally take the form of solutions, suspensions, emulsion, tablets, pills, capsules, powders, sustained release formulations and the like. The composition is optionally formulated as a suppository, with traditional binders and carriers such as triglycerides. Oral formulation illustratively includes standard carriers such as pharmaceutical grades of mannitol, lactose, starch, magnesium stearate, sodium saccharine, cellulose, magnesium carbonate, etc. Examples of suitable pharmaceutical carriers are described in "Remington's Pharmaceutical Sciences" by E. W. Martin. The formulation should suit the mode of administration.

[0198] In a preferred embodiment, the composition is formulated in accordance with routine procedures as a pharmaceutical composition adapted for intravenous administration to human beings. Typically, compositions for intravenous administration are solutions in sterile isotonic aqueous buffer. The composition also includes an optional solubilizing agent and a local anesthetic such as lignocaine to ease pain at the site of the injection. Generally, the ingredients are supplied either separately or mixed together in unit dosage form, for example, as a dry lyophilized powder or water-free concentrate in a hermetically sealed container such as an ampoule or sachette indicating the quantity of active agent. Where the composition is to be administered by infusion, it can be dispensed with an infusion bottle containing sterile pharmaceutical grade water or saline. Where the composition is administered by injection, an ampoule of sterile water for injection or saline is optionally provided so that the ingredients may be mixed prior to administration.

[0199] The pharmaceutical compositions of the invention are illustratively formulated as neutral or salt forms. Pharmaceutically acceptable salts illustratively include those formed with free amino groups such as those derived from hydrochloric, phosphoric, acetic, oxalic, tartaric acids, etc., and those formed with free carboxyl groups such as those derived from sodium, potassium, ammonium, calcium, ferric hydroxides, isopropylamine, triethylamine, 2 ethylamino ethanol, histidine, procaine, etc.

[0200] The amount of the pharmaceutical composition of the invention which will be effective in the treatment of a particular disorder or condition will depend on the nature of the disorder or condition, and can be determined by standard clinical techniques. In addition, in vitro assays are optionally employed to help identify optimal dosage ranges. The precise dose to be employed in the formulation will also depend on the route of administration, and the seriousness of the disease or disorder, and should be decided according to the judgment of the practitioner and each patient's circumstances. However, suitable dosage ranges for intravenous administration are generally about 20 to 500 micrograms of active compound per kilogram body weight. Suitable dosage ranges for intranasal administration are generally about 0.01 pg/kg body weight to 1 mg/kg body weight. Effective doses may be extrapolated from dose response curves derived from in vitro or animal model test systems.

[0201] Suppositories generally contain active ingredient in the range of 0.5% to 10% by weight; oral formulations preferably contain 10% to 95% active ingredient.

[0202] The invention also provides a pharmaceutical pack or kit including one or more containers filled with one or more of the ingredients of the pharmaceutical compositions of the invention. Optionally associated with such container(s) is a notice in the form prescribed by a governmental agency regulating the manufacture, use or sale of pharmaceuticals or biological products, which notice reflects approval by the agency of manufacture, use or sale for human administration. In a preferred embodiment, the kit contains an antiviral agent of the invention, e.g., an antibody specific for the polypeptides encoded by a nucleotide sequence of SEQ ID NOs: 1 or 10, or as shown in SEQ ID NOs: 2-9, 59, or 11-19, or any hEbola epitope, or a polypeptide or protein of the present invention, or a nucleic acid molecule of the invention, alone or in combination with adjuvants, antivirals, antibiotics, analgesic, bronchodilators, or other pharmaceutically acceptable excipients.

[0203] The present invention further encompasses kits including a container containing a pharmaceutical composition of the present invention and instructions for use.

Detection Assays

[0204] The present invention provides a method for detecting an antibody, which immunospecifically binds to the hEbola virus, in a biological sample, including for example blood, serum, plasma, saliva, urine, feces, etc., from a patient suffering from hEbola infection, and/or hemorrhagic fever. In a specific embodiment, the method including contacting the sample with the hEbola virus, for example, of Deposit Accession No. 200706291, or having a genomic nucleic acid sequence of SEQ ID NOs: 1 or 10, directly immobilized on a substrate and detecting the virus-bound antibody directly or indirectly by a labeled heterologous anti-isotype antibody. In another specific embodiment, the sample is contacted with a host cell which is infected by the hEbola virus, for example, of Deposit Accession No. 200706291, or having a genomic nucleic acid sequence of SEQ ID NOs: 1 or 10, and the bound antibody is optionally detected by immunofluorescent assay.

[0205] An exemplary method for detecting the presence or absence of a polypeptide or nucleic acid of the invention in a biological sample involves obtaining a biological sample from various sources and contacting the sample with a compound or an agent capable of detecting an epitope or nucleic acid (e.g., mRNA, genomic DNA) of the hEbola virus such that the presence of the hEbola virus is detected in the sample. A preferred agent for detecting hEbola mRNA or genomic RNA of the invention is a labeled nucleic acid probe capable of hybridizing to mRNA or genomic RNA encoding a polypeptide of the invention. The nucleic acid probe is, for example, a nucleic acid molecule including the nucleotide sequence of SEQ ID NOs: 1 or 10, a complement thereof, or a portion thereof, such as an oligonucleotide of at least 15, 20, 25, 30, 50, 100, 250, 500, 750, 1000 or more contiguous nucleotides in length and sufficient to specifically hybridize under stringent conditions to a hEbola mRNA or genomic RNA.

[0206] As used herein, the term "stringent conditions" describes conditions for hybridization and washing under which nucleotide sequences having at least 30%, 35%, 40%, 45%, 50%, 55%, 60%, 65%, 70%, 75%, 80%, 85%, 90%, or 95% identity to each other typically remain hybridized to

each other. Such hybridization conditions are described in, for example but not limited to, *Current Protocols in Molecular Biology*, John Wiley & Sons, N.Y. (1989), 6.3.1 6.3.6; *Basic Methods in Molecular Biology*, Elsevier Science Publishing Co., Inc., N.Y. (1986), pp. 75 78, and 84 87; and *Molecular Cloning*, Cold Spring Harbor Laboratory, N.Y. (1982), pp. 387 389, and are well known to those skilled in the art. A preferred, non-limiting example of stringent hybridization conditions is hybridization in 6× sodium chloride/sodium citrate (SSC), 0.5% SDS at about 68° C. followed by one or more washes in 2×SSC, 0.5% SDS at room temperature. Another preferred, non-limiting example of stringent hybridization conditions is hybridization in 6×SSC at about 45° C. followed by one or more washes in 0.2×SSC, 0.1% SDS at 50 to 65° C.

[0207] A nucleic acid probe, polynucleotide, oligonucleotide, or other nucleic acid is preferably purified. An “isolated” or “purified” nucleotide sequence is substantially free of cellular material or other contaminating proteins from the cell or tissue source from which the nucleotide is derived, or is substantially free of chemical precursors or other chemicals when chemically synthesized. The language “substantially free of cellular material” includes preparations of a nucleotide/oligonucleotide in which the nucleotide/oligonucleotide is separated from cellular components of the cells from which it is isolated or produced. Thus, a nucleotide/oligonucleotide that is substantially free of cellular material includes preparations of the nucleotide having less than about 30%, 20%, 10%, 5%, 2.5%, or 1%, (by dry weight) of contaminating material. When nucleotide/oligonucleotide is produced by chemical synthesis, it is preferably substantially free of chemical precursors or other chemicals, i.e., it is separated from chemical precursors or other chemicals which are involved in the synthesis of the protein. Accordingly, such preparations of the nucleotide/oligonucleotide have less than about 30%, 20%, 10%, or 5% (by dry weight) of chemical precursors or compounds other than the nucleotide/oligonucleotide of interest. In a preferred embodiment of the present invention, the nucleotide/oligonucleotide is isolated or purified.

[0208] In another preferred specific embodiment, the presence of hEbola virus is detected in the sample by a reverse transcription polymerase chain reaction (RT-PCR) using the primers that are constructed based on a partial nucleotide sequence of the genome of hEbola virus, for example, that of Deposit Accession No. 200706291, or having a genomic nucleic acid sequence of SEQ ID NOs: 1 or 10. In a non-limiting specific embodiment, preferred primers to be used in a RT-PCR method are the primers are described in detail herein.

[0209] In more preferred specific embodiment, the present invention provides a real-time quantitative PCR assay to detect the presence of hEbola virus in a biological sample by subjecting the cDNA obtained by reverse transcription of the extracted total RNA from the sample to PCR reactions using the specific primers described in detail herein, and a fluorescence dye, such as SYBR® Green I, which fluoresces when bound nonspecifically to double-stranded DNA. The fluorescence signals from these reactions are captured at the end of extension steps as PCR product is generated over a range of the thermal cycles, thereby allowing the quantitative determination of the viral load in the sample based on an amplification plot.

[0210] A preferred agent for detecting hEbola is an antibody that specifically binds a polypeptide of the invention or any hEbola epitope, preferably an antibody with a detectable label. Antibodies are illustratively polyclonal, or more preferably, monoclonal. An intact antibody, or a fragment thereof (e.g., Fab or F(ab')₂) is operable herein.

[0211] The term “labeled”, with regard to the probe or antibody, is intended to encompass direct labeling of the probe or antibody by coupling (i.e., physically linking) a detectable substance to the probe or antibody, optionally via a linker, as well as indirect labeling of the probe or antibody by reactivity with another reagent that is directly labeled. Examples of indirect labeling include detection of a primary antibody using a fluorescently labeled secondary antibody and end-labeling of a DNA probe with biotin such that it is detectable with fluorescently labeled streptavidin. The detection method of the invention is optionally used to detect mRNA, protein (or any epitope), or genomic RNA in a sample in vitro as well as in vivo. Exemplary in vitro techniques for detection of mRNA include northern hybridizations, in situ hybridizations, RT-PCR, and RNase protection. In vitro techniques for detection of an epitope of hEbola illustratively include enzyme linked immunosorbent assays (ELISAs), western blots, immunoprecipitations and immunofluorescence. In vitro techniques for detection of genomic RNA include northern hybridizations, RT-PCT, and RNase protection. Furthermore, in vivo techniques for detection of hEbola include introducing into a subject organism a labeled antibody directed against the polypeptide. In one embodiment, the antibody is labeled with a radioactive marker whose presence and location in the subject organism is detected by standard imaging techniques, including autoradiography.

[0212] In a specific embodiment, the methods further involve obtaining a control sample from a control subject, contacting the control sample with a compound or agent capable of detecting hEbola, e.g., a polypeptide of the invention or mRNA or genomic RNA encoding a polypeptide of the invention, such that the presence of hEbola or the polypeptide or mRNA or genomic RNA encoding the polypeptide is detected in the sample, and comparing the absence of hEbola or the polypeptide or mRNA or genomic RNA encoding the polypeptide in the control sample with the presence of hEbola, or the polypeptide or mRNA or genomic DNA encoding the polypeptide in the test sample.

[0213] The invention also encompasses kits for detecting the presence of hEbola or a polypeptide or nucleic acid of the invention in a test sample. The kit illustratively includes a labeled compound or agent capable of detecting hEbola or the polypeptide or a nucleic acid molecule encoding the polypeptide in a test sample and, in certain embodiments, a means for determining the amount of the polypeptide or mRNA in the sample (e.g., an antibody which binds the polypeptide or an oligonucleotide probe which binds to DNA or mRNA encoding the polypeptide). Kits optionally include instructions for use.

[0214] For antibody-based kits, the kit illustratively includes: (1) a first antibody (e.g., attached to a solid support) which binds to a polypeptide of the invention or hEbola epitope; and, optionally, (2) a second, different antibody which binds to either the polypeptide or the first antibody and is preferably conjugated to a detectable agent.

[0215] For oligonucleotide-based kits, the kit illustratively includes: (1) an oligonucleotide, e.g., a detectably labeled oligonucleotide, which hybridizes to a nucleic acid sequence

encoding a polypeptide of the invention or to a sequence within the hEbola genome; or (2) a pair of primers useful for amplifying a nucleic acid molecule containing an hEbola sequence. The kit optionally includes a buffering agent, a preservative, or a protein stabilizing agent. The kit optionally includes components necessary for detecting the detectable agent (e.g., an enzyme or a substrate). The kit optionally contains a control sample or a series of control samples which can be assayed and compared to the test sample contained. Each component of the kit is usually enclosed within an individual container and all of the various containers are within a single package along with instructions for use.

Screening Assays to Identify Antiviral Agents

[0216] The invention provides methods for the identification of a compound that inhibits the ability of hEbola virus to infect a host or a host cell. In certain embodiments, the invention provides methods for the identification of a compound that reduces the ability of hEbola virus to replicate in a host or a host cell. Any technique well known to the skilled artisan is illustratively used to screen for a compound useful to abolish or reduce the ability of hEbola virus to infect a host and/or to replicate in a host or a host cell.

[0217] In certain embodiments, the invention provides methods for the identification of a compound that inhibits the ability of hEbola virus to replicate in a mammal or a mammalian cell. More specifically, the invention provides methods for the identification of a compound that inhibits the ability of hEbola virus to infect a mammal or a mammalian cell. In certain embodiments, the invention provides methods for the identification of a compound that inhibits the ability of hEbola virus to replicate in a mammalian cell. In a specific embodiment, the mammalian cell is a human cell.

[0218] In another embodiment, a cell is contacted with a test compound and infected with the hEbola virus. In certain embodiments, a control culture is infected with the hEbola virus in the absence of a test compound. The cell is optionally contacted with a test compound before, concurrently with, or subsequent to the infection with the hEbola virus. In a specific embodiment, the cell is a mammalian cell. In an even more specific embodiment, the cell is a human cell. In certain embodiments, the cell is incubated with the test compound for at least 1 minute, at least 5 minutes, at least 15 minutes, at least 30 minutes, at least 1 hour, at least 2 hours, at least 5 hours, at least 12 hours, or at least 1 day. The titer of the virus is optionally measured at any time during the assay. In certain embodiments, a time course of viral growth in the culture is determined. If the viral growth is inhibited or reduced in the presence of the test compound, the test compound is identified as being effective in inhibiting or reducing the growth or infection of the hEbola virus. In a specific embodiment, the compound that inhibits or reduces the growth of the hEbola virus is tested for its ability to inhibit or reduce the growth rate of other viruses to test its specificity for the hEbola virus.

[0219] In one embodiment, a test compound is administered to a model animal and the model animal is infected with the hEbola virus. In certain embodiments, a control model animal is infected with the hEbola virus without the administration of a test compound. The test compound is optionally administered before, concurrently with, or subsequent to the infection with the hEbola virus. In a specific embodiment, the model animal is a mammal. In an even more specific embodiment, the model animal is, but is not limited to, a cotton rat, a mouse, or a monkey. The titer of the virus in the model animal

is optionally measured at any time during the assay. In certain embodiments, a time course of viral growth in the culture is determined. If the viral growth is inhibited or reduced in the presence of the test compound, the test compound is identified as being effective in inhibiting or reducing the growth or infection of the hEbola virus. In a specific embodiment, the compound that inhibits or reduces the growth of the hEbola in the model animal is tested for its ability to inhibit or reduce the growth rate of other viruses to test its specificity for the hEbola virus.

[0220] According to the method of the invention, a human or an animal is optionally treated for for EboBun or EboIC, other viral infection or bacterial infection by administering an effective amount of an inventive therapeutic composition. Preferably, a vaccine is administered prophylactically. An "effective amount" is an amount that will induce an immune response in a subject. Illustratively, an effective amount of the compositions of this invention ranges from nanogram/kg to milligram/kg amounts for young children and adults. Equivalent dosages for lighter or heavier body weights can readily be determined. The dose should be adjusted to suit the individual to whom the composition is administered and will vary with age, weight and metabolism of the individual. The exact amount of the composition required will vary from subject to subject, depending on the species, age, weight and general condition of the subject, the particular peptide or polypeptide used, its mode of administration and the like. An appropriate amount can be determined by one of ordinary skill in the art using only routine experimentation given the teachings herein. One skilled in the art will realize that dosages are best optimized by the practicing physician or veterinarian and methods for determining dose amounts and regimens and preparing dosage forms are described, for example, in Remington's Pharmaceutical Sciences, (Martin, E. W., ed., latest edition), Mack Publishing Co., Easton, Pa. Preferably, a single administration is operable to induce an immune response.

[0221] Methods involving conventional biological techniques are described herein. Such techniques are generally known in the art and are described in detail in methodology treatises such as *Molecular Cloning: A Laboratory Manual*, 2nd ed., vol. 1-3, ed. Sambrook et al., Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y., 1989; and *Current Protocols in Molecular Biology*, ed. Ausubel et al., Greene Publishing and Wiley-Interscience, New York, 1992 (with periodic updates). Immunological methods (e.g., preparation of antigen-specific antibodies, immunoprecipitation, and immunoblotting) are described, e.g., in *Current Protocols in Immunology*, ed. Coligan et al., John Wiley & Sons, New York, 1991; and *Methods of Immunological Analysis*, ed. Masseyeff et al., John Wiley & Sons, New York, 1992.

[0222] Embodiments of inventive compositions and methods are illustrated in the following detailed examples. These examples are provided for illustrative purposes and are not considered limitations on the scope of inventive compositions and methods.

EXAMPLES

Example 1

Newly Discovered Ebola Virus Associated with Hemorrhagic Fever Outbreak in Bundibugyo, Uganda

[0223] In late November 2007 HF cases were reported in the townships of Bundibugyo and Kikyo in Bundibugyo Dis-

trict, Western Uganda (FIG. 1A). These samples were assayed as described by Towner, JS, et al., *PLoS Pathog*, 2008 November; 4(11): e1000212, the contents of which are incorporated herein by reference for methods, results, reagents, and all other aspects of the publication. A total of 29 blood samples were initially collected from suspect cases and showed evidence of acute ebolavirus infection in eight specimens using a broadly reactive ebolavirus antigen capture assay known to cross-react with the different ebolavirus species¹ and an IgM capture assay based on Zaire ebolavirus reagents (Table 1). These specimens were negative when initially tested with highly sensitive real-time RT-PCR assays specific for all known Zaire and Sudan ebolaviruses and marburgviruses. However, further evidence of acute ebolavirus infection was obtained using a traditionally less sensitive (relative to the real-time RT-PCR assays) but more broadly reactive filovirus L gene-specific RT-PCR assay (1 specimen) (Table 1). Sequence analysis of the PCR fragment (400 bp of the virus L gene) revealed the reason for the initial failure of the real-time RT-PCR assays, as the sequence was distinct from that of the 4 known species of ebolavirus, although distantly related to Côte d'Ivoire ebolavirus. In total, 9 of 29 specimens showed evidence of ebolavirus infection, and all tests were negative for marburgvirus (data not shown).

[0224] Approximately 70% of the virus genome was rapidly sequenced from total RNA extracted from a patient serum (#200706291) using a newly established metagenomics pyrosequencing method (454 Life Sciences) which involves successive rounds of random DNA amplification⁸. Using the newly derived draft sequence, a real-time RT-PCR assay specific for the NP gene of this virus was quickly developed and evaluated. The assay was shown to have excellent sensitivity (Table 1), finding positive all the initial six samples that tested positive by either virus antigen capture (five specimens) or virus isolation assays (four specimens). The antigen-capture, IgM, IgG and newly designed real-time PCR assays were quickly transferred to the Uganda Virus Research Institute during the course of the outbreak to facilitate rapid identification and isolation of Ebola cases in the affected area for efficient control of the outbreak. The outbreak continued through late December 2007, and resulted in 149 suspected cases and 37 deaths⁹.

[0225] Table 1. Ebolavirus diagnostic results of initial 29 specimens obtained from Bundibugyo District with numerical specimen numbers assigned. RT-PCR refers to results obtained from conventional PCR using the broadly reactive Filo A/B primers¹³. Ag, IgM, and IgG refer to results from ELISA-based assays^{10, 11} with Zaire ebolavirus reagents while virus isolation refers to culture attempts on Vero E6 cells². Q-RT-PCR refers to results obtained using the optimized Bundibugyo ebolavirus specific real-time RT-PCR assay with cycle threshold (Ct) values of positive (Pos) samples indicated in the far right column. * Specimen #200706291 is the clinical sample from which prototype isolate #811250 was obtained.

TABLE 1

Sample No.	RT-PCR	Ag	IgM	IgG	Virus Isolation	Q- RT-PCR	Ct
200706288	neg	neg	neg	neg	neg	neg	40
200706289	neg	neg	neg	neg	neg	neg	40
200706290	neg	neg	neg	neg	neg	neg	40
200706291*	Pos	Pos	neg	neg	Pos	Pos	23.64

TABLE 1-continued

Sample No.	RT-PCR	Ag	IgM	IgG	Virus Isolation	Q- RT-PCR	Ct
200706292	neg	neg	neg	neg	neg	neg	40
200706293	neg	neg	neg	neg	neg	neg	40
200706294	neg	neg	neg	neg	neg	neg	40
200706295	neg	neg	neg	neg	neg	neg	40
200706296	neg	neg	Pos	Pos	neg	neg	40
200706297	neg	neg	Pos	Pos	neg	neg	40
200706298	neg	Pos	Pos	Pos	neg	Pos	34.83
200706299	neg	neg	Pos	Pos	neg	neg	40
200706300	neg	neg	neg	neg	neg	neg	40
200706301	neg	neg	neg	neg	neg	neg	40
200706302	neg	Pos	Pos	neg	neg	Pos	35.01
200706303	neg	neg	neg	neg	neg	neg	40
200706304	neg	neg	neg	neg	Pos	Pos	38.18
200706305	neg	neg	neg	neg	neg	neg	40
200706306	neg	neg	neg	neg	neg	neg	40
200706307	neg	neg	neg	neg	neg	neg	40
200706320	ND	Pos	neg	neg	Pos	Pos	30.24
200706321	ND	neg	neg	neg	neg	neg	40
200706322	ND	neg	neg	neg	neg	neg	40
200706323	ND	neg	neg	neg	neg	neg	40
200706324	ND	neg	neg	neg	neg	neg	40
200706325	ND	neg	neg	neg	neg	neg	40
200706326	ND	neg	neg	neg	neg	neg	40
200706327	ND	Pos	neg	neg	Pos	Pos	34.41
200706328	ND	neg	neg	neg	neg	neg	40

[0226] The entire genome sequence of this virus was completed using a classic primer walking sequencing approach on RNA. The complete genome of the Eb ebolavirus was not available, so it too was derived by a similar combination of random primed pyrosequencing and primer walking approaches. Acquisition of these sequences allowed for the first time the phylogenetic analysis of the complete genomes of representatives of all known species of Ebola and Marburg viruses. The analysis revealed that the newly discovered virus differed from the four existing ebolavirus species (FIG. 1), with approximately 32% nucleotide difference from even the closest relative, EboIC (Table 2). Similar complete genome divergence (35-45%) is seen between the previously characterized ebolavirus species.

[0227] Table 2. Identity matrix based on comparisons of full-length genome sequences of Zaire ebolaviruses 1976 (Genbank accession number NC_002549) and 1995 (Genbank accession number AY354458), Sudan ebolavirus 2000 (Genbank accession number NC_006432), Cote d'Ivoire ebolavirus 1994 (SEQ ID NO: 10), Reston ebolavirus 1989 (Genbank accession number NC_004161), and Bundibugyo ebolavirus 2007 (SEQ ID NO: 1).

TABLE 2

	Zaire '95	Sudan '00	EboIC '94	EboBun '07	Reston '89
Zaire '76	.988	.577	.630	.632	.581
Zaire '95		.577	.631	.633	.581
Sudan '00			.577	.577	.609
EboIC '94				.683	.575
EboBun '07					.576

[0228] The material and information obtained from the discovery of the new unique virus EboBun and the realization that together with EboIC these viruses represent a Glade of Bundibugyo-Ivory Coast Ebola virus species is valuable,

and makes possible the development of clinical, diagnostic and research tools directed to human hEbola infection.

Material and Methods

[0229] Ebolavirus Detection and Virus Isolation.

[0230] Several diagnostic techniques were used for each sample: (i) antigen capture, IgG, and IgM assays were performed as previously described¹¹ (ii) virus isolation attempts were performed on Vero E6 cells¹² and monitored for 14 days; (iii) RNA was extracted and tested for Zaire¹⁶ and Sudan ebolavirus and marburgvirus⁴ using real-time quantitative RT-PCR assays designed to detect all known species of each respective virus species the primers/probe for the Sudan ebolavirus assay were EboSudBMG 1(+) 5'-GCC ATG GIT TCA GGT TTG AG-3' (SEQ ID NO: 21), EboSudBMG 1(-) 5'-GGT IAC ATT GGG CAA CAA TTC A-3' (SEQ ID NO: 22) and Ebola Sudan BMG Probe 5'-FAM-AC GGT GCA CAT TCT CCT TTT CTC GGA-BHQ1 (SEQ ID NO: 23)]; (iv) the conventional RT-PCR was performed with the filo A/B primer set as previously described¹⁶ using Superscript III (Invitrogen) according to the manufacturer's instructions. The specimen 200706291 was selected as the reference sample for further sequence analysis.

[0231] Genome Sequencing.

[0232] Pyrosequencing was carried out utilizing the approach developed by 454 Life Sciences, and the method described by Cox-Foster et al.⁸ Subsequent virus whole genome primer walking was performed as previously described¹⁷ but using the primers specific for Bundibugyo ebolavirus RT-PCR amplification. In total, the entire virus genome was amplified in six overlapping RT-PCR fragments (all primers listed 5' to 3'): fragment A (predicted size 2.7 kb) was amplified using forward-GTGAGACAAAGAATCAT-TCTTG (SEQ ID NO: 24) with reverse-CATCAATTGCT-CAGAGATCCACC (SEQ ID NO: 25); fragment B (predicted size 3.0 kb) was amplified using forward-CCAACAACACTGCATGTAAGT (SEQ ID NO: 26) with reverse-AGGTCGCGTTAATCTTCATC (SEQ ID NO: 27); fragment C (predicted size 3.5 kb) was amplified using forward-GATGGTTGAGTTACTTTCCGG (SEQ ID NO: 28) with reverse-GTCTTGAGTCATCAATGCC (SEQ ID NO: 29); fragment D (predicted size 3.1 kb) was amplified using forward-CCACCAGCACCAAAGGAC (SEQ ID NO: 30) with reverse-CTATCGGCAATGTAACATTTGG (SEQ ID NO: 31); fragment E (predicted size 3.4 kb) was amplified using forward-GCCGTTGTAGAGGACACAC (SEQ ID NO: 32) with reverse-CACATTAAATTGTTCTAACATG-CAAG (SEQ ID NO: 33) and fragment F (predicted size 3.5 kb) was amplified using forward-CCTAGGTTATTTA-GAAGGGACTA (SEQ ID NO: 34) with reverse-GGT AGA TGT ATT GAC AGC AAT ATC (SEQ ID NO: 35).

[0233] The exact 5' and 3' ends of Bundibugyo ebolavirus were determined by 3' RACE from virus RNA extracted from virus infected Vero E6 cell monolayers using TriPure isolation reagent. RNAs were then polyadenylated in vitro using A-Plus poly(A) polymerase tailing kit (Epicenter Biotechnologies) following the manufacturer's instructions and then purified using an RNeasy kit (Qiagen) following standard protocols. Ten microliters of in vitro polyadenylated RNA were added as template in RT-PCR reactions, using SuperScript III One-Step RT-PCR system with Platinum Taq High Fidelity (Invitrogen) following the manufacturer's protocol. Two parallel RT-PCR reactions using the oligo(dT)-containing 3'RACE-AP primer (Invitrogen) mixed with 1 of 2 viral

specific primers, Ebo-U 692(-) ACAAAAAGCTATCTG-CACTAT (SEQ ID NO: 36) and Ebo-V18269(+) CTCA-GAAGCAAAATTAATGG (SEQ ID NO: 37), generated ~700 nt long fragments containing the 3' ends of either genomic and antigenomic RNAs. The resulting RT-PCR products were analyzed by agarose electrophoresis, and DNA bands of the correct sizes were purified using QIAquick Gel Extraction Kit (Qiagen) and sequenced using standard protocols (ABI).

[0234] The nucleotide sequence of the Côte d'Ivoire ebolavirus (EboIC) isolate RNA was initially determined using the exact same pyrosequencing strategy as that used for Bundibugyo ebolavirus described above. This method generated sequence for approximately 70% of the entire genome. This draft sequence was then used to design a whole genome primer walking strategy for filling any gaps and confirming the initial sequence. The following Côte d'Ivoire ebolavirus-specific primers were used to generate RT-PCR fragments, designated A-F, as follows: Fragment A (predicted size 3.0 kb) was amplified using forward-GTGTGCGAATAACTAT-GAGGAAG (SEQ ID NO: 38) and reverse-GTCTGTG-CAATGTTGATGAAGG (SEQ ID NO: 39); Fragment B (predicted size 3.2 kb) was amplified using forward-CAT-GAAAACCACACTCAACAAC (SEQ ID NO: 40) and reverse-GTTGCCTTAATCTTCATCAAGTTC (SEQ ID NO: 41); Fragment C (predicted size 3.0 kb) was amplified using forward-GGCTATAATGAATTTCTCCAG (SEQ ID NO: 42) and reverse-CAAGTGTAATTTGTGGTCTAGC (SEQ ID NO: 43); fragment D (predicted size 3.5 kb) was amplified using forward-GCTGGAATAGGAATCACAGG (SEQ ID NO: 44) and reverse-CGGTAGTCTACAGTTCTT-TAG (SEQ ID NO: 45); fragment E (predicted size 4.0 kb) was amplified using forward-GACAAAGAGATTAGATT-AGCTATAG (SEQ ID NO: 46) and reverse-GTAAT-GAGAAGGTGTCATTTGG (SEQ ID NO: 47); fragment F (predicted size 2.9 kb) was amplified using forward-CAC-GACTTAGTTGGACAATTGG (SEQ ID NO: 48) and reverse-CAGACACTAATTAGATCTGGAAG (SEQ ID NO: 49); fragment G (predicted size 1.3 kb) was amplified using forward-CGGACACACAAAAAGAAWRAA (SEQ ID NO: 50) and reverse-CGTTCTTGACCTTAGCAGTTTC (SEQ ID NO: 51); and fragment H (predicted size 2.5 kb) was amplified using forward-GCACTATAAGCTCGATGAAGTC (SEQ ID NO: 52) and reverse-TGGACACACAAAAARGA-RAA (SEQ ID NO: 53). A gap in the sequence contig was located between fragments C and D and this was resolved using the following primers to generate a predicted fragment of 1.5 kb: forward-CTGAGAGGATCCAGAAGAAAG (SEQ ID NO: 54) and reverse-GTGTAAGCGTTGATATAC-CTCC (SEQ ID NO: 55). The terminal ~20 nucleotides of the sequence were not experimentally determined but were inferred by comparing with the other known Ebola genome sequences.

[0235] Bundibugyo ebolavirus Real-Time RT-PCR Assay.

[0236] The primers and probe used in the Bundibugyo ebolavirus specific Q-RT-PCR assay were as follows: EboU965 (+): 5'-GAGAAAAGGCCTGTCTGGAGAA-3' (SEQ ID NO: 56), EboU1039(-): 5'-TCGGGTATTGAATCAGACCT-TGTT-3' (SEQ ID NO: 57) and EboU989 Prb: 5'-Fam-TTCAACGACAAATCCAAGTGCACGCA-3'-BHQ1 (SEQ ID NO: 58). Q-RT-PCR reactions were set up using Superscript III One-Step Q-RT-PCR (Invitrogen) according to the manufacturer's instructions and run for 40 cycles with a 58° C. annealing temperature.

[0237] Phylogenetic Analysis.

[0238] Modeltest 3.7¹⁸ was used to examine 56 models of nucleotide substitution to determine the model most appropriate for the data. The General Time Reversible model incorporating invariant sites and a gamma distribution (GTR+I+G) was selected using the Akaike Information Criterion (AIC). Nucleotide frequencies were A=0.3278, C=0.2101, G=0.1832, T=0.2789, the proportion of invariant sites=0.1412, and the gamma shape parameter=1.0593. A maximum likelihood analysis was subsequently performed in PAUP*4.0b10¹⁹ using the GTR+I+G model parameters. Bootstrap support values were used to assess topological support and were calculated based on 1,000 pseudoreplicates²⁰.

[0239] In addition, a Bayesian phylogenetic analysis was conducted in MrBayes 3.2²¹ using the GTR+I+G model of nucleotide substitution. Two simultaneous analyses, each with four Markov chains, were run for 5,000,000 generations sampling every 100 generations. Prior to termination of the run, the AWTY module was used to assess Markov Chain Monte Carlo convergence to ensure that the length of the analysis was sufficient²². Trees generated before the stabilization of the likelihood scores were discarded (burn in =40), and the remaining trees were used to construct a consensus tree. Nodal support was assessed by posterior probability values (>95=statistical support).

Example 2

Immunization against EboBun

[0240] To determine the capability of immunogens to elicit an immune response in non-human primates (NHP), 12 cynomolgus macaques, of which 10 are immunized with VSVΔG/EboBunGP either orally (OR; n=4), intranasally (IN; n=4) or intramuscularly (IM; n=2) in accordance with all animal control and safety guidelines and essentially as described by Qiu, X, et al., PLoS ONE. 2009; 4(5): e5547. The remaining 2 control animals are vaccinated intramuscularly with VSVΔG/MARVGP. VSVΔG/MARVGP does not provide heterologous protection against EboBun, therefore these NHPs succumb to EboBun infection. Animals are acclimatized for 14 days prior to infection. Animals are fed and monitored twice daily (pre- and post-infection) and fed commercial monkey chow, treats and fruit. Husbandry enrichment consists of commercial toys and visual stimulation.

[0241] The recombinant VSVΔG/EboBun vaccines are synthesized expressing the EboBun glycoprotein (GP) (SEQ ID NO: 9), soluble glycoprotein (sGP) (SEQ ID NO: 4), or nucleoprotein (NP) (SEQ ID NO: 3). Control VSVΔG/MARVGP vaccines represent the analogous proteins from Lake victoria marburgvirus (MARV) (strain Musoke). The following results for GP are similar for sGP and NP. Vaccines are generated using VSV (Indiana serotype) as described previously. Garbutt, M, et al., J Virol, 2004; 78(10):5458-5465; Schnell, M J, et al., PNAS USA, 1996; 93(21):11359-11365. EboBun challenge virus is passaged in Vero E6 cells prior to challenge, as described previously Jones, S M, et al., Nat Med. 2005; 11(7):786-790; Jahrling, P B, et al., J Infect Dis, 1999; 179 (Suppl 1):S224-34. An EboBun immunogen peptide pool consisting of 15mers with 11 amino acid overlaps (Sigma-Genosys) spanning the entire sequence of the EboBun immunogens and strain Mayinga 1976 GP are used.

