

Supplementary Figure 1: A) Multiple alignment of Cas1 representative sequence set; B) Multiple alignment of Cas2 representative sequence set; C) Multiple alignment of Cas4 representative sequence set; (Please see Supplementary Materials and Methods for details); Sequences are denoted by Genbank identifiers and complete organism name.

A

407462288_Candidatus_Nitrosopumilus_koreensis_AR1
 2122551_Methanosaarcina_mazei_GoI
 147921327_Methanococcus_arvoryzae_MR50
 124485266_Methanocorpusculum_labreanum_Z
 307352568_Methanoplanus_petrolearius_DSM_11571
 15415254_Methanocoregula_boonei_6A8
 219852754_Methanospaerula_palustris_El_9c
 126178701_Methanoculleus_marisnigri_JR1
 397780289_Methanoculleus_bourguense_MS2
 156937861_Lignicoccus_hospitalis_KIN4-I
 70607609_Sulfobolbus_acidocaldarius_DSM_639
 187736488_Akkermansia_muciniphila_ATCC_BAA-835
 436839745_Desulfovibrio_hydrotermalis_AM13_-DSM_14728
 71894593_Mycoplasma_synoviae_53
 193216853_Mycoplasma_arthritis_158L3-1
 187250661_Ellusimicrobium_minutum_Pein19
 325972002_Sphaerochaeta_globus_str_Buddy
 47458867_Mycoplasma_mobile_163K
 269123285_Streptobacillus_moniliformis_DSM_12112
 57865888_Staphylococcus_epidermidis_RP62A
 319957207_Nitratifractor_salsuginis_DSM_16511
 310286727_Bifidobacterium_bifidum_S17
 90961086_Lactobacillus_salivarius_UCC118
 406600270_Leuconostoc_gelidium_JB7
 386318631_Staphylococcus_pseudointermedius_ED99
 302336019_Olsenella_oli_DSM_7084
 15675042_Streptococcus_pyogenes_M1_GAS
 352684362_Acidaminococcus_intestini_RYC-MR95
 42525844_Treponema_denticola_ATCC_35405
 169823756_Finegoldia_magana_ATCC_29328
 117929157_Acidothermus_cellulolyticus_11B
 28344312_Bifidobacterium_bifidum_Bd1
 38232679_Corynebacterium_diphtheriae_NCTC_13129
 238924076_Eubacterium_rectale_ATCC_33656
 116627543_Streptococcus_thermophilus_LMD-9
 57238545_Campylobacter_jejuni_RM1221
 291276264_Helicobacter_mustelae_12198
 294086112_Candidatus_Punicispirillum_marinum_IMCC1322
 189485226 uncultured.Termite_group_1_bacterium.phylotype_Rs-D17
 222109284_Acidovorax_ebreus_TPSY
 15602991_Pasteurella_multocida_subsp._multocida_str_-Pm70
 310780383_Ilyobacter_polytropus_DSM_2926
 220936119_Clostridium_lavamentophilum_H10
 34557789_Wolinella_succinogenes_DSM_1740
 90425960_Gludoseudomonas_palustre_BisB18
 209542526_Gluconacetobacter_diazotrophicus_Pal_5
 121608212_Vermiphobacter_eiseniae_EF01-2
 60683388_Bacteroides_fragilis_NCTC_9343
 154250554_Parvibaculum_lavamentivorans_DS-1
 283778924_Pirellula_staleyi_DSM_6068
 154174048_Campylobacter_curvus_525-92
 292492938_Nitrosoalkaliphilus_nc4
 111224947_Frankia_alni_ACN14a
 385807970_Ignavibacterium_album_JCM_16511
 150021526_Thermosiphon_melanesiensis_B1429
 28211139_Clostridium_tetani_B88
 48470705_Picrophilus_torridus_DSM_9790
 46164818_Nanoarchaeum_equitans_Kin4-M
 55376276_Haloarcula_marismortui_ATCC_43049
 389844445_Mesotoga_prime_MesG1-Ag_4-2
 309062116_Thermococcus_sp_-Cl1
 13540937_Thermoplasmavolcanium_GSS1
 289596694_Aciduliprofundum_boonei_T469
 14590110_Pyrococcus_horikoshii_OT3
 84489236_Methanospaerula_stadtmanae_DSM_3091
 268317185_Rhodothermus_marinus_DSM_4252
 18977490_Pyrococcus_furiosus_DSM_3638
 427708213_Nostoc_sp_-PCC_7107
 119720226_Thermofilum_pendens_Hrk_5
 118497969_Fanciella_novicina_U112
 332616191_Haliscoenobacter_hydroassis_DSM_1100
 326793969_Marinomonas_mediterranea_MBb-1
 220934970_Thioalkalivibrio_sulfidophilus_HL-EBr7
 288947693_Allochromatium_vinosum_DSM_180
 431929780_Thioflavibacter_mobilis_8321
 206890539_Thermodesulfovibrio_yellowstonii_DSM_11347
 52425690_Mannheimia_succiniciproducens_MBEL55E
 30248142_Nitrosomonas_europaea_ATCC_19718
 121997419_Halorhodospira_halophila_SL1
 430760780_Thioalkalivibrio_nitratireducens_DSM_14787

-----NQKNHYNVKFLKGY-GHSISVFKDSKII---L---KSNHDPFSSPP-----EQE-EWFV-----KNMPYEKIVLSGK-GYVSTEALSSLSENNRNVILL--DNH-G-KP-----V
 -----MLKLILAH-GINHMVGDGAKL---I---K-DGRFSTE-----EPQ-EYFP-----SPKRIDIGIIYKLNTEALRIPHLIKHNHQVSIL---DWN-G-KL-----L
 -----MNPLLIQGF-GTRISVEKRLK---I---S-T---ET-----NDY-EYFP-----HQIDHDTIDVITGPTGNISFAEMRIMCHKINLTLL---DWN-G-NL-----L
 -----MIPWVPAVVG-Y-GAHIKSTKRL---I---R-Q---KG-----STT-Q-Y-----PLDDMHMRILIAOG---HSLHTSVRERLADRGIAVSPF---TAH-G-KP-----V
 -----MPSDPIWPVHWGFG-GAHIKSTKRL---I---L-N---KS-----DVE-E-I-----PIKSVDHVLLVGG-HNIHNSTTSFLHNRNGSYISPF---EDP-G-SP-----L
 -----MTAIPWVPAVVG-Y-GAHIKSTKRL---I---L-A---KG-----KVE-E-Y-----PLEIIONLIVVGG-HNIHNSTTSFLHNRNGSYISPF---EDP-G-SP-----L
 -----RSCSKLPWLWVSGY-GAHIKSTPLLI---V---Q-K---NG-----TTT-E-Y-----PIGDVHLLVGG-HTLHTSAVLQHMQAGNWVSPF---AAD-G-TP-----V
 -----OMAAETPLPVFGY-GGHIKATTRLI---I---A-R---GS-----DTR-R-Y-----PIQAVKHLLVGG-HTLHTSAVNLKAGAAITIF---DID-G-TP-----V
 -----MTATOPWLPVFGY-GGHIKATTRLI---I---A-H---GS-----TTR-R-Y-----PLQAVKHLLVGG-HTLHTSAVNLKAGAVITTF---DID-G-TP-----V
 -----MILVURGH-AFVGRKGYMIT---V---RYKK---DG-----KEV-TEAF-----PALDIEMAVFVGKGITVSTAALQILLEEQNPVTLPH---GVD-W-S-----F
 -----MSLVIKNA-TIMRKGSIDLQ---M---TWS---VG-----KNL-T-V-----SSLDELVUVVGNVRITSEVELFLSSLNIPVLVH---GKR-N-DV-----
 -----MSYHILSDAY-TCHLSCDKGQLR---C---A-D---GE-N-----SPR-T-I-----PLEDVGAVVLLSSPKATLTSNLIELARKRIGFVLC---ESY---RP-----A
 -----AQLKRITVVCNHH-FASLGVDIICLCA---V---K-K---NG-T-----AIA-R-I-----PFADTEIVVYVGG-SSVSTKLCKSYEKIPVTC---TPS-G-YY-----I
 -----MMARKVVEIKEY-ELYFLFHNLSLI---V---KKE---NN-----KIV---L-----PLSDIDTILSNFNYCTISVPLINIAVSNNNIIIC---NND-F-EP-----N
 -----MEKRVVIEISES-NNTYLLFLNNLV---V---N-K---DG-T-----KIT---I-----PTSQIDTFLFENDKLTLTFLDVLHDHGGINIIVC---GRN-H-MP-----K
 -----MWRLVPLPGD-GYHLCVNNNFS-AV---V---K-D---RE-----EKL-H-C-----LFDDINSIILYFLNNTISCTIQCKCLEHKVVPVFC---DKT-Y-NP-----A
 -----MWRLVPLGDT-GYHLCVNNNFS-AV---V---E-K---DG-E-----ESL---I-----FFDDIHDISIICH-GYHLCVNNFDSKCLHHKVVPVFC---DEK-H-IP-----L
 -----MGWKVLEINTD-EVHYLHVNLNY---I---K---KG-----RN---E-----EKI-L-L-----NIRDIDTILFLNPNQYTSIRLITFLAKNNNVIIPM---NEK-N-EP-----N
 -----MSFRQVLITQK-SYIHLENDNL---I---V---E-----KD-D-----KXL-S-V-----PISDISIIVFESLESYVSLRISELSLRNMTMIPC---DYR-H-MP-----
 -----MKDVYVENVH-YPTVAKTENKIFRN---I---D---KS-----EKF---Y-----LFEEIAIFIIDVHYSKSFHVLVCIENDIAIIPC---DKX-H-SP-----V
 -----MAAWKTRITRKP-CRLSINGNLV---V---E-----DE-K-----VRF-K-L-----SLSLTDISIIFEGDRFTLSAKVSLRSHKVKATLPC---DEY-Y-MP-----T
 -----MAWRNVVTRTH-CKISTKMLRL---I---Q-T---DD-----DVC-Q-L-----PMDDIGLMILMIETTRAVITCAIDDLRDKVIVPC---DDK-H-LP-----I
 -----MGRFRNVIITQH-SKLYSKNNNAM---V---Q-NK---DG-----INQ---I-----PLVDMIDLILSTQTQAVITALSVSITAELIKEAKVVIIPC---DNK-N-EP-----V
 -----MGRWTVUTKHK-SKLYSKNNH---V---O-T---AE-----KYQ-O-I-----PMEDIQILILITQAVITAVIAVETIKRNIK1IIFS---DIN-G-QP-----I
 -----MSFRVIIITKE-SKLSLRMNNQLI---V---K-S---E-----NLS-R-I-----PLNEILCLVIEHPNVMSMGTGHLMNALSDQKIVTILC---NTK-H-LP-----N
 -----MGRFRNVIILESP-CKCTYVFLNVL---V---R-K---ED-----EIV-K-V-----HLSLEISVSVLQSTRAFISAYLSELAKAK1SVPVVS---DEK-C-NP-----I
 -----MAGWRTVVNNTH-SKLYSKNNH---I---P---D---AY-----KTE-L-I-----HLSLEDILLETDLTIVMLVLRDENVILVIFC---DDK-R-LP-----T
 -----MS-RTVITVITQR-CKLDFCMVNVE---V---Q-T---AV-----SKR-R-I-----FIDEIKLTIETVNTAYTALVYVTSFVTSVLAELKIAEKVVIIPC---GKD-H-NT-----Q
 -----MSWRTVVIISNR-AKLDDHLNLHV---V---R-G-----KTQ-K-V-----FIEELSVIETIETAVITSIAALLNELIKQKVVKIFC---DEK-R-NP-----A
 -----MGWRTVVLISQR-AKLDDHLNLHV---I---R---KE-----EIV-K-I-----HPIEAVALVDTLMSVTMSFVTSVLAELKIAEKVVIIPC---GKD-H-NT-----I
 -----MTGPWRVLLSEL-GEVHVAQGALL---V---GD-----ERV-----PLDVDV-AMM1TGPVYSLHSGVIDRAAFAFGVGVHIC---DWR-G-VP-----V
 -----MSQSRVVIIDCTNL-DGTISYERGRKM---V---R-K---HQ-----ESE-SVCV-----PLAAQAAVVLIGR-VCCSVALLHEMAKAGVSVMLC---DWR-G-IP-----D
 -----MNPGRWVRLBLIDF-GDKVSYRQQLA---I---T---DS-----G-----ELRAT-L-----PLAQAVIILVNL-LLISGAVLVLKSESYDIAVILC---DWR-R-VP-----V
 -----MAPFRVILIENE-FTVTKLHLNLNLI---I---T---KC-G-----EDI-W-I-----PLDDISMIVLNLSNLTRMLCMOLABQGIGLMLY---NQO-H-LP-----T
 -----MTWRVWHSQS-EKMLRKLNLNLI---V---Q-----KM-G-----QEF-T-V-----PLSDISMIVLNLSNLTRMLCMOLABQGIGLMLY---NQO-H-LP-----T
 -----SYDEAFLTILLSSN-AKLNLHLNLHV---I---K-Q---DE-----NIA-K-L-----FLKDINIVLTSQLSVSIISALFARHKHILLITC---DET-H-SI-----N
 -----MPDEAFLRMLPSKP-AKLHKLHNNL---I---I-----HG-K-----EDI-G-I-----PLRDINTIVLTDITPQITLTSMLKELAQYKILVFFC---DDS-H-LP-----S
 -----MIEISQO-PHRISVGRGAI---V---S-D---NE-T-----ELG-Q-----DLDGVLVSIVTSGATVTPMITAEARANIP1IIC---NDR-F-QP-----V
 -----MSWRTVFLFSKPS-CLSLSVNSQNL---I---R-P---QD-S-----TVH-D-I-----PIEDISVLSNINTUSLSECSQNSVIVFSC---DSS-H-IP-----C
 -----MVWRSIVINRP-AKLKRHEPALV---V---E-----Q-----QSA-R-V-----PFEDIAVILVNRHEITLTHVLSLACGEGYIGGLYST---GDN-H-QP-----N
 -----GVIMSWRSILISQG-GKLSLQJNQML---I---Q---QS-N-----PNS-T-V-----PLEDIAVISETKELITAPLISVLAQAGYVGTLLTC---DEQ-F-LP-----C
 -----MGWRNPIFTS-TKISTRDGSLL---I---T-----KE-G-----GEV-N-I-----PLEDIISIIILNNNTSITVSLVLEIERSDIVLTS---DEK-H-LP-----N
 -----MAAWRVVIATTS-CHLRVRDRLQVL---I---E-S-----RE-G-----ERT-Q-L-----PIDDINTVLSVTSITVTSALLSALAEIDLALYSC---DGK-H-TP-----N
 -----MAWRLHILQTS-ARLSDRQDVC---V---R-Q---DA-E-----EVRAV---I-----PLADIGVIVENPQVTLISALLSALALQEGTALMSC---DSS-H-LP-----D
 -----MAWGRVHISHP-SRLTHRNQVL---V---A---Q---DG-----EVS-L-----ALEDIAV1DITQPTQALTSALMSAMDAGVILIFT---DER-H-TP-----S
 -----MIGRTYVITAD-RRHLFVNRGMV---IKDTEG---E-RK-----ELG-Q-V-----PLDDIAVIAHAGLHTYTNNLVLAERGAPVLC---GPN-H-NA-----V
 -----MIKTLYTFGPN-GRILSRLNAQV---I---K-L---PD-----EKA-T-V-----VLPETLKLQQAETVPIKIEDIVVLLNQKQTITGVSLEALENNCSVITC---DSR-S-MP-----V
 -----MIRKTVEFATP-GTRLSVANKQVL---I---E-R---PD-L-----PKA-T-L-----PIEDLVEVFLDLRATYQVAFELLEAGATVMT---GRD-H-LP-----A
 -----PGLTLAGPOIEELVHGBVQI---P---K-H---QD-----QR-----GIE-S-I-----PLHTLREVVVLGA-VSLSHVLVSLAQONAISSLLL---DES---AN-----R
 -----MOKSDRTHFLPSL-GRLRDRDNYY---P-----DKFDETGG---VTA-SKIL-----PINADEIYLITLRLVDTLTDPLADNNNLLHVF---SPF-Q-SP-----R
 -----METLFWGSR-EARLRRENTLIS---I---T-I-----EG-----KTR-A-F-----METLFWGSR-EARLRRENTLIS---I-----PLTGHVLLAEB-STLNLSKLTLCGKHGVRLSVF---DYY-G-YF-----K
 -----MPAAGRITYNLTIP---CRIREDNSLR---I---ERA---DS-----R-----PLTGHVLLAEB-STLNLSKLTLCGKHGVRLSVF---DYY-G-YF-----K
 -----PQGKMMNLYIIFSN-SILSRKDDSLV---L-----DSIPQTIDAIHETDFTYETEGI-L---PAKPTDGSQNPRIHAPAEVSBAYTITFGE-VRFNFTQFFKCVGFYIIPVHF---NYY-G-NI-----V
 -----METLYITFIDT---GRLRQSLSVL---I---K-P---KD-----NMX-PLPV-----PLKSVSSLIMFSE-VSMRMKFLDLSKLEGEPAFFY---GY-E-NF-----I
 -----MGSTRYITVS-GELKLRKNDSLC---P---R-K---NN-----KVN-Y-I-----PVENTKEIICMSE-VINNSKLSLDFLSQNNIIMHF---NYY-E-GY-----S
 -----MQTYVIIIS-SGTLNRMEDSLR---P---S-N-----ENK-I-I-----PLENIESIIISGN-VSITKPAISLSKNNPVFFM---SMY-D-NY-----I
 -----MKTIVYIISL-GKLYRKNGNLYF---I---N-K---DK-----KKS-P-I-----PLESIEKIILFNK-VSDTINYALKLADRNIP1PHFVENTKKGISYV---L
 -----MNDNYHVFTD-GTLLRHDNTIRL---V---T-E-----DQ-----EKK-Y-L-----PIENAELFLHQLQ-IDYNTRLVLSVFLGVVAHF---GWN-D-YY-----A
 -----MKKNFTYLFAS-QOLSRKDNTL---I---E-R---EG-----RRK-F-I-----PIEKEIYLITLRLVDTLTDPLADNNNLLHVF---SHN-G-WY-----S
 -----MKYPLFIDTH-GRLRQSLSVL---P-----VG-----RLE-KRTI-----PLAQVSCHVRLCAR-VSLSGAVELLSERRIPVPHY---STR-G-DY-----R
 -----MYNPHYITQD-GTITREGNTLFY---C---G-P---DF-----KRH---L-----PVMVNEOIIISAK-SLSWSLADYLSKLQIVVHII---STN-G-KY-----M
 -----MDSLVIITKE-AIIKREANTLY---I---L---V---R---KG-----EKR-S-I-----PIHNLDRITCAMP-VSFSSGAIKHLVLSNGVVFHF---DMY-G-NY-----E
 -----MKSDIYITQ-P-GILERKANTLFP---V---N-E---EM-----KRA-L-----PINTSEIYCFAD-PTLTSVGLAKLSDNDPVPHF---NKKY-G-YY-----R
 -----GCDTIEKPYIISN-GILYRKENTLKP---K-N---K---VS-----DTS---I-----PIHAINEINCYCG-VSLSRGAISL1QKEK1IINNF---NKKY-G-YY-----E
 -----MKRKYIFFSS-GRLRQRNQTLFF---EKAAGE-RIPDDQDETGVPSGPTGPTGEST-P-F-----PVEQVESLYFFGE-VDLNSKLTTFLARHDIPAHFY---DYY-G-NY-----T
 -----MRKSLTISD-GTLLRHDNTLF---E-N---VN-G-----RKP-L-----AIEGTYDYYIYGH-VNITSQALHLYIAQKGILLHF---NHY-G-YY-----D
 -----MSTIYITQD-DTVPQIHOYLK---V---F-Y---QQ-T-----QRI-C-I-----PIRNLQSFIPFQVNL-IPKEV1QIVRSNHIPLVYL---TQT-G-EY-----L
 -----MRLVLRGVDEVTVSRSRSTV---I---K---SG-N-----RVE-E-R-----ALRDVDAVLVVGSGKIKISSSLPFLVLAHGIPLSSL---AK---GG-----V
 -----NDIESKNIVFVNIFDVGKVLSSLSDLIV---I---K-D---KE-T-----SVHKVLAFLVFGN-MTMTSQSLLETCCKNAIQLVFM---KNS-F-RP-----Y
 -----MEIYIYIIS-QNLYDLYHECTIL---V---V-N---GE-----KRT-P-V-----AFKQKIERIWIIPS-TSIAQALLVEQVGEVLFIC---SGE-G-SP-----L
 -----EREKLGLTLLITIAQD-AMLSKSEKORL---V---E-Q---YD-----ELH-T-Y-----PWTATSLVILVGP-HHITTPALKSAMHPNVPHF---SQQ-G-RY-----Q
 -----QAAAEQTLILAGE-PARVGLDGRRL---V---Q---RG-----EAE-LLSA-----PLETLAGVFLVGP-HQSTVQTLALGDRGIPLALALA---TQG-G-RL-----R
 -----MNAQQLLVIDHRTQDQLTLDGGLARL---V---T---PG-T-----KPR-H-I-----PLGVGLVGVHGR-ALVGCDCWVRALARIGPAVMQ---PGR-G-----R
 -----MILVUDVRD-SGVLRHDSGVL---I---E-R---EG-G-----ETR-R-A-----PINOLELVLVVG---PLAETAVNRLGRHAAQGPTVFL---PVR-G-RD-----G
 -----MSTVFLDRK-DIEIRVDGNSL---P---YAK---GK-----KDG-S-L-----PLSLPKRUVVLYGNK-VKITSVLYKLVNHGIFTVL---TGT-L-KY-----S
 -----MPTLVIDR-TTELKVNGDVL---C---Y-E---KG---R-----RIA-T-I-----PLASVDRYLMKGD-INLQISLSSKLGKIGVGVFL---QCR-KNKP-----
 -----MTSLFVDR-GVELGLESQAVI---P---R-E---NG-E-----RIG-T-V-----PIAPLTVFLRVLGD-NVLPALAGLKLGERGVGVVIL---SRTS-RP-----
 -----MGTLYIDRR-RTRLELAHKALT---I---R-E---PE-A-----QPR-S-V-----PLSLIDRLLVIGO-VELSSGLVLTLaESGVSLVFM---PSR-G-QR-----R
 -----MATLFLDRA-QLELRDIDGNVLA---V---Y-E---GG-R-----RRG-T-V-----PLSLLERVVRVQGRQTRFESGVLLKLAESGTAFTVFM---GAR-S-SRQ-----V

