```
                    5        10        15        20        25        30        35        40        45        50        55        60        65        70        75        80
conservation        ********** ********************* ********************* ********************* ********** **********
SPP1˙AM             ATGAAGGCCGCCATTGTTTTCGTGCTCCTCTTTGCCATAGTCTACTGTCGTCCTCTGAAGCGTTCAGTGAGCAGCTCAGA  80
SPP1˙CI             ATGAAGACTCTTATCGTTTTAACGCTCCTCGTTGCCACAGTCTTCTGCCTACCTGTCAAGCGCTCAGCTAGCAGTTCAGA  80
SPP1˙DR             ATGAAATCTATTATTGTTTTAACACTCCTCGTCGCCACAGTCTTCTGTGTACCTGTGAAGCGCTCAGCAAGCAGTTCAGA  80
SPP1˙IP             ATGAAGGCTGTCATTGTTTTCGTGCTCCTTTTCGCAGCGGTCTATTGTCACCCTGTGAAACGTTCAGCCAGCAGCTCAGA  80
SPP1˙LW             ATGAAGACCGTTATCGTTTTAACCCTCCTCATCGCCACAGTCTTCTGCCTGCCTGTCAAGCGCTCAGCCAGCAGTTCAGA  80
SPP1˙PN             ATGAAGGCCGTCATGATTTTCGGGCTCCTCTTCTGCTGTAGTCTACTGCCGTCCTGTAAAACGCTCAGCAAGCAGCTCAGA  80
SPP1˙PP             ATGAAGACTGTTATCGTTTTAACGCTTCTCATCGCCACAGTCTTCTGCCTACCTGTCAAGCGCTCAGCCAGCAGTTCAGA  80
SPP1-C1˙CC          ATGAAGACTGTCATCGTTTTAACGCTCCTCATCGCCACAGTTTTCTGTCTACCTGTGAAGCGCTCAGCTAGCAGTTCAGA  80
SPP1-C2˙CC          ATGAAGACTGTTTTCGTTTTAACGCTCCTCATCGCCACAGTCTTCTGTCTACCTGTGAAGCGCTCAGCTAGCAGTTCAGA  80
SPP1-C1˙SG          ATGAAGACTGTCATCGTTTTAACGCTCCTCATCGCCACAGTTTTCTGTCTACCTGTGAAGCGCTCAGCTAGCAGCTCAGA  80
SPP1-C2˙SG          ATGAAGACTGTTATCGTTTTAACGCTCCTCATCGCCACAGTCTTCTGTCTGCCTGTGAAGCGCTCAGCTAGCAGTTCAGA  80
SPP1-C1˙SR          ATGAAGACTGTCATCGTTTTAACGCTCCTCATCGCCACAGTTTTCTGTCTACCTGTGAAGCGCTCAGCTAGCAGTTCAGA  80
SPP1-C2˙SR          ................................................................................  0
SPP1-C1˙SA          ATGAAGACTGTCATCGTTTTAACGCTCCTCATCGCCACAGTTTTCTGTCTACCTGTGAAGCGCTCAGCTAGCAGTTCAGA  80
SPP1-C2˙SA          ATGAAGACTGTTATCGTTTTAACACTCCTCATCGCCACAGTCTTCTGTCTGCCTGTGAAGCGCTCGGCTAGCAGTTCAGA  80

                    85        90        95        100       105       110       115       120       125       130       135       140       145       150       155
conservation        ********** ** * ** ***** ********** * ********** ********** ******* ********* ****
SPP1˙AM             GAGTTCTGAGGAACAAGCCGCTGCCAAACCGGCGCCCCCAATTCTGCGGAAAGCCCCTGCAAAAGCAGTGCAGGCA....  156
SPP1˙CI             GAGCTCAGAAGAGCTTGTTGCTGTTCAACGGCCACCTCCTATTCTCCGAAAAGCTGCTGCTAATATGTTTGAGGCAGAAC  160
SPP1˙DR             GAGCTCAGAAGAGCTTGTAATTGTTCAACGGCCACCTCCTATTCTTCGAAAAGCTGCTGCCAATGTGTTCCAGGTAGAGC  160
SPP1˙IP             GAGTGCTGACCTGCAGCCTGTTGATACTCAGAGC......ATCAATGGAAAAAGAATCACAGTAGCCATATTCACAAAGC  154