[0242] Twelve filovirus naïve cynomolgus monkeys randomized into four groups receive 2 ml of 1×10^7 PFU/ml of vaccine in Dulbecco's modified Eagle's medium (DMEM).

Animals in the three experimental groups are vaccinated with either: 1) 2 ml orally (OR) (n=4); 2) 1 ml dripped into each nostril, intranasally (IN) (n=4); or 3) 1 ml each into two sites intramuscularly (IM) (n=2). The two controls are injected intramuscularly with 2 ml of 1×10^7 PFU/ml of VSVΔG/MARVGP. All animals are challenged intramuscularly 28 days later with 1,000 PFU of EboBun.

[0243] Routine examination is conducted on 0, 2, 4, 6, 10, 14 and 21 days post-vaccination, then 0, 3, 6, 10, 14, 19, 26 days, 6 and 9 months after the EboBun challenge. For the examinations animals are anaesthetized by intramuscular injection with 10 mg/kg of ketaset (Ayerst). Examinations include haematological analysis, monitoring temperature (rectal), respiration rate, lymph nodes, weight, hydration, discharges and mucous membranes. Also, swabs (throat, oral, nasal, rectal, vaginal) and blood samples are collected (4 ml from femoral vein, 1 ml in EDTA vacutainer tube; 3 ml in serum separator vacutainer tube). Cynomolgus monkey PBMCs are isolated using BD CPT sodium citrate Vacutainers (Becton Dickinson) as per manufacturer's protocol.

[0244] All VSVΔG/EboBunGP immunized animals are protected from high dose challenge. These animals show no evidence of clinical illness after vaccination or EboBun challenge. Both control animals demonstrate typical symptoms associated with EboBun HF including fever, macular rashes, lethargy, and unresponsiveness. Continued infection requires euthanization. Hematology analyses at each examination date demonstrate increases in the platelet-crit in the OR and IN groups post-challenge, however, no significant changes are observed in any NHPs post-immunization or in the VSVΔG/EboBunGP immunized NHPs post-challenge.

[0245] EboBun antibody production from humoral antibody response to vaccination and challenge is examined by a virus like particle (VLP) based ELISA assay. Generation of EboBun VLPs is performed by the protocol for ZEBOV as described by Wahl-Jensen, V., et al., J Virol, 2005; 79(4): 2413-2419. ELISA is performed by the protocol described by Qiu, X, et al., PLoS ONE. 2009; 4(5): e5547.

[0246] The VSVΔG/MARVGP immunized animals do not develop a detectable antibody response to EboBun. In contrast, potent antibody responses are detected in all VSVΔG/EboBunGP immunized animals independent of immunization route. Between days 14 and 21 post-vaccination, all VSVΔG/EboBunGP immunized NHPs develop high levels of IgA, IgM, and IgG against EboBunGP. After challenge the IgM titres do not exceed the post-vaccination levels, however, IgG and IgA antibody titres are increased peaking 14 days post-challenge then slowly decreasing before maintaining a relatively high antibody titre up to 9 months.

[0247] The level of neutralization antibodies is detected by a EboBun-GFP flow cytometric neutralization assay in serum collected at days 0 and 21 post-vaccination. Samples are assayed in duplicate for their ability to neutralize an infection with EboBun-GFP in VeroE6 cells. Serially diluted serum samples are incubated with an equal volume of EboBun-GFP in DMEM, at 37° C., 5% CO₂ for 1 hr followed by addition of 150 µl per well of a confluent 12 well plate of VeroE6 cells (MOI=0.0005). After 2 hours at 37° C., 5% CO₂, 1 ml of DMEM, 2% fetal bovine serym (FBS), 100 U/ml penicillin, 100 µg/ml streptomycin is added per well and incubated for 5 days. Cells are harvested by removing the culture supernatant, washing with 1 ml PBS, 0.04% EDTA, then adding 800 µl of PBS 0.04% EDTA for 5 minutes at 37° C. before adding 8 ml PBS, 4% paraformaldehyde (PFA) and overnight incu-

bation. The cells are acquired (10,000 events) and analyzed with CellQuest Pro v3.3 on a Becton Dickinson FACSCalibur flow cytometer.

[0248] The OR and IN routes produce EboBunGP-specific neutralizing antibodies with the OR route producing the highest titres post-vaccination. The IM immunization produces detectable levels of neutralizing antibody. In comparison, 3/4 NHPs in the OR group demonstrate a 50% reduction in EboBun-GFP positive cells at a titre of 1:40. Similarly, the IN route results in a reduction of EboBun-GFP positive cells at the 1:40 dilution.

[0249] EboBunGP-specific effector cellular immune responses are determined using IL-2 and IFN- γ ELISPOT assays as described by Qin, X, et al., PLoS ONE. 2009; 4(5): e5547 to determine the number of IL-2 and IFN- γ secreting lymphocytes. Prior to challenge on days 10 to 14 post-vaccination there is a detectable EboBun immunogen-specific IFN- γ response in all immunized animals. The IM route is the most potent, inducing approximately 2-fold more IFN- γ secreting cells than OR ($p < 0.001$) or IN ($p = 0.043$) routes. A strong post-challenge secondary IFN- γ response is induced in all VSVΔG/EboBun immunized animals with the IM route producing the most IFN- γ cells at day 6. By day 10 the OR group demonstrates a stronger response. The IFN- γ in the IN group rises steadily, peaking at day 26 post-challenge with 4.3 and 2 fold more EboBun specific IFN- γ secreting cells than the IM ($p = 0.003$) and OR ($p = 0.075$) group, respectively. All three routes produce strong EboBun-specific IFN- γ responses.

[0250] Post-vaccination, the IM group also has more EboBunGP-specific IL-2 secreting cells than either of the mucosally immunized groups. Post-challenge, the IM route continues to dominate early after challenge peaking on day 10. This difference shows a trend when compared to the IN group ($p = 0.067$) and is significant when compared to the OR group ($p < 0.001$). Additionally, the IN group has more IL-2 producing cells than the OR group ($p = 0.090$) on day 10 post-challenge. By day 26 post-challenge all three routes continue to produce a EboBunGP-specific IL-2 response, however, the IN group response is strongest. At day 26 post-challenge the IN group has the most potent IFN- γ and IL-2 responses, as well as the highest IgA and IgG antibody titre, indicating this immunization route, followed by a EboBun challenge, results in the development of potent and sustained effector responses.

[0251] Absolute lymphocyte numbers for CD3⁺, CD4⁺, and CD8⁺ (CD3⁺CD4⁺) T cell populations are determined by flow cytometry. No decrease is observed in the lymphocyte populations for any of the VSVΔG/EboBunGP vaccinated NHPs. In contrast, control animals who are not protected from EboBun show lymphocyte numbers decreased by 28-57%.

[0252] Macrophage numbers are slightly increased in control animals. However, the number of CD14⁺ cells is greater in the VSVΔG/EboBunGP vaccinated groups with the IM route showing the most significant increases.

[0253] In order to determine the long term immune response after challenge, EboBunGP-specific CD4⁺ and CD8⁺ memory T-lymphocytes are examined for their ability to proliferate (CFSE⁺) or produce IFN- γ in response to EboBunGP peptides at 6 months post-vaccination. EboBunGP-specific memory responses are observed as a result of vaccination followed by a ZEBOV challenge. These responses persist for at least 6 months. The memory popula-

tions in OR and IN inoculation routes demonstrate the greatest potential for proliferation and IFN- γ production post-challenge.

[0254] Any patents or publications mentioned in this specification are incorporated herein by reference to the same extent as if each individual publication is specifically and individually indicated to be incorporated by reference.

[0255] The compositions and methods described herein are presently representative of preferred embodiments, exemplary, and not intended as limitations on the scope of the invention. Changes therein and other uses will occur to those skilled in the art. Such changes and other uses can be made without departing from the scope of the invention as set forth in the claims. All numerical ranges are inclusive of the whole integers and decimals between the endpoints, and inclusive of the endpoints.

REFERENCES

- [0256]** 1. Suzuki, Y., and Gojobori, T., (1997) The origin and evolution of Ebola and Marburg viruses. *Mol Bio Evol*, 14(8): 800-806.
- [0257]** 2. Sanchez, A., Geisbert, T. W., Feldmann, H. in *Fields Virology* (ed. Knipe, D. M., Howley, P. M.) 1409-1448 (Lippincott Williams and Wilkins, Philadelphia, 2007).
- [0258]** 3. Leroy, E. M. et al., (2005) Fruit bats as reservoirs of Ebola virus. *Nature*, 438, 575-6.
- [0259]** 4. Towner, J. S. et al., (2007) Marburg virus infection detected in a common African bat. *PLoS ONE*, 2(8), e764.
- [0260]** 5. Swanepoel, R. et al., (2007) Studies of reservoir hosts for Marburg virus. *Emerg Infect Dis*, 13(12), 1847-51.
- [0261]** 6. Le Guenno, B. et al., (1995) Isolation and partial characterization of a new species of Ebola virus. *Lancet*, 345(8960), 1271-4.
- [0262]** 7. Ksiazek, T. G. et al. (1999) Clinical virology of Ebola hemorrhagic fever (EHF): virus, virus antigen, IgG and IgM antibody findings among EHF patients in Kikwit, 1995. *J. Infect Dis* 179 (suppl 1), S177-S187.
- [0263]** 8. Cox-Foster, D. L. et al. (2007) A metagenomic survey of microbes in honey bee colony collapse disorder. *Science* 318, 283-7.
- [0264]** 9. World Health Organization (2008) Ebola outbreak contained in Uganda. Features, 22 February, www.who.int/features/2008/ebola_outbreak/en/.
- [0265]** 10. Sullivan, N. J., Sanchez, A., Rollin, P. E., Yang, Z.-Y. & Nabel, G. J. (2000) Development of a preventive vaccine for Ebola virus infection in primates. *Nature* 408, 605-609.
- [0266]** 11. Ksiazek, T. G., West, C. P., Rollin, P. E., Jahrling, P. B. & Peters, C. J. (1999) ELISA for the detection of antibodies to Ebola viruses. *J. Infect Dis* 179 (suppl 1), S191-S198.
- [0267]** 12. Rodriguez, L. et al. (1999) Persistence and genetic stability of Ebola virus during the outbreak in Kikwit, Zaire 1995. *J. Infect Dis* 179 (suppl 1), S170-S176.
- [0268]** 13. Sanchez, A. et al. Detection and molecular characterization of Ebola viruses causing disease in human and nonhuman primates. *J. Infect Dis* 179 (suppl 1), S164-S169 (1999).
- [0269]** 14. Jones, S. M. et al. (2005) Live attenuated recombinant vaccine protects nonhuman primates against Ebola and Marburg viruses. *Nat Med* 11, 786-90.

- [0270] 15. Geisbert, T. W. et al. (2008) Recombinant vesicular stomatitis virus vector mediates postexposure protection against Sudan Ebola hemorrhagic fever in non-human primates. *J Virol* 82, 5664-8.
- [0271] 16. Towner, J. S., Sealy, T. K., Ksiazek, T. & Nichol, S. T. (2007) High-throughput molecular detection of hemorrhagic fever virus threats with applications for outbreak settings. *J. Inf Dis* 196 (suppl 2), S205-212.
- [0272] 17. Towner, J. S. et al. (2006) Marburgvirus genomics and association with a large hemorrhagic fever outbreak in Angola. *J Virol* 80, 6497-516.
- [0273] 18. Posada, D. & Crandall, K. A. (1998) MODELTEST: testing the model of DNA substitution. *Bioinformatics* 14, 817-818.
- [0274] 19. Swofford, D. L. (2002) PAUP*: phylogenetic analysis using parsimony (*and other methods) version 4.0b10. Sinauer Assoc., Sunderland, Mass.
- [0275] 20. Felsenstein, J. (1985) Confidence limits on phylogenies: an approach using the bootstrap. *Evolution* 39, 783-791.
- [0276] 21. Ronquist, F. & Huelsenbeck, J. P. (2003) MRBAYES 3: Bayesian phylogenetic inference under mixed models. *Bioinformatics* 19, 1572-1574.
- [0277] 22. Nylander, J. A. A., Wilgenbusch, J. C., Warren, D. L. & Swofford, D. L. (2008) AWTY (are we there yet?): a system for graphical exploration of MCMC convergence in Bayesian phylogenetics. *Bioinformatics* 24, 581-583.

 SEQUENCE LISTING

<160> NUMBER OF SEQ ID NOS: 59

<210> SEQ ID NO 1

<211> LENGTH: 18940

<212> TYPE: DNA

<213> ORGANISM: Bundibugyo ebolavirus

<220> FEATURE:

<221> NAME/KEY: misc_feature

<223> OTHER INFORMATION: Full viral sequence

<400> SEQUENCE: 1

```

cggacacaca aaaagaatga aggatattga atctttattg tgtgcgagta actacgagga      60
agattaaaga ttttctcttc attgaaattg aaattgagat tctaactctg acggatcgat      120
ccccaaatacc aacactgaga attggcctga agaagtcata tgcctccttg caaaaccaag      180
agcaggcccca aagggccatt aggccacatc tgcctgagcct gcagaacacg caggacttac      240
ttagcagaag agagcgcgtg ccgaaaccag ccaacaaatt gacacagctg ctactctgta      300
ccctgaattc ataacaata ttaagttgac aacagagata ctaatccaat atttggatca      360
agaatcaaaa tagtgaaacg actgactatc cctccttaga attagcaaaag atccttttgt      420
agactattgt gctacattct ctatccaaga cctcaaaatg gatcctcgtc caatcagaac      480
ctggatgatg cataacacat ctgaagtga agcagactac cataagattc taactgccgg      540
attgtccgtc cagcaaggca ttgtgagaca aagaatcatt cctgtttacc aaatctcaaa      600
cctggaggaa gtatgtcaac tcatcataca ggcatcgag gctggcgctg acttccagga      660
tagtgagatg agctttttgt taatgctatg tctgcatcat gcctatcaag gggattataa      720
acaatttttg gaaagtaatg cggtaaaata ccttgaaggt catggattcc gttttgagat      780
gaagaaaaag gaaggtgtca agcgcctgga ggaactactc cctgctgctc cgagtggaaa      840
gaacatcaag agaacattgg ctgcaatgcc cgaggaggaa acaacagaag caaatgctgg      900
acaattttct tcatctgcta gtctgtttct cccaaaattg gttgtcggag aaaaggcctg      960
tctggagaag gttcaacgac aaatccaagt gcacgcagaa caaggtctga ttcaataccc     1020
gacatcttgg caatcggtgg gacatatgat ggtcatcttc agactaatgc gaaccaactt     1080
cctgattaag ttctctctaa tacatcaagg aatgcatatg gttgcagggc atgatgctaa     1140
tgatgccgtc attgccaact ctgtagetca agctcgtttc tccgatttgt tgatagtcaa     1200
aacagtgtct gatcatatcc tccaaaaaac agagcacgga gttcgctctc atcccttggc     1260
gcgaacagcc aaagtcaaaa atgaggtgag ctcttttaag gcgcgtttag cctcactagc     1320

```

-continued

acaacatgga gaatatgccc cgtttgctcg tctgctgaat ctatctgggg ttaataatct	1380
tgagcatggg cttttccctc aactttctgc aattgctttg ggagtagcaa ctgcacatgg	1440
gagcactctg gctggagtca atgtaggaga gcaataccaa caactgcgag aagcagccac	1500
tgaggccgaa aagcagttgc agaaatatgc tgaatctcgt gaacttgatc acctaggtct	1560
tgatgatcag gaaaagaaaa tcctaaaaga cttccatcag aaaaagaatg agatcagctt	1620
ccagcagacg acagccatgg tcacactgcg gaaagagaga ttggccaaat tgaccgaagc	1680
tattacttcc acctctatcc tcaaacacgg aaggcggat gatgatgaca atgacatacc	1740
ctttccaggg ccaatcaatg ataacgagaa ctctggtcag aacgatgacg atccaacaga	1800
ctcccaggat accacaatcc cggatgtaat aatcgatcca aacgatggtg ggtataataa	1860
ttacagcgat tatgcaaatg atgctgcaag tgctcctgat gacctagttc tttttgacct	1920
tgaggacgag gatgatgctg ataaccggcg tcaaacacg ccagaaaaaa atgatagacc	1980
agcaacaaca aagctgagaa atggacagga ccaggatgga aaccaaggcg aaactgcac	2040
cccacgggta gcccccaacc aatacagaga caagccaatg ccacaagtac aggacagatc	2100
cgaaaatcat gacaaaaccc ttcaaacaca gtccagggtt ttgactccta tcagcgagga	2160
agcagacccc agcgaccaca acgatggtga caatgaaaag attcctcccc tggaatcaga	2220
cgacgagggt agcactgata ctactgcagc agaaacaaag cctgccactg cacctccgc	2280
tcccgctcac cgaagtatct ccgtagatga ttctgtcccc tcagagaaca tccccgcaca	2340
gtccaatcaa acgaacaatg aggacaatgt caggaacaat gctcagtcgg agcaatccat	2400
tgcagaaatg tatcaacata tcttgaaaac acaaggacct tttgatgcca tctttacta	2460
ccatgatgat aaagaagagc ccacatcttt cagcactagt gatgggaagg agtatacata	2520
tccagactct cttgaagatg agtatccacc ctggctcagc gagaaggaag ccatgaacga	2580
agacaataga ttcataacca tggatggtca gcagttttac tggcctgtga tgaatcatag	2640
aaataaatc atggcaatcc tccagcatca caggtgatcc gacctctaaa actgagctcc	2700
taactacaag ctaccccatc actctgccgg aatgccagaa cctccctcca aaacagctcc	2760
acatcgagaa cctccgacgc ggtacacagg caagacaggc aacctaatga tgttcctgtt	2820
caccacaac cgcaaccaac acttgatcga cttccaagac aactacaacc cccttagcca	2880
actccaccac agaagcacca ccccataaca acaaccccaa accaacaaca ctgcatgtaa	2940
gtattgtctc accccaagat gatccctgga caccaacaac ccctaacct ccccaagttg	3000
tcattaagaa aaaatatatg atgaagatta aaaccttcat cagagctatt tcttctacgc	3060
ttggttagga ccagtattca caaactatct tacaatccct acccaatatg acctctaaca	3120
gagcaagggt gacttacaac ccaccaccaa caaccacagg cacacgatcg tgtgggccgg	3180
aactttccgg gtggatctct gagcaattga tgacaggcaa gattccgatt accgatatct	3240
tcaatgaaat tgaaacctta cctagtataa gtcctcgtat ccaactccaa atcaaaaccc	3300
caagtgttca aacacgcagt gtccagaccc aaactgaccc aaattgtaat catgattttg	3360
cagaggttgt gaaaatgcta acatctctaa cccttgtcgt acaaaaacaa acccttgcaa	3420
ctgaatcact tgagcaacgc attactgacc tggaaggtag cctgaaacca gtgtctgaga	3480
tcaccaagat tgtttctgca ctaaatagat cctgtgcaga gatggtggcc aaatatgatc	3540
ttctagtaat gacgactggt cgtgcaactg ccaactgctgc agctactgaa gcatactggg	3600

-continued

cagaacatgg	acgtcctcca	ccggggccct	cattgtacga	ggaggatgca	atcaggacta	3660
aaattggaaa	acaaggggat	atggtaccca	aggaagtgc	agaggccttc	cgtaatctgg	3720
atagtactgc	ccttctaacg	gaagagaatt	ttgggaaacc	agacatatcc	gcaaaagact	3780
tcgcgaatat	catgtatgat	cacctcccag	gttttggcac	agcatttcat	caactagtgc	3840
aagtatatctg	caagttaggg	aaggacaatt	cctcacttga	tgtaattcat	gcagaatttc	3900
aggccagcct	tgetgaagga	gactctcttc	agtgtgccct	gattcagata	accaaacgga	3960
ttctatttt	ccaagatgca	gcaccaccg	taatccatat	tcggtcacgc	ggtgatatac	4020
caaaggcgtg	tcaaaagagc	ctccgccctg	ttccaccatc	accaaagatt	gataggggtt	4080
gggtatgcat	attccagcta	caagacggaa	aaacactcgg	actcaaaatc	taagggtgaac	4140
aattgcgcaa	cctccacagt	cgcctatatt	gcttccttcc	ggaatcaggg	tatgatcgcg	4200
taaaaaataa	gcttccaaca	tattgataca	cgatccatat	ccataatgcc	atctccagga	4260
atatgagaac	gcaaggccat	atcaggaccc	gatctcaatt	ccaatgcaac	ctactgttaa	4320
gaataaaaata	accaatgtcc	tctagcctta	tatgttctca	aaaatacaag	tgatgaagat	4380
taagaaaaag	catcctttac	ttgagaggag	ctaattcttt	atacttcac	taatctttaa	4440
gtaagtgtgat	cactaccacc	atgaggaggg	caattctacc	tactgcaccg	ccagaatata	4500
tagaggctgt	ctacccaatg	agaacgggta	gtactagtat	caacagtact	gccagtggtc	4560
cgaactttcc	agcaccggat	gtaatgatga	gtgatacacc	ctccaactca	ctccgaccaa	4620
ttgctgatga	taacatcgat	catccaagtc	atacaccaac	cagtgtttca	tcagccttta	4680
tactcgaggc	aatgggtgaat	gtgatatcgg	ggccgaaggt	actaatgaag	caaattccta	4740
tatggctccc	cttgggtggt	gctgatcaaa	aaacatatag	ttttgactca	actacagctg	4800
caattatgct	cgcacgtgac	accatcactc	actttggcaa	aacctccaat	ccgcttgtga	4860
gaatcaatcg	acttggctct	gggatccccg	atcacccggt	gcggcttcta	agaataggaa	4920
atcaagcett	cttgcaagag	tttgtgtgct	ctccagttca	attgccgcag	tatttcactt	4980
ttgacctgac	ggctctaaag	ctgatcactc	aacctctccc	ggcagcaacc	tggaacggatg	5040
atactccgac	cggctctaca	ggaatacttc	gtcctggaat	ttcctttcat	ccaaaactga	5100
gacctatcct	attgccaggg	aagaccggga	aaagaggatc	cagctccgat	cttacttctc	5160
ctgataaaat	acaagcaata	atgaactttc	tccaagacct	caaactcgtg	ccgattgatc	5220
cagccaagaa	cattatgggt	attgaagtgc	cggaaactct	gggccacaga	ctaactggaa	5280
agaaaatcac	aacaaaaaat	ggtcaaccaa	taattcctat	tcttctacca	aagtatatgt	5340
gcatggatcc	catttctcag	ggagacctca	caatgggtcat	cactcaagac	tgtgacactt	5400
gccattctcc	tgctagtctt	cctccagtca	gcgagaaatg	agcatgaagt	ccgaggtctc	5460
ccggcccaca	cgacccccag	ggccttcgtc	cggctaccga	accaaccatc	cgaccttcat	5520
caaaacccaa	aaataccgcc	acgcgaaagc	taaaatgcag	gaccacaatc	caaccagcaa	5580
caccatccat	acacagggat	caattgggct	gccgcagcat	atagacccaa	tagcaagctg	5640
ctgtccagaa	aatagttccg	gaaagtaact	caaccatcgc	aagcccaatg	cagctttcag	5700
aaatccgcca	gcaacccaac	tccactgtac	ccccaatatt	aacctgaatc	gactaaccgc	5760
actttaattt	gaagtacatt	tgttcaatgg	gttcattatt	aacagtgttg	cttttagatt	5820
gtacctttgc	tcacagatag	taaattgtta	tggtatcaaa	tcttattaag	aaaaagaaca	5880

-continued

cgatgaagat taacgcgacc tagagcgctg ccttcacatc atcaatttaa cttgtcaata	5940
gagcaaccta gtttgtgatt actcatcttc cgtagttgac aaacactttg ctgggttaatt	6000
gtaaataatc cacagtcac atggttacat caggaattct acaattgccc cgtgaacgct	6060
tcagaaaaac atcatttttt gtttggttaa taatccattt tcacaaagtt ttcctatcc	6120
cattgggagt agttcacaac aacactctcc aggttaagtga tatagataaa ttggtgtgcc	6180
gggataaact ttctccaca agtcagctga aatcggtcgg gcttaacta gaaggtaatg	6240
gagttgccac agatgtacca acagcaacga agagatggg attccgagct ggtgtccac	6300
ccaaagtgt gaactacgaa gctggggagt gggctgaaaa ctgctacaac ctggacatca	6360
agaaagcaga tggtagcgaa tgcctacctg aagcccctga ggggtgaaga ggcttccctc	6420
gctgccgtta tgtgcacaag gtttctgaa cagggcctg ccctgaaggt tacgctttcc	6480
acaaagaagg cgctttcttc ctgtatgatc gactggcatc aacaatcatc tatcgaagca	6540
ccacgttttc agaaggtgtt gtggctttct tgatcctccc cgaaactaaa aaggactttt	6600
tccaatcgcc accactacat gaaccggcca atatgacaac agaccatcc agctactacc	6660
acacagtcac acttaattat gtggctgaca attttgggac caatatgact aactttctgt	6720
ttcaagtga tcatctaact tatgtgcaac ttgaaccaag attcacacca caattttctg	6780
tccaactcaa tgagaccatt tatactaatt ggctcgag caaccaccaca ggaacactaa	6840
tttgaaagt aaatcctact gttgacacc gcgtaggtga atgggccttc tgggaaaata	6900
aaaaaacttc aaaaaaacc tttcaagtga agagctgtct gtcatatattg taccaagagc	6960
ccaggatcca ggcagcaacc agaagacgaa ggtcactccc accagcttcg ccaacaacca	7020
aacctccaag aaccacgaag acttgggttc agaggatccc gcttcagtgg ttcaagtgcg	7080
agacctccag agggaaaaca cagtgcgcac cccaccccca gacacagtcc ccacaactct	7140
gatccccgac acaatggagg aacaaaccac cagccactac gaaccacca acatttccag	7200
aaaccatcaa gagaggaaca acaccgcaca ccccgaaact ctgcaccaaca atccccaga	7260
caacacaacc ccgtcgacac cacctcaaga cggtagcgg acaagttccc acacaacacc	7320
ctccccccg ccagtcacca ccagcacaat ccaccccacc acacgagaga ctacattcc	7380
caccacaatg acaacaagcc atgacaccga cagcaatcga ccaacccaa ttgacatcag	7440
cgagtctaca gagccaggac cactcaccaa caccacaaga ggggctgcaa atctgctgac	7500
aggctcaaga agaaccggaa gggaaatcac cctgagaaca caagccaaat gcaacccaaa	7560
cctacactat tggacaaccc aagatgaagg ggctgccatt ggtttagcct ggatacctta	7620
cttcgggccc gcagcagagg gaatttatc ggaagggata atgcacaatc aaaatgggct	7680
aatttgccgg ttgagcgagc tagcaaatga gacgactcaa gccctacagt tattcttgcg	7740
tgctaccacg gaattgcga ctttctctat attgaatcga aaagccatcg actttttact	7800
ccaaagatgg ggaggaacgt gccacatctt agggccagat tgctgtattg agcccatga	7860
ttggactaag aacattactg acaaaataga tcaaatcatt catgatttca ttgataaacc	7920
tctaccagat caaacagata atgacaattg gtggacaggg tggaggcaat gggttcctgc	7980
cgggatcggg atcacggggg taataatcgc agttatagca ctgctgtgta ttgcaaatt	8040
tctactctaa tctagtcga ctctgtacca gcataatggc ctctaaaata agcttttgct	8100
tctgcttctc atagttaata catttcagca aaaatcaact attaagtcaa aagaagatcc	8160

-continued

ctctaataat cctaattacc ttcaaaaatc tagaacttta ttaattctca gggatatttag	8220
aacagccaga tgacttgact aagtttgtac tgtaataaaa agatacttga tgaagattaa	8280
gaaaaagaca gtcttgtgat tgtcactaat cttcatctca aaacatatta ttttaccaga	8340
agctactata gctacacctc ttgacacata gcaaacctta ctcagtgtga taattgtttg	8400
cctgctatatt acatatattac taacttacaa aattatcttg gggatttctc tgaacatata	8460
atcagaattg gcatttataa cacaagttag tcctaattgga ctcatttcat gagagagggc	8520
gtagcagaac tattcgacag agtgcaagag atgggcccag tcatacaagta agaacaagat	8580
cactctccag agacagccac cgcagcgaat atcatacacc taggagctct tcccaagttc	8640
gagtcccgac tgtgtttcat cggaagcgta ctgattcttt gacagttcca ccagcaccaa	8700
aggacatatg tcctacctta aggaaaggat ttttgtgtga cagcaatttt tgtaaaaagg	8760
accatcaact agaaagttaa acagataggg agctgctttt gctgattgca cggaaaacct	8820
gcggctccct tgaacaacaa ttgaacatca ctgctcctaa agatacacga ttagcaaatc	8880
caattgcaga tgatttccaa caaaaagacg gcccaaaaat tacactattg acacttttgg	8940
agactgcgga gtattggtca aaacaagata tcaagggcat tgatgactca agactaagag	9000
cattactaac cctttgtgcc gtcattgacga ggaaattctc aaaatcccag cttagtctat	9060
tgtgtgagag tcatctacga cgagaagggc taggacagga tcaatcagaa tctgttcttg	9120
aagtgtatca gcgcttatc agcgacaaag gcggaaattt tgaggcagcc ctatggcaac	9180
aatgggaccg acagtcttg atcatgttta taacagcatt tcttaattatt gctttacaat	9240
taccctgtga aagttcatct gttgttattt caggattaag gctgctagtg cctcaatcag	9300
aagataccga gacctcaacc tacaccgaga cacgtgcatg gtcagaggaa ggtggcccc	9360
attaacatct tccacagtcg aatctacat aatttcctta ttcaacgcag ataagaatca	9420
gtactaaacc acaagtgcga aaattaacaa aacaccagca taagtgaat cctgtctgtg	9480
attagcaaca cgaatgatct tcaatcctgt tgcaattcgc cagtataat tgtattcaca	9540
ttgtggccac aatatactgt cttttcccat tgaaaaataa ggctgaatct attacgtac	9600
acaaacttac aggattagca ccacgacggc tcaatactat acctattggt cacggctcga	9660
tgtgttaatc acttatattg tattcatttg aaattactca ttaggcaaat actttgatta	9720
agaaaaata attggaaaac cagaaaatcc ctaggatttt aaattcctat ctccggagat	9780
ccgagataat taatcaagca atgaggggac aatggtgaac aacaacatat tgttgcccc	9840
tttagattgg tcagttccaa aaacaagtga tgaagattaa tgcagatgac caaggaacac	9900
atatttgtga tttaaactgt ccagtttagac tctgttcaag gatcttcac tttttagct	9960
ccactctgag tcacaacata attgagtttt tgctcagaac agttatcagg attaaattct	10020
ctcaataaac tgaaactact agcatcactc tcaatttcat tacttacgac aatcattatc	10080
tttaataatat ttctctaaat tactgactta attagcttgt aatcagataa tatcgaaacc	10140
aatttatcat aaggcataat ttgtataagt gatttaggat ttaccccgaga agtgaaataa	10200
ttcttagaat aaaagaccga ctagaatatc cttaaggctg tctaacgtgc cacacagcta	10260
gggttagcct gacatctgga acaagatcga tactaatata gggatttggt tcatactagc	10320
tctctgcaaa cacaatggct aaggcaacag gtaggtacaa cttgggttca cctaaaaagg	10380
acctcgagag ggggcttggt ttgagtgatt tgtgcacgtt tttagttgat cagactatcc	10440

-continued

```

aggggtggcg ggtgacttgg gttgggattg aatttgacat cggccagaaa gggatggctc 10500
tactgcatcg gttaaaaact gctgacttcg ctctgcatg gtcgatgaca aggaatttat 10560
ttcttcattt atttcaaaat tcaaattcta ctattgagtc tccctctctg gcattacgag 10620
tgattctggc agctggtatt caagaccagt taattgacca atccttggtg gaaccgttgg 10680
cgggagccct gagcttagtc tccgattggc ttcttacaac aaacacaaac cattttcaaa 10740
tgcgcacgca gcacgctaaa gagcaactga gcttgaagat gctatcatta gtgcgctcta 10800
atatcttgaa attcatcagt caattggacg cactacatgt cgtgaactac aatggactct 10860
tgagcagtat cgaaattggc actagaaatc ataccattat catcacaaaga accaaccatgg 10920
gtttcctggt agaattacag gagcctgata aatctgccat gaatcaaaag aaaccaggac 10980
cagtcaagtt cctccctctg catgaatcaa ccttcaaggc tctaatcaaa aaaccgcgaa 11040
ctaagatgca ggcttgatt ctggaattta acagctccct ggcaatatag tccaacgcta 11100
ccaaccatca ttttttgtaa ctgcatctct tttatttcct ttctaacttg atacaattat 11160
aatcaagatc cctaaccct tttgacgaag tgggctaatt tttgctcatt ctaataataa 11220
atcataacct gaataaaaga caccacaata ttataaccca ataacaccta gagaatttct 11280
gaattgctaa agattatata ctgcgactaa gagacaagtt aatcaatctt tacttaataa 11340
tatactaaat gctagatagc tctggctaac taacctgagt tgtggattac tccttttaaa 11400
agtctatcaa ttttaagctta tcaactaatat taaggaggac tttttaataa agagcaagtg 11460
ttatgtagtc ttactaagaa tgatttgagg aagattaaga aaaagtgcct gtggggtcct 11520
tccgtttag aggcacacg agcaaacctc ttctctaat tttaatatgg caactcaaca 11580
tacacaatat ccagatgcaa gattatcttc acccattgtc ttagatcaat gtgatcttgt 11640
caccctgtct tgcggtctgt attcttcata ctcattaat cctcagttga aaaattgtag 11700
actacaaaa catatttacc gcctcaaatt tgatgctacg gttacaaaat ttttaagcga 11760
tgttccaata gttacattgc cgatagatta cttgacccct ttacttttac gaactttatc 11820
cggggagggc ttatgcctg tcgaacccaa gtgcagccaa ttcttagatg aaatagtaag 11880
ttatgttttg caggatgcac gttttttaag atactatctt aggcattgtg gagtacacga 11940
tgacaatgtt ggaaaaaatt ttgagccaaa gattaaggct ttgatttatg ataataaatt 12000
tctgcaacaa ttgttttatt ggtacgattt agcaatccta acgcgtagag ggcgcctgaa 12060
tcgaggggat aaccgttcaa catggtttgc aatgacgat ttaatagaca ttctcgggta 12120
cggtgattat attttctgga aaataccgtt gtcattgttg tcaactcaaca cagaggggat 12180
tcctcatgca gctaaggact ggtatcacgc atcaatcttc aaagaagcgg ttcaaggatc 12240
cacacatata gtgtcagttt ccactgcaga tgttttaatt atgtgtaagg acatcataac 12300
ctgtcgtttc aataccacac tcattgcagc attggcaaat ttagaagatt ctatctgttc 12360
tgactatcca caacctgaaa caatctctaa tctgtataag gcaggggatt acttaatctc 12420
gatactgggt tcagaagggt ataaggatcat aaagttttta gaaccactat gtttagctaa 12480
gatecaattg tgctcaaatt acactgagag gaaagggaga ttcttactc aatgcattt 12540
ggcgttaat cacacacttg aagaacttat tgagggccgg ggattgaagt cacaacaaga 12600
ctggaagatg agggaatttc accgaatctt agtaaattta aagtcaacac cacaacaact 12660
ctgtgaattg ttttcagtgc aaaagcattg ggggcacctc gtgctacata gcgagaaggc 12720