292492856 *Nitrosococcus halophilus* *Nc4*
 34541577 *Porphyrimonas gingivalis* *W83*
 88602029 *Methanospirillum hungatei* *JF-1*
 32589070 *Nitrosomonas_sp* *AL212*
 269798874 *Veillonella parvula* *DSM_2008*
 332668016 *Halsicomicrobacter hydrossis* *DSM_1100*
 379728404 *Saprosira grandis* *str_Lewin*
 327314605 *Prevotella dentincola* *FW289*
 323295171 *Flexibacter litoralis* *DSM_6794*
 15038972 *Methanococcus vanneili* *SB*
 150401497 *Methanococcus aeolicus* *Nankai-3*
 204804851 *Spirosooma linguale* *DSM_74*
 257125862 *Leptotrichia buccalis* *C-1013-b*
 21226661 *Methanosarcina maezi* *Gol*
 321866577 *Candidatus Cloacamonas acidaminovorans* *str_Evry*
 319790504 *Thermovibrio ammonificans* *HB-1*
 327388662 *Thermosulfobacterium geofontis* *OPP15*
 20094748 *Methanopyrus kandleri* *AV19*
 386750121 *Helicobacter ceterorum* *MIT_00-7128*
 126465226 *Staphylothermus marinus* *F1*
 170290046 *Candidatus Korarchaeum cryptofilum* *OPF8*
 385805115 *Fervidococcus fontis* *Kam940*
 15898245 *Sulfolobus sulfataricus* *P2*
 156937921 *Ignicoccus hospitalis* *KIN4-I*
 124027534 *Hyperthermus butylicus* *DSM_5456*
 118431381 *Aeropyrum pernix* *K1*
 307595807 *Vulcanisaeta distributa* *DSM_14429*
 305663333 *Ignisphaera aggregans* *DSM_17230*
 347523053 *Pyrobolus fumarii* *1A*
 302347808 *Acidilobus saccharovorans* *345-15*
 21884093 *Desulfurococcus kamchatkensis* *122in*
 389861365 *Thermogladius cellulolyticus* *1633*
 15898286 *Sulfolobus sulfataricus* *P2*
 15922975 *Sulfolobus tokodaii* *str_7*
 18311772 *Pyrococcus aerophilum* *str_IM2*
 18311699 *Pyrococcus aerophilum* *str_IM2*
 4296139 *Legionella pneumophila* *str_Paris*
 34557933 *Francisella novicida* *U112*
 328954440 *Desulfovibacca acetoxidans* *DSM_1109*
 408403000 *Candidatus Nitrosphaera gargensis* *Ga9-2*
 330509017 *Methanosaeta concilii* *GP6*
 148656004 *Roseiflexus* *sp_RS-1*
 386854797 *Denicoccus gobiensis* *I-0*
 167628642 *Helicobacterium modesticaldum* *Icel*
 148642083 *Methanobrevibacter smithii* *ATCC_35061*
 288560430 *Methanobrevibacter ruminantium* *M1*
 1149462 *Archaeoglobus fulgidus* *DSM_4304*
 383319612 *Methanocella conradii* *B2254*
 29467623 *Rhodobacter capsulatus* *SB_1003*
 302343124 *Desulfarculus baarsii* *DSM_2075*
 389875622 *Tistrella mobilis* *KA081020-065*
 374998939 *Azospirillum lipofulferum* *4B*
 28857888 *Azospirillum* *sp_B510*
 116627764 *Streptococcus thermophilus* *LMD-9*
 16059954 *Mycobacterium tuberculosis* *H37Rv*
 348026605 *Megasphaera elsdenii* *DSM_20460*
 296133514 *Thermincula potens* *JR*
 114567264 *Syntrophomonas wolfei* *subsp_wolfei* *str_Goettingen*
 322965452 *Thermodesulfovibium narugense* *DSM_14796*
 218442809 *Cyanothecae* *sp_PCC_7424*
 172289611 *Nostoc* *sp_PCC_7120*
 119357853 *Chlororubrum phaeobacteroides* *DSM_266*
 119357846 *Chlororubrum phaeobacteroides* *DSM_266*
 36116789 *Microcylnatuns phosphovorus* *NM-1*
 37677204 *Vibrio vulnificus* *YU016*
 15425031 *Fervidobacterium nodosum* *Rt17-B1*
 38055760 *Synechocystis* *sp_PCC_6803*
 85392490 *Rhodopirellulum rubrum* *ATCC_11170*
 26141594 *Fibrobacter succinogenes* *subsp_succinogenes* *S85*
 86022266 *Natronomonas pharaonis* *DSM_2160*
 32114617 *Rhodomicrobium vannielii* *ATCC_17100*
 199638006 *Bifidobacterium animalis* *subsp_lactis* *AD011*
 116748794 *Syntrophobacter fumaroxidans* *MPOB*
 37051217 *Desulfuritalea indicum* *S5*
 328949009 *Treponema succinifaciens* *DSM_2489*
 333969612 *Treponema azotonutriens* *ZAS-9*
 136373559 *Calditerrivibrio nitroreducens* *DSM_19672*
 218961167 *Candidatus Cloacamonas acidaminovorans* *str_Evry*
 294828260 *Leptospira interrogans* *serovar Lai* *str_M5601*
 347756303 *Candidatus Chloracidobacterium thermophilum* *B*
 126457462 *Sorangium cellulosum* *So ce56*
 86742032 *Herpetosiphon aurantiacus* *DSM_785*
 433631663 *Mycobacterium canettii* *CIPT_140070010*
 15612904 *Bacillus halodurans* *C-125*
 108763792 *Mycoboccus xanthus* *DG_1622*
 344996859 *Caldicellulosiruptor lactoaceticus* *6A*
 85858452 *Syntrophobacter aciditrophicus* *SB*
 148657121 *Roseiflexus* *sp_RS-1*
 15898907 *Herpetosiphon aurantiacus* *DSM_785*
 127277877 *Nostoc* *sp_PCC_7120*
 268318378 *Rhodothermus marinus* *DSM_4252*
 55978328 *Thermus thermophilus* *HB8*
 297622675 *Truerepa radiovicinalis* *DSM_17093*
 389853000 *Desulfobacca acetoxidans* *DSM_11109*

337286709_Thermodesulfatator_indicus_DSM_15286
35853422_Desulfovobacca_acetoxidans_DSM_11109
38505684_Synechocystis_sp_PCC_6803
134299482_Desulfovotomaculum_reducens_MI-1
17229060_Nostoc_sp_PCC_7120
303931602_Acetoholobium_arabaticum_DSM_5501
159889755_Herpetosiphon_aromaticarius_DSM_785
320161859_Anabaenopsis_thermophila_UNI-1
156741961_Roseiflexus_castenholzii_DSM_13941
383762167_Caldilinea_aerophilica_DSM_14535_-NRBC_104270
46252657_Thermus_thermophilus_HB27
134101617_Saccharopolyspora erythraea_NRRL_2338
38719041_Gordonia_polyisoprenivorans_VH2
320120261_Issacharobacter_pallidus_ATCC_43644
628318441_Rhodothermus_marinus_DSM_4252
39995168_Geobacter_sulfurireducens_PCA
83591519_Rhodospirillum_rubrum_ATCC_11170
50085564_Acinetobacter_sp_AdP1
77162605_Nitrococcus_oceanii_ATCC_19707
15602176_Pasteurella_multocida_subsp._multocida_str_F
32834755_Corynebacterium_diphtheriae_NCTC_13129
21674788_Chlorobacter_tepidum_TLS
53804737_Methylcoccus_capsulatus_str_Bath
209966592_Rhodospirillum_centeratum_SW
8988534_Rhodoferrax_ferrireducens_T118
1613062_Escherichia_coli_str_R-K2_substr_MG1655
68535718_Corynebacterium_jeikeium_K411
68738746_Frankia_sp_Cc13
51891801_Symbiobacterium_thermophilum_IAM_14863
29834079_Streptomyces_vermisilensis_MA-4680

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407462288_Candidatus_Nitrosopumilus_koreensis_AR1
1229351_Methanoscarcina_mazei_Gol
147921327_Methanocella_arvoryzae_MR850
124485266_Methanocorpusculum_labreanum_Z
307352568_Methanoplanus_petrolearius_DSM_11571
154150254_Methanoregula_boonei_6A8
192852754_Methanospaeraula_palustris_E1-9c
126178701_Methanoculleus_marisnigri_JR1
397780289_Methanoculleus_bourgensis_MS2
156937861_Ignicoccus_hospitalis_KIN4-I
7607609_Sulfolobus_acidocaldarius_DSM_639
187736488_Akkermansia_muciniphila_ATCC_BAA-835
463893745_Desulfovibrio_hydrotermalis_AM13_-DSM_14728
71894593_Mycoplasma_synoviae_53
193216853_Mycoplasma_arthritis_158L3-1
187250661_Ellusimicrobium_minutum_Pei191
325972002_Sphaerochaeta_globus_str_Buddy
47458867_Mycoplasma_mobile_163K
269123825_Streptobacillus_moniformis_DSM_12112
57865886_Staphylococcus_epidermidis_RP62A
319957207_Nitriradicator_salsuginis_DSM_16511
310286727_Bifidobacterium_bifidum_S17
90961086_Lactobacillus_salivarius_UCC118
406600270_Leuconostoc_gelidum_JB7
386318631_Staphylococcus_pseudintermedius_ED99
302336019_Olsenleueria_uli_DSM_7084
15675042_Zeptoococcus_pyogenes_M1_GAS
52684362_Acidaminococcus_intestini_RyC-MR95
42525844_Treponema_denticola_ATCC_35405
169823756_Finegoldia_magna_ATCC_29328
117929157_Acidothermus_cellulolyticus_11B
283456134_Bifidobacterium_dentium_Bd1
38232679_Corynebacterium_diphtheriae_NCTC_13129
238924076_Bifidobacterium_rectale_ATCC_33656
116627543_Streptococcus_therophilus_LMD-9
57238545_Campylobacter_jejuni_RM1221
92176264_Helicobacter_mustelae_12198
294086112_Candidatus_Puniceispirillum_minarum_IMCC1322
189485226_uncultured_Termite_group_1_bacterium_phylotype_RS-D17
222109284_Acidovorax_ebreus_TPSY
15602991_Pasteurella_multocida_subsp._multocida_str_-Pm70
310780383_Ilyobacter_polytropus_DSM_2926
220930481_Clostridium_cellulolyticum_H10
43557789_Wolinella_succinogenes_DSM_1740
90425960_Rhodopseudomonas_palustris_BisB18
209542526_Glucuronacetobacter diazotrophicus_PAl_5
121608212_Verminephrobacter_eiseniae_EP01-2
60683388_Bacteroides_fragilis_NCTC_9343
154250554_Parvibaculum_lavamentivorans_DS-1
283778294_Pirellula_staleyi_DSM_6068
154174048_Campylobacter_cervus_525-92
292492938_Nitrosococcus_halophilus_Nc4
111224947_Frankia_alni_ACN14a
385089790_Ignavibacterium_album_JCM_16511
150021526_Thermosiphon_melanesiensis_BI429
28211139_Clostridium_retani_E88
48477075_Picrophilus_torridus_DSM_9790
41614813_Nanoarchaeum_egutians_Kin4-M
55376278_Haloarcula_marismortui_ATCC_43049