SPP1˙LW             GAGCTCAGAAGAAATGATTGCTGTTCAACGGCCACCTCCTCTTCTACAAAAAGCTGCTGCTGATATGTTCCAGGCAGAAT  160
SPP1˙PN             GAGCTCTGAGGAACAGGCAGTTGCCAAACTGCCACTCCCAGTTCTTGATAAAGGCCCTGCAGATCCACTAAAGGCAGCAC  160
SPP1˙PP             GAGCTCAGAAGAAATGATTGCCGTTCAACGGCCACCTCCTATTCTA...AAAGCTGCTGCTGATATGTTCCAGGCAGAAC  157
SPP1-C1˙CC          GAGCTCAGAAGAACTTGTGGTTGCTCAACAACCAACTCCTATTCTACGAAAAGCTGCTGCTGTTATGTTCAAGGCAGAAC  160
SPP1-C2˙CC          GAGCTCAGAAGAACTTGTGACTGTTCAACAACCACCTCCTATTCTAGAAAAAGCTGCTGCTGTTATGTACAAGGCAGAAC  160
SPP1-C1˙SG          GAGCTCAGAAGAACTTGTGGTTGCTCAACAACCACCTCCTATTCTACGAAAAGCTGCTGCTGTTATGTTCAAGGCAGAAC  160
SPP1-C2˙SG          GAGCTCAGAAGAACTTGTGGTTGTTCAACGGCCACCTCCTATTCTGCAAAAAGCTGCTGCTGTTATGTTCAAGGCAGAAC  160
SPP1-C1˙SR          GAGCTCAGAAGAACTTGTGGTTGCTCAACGGCCACCTCCTATTCTACGAAAAGCTGCTGCTGTTATGTTCAAGGCAGAAC  160
SPP1-C2˙SR          ................................................................................  0
SPP1-C1˙SA          GAGCTCAGAAGAACTTGTGGTTGCTCAACGGCCACCTCCTATTCTACGAAAAGCTGCTGCTGTTATGTTCAAGGCAGAAC  160
SPP1-C2˙SA          GAGCTCAGAAGAACTTGTGGTTGTTCAACGGCCACCTCCTATTCTGCAAAAAGCTGCTGCT...ATGTTCAAGGCAGAAC  157

                    160       165       170       175       180       185       190       195       200       205       210       215       220       225       230
conservation        ******!****** ***** *!********** *!******!******* ****** ********** ********
SPP1˙AM             ..GTAGAGGTTGTACCTGCTCAGGCTGTTAAAGCAGCTCCAGAAGCTCCAGCAGATCCAGAAGCAGCAGCAGCGGCAGCA  234
SPP1˙CI             CTACACAGACCACGCCAACAGAATCTAATGAAAGCACA...GACAGTGCAGATGACACTGAGGAGGCATATGAGGAATCT  237
SPP1˙DR             CTACACAGACCACGCCAACAGAATCGAATGAAAGCACA...GACAGCGCAGATGACACTGAGGAGGCAGATGAGGAATCT  237
SPP1˙IP             CTACCTAATTTCAAGTGTTTTATAATTACTACTCGGCA...CATACTGTATTGAAAGTCACTGCGGTTGCTGATGTTAGC  231
SPP1˙LW             CTACACAGACCACACCAGCAGATTCCAATGAAAGCACA...GACAGTGAAGATGACACTGAGGAGGCAGATGAGAAATCT  237
SPP1˙PN             CTGCACAGGACAAAAACACAGAATCAGACACAGAGGGA...GACAGTGCAGACAGCGCAGACACGCAAGACACGCAAGAC  237
SPP1˙PP             CTACACAGACCACACCAGCAGATTCCAATGAAAGCACA...GACAGTGAAGATGACACTGAGGAGGAAGATGAGGAATCT  234
SPP1-C1˙CC          CTACACAGTCCACGCCAGGAGAATCCAATGAAAGCACA...GACAGTGCAGATGACAGTGAGGGTGCAGATGAGGAATCT  237
SPP1-C2˙CC          CCACACAGACCACACCAGTAGATTCCAATGAAAGCACA...GACAGTGCAGATGACAGTGAGGATGCAGATGAGGAATCT  237
SPP1-C1˙SG          CTACACAGACCACGCCAGCTGAATCCAATGAAAGCACA...GACAGTGCAGATGACAGTGAGGATGCAGATGAGGAATCT  237
SPP1-C2˙SG          CTACACAGACCACGCCAGCTGAATCCAATGAAAGCACA...GACAGTGCAGATGACAGTGAGGATGCAGATGAGGAATCT  237