```

-continued

tattcagaaa	gtaaagaaac	atgcaaccgt	aataaaagca	ttgcgtcccg	taatcatctt	12780
tgagacatat	tgtgtgttca	agtacagcat	tgccaaacat	tattttgata	gccaaaggtc	12840
atggtatagt	gtaatctcag	ataaacatct	aacaccaggt	ttacactctt	acattaagag	12900
gaaccaatth	ccgccactgc	ctatgattaa	agacttattg	tgggaattct	atcaccttga	12960
tcatcctccc	ttattttcca	ccaagattat	tagtgacttg	agtattttca	ttaaggatcg	13020
cgctaccgca	gtggaaaaaa	catgttggga	tgcagttttc	gagcctaag	ttcttggata	13080
tagtcctcca	aacaagttct	caactaagag	ggttcctgaa	cagtttcttg	aacaagaaaa	13140
tttctcgatt	gatagtgttc	tcacttatgc	ccagcgcctg	gattatctac	ttccacaata	13200
ccggaattht	tctttctcac	ttaaggaaaa	agaattaaat	gtaggacgag	cttttggtaa	13260
gctaccttat	cctacacgta	atgttcaaac	tttatgtgaa	gccttatttg	cagatggatt	13320
agctaaagcc	tttctagta	acatgatggg	tgtaacagag	cgtgagcaga	aggaaagcct	13380
cttgaccag	gcgtcgtggc	accacacaag	tgaagatttc	ggtgagaatg	ccactgttag	13440
aggcagcagt	ttgtttaccg	acctagaaaa	atacaacttg	gcatttagat	atgagtttac	13500
agctccattt	attgaatact	gtaatcgatg	ttatgggtga	aaaaatttat	tcaattggat	13560
gcattatacg	ataccgcaat	gttatataca	tgtaatgtgat	tattataatc	ccctcatgg	13620
agtttcgcta	gaaaatcggg	aagatccccc	ggaaggccct	agctcttacc	gtggatcatct	13680
tgggggaatt	gagggactcc	aacagaaaact	ctggaccagc	atttcatgtg	cacaaatctc	13740
attagttgag	atcaagactg	gtttcaaatt	gagatctgcg	gtaatgggtg	ataatcaatg	13800
catcacagtt	ctttccgtat	ttcctctaga	gacagattcc	aatgagcaag	agcatagctc	13860
cgaggacaat	gctgctcgcg	tagcagccag	tttagccaaa	gtcacgagtg	cctgtggcat	13920
cttctctaaa	ccagatgaga	cttttgtgca	ttcaggcttt	atttatttcg	gtaagaagca	13980
atattttaat	ggcgttcaat	tgccacaatc	actcaagact	gctaccagga	ttgctccctt	14040
gtcagatgca	atctttgatg	accttcaggg	aactctggct	agtataggaa	cggeatttga	14100
gagatctata	tccgagacta	gacatgtata	cccttgccgg	gtggttgccg	cattccatac	14160
attctctctc	gttaggatcc	tccaatacca	ccaccttggg	ttcaacaaag	gaaccgatct	14220
aggtcaacta	tcactaagca	aaccgttgga	tttcggaact	atcactcttg	ctttagcggg	14280
acctcaagtt	ctaggagggt	tatcgthttt	aaaccagag	aatgtthttt	atcgcaacct	14340
tggagacccc	gtgacctccg	gcctattcca	acttaggact	tacctgcaa	tgatcaacat	14400
ggagacttta	tttctacctt	taattgcaa	gaacccggg	aactgtagt	caattgactt	14460
tgtactcaac	ccaagcggat	tgaatgtccc	tgggtcaca	gacctaacat	cttttttacg	14520
tcagatagtg	cgtagaaca	tcacattgag	tgcaaaaaat	aagctaata	acacattggt	14580
tcaactctca	gccgatttag	aagatgagat	ggtagttaa	tggctacttt	cttcaacacc	14640
tgtaatgagt	cggtttgctg	ctgatataat	ctctcgta	ccgagtggga	agcgttgca	14700
gatoctaggt	tattttaga	ggactagaac	cttgctagcc	tccaaagtca	tcaataacaa	14760
tgcagagact	cctattttag	ataggttgag	gaaaatcaca	ctgcagagat	ggagttttg	14820
gttttagctac	ctagaccact	gtgatcaggt	tctagcagat	gctttaata	aagtttcttg	14880
tcagattgat	ttggcgcaaa	ttttacgtga	atatacctgg	gcacacatac	tagagggaag	14940
acagctcatt	ggtgcaacac	ttccttgcat	gttagaaca	tttaattgtg	tttggctcaa	15000

-continued

atcgtagcaa	caatgcccta	aatgtgcaaa	atctagaaat	ccaaaaggag	agccatttgt	15060
gtcaattgca	attaagaaac	aagttgtgag	tgcatggccg	aatcagtcac	ggtaaattg	15120
gaccattggg	gacgggtgac	cttacatcgg	gtctcgaaca	gaggacaaga	ttgggcagcc	15180
agcaatcaag	cctaagtgtc	cctctgctgc	cttacgtgaa	gcaatagagt	tgacatctag	15240
actaacatgg	gttaccacaag	gtggtgccaa	tagtgatttg	ctagttaaac	cttttgtaga	15300
ggcacgagta	aacctgagtg	tgacaggagat	ccttcaaatg	acgccttctc	attattcagg	15360
gaacatcgta	catcggtata	atgaccaata	cagccctcat	tctttcatgg	caaatagaat	15420
gagtaattcc	gcgacgagat	tggtgggtgc	gacaaatact	ctcggggagt	tctcaggtgg	15480
ggggcaatca	gcaagggaca	gcaatatcat	ctttcaaat	gtaatcaatt	tttcggttgc	15540
cctatttgat	ttacgatttc	ggaacaccga	aacatcctcc	attcagcata	atcgtagcca	15600
tctccatctt	tcacagtgtt	gcacacggga	agtcacagct	caatacctaa	cctacacgtc	15660
tacgctttcc	ttgatctca	caaggtaccg	agagaatgag	ttaatttatg	ataacaatcc	15720
gttaaaagg	ggacttaatt	gcaacctatc	ctttgataat	ccacttttca	agggccaaaag	15780
gctcaatata	atagaggagg	atttgattag	atttcctcat	ctatctgggt	gggaacttgc	15840
gaaaaccatc	attcagtcca	ttatctcaga	cagcaataac	tcattccacag	acccatttag	15900
cagtggagaa	acacgatcat	tcacaactca	ctttctcaca	tatcctaagg	ttgggctcct	15960
ctatagtttc	ggcgccatcg	tgagttatta	cttagggaat	accattatta	ggacaaaaaa	16020
gctagacctc	agtcatttta	tgtattactt	aacaactcaa	atccataatt	tgccacatcg	16080
ctcgttgagg	atacttaagc	ccacctttaa	acatgttagt	gtgatataca	gactaatgag	16140
tattgatcct	catttttcaa	tctacatcgg	gggtacggca	ggtagatcag	ggctttcggga	16200
tgctaccaga	ctattccttc	gagtggccat	ttcttccttc	cttcaattta	tcaaaaaatg	16260
gatcggtgaa	tacaagacag	ctattcctct	gtgggttata	taccctttgg	agggacaaaa	16320
tccagatcca	attaatagct	ttctacatct	gattatagcc	ttactgcaaa	atgaatcccc	16380
tcaaaacaac	atccaattcc	aagaagacag	aaataatcaa	cagttgtccg	ataatctagt	16440
ttacatgtgc	aagagcactg	ccagtaattt	cttccatgca	tcacttgccct	attggaggag	16500
ccggcacaaa	ggacggccca	aaaatcgatc	gaccgaagaa	cagacagtta	aaccataacc	16560
atatgataat	tttcattctg	ttaaatgtgc	ctcaaaccce	ccaagcatcc	ccaaatctaa	16620
gtcaggaaat	caaggttcaa	gcgcattttt	tgagaaactt	gaatatgata	aagaaagaga	16680
attgccaaca	gcttccacac	cagccgaaca	atccaagacc	tatatcaagg	ccctatccag	16740
ccgaatttat	catggtaaaa	caccatccaa	tgccgcacaa	gatgattcaa	caacctccaa	16800
gggctgcgat	tccaaagaag	aaaatgccgt	tcaagcttca	caccgaattg	tcttaccatt	16860
ttttacattg	tcacagaacg	actacagaac	tcctcagct	aaaaagtcag	agtatataac	16920
tgaaatcacc	aaactaatte	gacaattaaa	ggcaattcca	gataccactg	tatactgtcg	16980
ctttacaggg	gttgatctct	caatgcatta	taagcttgat	gaggttctct	gggaattcga	17040
tagtttcaaa	actgctgtga	ctctagctga	aggagaaggg	tcaggtgcct	tattactact	17100
acaaaaatat	aaggtcagaa	caatcttttt	taacacttta	gctacagagc	atagcatcga	17160
ggcagaaata	gtttctggga	caaccacacc	tcgaatgtct	cttctgttaa	tgccaaaact	17220
tcgatgatgat	caaataaatg	taatattaaa	caattctgct	agccagggtta	ctgatatacc	17280

-continued

```

taacctgca tggttcactg accagaaatc tagaatcccc acacaagttg agattatgac 17340
tatggatgct gaaacgacag aaaatattaa tcgggtcaaaa ttatatgagg ctattcagca 17400
attaattggt tcacacattg atacaagggg gctaaagatt gttattataa aggttttttt 17460
aagtgatatt gaaggctccc tgtggcttaa tgaccatctt gcccccttat tcggatccgg 17520
ctatttaatt aaacctatta cttcgagtcc aaagtcaagc gaatgggtact tatgtctttc 17580
aaatttcctt tcagcctctc gacgacggcc tcatcagggt catgctacct gtatgcaagt 17640
catccaaaca gcgctacgac tccaagttca aaggagttca tactggctta gccatttagt 17700
gcaatatgct gatattaatt tgcacttgag ttatgttaat ttgggtttcc cttcattgga 17760
aaaggttctt taccatcgat ataacctagt tgattcacgg aagggtccac tggctctgat 17820
cctttacat ttaacacact tgcaagcaga gattagagaa ttagtggtg actataatca 17880
gcaacgacaa agtgcgaacc aaacatacca cttcatcaaa acgacaaagg gccggattac 17940
aaaattagtc aatgactacc ttaaatttta tctcgtagtg caagcactga agcataattg 18000
tctttggcag gaagaactca gaacacttcc tgacttaac aatgtttgca atcgatttta 18060
ccatataagg gactgctcat gtgaagatcg atttttaatt caaactcttt acttaacccg 18120
tatgcaagac tcagaagcaa aattaatgga gagattaacc gggtttctag gattgtatcc 18180
taatggtatt aacgcttaag atccccttag aggcacgca atatgactcc aaacattaaa 18240
tgatattgct gtcaatacat ctacctgacc gagagcaagg tttattataa aaaacctata 18300
cacatgactg caatgcgtaa tttataccga aacacagtga gggctgcaca tgcaggttcc 18360
tgttgagctt taaaagatca tgcaatataa aatgatattt gtataactaat catgttagta 18420
ctaactaaca gtactcactg catatactct atcaattaag aaaaattact gtggtttatg 18480
catttaaatg acatcacaga tggatataat atagttaatt cttacctaaa tgttgagtta 18540
tagtaatttg aagttataat tatgattagt gcttatacta taaataatag ctataccaag 18600
tatacacaag aagttatgat tttgtattca aattatatc acaggaactt gtgattaata 18660
ataaaagtct cagttgttgg ttgttgagtt gtaaaactcc cgttaaaaat ttattttcca 18720
cttataacta ataataatca tagatcagta tgagttgagg ctattcaaac cttagaaaaa 18780
ttgtgcgatg tttttacca tgtcaatctt gatttcaatg atattggagg gcttgctgat 18840
aaattcagta attaacatta agtcagtgtg gaacctcatt ggatatttga tcgtacacaa 18900
aatatcttta caaaattggt ttctcttttt tgtgtgtcca 18940

```

```

<210> SEQ ID NO 2
<211> LENGTH: 2210
<212> TYPE: PRT
<213> ORGANISM: Bundibugyo ebolavirus
<220> FEATURE:
<221> NAME/KEY: misc_feature
<223> OTHER INFORMATION: Bundibugyo ebolavirus L viral protein

```

```

<400> SEQUENCE: 2

```

```

Met Ala Thr Gln His Thr Gln Tyr Pro Asp Ala Arg Leu Ser Ser Pro
1           5           10           15

Ile Val Leu Asp Gln Cys Asp Leu Val Thr Arg Ala Cys Gly Leu Tyr
20          25          30

Ser Ser Tyr Ser Leu Asn Pro Gln Leu Lys Asn Cys Arg Leu Pro Lys
35          40          45

```

-continued

His	Ile	Tyr	Arg	Leu	Lys	Phe	Asp	Ala	Thr	Val	Thr	Lys	Phe	Leu	Ser	50	55	60
Asp	Val	Pro	Ile	Val	Thr	Leu	Pro	Ile	Asp	Tyr	Leu	Thr	Pro	Leu	Leu	65	70	75
Leu	Arg	Thr	Leu	Ser	Gly	Glu	Gly	Leu	Cys	Pro	Val	Glu	Pro	Lys	Cys	85	90	95
Ser	Gln	Phe	Leu	Asp	Glu	Ile	Val	Ser	Tyr	Val	Leu	Gln	Asp	Ala	Arg	100	105	110
Phe	Leu	Arg	Tyr	Tyr	Phe	Arg	His	Val	Gly	Val	His	Asp	Asp	Asn	Val	115	120	125
Gly	Lys	Asn	Phe	Glu	Pro	Lys	Ile	Lys	Ala	Leu	Ile	Tyr	Asp	Asn	Glu	130	135	140
Phe	Leu	Gln	Gln	Leu	Phe	Tyr	Trp	Tyr	Asp	Leu	Ala	Ile	Leu	Thr	Arg	145	150	155
Arg	Gly	Arg	Leu	Asn	Arg	Gly	Asn	Asn	Arg	Ser	Thr	Trp	Phe	Ala	Asn	165	170	175
Asp	Asp	Leu	Ile	Asp	Ile	Leu	Gly	Tyr	Gly	Asp	Tyr	Ile	Phe	Trp	Lys	180	185	190
Ile	Pro	Leu	Ser	Leu	Leu	Ser	Leu	Asn	Thr	Glu	Gly	Ile	Pro	His	Ala	195	200	205
Ala	Lys	Asp	Trp	Tyr	His	Ala	Ser	Ile	Phe	Lys	Glu	Ala	Val	Gln	Gly	210	215	220
His	Thr	His	Ile	Val	Ser	Val	Ser	Thr	Ala	Asp	Val	Leu	Ile	Met	Cys	225	230	235
Lys	Asp	Ile	Ile	Thr	Cys	Arg	Phe	Asn	Thr	Thr	Leu	Ile	Ala	Ala	Leu	245	250	255
Ala	Asn	Leu	Glu	Asp	Ser	Ile	Cys	Ser	Asp	Tyr	Pro	Gln	Pro	Glu	Thr	260	265	270
Ile	Ser	Asn	Leu	Tyr	Lys	Ala	Gly	Asp	Tyr	Leu	Ile	Ser	Ile	Leu	Gly	275	280	285
Ser	Glu	Gly	Tyr	Lys	Val	Ile	Lys	Phe	Leu	Glu	Pro	Leu	Cys	Leu	Ala	290	295	300
Lys	Ile	Gln	Leu	Cys	Ser	Asn	Tyr	Thr	Glu	Arg	Lys	Gly	Arg	Phe	Leu	305	310	315
Thr	Gln	Met	His	Leu	Ala	Val	Asn	His	Thr	Leu	Glu	Glu	Leu	Ile	Glu	325	330	335
Gly	Arg	Gly	Leu	Lys	Ser	Gln	Gln	Asp	Trp	Lys	Met	Arg	Glu	Phe	His	340	345	350
Arg	Ile	Leu	Val	Asn	Leu	Lys	Ser	Thr	Pro	Gln	Gln	Leu	Cys	Glu	Leu	355	360	365
Phe	Ser	Val	Gln	Lys	His	Trp	Gly	His	Pro	Val	Leu	His	Ser	Glu	Lys	370	375	380
Ala	Ile	Gln	Lys	Val	Lys	Lys	His	Ala	Thr	Val	Ile	Lys	Ala	Leu	Arg	385	390	395
Pro	Val	Ile	Ile	Phe	Glu	Thr	Tyr	Cys	Val	Phe	Lys	Tyr	Ser	Ile	Ala	405	410	415
Lys	His	Tyr	Phe	Asp	Ser	Gln	Gly	Ser	Trp	Tyr	Ser	Val	Ile	Ser	Asp	420	425	430
Lys	His	Leu	Thr	Pro	Gly	Leu	His	Ser	Tyr	Ile	Lys	Arg	Asn	Gln	Phe	435	440	445

-continued

Pro	Pro	Leu	Pro	Met	Ile	Lys	Asp	Leu	Leu	Trp	Glu	Phe	Tyr	His	Leu
450					455						460				
Asp	His	Pro	Pro	Leu	Phe	Ser	Thr	Lys	Ile	Ile	Ser	Asp	Leu	Ser	Ile
465					470					475					480
Phe	Ile	Lys	Asp	Arg	Ala	Thr	Ala	Val	Glu	Lys	Thr	Cys	Trp	Asp	Ala
				485					490					495	
Val	Phe	Glu	Pro	Asn	Val	Leu	Gly	Tyr	Ser	Pro	Pro	Asn	Lys	Phe	Ser
			500					505					510		
Thr	Lys	Arg	Val	Pro	Glu	Gln	Phe	Leu	Glu	Gln	Glu	Asn	Phe	Ser	Ile
		515					520					525			
Asp	Ser	Val	Leu	Thr	Tyr	Ala	Gln	Arg	Leu	Asp	Tyr	Leu	Leu	Pro	Gln
	530					535					540				
Tyr	Arg	Asn	Phe	Ser	Phe	Ser	Leu	Lys	Glu	Lys	Glu	Leu	Asn	Val	Gly
545					550					555					560
Arg	Ala	Phe	Gly	Lys	Leu	Pro	Tyr	Pro	Thr	Arg	Asn	Val	Gln	Thr	Leu
				565					570					575	
Cys	Glu	Ala	Leu	Leu	Ala	Asp	Gly	Leu	Ala	Lys	Ala	Phe	Pro	Ser	Asn
			580					585					590		
Met	Met	Val	Val	Thr	Glu	Arg	Glu	Gln	Lys	Glu	Ser	Leu	Leu	His	Gln
		595					600					605			
Ala	Ser	Trp	His	His	Thr	Ser	Asp	Asp	Phe	Gly	Glu	Asn	Ala	Thr	Val
	610					615					620				
Arg	Gly	Ser	Ser	Phe	Val	Thr	Asp	Leu	Glu	Lys	Tyr	Asn	Leu	Ala	Phe
625					630					635					640
Arg	Tyr	Glu	Phe	Thr	Ala	Pro	Phe	Ile	Glu	Tyr	Cys	Asn	Arg	Cys	Tyr
				645					650					655	
Gly	Val	Lys	Asn	Leu	Phe	Asn	Trp	Met	His	Tyr	Thr	Ile	Pro	Gln	Cys
		660					665						670		
Tyr	Ile	His	Val	Ser	Asp	Tyr	Tyr	Asn	Pro	Pro	His	Gly	Val	Ser	Leu
	675						680					685			
Glu	Asn	Arg	Glu	Asp	Pro	Pro	Glu	Gly	Pro	Ser	Ser	Tyr	Arg	Gly	His
	690				695						700				
Leu	Gly	Gly	Ile	Glu	Gly	Leu	Gln	Gln	Lys	Leu	Trp	Thr	Ser	Ile	Ser
705					710					715					720
Cys	Ala	Gln	Ile	Ser	Leu	Val	Glu	Ile	Lys	Thr	Gly	Phe	Lys	Leu	Arg
				725					730					735	
Ser	Ala	Val	Met	Gly	Asp	Asn	Gln	Cys	Ile	Thr	Val	Leu	Ser	Val	Phe
		740					745						750		
Pro	Leu	Glu	Thr	Asp	Ser	Asn	Glu	Gln	Glu	His	Ser	Ser	Glu	Asp	Asn
		755					760					765			
Ala	Ala	Arg	Val	Ala	Ala	Ser	Leu	Ala	Lys	Val	Thr	Ser	Ala	Cys	Gly
	770				775						780				
Ile	Phe	Leu	Lys	Pro	Asp	Glu	Thr	Phe	Val	His	Ser	Gly	Phe	Ile	Tyr
785					790					795					800
Phe	Gly	Lys	Lys	Gln	Tyr	Leu	Asn	Gly	Val	Gln	Leu	Pro	Gln	Ser	Leu
				805					810					815	
Lys	Thr	Ala	Thr	Arg	Ile	Ala	Pro	Leu	Ser	Asp	Ala	Ile	Phe	Asp	Asp
			820					825					830		
Leu	Gln	Gly	Thr	Leu	Ala	Ser	Ile	Gly	Thr	Ala	Phe	Glu	Arg	Ser	Ile
	835						840					845			
Ser	Glu	Thr	Arg	His	Val	Tyr	Pro	Cys	Arg	Val	Val	Ala	Ala	Phe	His

-continued

850	855	860
Thr Phe Phe Ser Val Arg Ile Leu Gln Tyr His His Leu Gly Phe Asn 865 870 875 880		
Lys Gly Thr Asp Leu Gly Gln Leu Ser Leu Ser Lys Pro Leu Asp Phe 885 890 895		
Gly Thr Ile Thr Leu Ala Leu Ala Val Pro Gln Val Leu Gly Gly Leu 900 905 910		
Ser Phe Leu Asn Pro Glu Lys Cys Phe Tyr Arg Asn Leu Gly Asp Pro 915 920 925		
Val Thr Ser Gly Leu Phe Gln Leu Arg Thr Tyr Leu Gln Met Ile Asn 930 935 940		
Met Asp Asp Leu Phe Leu Pro Leu Ile Ala Lys Asn Pro Gly Asn Cys 945 950 955 960		
Ser Ala Ile Asp Phe Val Leu Asn Pro Ser Gly Leu Asn Val Pro Gly 965 970 975		
Ser Gln Asp Leu Thr Ser Phe Leu Arg Gln Ile Val Arg Arg Thr Ile 980 985 990		
Thr Leu Ser Ala Lys Asn Lys Leu Ile Asn Thr Leu Phe His Ser Ser 995 1000 1005		
Ala Asp Leu Glu Asp Glu Met Val Cys Lys Trp Leu Leu Ser Ser 1010 1015 1020		
Thr Pro Val Met Ser Arg Phe Ala Ala Asp Ile Phe Ser Arg Thr 1025 1030 1035		
Pro Ser Gly Lys Arg Leu Gln Ile Leu Gly Tyr Leu Glu Gly Thr 1040 1045 1050		
Arg Thr Leu Leu Ala Ser Lys Val Ile Asn Asn Asn Ala Glu Thr 1055 1060 1065		
Pro Ile Leu Asp Arg Leu Arg Lys Ile Thr Leu Gln Arg Trp Ser 1070 1075 1080		
Leu Trp Phe Ser Tyr Leu Asp His Cys Asp Gln Val Leu Ala Asp 1085 1090 1095		
Ala Leu Ile Lys Val Ser Cys Thr Val Asp Leu Ala Gln Ile Leu 1100 1105 1110		
Arg Glu Tyr Thr Trp Ala His Ile Leu Glu Gly Arg Gln Leu Ile 1115 1120 1125		
Gly Ala Thr Leu Pro Cys Met Leu Glu Gln Phe Asn Val Phe Trp 1130 1135 1140		
Leu Lys Ser Tyr Glu Gln Cys Pro Lys Cys Ala Lys Ser Arg Asn 1145 1150 1155		
Pro Lys Gly Glu Pro Phe Val Ser Ile Ala Ile Lys Lys Gln Val 1160 1165 1170		
Val Ser Ala Trp Pro Asn Gln Ser Arg Leu Asn Trp Thr Ile Gly 1175 1180 1185		
Asp Gly Val Pro Tyr Ile Gly Ser Arg Thr Glu Asp Lys Ile Gly 1190 1195 1200		
Gln Pro Ala Ile Lys Pro Lys Cys Pro Ser Ala Ala Leu Arg Glu 1205 1210 1215		
Ala Ile Glu Leu Thr Ser Arg Leu Thr Trp Val Thr Gln Gly Gly 1220 1225 1230		
Ala Asn Ser Asp Leu Leu Val Lys Pro Phe Val Glu Ala Arg Val 1235 1240 1245		

Asn	Leu	Ser	Val	Gln	Glu	Ile	Leu	Gln	Met	Thr	Pro	Ser	His	Tyr
	1250					1255					1260			
Ser	Gly	Asn	Ile	Val	His	Arg	Tyr	Asn	Asp	Gln	Tyr	Ser	Pro	His
	1265					1270					1275			
Ser	Phe	Met	Ala	Asn	Arg	Met	Ser	Asn	Ser	Ala	Thr	Arg	Leu	Val
	1280					1285					1290			
Val	Ser	Thr	Asn	Thr	Leu	Gly	Glu	Phe	Ser	Gly	Gly	Gly	Gln	Ser
	1295					1300					1305			
Ala	Arg	Asp	Ser	Asn	Ile	Ile	Phe	Gln	Asn	Val	Ile	Asn	Phe	Ser
	1310					1315					1320			
Val	Ala	Leu	Phe	Asp	Leu	Arg	Phe	Arg	Asn	Thr	Glu	Thr	Ser	Ser
	1325					1330					1335			
Ile	Gln	His	Asn	Arg	Ala	His	Leu	His	Leu	Ser	Gln	Cys	Cys	Thr
	1340					1345					1350			
Arg	Glu	Val	Pro	Ala	Gln	Tyr	Leu	Thr	Tyr	Thr	Ser	Thr	Leu	Ser
	1355					1360					1365			
Leu	Asp	Leu	Thr	Arg	Tyr	Arg	Glu	Asn	Glu	Leu	Ile	Tyr	Asp	Asn
	1370					1375					1380			
Asn	Pro	Leu	Lys	Gly	Gly	Leu	Asn	Cys	Asn	Leu	Ser	Phe	Asp	Asn
	1385					1390					1395			
Pro	Leu	Phe	Lys	Gly	Gln	Arg	Leu	Asn	Ile	Ile	Glu	Glu	Asp	Leu
	1400					1405					1410			
Ile	Arg	Phe	Pro	His	Leu	Ser	Gly	Trp	Glu	Leu	Ala	Lys	Thr	Ile
	1415					1420					1425			
Ile	Gln	Ser	Ile	Ile	Ser	Asp	Ser	Asn	Asn	Ser	Ser	Thr	Asp	Pro
	1430					1435					1440			
Ile	Ser	Ser	Gly	Glu	Thr	Arg	Ser	Phe	Thr	Thr	His	Phe	Leu	Thr
	1445					1450					1455			
Tyr	Pro	Lys	Val	Gly	Leu	Leu	Tyr	Ser	Phe	Gly	Ala	Ile	Val	Ser
	1460					1465					1470			
Tyr	Tyr	Leu	Gly	Asn	Thr	Ile	Ile	Arg	Thr	Lys	Lys	Leu	Asp	Leu
	1475					1480					1485			
Ser	His	Phe	Met	Tyr	Tyr	Leu	Thr	Thr	Gln	Ile	His	Asn	Leu	Pro
	1490					1495					1500			
His	Arg	Ser	Leu	Arg	Ile	Leu	Lys	Pro	Thr	Phe	Lys	His	Val	Ser
	1505					1510					1515			
Val	Ile	Ser	Arg	Leu	Met	Ser	Ile	Asp	Pro	His	Phe	Ser	Ile	Tyr
	1520					1525					1530			
Ile	Gly	Gly	Thr	Ala	Gly	Asp	Arg	Gly	Leu	Ser	Asp	Ala	Thr	Arg
	1535					1540					1545			
Leu	Phe	Leu	Arg	Val	Ala	Ile	Ser	Ser	Phe	Leu	Gln	Phe	Ile	Lys
	1550					1555					1560			
Lys	Trp	Ile	Val	Glu	Tyr	Lys	Thr	Ala	Ile	Pro	Leu	Trp	Val	Ile
	1565					1570					1575			
Tyr	Pro	Leu	Glu	Gly	Gln	Asn	Pro	Asp	Pro	Ile	Asn	Ser	Phe	Leu
	1580					1585					1590			
His	Leu	Ile	I											

-continued

Leu Val 1625	Tyr Met Cys Lys Ser 1630	Thr Ala Ser Asn Phe 1635	Phe His Ala
Ser Leu 1640	Ala Tyr Trp Arg Ser 1645	Arg His Lys Gly Arg 1650	Pro Lys Asn
Arg Ser 1655	Thr Glu Glu Gln Thr 1660	Val Lys Pro Ile Pro 1665	Tyr Asp Asn
Phe His 1670	Ser Val Lys Cys Ala 1675	Ser Asn Pro Pro Ser 1680	Ile Pro Lys
Ser Lys 1685	Ser Gly Thr Gln Gly 1690	Ser Ser Ala Phe Phe 1695	Glu Lys Leu
Glu Tyr 1700	Asp Lys Glu Arg Glu 1705	Leu Pro Thr Ala Ser 1710	Thr Pro Ala
Glu Gln 1715	Ser Lys Thr Tyr Ile 1720	Lys Ala Leu Ser Ser 1725	Arg Ile Tyr
His Gly 1730	Lys Thr Pro Ser Asn 1735	Ala Ala Lys Asp Asp 1740	Ser Thr Thr
Ser Lys 1745	Gly Cys Asp Ser Lys 1750	Glu Glu Asn Ala Val 1755	Gln Ala Ser
His Arg 1760	Ile Val Leu Pro Phe 1765	Phe Thr Leu Ser Gln 1770	Asn Asp Tyr
Arg Thr 1775	Pro Ser Ala Lys Lys 1780	Ser Glu Tyr Ile Thr 1785	Glu Ile Thr
Lys Leu 1790	Ile Arg Gln Leu Lys 1795	Ala Ile Pro Asp Thr 1800	Thr Val Tyr
Cys Arg 1805	Phe Thr Gly Val Val 1810	Ser Ser Met His Tyr 1815	Lys Leu Asp
Glu Val 1820	Leu Trp Glu Phe Asp 1825	Ser Phe Lys Thr Ala 1830	Val Thr Leu
Ala Glu 1835	Gly Glu Gly Ser Gly 1840	Ala Leu Leu Leu Leu 1845	Gln Lys Tyr
Lys Val 1850	Arg Thr Ile Phe Phe 1855	Asn Thr Leu Ala Thr 1860	Glu His Ser
Ile Glu 1865	Ala Glu Ile Val Ser 1870	Gly Thr Thr Thr Pro 1875	Arg Met Leu
Leu Pro 1880	Val Met Ala Lys Leu 1885	His Asp Asp Gln Ile 1890	Asn Val Ile
Leu Asn 1895	Asn Ser Ala Ser Gln 1900	Val Thr Asp Ile Thr 1905	Asn Pro Ala
Trp Phe 1910	Thr Asp Gln Lys Ser 1915	Arg Ile Pro Thr Gln 1920	Val Glu Ile
Met Thr 1925	Met Asp Ala Glu Thr 1930	Thr Glu Asn Ile Asn 1935	Arg Ser Lys
Leu Tyr 1940	Glu Ala Ile Gln Gln 1945	Leu Ile Val Ser His 1950	Ile Asp Thr
Arg Val 1955	Leu Lys Ile Val Ile 1960	Ile Lys Val Phe Leu 1965	Ser Asp Ile
Glu Gly 1970	Leu Leu Trp Leu Asn 1975	Asp His Leu Ala Pro 1980	Leu Phe Gly
Ser Gly 1985	Tyr Leu Ile Lys Pro 1990	Ile Thr Ser Ser Pro 1995	Lys Ser Ser
Glu Trp	Tyr Leu Cys Leu Ser	Asn Phe Leu Ser Ala	Ser Arg Arg