389844445_Mesotoga_prima_MesG1-Ag-4-2
 390962116_Thermococcus_sp_CL1
 13540937_Thermoplasma_vulcanium_GSS1
 289596694_Aciduliprofundus_boonei_T469
 14590110_Pyrococcus_horikoshii_Ot3
 84489236_Methanospaera_stadtmanae_DSM_3091
 268317185_Rhodothermus_marinus_DSM_4252
 18977490_Pyrococcus_furius_DSM_3638
 427708213_Nostoc_sp_PCC_7107
 11720226_Thermofilum_pendens_Hrk_5
 118497969_Francisella_novicida_U112
 332661941_Haliscmenobacter_hydrossis_DSM_1100
 326793969_Marinomonas_mediterranea_MMB-1
 220934970_Theiakalivibrio_sulfidophilus_HL-EbGr7
 289847693_Allochromatium_vinosum_DSM_180
 43192970_Thioflavobacterium_mobilis_s821
 206890537_Thermodesulfobacter_yellowstonii_DSM_11347
 52425690_Hannheimia_succiniciproducens_MBL55E
 302481421_Nitrosomonas_europaea_ATCC_19718
 121997419_Halorhodospira_halophilica_SlL
 430760780_Thioalkalivibrio_nitratreducens_DSM_14787
 292492856_Nitrosococcus_halophilus_Nc4
 34541577_Porphyrmonas_gingivalis_W83
 88602029_Methanospirillum_hungatei_JF-1
 325980970_Nitrosomonas_sp_AL12
 26979874_Veillonella_parvula_DSM_2008
 332668016_Haliscmenobacter_hydrossis_DSM_1100
 379728404_Saprosphaera_grandis_str_-Lewin
 327314603_Pivotella_denticola_F0289
 392395717_Flexibacter_litoralis_DSM_6794
 150398872_Methanococcus_vannielii_SB
 15040149_Methanococcus_aeolicus_Nankai-3
 284040851_Spirosooma_linguae_DSM_74
 257125862_Leptotrichia_buccalis_C-1013-b
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 319790504_Thermovibrio_ammonificans_HB-1
 337288662_Thermodesulfobacterium_geofontis_OPF15
 20094748_Methanopyrus_kandleri_AV19
 386750121_Helicobacter_cetorum_MIT_00-7128
 126465226_Staphylothermus_marinus_F1
 170290046_Candidatus_Korarchaeum_cryptofilum_OPF8
 385805115_Fervidicoccus_fontis_Kam940
 15898245_Sulfobolus_solfataricus_P2
 156937921_Ignicoccus_hospitalis_KIN4-I
 124027534_Hyperthermus_butilyticus_DSM_5456
 118431380_Aeropyrum_pernix_K1
 307595807_Vulcanisaeta_distibuta_DSM_14429
 305663330_Ignisphaera_aggregans_DSM_17230
 347523053_Pyrolobus_fumarii_LA
 302347808_Acidilobus_saccharovorans_345-15
 218884093_Desulfurococcus_kanthakensis_1221n
 389861365_Thermogladius_cellulosilyticus_1633
 15898286_Sulfobolus_solfataricus_P2
 15922975_Sulfobolus_tokodaii_str_-7
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 18311692_Pyrobaculum_aerophilum_str_-IM2
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 11849735_Francisella_novicida_U112
 328954440_Desulfovobacteria_acetoxidans_DSM_11109
 408403000_Candidatus_Nitrososphaera_gargensis_Ga9-2
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 148656004_Roseiflexus_sp_RS-1
 386854797_Deinococcus_gobiensis_I-0
 167628642_Heliobacterium_modesticaldum_Icel
 148642083_Methanobrevibacter_smithii_ATCC_35061
 288560430_Methanobrevibacter_ruminantium_M1
 11499462_Archaeoglobus_fulgidus_DSM_4304
 383319162_Methanocella Conradii_H2254
 294676823_Rhodobacter_capsulatus_SB_1003
 302343124_Desulfarculus_baarsii_DSM_2075
 389875622_Tistrella_mobilis_PSA01020-065
 374998939_Azospirillum_lipofumerum_4B
 288957883_Azospirillum_sp_B510
 116627764_Streptococcus_thermophilus_LMD-9
 15609954_Mycobacterium_tuberculosis_H37Rv
 348026605_Megasphaera_eldsenii_DSM_20460
 296133514_Thermuncula_potens_JR
 114567264_Syntrophomonas_wolfei_subsp._wolfei_str_-Goettingen
 332296542_Thermodesulfobium_paragense_DSM_14796
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 17228961_Nostoc_sp_PCC_7120
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 119357846_Chlorobium_phaeobacteroides_DSM_266
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 154250301_Fervidobacterium_nodosum_Rt17-B1
 38505760_Synechocystis_sp_PCC_6803
 83592490_Rhodospirillum_rubrum_ATCC_11170
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 32114617_Rhodomicrombium_vannielii_ATCC_17100
 219683006_Bifidobacterium_animalis_subsp._lactis_AD011
 116748794_Syntrophobacter_fumaroxidans_MP08

317051217_Desulfurispirillum_indicum_SS
328949009_Treponema_succinifaciens_DSM_2489
33396012_Treponema_azotonutricium_ZAS-9
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308282860_Lepthospira_interrogans_serovar_Lai_str_M6601
347756303_Candidatus_Clarcobacterium_thermophilum_B
162457462_Sorangium_cellulosum_So_ce56
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16112904_Bacillus_halodurans_C-125
108763792_Mycobacterium_xanthus_DK_1622
34496859_Caldicellulosiruptor_lactoaceticus_6A
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146657121_Roseiflexus_sp_RS-1
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17227877_Nostoc_sp_PCC_7120
268318378_Rhodothermus_marinus_DSM_4252
55978328_Thermus_thermophilus_HB8
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328953422_Desulfobacca_acetoxidans_DSM_11109
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320102061_Isosphaera_pallida_ATCC_43644
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7716205_Nitrosococcus_oceanii_ATCC_19707
15602176_Pasteurella_multocida_subsp._multocida_str_Pm
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21674788_Chlorobium_tepidum_TLS
53804737_Methylococcus_capsulatus_str_Bath
209966592_Rhodospirillum_centenum_SW
89885843_Rhodoferax_ferrireducens_T118
16130624_Escherichia_coli_str_K-12_substr_MG1655
68535718_Corynebacterium_jeikeium_K411
68737846_Frankia_sp_Cc13
51891801_Symbiobacterium_thermophilum_LM_14863
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 221209284_Acidovorax_ebreus_TPSY
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 220930481_Clostridium_cellulolyticum_H10
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 121608212_Verminephrobacter_eiseniae_EF01-2
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 154250554_Parvibaculum_lavamentivorans_DS-1
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 154174048_Campylobacter_curvus_525-92
 292492938_Nitroscoccus_halophilus_Nc4
 112124497_Frankia_alni_ACN14a
 385809790_Ignavibacterium_album_JCM_16511
 150021526_Thermosiphophila_melanesiensis_B1429
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 41614813_Nanoarchaeum_equitans_Kin4-M
 55376278_Haloarcula_marismortui_ATCC_43049
 389844445_Mesotoga_prima_MesG1-Ag-4-2
 390962116_Thermococcus_sp_-C11
 13540937_Thermoplasma.volcanicum_GSS1
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 14590110_Pyroccoccus_horikoshii_OT3
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 268317185_Rhodothermus_marinus_DSM_4252
 18977490_Pyroccoccus_furiosus_DSM_3638
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 431929780_Thioflavoccocus_mobilis_8321
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 121997419_Halorhodospira_halophilica_SL1
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 327314063_Prevotella_denticola_F0289
 392395717_Flexibacter_litoralis_DSM_6794
 150398872_Methanococcus_vannielii_SB
 150401497_Methanococcus_aeolicus_Nankai-3
 284040511_Spirosooma_linguale_DSM_74
 257125862_Lepotrichia_buccalis_C-1013-b
 21226661_Methanoscarcina_mazei_Gol
 218961577_Candidatus_Cloacamonas_acidimaminovorans_str_-Evry
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 386750121_Helicobacter_cetorum_MIT_00-7128
 126465226_Staphylothermus_marinus_F1
 170290046_Candidatus_korarchaeum_cryptofillum_OPF8
 385805115_Perfrividococcus_fontis_Kam940
 15898245_Sulfolobus_solfataricus_P2
 156937921_Ignicoccus_hospitalis_KIN4-I
 124027534_Hyperthermus_butyllicus_DSM_5456
 118431388_Aeropyrum_perkinii_XI
 307595807_Vulcanisaeta_distributa_DSM_14429
 305663330_Ignisphaera_aggregans_DSM_17230
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 389861365_Thermogladius_cellulolyticus_1633
 15898286_Sulfolobus_solfataricus_P2
 15922975_Sulfolobus_tokodaii_str_-7
 18311772_Pyrococcus_aerophilum_str_-IM2
 18311699_Pyrococcus_aerophilum_str_-IM2
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 288560430_Methanobrevibacter_ruminantium_M1
 11494962_Archaeoglobus_fulgidus_DSM_4304
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 37498933_Azospirillum_lipoferon_4B

288957883_Azospirillum_sp_-B510
 11626764_Streptococcus_thermophilus_LMD-9
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 348026605_Megasphaera_elsenheimeri_DSM_20460
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 33396012_Treponeema_azotonutricium_ZAaS-9
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 294828260_Leptospira_interrogans_serovar_Lai_str_-56601
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 86742032_Frankia_sp_-Cc13
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 383762167_Cardilinea_aerophila_DSM_14353_-NBRC_104270
 46255267_Thermus_thermophilus_HB27
 134101617_Saccaropolyspora_eurythraea_NRRL_2338
 378719041_Gordonia_polyisoprenivorans_VH2
 320161661_Isochrysa_pallida_ATCC_43644
 268318441_Rhodothermus_marinus_DSM_4252
 39995165_Geobacter_sulfurreducens_PCA
 83591518_Rhodospirillum_rubrum_ATCC_11170
 50085564_Acinetobacter_sp_-AdP1
 77166205_Nitrosococcus_oceanii_ATCC_19707
 15602176_Pasteurella_multocida_subsp._multocida_str_-Pm70
 38234755_Corynebacterium_diphtheriae_NCTC_13129
 21674788_Chlorobium_tepidum_TLS
 53804737_Methylococcus_capsulatus_str_-Bath
 209966592_Rhodospirillum_centenum_Sw
 89885834_Rhodotherax_ferrireducens_T118
 16130662_Escherichia_coli_str_-K_12_substr_-MG1655
 68535718_Corynebacterium_jeikeium_K411
 86738746_Frankia_sp_-Cc13
 51891801_Symbiobacterium_thermophilum_IAM_14863
 29834079_Streptomyces_avermitilis_MA_4680