SPP1-C1˙SR          CTACACAGACCACGCCAGCTGAATCCAATGAAAGCACA...GAAAGTGCAGATGACAGTGAGGATGCAGATGAGGAATCT  237
SPP1-C2˙SR          .....CAGACCACGCCAGCTGAATCCAATGAAAGCACA...GACAGTTCAGATGACAGTGAGGATGCAGATGAGGAATCT  72
SPP1-C1˙SA          CTACACAGACCACGCCAGCTGAATCCAATGAAAGCACA...GACAGTGCAGATGACAGTGAGGATGCAGATGAGGAATCT  237
SPP1-C2˙SA          CTACACAGACCACGCCAGTAGAATCCAATGAAAGCACA...GACAGTGCAGATGACAGTGAGGATGCAGATGAGGAATCT  234
```

```
                    235       240       245       250       255       260       265       270       275       280       285       290       295       300       305       310
conservation        *****!******* *!* * ****! ** ***!!***!**!!!***!  *!*****!!*!!* !** ** ******* **
SPP1˙AM             TCATCGGATGAGGACGACTCAGATGAAGCTGAGGACTCCAGTGAGAGTGCTGAGTCTGGAGAAACTGAGGCCACCACCAC  314
SPP1˙CI             GAAACAGATGAGAACAAAGAGGAGAACGACACTGATTCCAGTGAAAGT...GAATCTGGAGAAACTGAGACCACCGCAGT  314
SPP1˙DR             GAAACAGATGAGAAGGAAGAGGAGAATGAGACTGACTCCAGTGAGAGT...GAATCTGGAGAATCTGATACCACCATCAT  314
SPP1˙IP             TCTGAAGAGGATGATGATGATGATGATGACAGTGACAGCGATGAGTCTGATGAGAATGGCGAGCCCACCAGTTCT.....  306
SPP1˙LW             GAAACAGATGAGGATGAG...GAGAATGAGACTGACTCCAGTGAGAGT...AATTCTGGAGAAACTCAGAGCACCGTAGT  311
SPP1˙PN             ACTATAGATTTGTGCCAGGATTGGGACACAGATGACTCCAGTGAGAGTGCTGAATCAGGAGAGACTGAGGCCACTACCGT  317
SPP1˙PP             AAAACAGATGAGGATGAG...GAGAACGAGACTGACTCCAGTGAGAGT...AATTCTGGGGAAACTCAGAGCACCGTCGT  308
SPP1-C1˙CC          GAAATGGATGAGAACGAAAAGGAGAACGATACTGACTCCAGTGAGAGT...GAATCTGGAGAACCTGCGACCACCGTTGT  314
SPP1-C2˙CC          GAAACAGATGAGAAGGAAGATGAGAATGATACTGATACCAGTGAGAGT...GAATCTGGAGAAACTGCGATCACCGTTGT  314
SPP1-C1˙SG          AAAACAGTTGAGAACGAAAAGGAGAACGACACTGACTCCAGTGAGAGT...GAATCTGGAGAATCTGCGACCATCGTTGT  314
SPP1-C2˙SG          GAAACAGATGAGAGGGAAGATGAGAATGATACTGATTCCAGTGAGAGT...GAATCTGGAGAATCTGCGATCACCGTTGT  314
SPP1-C1˙SR          GAAACAGTTGAGAACGAAAAGGAGAACGATACTGACTCCAGTGAGAGT...GAATCTGGAGAATCTGCGACCGCCGTTGT  314
SPP1-C2˙SR          GAAACAGATGAGAGGGAAGATGAGAATGATACTGATACCAGTGAGAGT...GAATCTGGAGAATCTGCGATCACCGTTGT  149
SPP1-C1˙SA          AAAACAGCTGAGAATGAAAAGGAGAACGATACTGACTCCAGTGAGAGT...GAATCTGGAGAATCTGCGACCACCGTTGT  314
SPP1-C2˙SA          GAAACAGATGAGAGGGAAGATGAGAATGATACTGATTCCAGTGAGAGT...GAATCTGGAGAATCTGCGATCACCGTTGT  311

                    315       320       325       330       335       340       345       350       355       360       365       370       375       380       385       390
conservation        *   !! ***!**** !! *!!*! !!****!!**!*!!**!****!!*!!*!!!!!!!!***!*!*!!**!**!!*!!!!