-continued

2000	2005	2010
Arg Pro His Gln Gly His	Ala Thr Cys Met Gln Val	Ile Gln Thr
2015	2020	2025
Ala Leu Arg Leu Gln Val	Gln Arg Ser Ser Tyr Trp	Leu Ser His
2030	2035	2040
Leu Val Gln Tyr Ala Asp	Ile Asn Leu His Leu Ser	Tyr Val Asn
2045	2050	2055
Leu Gly Phe Pro Ser Leu	Glu Lys Val Leu Tyr His	Arg Tyr Asn
2060	2065	2070
Leu Val Asp Ser Arg Lys	Gly Pro Leu Val Ser Ile	Leu Tyr His
2075	2080	2085
Leu Thr His Leu Gln Ala	Glu Ile Arg Glu Leu Val	Cys Asp Tyr
2090	2095	2100
Asn Gln Gln Arg Gln Ser	Arg Thr Gln Thr Tyr His	Phe Ile Lys
2105	2110	2115
Thr Thr Lys Gly Arg Ile	Thr Lys Leu Val Asn Asp	Tyr Leu Lys
2120	2125	2130
Phe Tyr Leu Val Val Gln	Ala Leu Lys His Asn Cys	Leu Trp Gln
2135	2140	2145
Glu Glu Leu Arg Thr Leu	Pro Asp Leu Ile Asn Val	Cys Asn Arg
2150	2155	2160
Phe Tyr His Ile Arg Asp	Cys Ser Cys Glu Asp Arg	Phe Leu Ile
2165	2170	2175
Gln Thr Leu Tyr Leu Thr	Arg Met Gln Asp Ser Glu	Ala Lys Leu
2180	2185	2190
Met Glu Arg Leu Thr Gly	Phe Leu Gly Leu Tyr Pro	Asn Gly Ile
2195	2200	2205
Asn Ala		
2210		

<210> SEQ ID NO 3
 <211> LENGTH: 739
 <212> TYPE: PRT
 <213> ORGANISM: Bundibugyo ebolavirus
 <220> FEATURE:
 <221> NAME/KEY: misc_feature
 <223> OTHER INFORMATION: Bundibugyo ebolavirus NP viral protein

<400> SEQUENCE: 3

Met Asp Pro Arg Pro Ile Arg Thr Trp Met Met His Asn Thr Ser Glu		
1	5	10 15
Val Glu Ala Asp Tyr His Lys Ile Leu Thr Ala Gly Leu Ser Val Gln		
	20	25 30
Gln Gly Ile Val Arg Gln Arg Ile Ile Pro Val Tyr Gln Ile Ser Asn		
	35	40 45
Leu Glu Glu Val Cys Gln Leu Ile Ile Gln Ala Phe Glu Ala Gly Val		
	50	55 60
Asp Phe Gln Asp Ser Ala Asp Ser Phe Leu Leu Met Leu Cys Leu His		
	65	70 75 80
His Ala Tyr Gln Gly Asp Tyr Lys Gln Phe Leu Glu Ser Asn Ala Val		
	85	90 95
Lys Tyr Leu Glu Gly His Gly Phe Arg Phe Glu Met Lys Lys Lys Glu		
	100	105 110

-continued

Gly	Val	Lys	Arg	Leu	Glu	Glu	Leu	Leu	Pro	Ala	Ala	Ser	Ser	Gly	Lys
		115					120					125			
Asn	Ile	Lys	Arg	Thr	Leu	Ala	Ala	Met	Pro	Glu	Glu	Glu	Thr	Thr	Glu
	130					135					140				
Ala	Asn	Ala	Gly	Gln	Phe	Leu	Ser	Phe	Ala	Ser	Leu	Phe	Leu	Pro	Lys
145					150					155					160
Leu	Val	Val	Gly	Glu	Lys	Ala	Cys	Leu	Glu	Lys	Val	Gln	Arg	Gln	Ile
			165						170					175	
Gln	Val	His	Ala	Glu	Gln	Gly	Leu	Ile	Gln	Tyr	Pro	Thr	Ser	Trp	Gln
		180						185					190		
Ser	Val	Gly	His	Met	Met	Val	Ile	Phe	Arg	Leu	Met	Arg	Thr	Asn	Phe
		195					200					205			
Leu	Ile	Lys	Phe	Leu	Leu	Ile	His	Gln	Gly	Met	His	Met	Val	Ala	Gly
	210					215					220				
His	Asp	Ala	Asn	Asp	Ala	Val	Ile	Ala	Asn	Ser	Val	Ala	Gln	Ala	Arg
225					230					235					240
Phe	Ser	Gly	Leu	Leu	Ile	Val	Lys	Thr	Val	Leu	Asp	His	Ile	Leu	Gln
			245						250					255	
Lys	Thr	Glu	His	Gly	Val	Arg	Leu	His	Pro	Leu	Ala	Arg	Thr	Ala	Lys
		260					265						270		
Val	Lys	Asn	Glu	Val	Ser	Ser	Phe	Lys	Ala	Ala	Leu	Ala	Ser	Leu	Ala
		275					280					285			
Gln	His	Gly	Glu	Tyr	Ala	Pro	Phe	Ala	Arg	Leu	Leu	Asn	Leu	Ser	Gly
	290					295					300				
Val	Asn	Asn	Leu	Glu	His	Gly	Leu	Phe	Pro	Gln	Leu	Ser	Ala	Ile	Ala
305					310					315					320
Leu	Gly	Val	Ala	Thr	Ala	His	Gly	Ser	Thr	Leu	Ala	Gly	Val	Asn	Val
			325						330					335	
Gly	Glu	Gln	Tyr	Gln	Gln	Leu	Arg	Glu	Ala	Ala	Thr	Glu	Ala	Glu	Lys
		340						345					350		
Gln	Leu	Gln	Lys	Tyr	Ala	Glu	Ser	Arg	Glu	Leu	Asp	His	Leu	Gly	Leu
		355					360					365			
Asp	Asp	Gln	Glu	Lys	Lys	Ile	Leu	Lys	Asp	Phe	His	Gln	Lys	Lys	Asn
	370					375					380				
Glu	Ile	Ser	Phe	Gln	Gln	Thr	Thr	Ala	Met	Val	Thr	Leu	Arg	Lys	Glu
385					390					395					400
Arg	Leu	Ala	Lys	Leu	Thr	Glu	Ala	Ile	Thr	Ser	Thr	Ser	Ile	Leu	Lys
			405						410					415	
Thr	Gly	Arg	Arg	Tyr	Asp	Asp	Asp	Asn	Asp	Ile	Pro	Phe	Pro	Gly	Pro
		420						425					430		
Ile	Asn	Asp	Asn	Glu	Asn	Ser	Gly	Gln	Asn	Asp	Asp	Asp	Pro	Thr	Asp
		435					440					445			
Ser	Gln	Asp	Thr	Thr	Ile	Pro	Asp	Val	Ile	Ile	Asp	Pro	Asn	Asp	Gly
		450				455					460				
Gly	Tyr	Asn	Asn	Tyr	Ser	Asp	Tyr	Ala	Asn	Asp	Ala	Ala	Ser	Ala	Pro
465				470						475					480
Asp	Asp	Leu	Val	Leu	Phe	Asp	Leu	Glu	Asp	Glu	Asp	Asp	Ala	Asp	Asn
			485					490					495		
Pro	Ala	Gln	Asn	Thr	Pro	Glu	Lys	Asn	Asp	Arg	Pro	Ala	Thr	Thr	Lys
		500						505					510		
Leu	Arg	Asn	Gly	Gln	Asp	Gln	Asp	Gly	Asn	Gln	Gly	Glu	Thr	Ala	Ser

-continued

515					520					525					
Pro	Arg	Val	Ala	Pro	Asn	Gln	Tyr	Arg	Asp	Lys	Pro	Met	Pro	Gln	Val
530					535					540					
Gln	Asp	Arg	Ser	Glu	Asn	His	Asp	Gln	Thr	Leu	Gln	Thr	Gln	Ser	Arg
545					550					555					560
Val	Leu	Thr	Pro	Ile	Ser	Glu	Glu	Ala	Asp	Pro	Ser	Asp	His	Asn	Asp
				565					570					575	
Gly	Asp	Asn	Glu	Ser	Ile	Pro	Pro	Leu	Glu	Ser	Asp	Asp	Glu	Gly	Ser
			580					585					590		
Thr	Asp	Thr	Thr	Ala	Ala	Glu	Thr	Lys	Pro	Ala	Thr	Ala	Pro	Pro	Ala
			595				600					605			
Pro	Val	Tyr	Arg	Ser	Ile	Ser	Val	Asp	Asp	Ser	Val	Pro	Ser	Glu	Asn
	610					615					620				
Ile	Pro	Ala	Gln	Ser	Asn	Gln	Thr	Asn	Asn	Glu	Asp	Asn	Val	Arg	Asn
625					630					635					640
Asn	Ala	Gln	Ser	Glu	Gln	Ser	Ile	Ala	Glu	Met	Tyr	Gln	His	Ile	Leu
				645					650					655	
Lys	Thr	Gln	Gly	Pro	Phe	Asp	Ala	Ile	Leu	Tyr	Tyr	His	Met	Met	Lys
			660					665					670		
Glu	Glu	Pro	Ile	Ile	Phe	Ser	Thr	Ser	Asp	Gly	Lys	Glu	Tyr	Thr	Tyr
		675					680					685			
Pro	Asp	Ser	Leu	Glu	Asp	Glu	Tyr	Pro	Pro	Trp	Leu	Ser	Glu	Lys	Glu
	690					695					700				
Ala	Met	Asn	Glu	Asp	Asn	Arg	Phe	Ile	Thr	Met	Asp	Gly	Gln	Gln	Phe
705					710					715					720
Tyr	Trp	Pro	Val	Met	Asn	His	Arg	Asn	Lys	Phe	Met	Ala	Ile	Leu	Gln
				725					730					735	

His His Arg

<210> SEQ ID NO 4
 <211> LENGTH: 373
 <212> TYPE: PRT
 <213> ORGANISM: Bundibugyo ebolavirus
 <220> FEATURE:
 <221> NAME/KEY: misc_feature
 <223> OTHER INFORMATION: Bundibugyo ebolavirus SGP viral protein

<400> SEQUENCE: 4

Met	Val	Thr	Ser	Gly	Ile	Leu	Gln	Leu	Pro	Arg	Glu	Arg	Phe	Arg	Lys
1				5					10					15	
Thr	Ser	Phe	Phe	Val	Trp	Val	Ile	Ile	Leu	Phe	His	Lys	Val	Phe	Pro
		20					25					30			
Ile	Pro	Leu	Gly	Val	Val	His	Asn	Asn	Thr	Leu	Gln	Val	Ser	Asp	Ile
	35					40						45			
Asp	Lys	Leu	Val	Cys	Arg	Asp	Lys	Leu	Ser	Ser	Thr	Ser	Gln	Leu	Lys
	50					55					60				
Ser	Val	Gly	Leu	Asn	Leu	Glu	Gly	Asn	Gly	Val	Ala	Thr	Asp	Val	Pro
65				70					75					80	
Thr	Ala	Thr	Lys	Arg	Trp	Gly	Phe	Arg	Ala	Gly	Val	Pro	Pro	Lys	Val
			85					90						95	
Val	Asn	Tyr	Glu	Ala	Gly	Glu	Trp	Ala	Glu	Asn	Cys	Tyr	Asn	Leu	Asp
	100						105						110		
Ile	Lys	Lys	Ala	Asp	Gly	Ser	Glu	Cys	Leu	Pro	Glu	Ala	Pro	Glu	Gly

-continued

115	120	125
Val Arg Gly Phe Pro Arg Cys Arg Tyr Val His Lys Val Ser Gly Thr		
130	135	140
Gly Pro Cys Pro Glu Gly Tyr Ala Phe His Lys Glu Gly Ala Phe Phe		
145	150	155
Leu Tyr Asp Arg Leu Ala Ser Thr Ile Ile Tyr Arg Ser Thr Thr Phe		
165	170	175
Ser Glu Gly Val Val Ala Phe Leu Ile Leu Pro Glu Thr Lys Lys Asp		
180	185	190
Phe Phe Gln Ser Pro Pro Leu His Glu Pro Ala Asn Met Thr Thr Asp		
195	200	205
Pro Ser Ser Tyr Tyr His Thr Val Thr Leu Asn Tyr Val Ala Asp Asn		
210	215	220
Phe Gly Thr Asn Met Thr Asn Phe Leu Phe Gln Val Asp His Leu Thr		
225	230	235
Tyr Val Gln Leu Glu Pro Arg Phe Thr Pro Gln Phe Leu Val Gln Leu		
245	250	255
Asn Glu Thr Ile Tyr Thr Asn Gly Arg Arg Ser Asn Thr Thr Gly Thr		
260	265	270
Leu Ile Trp Lys Val Asn Pro Thr Val Asp Thr Gly Val Gly Glu Trp		
275	280	285
Ala Phe Trp Glu Asn Lys Lys Thr Ser Gln Lys Pro Phe Gln Val Lys		
290	295	300
Ser Cys Leu Ser Tyr Leu Tyr Gln Glu Pro Arg Ile Gln Ala Ala Thr		
305	310	315
Arg Arg Arg Arg Ser Leu Pro Pro Ala Ser Pro Thr Thr Lys Pro Pro		
325	330	335
Arg Thr Thr Lys Thr Trp Phe Gln Arg Ile Pro Leu Gln Trp Phe Lys		
340	345	350
Cys Glu Thr Ser Arg Gly Lys Thr Gln Cys Arg Pro His Pro Gln Thr		
355	360	365
Gln Ser Pro Gln Leu		
370		

<210> SEQ ID NO 5
 <211> LENGTH: 251
 <212> TYPE: PRT
 <213> ORGANISM: Bundibugyo ebolavirus
 <220> FEATURE:
 <221> NAME/KEY: misc_feature
 <223> OTHER INFORMATION: Bundibugyo ebolavirus VP24 viral protein

<400> SEQUENCE: 5

Met Ala Lys Ala Thr Gly Arg Tyr Asn Leu Val Ser Pro Lys Lys Asp
1 5 10 15
Leu Glu Arg Gly Leu Val Leu Ser Asp Leu Cys Thr Phe Leu Val Asp
20 25 30
Gln Thr Ile Gln Gly Trp Arg Val Thr Trp Val Gly Ile Glu Phe Asp
35 40 45
Ile Ala Gln Lys Gly Met Ala Leu Leu His Arg Leu Lys Thr Ala Asp
50 55 60
Phe Ala Pro Ala Trp Ser Met Thr Arg Asn Leu Phe Pro His Leu Phe
65 70 75 80

Gln	Asn	Ser	Asn	Ser	Thr	Ile	Glu	Ser	Pro	Leu	Trp	Ala	Leu	Arg	Val
			85						90				95		
Ile	Leu	Ala	Ala	Gly	Ile	Gln	Asp	Gln	Leu	Ile	Asp	Gln	Ser	Leu	Val
		100						105					110		
Glu	Pro	Leu	Ala	Gly	Ala	Leu	Ser	Leu	Val	Ser	Asp	Trp	Leu	Leu	Thr
		115					120					125			
Thr	Asn	Thr	Asn	His	Phe	Gln	Met	Arg	Thr	Gln	His	Ala	Lys	Glu	Gln
	130					135					140				
Leu	Ser	Leu	Lys	Met	Leu	Ser	Leu	Val	Arg	Ser	Asn	Ile	Leu	Lys	Phe
145					150					155					160
Ile	Ser	Gln	Leu	Asp	Ala	Leu	His	Val	Val	Asn	Tyr	Asn	Gly	Leu	Leu
				165					170					175	
Ser	Ser	Ile	Glu	Ile	Gly	Thr	Arg	Asn	His	Thr	Ile	Ile	Ile	Thr	Arg
		180						185					190		
Thr	Asn	Met	Gly	Phe	Leu	Val	Glu	Leu	Gln	Glu	Pro	Asp	Lys	Ser	Ala
		195					200					205			
Met	Asn	Gln	Lys	Lys	Pro	Gly	Pro	Val	Lys	Phe	Ser	Leu	Leu	His	Glu
	210					215					220				
Ser	Thr	Phe	Lys	Ala	Leu	Ile	Lys	Lys	Pro	Ala	Thr	Lys	Met	Gln	Ala
225					230					235					240
Leu	Ile	Leu	Glu	Phe	Asn	Ser	Ser	Leu	Ala	Ile					
				245					250						
<210> SEQ ID NO 6															
<211> LENGTH: 289															
<212> TYPE: PRT															
<213> ORGANISM: Bundibugyo ebolavirus															
<220> FEATURE:															
<221> NAME/KEY: misc_feature															
<223> OTHER INFORMATION: Bundibugyo ebolavirus VP30 viral protein															
<400> SEQUENCE: 6															
Met	Asp	Ser	Phe	His	Glu	Arg	Gly	Arg	Ser	Arg	Thr	Ile	Arg	Gln	Ser
1				5					10					15	
Ala	Arg	Asp	Gly	Pro	Ser	His	Gln	Val	Arg	Thr	Arg	Ser	Ser	Ser	Arg
			20					25					30		
Asp	Ser	His	Arg	Ser	Glu	Tyr	His	Thr	Pro	Arg	Ser	Ser	Ser	Gln	Val
		35					40					45			
Arg	Val	Pro	Thr	Val	Phe	His	Arg	Lys	Arg	Thr	Asp	Ser	Leu	Thr	Val
	50					55					60				
Pro	Pro	Ala	Pro	Lys	Asp	Ile	Cys	Pro	Thr	Leu	Arg	Lys	Gly	Phe	Leu
65			</												

-continued

Met Thr Arg Lys Phe Ser Lys Ser Gln Leu Ser Leu Leu Cys Glu Ser
 180 185 190
 His Leu Arg Arg Glu Gly Leu Gly Gln Asp Gln Ser Glu Ser Val Leu
 195 200 205
 Glu Val Tyr Gln Arg Leu His Ser Asp Lys Gly Gly Asn Phe Glu Ala
 210 215 220
 Ala Leu Trp Gln Gln Trp Asp Arg Gln Ser Leu Ile Met Phe Ile Thr
 225 230 235 240
 Ala Phe Leu Asn Ile Ala Leu Gln Leu Pro Cys Glu Ser Ser Ser Val
 245 250 255
 Val Ile Ser Gly Leu Arg Leu Leu Val Pro Gln Ser Glu Asp Thr Glu
 260 265 270
 Thr Ser Thr Tyr Thr Glu Thr Arg Ala Trp Ser Glu Glu Gly Gly Pro
 275 280 285
 His

 <210> SEQ ID NO 7
 <211> LENGTH: 341
 <212> TYPE: PRT
 <213> ORGANISM: Bundibugyo ebolavirus
 <220> FEATURE:
 <221> NAME/KEY: misc_feature
 <223> OTHER INFORMATION: Bundibugyo ebolavirus VP35 viral protein

 <400> SEQUENCE: 7

 Met Thr Ser Asn Arg Ala Arg Val Thr Tyr Asn Pro Pro Pro Thr Thr
 1 5 10 15
 Thr Gly Thr Arg Ser Cys Gly Pro Glu Leu Ser Gly Trp Ile Ser Glu
 20 25 30
 Gln Leu Met Thr Gly Lys Ile Pro Ile Thr Asp Ile Phe Asn Glu Ile
 35 40 45
 Glu Thr Leu Pro Ser Ile Ser Pro Ser Ile His Ser Lys Ile Lys Thr
 50 55 60
 Pro Ser Val Gln Thr Arg Ser Val Gln Thr Gln Thr Asp Pro Asn Cys
 65 70 75 80
 Asn His Asp Phe Ala Glu Val Val Lys Met Leu Thr Ser Leu Thr Leu
 85 90 95
 Val Val Gln Lys Gln Thr Leu Ala Thr Glu Ser Leu Glu Gln Arg Ile
 100 105 110
 Thr Asp Leu Glu Gly Ser Leu Lys Pro Val Ser Glu Ile Thr Lys Ile
 115 120 125
 Val Ser Ala Leu Asn Arg Ser Cys Ala Glu Met Val Ala Lys Tyr Asp
 130 135 140
 Leu Leu Val Met Thr Thr Gly Arg Ala Thr Ala Thr Ala Ala Thr
 145 150 155 160
 Glu Ala Tyr Trp Ala Glu His Gly Arg Pro Pro Pro Gly Pro Ser Leu
 165 170 175
 Tyr Glu Glu Asp Ala Ile Arg Thr Lys Ile Gly Lys Gln Gly Asp Met
 180 185 190
 Val Pro Lys Glu Val Gln Glu Ala Phe Arg Asn Leu Asp Ser Thr Ala
 195 200 205
 Leu Leu Thr Glu Glu Asn Phe Gly Lys Pro Asp Ile Ser Ala Lys Asp
 210 215 220

-continued

```

Leu Arg Asn Ile Met Tyr Asp His Leu Pro Gly Phe Gly Thr Ala Phe
225                230                235                240

His Gln Leu Val Gln Val Ile Cys Lys Leu Gly Lys Asp Asn Ser Ser
                245                250                255

Leu Asp Val Ile His Ala Glu Phe Gln Ala Ser Leu Ala Glu Gly Asp
                260                265                270

Ser Pro Gln Cys Ala Leu Ile Gln Ile Thr Lys Arg Ile Pro Ile Phe
                275                280                285

Gln Asp Ala Ala Pro Pro Val Ile His Ile Arg Ser Arg Gly Asp Ile
                290                295                300

Pro Lys Ala Cys Gln Lys Ser Leu Arg Pro Val Pro Pro Ser Pro Lys
305                310                315                320

Ile Asp Arg Gly Trp Val Cys Ile Phe Gln Leu Gln Asp Gly Lys Thr
                325                330                335

Leu Gly Leu Lys Ile
                340

<210> SEQ ID NO 8
<211> LENGTH: 326
<212> TYPE: PRT
<213> ORGANISM: Bundibugyo ebolavirus
<220> FEATURE:
<221> NAME/KEY: misc_feature
<223> OTHER INFORMATION: Bundibugyo ebolavirus VP40 viral protein

<400> SEQUENCE: 8

Met Arg Arg Ala Ile Leu Pro Thr Ala Pro Pro Glu Tyr Ile Glu Ala
1                5                10                15

Val Tyr Pro Met Arg Thr Val Ser Thr Ser Ile Asn Ser Thr Ala Ser
                20                25                30

Gly Pro Asn Phe Pro Ala Pro Asp Val Met Met Ser Asp Thr Pro Ser
                35                40                45

Asn Ser Leu Arg Pro Ile Ala Asp Asp Asn Ile Asp His Pro Ser His
50                55                60

Thr Pro Thr Ser Val Ser Ser Ala Phe Ile Leu Glu Ala Met Val Asn
65                70                75                80

Val Ile Ser Gly Pro Lys Val Leu Met Lys Gln Ile Pro Ile Trp Leu
                85                90                95

Pro Leu Gly Val Ala Asp Gln Lys Thr Tyr Ser Phe Asp Ser Thr Thr
100                105                110

Ala Ala Ile Met Leu Ala Ser Tyr Thr Ile Thr His Phe Gly Lys Thr
115                120                125

Ser Asn Pro Leu Val Arg Ile Asn Arg Leu Gly Pro Gly Ile Pro Asp
130                135                140

His Pro Leu Arg Leu Leu Arg Ile Gly Asn Gln Ala Phe Leu Gln Glu
145                150                155                160

Phe Val Leu Pro Pro Val Gln Leu Pro Gln Tyr Phe Thr Phe Asp Leu
165                170                175

Thr Ala Leu Lys Leu Ile Thr Gln Pro Leu Pro Ala Ala Thr Trp Thr
180                185                190

Asp Asp Thr Pro Thr Gly Pro Thr Gly Ile Leu Arg Pro Gly Ile Ser
195                200                205

Phe His Pro Lys Leu Arg Pro Ile Leu Leu Pro Gly Lys Thr Gly Lys

```

-continued

210	215	220
Arg Gly Ser Ser Ser	Asp Leu Thr Ser Pro	Asp Lys Ile Gln Ala Ile
225	230	235 240
Met Asn Phe Leu Gln	Asp Leu Lys Leu Val	Pro Ile Asp Pro Ala Lys
245	250	255
Asn Ile Met Gly Ile	Glu Val Pro Glu Leu Leu	Val His Arg Leu Thr
260	265	270
Gly Lys Lys Ile Thr	Thr Lys Asn Gly Gln Pro	Ile Ile Pro Ile Leu
275	280	285
Leu Pro Lys Tyr Ile	Gly Met Asp Pro Ile Ser	Gln Gly Asp Leu Thr
290	295	300
Met Val Ile Thr Gln	Asp Cys Asp Thr Cys His	Ser Pro Ala Ser Leu
305	310	315 320
Pro Pro Val Ser Glu	Lys	
	325	

<210> SEQ ID NO 9
 <211> LENGTH: 676
 <212> TYPE: PRT
 <213> ORGANISM: Bundibugyo ebolavirus
 <220> FEATURE:
 <221> NAME/KEY: misc_feature
 <223> OTHER INFORMATION: Bundibugyo ebolavirus GP viral protein

<400> SEQUENCE: 9

Met Val Thr Ser Gly	Ile Leu Gln Leu Pro	Arg Glu Arg Phe Arg Lys
1	5	10 15
Thr Ser Phe Phe Val	Trp Val Ile Ile Leu Phe	His Lys Val Phe Pro
20	25	30
Ile Pro Leu Gly Val	Val His Asn Asn Thr	Leu Gln Val Ser Asp Ile
35	40	45
Asp Lys Leu Val Cys	Arg Asp Lys Leu Ser	Ser Thr Ser Gln Leu Lys
50	55	60
Ser Val Gly Leu Asn	Leu Glu Gly Asn Gly	Val Ala Thr Asp Val Pro
65	70	75 80
Thr Ala Thr Lys Arg	Trp Gly Phe Arg Ala	Gly Val Pro Pro Lys Val
85	90	95
Val Asn Tyr Glu Ala	Gly Glu Trp Ala Glu	Asn Cys Tyr Asn Leu Asp
100	105	110
Ile Lys Lys Ala Asp	Gly Ser Glu Cys Leu	Pro Glu Ala Pro Glu Gly
115	120	125
Val Arg Gly Phe Pro	Arg Cys Arg Tyr Val	His Lys Val Ser Gly Thr
130	135	140
Gly Pro Cys Pro Glu	Gly Tyr Ala Phe His	Lys Glu Gly Ala Phe Phe
145	150	155 160
Leu Tyr Asp Arg Leu	Ala Ser Thr Ile Ile	Tyr Arg Ser Thr Thr Phe
165	170	175
Ser Glu Gly Val Val	Ala Phe Leu Ile Leu	Pro Glu Thr Lys Lys Asp
180	185	190
Phe Phe Gln Ser Pro	Pro Leu His Glu Pro	Ala Asn Met Thr Thr Asp
195	200	205
Pro Ser Ser Tyr Tyr	His Thr Val Thr Leu	Asn Tyr Val Ala Asp Asn
210	215	220

-continued

Phe	Gly	Thr	Asn	Met	Thr	Asn	Phe	Leu	Phe	Gln	Val	Asp	His	Leu	Thr	225	230	235	240
Tyr	Val	Gln	Leu	Glu	Pro	Arg	Phe	Thr	Pro	Gln	Phe	Leu	Val	Gln	Leu	245	250	255	
Asn	Glu	Thr	Ile	Tyr	Thr	Asn	Gly	Arg	Arg	Ser	Asn	Thr	Thr	Gly	Thr	260	265	270	
Leu	Ile	Trp	Lys	Val	Asn	Pro	Thr	Val	Asp	Thr	Gly	Val	Gly	Glu	Trp	275	280	285	
Ala	Phe	Trp	Glu	Asn	Lys	Lys	Asn	Phe	Thr	Lys	Thr	Leu	Ser	Ser	Glu	290	295	300	
Glu	Leu	Ser	Val	Ile	Phe	Val	Pro	Arg	Ala	Gln	Asp	Pro	Gly	Ser	Asn	305	310	315	320
Gln	Lys	Thr	Lys	Val	Thr	Pro	Thr	Ser	Phe	Ala	Asn	Asn	Gln	Thr	Ser	325	330	335	
Lys	Asn	His	Glu	Asp	Leu	Val	Pro	Glu	Asp	Pro	Ala	Ser	Val	Val	Gln	340	345	350	
Val	Arg	Asp	Leu	Gln	Arg	Glu	Asn	Thr	Val	Pro	Thr	Pro	Pro	Pro	Asp	355	360	365	
Thr	Val	Pro	Thr	Thr	Leu	Ile	Pro	Asp	Thr	Met	Glu	Glu	Gln	Thr	Thr	370	375	380	
Ser	His	Tyr	Glu	Pro	Pro	Asn	Ile	Ser	Arg	Asn	His	Gln	Glu	Arg	Asn	385	390	395	400
Asn	Thr	Ala	His	Pro	Glu	Thr	Leu	Ala	Asn	Asn	Pro	Pro	Asp	Asn	Thr	405	410	415	
Thr	Pro	Ser	Thr	Pro	Pro	Gln	Asp	Gly	Glu	Arg	Thr	Ser	Ser	His	Thr	420	425	430	
Thr	Pro	Ser	Pro	Arg	Pro	Val	Pro	Thr	Ser	Thr	Ile	His	Pro	Thr	Thr	435	440	445	
Arg	Glu	Thr	His	Ile	Pro	Thr	Thr	Met	Thr	Thr	Ser	His	Asp	Thr	Asp	450	455	460	
Ser	Asn	Arg	Pro	Asn	Pro	Ile	Asp	Ile	Ser	Glu	Ser	Thr	Glu	Pro	Gly	465	470	475	480
Pro	Leu	Thr	Asn	Thr	Thr	Arg	Gly	Ala	Ala	Asn	Leu	Leu	Thr	Gly	Ser	485	490	495	
Arg	Arg	Thr	Arg	Arg	Glu	Ile	Thr	Leu	Arg	Thr	Gln	Ala	Lys	Cys	Asn	500	505	510	
Pro	Asn	Leu	His	Tyr	Trp	Thr	Thr	Gln	Asp	Glu	Gly	Ala	Ala	Ile	Gly	515	520	525	
Leu	Ala	Trp	Ile	Pro	Tyr	Phe	Gly	Pro	Ala	Ala	Glu	Gly	Ile	Tyr	Thr	530	535	540	
Glu	Gly	Ile	Met	His	Asn	Gln	Asn	Gly	Leu	Ile	Cys	Gly	Leu	Arg	Gln	545	550	555	560
Leu	Ala	Asn	Glu	Thr	Thr	Gln	Ala	Leu	Gln	Leu	Phe	Leu	Arg	Ala	Thr	565	570	575	
Thr	Glu	Leu	Arg	Thr	Phe	Ser	Ile	Leu	Asn	Arg	Lys	Ala	Ile	Asp	Phe	580	585	590	
Leu	Leu	Gln	Arg	Trp	Gly	Gly	Thr	Cys	His	Ile	Leu	Gly	Pro	Asp	Cys	595	600	605	
Cys	Ile	Glu	Pro	His	Asp	Trp	Thr	Lys	Asn	Ile	Thr	Asp	Lys	Ile	Asp	610	615	620	
Gln	Ile	Ile	His	Asp	Phe	Ile	Asp	Lys	Pro	Leu	Pro	Asp	Gln	Thr	Asp				

-continued

625	630	635	640
Asn Asp Asn Trp Trp Thr Gly Trp Arg Gln Trp Val Pro Ala Gly Ile	645	650	655
Gly Ile Thr Gly Val Ile Ile Ala Val Ile Ala Leu Leu Cys Ile Cys	660	665	670
Lys Phe Leu Leu	675		

<210> SEQ ID NO 10
 <211> LENGTH: 18935
 <212> TYPE: DNA
 <213> ORGANISM: Cote d'Ivoire ebolavirus

<400> SEQUENCE: 10

```

cggacacaca aaaagaaaga aggttttttg atctttattg tgtgcgaata actatgagga      60
agattaataa ttttcctctc attgacactt acattaagat taagattctc attgatctgt    120
tacttactct gaggataata attggtgttc agaagtaccc cattccccag tgggggcaaa    180
gacagtccaa aagactcaac ttgtcctatt caactaatct gttttgtctc agtagttcac    240
atattgatca taccaggag ttggacctaa ttcaaagct tagagtggga cctagtgtat    300
cctcggggct gtaatataat cagccattta acacataaca agccctactg ttttcttggt    360
ttgccgtgca tttagaataa gagacaactt aaacctccga ttccggcaaca cagggaataa    420
tctcaccaga cccggcagtg tcttcaggct tcatagcccc aagatggaga gtcgggcccc    480
caaagcatgg atgacgcaca ccgcatcagg ttctgaaaca gattaccata agattttaac    540
agcaggattg tcagtccaac aaggcattgt gagacaacgg gtcattcaag tccaccaggt    600
tacaaaccta gaagaaatat gccaatgtat cattcaagcc tttgaagctg gtgttgattt    660
tcaagagagt gcagacagtt tcttgctgat gctatgttta catcatgctt atcagggtga    720
ctacaagcaa ttcttggaag gcaatgcagt caagtacctt gagggcatg gctttcgctt    780
tgaggtcagg aaaaaggaag gagtcaagcg actcgaagaa ttgcttcttg ctgcatccag    840
tggaagagc atcaggagaa cactggctgc aatgcctgaa gaggagacaa cagaagcaaa    900
tgccggacag ttcctctctt ttgctagctt atttcttctt aagctagttg tcggagaaaa    960
agcctgtcta gaaaagggtc agcggcaaat tcaagttcat tctgagcagg gattgatcca   1020
ataccacaca gcctggcagt cagttggaca catgatggtc attttcagac tgatgagaac   1080
aaattttcta attaagttcc tccttataca tcaagggatg catatggtag caggacacga   1140
tgctaacgat gctgtcatcg caaactctgt agctcaagca cgtttttcag gattattgat   1200
cgttaaaaca gtgctagatc acatccttca gaaaacagag cacggagtgc gtcttcatcc   1260
tttggcaaga actgctaagg tcaagaacga agtaaatcc ttttaaggctg ccttagctc    1320
gctagacaaa catggagagt atgctccttt tgctcgcttg ctgaatcttt ctggagtc    1380
caatctcgag cacggactgt ttoctcagct ttctgcaatt gccctaggtg tcgcaacggc    1440
acacggcagt accctggcag gagtaaatgt gggggaacag tatcagcaac tacgagaagc    1500
agccactgag gcagaaaaac aattgcagaa atacgctgaa tctcgcgagc ttgaccatct    1560
aggtctcgat gatcaagaga agaagatctt gaaagacttc catcagaaga aaaatgaaat    1620
cagcttccag cagacaacag ccatgggtcac actacggaag gaaaggctag ccaagctcac    1680
tgaggcaatc acctccacat cccttctcaa gacaggaaaa cagtatgatg atgacaacga    1740