AAAAYWPALGR-CL---EQ---G-----WR--LER--R--RR----RPP-PDPVNVLISYLS-AVLHRDLAAFVVRHGLPGFALH---SARD-GAQACVSDLIEEFRPL---VEGLAVLFN-NR---
 LAKSYFVYLN-LV---PD-D---FH--FNG---R---SR---FA---FQG---R---ST---FA---FEK---R---SK---RPP-LDPAFNSMVSILCY-SILKYKN1GAIERHSN1AGFHL---KHH0-HHATLSDLMEEWNPPI---VDTNLMLER--N---
 AAKYFTALGH-LV---PQ---E---FA---FQG---R---ST---FA---FEK---R---SK---RPP-LDPPNLSLSSFY-TMLMEYLTAITNEGLPHYPVGFLH---TLKE-HIPALASDLMEEWRARL---ADSFVMSLVQ---H---
 ASKHYFKALSL-MV---DR---R---FA---FSG---R---NR---RPP-LDPPNLSLSSFY-TMLLYTEVTTAVNNKGLPHYALGMLH---RDRO-GHPALASDLMEEWRPV---VDSLVM5IV0---G---
 AARYFQGLAE-LV---EP---D---FA---FSG---R---NR---RPP-LDPPNLSLSSFY-TMLMLDYTAVANRGLNPYASFLH---SIRR-GHPALASDLMEEWRRAIL---ADSLALYVTTS-K---
 IASIFEFALN-LP---IH---K-E---PK---FPG---R---NR---RPP-LDPPNLSLSSFY-VLLYORVESVLSYCEPSLDPYLGFPFH---TTDY-SKPSLALDLMEEFRAL---VDSLVLRLIN-K---
 AAARYFPAFGQ-LI---TNAA---FS---FSL---R---NR---RPP-TDPVNLSLSSFY-TLLFNVNLSSLIISSEGLSPVYFGNPH---YGER-DKPYLAFLDMEEFRAII---VDMVLVRIN-N---
 AATLYFQQLGS-LP---TG---A---PV---FEK---R---TK---RPP-TDPVNLSLSSFY-TLLSQNFVSPVQVIGLHTMFGNLH---VPRD-NUPVLSLDMEEFRQAL---VDSLVAYLIN-S---
 GAATYFQVFED-FP---DT---G-G---PI---FRE---R---VK---RPP-TDPVNLSLSSFY-SLLFNNFISMARLHRLHPYVGFLH---ADKP-AHPALVSDLIEEFRTL---VDGLVIALIN-K---
 ASALYFECIGM-LF---SK---NLP---FH---TRSLFLR---VR---RPP-TDPVNLSLSSFY-TMLHTNIFSMVQASGLNPYIGFLH---AERK-GNPALVNDLVEEFRTI---VDSLVLITLN-R---
 AARVIFPCLQG-LF---PA---E---LQ---FTL---R---SR---RPP-TDPVNLSLSSFY-TMLLNFVNLSSLF---MDIANASALSYLY-TLLTAECVTLVAAGLDPAGFLH---ADDK-GRPSLALDLMEEFRPHV---ADQVVITAAR-N---
 AAREYFQAFQGQ-FQ---SA---D---WG---FNS---R---NR---RPP-TDPVNLSLSSFY-TMLLNFVNLSSLF---MDIANASALSYLY-TLLTFNIRHALVEARGLPHYLGFLH---GSQP-RQPSPVLDLMEGRPWL---VDELVLQLAG-K---
 GSKYIFSGFKK-CI---RT---EE---LT---FNG---R---TA---RPP-KDEINAMLSLCY-YFLYFEMELLAIAVAGLDPYFGNHL---TIDV-SKQSLFLDLEEFRCVI---IDFNVLNLIN-L---
 AANLYFQAWAD-CI---E-NSN---FI---FPG---R---SR---RPP-GNPVNFMSLFCY-TLLWNHLNLSSLF---VDSLVMYMIN-R---
 GAHQYFAGLRT-AP---VG---G---IP---FLG---R---AQ---RPP-DPDANSLLSFCY-VLLNGRNRLTGLEAVGDPVCFGPFLH---DLPQ-GRPSLALDLMEEFRAL---VDRALRLICN-L---
 GARIYFQFGR-CL---AC---G---LD---FPG---R---KY---RPP-TDPVNLSLSSFY-TLLTARSLSSLYSGLDPMFPRHPL---EDPS-GRNLSQADLMEEFRTPL---VDRIVLFLFN-R---
 AAERYFPRDLDE-TL---VD---G---WT---FEK---R---TK---RPP-EDHNSLSSLY-VVNEKNEVMSALRQYQNLDPFLGVHLH---ADRQ-GRPSLALDLMEEFRP1F---CDALVTRLVN-R---
 ATAFYFSPAMPH-MP---AA---RAGD---VE---FEV---R---SR---RPP-QNAVATNLSAYA-AVLASECVAALAVGLDPRLGVHLH---QPRPS-GRASLALDLMEEFRPLI---ADQAVLTGFN-T---
 AAALYFQSOFH-LI---SV---NKRNE---WP---WLE---MR---RPP-KDPDNATLSSFY-SLLNSDCIRAVSCLDAHGFLH---SSKR-NKPALALDLMEEFRAPI---ADSVQQTVN-N---
 AANLYFQFVPL-LV---RV---PG---F---FNG---R---NR---RPP-LDPPNLSLSSFY-TLLTOEVLTAIKVVGFLYLGFLH---AVDY-GRPSLALDLMEEWRTPF---GDELVLALVN-R---
 CAATYFQAFGS-IP---EDRE---FQ---FRK---R---SR---RPP-EDPDLNALSFCY-TLLYRDELTALVQSTGDLAVGVYGLH---SAEY-GRPSLALDLMEEFRFV---VDELVVLKVN-L---
 AARKYFQFVKF-NL---OD---E---WA---EFKT---R---SK---RPP-RTNVAFLSFLY-TLLMRYVESIAQSGLDSCCGNLH---AVNY-GTKALVFLDLMEEFRSPV---CDSVCCSLFN-L---
 AARNF59612_Treponeema_azotonutricium_ZAaS-9
 1-IP---E---WA---QFR---R---SK---RPP-LTNINSVLSSFLY-TLLNMRVETAIETTGLDPYAGFLH---AAEY-GKNALVFLDLMEEFRTP1---ADTLCALCN-LG---
 ISSIYFSAFDCLLN---KE---IK---FEK---R---TR---RPP-EDHNSLSSLY-TLLNSLIRTEVITIGLDPYFGVHLH---AEDY-GRPSLALDLMEEFRALA---VDMPAINAVN-K---
 AAKYFQFAPD-LI---ANPD---FP---FSG---R---NK---RPP-KDEVNAMLSLFCY-TLLMNQVMCAIYCGLDPYAGFLH---DLDY-GRQSLVLDLIMEEFRPL---IDNMVILSLIN-R---
 SARAYFSGLPA-LK---NKNPDS---LV---PNG---R---SK---RPP-KDPDNATLSSFY-SLLNSDCIRAVSCLDAHGFLH---CDMTLIGSIN-R---
 AQQYFQAFAS-LI---RT---SE-PPSEPMSGPAG---FE---FVG---R---NR---RPP-RDPRVNALLSSFY-TLLYRDELTALVQSTGDLAVGVYGLH---TFRS-AAHPLALDLMELFVPL---ADMVVASIN-R---
 AARYGFOCALAALVH---PDAGDA---LR---PRG---R---SR---RPP-EDPRVNALLSSFY-TLLYRDELTALVQSTGDLAVGVYGLH---QFRS-AAPFLALDLMELFVPL---VDMAVLAVN-R---
 GARRYFQATWPH-L---LL---TP---HATV---TA---PAG---R---TS---RPP-TDPVNLSLSSFY-TLLMRYVESIAQSGLDSCCGNLH---GIRD-GKPALALDLMEEFRALL---VDRIVLFTAFN-Q---
 GAKRYFAEWSL---S-PPQG---FF---FTA---R---IR---RPP-MDPVNALLSSFY-TLLYRDELTALVQSTGDLAVGVYGLH---PVRP-GRPSLALDLMEEFRAPF---VDRIVLTLIN-R---
 FA---FHG---R---SR---RPP-KDKNVANLSSFY-TLLANDVAALAEVGLDPYAGFLH---QDRP-GRASLALDLMEEFRPL---ADRFVLSLIN-R---
 AARDYFESFPA-L---LKKSAQG---FE---FDQ---R---NR---RPP-RNPLNALSFCY-ALLQADCAAGLTVGVLDPYAGFLH---EDRP-GRSLALDLMEEFRAPV---VDRIVLFLVN-R---
 ISKIYFEGFKK-VL---PP---E---PC---FEE---R---NR---RPP-RDPPVNALLSSFY-TTFLQCTIGAPLFGPYAGFLH---SNVY-GRPALALDLMEEFRP1---ADAFVLSLIR-N---
 GSALYFEEAFGR-GF---IE---N---QG---FPR---R---VR---RPP-TDPVNLSLSSFY-TPLMRVVAAVVAGLDPYAGFLH---ALDY-GRYSLVLDLIMEEFRTII---ADTTLTSLFN-L---
 AAAAYFQANRA-SL---PP---V---WG---FGG---R---AF---RPP-PPDPNALSSFY-TLLYRDELTALVQSTGDLAVGVYGLH---VIEP-GRPSLALDLMEEFRPLI---VDRIVLTLR-T---
 GAAWCNTWSA-VF---KT---A---WG---FAG---R---AY---RPP-EDPDLNALSFCY-TLLNLDLMTAVQALSDFVPLGVFLH---TQVF-GRPSLALDLMEEFRPCI---VDRMVLVLDL-AG---
 CARTYQALKH-WF---PT---O---WN---FNG---R---NR---RPP-LDPPNALLSSFY-GVLLARVPSACVQAGLDPYAGFLH---AIEP-YEPNLVLDLIMEEFRTV---VDRIVLTLIN-R---
 ASRKFVSPGE-ML---CH---E---TG---R---FR---RPP-TDPVNALLSSFY-GILYRIVLAVGDPHIGFLH---GIRD-GKPALALDLMEEFRPL---VDRIVLFTAFN-Q---
 TGWPAFPG---R---HR---RPP-TDPVNALLSSFY-TLLANDVVAACALGVDPYAGFLH---APAY-GRPSLALDLMEEFRAPF---ADRLAFLRLN-R---
 GSRVYFQGLAR---LL---GP---Y---FGG---R---TR---RPP-TDPVNALLSSFY-YALLLGNRLVAVLAVGLDPYAGFLH---AEGR-RSPALALDLMEEFRFV---VDRIVLTLIN-R---
 CARLYFQGLAR-PL---AA---R---Y---FTG---R---NR---RPP-TDPVNALLSSFY-YALLLGNRLVAVLAVGLDPYAGFLH---AERD-RSPALALDLMEEFRFV---VDRIVLTLIN-R---
 AAHTYFQGLAR-MV---RH---G---FG---FSG---R---QH---RPP-TDPVNALLSSFY-TLLVNMVVAASLTDLIMEEFRTV---QPRF-GHATLASLDEEFRAL---VDRITLTLSLIN-N---
 AAQVYFQFVGK-LF---QV---EG---LK---FPG---R---IR---RPP-PDINALLSSFY-TLLAQDVCVAQVAGLDPYAGFLH---PEVEY-GRPSLVLDAEWRPL---VDSLVVRLRN-WK---
 GAAQYFCFCQH-LF---RV---DD---IT---FSG---R---NR---RPP-QDPVNLLVLSFCY-TLLANFAQ1VSVAESTGDPYAGFLH---SVEY-GRPSLVLDAEWRPL---VDAVVVQVN-KR---
 AAREYFQAFQE-ML---CH---E---WT---FTG---R---FR---RPP-TDPVNALLSSFY-TLLYRDELTALVQSTGDLAVGVYGLH---ETTR-QPAMALDLMEEFRPV---RSDP-GRPALALDIEEFRTV---ADSVVLTVLK-QR---
 GSRDFVSPVGL-LIKDRVP---FD---FNK---R---SR---RPP-EDPANALLSSFY-SLLKLNDLMTAVQALSDFVPLGVFLH---RSDP-GRPALALDIEEFRTV---ADSVVLTALN-KG---
 GSAQAFQCNQ-LI---RVDN---FS---FHT---R---NR---RPP-IDPVNLLSFCY-SLLRLLNQFLGALVAINVGFDPYAGFLH---TERY-GRPSLALDLMEEFRPLI---VDAVVLTAIN-R---
 GSRVYFQFNL---DS---FN---FNG---R---NR---RPP-GKPNVNCLLSFCY-SLLNNDVLAAALYVGFDPYAGFLH---SSQY-GKPALALDLMEEFRPVI---VDSVSVKSVN-KK---
 GSAQAFQCNQWCM-LL---NO---P---WE---WHG---R---RR---RPP-PDPPVNALLSSFY-SLLYTSQVLQSLAIVQAGLDPYAGFLH---QSSY-GKPALALDLMEEFRPVI---VDSVSVLWN-TK---
 ATAIFYFSGLWM---LP---NP---E---WK---FDA---R---LR---RPP-PDPPVNLLVLSFCY-TLLAQDCVVAQVAGLDPYAGFLH---EVAY-NRPALGLDLMEEFRPLI---VDGVLVICH-SG---
 ATARYFQAVRA-LI---GA---E---WR---FEA---R---IR---RPP-PDRVNALLSSFY-TLLVHKMLGVAEAGAFLDPYAGFLH---HIDY-GRPSLALDLMEEFRP1---VDSVLVIRCCN-DG---
 GTAIYFQANKS-LI---KK---P---WQ---FHT---R---QR---RPP-PPDVNLLSFCY-TLLSNQNLVAGLAVGDPYAGFLH---OLSY-NRPNLALDLMEEFRPV---VDSVVLRCRN-NS---
 AADLYFRAEFA-L---PE---E---FA---FGE---R---SR---RPP-RDPAFNLLSFCY-TLLAKECESALVAGLDPYAGFLH---EVRY-GRPSLALDLMEEFRSVL---ADSVVLSLNN-N---
 AARTYFQALGE-FT---TR---DHRLLGP---AID---TMG---R---TR---RPP-DRAVSLLFCY-CLLKDITTAQYALGDPYAGFLH---QFRH-GRPAITLDAEFEFRPLI---ADSTAFLTLIN-N---
 AARYVFNENFS-G---LL---RG---ALDQJIAS---EFD---TNG---R---KR---RPP-PPDVNALLSSFY-YALLLGNRLVAVLAVGLDPYAGFLH---ADSAVVFMLN-N---
 IARLYFQPNFG-LI---KP---DDNHTPHQPPSDDFRPLTFH---R---NR---RPP-RDPPVNALLSSFY-SLLRLLNQFLGALVAINVGFDPYAGFLH---GCRCLQALDLMEEFRPLI---ADSAVLFAIN-T---
 AARLYFQFQSLR---ML---RH---RIAPD---FKN---R---NR---RPP-KDPDNALSSFY-ALLLKDAMCALMATGLDPYAGFLH---QMRP-GRPSLALDLMEEFRPLI---ADSVVLRVN-TG---
 SAGRYFQFSR---ML---RG---GDGEOM---GFD---FTT---R---NR---RPP-KDPVNALLSSFY-AMYLARVLAVALAVGLDPYAGFLH---QPRP-GRPALALDMMEEFRPLI---ADSTVLMAIN-NG---
 AAAVYFRAFQFQ-LP---TQ---AVTTLPL---APF---PER---R---NR---RPP-ADPVNALLSFCY-AVLRTLTSALLSIAQGLDPYAGFLH---TERP-GRPALALDIESFRPV---ADSTVLMVN-NG---
 WAKALYKELAKQFN---ID---FKA---DEGNKH---DSL-TDIANNYLDHGQ-YIAYGAYVAVHANGMCGISFALILH---GKTT-RGGGLVPLDADLVKDAF---VMPFAFICAS-K---
 FAKLYKLYFWAN-CT---ET---EN---FTR---D---PG---KRDNPNDPNSYLLDQH---YLVYQIAAVALWGLWGPJHSPLVTH---GTT-RGGVLFVDPADILKDTC---VMPIAFHOAA---
 LITKLKYLANQATN---NEPD---FKR---SE---RKGKSIDLNQFLDQH---YIAYGQIAAVALWGLWGPJHSPLVTH---GKT-RGGGLVPLDADILKDSM---ILPQAFISAQO---
 IMRNYTENAKKAG---IRG---FKR---DTKAADPVNVGLNISN---R---Y---SILYQAAVATCTAIGVNPALGIH---DGT-RDTRSLFLDADLYKQVQ---SIDLAPSLTA-DM---
 RVRKLYEEMAVKY---VG---WKGRPFEP---GK---FEM---SDTNTKLLTASN-AALYSIIISLVAHSGMYSPHIGFIH---SASP---LPFIYDADLYKQVQ---SIDLAPSLTA-DM---
 53804737_Methylococcus_capsulatus_str_-Bath
 209966592_Rhodospirillum_centenum_SW
 89885834_Rhodotherax_ferrireducens_T118
 16130662_Escherichia_coli_str_-K_12_substr_-MG1655
 68535718_Corynebacterium_jeikeium_K411
 86738746_Frankia_sp_-Cc13
 51891801_Symbiobacterium_thermophilum_IAM_14863
 29834079_Streptomyces_avermitilis_MA_4680

QR---IKLKEYSPTKDG---T-VV---LEYSLIKRFLEMLERQFSQERKYSFRHGKKTQDGLKSQVQIIVVKKIIQ---NLVEYS-TGK---QKSLES
 -G---AMESK-DFIRTE---N-Y---N-LR---LKPTGARKIVNEFSNT-LNKKVSYQK---ESTWSVYIFPLKVR---ELAHYL-TSK-KEKLDFTK
 -K---KLKNN-DFIVTE---N-Y---H-LR---LREATAKKLIERIKLN-FNLKAPYKNQ---NYTYENILIDQVQ---QFANFI-QDK-NKTVFETT
 -G---RLDG-AVEVIT---R-CL---LSNEKEFMKRHLGS-IN---ITLIEENNV---RYAESV-KDG-RDIVFH-
 -B---YLTDDMYETNTK---R-CH---LDQGTRLRLSNAKLS-ID---QKRLCQVH---NPLQAL-SNK-EDLTIL
 -S---LNQVG-DPEITS---R-N---C-CT---LSDN1RMLMHLLQTS-ID---VTAIGROVE---MLADAL-IRN-REFQILD
 -G---ISSG-DYECIGE---S---R-CT---LGGDLSRRAALARLDS-ID---QARIDAQVR---IVRDAL-LAN---
 -E---ISSE-DSYECSE---K---R-CY---LSDALRMLHLLQTS-ID---QSRVDTQVOC---ILRDAL-LTHN
 -E---EISED-DSYECSE---K---R-CY---LNEKSLQVTLFQFESK---KYSRVKWWYAKAW---QLRDAI-ESGK
 -L---KLKDONS---GY---LNEKSLQVTLFQFESK---KYSRVKWWYAKAW---QLRDAI-ESGK
 -G---KTEER---A-GE---ITREFRQHITATLQAS-VMYRDK---QLPLKAAVEAVCR---SFRKAV-LAG-QSE---
 -N---IITAK-DPEPVT---AK---K-FK---LNGRSMGKILESFKE-MCIRIATE---AGTQAQLDAQVQ---NLRNWV-MDR-DELKFFR
 -IIEKNYDF---P---IINFKKVPLIFEK---ILLNKS---PISVNEYI---CK-LIENFI-NKD---
 -ISRGEPNF---QEFKEELTIFYKHF-FNCQGE---NLMLIDCIDKLVV---LVLSDH-DIKG---

Continued

407462288_Candidatus_Nitrosopumilus_koreensis_AR1
 21229351_Methanosaeca_mazei_Gol
 147921327_Methanocella_arvoryzae_MRE50
 124485266_Methanococcus_corpusculum_labreanum_Z
 307352568_Methanoplanus_petrolearius_DSM_11571
 154150254_Methanoregula_boonei_GA8
 219852754_Methanospaera_palustris_El-9c
 126178701_Methanococcus_marisnigri_JR1
 397780289_Methanococcus_bougeriensis_MS2
 156937861_Ignicoccus_hospitalis_KIN4-I
 70607609_Sulfolobus_acidocaldarius_DSM_639
 187736488_Akkermansia_muciniphila_ATCC_BAA-835
 436839745_Desulfovibrio_hydrothermialis_AM13_-DSM_14728
 71894593_Mycoplasma_synoviae_53
 193216853_Mycoplasma_arthritidis_158L3-1

187250661_Elusimicrobium_minutum_Pei191
 325972002_Sphaerochaeta_globus_str_Buddy
 47458867_Mycoplasma_mobile_16K
 269123825_Streptobacillus_munilitoris_DSM_12112
 57865880_Staphylococcus_epidermidis_RP62a
 319957207_Nitratireducens_salsuginis_DSM_16511
 310286727_Bifidobacterium_bifidum_S17
 90961086_Lactobacillus_salivarius_UCC118
 406600270_Leuconostoc_gelidum_JB7
 38618631_Staphylococcus_pseudointermedius_ED99
 302336019_Olsenella_oli DSM_7084
 15675042_Streptococcus_pyogenes_M1_GAS
 352684362_Acidaminococcus_intestini_TYRC-MR95
 42525844_Treponema_denticola_ATCC_35405
 169823756_Finegoldia_magna_ATCC_29328
 117929157_Acidothermus_cellulolyticus_llb
 283456134_Bifidobacterium_dentinum_Bd1
 38232679_Corynebacterium_diphtheriae_NCTC_13129
 238924076_Eubacterium_rectale_ATCC_33656
 116627543_Streptococcus_thermophilus_LMD-9
 57238545_Campylobacter_jejuni_RM1221
 291276264_Helicobacter_mustelae_12198
 294086112_Candidatus_Punicispirillum_marinum_IMCC1322
 189485226 uncultured_Termite_group_I_bacterium_phenotype_Rs-D17
 222109284_Acidovorax_ebreus_TPSY
 15602991_Pasteurella_multocida_subsp._multocida_str_-Pm70
 310780383_Ilyobacter_polytropus_DSM_2926
 220930481_Clostridium_cellulolyticum_H10
 34557789_Wolinella_succinogenes_DSM_1740
 90425960_Rhodopseudomonas_palustris_BisB18
 209542526_Glucanacetobacter_diazotrophicus_Pal_5
 121608212_Vermiphobacter_eisenii_EF01-2
 60683388_Bacteroides_fragilis_NCTC_9343
 154250554_Parvibaculum_lavamentivorans_DS-1
 283778924_Pirellula_staleyi_DSM_6068
 154174048_Campylobacter_curvus_525-92
 292492938_Nitrosoococcus_halophilus_Nc4
 111224947_Frankia_aini_ACN14c
 385807970_Ignavia bacterium_JCM_16511
 150021526_Thermosiphon_melanesiensis_B1429
 28211139_Clostridium_tetani_B88
 48477075_Picrophilus_torridus_DSM_9790
 41614813_Nanoarchaeum_equitans_Kin4-M
 55376278_Haloarcula_marismortui_ATCC_43049
 389844445_Mesotoga_prima_MesG1-Ag-4-2
 309062116_Thermococcus_sp_-Cl1
 13540937_Thermoplasmata_vulcanium_GSS1
 289596694_Aciduliprofundum_boonei_T469
 14590110_Pyrococcus_horikoshii_OT3
 84489236_Methanospaera_stadtmanae_DSM_3091
 268317185_Rhodothermus_marinus_DSM_4252
 18977490_Pyrococcus_furiosus_DSM_3638
 427708213_Nostoc_sp_-PCC_7107
 119720226_Thermoflum_pendens_Hrk_5
 118497969_Fanciella_novicida_U112
 332661941_Haliscmenobacter_hydrossis_DSM_1100
 326793369_Marinomonas_mediterranea_MMB-1
 220934797_Thioalkalivibrio_sulfidophilus_HL-EbGr7
 288947693_Allochromatium_vinosum_DSM_180
 431929780_Thioflavicoccus_mobilis_8321
 206890539_Thermodesulfobacter_fibrio yellowstonii_DSM_11347
 52425690_Mannheimia_succiniciproducens_MBE155E
 30248142_Nitrosomonas_europaea_ATCC_19718
 12197419_Halorhodospira_halophila_SL1
 430760780_Thioalkalivibrio_nitratireducens_DSM_14787
 292492856_Nitrosoococcus_halophilus_Nc4
 34541577_Porphyromonas_gingivalis_W83
 88602029_Methanospirillum_hungatei_JF-1
 325980970_Nitrosomonas_sp_-AL212
 269798874_Veilleonella_parvula_DSM_2008
 332668016_Haliscmenobacter_hydrossis_DSM_1100
 377928404_Saprosira_grandis_str_-Lewin
 327314603_Prevotella_denticola_F0289
 392395717_Flexibacter_litoralis_DSM_6794
 150398872_Methanococcus_vannielii_SB
 150401497_Methanococcus_aeolicus_Nankai-3
 284040851_Spirosooma_linguale_DSM_74
 257125862_Leptotrichia_buccalis_C-1013-b
 21226661_Methanosparsina_mazei_Go1
 218961577_Candidatus_Cloacamonas_acidaminovorans_str_-Evry
 319790540_Thermococcus_ammonificans_HB-1
 337288662_Thermodesulfobacterium_geofontis_OPF15
 20094748_Methanopyrus_kandleri_AV19
 386750121_Helicobacter_cetorum_MIT_00-7128
 126465226_Staphylothermus_marinus_F1
 170290046_Candidatus_Korarchaeum_cryptofilum_OPF8
 385805115_Fervidicoccus_fontis_Kam940
 15898245_Sulfolobus_solfataricus_P2
 156937921_Ignicoccus_hospitalis_KIN4-I
 124027534_Hyperthermus_butylicus_DSM_5456
 118431380_Aeropyrum_permix_K1
 307595807_Vulcanisaeta_distributa_DSM_14429
 305663330_Ignisphaera_aggregans_DSM_17230
 347523053_Pyrolobus_fumarii_1A
 302347808_Acidilobus_saccharovorans_345-15