SPP1˙AM             CCCACCTTTCACCGAGGAGCCCACCCTGGGCCCTATTATTAACGACGGCCGCGGAGACAACATGGGCTACCCCAGCGACT  394
SPP1˙CI             C...CCATCCACAGTAGAGCCCTCACTGGACCCCATCATCAACACAGGCCGGGGAGACAGCATGGGCTACCCTAGCGACT  391
SPP1˙DR             C...CCAGTCACAGTCGATCCCACGCTGGGTCCCATTATCAACACAGGCCGGGGAGACAGTTTGGGCTACCCTAGCGACT  391
SPP1˙IP             ....CCTCCTATCATAGAGACCACTCTGATCCCTGTTATTAATT...GGCCGTGGAGACAGCATTGGCTTCCAGAGTGACT  379
SPP1˙LW             C...CCATCCACAGTCGAGCCCTCGCTGGACCCCATCATCAACACAGGCCGGGGAGACAGCATGGGCTACCCCAGCGACT  388
SPP1˙PN             CCCACCTGTAACCGTGGAGCCCACCCTGGACCCTATTATTGACAACGGTCGTGGAGACAGCATGGGCTACCCCAGCGACT  397
SPP1˙PP             C...CCATCCACAGTCGAGCCCTCGCTGGACCCCATTATCCACACAGGCCGGGGAGACAACATGGGCTACCCAAGCGACT  385
SPP1-C1˙CC          C...CCATTCACAGTGGAACCCCCACTGGACCCCATCATCAACACAGGCCGGGGAGACAACATGGACTATCCTAGCGACT  391
SPP1-C2˙CC          C...CCCTCCACAGTAGAACCCTCACTAGATCCCATCATCAACACAGGCCGGGGAGACAACATGGGCTACCCTAGCGACT  391
SPP1-C1˙SG          C...CCCTCCACAGTGGAACCCTCACTGGACCCCATCATCAACACAGGCCGGGGAGACAGCATGGGCTATCCTAGCGACT  391
SPP1-C2˙SG          C...CCCTCCACAGTAGAACCCTCACTGGACCCCATCATCAACACAGGCCGGGGAGACAGCATGGGCTACCCTAGCGACT  391
SPP1-C1˙SR          C...CCCTCCACAGTGGAACCCTCACTGGACCCCATCATCAACACTGGCCGGGGAGACAGCATGGGCTATCCTAGCGACT  391
SPP1-C2˙SR          C...CCCTCCACAGTAGAACCCTCACTGGACCCCATCATCAACACAGGCCGGGGAGACAGCATGGGCTACCCTAGCGACT  226
SPP1-C1˙SA          C...CCCTCCACAGTGGAACCCTCACTGGACCCCATCATCAACACAGGCCGGGGAGACAGCATGGGCTATCCTAGCGACT  391
SPP1-C2˙SA          C...CCCTCCACAGTAGAACCCTCACTGGACCCCATCATCAACACAGGCCGGGGAGACAGCATGGGCTACCCTAGCGACT  388
```

```
                                                
```

```
                                                
```