```

-continued

tatccccctt	cctgggcccc	tcaatgataa	cgaaaactca	gaacagcaag	acgatgatcc	1800
aacagattct	caggacacta	ccatccctga	tatcattggt	gacccggatg	atggcagata	1860
caacaattat	ggagactatc	ctagtggagc	ggcgaatgcc	cctgaagacc	ttgttctttt	1920
tgaccttgaa	gatggtgacg	aggatgatca	ccgaccgtca	agttcatcag	agaacaacaa	1980
caaacacagt	cttacaggaa	ctgacagtaa	caaaaacaagt	aactggaatc	gaaacccgac	2040
taatatgcca	aagaaagact	ccacacaaaa	caatgacaat	cctgcacagc	gggctcaaga	2100
atacgccagg	gataacatcc	aggatacacc	aacaccccat	cgagctctaa	ctcccatcag	2160
cgaagaaacc	ggctccaatg	gtcacaatga	agatgacatt	gatagcatcc	ctcctttgga	2220
atcagacgaa	gaaaacaaca	ctgagacaac	cattaccacc	acaaaaata	ccactgctcc	2280
accagcacct	gtttatcgga	gtaattcaga	aaaggagccc	ctcccgaag	aaaaatccca	2340
gaagcaacca	aaccaagtga	gtggtagtga	gaataccgac	aataaacctc	actcagagca	2400
atcagtggaa	gaaatgtatc	gacacatcct	ccaaacacaa	ggaccatttg	atgccatcct	2460
atactattac	atgatgacgg	aggagccgat	tgtcttttagc	actagtgatg	ggaagaata	2520
cgtataccct	gattctcttg	aagggggagc	tccaccgtgg	ctcagtgaag	aagaggcctt	2580
gaatgaggac	aataggttta	tcacaatgga	tgatcaacaa	ttctactggc	ctgtaatgaa	2640
tcacaggaac	aaattcatgg	ctatccttca	gcaccacaag	taatttcttc	ataatgacag	2700
atcattgtaa	ggttattacc	accatccctg	caacaagca	tgaaaaccac	actcaacaac	2760
gccttaccac	aggatacctt	ggagaccata	caccaagatc	agcagctgtg	caaccacccc	2820
catgccaatc	caccaccaca	accaccaaac	aataatccca	agaccaaacc	gcacacatcc	2880
agatcaaccc	aaacctcaa	acaccacccc	actccgcat	cccagaccaa	actccgcccc	2940
agacaagcac	cccaccatc	ccagaaaccg	cacggccgag	aatcgatccc	cagcattcaa	3000
aatgcgttat	taagaaaaaa	catatgatga	agattaaaac	cttcatcaac	attgcacaga	3060
ctttgatcct	taggagttta	ttctagctat	ctacaaaacg	gggtccaaac	ggaatgattt	3120
ccactagggc	tgacgaatc	aatgatcctt	cattaccaat	cagaaaccag	tgtacacgtg	3180
gccctgaact	atcaggatgg	atctccgaac	aattaatgac	aggcaaaatt	cgggtacatg	3240
aaatcttcaa	cgacactgag	ccccacataa	gctcagggtc	cgactgcctt	cccagaccca	3300
aaaaacacgc	cccccgact	cgcaacaccc	agacacagac	cgatccggtt	tgcaatcaca	3360
attttgaaga	cgttacacaa	gcactaacat	cattaaccaa	tgtcatacaa	aaacaggctc	3420
ttaacttaga	gtctctcgaa	caacgcacac	tagatctaga	gaatggctta	aagccaatgt	3480
atgacatggc	taaagtcatt	tctgcattga	atagatcttg	tgctgagatg	gtagcaaaat	3540
atgatctcct	ggtgatgaca	actggccgag	caaccggcac	cgcgctgca	actgaggctt	3600
attgggagga	acatggacaa	ccaccacctg	gaccatcact	ttatgaagag	agtgcgatta	3660
gaggcaagat	taacaagcaa	gaggataaag	tacctaagga	agttcaagaa	gcttttcgta	3720
atctggacag	taccagctca	ctaacagaag	agaactttgg	caagccagat	atatctgcaa	3780
aggacctacg	agacatcatg	tatgaccacc	taccaggctt	cgggtacggc	tttcccaaac	3840
tggtccaggt	aatttgcaag	ctaggaaaag	acaattctgc	attggacatt	attcatgctg	3900
agttccaagc	cagccttgct	gaaggtgatt	ctccccaatg	tgccctgac	caaataacaa	3960
aacggatccc	catcttcag	gatgccactc	cgcccacaat	tcacatccgc	tctcgtggtg	4020

-continued

acatcccacg	tgccctgcaa	aaaagtctcc	gtccagttcc	tccatcacca	aaaatagaca	4080
gaggttgggt	ttgcattttc	caattgcagg	acgggaagac	acttgggctc	aagatatagg	4140
gtccccagc	caaagacacg	tgccgtccca	tcctccctca	ccttcagaca	tcaacgcacg	4200
gcagtcacca	acaccgggtg	gggaggcgcc	cggcgacaac	acatgatgat	aggctgatct	4260
tcgggataag	agacatgaaa	aacccaaaaa	ccgtttacat	ccagatccaa	gatcaagagt	4320
ggcttggaag	taaggggcac	ttgttctttg	tctcaaagga	cttacaaaaa	caagggtgat	4380
gaagattaag	aaaaagccct	cttcagttgc	aaggagctaa	ttcttaaaac	ttcatctaga	4440
ctaaggataa	atcgattcca	atcacgatga	ggagaatcat	cctaccacacg	gcaccacctg	4500
aatacatgga	ggctgtttac	ccaatgagaa	caatgaattc	tggtgcagac	aacactgcca	4560
gtggccctaa	ttacacaaca	actggtgtga	tgacaaatga	tactccctct	aattcactcc	4620
gaccagttgc	agatgataat	attgatcatc	cgagccacac	gcctaacagt	gttgccctctg	4680
catttatatt	ggaagctatg	gtgaatgtaa	tatctggccc	gaaagtgtctg	atgaagcaaa	4740
tcccaatctg	gcttctctctg	ggtgtctctg	accagaagac	atatagcttt	gattcaacca	4800
ctgctgccat	tatgctagca	tcataataca	tactcattt	tggcaaaacc	tcaaatcccc	4860
ttgtgagaat	caaccgactt	ggtcctggca	tacctgatca	cccactacga	ctcctaagaa	4920
taggaaatca	agccttccta	caagagtttg	tgctacctcc	tgtacaactg	ccacaatact	4980
tacttttga	tctgacacg	ctgaagctga	tcaccagacc	actcccagcg	gcaacctgga	5040
cagatgaaac	tccagctgtg	tcaactggca	cgctccgccc	agggatctca	ttccatccca	5100
aattaaggcc	tatcctgcta	ccaggaagag	ctggaaagaa	gggctccaac	tccgatctaa	5160
catctctga	caaaatccag	gctataatga	atttcctaca	agacctcaaa	attgtaccaa	5220
tcgatccaac	caagaatata	atgggtattg	aagtgccaga	actcctgggt	cacaggetga	5280
ctgggaagaa	gacaactacc	aagaatggtc	aaccaatcat	tccaattctg	ctaccaaggt	5340
acattgggtc	tgatctctca	tctcaagggt	atctcacaat	ggatgatcact	caggactgtg	5400
attcctgcca	ctccccggcc	agtcttcccc	cagtcaatga	aaaatgacca	tgagactcaa	5460
catcacactg	ccagagcacc	tcaccgcaag	tctatacaac	aatcaacccc	ggcatctaca	5520
acctgcaaaa	accagcccat	ctgatactcc	tggtatcggt	ggcaagacaa	ggcagccaag	5580
cagcagcccc	cgagccgagc	ccaaacccat	tacaccgag	cccaaccccc	atccagcaac	5640
ccacaaccgt	caaacgcaca	gatggacaag	caaagaacat	caagccagga	gcaaacacaga	5700
ccccagttct	aagctgatca	acccctcccg	caatcccacc	aacgccagca	aaaatcccc	5760
aactcgatac	caaccccaag	caaatcagct	caaaccgtct	atctctcccc	gcttcaactcc	5820
acaccccgag	ttcagcaaac	gatcaacgca	cttcttatgc	cacagcttat	attaagaaaa	5880
agaacttgat	gaagattaag	gcaaccagtg	gtgctatctt	catctctttg	atttgagtct	5940
taagtgaata	cacaggttct	aatactgttc	ttctgtccaa	cggtataatt	cagccaggcc	6000
taagacagta	gctaatacaca	gtcatcatgg	gagcgtcagg	gattctgcaa	ttgccccgtg	6060
agcgttctag	gaaaacatct	ttctttgttt	gggtaataat	cctattccat	aaagtctttt	6120
caatcccgtt	gggggttgta	cacaacaata	ccctacaagt	gagtgatatt	gacaagtttg	6180
tgtgccgaga	caaatctctc	tcaactagcc	aattgaagtc	agtcgggttg	aacttggagg	6240
gcaatggagt	agcaactgat	gtaccaacgg	caacccaaaag	atgggggtttt	cgagctggtg	6300

-continued

ttccacaaaa	ggtggtaaat	tgcgaagctg	gagaatgggc	tgagaactgt	tataacctgg	6360
ctataaagaa	agttgatggg	agtgagtgcc	taccagaagc	ccctgagggg	gtgaggggatt	6420
ttccccgttg	ccgctatgta	cacaaagtct	caggaactgg	accatgcccc	ggaggactcg	6480
cctttcacaa	agaaggagcc	ttcttctctg	atgaccgact	cgcatacaaa	atcattttatc	6540
ggggtacaac	ctttgccgaa	ggagttattg	catttctgat	cttgccctaag	gcgcgaaagg	6600
attttttcca	gtctcctcca	ttgcatgagc	ctgccaaat	gaccacggat	ccctccagtt	6660
actatcacac	gacaacaata	aactacgtgg	ttgataattt	tggaaccaac	accacagagt	6720
ttctgttcca	agtcgatcat	ttgacgtatg	tgcagctcga	ggcaagattc	acaccacaat	6780
tccttgctct	cctaaatgaa	accatctact	ctgataaccg	cagaagtaac	acaacaggaa	6840
aactaatctg	gaaaaataat	cccactgttg	ataccagcat	gggtgagtgg	gctttctggg	6900
aaaaataaaa	aacttcacaa	aaacccttcc	aagtgaagag	ttgtctttcg	tacctgtacc	6960
agaaacccag	aaccaggtcc	ttgacacgac	agcgacggtc	tctcctccca	tctccgcccc	7020
caaccacgca	gccgaagacc	acaaagaatt	ggtttcagag	gattccactc	cagtggttca	7080
gatgcaaaa	atcaagggaa	aggacacaat	gccaaccaca	gtgacgggtg	taccaacaac	7140
cacacctct	ccattttcaa	tcaatgctcg	caacactgat	cataccaaat	cattttatcgg	7200
cctggagggg	ccccagaag	accacagcac	cacacagcct	gccaaagcca	ccagccaacc	7260
aaccaacagc	acagaatcga	cgacactaaa	cccaacatca	gagccctcca	gtagaggcac	7320
gggaccatcc	agccccacgg	tccccaacac	cacagaaaagc	cacgccgaac	ttggcaagac	7380
aacccccacc	acactcccag	aacagcacac	tgccgccagt	gccattccaa	gagccgtgca	7440
ccccgacgaa	ctcagtgga	ctggcttctc	gacgaacaca	atacgggggg	ttacaaatct	7500
cctgacagga	tccagaagaa	agcgaaggga	tgtcactccc	aatacacaac	ccaaatgcaa	7560
cccaaacctg	cactatttga	cagccttga	tgagggtgct	gccatagggt	tagcctggat	7620
accatacttc	gggccagcag	ctgaggggaat	ttacactgaa	ggcataatgg	agaatcaaaa	7680
tggattgac	tgtgatttga	ggcagctggc	caacgaaaag	acacaagctc	ttcaattgtt	7740
cttaagggca	actactgagt	tgcgtacatt	ctctatacta	aatcggaag	caatagactt	7800
cttgctccaa	agatggggag	gaacatgtca	cattctaggg	cctgattgtt	gcattgaacc	7860
ccaagattgg	acaaaaata	tactgataa	aattgatcaa	ataatccatg	actttgtcga	7920
taataatctt	ccaaatcaga	atgatggcag	caactggtgg	actggatgga	aacaatgggt	7980
tctgctgga	ataggaatca	caggagtaat	cattgctatt	attgctttgc	tgtgcatttg	8040
caaattcatg	ctttgaacta	atatagcatc	atactttcta	atattccccc	aatatgaatt	8100
ttgtttttcg	attttattta	atgatataac	ctctgtatac	ctcactaatg	tactcgagca	8160
taatttccct	gatagacttg	attgtatttg	atgattaagg	acctcacaaa	attcctgggg	8220
attgaaaaga	actggataac	tcaataaatt	ttatgctagg	accacaaata	cacttgatga	8280
agattaagaa	aaagataatc	ttatgattat	cattgatctt	catctatacc	ttaaatactc	8340
tattcaagga	gagtatgaca	aaaccaagta	gtattggata	aacttgctct	gcattcaaat	8400
ctgaagacat	acggcttata	tattcactat	tgtattagaa	aatctagggg	atatcatttg	8460
aaactaatta	gtgactaaaa	cacacaactc	aagtcggcca	gaatgggaag	tgttcataaa	8520
agaggtcgct	ccaggatctc	ccgacaaaa	acaagggatg	gacctagtca	tttagtacgg	8580

-continued

gcgagatcat cctctcgagc tagttatcga agtgaatacc atacaccaag gagtgcctcg	8640
cagatccgtg tccccactgt ctttcacgga aaaaagacag atttattgac agttccacca	8700
gcacctaag atgtatgccc gactttaag aaagggttcc tatgtgacag caatttctgt	8760
aaaaaggatc accaacttga aagcttaaca gatagagagt tactcttgct gattgcacgc	8820
aagacatgtg gatccacgga acaacaacta agcatagtgt ctccaaaaga ttcacgtctg	8880
gctaataccta ttgctgagga ttccaacaa aaagatgggc ctaaggtaac actgtcgatg	8940
cttatagaga cagcagagta ttggtccaaa caggacatta agaacatcga tgattcaaga	9000
ttaagagctt tattgacctt ttgtgctgtt atgacgcgca aattttcaaa atctcaactt	9060
agcttgctat gtgaaagcca cttacggcga gaaggacttg gtcaagacca atcagagtca	9120
gttctggagg tatatcaacg cttacacagc gataaagggt ggaatttcga ggcagcacta	9180
tggcagcagt gggatcggca atcattgata atgttcataa cagcattttt aaatattgca	9240
ttacaattac catgtgagag ttcactgttt gttatttcag gtttgagaat gctgataccc	9300
cagtcggaag ccactgaggt tgtaaccccc tccgaaacct gcacatgggc agaaggagga	9360
agttcccat gaagcccaa atcacaaggc gagctaaaaa atcccttttg aacatgcata	9420
acatcacata caatttcaaa ggcattggaa taaatggtga tttcaggaag attagtgttt	9480
gccctcaaaa tcagatccga gcaataatca tctactctac agccagttaa tttctaatat	9540
aaagggttaa aaaatgctgc aggcagccta ttgttcaca ggtcccaatt cttctgttta	9600
aattgtagga gctagcaca gtgatgcaat taaatgatac tagtatatac aatgccacca	9660
acttaattct aagattttgt atatctcgga aattcaaaat taaatgctac gttattgatt	9720
caattaagaa aaagacaatg gaccatcaaa attagttaa tacctgaact aatgcactta	9780
tagaaacagg agaaccagcc agacagcaga caaataacaa tgaaccacaa tatgttactg	9840
ctataatgaa gttcgtaat tcaaaaacaa atgatgaaga ttaatgcaga tgtctaaagg	9900
ataaacactc catgcatcag tgttataatt gggctctgta gaaaatcttc atctctcca	9960
acctacctca aagaaggatt ttaccgcat tgggagttat aacgacaata gggacaacca	10020
cctttgacac tagccaagct tgctgtgggc acacagcatt ttatcttgca acgtcgacat	10080
tcccatcaat ctgaggagta acagctatca aaacaacgca tatgtagaca ttgtcggtta	10140
tagtactgcc taagacaact atttataata acagttggaa ttcatttttt caccgaagct	10200
attctcaagt taacagttga aacaggactc gaccagagc aactccgat acgtaacata	10260
agaaaagaac aacccttgac ccagagtga caagctcata ctatcaaggc taatcctcgg	10320
gcctgcctgg agtcacaat ggccaaggct actgggaggt acaaccttat cccccaaag	10380
aaagatcttg aaaaagggtt ggttctgaat gacctttgca ctctctcagt ggcccagacg	10440
gtccagggat ggaagggtac ctgggctggg attgaatttg atgttacaca gaaagggtg	10500
gccttattgc acaggctcaa gaccagtgat tttgctccag cctgggtcaat gaccaggaac	10560
ttatttcac atctctttca aaaccgaac totacaattg agtcgccact ttgggcactg	10620
cgggtcatac tagcagcagg tattcaagat cagctaattg atcaatcgtt gatcgaaacc	10680
ttggcaggag cgctaggctt aattgctgat tggcttctta ctactggaac aaaccacttt	10740
caaatgcga cacaacaggc taaggagcaa ctaagtctaa aaatgttgct cctggtgcga	10800
tcaaactcc taaagttcat caaccaacta gatgcactac atgttgtgaa ttacaatgga	10860

-continued

cttctcagta	gcattgaaat	tggcaccaaa	agccatacaa	ttataattac	cgggacaaat	10920
atggggtttt	tggtagagtt	gcaagagcct	gacaaatcag	ccatgaacac	cagaaaacca	10980
ggaccagtca	aattctccct	cctccatgaa	tcaaccttga	agacacttgc	taaaaaacct	11040
gcgaccacga	tgcaagcact	aatcttagaa	ttcaatagtt	ctctcgctat	ttaactcaac	11100
tcatcaaaat	gctaacttgt	gatccttaag	ctgcacctta	gacttttgat	aagaatacta	11160
actattgatg	attgtctttg	acatgaggat	aagaacactg	cccattagat	agatgggggt	11220
caccattaat	acacaattac	ccaatcatgt	taacagcagt	tagatccctc	aagtatatca	11280
agttcattct	accctttgca	ttgtcactct	aattaaatca	cctgatacaa	ttatgttaat	11340
tagctagatt	ctctcatttt	tagacttggt	tgctagaata	attgatcatc	cacttgatta	11400
cacatccaac	taggggtctag	ttcatagatt	gctaataatc	tttagttcaa	tactaatgac	11460
aaagagatta	gattagctat	agcttgagga	agattaagaa	aaagtgtctg	tggggctctt	11520
ccgtgtagaa	gggcacacag	ccataattct	tcctctttat	acaacatggc	tacacaacat	11580
acgcaatatc	cagacgcaag	gttatcatca	cctatagttt	tagatcagtg	tgatcttgtc	11640
actcgtgctt	gtggattgta	ttccgcatac	tccttaaatc	cccaactaaa	gaactgtaga	11700
ctaccgaaac	atataaccg	actaaaatat	gacaccactg	ttacagagtt	tttgagtgat	11760
gtgccggtag	caacattgcc	agcggatttt	ttagtaccta	catttcttag	gactctatca	11820
ggaaatgggt	cttgtccaat	tgatccaaaa	tgcagtcaat	ttttagaaga	aattgtcaat	11880
tatactctac	aagatatctg	cttcctaaac	tattacctca	atcgagccgg	agtgcataac	11940
gatcatgtgg	atagggattt	tggacaaaaa	attcgcaatc	taatttgcca	caatgagggt	12000
ttacatcaaa	tgtttcactg	gtatgatctt	gcaattctag	cacgtagagg	gcgactaaat	12060
agaggggaata	atcgctcaac	atggtttgca	agtgataatt	tggtagatat	cctagggttat	12120
ggagattata	ttttttggaa	aataccatta	tcactactac	cagtggatac	acaaggcctc	12180
ccacatgcag	ccaaggactg	gtatcatgaa	tcggttttca	aggaggctat	tcaaggccat	12240
acacacatcg	tgtccatctc	tacagcagat	gtcttaatca	tgtgtaagga	cataatcacc	12300
tgtogattta	atactttact	gattgctgct	gtggcaaatc	tagaggattc	agttcattca	12360
gattaccctt	taccagaaac	agtgctctgac	ctatacaaag	caggagatta	tttaatctca	12420
ttgctaggat	cagaaggtta	caaagtcata	aaattccttg	agccgttatg	cttagcaaaag	12480
atccaactct	gctcaaatta	cactgagagg	aaaggaagat	tcctcactca	aatgcattta	12540
gctgtaaate	atacacttga	ggaacttaca	gggtcccag	aattaaggcc	acaacagatt	12600
cggagaggtaa	gggaattcca	tcaaagtctg	ataaacctta	aggcaactcc	tcaacaactc	12660
tgtgagttgt	tttcagtgcg	aaagcattgg	gggcacctg	tcttgcatag	cgaagggt	12720
atccaaaaag	taaagaagca	tgcaacagtg	ataaaagcat	tgcgccaat	aataatcttt	12780
gaaacatatt	gtgtgtttta	atacagcatt	gcaaaacatt	attttgatag	tcagggtacg	12840
tggtacagtg	tgacttctga	cagatgctta	acaccaggcc	tttctcttta	catcaaaaga	12900
aaccaatttc	ctccactacc	tatgatcaaa	gaacttttgt	gggaatttta	tcacttagat	12960
catcctccgt	tattctccac	caaagtgatt	agtgatttga	gtatctttat	taaagatcgt	13020
gctactgcag	tcgagaaaac	atgctgggac	gcagtttttg	aaccaaatgt	tcttggttat	13080
aaccaccgga	ataaatttgc	tacaaaaagg	gtacctgagc	aattccttga	acaggagaaat	13140

-continued

ttctcaatag	agagtgtcct	acattatgct	caacgtctgg	aatatcttct	cccggagtac	13200
cggaacttct	ctttttcact	caaggagaag	gagttaaaca	ttggacgagc	ttttgggaaa	13260
ttgccatadc	caacacgcaa	tgttcaaact	ctgtgcgaag	ctttgttagc	agatgggttg	13320
gcgaaagcat	tcccaagcaa	tatgatggtt	gtgacagagc	gcgagcaaaa	agaaagcctt	13380
ttgcatcaag	cgtcttgcca	tcacacaagt	gatgatattg	gtgagaatgc	tactgttaga	13440
ggcagtagtt	ttgtaacaga	cttggaaaaa	tacaatttag	cattccgata	tgagtttaca	13500
gctcctttta	ttgaatactg	taatcggtgt	tacgggtgaa	gaaatttggt	taattggatg	13560
cactacacta	taccacagtg	ttatatacat	gtgagtgatt	attataaccc	cccacatgga	13620
gtctctctcg	aaaaccgaga	aaatccacca	gaaggcca	gctcttaccg	tggtcatcta	13680
ggcgggattg	agggaactta	acaaaaactc	tggaacaagc	tctcatgtgc	acagatttca	13740
ttagttgaaa	tcaaaaccgg	ttttaaaactg	cgatctgcgg	taatgggtga	caatcaatgt	13800
ataactgtac	tctctgtatt	tcccctcgaa	actgagtcta	gtgagcaaga	attaagttct	13860
gaagataaat	ccgctagagt	agctgctagc	ttagcaaaag	tcacaagtgc	ctgcggcatc	13920
tttttaaaac	ctgatgaaac	ttttgttcac	tcagggttca	tttatgttgg	caaaaaacaa	13980
tatttgaatg	gagtacaatt	acctcaatca	ctgaaaactg	ctactagaat	tgacccttg	14040
tcagatgcta	tctttgatga	tcttcaaggg	acactagcta	gcataggcac	ggcttttgaa	14100
agatctatct	ccgaaactag	gcacgtagtc	ccttgtagag	tagcagctgc	attccatacc	14160
ttttttccg	taagaatctt	acaatatcat	catcttggtc	tcaacaaggg	aacagacctg	14220
ggteaattgt	cattaagcaa	gccattagat	tttggaacta	taactttggc	cttggcagta	14280
ccacaagtct	tgggtggctt	atcattccta	aatccagaaa	aatgttttta	tagaaatctg	14340
ggtagatcctg	ttacttcagg	gctgtttcag	ctcaagacat	atcttcaa	gatccacatg	14400
gatgatattg	ttttaccttt	gatcgcaaag	aaccagggga	actgtagcgc	aattgacttt	14460
gtgttaaaac	ctagtgggtt	aaacgtaccg	gggtcacagg	atttgacatc	cttctacgt	14520
cagatagtgc	gccgaacaat	tactctaagt	gctaaaaata	aattaataaa	cactttgttc	14580
cattcttctg	ctgatattaga	agatgaaatg	gtttgcaaat	ggttgctttc	ttctacacca	14640
gtcatgagta	ggtttgccgc	cgatatat	tctcgactc	ccagtgggaa	acgtttacag	14700
atcttaggtt	accttgaagg	gactagaaca	ttgttagcct	ctaaaattat	aaatcataat	14760
actgagacac	ctatcctaga	tcgattgagg	aaaattacgc	tgcaaagggtg	gagcctgtgg	14820
tttagttatc	tcgaccactg	tgatcaagtt	ctggctgatg	ccctaactca	gataacctgc	14880
actgtggact	tagcacagat	tcttcgcgag	tacacctggg	cacacatact	agagggaagg	14940
cagctcattg	gagcaacact	tccttgata	ctagaacaac	taaatgtcat	ctggctcaaa	15000
ccatagagc	attgccctaa	atgtgcaaag	tcagcaaacc	ctaaagggga	accttttgtt	15060
tctattgcaa	ttaaaaaaca	tgtagtaagt	gcttggcctg	atcaatcacg	acttagttgg	15120
acaattggag	atggcatccc	ttatatcgga	tctcgaaacg	aggataagat	tgggcagcca	15180
gccatcaaac	caaaatgccc	ttcagcagcc	ttactgtgaag	caattgagtt	gacatcaaga	15240
ttgacttggg	ttactcaagg	tggagcaaac	agcgacttac	tagttaaaac	cttcatagaa	15300
gcacgagtaa	atttaagcgt	acaggaaatt	ctccaaatga	caccttctca	ttactccggc	15360
aacattgtgc	atcgatataa	tgatcaatat	agtccacact	catttatggc	aaataggatg	15420

-continued

```

agtaattctg ctactagggt agttgtttcg acaaacactc ttggagaatt ttcaggagga 15480
ggtcagtcag caagagatag taatattatc ttccagaatg tcattaatth tgctgttgca 15540
ctttttgatc tacgatttag gaacgtggct actttctcta tacaacatca tcggggtcat 15600
cttcatttgt caaagtgttg cacgcgagag gttccagccc aatatttagt ttatacatca 15660
acattgccat tggaccttac acggtatcgg gataatgagt tgatttacga tgacaatcca 15720
ttaagagggt gtttaaattg caatctttct tttgataatc cgcttttcaa gggccagaga 15780
cttaacataa ttgaagaaga cttgattaga ctaccttact tatcaggatg ggagctagct 15840
aaaaactgta tccaatctat aatttctgac agcaacaatt catcaacgga tccaatcagt 15900
agtggggaaa cagcatcatt caccactcac ttcttgacat atcctaagat tggactacta 15960
tatagttttg gtgcactcat cagttattat ctaggcaaca ccattattag aacaaaaaaa 16020
ttgactctta acaacttcat atattaccta gctactcaa tacataatth acctcatcgc 16080
tcgttgagaa tccttaaac tactttgaaa cacgctagtg ttatctcgag attaataagt 16140
attgactctc acttctcaat ttatatgga ggaactgctg gtgatcgagg actttccgat 16200
gcggcaagat tgtttcttag aactgccatt actgtcttcc ttcaattcgt tagaaagtgg 16260
atagtgaac gcaagacagc tattccactg tgggtcatct acctctaga aggtcaaagt 16320
cctagtccga tcaacagttt tctacaccac gtcacgcat tgttgcaaca tgagtctctc 16380
cacgatcatg tttgtctgc agaagcccac agtcgagtgg agacattga taatttagtt 16440
tatatgtgta aaagcacagc aagtaacttc tttcatgctt cattagcata ctggagaagt 16500
cgatctaaaa atcaagacaa aagagagatg acaagatat tatctttgac gcaaacggaa 16560
aagaaaaatt cattcggtta tacagcacat ccagaaagca ctgctgttct tggttccctc 16620
cagaccagcc ttgctccacc tccatctgct gacgaggcta catatgatag gaaaaacaaa 16680
gttttgaaag cttccagacc tggcaagtat tcccagaata caaccaaac cccacccaac 16740
caaacaggtt gtcgcatagt atctcccaat atcacaggea cagatgggtg ccttctgccc 16800
aatgagggtt ctaacagcaa taacaataat ttagtctcgc acagaattgt actgccgttt 16860
ttacatttgt ctcataatta taacgaaaga cctctatca gaaagtctga ggggacaaca 16920
gagattgtaa ggcttactcg gcagctgagg gcaataccag acaccacaat atattgccgc 16980
ttcacgggaa tagtttcttc aatgcactat aagctcgatg aagtccttg ggaatttgat 17040
aattttaagt ctgctataac acttgccgaa ggtgaagggt cgggtgcatt actcttatta 17100
caaaaatata aagtagaaac cttgtttttt aatacactag ccacagaaca cagcattgaa 17160
gcagaaatta tttctggaat aactacacca agaatgcttc tccctattat gtctaggttc 17220
catggtggac aaataaaagt cactttaaac aattctgcaa gccagattac cgatattact 17280
aatcaaagt ggttggcaga ccaaaaatct aggatcccta agcaagtaga gattataacc 17340
atggatgctg aaacaacaga aacattaat cggcctaaaat tgtacgaagc agtccaacag 17400
ctgattgtct cacatattga tccgaatgca ctcaaagttg tggttcttaa agttttctta 17460
agtgcattg atggaatcct atggctgaat gataacctta cccctttgtt tgggctgggt 17520
tacttgatca agccgatcac ctctagccca aaatctagtg agtggtagct atgtctctca 17580
aacctctctt caactcaag acgattacct catcagagtc atactacttg catgcattgt 17640
attcaaacag cactccagct acaaatcag aggagctcat attggcttag ccacctgttc 17700

```

-continued

```

cagtatgcc aacataatgt gcatttagat tatattaatc tcggtttccc ttcattggag 17760
aggggtttat accatagata caatttagtc gattctcaga aaggcccttt gacttccatt 17820
gtccaacatc tagcgcacct gcagaccgag attagggagt tggtaaatga ctataatcaa 17880
caaagacaaa gtcgaaccca aacatatcat ttcattaaaa caataaaagg tcgtattaca 17940
aaattggtaa atgattacct taagtctttt ctaataatac aagccttaaa gcacaattgc 18000
acatggcaag aggaactaag agctcttcca gatctaatta gtgtctgcac tcgattctat 18060
catactcgaa actgttcctg tgaaccacgg ttcctagtac agactttata cttatcacgc 18120
atgcaggatt cggaaatcaa actaatagat agattgaccg gccttcttag tctatgtcca 18180
aatgggtttt ttcggttaagg actcttgacg tacaaactcc acatagttat acaatggtag 18240
caggacacta tatgtaaat gaccctaaga aagagtaatt cgacacacag agttctcaag 18300
tgaaccacct catctcagat tatctgtggt tgcaattcta ataccgatt gttaccccg 18360
gagtataact ccagattaat ataagaaat accttttgtc ctgcaaatat atcttaaat 18420
caagtacata cgctccaaat cgtataaaat attaagaaaa agttaatctg cttgctttta 18480
ttataacttt aatattcgac aaatagttta cggctctatc actcaaaaat ttcattaaca 18540
aaagaagtac tctgagtata ttcacatc atatgtgatt aacatataag caacgcatga 18600
tgcgccttcc tcttacttat tgtgtgtgca cgcagtcggt gtactacctc gaaaattcca 18660
aacaataaat cgtgtctatc ccgcatctag tgtctttaat ttaagatctc aaatccaaaa 18720
aactgggttt atgttgatgt aaatcaataa taccgaaatt gcttgatatt aaaataaagc 18780
ttaaaggatt tttccttaaa cggatgatgt aggtatatag gaaagctcga tcacgatgtc 18840
ccttactcag aaaaagaaaa acggaagccc tattggccat ttaatcgtac acaaaaaatat 18900
ctttacaaaa ttgttttctc ttttttgtgt gtcca 18935

```

```

<210> SEQ ID NO 11
<211> LENGTH: 739
<212> TYPE: PRT
<213> ORGANISM: Bundibugyo ebolavirus
<220> FEATURE:
<221> NAME/KEY: misc_feature
<223> OTHER INFORMATION: Cote d'Ivoire ebolavirus NP protein

```

```

<400> SEQUENCE: 11

```

```

Met Glu Ser Arg Ala His Lys Ala Trp Met Thr His Thr Ala Ser Gly
 1             5             10             15
Phe Glu Thr Asp Tyr His Lys Ile Leu Thr Ala Gly Leu Ser Val Gln
 20            25            30
Gln Gly Ile Val Arg Gln Arg Val Ile Gln Val His Gln Val Thr Asn
 35            40            45
Leu Glu Glu Ile Cys Gln Leu Ile Ile Gln Ala Phe Glu Ala Gly Val
 50            55            60
Asp Phe Gln Glu Ser Ala Asp Ser Phe Leu Leu Met Leu Cys Leu His
 65            70            75            80
His Ala Tyr Gln Gly Asp Tyr Lys Gln Phe Leu Glu Ser Asn Ala Val
 85            90            95
Lys Tyr Leu Glu Gly His Gly Phe Arg Phe Glu Val Arg Lys Lys Glu
100           105           110
Gly Val Lys Arg Leu Glu Glu Leu Leu Pro Ala Ala Ser Ser Gly Lys