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 15922975_Sulfolobus_tokodaii_str_-7
 18311772_Pyrococcus_aerophilum_str_-IM2
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 34557933_Wolinella_succinogenes_DSM_1740
 118497353_Francisella_novicida_U12
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 408403000_Candidatus_Nitrososphaera_gargensis_Ga9-2
 330509017_Methanosaetae_conciliii_GF6
 148656004_Roseiflexus_sp_RS-1
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 383319612_Methanococcus_conradii_H254
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 296133514_Rhynchococcus_potens_JR
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 332296542_Thermodesulfovibacca_narugense_DSM_14796
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 172228961_Nostoc_sp_PCC_7120
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 83592490_Rhodospirillum_rubrum_ATCC_11170
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 317051217_Desulfurispirillum_indicum_SS
 328949009_Teponeuma_succinifaciens_DSM_2489
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 313673559_Caliditerrivibrio_nitroreducens_DSM_19672
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 294828260_Leptospira_interrogans_serovar_Laii_str_56601
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 148657121_Roseiflexus_sp_RS-1
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 IDF-----EPFVKPKQ-----
 -PY-----EPWLMTTIKWAG-----
 SGE-----RNLSLGEVEII-----
 FNF-----ESLEIDWNKK-----
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 IIF-----PAILK-----
 IOP-----PKYBYAWAV-----
 -SW-----EWELND-----
 PEI-----IKIDFYDLFKEL-----
 PTYLL-----LPKMIEVG-----
 IDF-----GALRECF-----
 PELGFVRCGTELCE-----
 NEV-----PNNAINDNNI-----
 LKL-----PEWRPEI-----
 IKI-----PTWTY-----
 GEI-----EPFDIV-----
 KGV-----PEFRI-----
 -----SKIKIYEI-----
 SQI-----NFYRNEL-----
 IKP-----IDYEL-----
 RSL-----RPAPAMELSHA-----
 PEYR-----GDYEKG-----
 PIWE-----GPFDAER-----
 YNL-----IGEEYG-----
 SKFHPVVSSELRWK-----
 -EL-----KIVKFD-----
 ESNGREVYESFSDV-----
 MPM-----PQLIPPELEIAVMSEC-----
 LLL-----PEFKTA-----
 PRLOLQRQHVFEI-----
 LPL-----KEHOYE-----
 IPI-----QEHRYY-----
 PEI-----IPLEVHANE-----
 VQG-----EVYEADGDV-----
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 ISY-----PER-----
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 -RF-----EPFRHP-----
 MPY-----VGFWWG-----
 AEW-----ESFKRKV-----
 TPY-----TPFRPPW-----
 KSY-----DPYRAN-----
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 EDY-----KALHNMW-----
 ETY-----GPFRPW-----
 EDY-----KSFKRS-----
 KRY-----RAFVRWW-----
 KEY-----KSIRRAW-----
 KTY-----ESFKTW-----
 EPY-----RGLHMWW-----
 EY-----SPIVAWF-----
 EFY-----RLALKFPHANSDFNTIKPQTVPPLKLVR-----
 LAP-----TWDKRYIIHE-----
 AEW-----PIFCYETRRVVVDSQL-----
 -HH-----LPID-----
 --F-----IAWSPKR-----
 PAF-----EPFRTR-----
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 18311699_Pyrobaculum_aerophilum_str_-IM2
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 368654797_Deinococcus_gobiensis_I-0
 167628642_Heliobacterium_modesticalm_Ice1
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 15609954 Mycobacterium_tuberculosis_H37Rv
 348026605 Megaspheara_elsdenii_DSM_20460
 296133514 Thermuncola_potens_JR
 114567264 Syntrophomonas_wolfei_subsp_-wolfei_str_-Goettingen
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 119357853 Chlorobium_phaeobacteroides_DSM_266
 119357846 Chlorobium_phaeobacteroides_DSM_266
 336116789 Microlunatus_phosphorus_NM-1
 37677204 Vibrio_vulnificus_YJ016
 154250301 Fervidobacterium_nodosum_Rt17-B1
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 261415948 Fibrobacter_succinogenes_subsp_-succinogenes_S85
 76802276 Natronomicrobium_pharaonis_DSM_2160
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 116748794 Syntrophobacter_fumaroxidans_MPOB
 317051217 Desulfurispirillum_indicum_SS
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 33396012 Teponema_azotonutricium_ZAS-9
 313673559 Calditerrivibrio_nitroreducens_DSM_19672
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 294828260 Leptospira_interrogans_serovar_Lai_str_-56601
 347756303 Candidatus_Chloracidobacterium_thermophilum_B
 1624257462 Sorangium_cellulosum_So_ce56
 86742032 Frankia_sp_CcI3
 433631663 Mycobacterium_cantellii_CIPT_140070010
 15612904 Bacillus_halodurans_C-125
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 85858452 Syntrophobacter_aciditrophicus_SS
 148657121 Roseiflexus_sp_RS-1
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 268318378 Rhodothermus_marinus_DSM_4252
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 328953000 Desulfobacca_acetoxidans_DSM_11109
 337286709 Thermo desulfatator indicus_DSM_15286
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 134299482 Desulfotomaculum_reducens_MI-1
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 320161859 Anaerolinea_thermophila_UNI-1

156741961_Solefplexus_castenholzii_DSM_13941
387362167_Caldilinea_aerophila_DSM_14553__NBRC_104270
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134101617_Saccharopolyspora erythraea_NRRL_2338
378719041_Gordonia_polyisoprenivorans_VH2
320102061_Isosphaera_pallida_ATCC_43644
268318441_Rhodothermus_marinus_DSM_4252
33995168_Geobacter_sulfurreducens_PCA
83591519_Rhodospirillum_rubrum_ATCC_11170
50085564_Acinetobacter_sp._ADP1
7716205_Nitrosococcus_oceanii_ATCC_19707
15602176_Pasteurella_multocida_subsp._multocida_str._Pm70
27384755_Corynebacterium_diphtheriae_NCTC_13129
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53804737_Methyloccocus capsulatus_str._Bath
209966592_Rhodospirillum_centeratum_SW
39885834_Rhodoferrax_ferrireducens_T118
16130662_Escherichia_coli_str._K-12_substr._MG1655
68535718_Corynebacterium_jeikeium_K411
68737846_Frankia_sp._CC13
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2834079_Streptomyces_avermitilis_MA_4680

ADY----EPPAVR
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-RY----EPPAVR
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-DY----PVFTTR
PRY---IPVTR
PAY---PNFVTR
PRY---PHYCPR

-QL--
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92176263|Helicobacter_mustelae_l2198
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42525845|Treponema_denticola_ATCC_35405
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88602028|Methanospirillum_hungatei_JF-1
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124027532|Hyperthermus_butyllicus_DSM_5456
18311698|Pyrococcus_aerophilum_str_-IM2
150398873|Methanococcus_vannielii_SB
712572561|Leptotrichia_buccalis_C-1013-b
150401498|Methanococcus_aeolicus_Nankai-3
21226660|Methanosaeta_mazei_GoI
218961576|Candidatus_Cloacanomas_acidaminovorans_str_-Evry
284040852|Spirosoma_lingue DSM_74
15898244|Sulfolobus_solfataricus_P2

332668018 | *Haliscomenobacter hydrossis* DSM_1100
 386854796 Deinococcus_gobiensis_I-0
 288560432 *Methanobrevibacter ruminantium* Mi
 330509016 *Methanosaeta concilii* GP6
 20094746 | *Methanopyrus kandleri* AV19
 11499460 | *Archaeoglobus fulgidus* DSM_4304
 383319614 | *Methanocella conradii* H2254
 148656005 | *Roseiflexus* sp_-RS_1
 408403001 | *Candidatus Nitrososphaera gargensis* Ga9-2
 167628643 | *Helio bacterium modesticaldum* Icel
 170290048 | *Candidatus Korarchaeum cryptofilum* OPF8
 126465228 | *Staphylothermus marinus* Fl
 327314605 | *Prevotella denticola* F0289
 392395715 | *Flexibacter litoralis* DSM_6794
 34541576 | *Porphyromonas gingivalis* W83
 288947692 | *Allochromatium vinosum* DSM_180
 379728402 | *Saprosira grandis* str_-Lewin
 413929778 | *Thioflavoccocus mobilis* 8321
 288947691 | *Allochromatium vinosum* DSM_180
 269798875 | *Velilonella parvula* DSM_2008
 30248143 | *Nitrosomonas europaea* ATCC_19718
 206884968 | *Thermodesulfovibrio yellowstonii* DSM_11347
 121997422 | *Halorhodospira halophila* SLL
 292492855 | *Nitroscoccus halophilus* NC_4
 430760779 | *Thioalkalivibrio nitratireducens* DSM_14787
 317051220 | *Desulfurispirillum indicum* S5
 118497968 | *Fuscicella novicida* U12
 76802277 | *Natronomonas pharaonis* DSM_2160
 162457463 | *Sorangium cellulosum* So_ce56
 347756304 | *Candidatus Chloracidobacterium thermophilum* B
 24215881 | *Lepospira interrogans* serovar_Lai_str_-56601
 312114616 | *Rhombicrobium vannielii* ATCC_17100
 218884092 | *Desulfurococcus kanchatkensis* 1221n
 413929779 | *Thioflavoccucus mobilis* 8321
 39995169 | *Geobacter sulfurireducens* PCA
 83591518 | *Rhodospirillum rubrum* ATCC_11170
 332661940 | *Haliscomenobacter hydrossis* DSM_1100
 261415949 | *Fibrobacter succinogenes* subsp_-succinogenes S85
 121997422 | *Halorhodospira halophila* SLL
 302343125 | *Sulfurarcus baarsii* DSM_2075
 389875623 | *Tistrella mobilis* KA081020-065
 374998938 | *Azospirillum lipoferum* 4B
 294676822 | *Rhodobacter capsulatus* SB_1003
 288957882 | *Azospirillum* sp_B510
 319790503 | *Thermovibrio ammonificans* HB-1
 108759816 | *Myxococcus xanthus* DK_1622
 15612905 | *Bacillus halodurans* C-125
 189485224 | uncultured Termitote_group_I_bacterium_phylotype_Rs-D17
 86742033 | *Frankia* sp_Cc13
 433631662 | *Mycobacterium canettii* CIPT_140070010
 33396013 | *Trepomema azotonutricium* ZAS-9
 328949008 | *Trepomema succinifaciens* DSM_2489
 337288661 | *Thermodesulfovibrio geofontis* OPF15
 319790501 | *Thermovibrio ammonificans* HB-1
 283778925 | *Pirellula staleyi* DSM_6068
 386751020 | *Helicobacter ceterorum* MIT_00-7128
 220934971 | *Thioalkalivibrio sulfidophilus* HL_EBgR7
 15609953 | *Mycobacterium tuberculosis* H37Rv
 85858453 | *Synthrophus acidi trophicus* SB
 268318440 | *Rhodothermus marinus* DSM_4252
 320102062 | *Iso phasma pallida* ATCC_43644
 378719040 | *Gordonia polysiphonivorans* VH2
 134101618 | *Saccharopolyspora erythraea* NRRL_2338
 336116790 | *Microlunatus phosphorovorus* NM-1
 268318377 | *Rhodothermus marinus* DSM_4252
 83592489 | *Rhodospirillum rubrum* ATCC_11170
 206890293 | *Thermodesulfovibrio yellowstonii* DSM_11347
 52425689 | *Mannheimia succiniciproducens* MBL55E
 302481411 | *Nitrosomonas europaea* ATCC_19718
 292492858 | *Nitrosococcus halophilus* NC_4
 430760782 | *Thioalkalivibrio nitratireducens* DSM_14787
 348026606 | *Megaspasma eldenii* DSM_20460
 317051218 | *Desulfurispirillum indicum* S5
 337286710 | *Thermodesulfatator indicus* DSM_15286
 85858454 | *Synthrophus acidi trophicus* SB
 436839746 | *Desulfovibrio hydro thermalis* AM13_-DSM_14728
 134299483 | *Desulfotomaculum reducens* MI-1
 116627765 | *Streptococcus thermophilus* IMD-9
 37677206 | *Vibrio vulnificus* YJ016
 37677207 | *Vibrio vulnificus* YJ016
 427708212 | *Nostoc* sp_PCC_7107
 46255266 | *Thermus thermophilus* HB27
 159898908 | *Herpetosiphon aurantiacus* DSM_785
 114567263 | *Synthrophomonas wolfei* subsp_-wolfei str_-Goettingen
 296133513 | *Therminicola potens* JR
 326793968 | *Marinimonas mediterranea* MMB-1
 302391603 | *Acetobacterium arabaticum* DSM_5501
 75910375 | *Anabena variabilis* ATCC_29413
 119357847 | *Chlorobium phaeobacteroides* DSM_266
 38505685 | *Synechocystis* sp_PCC_6803
 337286707 | *Thermodesulfator indicus* DSM_15286
 119357852 | *Chlorobium phaeobacteroides* DSM_266
 344966858 | *Caldicellulosiruptor lactoaceticus* 6A
 218442810 | *Cyanobacter* sp_PCC_7424
 17227878 | *Nostoc* sp_PCC_7120

159898756 | *Herpetosiphon_aurantiacus_DSM_785*
 218961160 | *Candidomas_acidaminovorans_str_Evry*
 148657119 | *Rozeiflexus_sp_RS-1*
 320161860 | *Anaeolinea_thermophila_UNI-1*
 297622674 | *Truepera_radiovictrix_DSM_17093*
 383762166 | *Caldinella_aerophila_DSM_14535_NBRC_104270*
 156741962 | *Rozeiflexus_castenholzii_DSM_13941*
 38505761 | *Synechocystis_sp_PCC_6803*
 116748795 | *Syntrophobacter_fumaroxidans_MP08*
 322965451 | *Thermodesulfobium_narugense_DSM_14796*
 313673557 | *Calditerrivibrio_nitroreducens_DSM_19672*
 328953001 | *Desulfobacca_acetoxidans_DSM_11109*
 328953423 | *Desulfobacca_acetoxidans_DSM_11109*
 118479354 | *Fusciellula_novicida_U112*
 54296140 | *Legionella_pneumophila_str_Paris*
 34557934 | *Wolinella_succinogenes_DSM_1740*
 292492939 | *Nitrosococcus_halophilus_Nc_4*
 337288663 | *Thermodesulfobacterium_geofonitis_OPP15*
 52425691 | *Mannheimia_succiniciproducens_MBEL55B*
 154174506 | *Campylobacter_curvus_525-92*
 48477074 | *Picrophilus_torridus_DSM_9790*
 328949007 | *Treponema_succinifaciens_DSM_2489*
 11224948 | *Frankia_alni_ACN14a*
 319790502 | *Thermovibrio_ammonificans_HB-1*
 55376279 | *Halocarcula_marismortui_ATCC_43049*
 268317184 | *Rhodothermus_marinus_DSM_4252*
 150021525 | *Thermosiphon_melanesiensis_BI429*
 38580911 | *Ignicoccus_lignum_JCM_16511*
 41614812 | *Nanoarchaeum_equitans_Kin4-M*
 389844442 | *Mesotoga_prime_MesG1-Ag-4-2*
 390962115 | *Thermococcus_sp_CL1*
 13540936 | *Thermoplasma_vulcanium_GSS1*
 289596693 | *Acidiplofundus_boonei_T469*
 33359262 | *Pyrococcus_horikoshii_OT3*
 150021527 | *Thermosiphon_melanesiensis_BI429*
 84489237 | *Methanospaera_stadtmanae_DSM_3091*
 33359542 | *Pyrococcus_furiosus_DSM_3638*