```

-continued

115					120					125					
Ser	Ile	Arg	Arg	Thr	Leu	Ala	Ala	Met	Pro	Glu	Glu	Thr	Thr	Glu	
130						135					140				
Ala	Asn	Ala	Gly	Gln	Phe	Leu	Ser	Phe	Ala	Ser	Leu	Phe	Leu	Pro	Lys
145					150					155					160
Leu	Val	Val	Gly	Glu	Lys	Ala	Cys	Leu	Glu	Lys	Val	Gln	Arg	Gln	Ile
				165					170					175	
Gln	Val	His	Ser	Glu	Gln	Gly	Leu	Ile	Gln	Tyr	Pro	Thr	Ala	Trp	Gln
			180					185					190		
Ser	Val	Gly	His	Met	Met	Val	Ile	Phe	Arg	Leu	Met	Arg	Thr	Asn	Phe
		195					200					205			
Leu	Ile	Lys	Phe	Leu	Leu	Ile	His	Gln	Gly	Met	His	Met	Val	Ala	Gly
	210					215					220				
His	Asp	Ala	Asn	Asp	Ala	Val	Ile	Ala	Asn	Ser	Val	Ala	Gln	Ala	Arg
225					230					235					240
Phe	Ser	Gly	Leu	Leu	Ile	Val	Lys	Thr	Val	Leu	Asp	His	Ile	Leu	Gln
			245						250					255	
Lys	Thr	Glu	His	Gly	Val	Arg	Leu	His	Pro	Leu	Ala	Arg	Thr	Ala	Lys
			260					265					270		
Val	Lys	Asn	Glu	Val	Asn	Ser	Phe	Lys	Ala	Ala	Leu	Ser	Ser	Leu	Ala
		275					280					285			
Gln	His	Gly	Glu	Tyr	Ala	Pro	Phe	Ala	Arg	Leu	Leu	Asn	Leu	Ser	Gly
	290					295					300				
Val	Asn	Asn	Leu	Glu	His	Gly	Leu	Phe	Pro	Gln	Leu	Ser	Ala	Ile	Ala
305					310					315					320
Leu	Gly	Val	Ala	Thr	Ala	His	Gly	Ser	Thr	Leu	Ala	Gly	Val	Asn	Val
			325						330					335	
Gly	Glu	Gln	Tyr	Gln	Gln	Leu	Arg	Glu	Ala	Ala	Thr	Glu	Ala	Glu	Lys
			340					345					350		
Gln	Leu	Gln	Lys	Tyr	Ala	Glu	Ser	Arg	Glu	Leu	Asp	His	Leu	Gly	Leu
		355					360					365			
Asp	Asp	Gln	Glu	Lys	Lys	Ile	Leu	Lys	Asp	Phe	His	Gln	Lys	Lys	Asn
	370					375					380				
Glu	Ile	Ser	Phe	Gln	Gln	Thr	Thr	Ala	Met	Val	Thr	Leu	Arg	Lys	Glu
385					390					395					400
Arg	Leu	Ala	Lys	Leu	Thr	Glu	Ala	Ile	Thr	Ser	Thr	Ser	Leu	Leu	Lys
			405						410					415	
Thr	Gly	Lys	Gln	Tyr	Asp	Asp	Asp	Asn	Asp	Ile	Pro	Phe	Pro	Gly	Pro
			420					425					430		
Ile	Asn	Asp	Asn	Glu	Asn	Ser	Glu	Gln	Gln	Asp	Asp	Asp	Pro	Thr	Asp
		435					440					445			
Ser	Gln	Asp	Thr	Thr	Ile	Pro	Asp	Ile	Ile	Val	Asp	Pro	Asp	Asp	Gly
		450				455					460				
Arg	Tyr	Asn	Asn	Tyr	Gly	Asp	Tyr	Pro	Ser	Glu	Thr	Ala	Asn	Ala	Pro
465					470					475					480
Glu	Asp	Leu	Val	Leu	Phe	Asp	Leu	Glu	Asp	Gly	Asp	Glu	Asp	Asp	His
			485					490						495	
Arg	Pro	Ser	Ser	Ser	Ser	Glu	Asn	Asn	Asn	Lys	His	Ser	Leu	Thr	Gly
			500					505					510		
Thr	Asp	Ser	Asn	Lys	Thr	Ser	Asn	Trp	Asn	Arg	Asn	Pro	Thr	Asn	Met
	515						520					525			

-continued

Pro Lys Lys Asp Ser Thr Gln Asn Asn Asp Asn Pro Ala Gln Arg Ala
 530 535 540
 Gln Glu Tyr Ala Arg Asp Asn Ile Gln Asp Thr Pro Thr Pro His Arg
 545 550 555 560
 Ala Leu Thr Pro Ile Ser Glu Glu Thr Gly Ser Asn Gly His Asn Glu
 565 570 575
 Asp Asp Ile Asp Ser Ile Pro Pro Leu Glu Ser Asp Glu Glu Asn Asn
 580 585 590
 Thr Glu Thr Thr Ile Thr Thr Thr Lys Asn Thr Thr Ala Pro Pro Ala
 595 600 605
 Pro Val Tyr Arg Ser Asn Ser Glu Lys Glu Pro Leu Pro Gln Glu Lys
 610 615 620
 Ser Gln Lys Gln Pro Asn Gln Val Ser Gly Ser Glu Asn Thr Asp Asn
 625 630 635 640
 Lys Pro His Ser Glu Gln Ser Val Glu Glu Met Tyr Arg His Ile Leu
 645 650 655
 Gln Thr Gln Gly Pro Phe Asp Ala Ile Leu Tyr Tyr Tyr Met Met Thr
 660 665 670
 Glu Glu Pro Ile Val Phe Ser Thr Ser Asp Gly Lys Glu Tyr Val Tyr
 675 680 685
 Pro Asp Ser Leu Glu Gly Glu His Pro Pro Trp Leu Ser Glu Lys Glu
 690 695 700
 Ala Leu Asn Glu Asp Asn Arg Phe Ile Thr Met Asp Asp Gln Gln Phe
 705 710 715 720
 Tyr Trp Pro Val Met Asn His Arg Asn Lys Phe Met Ala Ile Leu Gln
 725 730 735
 His His Lys

<210> SEQ ID NO 12
 <211> LENGTH: 341
 <212> TYPE: PRT
 <213> ORGANISM: Bundibugyo ebolavirus
 <220> FEATURE:
 <221> NAME/KEY: misc_feature
 <223> OTHER INFORMATION: Cote d'Ivoire ebolavirus VP35 NP protein

<400> SEQUENCE: 12

Met Ile Ser Thr Arg Ala Ala Ala Ile Asn Asp Pro Ser Leu Pro Ile
 1 5 10 15
 Arg Asn Gln Cys Thr Arg Gly Pro Glu Leu Ser Gly Trp Ile Ser Glu
 20 25 30
 Gln Leu Met Thr Gly Lys Ile Pro Val His Glu Ile Phe Asn Asp Thr
 35 40 45
 Glu Pro His Ile Ser Ser Gly Ser Asp Cys Leu Pro Arg Pro Lys Asn
 50 55 60
 Thr Ala Pro Arg Thr Arg Asn Thr Gln Thr Gln Thr Asp Pro Val Cys
 65 70 75 80
 Asn His Asn Phe Glu Asp Val Thr Gln Ala Leu Thr Ser Leu Thr Asn
 85 90 95
 Val Ile Gln Lys Gln Ala Leu Asn Leu Glu Ser Leu Glu Gln Arg Ile
 100 105 110
 Ile Asp Leu Glu Asn Gly Leu Lys Pro Met Tyr Asp Met Ala Lys Val
 115 120 125

-continued

```

Ile Ser Ala Leu Asn Arg Ser Cys Ala Glu Met Val Ala Lys Tyr Asp
 130                135                140

Leu Leu Val Met Thr Thr Gly Arg Ala Thr Ala Thr Ala Ala Thr
145                150                155                160

Glu Ala Tyr Trp Glu Glu His Gly Gln Pro Pro Pro Gly Pro Ser Leu
      165                170                175

Tyr Glu Glu Ser Ala Ile Arg Gly Lys Ile Asn Lys Gln Glu Asp Lys
      180                185                190

Val Pro Lys Glu Val Gln Glu Ala Phe Arg Asn Leu Asp Ser Thr Ser
      195                200                205

Ser Leu Thr Glu Glu Asn Phe Gly Lys Pro Asp Ile Ser Ala Lys Asp
      210                215                220

Leu Arg Asp Ile Met Tyr Asp His Leu Pro Gly Phe Gly Thr Ala Phe
225                230                235                240

His Gln Leu Val Gln Val Ile Cys Lys Leu Gly Lys Asp Asn Ser Ala
      245                250                255

Leu Asp Ile Ile His Ala Glu Phe Gln Ala Ser Leu Ala Glu Gly Asp
      260                265                270

Ser Pro Gln Cys Ala Leu Ile Gln Ile Thr Lys Arg Ile Pro Ile Phe
      275                280                285

Gln Asp Ala Thr Pro Pro Thr Ile His Ile Arg Ser Arg Gly Asp Ile
      290                295                300

Pro Arg Ala Cys Gln Lys Ser Leu Arg Pro Val Pro Pro Ser Pro Lys
305                310                315                320

Ile Asp Arg Gly Trp Val Cys Ile Phe Gln Leu Gln Asp Gly Lys Thr
      325                330                335

Leu Gly Leu Lys Ile
      340

<210> SEQ ID NO 13
<211> LENGTH: 326
<212> TYPE: PRT
<213> ORGANISM: Bundibugyo ebolavirus
<220> FEATURE:
<221> NAME/KEY: misc_feature
<223> OTHER INFORMATION: Cote d'Ivoire ebolavirus VP40 NP protein

<400> SEQUENCE: 13

Met Arg Arg Ile Ile Leu Pro Thr Ala Pro Pro Glu Tyr Met Glu Ala
 1                5                10                15

Val Tyr Pro Met Arg Thr Met Asn Ser Gly Ala Asp Asn Thr Ala Ser
      20                25                30

Gly Pro Asn Tyr Thr Thr Thr Gly Val Met Thr Asn Asp Thr Pro Ser
      35                40                45

Asn Ser Leu Arg Pro Val Ala Asp Asp Asn Ile Asp His Pro Ser His
      50                55                60

Thr Pro Asn Ser Val Ala Ser Ala Phe Ile Leu Glu Ala Met Val Asn
      65                70                75                80

Val Ile Ser Gly Pro Lys Val Leu Met Lys Gln Ile Pro Ile Trp Leu
      85                90                95

Pro Leu Gly Val Ser Asp Gln Lys Thr Tyr Ser Phe Asp Ser Thr Thr
      100                105                110

Ala Ala Ile Met Leu Ala Ser Tyr Thr Ile Thr His Phe Gly Lys Thr

```


-continued

115	120	125
Ser Asn Pro Leu Val Arg Ile Asn Arg Leu Gly Pro Gly Ile Pro Asp 130 135 140		
His Pro Leu Arg Leu Leu Arg Ile Gly Asn Gln Ala Phe Leu Gln Glu 145 150 155 160		
Phe Val Leu Pro Pro Val Gln Leu Pro Gln Tyr Phe Thr Phe Asp Leu 165 170 175		
Thr Ala Leu Lys Leu Ile Thr Gln Pro Leu Pro Ala Ala Thr Trp Thr 180 185 190		
Asp Glu Thr Pro Ala Val Ser Thr Gly Thr Leu Arg Pro Gly Ile Ser 195 200 205		
Phe His Pro Lys Leu Arg Pro Ile Leu Leu Pro Gly Arg Ala Gly Lys 210 215 220		
Lys Gly Ser Asn Ser Asp Leu Thr Ser Pro Asp Lys Ile Gln Ala Ile 225 230 235 240		
Met Asn Phe Leu Gln Asp Leu Lys Ile Val Pro Ile Asp Pro Thr Lys 245 250 255		
Asn Ile Met Gly Ile Glu Val Pro Glu Leu Leu Val His Arg Leu Thr 260 265 270		
Gly Lys Lys Thr Thr Thr Lys Asn Gly Gln Pro Ile Ile Pro Ile Leu 275 280 285		
Leu Pro Lys Tyr Ile Gly Leu Asp Pro Leu Ser Gln Gly Asp Leu Thr 290 295 300		
Met Val Ile Thr Gln Asp Cys Asp Ser Cys His Ser Pro Ala Ser Leu 305 310 315 320		
Pro Pro Val Asn Glu Lys 325		

<210> SEQ ID NO 14
 <211> LENGTH: 676
 <212> TYPE: PRT
 <213> ORGANISM: Bundibugyo ebolavirus
 <220> FEATURE:
 <221> NAME/KEY: misc_feature
 <223> OTHER INFORMATION: Cote d'Ivoire ebolavirus GP NP protein

<400> SEQUENCE: 14

Met Gly Ala Ser Gly Ile Leu Gln Leu Pro Arg Glu Arg Phe Arg Lys 1 5 10 15
Thr Ser Phe Phe Val Trp Val Ile Ile Leu Phe His Lys Val Phe Ser 20 25 30
Ile Pro Leu Gly Val Val His Asn Asn Thr Leu Gln Val Ser Asp Ile 35 40 45
Asp Lys Phe Val Cys Arg Asp Lys Leu Ser Ser Thr Ser Gln Leu Lys 50 55 60
Ser Val Gly Leu Asn Leu Glu Gly Asn Gly Val Ala Thr Asp Val Pro 65 70 75 80
Thr Ala Thr Lys Arg Trp Gly Phe Arg Ala Gly Val Pro Pro Lys Val 85 90 95
Val Asn Cys Glu Ala Gly Glu Trp Ala Glu Asn Cys Tyr Asn Leu Ala 100 105 110
Ile Lys Lys Val Asp Gly Ser Glu Cys Leu Pro Glu Ala Pro Glu Gly 115 120 125

-continued

Val	Arg	Asp	Phe	Pro	Arg	Cys	Arg	Tyr	Val	His	Lys	Val	Ser	Gly	Thr
130						135					140				
Gly	Pro	Cys	Pro	Gly	Gly	Leu	Ala	Phe	His	Lys	Glu	Gly	Ala	Phe	Phe
145					150					155					160
Leu	Tyr	Asp	Arg	Leu	Ala	Ser	Thr	Ile	Ile	Tyr	Arg	Gly	Thr	Thr	Phe
				165						170					175
Ala	Glu	Gly	Val	Ile	Ala	Phe	Leu	Ile	Leu	Pro	Lys	Ala	Arg	Lys	Asp
			180					185					190		
Phe	Phe	Gln	Ser	Pro	Pro	Leu	His	Glu	Pro	Ala	Asn	Met	Thr	Thr	Asp
		195					200					205			
Pro	Ser	Ser	Tyr	Tyr	His	Thr	Thr	Thr	Ile	Asn	Tyr	Val	Val	Asp	Asn
		210				215					220				
Phe	Gly	Thr	Asn	Thr	Thr	Glu	Phe	Leu	Phe	Gln	Val	Asp	His	Leu	Thr
225					230					235					240
Tyr	Val	Gln	Leu	Glu	Ala	Arg	Phe	Thr	Pro	Gln	Phe	Leu	Val	Leu	Leu
			245						250					255	
Asn	Glu	Thr	Ile	Tyr	Ser	Asp	Asn	Arg	Arg	Ser	Asn	Thr	Thr	Gly	Lys
		260						265					270		
Leu	Ile	Trp	Lys	Ile	Asn	Pro	Thr	Val	Asp	Thr	Ser	Met	Gly	Glu	Trp
		275					280					285			
Ala	Phe	Trp	Glu	Asn	Lys	Lys	Asn	Phe	Thr	Lys	Thr	Leu	Ser	Ser	Glu
		290				295					300				
Glu	Leu	Ser	Phe	Val	Pro	Val	Pro	Glu	Thr	Gln	Asn	Gln	Val	Leu	Asp
305					310					315					320
Thr	Thr	Ala	Thr	Val	Ser	Pro	Pro	Ile	Ser	Ala	His	Asn	His	Ala	Ala
			325						330					335	
Glu	Asp	His	Lys	Glu	Leu	Val	Ser	Glu	Asp	Ser	Thr	Pro	Val	Val	Gln
		340						345					350		
Met	Gln	Asn	Ile	Lys	Gly	Lys	Asp	Thr	Met	Pro	Thr	Thr	Val	Thr	Gly
		355					360					365			
Val	Pro	Thr	Thr	Thr	Pro	Ser	Pro	Phe	Pro	Ile	Asn	Ala	Arg	Asn	Thr
		370				375					380				
Asp	His	Thr	Lys	Ser	Phe	Ile	Gly	Leu	Glu	Gly	Pro	Gln	Glu	Asp	His
385					390					395					400
Ser	Thr	Thr	Gln	Pro	Ala	Lys	Thr	Thr	Ser	Gln	Pro	Thr	Asn	Ser	Thr
			405						410					415	
Glu	Ser	Thr	Thr	Leu	Asn	Pro	Thr	Ser	Glu	Pro	Ser	Ser	Arg	Gly	Thr
		420						425					430		
Gly	Pro	Ser	Ser	Pro	Thr	Val	Pro	Asn	Thr	Thr	Glu	Ser	His	Ala	Glu
		435					440					445			
Leu	Gly	Lys	Thr	Thr	Pro	Thr	Thr	Leu	Pro	Glu	Gln	His	Thr	Ala	Ala
		450				455					460				
Ser	Ala	Ile	Pro	Arg	Ala	Val	His	Pro	Asp	Glu	Leu	Ser	Gly	Pro	Gly
465					470				475						480
Phe	Leu	Thr	Asn	Thr	Ile	Arg	Gly	Val	Thr	Asn	Leu	Leu	Thr	Gly	Ser
			485					490						495	
Arg	Arg	Lys	Arg	Arg	Asp	Val	Thr	Pro	Asn	Thr	Gln	Pro	Lys	Cys	Asn
		500						505					510		
Pro	Asn	Leu	His	Tyr	Trp	Thr	Ala	Leu	Asp	Glu	Gly	Ala	Ala	Ile	Gly
		515					520					525			
Leu	Ala	Trp	Ile	Pro	Tyr	Phe	Gly	Pro	Ala	Ala	Glu	Gly	Ile	Tyr	Thr

-continued

530	535	540			
Glu Gly Ile Met Glu Asn Gln Asn Gly Leu Ile Cys Gly Leu Arg Gln					
545	550	555			560
Leu Ala Asn Glu Thr Thr Gln Ala Leu Gln Leu Phe Leu Arg Ala Thr					
	565	570			575
Thr Glu Leu Arg Thr Phe Ser Ile Leu Asn Arg Lys Ala Ile Asp Phe					
	580	585			590
Leu Leu Gln Arg Trp Gly Gly Thr Cys His Ile Leu Gly Pro Asp Cys					
	595	600			605
Cys Ile Glu Pro Gln Asp Trp Thr Lys Asn Ile Thr Asp Lys Ile Asp					
	610	615			620
Gln Ile Ile His Asp Phe Val Asp Asn Asn Leu Pro Asn Gln Asn Asp					
	625	630			635
Gly Ser Asn Trp Trp Thr Gly Trp Lys Gln Trp Val Pro Ala Gly Ile					
	645	650			655
Gly Ile Thr Gly Val Ile Ile Ala Ile Ile Ala Leu Leu Cys Ile Cys					
	660	665			670
Lys Phe Met Leu					
	675				

<210> SEQ ID NO 15
 <211> LENGTH: 365
 <212> TYPE: PRT
 <213> ORGANISM: Bundibugyo ebolavirus
 <220> FEATURE:
 <221> NAME/KEY: misc_feature
 <223> OTHER INFORMATION: Cote d'Ivoire ebolavirus SGP NP protein

<400> SEQUENCE: 15

Met Gly Ala Ser Gly Ile Leu Gln Leu Pro Arg Glu Arg Phe Arg Lys					
1	5	10			15
Thr Ser Phe Phe Val Trp Val Ile Ile Leu Phe His Lys Val Phe Ser					
	20	25			30
Ile Pro Leu Gly Val Val His Asn Asn Thr Leu Gln Val Ser Asp Ile					
	35	40			45
Asp Lys Phe Val Cys Arg Asp Lys Leu Ser Ser Thr Ser Gln Leu Lys					
	50	55			60
Ser Val Gly Leu Asn Leu Glu Gly Asn Gly Val Ala Thr Asp Val Pro					
	65	70			75
Thr Ala Thr Lys Arg Trp Gly Phe Arg Ala Gly Val Pro Pro Lys Val					
	85	90			95
Val Asn Cys Glu Ala Gly Glu Trp Ala Glu Asn Cys Tyr Asn Leu Ala					
	100	105			110
Ile Lys Lys Val Asp Gly Ser Glu Cys Leu Pro Glu Ala Pro Glu Gly					
	115	120			125
Val Arg Asp Phe Pro Arg Cys Arg Tyr Val His Lys Val Ser Gly Thr					
	130	135			140
Gly Pro Cys Pro Gly Gly Leu Ala Phe His Lys Glu Gly Ala Phe Phe					
	145	150			155
Leu Tyr Asp Arg Leu Ala Ser Thr Ile Ile Tyr Arg Gly Thr Thr Phe					
	165	170			175
Ala Glu Gly Val Ile Ala Phe Leu Ile Leu Pro Lys Ala Arg Lys Asp					
	180	185			190

-continued

```

Phe Phe Gln Ser Pro Pro Leu His Glu Pro Ala Asn Met Thr Thr Asp
   195                               200                               205

Pro Ser Ser Tyr Tyr His Thr Thr Thr Ile Asn Tyr Val Val Asp Asn
   210                               215                               220

Phe Gly Thr Asn Thr Thr Glu Phe Leu Phe Gln Val Asp His Leu Thr
  225                               230                               235                               240

Tyr Val Gln Leu Glu Ala Arg Phe Thr Pro Gln Phe Leu Val Leu Leu
          245                               250                               255

Asn Glu Thr Ile Tyr Ser Asp Asn Arg Arg Ser Asn Thr Thr Gly Lys
          260                               265                               270

Leu Ile Trp Lys Ile Asn Pro Thr Val Asp Thr Ser Met Gly Glu Trp
          275                               280                               285

Ala Phe Trp Glu Asn Lys Lys Thr Ser Gln Lys Pro Phe Gln Val Lys
   290                               295                               300

Ser Cys Leu Ser Tyr Leu Tyr Gln Lys Pro Arg Thr Arg Ser Leu Thr
  305                               310                               315                               320

Arg Gln Arg Arg Ser Leu Leu Pro Ser Pro Pro Thr Thr Thr Gln Pro
          325                               330                               335

Lys Thr Thr Lys Asn Trp Phe Gln Arg Ile Pro Leu Gln Trp Phe Arg
          340                               345                               350

Cys Lys Thr Ser Arg Glu Arg Thr Gln Cys Gln Pro Gln
   355                               360                               365

```

```

<210> SEQ ID NO 16
<211> LENGTH: 302
<212> TYPE: PRT
<213> ORGANISM: Bundibugyo ebolavirus
<220> FEATURE:
<221> NAME/KEY: misc_feature
<223> OTHER INFORMATION: Cote d'Ivoire ebolavirus SSGP NP protein

```

```

<400> SEQUENCE: 16

```

```

Met Gly Ala Ser Gly Ile Leu Gln Leu Pro Arg Glu Arg Phe Arg Lys
  1                               5                               10                               15

Thr Ser Phe Phe Val Trp Val Ile Ile Leu Phe His Lys Val Phe Ser
          20                               25                               30

Ile Pro Leu Gly Val Val His Asn Asn Thr Leu Gln Val Ser Asp Ile
   35                               40                               45

Asp Lys Phe Val Cys Arg Asp Lys Leu Ser Ser Thr Ser Gln Leu Lys
   50                               55                               60

Ser Val Gly Leu Asn Leu Glu Gly Asn Gly Val Ala Thr Asp Val Pro
  65                               70                               75                               80

Thr Ala Thr Lys Arg Trp Gly Phe Arg Ala Gly Val Pro Pro Lys Val
          85                               90                               95

Val Asn Cys Glu Ala Gly Glu Trp Ala Glu Asn Cys Tyr Asn Leu Ala
   100                               105                               110

Ile Lys Lys Val Asp Gly Ser Glu Cys Leu Pro Glu Ala Pro Glu Gly
   115                               120                               125

Val Arg Asp Phe Pro Arg Cys Arg Tyr Val His Lys Val Ser Gly Thr
   130                               135                               140

Gly Pro Cys Pro Gly Gly Leu Ala Phe His Lys Glu Gly Ala Phe Phe
  145                               150                               155                               160

Leu Tyr Asp Arg Leu Ala Ser Thr Ile Ile Tyr Arg Gly Thr Thr Phe
          165                               170                               175

```

-continued

Ala Glu Gly Val Ile Ala Phe Leu Ile Leu Pro Lys Ala Arg Lys Asp
180 185 190

Phe Phe Gln Ser Pro Pro Leu His Glu Pro Ala Asn Met Thr Thr Asp
195 200 205

Pro Ser Ser Tyr Tyr His Thr Thr Thr Ile Asn Tyr Val Val Asp Asn
210 215 220

Phe Gly Thr Asn Thr Thr Glu Phe Leu Phe Gln Val Asp His Leu Thr
225 230 235 240

Tyr Val Gln Leu Glu Ala Arg Phe Thr Pro Gln Phe Leu Val Leu Leu
245 250 255

Asn Glu Thr Ile Tyr Ser Asp Asn Arg Arg Ser Asn Thr Thr Gly Lys
260 265 270

Leu Ile Trp Lys Ile Asn Pro Thr Val Asp Thr Ser Met Gly Glu Trp
275 280 285

Ala Phe Trp Glu Asn Lys Lys Leu His Lys Asn Pro Phe Lys
290 295 300

<210> SEQ ID NO 17
<211> LENGTH: 289
<212> TYPE: PRT
<213> ORGANISM: Bundibugyo ebolavirus
<220> FEATURE:
<221> NAME/KEY: misc_feature
<223> OTHER INFORMATION: Cote d'Ivoire ebolavirus VP30 NP protein

<400> SEQUENCE: 17

Met Glu Val Val His Glu Arg Gly Arg Ser Arg Ile Ser Arg Gln Asn
1 5 10 15

Thr Arg Asp Gly Pro Ser His Leu Val Arg Ala Arg Ser Ser Ser Arg
20 25 30

Ala Ser Tyr Arg Ser Glu Tyr His Thr Pro Arg Ser Ala Ser Gln Ile
35 40 45

Arg Val Pro Thr Val Phe His Arg Lys Lys Thr Asp Leu Leu Thr Val
50 55 60

Pro Pro Ala Pro Lys Asp Val Cys Pro Thr Leu Lys Lys Gly Phe Leu
65 70 75 80

Cys Asp Ser Asn Phe Cys Lys Lys Asp His Gln Leu Glu Ser Leu Thr
85 90 95

Asp Arg Glu Leu Leu Leu Leu Ile Ala Arg Lys Thr Cys Gly Ser Thr
100 105 110

Glu Gln Gln Leu Ser Ile Val Ala Pro Lys Asp Ser Arg Leu Ala Asn
115 120 125

Pro Ile Ala Glu Asp Phe Gln Gln Lys Asp Gly Pro Lys Val Thr Leu
130 135 140

Ser Met Leu Ile Glu Thr Ala Glu Tyr Trp Ser Lys Gln Asp Ile Lys
145 150 155 160

Asn Ile Asp Asp Ser Arg Leu Arg Ala Leu Leu Thr Leu Cys Ala Val
165 170 175

Met Thr Arg Lys Phe Ser Lys Ser Gln Leu Ser Leu Leu Cys Glu Ser
180 185 190

His Leu Arg Arg Glu Gly Leu Gly Gln Asp Gln Ser Glu Ser Val Leu
195 200 205

Glu Val Tyr Gln Arg Leu His Ser Asp Lys Gly Gly Asn Phe Glu Ala

-continued

210	215	220
Ala Leu Trp Gln Gln Trp Asp Arg Gln Ser Leu Ile Met Phe Ile Thr		
225	230	235 240
Ala Phe Leu Asn Ile Ala Leu Gln Leu Pro Cys Glu Ser Ser Ser Val		
	245	250 255
Val Ile Ser Gly Leu Arg Met Leu Ile Pro Gln Ser Glu Ala Thr Glu		
	260	265 270
Val Val Thr Pro Ser Glu Thr Cys Thr Trp Ser Glu Gly Gly Ser Ser		
	275	280 285
His		
<210> SEQ ID NO 18		
<211> LENGTH: 251		
<212> TYPE: PRT		
<213> ORGANISM: Bundibugyo ebolavirus		
<220> FEATURE:		
<221> NAME/KEY: misc_feature		
<223> OTHER INFORMATION: Cote d'Ivoire ebolavirus VP24 NP protein		
<400> SEQUENCE: 18		
Met Ala Lys Ala Thr Gly Arg Tyr Asn Leu Ile Ser Pro Lys Lys Asp		
1	5	10 15
Leu Glu Lys Gly Leu Val Leu Asn Asp Leu Cys Thr Leu Ser Val Ala		
	20	25 30
Gln Thr Val Gln Gly Trp Lys Val Thr Trp Ala Gly Ile Glu Phe Asp		
	35	40 45
Val Thr Gln Lys Gly Met Ala Leu Leu His Arg Leu Lys Thr Ser Asp		
	50	55 60
Phe Ala Pro Ala Trp Ser Met Thr Arg Asn Leu Phe Pro His Leu Phe		
65	70	75 80
Gln Asn Pro Asn Ser Thr Ile Glu Ser Pro Leu Trp Ala Leu Arg Val		
	85	90 95
Ile Leu Ala Ala Gly Ile Gln Asp Gln Leu Ile Asp Gln Ser Leu Ile		
	100	105 110
Glu Pro Leu Ala Gly Ala Leu Gly Leu Ile Ala Asp Trp Leu Leu Thr		
	115	120 125
Thr Gly Thr Asn His Phe Gln Met Arg Thr Gln Gln Ala Lys Glu Gln		
	130	135 140
Leu Ser Leu Lys Met Leu Ser Leu Val Arg Ser Asn Ile Leu Lys Phe		
145	150	155 160
Ile Asn Gln Leu Asp Ala Leu His Val Val Asn Tyr Asn Gly Leu Leu		
	165	170 175
Ser Ser Ile Glu Ile Gly Thr Lys Ser His Thr Ile Ile Ile Thr Arg		
	180	185 190
Thr Asn Met Gly Phe Leu Val Glu Leu Gln Glu Pro Asp Lys Ser Ala		
	195	200 205
Met Asn Thr Arg Lys Pro Gly Pro Val Lys Phe Ser Leu Leu His Glu		
	210	215 220
Ser Thr Leu Lys Thr Leu Ala Lys Lys Pro Ala Thr Gln Met Gln Ala		
225	230	235 240
Leu Ile Leu Glu Phe Asn Ser Ser Leu Ala Ile		
	245	250

-continued

```

<210> SEQ ID NO 19
<211> LENGTH: 2210
<212> TYPE: PRT
<213> ORGANISM: Bundibugyo ebolavirus
<220> FEATURE:
<221> NAME/KEY: misc_feature
<223> OTHER INFORMATION: Cote d'Ivoire ebolavirus L NP protein

<400> SEQUENCE: 19

Met Ala Thr Gln His Thr Gln Tyr Pro Asp Ala Arg Leu Ser Ser Pro
1          5          10          15

Ile Val Leu Asp Gln Cys Asp Leu Val Thr Arg Ala Cys Gly Leu Tyr
20        25        30

Ser Ala Tyr Ser Leu Asn Pro Gln Leu Lys Asn Cys Arg Leu Pro Lys
35        40        45

His Ile Tyr Arg Leu Lys Tyr Asp Thr Thr Val Thr Glu Phe Leu Ser
50        55        60

Asp Val Pro Val Ala Thr Leu Pro Ala Asp Phe Leu Val Pro Thr Phe
65        70        75        80

Leu Arg Thr Leu Ser Gly Asn Gly Ser Cys Pro Ile Asp Pro Lys Cys
85        90        95

Ser Gln Phe Leu Glu Glu Ile Val Asn Tyr Thr Leu Gln Asp Ile Arg
100       105       110

Phe Leu Asn Tyr Tyr Leu Asn Arg Ala Gly Val His Asn Asp His Val
115       120       125

Asp Arg Asp Phe Gly Gln Lys Ile Arg Asn Leu Ile Cys Asp Asn Glu
130       135       140

Val Leu His Gln Met Phe His Trp Tyr Asp Leu Ala Ile Leu Ala Arg
145       150       155       160

Arg Gly Arg Leu Asn Arg Gly Asn Asn Arg Ser Thr Trp Phe Ala Ser
165       170       175

Asp Asn Leu Val Asp Ile Leu Gly Tyr Gly Asp Tyr Ile Phe Trp Lys
180       185       190

Ile Pro Leu Ser Leu Leu Pro Val Asp Thr Gln Gly Leu Pro His Ala
195       200       205

Ala Lys Asp Trp Tyr His Glu Ser Val Phe Lys Glu Ala Ile Gln Gly
210       215       220

His Thr His Ile Val Ser Ile Ser Thr Ala Asp Val Leu Ile Met Cys
225       230       235       240

Lys Asp Ile Ile Thr Cys Arg Phe Asn Thr Leu Leu Ile Ala Ala Val
245       250       255

Ala Asn Leu Glu Asp Ser Val His Ser Asp Tyr Pro Leu Pro Glu Thr
260       265       270

Val Ser Asp Leu Tyr Lys Ala Gly Asp Tyr Leu Ile Ser Leu Leu Gly
275       280       285

Ser Glu Gly Tyr Lys Val Ile Lys Phe Leu Glu Pro Leu Cys Leu Ala
290       295       300

Lys Ile Gln Leu Cys Ser Asn Tyr Thr Glu Arg Lys Gly Arg Phe Leu
305       310       315       320

Thr Gln Met His Leu Ala Val Asn His Thr Leu Glu Glu Leu Thr Gly
325       330       335

Ser Arg Glu Leu Arg Pro Gln Gln Ile Arg Lys Val Arg Glu Phe His
340       345       350