-----MSDSTLYIIAYDI-----PNDKRRTKVHKLLCGY-GWTQYSFLFC-WLTKRHLVE--LRAKI---NR-LV-D-A-SL---DTIRLYRVC----GAC-VE-QA-ITI---
 -----MSLNKHKVLLQVAPDI-----VSDFKRYRPLBD-QORVKVQFCE-ILNLDKMLC---DN-T-N-P-EI---DSRYIYLLC---ANC-EK-NI-SVS---
 -----MPTIISYDI-----VDDDRRTNTVMKYLKGW-QTRVQSVFCE---ILNLDKMLC---DN-T-N-P-EI---DSRYIYLLC---ANC-EK-NI-SVS---
 -----MNDGRSPFLVAYDI-----VDDDRRLKLARLIMES-KF-GERVQGSVFE-CWLTATELER---SK-IL-K-C-EE---DSRYIYLLC---TAC-RE-NI-RTE---
 -----MAGLYTVSYDI-----SDDDRRLKLARLIMES-KF-GERVQGSVFE-CWLTATELER---SK-IL-K-C-EE---DSRYIYLLC---TAC-RE-NI-RTE---
 -----MFVIVSYDI-----PNDNRRKLVLANLKSF---GERVQGSVFE-CWLEPRELRE-LKRL---EA-RL-N-E-RE---DSRVFVPLC---SPV---EVL---
 -----MLYVIAYDI-----ADDRRRLKVQTLKLDFGGEVORVSEF-YTERNEYR---IQLRGL---RR-HL-D-I-EE---DSRVFVPLC---EHC-RA-QV-NLW---
 -----MFLYVIAYDI-----PDDARRKLVLNLBGP---GERVQGSVFE-CWLTEREYRA-LKKV---ERVNVL-N-EG---DSRVFVPLC---GAC-VA-NV-DVR---
 -----MPVIVSYDI-----PDDDRRLKKMADLLEGY-GORVQGSVFE-CWLTEREYRA-LKKV---ERVNVL-N-EG---DSRVFVPLC---GAC-VA-NV-DVR---
 -----MPVIVSYDI-----SDDDRVYRVAVLK---GERVQGSVFE-CWLTEREYRA-LKKV---ERVNVL-N-EG---DSRVFVPLC---GAC-VA-NV-DVR---
 -----MSIKAYSFVIVSYDI-----VEDRKRYKIAKLLBKG-GTRVQKSVFEC-YLTEKELIK---LNKEI---MK-KI-N-P-ET---DSRVFVPLC---KRC-KA-AI-SVL---
 -----MEKFVIVSYDI-----ADDRRKYRVDKLKLKNM-GMRVQKSVFEC-RLTDAQFLT---LKML---GR-IIN-Y-NT---DSRVFVPLC---KHC-VD-MI-EYI---
 -----MMFYAISYDI-----RDNNRRLRVLVAKILKDY-GERVQLSVFED-DLDEKSLAR---EK-CL-D-L-TA---DGLRLYPLC---GAC-RP-RI-EIM---
 -----MPTIISYDI-----TDNNRRLRVLVAKILKDY-GERVQLSVFED-DLDEKSLAR---EK-CL-D-L-TA---DGLRLYPLC---GAC-RP-RI-EIM---
 -----MQLRKLEYIAYDI-----EDNKTTRTIYIYQLLAGLKAQVKSFWG-YVSIASIELNA---IKRLF---DS-SL-T---IS---DKVFTTRVN---MHEQKLDSYF---
 -----MQIRLEYIAYDI-----RDNSRRTIYERLIGLTPVQKSFWG-YLNEQVY---IKRFL---DS-SL-T---IS---DKVFTTRVN---MHEQKLDSYF---
 -----MLKLFEVIVSYDI-----EDRNRRNKKLFEELKLMGLSP1QKSFWG-HLKLAIEKI---LPNL---CK-YC-S-DG---DKAFVVFKAR---LSSEI-VKN---
 -----MYMLITYDV-----EDRNRRNKKLFEELKLMGLSP1QKSFWG-HLKLAIEKI---LPNL---CK-YC-S-DG---DKAFVVFKAR---LSSEI-VKN---
 -----MRVIVYDV-----ETEKIREKVKRNLKKL-GVHAQWSVFS-LESYENLKV---LEEEE---G-EK-YRVAIFKIN---PKG---EI-RKI---
 -----MRDRYLVYDV-----SSSKRRNKKLFRMLRQVKSFWG-DITEAEAIR---LRLREL---SK-EL-M-V-PQ---DRVTFAA---NRR-NI-NV-THL---
 -----MAYILFYDV-----AGAQEOKKNNANRIRKAVAKEPK-LPVPQFSVPEG-EIRASDFKK---LTLNL---QX-ECVQ---EL-DSIVIYTFN---SLK-YS-ER-IVI---
 -----MVNVILYDV-----EKSQKQVLSVFPQG-ELTAKQQQS---LKDDEL---LE-II-N-Q-EK---DSIIFYFAE---EQK-HI-K-X-KIL---
 -----MEKHMLVYDV-----RDTKRLAKISKCVESY-GMRVQKSVFES-TASDETIGO---LKSRL---EX-IM-D-E-EE---DFVFLFVEVC---ERD-KG-KI-IEL---
 -----MPVFLVYDV-----AAERNPNALRTCRKY-LHWVQRSVPEG-ELSAAQYRA-LMTTL---RD-QL-D-L-TX---DSIRVYRTR---SPA-LV-ET-EWL---
 -----MPVFLVYDV-----MRBKGRKNSTVVIYDIFNPPEEYQGAWSVSDYTRRCKIAKLLK-Y-GVRIQSRVFE-I-EISKQTIP---LKAT---QX-YC-S-KR-DKFIFYLLE---DKV-KS-RV-KER---
 -----MVYAIIVYDV-----QADRTPKFLYLRYY-LTHWQNSVPEG-ELTEGTILVE---VKETL---QS-ML-E-EE---ESVMYMRD---SGS-YV-ER-SVY---
 -----MPYYIAIVYDV-----NVRVAKMLKLFRRY-LTWVQRSVPEG-ELTEAQFQR---LQHEA---GR-IM-N-P-DE---DVVIFYELR---DVR-YS-RR-LVL---
 -----MVIVYDV-----NKKRVRQVQLTARKY-LKWQVKSFWG-KITEKNLKK---LRDQL---LE-LI-N-Q-EE---DSIIVYIME---PEF-KP-YR-KIL---
 -----MHTEPISQVAVYDV-----TNQKRLPKVLKTRKYY-MNWVQNSAFEG-ELTAKQQQS---LKDDEL---LE-II-N-K-EK---DSIIFYFAE---EQK-HI-K-X-KIL---
 -----MQYKINMYAIVYDV-----NVSQRNQIREFLRY-LWVQVNSVPEG-EISPSPSLYY---MKKIL---QS-YI-G-ET---DSIIVYVLR---DKSCLMD-KI-VL---
 -----MVVIIYDV-----NEKRVAKVVLKILRAY-LWVQVNSVPEG-ELTYAELKK---VKSGL---KR-IM-N-T-QE---DSVIFPSS---LSN-DY-K-KEI---
 -----MARYIIVYDV-----NEKRVAKVVLKILRAY-LWVQVNSVPEG-ELTYAELKK---VKSGL---KR-IM-N-T-QE---DSVIFPSS---LSN-DY-K-KEI---
 -----NEKRVAKVVLKILRAY-LWVQVNSVPEG-WLDGNEVAB---LKRKL---SR-AI-N-E-EE---DSIIFYSLP---SDS-NL-V-ETFH---
 -----MLVYIVYDV-----DESRVKVNVKYLKYR-YLWVQVNSVPEG-EISMGK---MIKG---TK-II-K-C-DE---DSIIFYKPK---VKS-YI-DN-III---
 -----MBVIVYDV-----AEERVKVNFKLRSY-LHWI0NSVFRG-DITQSQLQA---IKDGI---KG-I-E-E-DY---DSVIIYVLP---SEK-YL-KV-INI---
 -----MVWVIVYDV-----NVERVRVHKLMLRYY-LWVQVNSVPEG-ELSKAQLYB---LEMRL---KR-IV-K-C-ED---DSVIIYVLP---SEK-YL-KV-INI---
 -----MRSKIXYILVYDV-----NEKRVKVKVLKKY-IIWQONSTPEG-NVTQANIKR---MUYEL---NK-KI-N-S-KE---DSIIVYFFN---SKT-VI-RK-EVD---
 -----MGVIYILVYDI-----KIERINKVKFLKTY-LWIQNSVPEG-EITKAQVKT---MIKL---KE-LT-D-E-DE---DSIIVYKI---PKK-YL-NK-EVL---
 -----MFHMYIVVIVYDV-----GVERVNKVKFLRMH-LWVQVNSVPEG-EVTLAEEF---IKEGL---KK-IIIDE---NS-DSVIIYKLR---SMP-PR-ETL---

Continued

50085558 | *Acinetobacter_sp_AdP1*
 77166204 | *Nitrosococcus_oceanii_ATCC_19707*
 15602177 | *Pasteurella_multocida_subsp_multocida_str_Pm70*
 209966591 | *Rhodospirillum_centeren_SW*
 21674789 | *Chlorobium_tepidum_TLS*
 86737847 | *Frankia_sp_Cc13*
 38234756 | *Corynebacterium_diphtheriae_NCTC_13129*
 90111482 | *Escherichia_coli_str_K-12_substr_MG1655*
 77128445 | *Methyllococcus_capsulatus_str_Bath*
 57865885 | *Staphylococcus_epidermidis_RP62A*
 193216854 | *Mycoplasma_arthritidis_158L3-1*
 71894594 | *Mycoplasma_synoviae_53*
 189440766 | *Bifidobacterium_longum_DJ10A*
 38232680 | *Corynebacterium_diphtheriae_NCTC_13129*
 187250662 | *Elusimicrobium_minutum_Pe119*
 117929156 | *Elusimicrobium_cellulolyticus_11B*
 121608213 | *Vermiphrobacter_eiseniae_EP01-2*
 325972001 | *Sphaerochaeta_globosa_str_Buddy*
 319957208 | *Nitratireductor_salsuginis_DSM_16511*
 294086113 | *Candidatus_Punicespisirillum_marinum_IMCC1322*
 60683387 | *Bacteroides_fragilis_NCTC_9343*
 154250553 | *Parvibaculum_lavamentivorans_DS-1*
 187736487 | *Akkermansia_muciniphila_ATCC_BAA-835*
 116627544 | *Streptococcus_thermophilus_LMD-9*
 57238544 | *Campylobacter_jejuni_RM1221*
 291276263 | *Helicobacter_mustelae_12198*
 220930480 | *Clostridioides_cellulolyticum_H10*
 209542527 | *Gluconacetobacter_diazotrophicus_Pal_5*
 90425959 | *Rhodospseudomonas_palustris_BisB18*
 189485227 | *uncultured_Termite_group_1_bacterium_phylotype_Rs-D17*
 310780382 | *Ilyobacter_polytropus_DSM_2926*
 15602990 | *Pasteurella_multocida_subsp_multocida_str_Pm70*
 222109283 | *Acidovorax_ebreus_TPSY*
 238924077 | *Eubacterium_rectale_ATCC_33656*
 406600269 | *Leuconostoc_gelidum_JB7*
 90961087 | *Lactobacillus_salivarius_UCC118*
 269123824 | *Streptobacillus_moniliiformis_DSM_12112*
 34557788 | *Wolinella_succinogenes_DSM_1740*
 331702226 | *Lactobacillus_buchneri_NRRL_B-30929*
 15675043 | *Streptococcus_pyogenes_M1_GAS*
 374307736 | *Filifactor_olocis_ATCC_35896*
 302336018 | *Olsenella_uili_DSM_7084*
 169823757 | *Finegoldia_magana_ATCC_29328*
 42525845 | *Treponema_denticola_ATCC_35405*
 148642084 | *Methanobrevibacter_smithii_ATCC_35061*
 88602028 | *Methanospirillum_hungatei_JF-1*
 119720227 | *Thermofilum_pendens_Hrk_5*
 70607607 | *Sulfolobus_acidocaldarius_DSM_639*
 156937922 | *Ignicoccus_hospitalis_KIN4-I*
 18311771 | *Pyrococcus_aerophilum_str_IM2*

-----GAK-----HK---FAHDGAYPVASIKQQKLLMLD---WVKASSL-----
 -----GSR-----RH---FDSRGEFAVFL---HSATLPVSNVEAAA-PVRLNNH---
 -----GDR-----RQ---FDNQGRVPTN---RTQRNIRLAEEMAWQDAYSLQI---
 -----GAP-----KR---TIVDQDHY---ALRKATE---
 -----GDT-----RK---EITEISGLPLVIEKSAASPPENQNRLTEPAKPVQ---
 -----GRN-----RT---QPVYDGLL---LFARSRR---
 -----GDS-----TL---QVLDAGDL---VLASRPGRAAVKLHGLQ---
 -----GLN-----RR---TPVLDLCLR---LVSFLPV---
 -----GLP-----PV---KFVELDGIV---LTRP---
 -----GKE-----VRLA---VNKEPKL---FI---
 -----GSK-----SLNE---KYNDNERYI---KL---
 -----GEK-----SLNE---IYNERDKYI---EL---
 -----SGK-----REIE---EETPDFLIT---LF---
 -----SSS-----AEVSGEEKPDQLAIR---
 -----GRL-----KVAQAREPDFF---DTLL---
 -----GER-----LVP---EDDPQQLA---IPF---
 -----GRS-----RQOPT---OKAPQDFD---LP---
 -----GQO-----VNEBCPQEQLL---FE---
 -----GEO-----SSQE---EITQPTLPL---DF---
 -----GRH-----SKRP---INADQLALP---
 -----GKJ-----VOD---VNTPFGQOLE---LF---
 -----GKK-----RQP---TTQSPSQLO---LF---
 -----GRS-----CIPRO---EVSLPQOMI---LN---
 -----GKJ-----TKQ---EIVSDNKLV---AI---
 -----GGI-----VFN---VKNNETNL---LFDIDSHGEFKYKNSNNEEQLNNKQKEQYHQNLPEF---
 -----GEI-----SQAHQ---KVQKPTPLT---LF---
 -----GKX-----TSNEWIDKMMIEQGM---L---
 -----GLA-----PTERIGPSQMV---LL---
 -----GEA-----TNEKQKAPQQLV---LL---
 -----GKX-----QLAB---Q-IDSRQVL---FF---
 -----SKE-----RSPE---EKIGTSKTI---II---
 -----GRL-----RQ---KKVNNAMQI---LP---
 -----GLP-----LPQE---KKVNAQIA---LP---
 -----GSR-----QTS---TAIGTERLI---II---
 -----GTP-----SDD---VRNSDEKMV---VI---
 -----GKX-----IKE---VRNSAERTI---IL---
 -----VKE-----PKLEIJKLSPNPLV---LP---
 -----GGP-----KSO---ERIGAKQVL---LP---
 -----GKP-----KED---VRNSSERTI---IL---
 -----GER-----NNCI---ANSDERLV---FLGEAFDES---
 -----GEK-----RSNN---VDSDDRLV---IL---
 -----GNR-----SVYD---EVDTMEELL---II---
 -----GEN-----KSS---MLDITDEGLI---IL---
 -----GES-----KTE---YLNTDERLV---VL---
 -----GKE-----ITLBEKKYK---IL---
 -----HE-----EL---PKVNRQFPEV---I---
 -----GRD-----YYI---VSERKYEVY---
 -----TON-----KISKNELQIYVAGEDI---
 -----AVE-----GKVG---
 -----GFP-----P---PARKNVDLY---

385805116 | Fervidicoccus_fontis_Kam940
 15922976 | Sulfolobus_tokodaii_str_-7
 347523052 | Pyrolobus_fumarii_LA
 118431379 | Aeropyrus_permix_KI
 305663329 | Ignisphaera_aggregans_DSM_17230
 307595806 | Vulcanisaeta_distributa_DSM_14429
 302347807 | Acidilobus_saccharovorans_345-15
 124027532 | Hyperthermus_butylicus_DSM_5456
 18311698 | Pyrococcus_aerophilum_str_-IM2
 150398873 | Methanococcus_vannielii_SB
 257125861 | Leptotrichia_buccalis_C-1013-b
 150401498 | Methanococcus_aeolicus_Nankai-3
 21226660 | Methanosaeca_mazei_Gol
 218961576 | Candidatus_Cloacamonas_acidaminovorans_str_-Ervy
 284040852 | Spirosoma_linguae_DSM_74
 15898244 | Sulfolobus_solfataricus_P2
 332668018 | Haliscomenobacter_hydroassis_DSM_1100
 368854796 | Deinococcus_gobiensis_I-0
 288560432 | Methanobrevibacter_ruminantium_Ml
 330509016 | Methanosaetae_conciliif_GP6
 20094746 | Methanopyrus_kandleri_AV19
 11499460 | Archaeoglobus_fulgidus_DSM_4304
 383319614 | Methanocella_conradii_HZ254
 148656005 | Roseiflexus_sp_-RS-1
 408403001 | Candidatus_Nitrosphaera_gargensis_Ga9-2
 167628643 | Helio bacterium_modestocaldum_Icel
 170290048 | Candidatus_Korarchaeum_cryptofilum_OPF8
 126465228 | Staphylothermus_marinus_F1
 327314605 | Pectenota_denticola_F0289
 392395715 | Flexibacter_litoralis_DSM_6794
 34541576 | Porphyromonas_gingivalis_W83
 288947692 | Allochromatium_vinosum_DSM_180
 379728402 | Sphaerotilis_grandis_str_-Lewin
 431929778 | Thioflavicoccus_mobilis_8321
 288947691 | Allochromatium_vinosum_DSM_180
 269798875 | Veillonella_parvula_DSM_2008
 30248143 | Nitrosomonas_europaea_ATCC_19718
 206889468 | Thermodesulfovibrio_yellowstonii_DSM_11347
 121997422 | Halorhodospira_halophila_SL1
 292492855 | Nitrosococcus_halophilus_Nc_4
 430760779 | Thioalkalivibrio_nitratreducens_DSM_14787
 317051220 | Desulfurispirillum_indicum_SS
 118497968 | Franciscella_novicida_U112
 76802277 | Natronomonas_pharaonis_DSM_2160
 162457463 | Sorangium_cellulosum_So_ce56
 347756304 | Candidatus_Chloracidobacterium_thermophilum_B
 24215881 | Leptospira_interrogans_serovar_Lai_str_-56601
 312114616 | Rhodomicrobium_vannielii_ATCC_17100
 218884092 | Desulfurococcus_kamchatkensis_1221n
 431929779 | Thioflavicoccus_mobilis_8321
 39995169 | Geobacter_sulfureducens_PCA
 83591518 | Rhodospirillum_rubrum_ATCC_11170
 332661940 | Haliscomenobacter_hydroassis_DSM_1100
 261415949 | Fibrobacter_succinogenes_subsp__succinogenes_S85
 121997420 | Halorhodospira_halophila_SL1
 302343125 | Desulfarculus_baarsii_DSM_2075
 389875623 | Tistrella_mobilis_KA081020-065
 374998938 | Azospirillum_lipofermum_4B
 294676822 | Rhodobacter_capsulatus_SB_1003
 288957882 | Azospirillum_sp_-B510
 319790503 | Thermovibrio_ammonificans_HB-1
 108759816 | Myxococcus_xanthus_DK_1622
 15612905 | Bacillus_halodurans_C-125
 189485224 | uncultured_Termite_group_1_bacterium_phylotype_Rs-D17
 86742033 | Frankia_sp_-Cc13
 433631662 | Mycobacterium_cantellii_CIPT_140070010
 333996013 | Treponema_azotonutricium_ZAS-9
 328949008 | Treponema_succinifaciens_DSM_2489
 337288661 | Thermodesulfovibacter_geofonitis_OPF15
 319790501 | Thermovibrio_ammonificans_HB-1
 283778925 | Pirellulace_staleyi_DSM_6068
 386750120 | Helicobacter_cetorum_MIT_00-7128
 220934971 | Thioalkalivibrio_sulfidophilus_HL-EbGr7
 15609953 | Mycobacterium_tuberculosis_H37Rv
 85858453 | Syntrophus_aciditrophicus_SB
 268318440 | Rhodothermus_marinus_DSM_4252
 320102062 | Isosphaera_pallida_ATCC_43644
 378719040 | Gordonia_polyisoprenivorans_VH2
 134101618 | Saccharopolyspora_erythraea_NRRL_2338
 336116790 | Microbacterium_phosphovorus_NM-1
 268318377 | Rhodothermus_marinus_DSM_4252
 83592489 | Rhodospirillum_rubrum_ATCC_11170
 206890293 | Thermodesulfovibrio_yellowstonii_DSM_11347
 52425689 | Mannheimia_succiniciproducens_MBEL55E
 30248141 | Nitrosomonas_europaea_ATCC_19718
 292492858 | Nitrosococcus_halophilus_Nc_4
 430760782 | Thioalkalivibrio_nitratreducens_DSM_14787
 348026606 | Megasphaera_eldenii_DSM_20460
 317051218 | Desulfurispirillum_indicum_SS
 337286710 | Thermodesulfatator_indicus_DSM_15286
 85858454 | Syntrophus_aciditrophicus_SB
 436839746 | Desulfovibrio_hydrothermalis_AM13_-DSM_14728
 134299483 | Desulfotomaculum_reducens_MI-1
 116627765 | Streptococcus_thermophilus_LMD-9
 37677206 | Vibrio_vulnificus_YJ016

37677205 | *Vibrio_vulnificus* YJ016
 427708212 | *Nostoc_sp_-PCC_7107*
 46255266 | *Thermus_thermophilus* HB27
 159898908 | *Herpetosiphon_aurantiacus* DSM_785
 114567263 | *Syntrophomonas_wolfei* subsp._*wolfei* str._Goettingen
 296133513 | *Thermochlora_potens* JR
 326793968 | *Marinomonas_mediterranea* MMB-1
 302391603 | *Acetohalobium_arabaticum* DSM_5501
 75910375 | *Anabaena_variabilis* ATCC_29413
 1139857847 | *Chlorobium_phaeobacteroides* DSM_266
 38505685 | *Synechocystis_sp_-PCC_6803*
 337286707 | *Thermodesulfatator_indicus* DSM_15286
 119357852 | *Chlorobium_phaeobacteroides* DSM_266
 344996858 | *Caldicellulosiruptor_lactoaceticus* 6A
 218442810 | *Cyanotechce_sp_-PCC_7424*
 17227878 | *Nostoc_sp_-PCC_7120*
 159898756 | *Herpetosiphon_aurantiacus* DSM_785
 218961168 | *Candidatus_Cloacamonas_acidaminovorans* str._Evry
 148657119 | *Roseiflexus_sp_-RS-1*
 320161860 | *Aerolinea_thermophila* UNI-1
 297622674 | *Truepera_radiotolerans* DSM_17093
 383762166 | *Caldilinea_aerophila* DSM_14535 - NBRC_104270
 156741962 | *Roseiflexus_castenholzii* DSM_13941
 38505761 | *Synechocystis_sp_-PCC_6803*
 116748795 | *Syntrophobacter_fumaroxidans* MPOB
 332296541 | *Thermodesulfobium_narugense* DSM_14796
 313673557 | *Caliditerrivibrio_nitroreducens* DSM_19672
 328953001 | *Desulfobacca_acetoxidans* DSM_11099
 328953423 | *Desulfobacca_acetoxidans* DSM_11099
 118497354 | *Francisella_novicida* U112
 54296140 | *Legionella_pneumophila* str._Paris
 34557934 | *Wolinella_succinogenes* DSM_1740
 292492939 | *Nitrosococcus_halophilus* Nc_4
 337288663 | *Thermodesulfobacterium_geofontis* OPF15
 52425691 | *Mannheimia_succiniciproducens* MBEL55E
 154174506 | *Campyllobacter_curvus* 525-92
 48477074 | *Picrophilus_torridus* DSM_9790
 328949007 | *Treponema_succinifaciens* DSM_2489
 111224948 | *Frankia_ainii* ACN14a
 319790502 | *Thermovibrio_ammonificans* HB-1
 55376279 | *Halococcus_marismortui* ATCC_43049
 268317184 | *Rhodothermus_marinus* DSM_4252
 150021525 | *Thermisipho_melanesiensis* BI429
 385809791 | *Ignavibacterium_album* JCM_16511
 41614812 | *Nanocarchaeum_equitans* Kin4-M
 389844442 | *Mesotoga* prima MesG1-Ag-4-2
 390962115 | *Thermococcus_sp_-ClI*
 13540936 | *Thermoplasma_vulcanium* GSS1
 289596693 | *Aciduliprofundus_boonei* T469
 33359262 | *Pyrococcus_horikoshii* OT3
 150021527 | *Thermosipho_melanesiensis* BI429
 84489237 | *Methanospaera_stadtmanae* DSM_3091
 33359542 | *Pyrococcus_furius* DSM_3638

C

11499463 | *Archaeoglobus_fulgidus* DSM_4304
 383319613 | *Methanocella_conradii* H254
 330509014 | *Methanosaeta conciliata* GP6
 40842009 | *Candidatus_Nitrososphaera_gargensis* Ga9-2
 167628645 | *Helio bacterium modesticaldum* Icel
 170220370 | *Candidatus_korarchaeum_cryptofilum* OPF8
 126465227 | *Staphylothermus_marinus* F1
 156937920 | *Ignococcus_kleinii* KIN4-I
 124027533 | *Hyperthermus_butylicus* DSM_5456
 118431181 | *Aeropyrum_pernix* K1
 307595808 | *Vulcanisaeta_distributa* DSM_14429
 305663331 | *Ignisphaera_aggregans* DSM_17230
 18311773 | *Pyrococcus_aerophilum* str._IM2
 18311700 | *Pyrococcus_aerophilum* str._IM2
 38505114 | *Fervidicoccus_fontis* Kam940
 15898287 | *Sulfolobus_solfataricus* P2
 15922974 | *Sulfolobus_tokodaii* str._7
 347523054 | *Pyrolobus_fumarii* 1A
 218884094 | *Dysulfurococcus_kamchatkensis* 1221n
 389861364 | *Thermogladius_cellulolyticus* 1633
 332296539 | *Thermodesulfobium_narugense* DSM_14796
 167627879 | *Francisella_philotragia* ATCC_25017
 569653555 | *Bacillus_clausii* KSM-K16
 76802275 | *Natronomonas_pharaonis* DSM_2160
 189485278 | uncultured_Termite_l_bacterium_Rs-D17
 134101617 | *Saccharopolyspora_erythraea* NRRL_2338
 134299482 | *Desulfovomaculum_reducens* MI-1
 433631664 | *Mycobacterium_cannetti* CIPT_140070010
 86742031 | *Frankia_sp_-Cct3*
 344996860 | *Caldicellulosiruptor_lactoaceticus* 6A
 219683006 | *Bifidobacterium_animalis* subsp._*lacticis* AD011
 347756303 | *Candidatus_Clarcobacterium_thermophilum* B
 294828260 | *Leptospira_interrogans* serovar_Lai_str._56601
 268318441 | *Rhodothermus_marinus* DSM_4252

-----GAM-----VAWQSQPF-----YID-----
 -----GTQ-----PE---FVFDAAFI-----V-----
 -----GHE-----GL---KRLLEDPDFV-----
 -----GHG-----VVTTHDPSFY-----WTD-----
 -----GTV-----ELLEEEVV-----IL-----
 -----GSV-----EKTVEDEV1-----IV-----
 -----GNP-----IAVFPAAVIL-----
 -----GSE-----SSPAQSII-----I-----
 -----GEM-----P1DTTVPFA-----
 -----GNR-----SPEIASPF1-----IT-----
 -----GTP-----QFEKPGSII-----I-----
 -----GRD-----KKFLRKREFE-----IL-----
 -----GOG-----TV---S-RDEPYY-----IA-----
 -----GQM-----RYYYEEDII-----
 -----GGE-----SPSPPNYY-----LI-----
 -----GGE-----PPYEPDPYL-----IL-----
 -----GSP-----EPREAVTII-----I-----
 -----GIG-----LYSYFEDLL-----IV-----
 -----GIG-----AVSKDPTHY-----LV-----
 -----GKG-----KITPPPQV1-----VI-----
 -----GSG-----SPTEDARLP-----MF-----
 -----GVA-----RP---IERPGLRII-----
 -----GEG-----PP---VEKSVDIV-----IV-----
 -----GEP-----PLTKPGSV-----IV-----
 -----GAG-----ELPATTKFS-----VV-----
 -----GNG-----VP---QELSEYDIL-----
 -----GNS-----VFLERKNDL-----II-----
 -----GOG-----VVSQDPDY1-----IL-----
 -----GWG-----VVRDDEDVI-----VV-----
 -----GYD-----DKTF---KDWEDEYGH1-----
 -----ELD-----ELRKPDGH1-----IV-----
 -----GFG-----YTSKIFIEKEYDII-----
 -----SKDSGKGEKMRVPVEDHRRDFSVL-----
 -----GON-----WEKIKFIP-----
 -----GCA-----SLP---KFNPQYLL1-----
 -----GQD-----KNGALFS-----
 -----GDV-----SHKDNL1-----
 -----GKN-----IEKD---DKITDEDIV-----IL-----
 -----GVP-----LGQDSV1-----
 -----GTP-----FTPDRDQIL-----
 -----GDD-----PMDDQQFL-----
 -----GTE-----QGRHRDRL-----
 -----GKS-----ENISNL1-----
 -----GVE-----EITQFF-----
 -----GED-----KDLQIY-----
 -----GLD-----K-A-KETOIL-----
 -----GRP-----P---D-EFDNVI-----
 -----GND-----KGNTSNMT-----
 -----GIE-----KGTTDMIP-----
 -----GRD-----KSPVEMIL-----
 -----GIE-----KW---N-VSSNFI-----
 -----GIE-----K---N-PIEFIL-----
 -----GIE-----KNPIEEII-----
 -----LPMLI-CSYCPTS---RDVYL-TMVKEVRG-----EESFPVSLGKAIHEAVAKAIREAKSKNFS-----
 -----PEKWKYESQGFKPPPTN-----DLVKKYSR1VWEHVLANAECSRREAASSQQPYSEEEDDIMTAAPP-----
 -----MPVWEV-SDRYCTEG---RDVYL-RHVQKVSV-----PASEENVAGRIYHQAMALEYPTAKRLLYKEGMS1CAELPG1ILMERLDDA1VANEAEQLSQTKFSPSEAEAG---QS5VSDNLRKLWSFVRS1VSS1PAYLAKYVLSDDSLINHALPF-----
 -----LGVEV-ASQVCSQG---RDVYL-RRIDGKIG---VPSKDSMVRGLVYHAYADEI1SVAKAYMVKQGVLAAGLYEHLKAKSEERFASPYMLTHQEYFREAPTEADIEEVKMLNDLK1LNGYEASQMASIKAILSKQPRIGIDALVHTA1PV-----
 -----LGVEV-ASQYCSSL---RDVYL-RVREGARS-----PKNGMFQPSLHLLARF1VATKGLYQEG---WDGYREILRQLQEMTLSELLEDEGLFLVFLASLAEADREELERKSRLLEFERGRLLIARLQEALVQPYIGEDALATALAPV-----
 -----LGVEWE-ASCIPTG---RDVWLIRHKVLRV-----PTTPQIAKIGVHVRLLVSSMLKAKKMMYMGK-----YELRDLITESESGIVERELEMNRKVYKLKPDE---EGLRGFCRRIARWEATIREGKVMVRAKYFPLNEESLQVIAFPV-----
 -----LPIWEI-AGSICPTY---RDWVIRHKVLRK1L-----FTTSQKAEGVIIH1K1VSHVFEKAALKL1YSGD-----YSDLKNALLPGVKELVUNEVKY1SELENIDA---NKIEFALQVADWEMLRLIEGRINTVKAKYV1DEEGLVSLAPV-----
 -----ARPSPVSDTDL-YTSDV-RKVLGVR1-----ESEAMASGRVHAEFMAPPKVGKVELETEDP1-----
 -----ARPRLSDVDTYVCPVGC---RDVYL-RVRLHTV-----NGLLRLRGLVHAEFALPFPKAHRVGDAYRML-----
 -----YRPSVSEVASVCPSPH---RDAYL-QHVRGQLP---PRLSPRAQYGV1VHFLEPFLPFLRVDGADSLE-----
 -----VRPTVSEVASVCPCTP---RDAYL-RVAVGATVMT-----EDESLSRLGRMVEHVEFLTPKESIPPSQLDN-----
 -----ARPSVSEVASVCPCTP---RDVYL-RDVLVHTV-----DQSVLRLGRAVHEVFLPPFRYGRDIEWLY-----
 -----LALSDF-AYVGCPCTG---RNLYL-KVGLGERG---AAVKALVEGQALHQLAKFALEDYKKVYASGH-----
 -----LSVSD1-VGCCGCSR---RDLYL-KRVLGVR1-----EPNCMKVFGAYIHEVFKRS1LAELSL1EGCA-----VKGWELVQSFNABAIREAA1A---EAEADPVGELARYLAIQVARVDEI1ASRSADSLS1AAARAPV-----
 -----LGVEV-AIKCCTK---RDLW1-RKXVGEV1-----LESSPL1YMGK1LHSMLN1APRDA51VS1RNL-----GPLESIRQELLSKRASLTSK1YRKAEMF1MTWLS1ST1L1QSL1INDPLSY1-----
 -----LS1SDV1-AYKCCTK---RDVYL-RKVLK1QGQ---EQRTPLLGGQATH1EVRNVSNEVVKLSSGY-----
 -----LTMGAE1-AYDCPCTK---RDVYY-RVNNLQKG---SERALQVLPFGQAI1FVFSKALNDVR1VSYL1GK-----DPYQ1VKESYLESFECPPQIESEYCRVYVGNL1LWFSSW1LAEAKAFYGDGSVGFPLW-----
 -----LGVEV-AYYCPCT---RDWLW-RVRLGRG-----QPTBA1MLGR1VHEAFLRAARLSDRL1LEG-----
 -----LGVEV-AYQYCPCT---RDWLW-RKRMGLAD-----ETTEPLAKGR1VHEAVTAARVCA1SRGSLRSGW-----KSTDADAVLBVGARERASVILEASAGIECDGDI1AVFAARVYKMMFWAQAWEETGAPP-----
 -----LGVEV-AGKCPTR---RDWLW-RKRMGLAD-----FAYKLKYIDRVQD---FKNINTIAGSAFH1ALSKYF1TNRYDFESSW-----EVNSVASEAAKEWR1R1D7GSRENARLVEK1YKST1L1S1GEAEAYEALYGSRRPPL-----
 -----LSPSPSLNLL1TCP---FAYKLKYIDRVQD---FKNINTIAGSAFH1ALSKYF1TNRYDFESSW-----ETWDILTYCLGRFRPQNGDNTATALKAYK1ST1L1S1GEAEAYEALYGSRRPPL-----
 -----LILTWLNDF1PCPSY1LHNHY---SNASDATY---YSSOTKGDAH1S1DK1Y1G1STKDD-----KEYRNNPVEVLSLSPNEEDK1KNNLFAMLK1Y1EPFARDLKVIS-----
 -----MM1SLQHOFQFCK---RQWAL-IH1QOWE---ENVRT1IEGQH1LKKADQF1PEIKEKRSKLT-----
 -----VHVSALNEYLYCPD---RFPYF-QRYHDEMG---TPYELVPGRSKHENA1QARDW-----
 -----LH1RQVQOLSYCK---RRCAL-LF1RQOQS---DNFPTAKG1VIMD1KAHAEK1EHKKKG1V1-----
 -----IGARALSDFRCPD---RLPH1-EHAEGYRF---DSEEMRLGRH1VHASVDTSGRERPAQ1R-----
 -----FPIIS1VLYCPD---RNPYY-RVVEGAED---SNHHLLEGK1LQEEERRDRORL1VREGYRQ-----
 -----VLLSE1EHWIYCPD---RQWAI-IH1HQFHT---DNDDTTRCHLEHQVRDVA1CHERSHGVQV-----
 -----LP1SALEMHHACP---RQAAL-IH1L0N1VFA---DN1ETMRQNVAHAVHHPSPAVPTPQGNR1IQ-----
 -----IP1ISSINTYV1C---RFPY1-EGFLGLRE---DNQY1LEGRN1QSE1QFQYINGFY-----
 -----LP1SLICDF1CPD---RRAWL-EIQGK1---ESLQMERGFDH1RAVDDANGRGTDT-----
 -----TRVMALHALAYC1---RFLY1-EVEE1R1---ANDRVY1EGGRVH1QD1QISE1R1-----
 --------MG1HSLLYCPD---RFLY1-EVEE1G1V---ADDRVY1AGRTL1HEE1PNE1DSSGR-----
 -----IPARMLNEFVYCPD---RLFYL-EWVQGEWE---DNADTLEGGSIH1KRTDSASGTL1PEIDEAFG-----EHA-----