```

-continued

Gln Met	Leu Ile	Asn Leu	Lys Ala	Thr Pro	Gln Gln	Leu Cys	Glu Leu		
	355		360			365			
Phe Ser	Val Gln	Lys His	Trp Gly	His Pro	Val Leu	His Ser	Glu Lys		
	370		375			380			
Ala Ile	Gln Lys	Val Lys	Lys His	Ala Thr	Val Ile	Lys Ala	Leu Arg		
	385		390			395		400	
Pro Ile	Ile Ile	Phe Glu	Thr Tyr	Cys Val	Phe Lys	Tyr Ser	Ile Ala		
		405		410			415		
Lys His	Tyr Phe	Asp Ser	Gln Gly	Thr Trp	Tyr Ser	Val Thr	Ser Asp		
	420			425			430		
Arg Cys	Leu Thr	Pro Gly	Leu Ser	Ser Tyr	Ile Lys	Arg Asn	Gln Phe		
	435		440			445			
Pro Pro	Leu Pro	Met Ile	Lys Glu	Leu Leu	Trp Glu	Phe Tyr	His Leu		
	450		455			460			
Asp His	Pro Pro	Leu Phe	Ser Thr	Lys Val	Ile Ser	Asp Leu	Ser Ile		
	465		470			475		480	
Phe Ile	Lys Asp	Arg Ala	Thr Ala	Val Glu	Lys Thr	Cys Trp	Asp Ala		
		485		490			495		
Val Phe	Glu Pro	Asn Val	Leu Gly	Tyr Asn	Pro Pro	Asn Lys	Phe Ala		
	500			505			510		
Thr Lys	Arg Val	Pro Glu	Gln Phe	Leu Glu	Gln Glu	Asn Phe	Ser Ile		
	515		520			525			
Glu Ser	Val Leu	His Tyr	Ala Gln	Arg Leu	Glu Tyr	Leu Leu	Pro Glu		
	530		535			540			
Tyr Arg	Asn Phe	Ser Phe	Ser Leu	Lys Glu	Lys Glu	Leu Asn	Ile Gly		
	545		550		555		560		
Arg Ala	Phe Gly	Lys Leu	Pro Tyr	Pro Thr	Arg Asn	Val Gln	Thr Leu		
		565		570			575		
Cys Glu	Ala Leu	Leu Ala	Asp Gly	Leu Ala	Lys Ala	Phe Pro	Ser Asn		
	580		585			590			
Met Met	Val Val	Thr Glu	Arg Glu	Gln Lys	Glu Ser	Leu Leu	His Gln		
	595		600			605			
Ala Ser	Trp His	His Thr	Ser Asp	Asp Phe	Gly Glu	Asn Ala	Thr Val		
	610		615			620			
Arg Gly	Ser Ser	Phe Val	Thr Asp	Leu Glu	Lys Tyr	Asn Leu	Ala Phe		
	625		630		635		640		
Arg Tyr	Glu Phe	Thr Ala	Pro Phe	Ile Glu	Tyr Cys	Asn Arg	Cys Tyr		
		645		650			655		
Gly Val	Arg Asn	Leu Phe	Asn Trp	Met His	Tyr Thr	Ile Pro	Gln Cys		
	660		665				670		
Tyr Ile	His Val	Ser Asp	Tyr Tyr	Asn Pro	Pro His	Gly Val	Ser Leu		
	675		680			685			
Glu Asn	Arg Glu	Asn Pro	Pro Glu	Gly Pro	Ser Ser	Tyr Arg	Gly His		
	690		695			700			
Leu Gly	Gly Ile	Glu Gly	Leu Gln	Gln Lys	Leu Trp	Thr Ser	Ile Ser		
	705		710		715		720		
Cys Ala	Gln Ile	Ser Leu	Val Glu	Ile Lys	Thr Gly	Phe Lys	Leu Arg		
		725		730			735		
Ser Ala	Val Met	Gly Asp	Asn Gln	Cys Ile	Thr Val	Leu Ser	Val Phe		
	740		745				750		
Pro Leu	Glu Thr	Glu Ser	Ser Glu	Gln Glu	Leu Ser	Ser Glu	Asp Asn		

-continued

755					760					765					
Ala	Ala	Arg	Val	Ala	Ala	Ser	Leu	Ala	Lys	Val	Thr	Ser	Ala	Cys	Gly
770					775					780					
Ile	Phe	Leu	Lys	Pro	Asp	Glu	Thr	Phe	Val	His	Ser	Gly	Phe	Ile	Tyr
785					790					795					800
Phe	Gly	Lys	Lys	Gln	Tyr	Leu	Asn	Gly	Val	Gln	Leu	Pro	Gln	Ser	Leu
				805					810					815	
Lys	Thr	Ala	Thr	Arg	Ile	Ala	Pro	Leu	Ser	Asp	Ala	Ile	Phe	Asp	Asp
				820					825					830	
Leu	Gln	Gly	Thr	Leu	Ala	Ser	Ile	Gly	Thr	Ala	Phe	Glu	Arg	Ser	Ile
				835					840					845	
Ser	Glu	Thr	Arg	His	Val	Val	Pro	Cys	Arg	Val	Ala	Ala	Ala	Phe	His
				850					855					860	
Thr	Phe	Phe	Ser	Val	Arg	Ile	Leu	Gln	Tyr	His	His	Leu	Gly	Phe	Asn
865					870					875					880
Lys	Gly	Thr	Asp	Leu	Gly	Gln	Leu	Ser	Leu	Ser	Lys	Pro	Leu	Asp	Phe
				885					890					895	
Gly	Thr	Ile	Thr	Leu	Ala	Leu	Ala	Val	Pro	Gln	Val	Leu	Gly	Gly	Leu
				900					905					910	
Ser	Phe	Leu	Asn	Pro	Glu	Lys	Cys	Phe	Tyr	Arg	Asn	Leu	Gly	Asp	Pro
				915					920					925	
Val	Thr	Ser	Gly	Leu	Phe	Gln	Leu	Lys	Thr	Tyr	Leu	Gln	Met	Ile	His
				930					935					940	
Met	Asp	Asp	Leu	Phe	Leu	Pro	Leu	Ile	Ala	Lys	Asn	Pro	Gly	Asn	Cys
945					950					955					960
Ser	Ala	Ile	Asp	Phe	Val	Leu	Asn	Pro	Ser	Gly	Leu	Asn	Val	Pro	Gly
				965					970					975	
Ser	Gln	Asp	Leu	Thr	Ser	Phe	Leu	Arg	Gln	Ile	Val	Arg	Arg	Thr	Ile
				980					985					990	
Thr	Leu	Ser	Ala	Lys	Asn	Lys	Leu	Ile	Asn	Thr	Leu	Phe	His	Ser	Ser
				995					1000					1005	
Ala	Asp	Leu	Glu	Asp	Glu	Met	Val	Cys	Lys	Trp	Leu	Leu	Ser	Ser	
				1010					1015					1020	
Thr	Pro	Val	Met	Ser	Arg	Phe	Ala	Ala	Asp	Ile	Phe	Ser	Arg	Thr	
				1025					1030					1035	
Pro	Ser	Gly	Lys	Arg	Leu	Gln	Ile	Leu	Gly	Tyr	Leu	Glu	Gly	Thr	
				1040					1045					1050	
Arg	Thr	Leu	Leu	Ala	Ser	Lys	Ile	Ile	Asn	His	Asn	Thr	Glu	Thr	
				1055					1060					1065	
Pro	Ile	Leu	Asp	Arg	Leu	Arg	Lys	Ile	Thr	Leu	Gln	Arg	Trp	Ser	
				1070					1075					1080	
Leu	Trp	Phe	Ser	Tyr	Leu	Asp	His	Cys	Asp	Gln	Val	Leu	Ala	Asp	
				1085					1090					1095	
Ala	Leu	Thr	Gln	Ile	Thr	Cys	Thr	Val	Asp	Leu	Ala	Gln	Ile	Leu	
				1100					1105					1110	
Arg	Glu	Tyr	Thr	Trp	Ala	His	Ile	Leu	Glu	Gly	Arg	Gln	Leu	Ile	
				1115					1120					1125	
Gly	Ala	Thr	Leu	Pro	Cys	Ile	Leu	Glu	Gln	Leu	Asn	Val	Ile	Trp	
				1130					1135					1140	
Leu	Lys	Pro	Tyr	Glu	His	Cys	Pro	Lys	Cys	Ala	Lys	Ser	Ala	Asn	
				1145					1150					1155	

-continued

Pro Lys Gly Glu Pro Phe Val	Ser Ile Ala Ile Lys Lys His Val
1160	1165 1170
Val Ser Ala Trp Pro Asp Gln	Ser Arg Leu Ser Trp Thr Ile Gly
1175	1180 1185
Asp Gly Ile Pro Tyr Ile Gly	Ser Arg Thr Glu Asp Lys Ile Gly
1190	1195 1200
Gln Pro Ala Ile Lys Pro Lys	Cys Pro Ser Ala Ala Leu Arg Glu
1205	1210 1215
Ala Ile Glu Leu Thr Ser Arg	Leu Thr Trp Val Thr Gln Gly Gly
1220	1225 1230
Ala Asn Ser Asp Leu Leu Val	Lys Pro Phe Ile Glu Ala Arg Val
1235	1240 1245
Asn Leu Ser Val Gln Glu Ile	Leu Gln Met Thr Pro Ser His Tyr
1250	1255 1260
Ser Gly Asn Ile Val His Arg	Tyr Asn Asp Gln Tyr Ser Pro His
1265	1270 1275
Ser Phe Met Ala Asn Arg Met	Ser Asn Ser Ala Thr Arg Leu Val
1280	1285 1290
Val Ser Thr Asn Thr Leu Gly	Glu Phe Ser Gly Gly Gly Gln Ser
1295	1300 1305
Ala Arg Asp Ser Asn Ile Ile	Phe Gln Asn Val Ile Asn Phe Ala
1310	1315 1320
Val Ala Leu Phe Asp Leu Arg	Phe Arg Asn Val Ala Thr Ser Ser
1325	1330 1335
Ile Gln His His Arg Ala His	Leu His Leu Ser Lys Cys Cys Thr
1340	1345 1350
Arg Glu Val Pro Ala Gln Tyr	Leu Val Tyr Thr Ser Thr Leu Pro
1355	1360 1365
Leu Asp Leu Thr Arg Tyr Arg	Asp Asn Glu Leu Ile Tyr Asp Asp
1370	1375 1380
Asn Pro Leu Arg Gly Gly Leu	Asn Cys Asn Leu Ser Phe Asp Asn
1385	1390 1395
Pro Leu Phe Lys Gly Gln Arg	Leu Asn Ile Ile Glu Glu Asp Leu
1400	1405 1410
Ile Arg Leu Pro Tyr Leu Ser	Gly Trp Glu Leu Ala Lys Thr Val
1415	1420 1425
Ile Gln Ser Ile Ile Ser Asp	Ser Asn Asn Ser Ser Thr Asp Pro
1430	1435 1440
Ile Ser Ser Gly Glu Thr Arg	Ser Phe Thr Thr His Phe Leu Thr
1445	1450 1455
Tyr Pro Lys Ile Gly Leu Leu	Tyr Ser Phe Gly Ala Leu Ile Ser
1460	1465 1470
Tyr Tyr Leu Gly Asn Thr Ile	Ile Arg Thr Lys Lys Leu Thr Leu
1475	1480 1485
Asn Asn Phe Ile Tyr Tyr Leu	Ala Thr Gln Ile His Asn Leu Pro
1490	1495 1500
His Arg Ser Leu Arg Ile Leu	Lys Pro Thr Leu Lys His Ala Ser
1505	1510 1515
Val Ile Ser Arg Leu Ile Ser	Ile Asp Ser His Phe Ser Ile Tyr
1520	1525 1530

-continued

Ile Gly 1535	Gly Thr 1535	Ala Gly 1535	Asp Arg 1540	Gly Leu 1540	Ser Asp 1545	Ala Ala 1545	Arg
Leu Phe 1550	Leu Arg 1550	Thr Ala 1555	Ile Thr 1555	Val Phe 1560	Leu Gln 1560	Phe Val 1560	Arg
Lys Trp 1565	Ile Val 1570	Glu Arg 1570	Lys Thr 1570	Ala Ile 1575	Pro Leu 1575	Trp Val 1575	Ile
Tyr Pro 1580	Leu Glu 1585	Gly Gln 1585	Ser Pro 1590	Ser Pro 1590	Ile Asn 1590	Ser Phe 1590	Leu
His His 1595	Val Ile 1600	Ala Leu 1600	Leu Gln 1605	His Glu 1605	Ser Ser 1605	His Asp 1605	His
Val Cys 1610	Ala Ala 1615	Glu Ala 1615	His Ser 1620	Arg Val 1620	Glu Thr 1620	Phe Asp 1620	Asn
Leu Val 1625	Tyr Met 1630	Cys Lys 1630	Ser Thr 1635	Ala Ser 1635	Asn Phe 1635	Phe His 1635	Ala
Ser Leu 1640	Ala Tyr 1645	Trp Arg 1645	Ser Arg 1650	Ser Lys 1650	Asn Gln 1650	Asp Lys 1650	Arg
Glu Met 1655	Thr Lys 1660	Ile Leu 1660	Ser Leu 1665	Thr Gln 1665	Thr Glu 1665	Lys Lys 1665	Asn
Ser Phe 1670	Gly Tyr 1675	Thr Ala 1675	His Pro 1680	Glu Ser 1680	Thr Ala 1680	Val Leu 1680	Gly
Ser Leu 1685	Gln Thr 1690	Ser Leu 1690	Ala Pro 1695	Pro Pro 1695	Ser Ala 1695	Asp Glu 1695	Ala
Thr Tyr 1700	Asp Arg 1705	Lys Asn 1705	Lys Val 1710	Leu Lys 1710	Ala Ser 1710	Arg Pro 1710	Gly
Lys Tyr 1715	Ser Gln 1720	Asn Thr 1720	Thr Lys 1725	Ala Pro 1725	Pro Asn 1725	Gln Thr 1725	Ser
Cys Arg 1730	Asp Val 1735	Ser Pro 1735	Asn Ile 1740	Thr Gly 1740	Thr Asp 1740	Gly Cys 1740	Pro
Ser Ala 1745	Asn Glu 1750	Gly Ser 1750	Asn Ser 1755	Asn Asn 1755	Asn Asn 1755	Leu Val 1755	Ser
His Arg 1760	Ile Val 1765	Leu Pro 1765	Phe Phe 1770	Thr Leu 1770	Ser His 1770	Asn Tyr 1770	Asn
Glu Arg 1775	Pro Ser 1780	Ile Arg 1780	Lys Ser 1785	Glu Gly 1785	Thr Thr 1785	Glu Ile 1785	Val
Arg Leu 1790	Thr Arg 1795	Gln Leu 1795	Arg Ala 1800	Ile Pro 1800	Asp Thr 1800	Thr Ile 1800	Tyr
Cys Arg 1805	Phe Thr 1810	Gly Ile 1810	Val Ser 1815	Ser Ser 1815	Met His 1815	Tyr Lys 1815	Leu Asp
Glu Val 1820	Leu Trp 1825	Glu Phe 1825	Asp Asn 1830	Phe Lys 1830	Ser Ala 1830	Ile Thr 1830	Leu
Ala Glu 1835	Gly Glu 1840	Gly Ser 1840	Gly Ala 1845	Leu Leu 1845	Leu Leu 1845	Gln Lys 1845	Tyr
Lys Val 1850	Glu Thr 1855	Leu Phe 1855	Phe Asn 1860	Thr Leu 1860	Ala Thr 1860	Glu His 1860	Ser
Ile Glu 1865	Ala Glu 1870	Ile Ile 1870	Ser Gly 1875	Ile Thr 1875	Thr Pro 1875	Arg Met 1875	Leu
Leu Pro 1880	Ile Met 1885	Ser Arg 1885	Phe His 1890	Gly Gly 1890	Gln Ile 1890	Lys Val 1890	Thr
Leu Asn 1895	Asn Ser 1900	Ala Ser 1900	Gln Ile 1905	Thr Asp 1905	Ile Thr 1905	Asn Pro 1905	Ser
Trp Leu 1910	Ala Asp 1915	Gln Lys 1915	Ser Arg 1920	Ile Pro 1920	Lys Gln 1920	Val Glu 1920	Ile

-continued

1910	1915	1920
Ile Thr Met Asp Ala Glu Thr	Thr Glu Asn Ile Asn	Arg Ser Lys
1925	1930	1935
Leu Tyr Glu Ala Val Gln Gln	Leu Ile Val Ser His	Ile Asp Pro
1940	1945	1950
Asn Ala Leu Lys Val Val Val	Leu Lys Val Phe Leu	Ser Asp Ile
1955	1960	1965
Asp Gly Ile Leu Trp Leu Asn	Asp Asn Leu Thr Pro	Leu Phe Gly
1970	1975	1980
Leu Gly Tyr Leu Ile Lys Pro	Ile Thr Ser Ser Pro	Lys Ser Ser
1985	1990	1995
Glu Trp Tyr Leu Cys Leu Ser	Asn Leu Leu Ser Thr	Ser Arg Arg
2000	2005	2010
Leu Pro His Gln Ser His Thr	Thr Cys Met His Val	Ile Gln Thr
2015	2020	2025
Ala Leu Gln Leu Gln Ile Gln	Arg Ser Ser Tyr Trp	Leu Ser His
2030	2035	2040
Leu Val Gln Tyr Ala Asn His	Asn Leu His Leu Asp	Tyr Ile Asn
2045	2050	2055
Leu Gly Phe Pro Ser Leu Glu	Arg Val Leu Tyr His	Arg Tyr Asn
2060	2065	2070
Leu Val Asp Ser Gln Lys Gly	Pro Leu Thr Ser Ile	Val Gln His
2075	2080	2085
Leu Ala His Leu Gln Thr Glu	Ile Arg Glu Leu Val	Asn Asp Tyr
2090	2095	2100
Asn Gln Gln Arg Gln Ser Arg	Thr Gln Thr Tyr His	Phe Ile Lys
2105	2110	2115
Thr Ile Lys Gly Arg Ile Thr	Lys Leu Val Asn Asp	Tyr Leu Lys
2120	2125	2130
Phe Phe Leu Ile Ile Gln Ala	Leu Lys His Asn Cys	Thr Trp Gln
2135	2140	2145
Glu Glu Leu Arg Ala Leu Pro	Asp Leu Ile Ser Val	Cys Thr Arg
2150	2155	2160
Phe Tyr His Thr Arg Asn Cys	Ser Cys Glu Asn Arg	Phe Leu Val
2165	2170	2175
Gln Thr Leu Tyr Leu Ser Arg	Met Gln Asp Ser Glu	Ile Lys Leu
2180	2185	2190
Ile Asp Arg Leu Thr Gly Leu	Leu Ser Leu Cys Pro	Asn Gly Phe
2195	2200	2205
Phe Arg		
2210		

<210> SEQ ID NO 20

<211> LENGTH: 18959

<212> TYPE: DNA

<213> ORGANISM: Zaire ebolavirus

<220> FEATURE:

<221> NAME/KEY: misc_feature

<223> OTHER INFORMATION: Full viral sequence

<400> SEQUENCE: 20

cggacacaca aaaagaaga agaattttta ggatcttttg tgtcggaata actatgagga 60

agattaataa ttttcctctc attgaaattt atatcggaat ttaaattgaa attgttactg 120

-continued

taatcacacc	tggtttgttt	cagagccaca	tcacaaagat	agagaacaac	ctaggtctcc	180
gaagggagca	agggcatcag	tgtgctcagt	tgaaaatccc	ttgtcaacac	ctaggtctta	240
tcacatcaca	agttccacct	cagactctgc	agggtgatcc	aacaacctta	atagaaacat	300
tattgttaaa	ggacagcatt	agttcacagt	caaacaagca	agattgagaa	ttaaccttgg	360
ttttgaactt	gaacacttag	gggattgaag	attcaacaac	cctaaagctt	ggggtaaaac	420
attggaaata	gttaaaagac	aaattgctcg	gaatcacaaa	attccgagta	tggattctcg	480
tcctcagaaa	atctggatgg	cgccgagtct	cactgaatct	gacatggatt	accacaagat	540
cttgacagca	ggtctgtccg	ttcaacaggg	gattgttcgg	caaagagtca	tcccagtgtg	600
tcaagtaaac	aatcttgaag	aaatttgcca	acttatcata	caggcctttg	aagcaggtgt	660
tgattttcaa	gagagtgcgg	acagtttctt	tctcatgctt	tgtcttcac	atgcgtacca	720
gggagattac	aaacttttct	tggaaagtgg	cgcagtcaag	tatttgaag	ggcacgggtt	780
ccgttttgaa	gtcaagaagc	gtgatggagt	gaagcgcctt	gaggaattgc	tgccagcagt	840
atctagtgga	aaaaacatta	agagaacact	tgctgccatg	ccggaagagg	agacaactga	900
agctaagtc	ggtcagtttc	tctcctttgc	aagtctattc	cttccgaaat	tggtagtagg	960
agaaaaggct	tgcttgaga	aggttcaaag	gcaaattcaa	gtacatgcag	agcaaggact	1020
gatacaatat	ccaacagctt	ggcaatcagt	aggacacatg	atggtgattt	tccgtttgat	1080
gcgaacaaat	tttctgatca	aatttctcct	aatacaccaa	gggatgcaca	tggttgccgg	1140
gcatgatgcc	aacgatgctg	tgatttcaaa	ttcagtggct	caagctcggt	tttcaggctt	1200
attgattgtc	aaaacagtac	ttgatcatat	cctacaaaag	acagaacgag	gagttcgtct	1260
ccatcctctt	gcaaggaccg	ccaaggtaaa	aatgaggtg	aactccttta	aggetgcact	1320
cagctccctg	gccaagcatg	gagagtatgc	tcctttcgcc	cgacttttga	acctttctgg	1380
agtaaataat	cttgagcatg	gtcttttccc	tcaactatcg	gcaattgcac	tccgagtcgc	1440
cacagcacac	gggagtaccc	tcgcaggagt	aatgttgga	gaacagtatc	aacaactcag	1500
agaggctgcc	actgaggctg	agaagcaact	ccaacaatat	gcagagtctc	gcgaacttga	1560
ccatcttgga	cttgatgatc	aggaaaagaa	aattcttatg	aacttccatc	agaaaaagaa	1620
cgaaatcagc	ttccagcaaa	caaacgctat	ggtaactcta	agaaaagagc	gcctggccaa	1680
gctgacagaa	gctatcactg	ctgcgtcact	gccccaaa	agtggacatt	acgatgatga	1740
tgacgacatt	ccctttccag	gacccatcaa	tgatgacgac	aatcctggcc	atcaagatga	1800
tgatccgact	gactcacagg	atcagaccat	tccgatgtg	gtggttgatc	ccgatgatgg	1860
aagctacggc	gaataccaga	gttactcgga	aaacggcatg	aatgcaccag	atgacttggt	1920
cctattcgat	ctagacgagg	acgacgagga	cactaagcca	gtgcctaata	gatcgaccaa	1980
gggtggacaa	cagaagaaca	gtcaaaagg	ccagcatata	gagggcagac	agacacaatc	2040
caggccaatt	caaaatgtcc	caggccctca	cagaacaatc	caccacgcca	gtgcgccact	2100
cacggacaat	gacagaagaa	atgaaccctc	cggtcaacc	agcctcgcga	tgctgacacc	2160
aattaacgaa	gaggcagacc	cactggacga	tgccgacgac	gagacgtcta	gccttccgcc	2220
cttgaggtca	gatgatgaag	agcaggacag	ggacggaact	tccaaccgca	caccactgt	2280
cgccccaccg	gctcccgat	acagagatca	ctctgaaaag	aaagaactcc	cgcaagacga	2340
gcaacaagat	caggaccaca	ctcaagaggc	caggaaccag	gacagtgaca	acaccagtc	2400

-continued

agaacactct	tttgaggaga	tgtatcgcca	cattctaaga	tcacaggggc	catttgatgc	2460
tgttttgtag	tatcatatga	tgaaggatga	gcctgtagtt	ttcagtagca	gtgatggcaa	2520
agagtacacg	tatccagact	cccttgaaga	ggaatatcca	ccatggctca	ctgaaaaaga	2580
ggctatgaat	gaagagaata	gatttggtac	attggatggg	caacaatttt	attggccggg	2640
gatgaatcac	aagaataaat	tcattggcaat	cctgcaacat	catcagtga	tgagcatgga	2700
acaatgggat	gattcaaccg	acaaatagct	aacattaagt	agtcaaggaa	cgaaaacagg	2760
aaagaatttt	gatgtctaag	gtgtgaatta	ttatcacaat	aaaagtgatt	cttatttttg	2820
aatttaaagc	tagcttatta	ttactagccg	tttttcaaag	ttcaatttga	gtcttaatgc	2880
aaataggcgt	taagccacag	ttatagccat	aattgtaact	caatattcta	actagcgatt	2940
tatctaaatt	aaattacatt	atgcttttat	aacttaccta	ctagcctgcc	caacatttac	3000
acgatcggtt	tataattaag	aaaaaactaa	tgatgaagat	taaaaccttc	atcatcctta	3060
cgtaattga	attctctagc	actcgaagct	tattgtcttc	aatgtaaaag	aaaagctggg	3120
ctaacaagat	gacaactaga	acaaagggca	ggggccatac	tgccggccacg	actcaaacg	3180
acagaatgcc	aggccctgag	ctttcgggct	ggatctctga	gcagctaata	accggaagaa	3240
ttcctgtaag	cgacatcttc	tgtgatattg	agaacaatcc	aggattatgc	tacgcatccc	3300
aatgcaaca	aacgaagcca	aaccggaaga	cgcgaacag	tcaaacccaa	acggacccaa	3360
tttgcaatca	tagttttgag	gaggtagtac	aaacattggc	ttcattgggt	actgttgtgc	3420
aacaacaaac	catcgcatca	gaatcattag	aacaacgcat	tacgagctct	gagaatgggc	3480
taaagccagt	ttatgatatg	gcaaaaacaa	tctcctcatt	gaacaggggt	tgtgctgaga	3540
tggttgcaaa	atatgatctt	ctggatgatga	caaccgggtc	ggcaacagca	accgctgcgg	3600
caactgaggc	ttattgggcc	gaactgggtc	aaccaccacc	tggaccatca	ctttatgaag	3660
aaagtgcgat	tggggtaag	attgaatcta	gagatgagac	cgtccctcaa	agtgttaggg	3720
aggcattcaa	caatctaaac	agtaccactt	cactaactga	ggaaaatttt	gggaaacctg	3780
acatttcggc	aaaggatttg	agaaacatta	tgtatgatca	cttgccctgg	tttggaactg	3840
ctttccacca	attagtagaa	gtgatttgta	aattgggaaa	agatagcaac	tcattggaca	3900
tcattcatgc	tgagttccag	gccagcctgg	ctgaaggaga	ctctctctca	tgtgccttaa	3960
ttcaaattac	aaaaagagtt	ccaatcttcc	aagatgctgc	tccacctgtc	atccacatcc	4020
gctctcgagg	tgacattccc	cgagcttgcc	agaaaagctt	gcgtccagtc	ccaccatcgc	4080
ccaagattga	tcgaggttgg	gtatgtgttt	ttcagcttca	agatggtaaa	acacttgagc	4140
tcaaaatttg	agccaatctc	ccttcctctc	gaaagaggcg	aataatagca	gaggcttcaa	4200
ctgctgaact	ataggttagc	ttacattaat	gatacacttg	tgagtatcag	cctgggataa	4260
tataagtcaa	ttaaagcacc	aagataaaat	tgttcataat	tcgctagcag	cttaaaatat	4320
aaatgaata	ggagctatat	ctctgacagt	attataatca	attgttatta	agtaacccaa	4380
acaaaaagtg	atgaagatta	agaaaaacct	acctcggtcg	agagagtgtt	ttttcattaa	4440
ccttcactct	gtaaacgttg	agcaaaattg	ttaaaaatat	gaggcgggtt	atattgccta	4500
ctgctcctcc	tgaatatatg	gaggccatat	acctgttcag	gtcaaattca	acaattgcta	4560
gagggtggca	cagcaatata	ggcttctctg	caccggagtc	agtcaatggg	gacactccat	4620
cgaatccact	caggccaatt	gccgatgaca	ccatcgacca	tgccagccac	acaccaggca	4680

-continued

gtgtgtcatc agcattcatc cttgaagcta tggatgaatgt catatcgggc cccaaagtgc	4740
taatgaagca aattccaatt tggcttctc taggtgtcgc tgatcaaaag acctacagct	4800
ttgactcaac tacggccgcc atcatgcttg cttcatacac tatcacccat ttcggcaagg	4860
caaccaatcc acttgtcaga gtcaatcggc tgggtcctgg aatcccgat catccctca	4920
ggctcctgcg aattggaac caggctttcc tccaggagt cgttcttcg ccagtccaac	4980
tacccagta tttcaccttt gatttgacag cactcaaact gatcaccaa ccactgctg	5040
ctgcaacatg gaccgatgac actccaacag gatcaaatgg agcgttgcgt ccagggaattt	5100
catttcatcc aaaacttcgc ccattcttt tacccaacaa agtggggaag aaggggaaca	5160
gtgccgatct aacatctccg gagaaaatcc aagcaataat gacttcactc caggacttta	5220
agatcgttcc aattgatcca accaaaaata tcatgggaat cgaagtgcc gaaactctg	5280
tccacaagct gaccggaag aaggtgactt ctaaaaatgg acaaccaatc atccctgttc	5340
ttttgccaaa gtacattggg ttggaccgg tggctccagg agacctcacc atggtaatca	5400
cacaggattg tgacacgtgt cattctctg caagtcttcc agctgtgatt gagaagtaat	5460
tgcaataatt gactcagatc cagttttata gaatcttctc agggatagtg ataactcta	5520
tttagtaatc cgtccattag aggagacact ttttaattgat caatatacta aagggtgctt	5580
acaccattgt cttttttctc tcctaaatgt agaacttaac aaaagactca taatatactt	5640
gtttttaaag gattgattga tgaaagatca taactaataa cattacaaat aatcctacta	5700
taatcaatac ggtgattcaa atgttaatct ttctcattgc acatactttt tgcccttacc	5760
ctcaaattgc ctgcattgct acatctgagg atagccagtg tgacttggtg tggaaatgtg	5820
gagaaaaaat cgggaccat ttctaggttg ttcacaatcc aagtacagac attgcccttc	5880
taattaagaa aaaatcggcg atgaagatta agccgacagt gagcgtaatc ttcattcttc	5940
ttagattatt tgttttcag agtaggggtc gtcagggtcct tttcaatcgt gtaacaaaa	6000
taaaactccac tagaaggata ttgtggggca acaacacaaat gggcggttaca ggaatattgc	6060
agttacctcg tgatcgatcc aagaggacat cattctttct tgggtaatt atccttttcc	6120
aaagaacatt ttccatccca cttggagtca tccacaatag cacattacag gttagtgatg	6180
tcgacaaaact agtttgcgt gacaaaactgt catccacaaa tcaattgaga tcagttggac	6240
tgaatctcga agggaatgga gtggcaactg acgtgccatc tgcaactaaa agatggggct	6300
tcaggtcggg tgtcccacca aaggtggtca attatgaagc tggatgaatg gctgaaaact	6360
gtacaatct tgaaatcaaa aaactgacg ggagtgtgag tctaccagca gcgcagacg	6420
ggattcgggg cttcccccg tgccggtatg tgcacaaagt atcaggaacg ggaccgtgtg	6480
cggagactt tgcttccat aaagagggtg ctttcttct gtatgatcga cttgcttcca	6540
cagttatcta ccgaggaacg actttcgtg aaggtgtcgt tgcatttctg ataactgccc	6600
aagctaagaa ggactcttc agctcacac ccttgagaga gccggtcaat gcaacggagg	6660
accggtctag tggctactat tctaccacaa ttagatatca ggctaccggt tttggaacca	6720
atgagacaga gtacttgttc gaggttgaca atttgaccta cgtccaactt gaatcaagat	6780
tcacaccaca gtttctgctc cagctgaatg agacaatata tacaagtggg aaaaggagca	6840
ataccacggg aaaactaatt tggaaggta accccgaaat tgatacaaca atcggggagt	6900
gggccttctg ggaaactaaa aaaacctcac tagaaaaatt cgcagtgaag agttgtcttt	6960

-continued

cacagttgta	tcaaacggag	ccaaaaacat	cagtggtcag	agtccggcgc	gaacttcttc	7020
cgacccagg	accaacacaa	caactgaaga	ccacaaaatc	atggcttcag	aaaattcctc	7080
tgcaatggtt	caagtgcaca	gtcaaggaag	ggaagctgca	gtgtcgcatc	taacaaccct	7140
tgccacaatc	tccacgagtc	cccaatccct	cacaaccaa	ccaggctcgg	acaacagcac	7200
ccataatata	cccgtgtata	aacttgacat	ctctgaggca	actcaagttg	aacaacatca	7260
ccgcagaaca	gacaacgaca	gcacagcctc	cgacactccc	tctgccacga	ccgcagccgg	7320
acccccaaaa	gcagagaaca	ccaacacgag	caagagcact	gacttcctgg	acccccccac	7380
cacaacaagt	ccccaaaacc	acagcgagac	cgctggcaac	aacaacactc	atcaccaaga	7440
taccggagaa	gagagtgcc	gcagcgggaa	gctaggctta	attaccaata	ctattgctgg	7500
agtcgcagga	ctgatcacag	gcgggagaag	aactcgaaga	gaagcaattg	tcaatgctca	7560
acccaaatgc	aaccctaatt	tacattactg	gactactcag	gatgaagggtg	ctgcaatcgg	7620
actggcctgg	ataccatatt	tccggccagc	agccgaggga	atttacatag	aggggcta	7680
gcacaatcaa	gatggtttaa	tctgtgggtt	gagacagctg	gccaacgaga	cgactcaagc	7740
tcttcaactg	ttcctgagag	ccacaactga	gctacgcacc	ttttcaatcc	tcaaccgtaa	7800
ggcaattgat	ttcttgctgc	agcgatgggg	cggcacatgc	cacattcttg	gaccggactg	7860
ctgtatcgaa	ccacatgatt	ggaccaagaa	cataacagac	aaaattgatc	agattattca	7920
tgattttgtt	gataaaaccc	ttccggacca	gggggacaat	gacaattggt	ggacaggatg	7980
gagacaatgg	ataccggcag	gtattggagt	tacaggcggt	ataattgcag	ttatcgcttt	8040
attctgtata	tgcaaatgtg	tcttttagtt	tttcttcaga	ttgcttcattg	gaaaagctca	8100
gcctcaaadc	aatgaaacca	ggatttaatt	atatggatta	cttgaatcta	agattacttg	8160
acaaatgata	atataatata	ctggagcttt	aaacatagcc	aatgtgattc	taactccttt	8220
aaactcacag	ttaatcataa	acaaggtttg	acatcaatct	agttatctct	ttgagaatga	8280
taaaacttgat	gaagattaag	aaaaaggtaa	tctttcgatt	atctttaate	ttcatccttg	8340
attctacaat	catgacagtt	gtcttttagtg	acaaggga	gaagcctttt	tattaagttg	8400
taataatcag	atctgcgaac	cggtagagtt	tagttgcaac	ctaacacaca	taaagcattg	8460
gtcaaaaagt	caatagaaat	ttaaacagtg	agtgagagaca	acttttaaat	ggaagcttca	8520
tatgagagag	gacgcccacg	agctgccaga	cagcattcaa	gggatggaca	cgaccaccat	8580
gttcgagcac	gatcatcatc	cagagagaat	tatcgagggtg	agtaccgtca	atcaaggagc	8640
gcctcacaag	tgccgcttcc	tactgtatct	cataagaaga	gagttgaacc	attaacagtt	8700
cctccagcac	ctaaagacat	atgtccgacc	ttgaaaaaag	gatttttggtg	tgacagtagt	8760
ttttgcaaaa	aagatcacca	gttgagaggt	ttaactgata	gggaattact	cctactaatc	8820
gcccgtgaaga	cttgtggatc	agtagaaca	caattaaata	taactgcacc	caaggactcg	8880
cgcttagcaa	atccaacggc	tgatgatttc	cagcaagagg	aaggtccaaa	aattaccttg	8940
ttgacactga	tcaagacggc	agaacactgg	gcgagacaag	acatcagaac	catagaggat	9000
tcaaaattaa	gagcattggt	gactctatgt	gctgtgatga	cgaggaaatt	ctcaaaatcc	9060
cagctgagtc	ttttatgtga	gacacaccta	aggcgcgagg	ggcttgggca	agatcaggca	9120
gaaccggttc	tcgaagtata	tcaacgatta	cacagtgata	aaggaggcag	ttttgaagct	9180
gcactatggc	aacaatggga	ccgacaatcc	ctaattatgt	ttatcactgc	attcttgaat	9240

-continued

attgctctcc agttaccgtg tgaaagtctt gctgtcgttg tttcaggggt aagaacattg	9300
gttctctcaat cagataatga ggaagcttca accaaccctgg ggacatgctc atgggtctgat	9360
gagggtagccc cttaataagg ctgactaaaa cactatataa ccttctactt gatcacaata	9420
ctccgtatac ctatcatcat atatttaatc aagacgatat cctttaaaac ttattcagta	9480
ctataatcac tctcgtttca aattaataag atgtgcatga ttgccctaat atatgaagag	9540
gtatgatata accctaacag tgatcaaaga aaatcataat ctcgatcgcc tcgtaatata	9600
acctgccaa catacctctt gcacaaagtg attcttgtac acaataatg ttttactcta	9660
caggaggtag caacgatcca tcccacaaa aaataagtat ttcattgactt actaatgatc	9720
tcttaaaata ttaagaaaaa ctgacggaac ataaattctt tatgcttcaa gctgtggagg	9780
aggtgtttgg tattggctat tgttatatta caatcaataa caagcttgta aaaatattgt	9840
tcttgtttca agaggtagat tgtgaccgga aatgctaaac taatgatgaa gattaatgag	9900
gaggtctgat aagaataaac cttattatct agattaggcc ccaagaggca ttcttcatct	9960
ccttttagca agtactatt tcagggtagt ccaattagtg gcacgtcttt tagctgtata	10020
tcagtcgccc ctgagatacg ccacaaaagt gtctctaagc taaattgggc tgtacacatc	10080
ccatacattg tattaggggc aataatatct aattgaactt agccgtttta aatttagtgc	10140
ataaatctgg gtaaacacca ccaggtaaac tccattggct gaaaagaagc ttacctacaa	10200
cgaacatcac tttgagcgcc ctccacatta aaaaatagga acgtcgttcc aacaatcgag	10260
cgcaagggtt caaggttgaa ctgagagtgt ctagacaaca aaatattgat actccagaca	10320
ccaagcaaga cctgagaaaa aacctaggct aaagctacgg gacgatataa tctaatatcg	10380
cccaaaaagg acctggagaa aggggttgtc ttaagcgacc tctgtaactt cttagttagc	10440
caaaactatc aggggttgaa ggtttattgg gctgggtatt agtttgatgt gactcacaaa	10500
ggaatggccc tattgcatag actgaaaact aatgactttg ccctgcatg gtcaatgaca	10560
aggaatctct ttcctctatt atttcaaaat ccgaattcca caattgaatc accgtgtggg	10620
gcattgagag tcatccttgc agcagggata caggaccagc tgattgacca gtctttgatt	10680
gaacccttag caggagccct tggctctgat tctgattggc tgctaacaac caacactaac	10740
catttcaaca tgcgaacaca acgtgtcaag gaacaattga gcctaaaaat gctgtcgttg	10800
attcgatcca atattctcaa gtttattaac aaattggatg ctctacatgt cgtgaactac	10860
aacggattgt tgagcagtat tgaaattgga actcaaaatc atacaatcat cataactcga	10920
actaacatgg gttttctggt ggagctccaa gaaccgaca aatcggaat gaaccgcatg	10980
aagcctgggc cggcgaaatt tccctcctt catgagtcca cactgaaagc atttacacaa	11040
ggatcctcga cagcaatgca aagtttgatt cttgaattta atagctctct tgcctatctaa	11100
ctaaggtaga atacttcata ttgagctaac tcatatatgc tgactcaata gttatcttga	11160
catctctgct ttcataatca gatataatg cataataaat aaatactcat atttcttgat	11220
aatgtgttta accacagata aatcctcact gtaagccagc ttccaagttg acacccttac	11280
aaaaaccagg actcagaatc cctcaacaaa gagattccaa gacaacatca tagaattgct	11340
ttatttatat aataagcatt ttatcaccag aaatcctata tactaaatgg ttaattgtaa	11400
ctgaaccgcg aggtcacatg tgttaggttt cacagattct atatatattc aactctatac	11460
tcgtaattaa cattagataa gtagattaag aaaaaagcct gaggaagatt aagaaaaact	11520

-continued

```

gcttattggg tctttccgtg ttttagatga agcagttgaa attcttcctc ttgatattaa 11580
atgggtacac aacataccca ataccagac gctaggttat catcaccaat tgtattggac 11640
caatgtgacc tagtcactag agcttgccgg ttatattcat catactccct taatccgcaa 11700
ctacgcaact gtaaactccc gaaacatatc taccgtttga aatacgaagt aactgttacc 11760
aagttcttga gtgatgtacc agtggcgaca ttgcccatag atttcatagt cccagttctt 11820
ctcaaggcac tgtcaggcaa tggattctgt cctgttgagc cgcggtgcca acagttctta 11880
gatgaaatca ttaagtacac aatgcaagat gctctcttct tgaatatata tctcaaaaat 11940
gtgggtgctc aagaagactg tgttgatgaa cactttcaag agaaaatctt atcttcaatt 12000
cagggcaatg aatttttaca tcaaatgttt ttctggtagt atctggctat tttaactcga 12060
aggggtagat taaatcgagg aaactctaga tcaacatggt ttgttcata tgatttaata 12120
gacatcttag gctatgggga ctatgttttt tggaagatcc caatttcaat gttaccactg 12180
aacacacaag gaatccccc tgctgctatg gactggatc aggcatcagt attcaaagaa 12240
gcggttcaag ggcatacaca cattgtttct gtttctactg ccgacgtctt gataatgtgc 12300
aaagatttaa ttacatgtcg attcaacaca actctaactt caaaaatagc agagattgag 12360
gatccagttt gttctgatta tcccaatttt aagatttgtt ctatgcttta ccagagcgga 12420
gattacttac tctccatatt agggctctgat ggggtataaaa ttattaagtt cctcgaaacca 12480
ttgtgcttgg ccaaaattca attatgctca aagtacactg agaggaaggg ccgattctta 12540
acacaaatgc atttagctgt aaatcacacc ctagaagaaa ttacagaaat gcgtgcacta 12600
aagccttcac aggcctcaaaa gatccgtgaa ttccatagaa cattgataag gctggagatg 12660
acgcacaaac aacttttgta gctattttcc attcaaaaac actgggggca tctgtgcta 12720
catagtgaag cagcaatcca aaaagttaaa aaacatgcta cgggtgctaaa agcattacgc 12780
cctatagtga ttttcgagac atactgtgtt tttaaatata gtattgcaa acattatttt 12840
gatagtcaag gatcttggtg cagtgttact tcagatagga atctaaccac ggttcttaat 12900
tcttatatca aaagaaatca attccctccg ttgccaatga ttaaagaact actatgggaa 12960
ttttaccacc ttgaccaccc tccacttttc tcaacaaaa ttattagtga cttaagtatt 13020
ttataaaaag acagagctac cgcagtagaa aggacatgct gggatgcagt attcgagcct 13080
aatgttctag gatataatcc acctcacaaa tttagtacta aacgtgtacc ggaacaattt 13140
ttagagcaag aaaacttttc tattgagaat gttctttctc acgcacaaaa actcgagtat 13200
ctactaccac aatatcgga cttttctttc tcattgaaag agaaagagtt gaatgtaggt 13260
agaaccttcg gaaaattgcc ttatccgact cgcaatgttc aaacacttg tgaagctctg 13320
ttagctgatg gtcttgctaa agcattttct agcaatatga tggtagttac ggaacgtgag 13380
caaaaagaaa gcttattgca tcaagcatca tggcaccaca caagtgatga ttttggtgaa 13440
catgccacag ttagagggag tagctttgta actgatttag agaaatacaa tcttgcatth 13500
agatatgagt ttacagcacc ttttatagaa tattgcaacc gttgctatgg tgtaagaat 13560
gtttttaatt ggatgcatta tacaatccca cagtgttata tgcattgcag tgattattat 13620
aatccaccac ataacctcac actggagaat cgagacaacc cccccgaagg gcctagttca 13680
tacaggggtc atatgggagg gattgaagga ctgcaacaaa aactctggac aagtatttca 13740
tgtgctcaaa tttctttagt tgaaattaag actggtttta agttacgctc agctgtgatg 13800

```

-continued

```

ggtgacaatc agtgcattac tgttttatca gtcttccctc tagagactga cgcagacgag 13860
caggaacaga gcgcgaaga caatgcagcg aggggtggccg ccagcctagc aaaagttaca 13920
agtgcctgtg gaatcttttt aaaacctgat gaaacatttg tacattcagg ttttatctat 13980
tttgaaaaaa aacaatattt gaatggggtc caattgcctc agtcccttaa aacggctaca 14040
agaatggcac cattgtctga tgcaattttt gatgatcttc aagggaccct ggctagtata 14100
ggcactgctt ttgagcgatc catctctgag acacgacata tctttccttg caggataacc 14160
gcagctttcc atacgttttt ttcggtgaga atcttgcaat atcatcatct cgggttcaat 14220
aaaggttttg accttggaac gttaacactc ggcaaacctc tggatttcgg aacaatatca 14280
ttggcactag cggtagcgca ggtgcttgga gggttatcct tcttgaatcc tgagaaatgt 14340
ttctaccgga atctaggaga tccagttacc tcaggcttat tccagttaaa aacttatctc 14400
cgaatgattg agatggatga tttattctta cctttaattg cgaagaacctc tgggaactgc 14460
actgccattg actttgtgct aaatcctagc ggattaaatg tcctggggtc gcaagactta 14520
acttcatttc tgcgccagat tgtacgcagg accatcacct taagtgcgaa aaacaaactt 14580
attaatacct tatttcagtc gtcagctgac ttcgaagacg aaatggtttg taaatggcta 14640
ttatcatcaa ctctgttat gagtcgtttt gcggccgata tcttttcacg cagcccgagc 14700
gggaagcgat tgcaattctt aggatacctg gaaggaacac gcacattatt agcctctaag 14760
atcatcaaca ataatacaga gacaccggtt ttggacagac tgaggaaaat aacattgcaa 14820
aggtggagcc tatggtttag ttatcttgat cattgtgata atatcctggc ggaggcttta 14880
acccaaataa ctgacacagt tgatttagca cagattctga gggaaattc atgggctcat 14940
attttagagg gaagacctct tattggagcc aactcccat gtatgattga gcaattcaaa 15000
gtgttttggc tgaaccctta cgaacaatgt ccgcagtggt caaatgcaaa gcaaccaggt 15060
gggaaaccat tcgtgtcagt ggcagtcaag aaacatattg ttagtgcatg gccgaacgca 15120
tcccgaataa gctggactat cggggatgga atcccataca ttggatcaag gacagaagat 15180
aagataggac aacctgctat taaacaaaaa tgtccttcgc cagccttaag agaggccatt 15240
gaattggcgt ccggttaaac atgggtaact caaggcagtt cgaacagtga cttgctaata 15300
aaaccatttt tgggaagcac agtaaattta agtgttcaag aaatacttca aatgacctc 15360
tcacattact caggaaatat tgttcacagg tacaacgac aatacagtcc tcattctttc 15420
atggccaatc gtatgagtaa ttcagcaacg cgattgattg tttctacaaa cactttaggt 15480
gagttttcag gaggtggcca gtctgcacgc gacagcaata ttattttcca gaatgttata 15540
aattatgcag ttgcactggt cgatattaaa tttagaaaca ctgaggctac agatatccaa 15600
tataatcggt ctacacctta tctaactaag tgttgacccc ggggaagtacc agctcagtat 15660
ttaacataca catctacatt ggatttagat ttaacaagat accgagaaaa cgaattgatt 15720
tatgacagta atcctctaaa aggaggactc aattgcaata tctcattcga taatccattt 15780
ttccaaggta aacggctgaa cattatagaa gatgatctta ttcgactgcc tcacttatct 15840
ggatgggagc tagccaagac catcatgcaa tcaattattt cagatagcaa caattcatct 15900
acagacccaa ttagcagtgg agaaacaaga tcattcacta ccattttctt aacttatccc 15960
aagataggac ttctgtacag ttttggggcc tttgtaagtt attatcttgg caatacaatt 16020
cttcggacta agaaattaac acttgacaat tttttatatt acttaactac tcaaattcat 16080

```

-continued

aatctaccac atcgctcatt gcgaatactt aagccaacat tcaaacatgc aagcgttatg	16140
tcacgggttaa tgagtattga tcttcatttt tctattttaca taggcgggtgc tgcagggtgac	16200
agaggactct cagatgcggc cagggtattt ttgagaacgt ccattttcatc ttttcttaca	16260
tttgtaaaag aatggataat taatcgcgga acaattgtcc ctttatggat agtatatccg	16320
ctagagggtc aaaacccaac acctgtgaat aattttctct atcagatcgt agaactgctg	16380
gtgcatgatt catcaagaca acaggctttt aaaactacca taagtgatca tgtacatcct	16440
cacgacaatc ttgtttacac atgtaagagt acagccagca atttcttcca tgcacattg	16500
gcgtactgga ggagcagaca cagaacagc aaccgaaaat acttggaag agactcttca	16560
actggatcaa gcacaaacaa cagtgatggt catattgaga gaagtcaaga acaaaccacc	16620
agagatccac atgatggcac tgaacggaat ctagtccctac aaatgagcca tgaataaaaa	16680
agaacgacaa ttccacaaga aaacacgcac cagggtccgt cgttcagtc ctttctaagt	16740
gactctgctt gtggtacagc aaatccaaaa ctaaatttcg atcgatcgag acacaatgtg	16800
aaatttcagg atcataactc ggcatccaag agggaaggtc atcaataat ctcacaccgt	16860
ctagtccctac ctttctttac attatctcaa gggacacgcc aattaacgtc atccaatgag	16920
tcacaaacc aagacgagat atcaaagta ttacggcaat tgagatccgt cattgatacc	16980
acagtttatt gtagatttac cggatatagtc tcgccatgc attacaaact tgatgaggtc	17040
ctttgggaaa tagagagttt caagtcggct gtgacgctag cagagggaga aggtgctggt	17100
gccttactat tgattcagaa ataccaagtt aagaccttat tttcaacac gctagctact	17160
gagtcagta tagagtcaga aatagtatca ggaatgacta ctctaggat gcttctacct	17220
gttatgtcaa aattccataa tgaccaaatt gagattatc ttaacaactc agcaagccaa	17280
ataacagaca taacaaatcc tacttggtt aaagaccaa gagcaaggct acctaaagcaa	17340
gtcagggta taacctgga tgcagagaca acagagaata taaacagatc gaaattgtac	17400
gaagctgtat ataaattgat cttacaccat attgatccta gcgtattgaa agcagtggtc	17460
cttaaagtct ttctaagtga tactgaggt atgttatggc taaatgataa tttagccccg	17520
ttttttgcc ctggttattt aattaagcca ataacgtcaa gtgctagatc tagtgagtgg	17580
tatctttgtc tgacgaact cttatcaact acacgtaaga tgccacacca aaacctctc	17640
agttgtaaac aggtaatact tacggcattg caactgcaa ttcaacgaag ccataactgg	17700
ctaagtcatt taactcagta tgctgactgt gagttacatt taagttatat ccgccttgg	17760
tttccatcat tagagaaagt actataccac aggtataacc tgcgcatc aaaaagaggt	17820
ccactagtct ctatcactca gcacttagca catcttagag cagagattcg agaattaact	17880
aatgattata atcaacacgc acaaagtcgg actcaaacat atcactttat tctgactgca	17940
aaaggacgaa tcacaaaact agtcaatgat tatttaaaat tctttcttat tgtgcaagca	18000
ttaaaacata atgggacatg gcaagctgag ttaagaaat taccagagtt gattagtgtg	18060
tgcaataggt tctaccatat tagagattgc aattgtgaag aacgtttctt agttcaaac	18120
ttatatttac atagaatgca ggattctgaa gtaagctta tcgaaaggct gacagggtt	18180
ctgagtttat ttccggatgg tctctacagg tttgattgaa ttaccgtgca tagtatcctg	18240
atacttgcaa aggttggtta ttaacatata gattataaaa aactcataaa ttgctctcat	18300
acatcatatt gatctaactc caataaaca ctatttaaat aacgaaagga gtcctatat	18360

-continued

tatatactat atttagcctc tctccctgcg tgataatcaa aaaattcaca atgcagcatg 18420
tgtgacatat tactgccgca atgaatttaa cgcaacataa taaactctgc actctttata 18480
attaagcttt aacgaaaggt ctgggctcat attgttattg atataataat gttgtatcaa 18540
tatectgtca gatggaatag tgttttgggt gataacacaa cttcttaaaa caaaattgat 18600
ctttaagatt aagtttttta taattatcat tactttaatt tgcgtttta aaaacgggtga 18660
tagccttaat ctttgtgtaa aataagagat taggtgtaat aaccttaaca ttttgttcta 18720
gtaagctact atttcataca gaatgataaa attaaaagaa aaggcaggac tgtaaaatca 18780
gaaatacctt ctttacaata tagcagacta gataataatc ttcgtgttaa tgataattaa 18840
gacattgacc acgctcatca gaaggctcgc cagaataaac gttgcaaaaa ggattcctgg 18900
aaaaatgggc gcacacaaaa atttaaaat aaatctatct cttctttttt gtgtgtcca 18959

<210> SEQ ID NO 21
<211> LENGTH: 20
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for Sudan ebola BMG
<220> FEATURE:
<221> NAME/KEY: modified_base
<222> LOCATION: (8)..(8)
<220> FEATURE:
<221> NAME/KEY: misc_feature
<222> LOCATION: (8)..(8)
<223> OTHER INFORMATION: n is a, c, g, or t

<400> SEQUENCE: 21

gccatggntt caggtttgag 20

<210> SEQ ID NO 22
<211> LENGTH: 22
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Sudan ebola BMG
<220> FEATURE:
<221> NAME/KEY: modified_base
<222> LOCATION: (4)..(4)
<223> OTHER INFORMATION: I
<220> FEATURE:
<221> NAME/KEY: misc_feature
<222> LOCATION: (4)..(4)
<223> OTHER INFORMATION: n is a, c, g, or t

<400> SEQUENCE: 22

ggtnacattg ggcaacaatt ca 22

<210> SEQ ID NO 23
<211> LENGTH: 26
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR probe for Sudan ebola BMG
<220> FEATURE:
<221> NAME/KEY: misc_feature
<222> LOCATION: (1)..(1)
<223> OTHER INFORMATION: Fluorescein (FAM)
<220> FEATURE:
<221> NAME/KEY: misc_feature
<222> LOCATION: (26)..(26)
<223> OTHER INFORMATION: Black hole quencher dye (BHQ1)

-continued

<400> SEQUENCE: 23

acggtgcaca ttctcctttt ctcgga

26

<210> SEQ ID NO 24

<211> LENGTH: 22

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR forward primer for Ebola Bundibugyo
fragment A

<400> SEQUENCE: 24

gtgagacaaa gaatcattcc tg

22

<210> SEQ ID NO 25

<211> LENGTH: 23

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR reverse primer for Ebola Bundibugyo
fragment A

<400> SEQUENCE: 25

catcaattgc tcagagatcc acc

23

<210> SEQ ID NO 26

<211> LENGTH: 21

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR forward primer for Ebola Bundibugyo
fragment B

<400> SEQUENCE: 26

ccaacaacac tgcattgtaag t

21

<210> SEQ ID NO 27

<211> LENGTH: 20

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR reverse primer for Ebola Bundibugyo
fragment B

<400> SEQUENCE: 27

aggctgcggtt aatcttcac

20

<210> SEQ ID NO 28

<211> LENGTH: 21

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR forward primer for Ebola Bundibugyo
fragment C

<400> SEQUENCE: 28

gatggttgag ttactttccg g

21

<210> SEQ ID NO 29

<211> LENGTH: 20

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR reverse primer for Ebola Bundibugyo

-continued

fragment C

<400> SEQUENCE: 29

gtcttgagtc atcaatgccc

20

<210> SEQ ID NO 30

<211> LENGTH: 18

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR forward primer for Ebola Bundibugyo
fragment D

<400> SEQUENCE: 30

ccaccagcac caaaggac

18

<210> SEQ ID NO 31

<211> LENGTH: 22

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR reverse primer for Ebola Bundibugyo
fragment D

<400> SEQUENCE: 31

ctatcgga tgtaactatt gg

22

<210> SEQ ID NO 32

<211> LENGTH: 19

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR forward primer for Ebola Bundibugyo
fragment E

<400> SEQUENCE: 32

gccgttgtag aggacacac

19

<210> SEQ ID NO 33

<211> LENGTH: 26

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR reverse primer for Ebola Bundibugyo
fragment E

<400> SEQUENCE: 33

cacattaaat tgttctaaca tgcaag

26

<210> SEQ ID NO 34

<211> LENGTH: 23

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

<220> FEATURE:

<223> OTHER INFORMATION: PCR forward primer for Ebola Bundibugyo
fragment F

<400> SEQUENCE: 34

cctaggttat ttagaaggga cta

23

<210> SEQ ID NO 35

<211> LENGTH: 24

<212> TYPE: DNA

<213> ORGANISM: Artificial Sequence

-continued

<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Ebola Bundibugyo fragment F

<400> SEQUENCE: 35

ggtagatgta ttgacagcaa tatc 24

<210> SEQ ID NO 36
<211> LENGTH: 21
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR primer for Ebola Uganda 692(-)

<400> SEQUENCE: 36

acaaaaagct atctgcacta t 21

<210> SEQ ID NO 37
<211> LENGTH: 20
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR primer for Ebola Uganda 18269(+)

<400> SEQUENCE: 37

ctcagaagca aaattaatgg 20

<210> SEQ ID NO 38
<211> LENGTH: 23
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for Cote d'Ivoire ebola virus fragment A

<400> SEQUENCE: 38

gtgtgcgaat aactatgagg aag 23

<210> SEQ ID NO 39
<211> LENGTH: 22
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Cote d'Ivoire ebola virus fragment A

<400> SEQUENCE: 39

gtctgtgcaa tgttgatgaa gg 22

<210> SEQ ID NO 40
<211> LENGTH: 22
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for Cote d'Ivoire ebola virus fragment B

<400> SEQUENCE: 40

catgaaaacc acactcaaca ac 22

<210> SEQ ID NO 41
<211> LENGTH: 24
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence

-continued

<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Cote d'Ivoire ebola virus
fragment B

<400> SEQUENCE: 41

gttgccttaa tcttcatcaa gttc 24

<210> SEQ ID NO 42
<211> LENGTH: 22
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for Cote d'Ivoire ebola virus
fragment C

<400> SEQUENCE: 42

ggctataatg aatttcctcc ag 22

<210> SEQ ID NO 43
<211> LENGTH: 22
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for ebola cote d'Ivoire virus
fragment C

<400> SEQUENCE: 43

caagtgtatt tgtggccta gc 22

<210> SEQ ID NO 44
<211> LENGTH: 20
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Cote d'Ivoire ebola virus
fragment C

<400> SEQUENCE: 44

gctggaatag gaatcacagg 20

<210> SEQ ID NO 45
<211> LENGTH: 21
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Cote d'Ivoire ebola virus
fragment D

<400> SEQUENCE: 45

cggtagtcta cagttcttta g 21

<210> SEQ ID NO 46
<211> LENGTH: 25
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for Cote d'Ivoire ebola virus
fragment E

<400> SEQUENCE: 46

gacaaagaga ttagattagc tatag 25

<210> SEQ ID NO 47
<211> LENGTH: 22

-continued

<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Cote d'Ivoire ebola virus
fragment E

<400> SEQUENCE: 47

gtaatgagaa ggtgtcattt gg 22

<210> SEQ ID NO 48
<211> LENGTH: 22
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for Cote d'Ivoire ebola virus
fragment F

<400> SEQUENCE: 48

cacgacttag ttggacaatt gg 22

<210> SEQ ID NO 49
<211> LENGTH: 23
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Cote d'Ivoire ebola virus
fragment F

<400> SEQUENCE: 49

cagacactaa ttagatctgg aag 23

<210> SEQ ID NO 50
<211> LENGTH: 21
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for Cote d'Ivoire ebola virus
fragment G

<400> SEQUENCE: 50

cggacacaca aaaagaawra a 21

<210> SEQ ID NO 51
<211> LENGTH: 21
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Cote d'Ivoire ebola virus
fragment G

<400> SEQUENCE: 51

cgttcttgac cttagcagtt c 21

<210> SEQ ID NO 52
<211> LENGTH: 22
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for Cote d'Ivoire ebola virus
fragment H

<400> SEQUENCE: 52

gcactataag ctcgatgaag tc 22

-continued

<210> SEQ ID NO 53
<211> LENGTH: 20
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Cote d'Ivoire ebola virus
fragment H

<400> SEQUENCE: 53

tggacacaca aaaargaraa 20

<210> SEQ ID NO 54
<211> LENGTH: 21
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for Cote d'Ivoire ebola virus
gap between fragments C and D

<400> SEQUENCE: 54

ctgagaggat ccagaagaaa g 21

<210> SEQ ID NO 55
<211> LENGTH: 22
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR reverse primer for Cote d'Ivoire ebola
virus gap between fragments C and D

<400> SEQUENCE: 55

gtgtaagcgt tgatatacct cc 22

<210> SEQ ID NO 56
<211> LENGTH: 22
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for ebola uganda virus
EboU965(+)

<400> SEQUENCE: 56

gagaaaaggc ctgtctggag aa 22

<210> SEQ ID NO 57
<211> LENGTH: 24
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR forward primer for ebola uganda virus
EboU1039(-)

<400> SEQUENCE: 57

tcgggtattg aatcagacct tggt 24

<210> SEQ ID NO 58
<211> LENGTH: 26
<212> TYPE: DNA
<213> ORGANISM: Artificial Sequence
<220> FEATURE:
<223> OTHER INFORMATION: PCR probe for ebola uganda virus EboU989

<400> SEQUENCE: 58

ttcaacgaca aatccaagtg cacgca 26

-continued

<210> SEQ ID NO 59
<211> LENGTH: 302
<212> TYPE: PRT
<213> ORGANISM: Bundibugyo ebolavirus
<220> FEATURE:
<221> NAME/KEY: misc_feature
<223> OTHER INFORMATION: SSGP viral protein

<400> SEQUENCE: 59

Met Val Thr Ser Gly Ile Leu Gln Leu Pro Arg Glu Arg Phe Arg Lys
1 5 10 15

Thr Ser Phe Phe Val Trp Val Ile Ile Leu Phe His Lys Val Phe Pro
20 25 30

Ile Pro Leu Gly Val Val His Asn Asn Thr Leu Gln Val Ser Asp Ile
35 40 45

Asp Lys Leu Val Cys Arg Asp Lys Leu Ser Ser Thr Ser Gln Leu Lys
50 55 60

Ser Val Gly Leu Asn Leu Glu Gly Asn Gly Val Ala Thr Asp Val Pro
65 70 75 80

Thr Ala Thr Lys Arg Trp Gly Phe Arg Ala Gly Val Pro Pro Lys Val
85 90 95

Val Asn Tyr Glu Ala Gly Glu Trp Ala Glu Asn Cys Tyr Asn Leu Asp
100 105 110

Ile Lys Lys Ala Asp Gly Ser Glu Cys Leu Pro Glu Ala Pro Glu Gly
115 120 125

Val Arg Gly Phe Pro Arg Cys Arg Tyr Val His Lys Val Ser Gly Thr
130 135 140

Gly Pro Cys Pro Glu Gly Tyr Ala Phe His Lys Glu Gly Ala Phe Phe
145 150 155 160

Leu Tyr Asp Arg Leu Ala Ser Thr Ile Ile Tyr Arg Ser Thr Thr Phe
165 170 175

Ser Glu Gly Val Val Ala Phe Leu Ile Leu Pro Glu Thr Lys Lys Asp
180 185 190

Phe Phe Gln Ser Pro Pro Leu His Glu Pro Ala Asn Met Thr Thr Asp
195 200 205

Pro Ser Ser Tyr Tyr His Thr Val Thr Leu Asn Tyr Val Ala Asp Asn
210 215 220

Phe Gly Thr Asn Met Thr Asn Phe Leu Phe Gln Val Asp His Leu Thr
225 230 235 240

Tyr Val Gln Leu Glu Pro Arg Phe Thr Pro Gln Phe Leu Val Gln Leu
245 250 255

Asn Glu Thr Ile Tyr Thr Asn Gly Arg Arg Ser Asn Thr Thr Gly Thr
260 265 270

Leu Ile Trp Lys Val Asn Pro Thr Val Asp Thr Gly Val Gly Glu Trp
275 280 285

Ala Phe Trp Glu Asn Lys Lys Leu His Lys Asn Pro Phe Lys
290 295 300

1. An isolated hEbola virus comprising a nucleic acid molecule comprising a nucleotide sequence of:

- a) a nucleotide sequence set forth in SEQ ID NOS: 1 or 10;
- b) a nucleotide sequence hybridizing under stringent conditions to SEQ ID NOS: 1 or 10; or
- c) a nucleotide sequence of at least 70%-99% identity to the SEQ ID NOS: 1 or 10, with the proviso that said nucleotide sequence is not SEQ ID NO: 20.

2. An isolated hEbola virus having Centers for Disease Control Deposit Accession No. 200706291.

3. The hEbola virus of claim 1 which is killed.

4. The hEbola virus of claim 1 which is an attenuated hEbola virus.

5. The virus of claim 4 wherein at least one property of the attenuated hEbola virus is reduced from among infectivity, replication ability, protein synthesis ability, assembling ability or cytopathic effect.

6. An isolated nucleic acid molecule comprising the nucleotide sequence of SEQ ID NOS: 1 or 10 or a complement thereof, or a fragment thereof wherein said fragment comprises a nucleotide sequence of between 4 and 4900 contiguous nucleotides of the nucleotide sequence of SEQ ID NOS: 1 or 10, or a complement thereof; with the proviso that said nucleotide sequence is not comprised by the nucleotide sequence set forth in SEQ ID NO: 20; or between 5500 and 6600 contiguous nucleotides of the nucleotide sequence of SEQ ID NOS: 1 or 10, or a complement thereof.

7. The isolated nucleic acid molecule of claim 6 comprising a nucleotide sequence of between 4 and 4900 contiguous nucleotides of the nucleotide sequence of SEQ ID NOS: 1 or 10, or a complement thereof; with the proviso that said nucleotide sequence is not comprised by the nucleotide sequence set forth in SEQ ID NO: 20; or between 5500 and 6600 contiguous nucleotides of the nucleotide sequence of SEQ ID NOS: 1 or 10, or a complement thereof.

8. The isolated nucleic acid molecule of claim 7 comprising a nucleotide sequence that encodes the amino acid sequence of SEQ ID NO: 2-9, 59, or SEQ ID NO: 11-19 or a complement thereof.

9. An isolated RNA or DNA nucleic acid molecule which hybridizes under stringent conditions to a nucleic acid molecule having the nucleotide sequence of SEQ ID NOS: 1 or 10 or a complement thereof.

10. An isolated polypeptide encoded by the nucleic acid molecule of claim 7.

11. The polypeptide of claim 10 comprising the amino acid of:

- a) an amino acid sequence set forth in any of SEQ ID NOS: 2-19, or 59; or
- b) an amino acid sequence that has 70%-99% homology to the amino acid sequence of (a).

12. The polypeptide of claim 10 wherein the amino acid sequence has

- 5 to 250 contiguous amino acid residues of the amino acid sequence of SEQ ID NOS: 5 or 18 (VP24);
- 5 to 280 contiguous residues of the amino acid sequence of SEQ ID NOS: 6 or 17 (VP30);
- 5 to 320 contiguous residues of the amino acid sequence of SEQ ID NOS: 8 or 13 (VP40);
- 5 to 340 contiguous residues of the amino acid sequence of SEQ ID NOS: 7 or 12 (VP35);
- 5 to 370 contiguous residues of the amino acid sequence of SEQ ID NOS: 4 or 15 (SGP);
- 5 to 370 contiguous residues of the amino acid sequence of SEQ ID NOS: 59 or 16 (SSGP);
- 5 to 670 contiguous residues of the amino acid sequence of SEQ ID NOS: 9 or 14 (GP);
- 5 to 730 contiguous residues of the amino acid sequence of SEQ ID NOS: 3 or 11 (NP); or
- 5 to 2200 contiguous residues of the amino acid sequence of SEQ ID NOS: 2 or 19 (L).

13. (canceled)

14. (canceled)

15. (canceled)

16. (canceled)

17. (canceled)

18. (canceled)

19. (canceled)

20. The hEbola virus of claims 3 or 4, or a protein extract therefrom, and a pharmaceutically acceptable carrier.

21. (canceled)

22. The nucleic acid molecule of claims 6 or 9, and a pharmaceutically acceptable carrier.

23. (canceled)

24. (canceled)

25. (canceled)

26. (canceled)

27. (canceled)

28. (canceled)

29. (canceled)

30. (canceled)

* * * * *