378719041 | *Gordonia_polyisoprenivorans*_VH2
 320102061 | *Iosphaera_pallida*_ATCC_43644
 39995168 | *Geobacter_sulfurreducens*_PCA
 83591519 | *Rhodospirillum_rubrum*_ATCC_11170
 108757432 | *Mycococcus_xanthus*_DK_1622
 46255268 | *Thermus_thermophilus*_HB27
 159898754 | *Herpetosiphon_auranitacius*_DSM_785
 38505683 | *Synechocystis_sp_-PCC_6803*
 17229059 | *Nostoc_sp_-PCC_7120*
 302391601 | *Acetohalobium_arabaticum*_DSM_5501
 48477076 | *Picrophilus_torridus*_DSM_9790
 154174706 | *Campylobacter_curvus*_525-92
 111224946 | *Frankia_alini*_ACN14A
 389844446 | *Mesotoga_prime*_MesG1-Ag-4-2
 268317193 | *Rhodothermus_marinus*_DSM_4252
 28211140 | *Clostridium_tetani*_E88
 41614818 | *Nanoarchaeum_equitans*_Kin4-M
 289596695 | *Aciduliprofundum_boonei*_T469
 189774941 | *Pyrococcus_furius*_DSM_3638
 14590111 | *Pyrococcus_horikoshii*_OT3
 55376277 | *Haloarcula_marismortui*_ATCC_43049
 84489235 | *Methanospaera_stadtmanae*_DSM_3091
 29249237 | *Nitrosococcus_halophilus*_Ng_4
 38505117 | *Fervidococcus_fontis*_Kam940
 347523047 | *Pyrolobus_fumarii*_IA
 15898285 | *Sulfolobus_solfataricus*_P2
 15922977 | *Sulfolobus_tokodaii*_str_-7
 302347806 | *Acidilobus_saccharovorans*_345-15
 218884091 | *Desulfurococcus_kamchatkensis*_1221n
 389861366 | *Thermogladius_cellulolyticus*_1633
 156937923 | *Ignicoccus_hospitalis*_KIN4-I
 15896973 | *Sulfolobus_solfataricus*_P2
 18311697 | *Pyrococcus_aerophilum*_str_-IM2
 170290049 | *Candidatus_Korarchaeum_cryptofilum*_OPP8
 126465229 | *Staphylothermus_marinus*_Fl
 257125860 | *Leptothrixia_buccalis*_C-1013-b
 124027531 | *Hyperthermus_butyllicus*_DSM_5456
 18311770 | *Pyrococcus_aerophilum*_str_-IM2
 28040853 | *Spirosooma_linguae*_DSM_74
 21226659 | *Methanoscarcina_mazei*_Gol
 218961575 | *Candidatus_Claeconomons_acidaminovorans*_str_-Evry
 150401499 | *Methanococcus_aeolicus*_Nankai-3
 118497355 | *Francisella_novicida*_U12
 34557935 | *Wolinella_succinogenes*_DSM_1740
 54296141 | *Legionella_pneumophila*_str_-Paris
 167628644 | *Helio bacterium_modesticaldum*_Icel
 368854795 | *Deinococcus_gobiensis*_I-0
 148656006 | *Roseiflexus_sp_-RS_1*
 118431378 | *Aeropyrum_pernix*_K1
 307595805 | *Vulcanisaeta_distributa*_DSM_14429
 305663328 | *Ignisphaera_aggregans*_DSM_17230
 408403002 | *Candidatus_Nitrosospaera_gargensis*_Ga9-2
 330509015 | *Methanosaeta conciliata*_GF6
 11499461 | *Archaeoglobus_fulgidus*_DSM_4304
 383319611 | *Methanocella_conradii*_H2254
 33059542 | *Pyrococcus_furiosus*_DSM_3638

LPARMLNEYVYCP---RLFYF-EWDQQRWV---DSDHTEQGHFTHRADVDR-KTSLPEPES---GDPAK
 MPARMLNEYVYCP---RLFYF-EWDQQRWV---ENLDVVEGRHRTRRVDCKTDLPSQVQE---GEAER
 IPRMLNEYVYCP---RLAYL-MWVGQEF---HENEFTPGVJVRHRRVDDAGGLVPLSETQEDS---RI
 VPASPMWNAYVCP---RLAVL-EWGRGEKA---RSVLIAJLAGRAHQATESGTPALPDQMMLR---EDQSL
 VALSALQHLYCPE---RQAAL-IHVERQR---EDVNTASGRLLHERVLDLPGHDARLGRV
 LPIPLKVNTPVVCPE---RRPYL-EVLGEMI---ANHHLIEGHYLHERAYTVEGPE---
 IPISMLNAYLACP---RPPGY-EYLQAEML---INEFVAQDGQILHSSVDAQRTWTIAEGVO
 LPLAYLNAYEYCH---RRPYL-EVLGEMI---INEHIIRGNHILHRNINEACTSVGDTRI
 IPIALNQAYCS---HRCWR-MECAGEFI---DNQYILECTLMDRVTSEQRRETWO
 VPLSAINAYNCP---YRVYL-EVLGEMI---DNTHTIEGILKHDRASHERRYSNRIQ
 RYGLDVSIVNICM---RRAWLHSLEIPTV---DSYVSDGAYLADIN
 ITGTLVNYFTMKC---REAWLYAHHTHADQ---DDENVLMGKALADIKEKD
 VGGVH1KYLLHACP---RQLWLWYMRGYPKG---SSDLVSPGEVDETTFTTR
 IIGNAINYLYHCK---TQLWFLWLHNISMES---TSESVQLGKLQEEYQSQE
 LPLAYLNAYEYCH---RKAWLAVHGLWMEQ---ESEAVALGRLRDETTYYAELO
 VNGTLVNYVYFHKC---RQCWLHGNRNLINED---NSQDVKIGKAIEVKEKEKG
 IGGAYIYYYFVCK---RKLWFWYANKTMIEH---ESDLVQIGKYLEIFKEKKY
 FTGSLVHEVYVCK---RAAWY-SYRGVRL---ENEYIILGKMLHESKYG
 VRGIEINYLYVCP---TKLWFPTTHGTEMQ---ESEWVDLGFKHLERYEN
 VTGLMVQYQYFTCK---RELWFSRGINPDF---ENEDMIIGRLHAEAYEDS
 VTGVMQYQYEVCK---RELWFSRHLLEIDR---DNTAVRGTVODESAYDDR
 ITGVMQYQYKLCK---RELWFWYMNQNLMNY---NNNDIEGLRHENSYKR
 LHGLHFQHVRCLC---RRAWMHHLG1SFAO---WNRHVQIGAHOHTSYARD
 ISLSSLRNLPCB---RMWF-KLNSDIE---PPTPSMILSERHDLKRIAESLMLPQPM
 FGGVVKEFVWCDB---VAAWL-SVNGLVSP---PVTRMLLEGVEGHRSVLEVIHGLGFD
 VSVDLKLKDYSLCK---VIPWI-RKMGKNGKEPI---TNSQRAKVNVLKEIVRDFEP
 VITVSDVKQHFCK---AIPWI-NVYMGRE---PSTSMEEGKKISYEVVKSLNLT
 VITGTAVKDYSYCP---ALPWL-KVRLGE---PETWGRAGRSLRASDVAVYKLGLERPW
 VSPSLVKEYYCP---VIPWI-QANLVMAB---PMITSMLGESEAMKPTG
 LSPSMVKEMVYCP---VIAWL-KARHILQE---PATDSMLAGASAGGGGLYARA
 ---MVNAYALCP---AFAYP-KYHPE---PQTSMLGEKLDVRELVEKRPVKEV
 ENTYVTDLVRCP---RRVRY-ESEYKELAISQ-VIAPSAIQLDGIHLGLESV
 PTPWDVLELCP---RYLWL-SKRHG---PMPTPNAAGRGAEAGLRLFLAKALGAE
 LSVRVEEYFFCP---LLPFY-KNLYGIDT---QEGFWAYLGRKAEEAEEERRKRF
 PTFRMVEEYIFCP---LPVVF-KVGLGIKK---PPSLSWSNLRQVEEELSRVYEBHYNV
 ISIFLVEFLDVFICP---RILYV-ENVLKLMGEK---KEDSNKKEERVVKGKNSLNKWNIDRKLK
 IP1VLUKEYYCP---VAAWH-KLWLWAE---PTFSMEAGLSEGETRWRLLERLEVEPARWR
 INPSLIRQYLCP---AAAYY-IMTGAAFP---PTERNRACKEVORBAEAVAKALGAEK
 LTPSHIETYLFCP---RFTYF-EVLAIPQYE---EKNWKVMRGMLNIDERLENRKDYLRKRLGP
 IRISDVELEYLFCPS---RFIYV-MYCLDIPQHE---EKRFRVFKGRHEVTRKLNTDVRVKKLNIRK
 ITPSVEMYLFCP---RFVYY-MNLLKEIQLHE---HRTLWNGKGRDHLKVMQKEYLRLKAGAIDK
 ITTSDVVEYMYCP---RFVYF-ERVLGIPQHE---EKREKVIGNITHRELEKINKGYIRSKINAKS
 IPVNLIRQWQFCP---RIVYY-QELLALPK---NKPPLWVAQEEPHKVKQLEKKRSPSYCLE
 LPINLIRQYLCP---RVVYF-LEVNLNPK---ASPFWVKEGSNHRQTELLEKERTLTLARFH
 LPVNMIRQYFCP---RIVYY-HMVVGKLV---AYPGWQVGEYEOHKQULLKERTRTLSKFKP
 LRVTDVKQVYCP---RIPYF-TVOPOLKP---RPTAKMDFGDKHARLLEKERTLAAVYEL
 IPTPLRQHVCYCP---RVTFF-ERCPVYR---RETIIMTHGREKHQTELVRERRRTLRSYELT
 FVTFLDRQHVCYCP---VVVYY-QVCLPDIR---PVTKLMEAGQEHRAESGREERSLRTYGLK
 RPVSMLKEVAYACP---RVAYY-MEVLRPSY---RPTEPNMLSRSEIYSDVHGRILRSSGFRV
 IPISMKEVYTCP---RQAYL-KIMMMEP---PTESMKYAKQTONTETITTKTKELG1QCEV
 IPIVLUKEYYCP---RIAFY-KYFNWEP---PTESMKFARYSKIDILKIVRGY1LDGDI
 ISVTDIKHYDFYCP---KITYF-DKVMHADA---VLSQQEDAKTHKKEKEKKDRKRTMFYEQ
 ---MKQYLYCP---KITYF-DHVLH1PK---PDQKLQGTGIMHDSITAKEKRRGAIYFDP
 MDVDSIDVQYLYCP---RKLYF-LKIAGIRI---SKPKNMEEGKAQEARGLKRSFARKLKGEL
 LQVSDITQYFYCP---RKVYF-MKTMG1KV---KPRPKMDLGKEEHEREHRRVKERKTIYGFSE
 ---MFHMYIVVVYDV-----GVERVNKVKKFLRMH-LNWNQNSVFEV-EGTLEAEPER--IKEGL---KK-IIDE---NS---DSVIIYKLR---SMP---PR-ETL---

Continued

11499463 | *Archaeoglobus_fulgidus*_DSM_4304
 383319613 | *Methanocella_conradii*_H2254
 330509014 | *Methanosaeta conciliata*_GF6
 408402999 | *Candidatus_Nitrosospaera_gargensis*_Ga9-2
 167628645 | *Helio bacterium_modesticaldum*_Icel
 170290047 | *Candidatus_Korarchaeum_cryptofilum*_OPP8
 126465227 | *Staphylothermus_marinus*_Fl
 156937920 | *Ignicoccus_hospitalis*_KIN4-I
 124027533 | *Hyperthermus_butyllicus*_DSM_5456
 118431378 | *Aeropyrum_pernix*_K1
 307595808 | *Vulcanisaeta_distributa*_DSM_14429
 305663331 | *Ignisphaera_aggregans*_DSM_17230
 408403002 | *Candidatus_Nitrosospaera_gargensis*_Ga9-2
 330509015 | *Methanosaeta conciliata*_GF6
 11499461 | *Archaeoglobus_fulgidus*_DSM_4304
 383319611 | *Methanocella_conradii*_H2254
 33059542 | *Pyrococcus_furiosus*_DSM_3638

-LVEHMRDGS---LGCGSLIS-VDCYDYL---NVVFDLKVGFEFNE---EMRLYTTGAYALVLE---SIHEIPVDPVGC1---ISITFNGER---MRVSKDLHYI
 -MIEHKISGNI---LGCGSILS-VDCFDYLH---NIIFDVKTATRKGENS---FVYR LATGAYALVFE---SVYEVPDIGCT---VVLNFKDDR---LLIERDPIH
 -VEHKLQDCKY---LGGSRLNLS-ADAINNMG---MLICLDTQKGKRD---FHRLSATGAYALVLE---SLYVEPVINIGCT---VVLSPFKHNPT-PHIERDPFII
 -IVEQKLDGSA---LGSLSTHS-LDGFASFD---SIIIDLKTGBEKE---FPHKLSTTCYALVY---SIFYPEVINVGCV---VYVNFNSNNSPV-PIIKRPHLI
 -VQEQLKLDGSF---LGLSAHLHS-ADAYNIGQ---PMIVFVKEFGEPRD---FPHRLATGAYALVME---AVYEHVPDVGLC---VCVTFKGD---VIIRDIHIL
 -ATELAIDGSL---LGSLQYLR-ADALWFG---GILFDVTKDRKD---WHR1QLAGAYALVLE---SFFPMPDVIGCV---VIVSEVPBG---LVRVFRDPFII
 -SLEMPVGDGL---LGSSLMLR-VDASLWPG---GLIYEVTKGYREK---WHLKLQVAGAYALB---SIYERPVDIGVV---VVVNRAVEG---IRVNDRIPVV
 -EVPEVPIPGHS---VGLGSLDT---FPIPVVYVYNGLE---RKKLAVTGYALALE---SWIGHPVEVGVIV---VSVDT---KKTYWNVVR
 -VGLDSALT---VGLGSLDT---FPIPVVTAAGPGTYGS---KKEKLQLAGAYALB---AWTGPVVDYGVVV---VVYTRRSDE---PRPEWVV
 -LRVPEPVGCP---VGLGSLDT---FPIPVVTAAGPGTYGS---FPIPVVTAAGPGTYGS---RKELAIAQYALVLE---RKELAIAQYALVLE---AWTGPVVDYGVVV---VGLRVDNGD---LKI1WRLV
 -AEPVNPVGD---VGLGSLDT---FPIPVVTAAGPGTYGS---FPIPVVTAAGPGTYGS---RKELAIAQYALVLE---RKELAIAQYALVLE---AWTGPVVDYGVVV---VGLRVDNGD---LKI1WRLV
 -SVEPAIPEGAP---IGLSLIVK-PDILVG---FPIVVELUTSSQFTEELTS---RKEDIAVTAQYALB---ANWIGHPVDIGV---LHISINGK---PRLTWV
 -ATOVAVDGS---LGMSYVW-ADGVAL---FMPIDITLGSLGSNLE---RKELAIAQYALVLE---AUSNGNPVDPIDIGT---IGISINS---VRFQWRV
 -LAAYVILDGR---LGTLTIVR-ADALYY---NNUVVKVGSYSE---RHALALQAGAYALB---ADEEVPVDAGLL---ISVAFNNGG---VKLRAYPV
 -IKTEFVLDGP---VGLGSLNL---GVIIEYKLGNGNLPPLTE---RHSVGLAGAYALB---SSLDIPIYGA---VVISELSSR---PSIKV
 -VGLGSLNL---GVIIEYKLGNGNLPPLTE---RHSVGLAGAYALB---SSLDIPIYGA---VVISELSSR---PSIKV
 -IESEFKVGDSP---LGGSRLS-VD1AQL---SLVNMKVMGSVQE---FPHKLALQAGAYALB---SALELPIDFGAL---IIVNGNINTS-SPEFKV
 -AT-EVTRVDSGA---IGLSLIVK-PDILVG---FPIVVELUTSSQFTEELTS---FPHKLALQAGAYALB---SPEFKV
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 -ATOVAVDGS---LGMSYVW-ADGVAL---GAVIEFPGPQA---NVDAAQYALB---ADWGPVVDYGVII---VOITVNST---VEYKATA
 -LAAYVILDGR---LGTLTIVR-ADALYY---NNUVVKVGSYSE---RHALALQAGAYALB---ADEEVPVDAGLL---ISVAFNNGG---VKLRAYPV
 -IPEVPIPGHS---VGLGSLDT---FPIPVVYVYNGLE---RKKLAVTGYALALE---SWIGHPVEVGVIV---VSVDT---KKTYWNVVR
 -VGLDSALT---VGLGSLDT---FPIPVVTAAGPGTYGS---KKEKLQLAGAYALB---AWTGPVVDYGVVV---VVYTRRSDE---PRPEWVV
 -LRVPEPVGCP---VGLGSLDT---FPIPVVTAAGPGTYGS---FPIPVVTAAGPGTYGS---RKELAIAQYALVLE---RKELAIAQYALVLE---AWTGPVVDYGVVV---VGLRVDNGD---LKI1WRLV
 -SVEPAIPEGAP---IGLSLIVK-PDILVG---FPIVVELUTSSQFTEELTS---RKEDIAVTAQYALB---ANWIGHPVDIGV---LHISINGK---PRLTWV
 -ATOVAVDGS---LGMSYVW-ADGVAL---FMPIDITLGSLGSNLE---RKELAIAQYALVLE---RKELAIAQYALVLE---AWTGPVVDYGVVV---VGLRVDNGD---LKI1WRLV
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 -VGLGSLNL---GVIIEYKLGNGNLPPLTE---RHSVGLAGAYALB---SSLDIPIYGA---VVISELSSR---PSIKV
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 -VSEPVYFPG---IGLSLIVK-PDILVG---FPIVVELUTSSQFTEELTS---FPHKLALQAGAYALB---SPEFKV
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 -VSEPVYFPG---IGLSLIVK-PDILVG---FPIVVELUTSSQFTEELTS---FPH

294828460 | *Lepotispora_interrogans_serovar_Lai*_str_-56601
 628318441 | *Rhodothermus_marinus*_DSM_4252
 378719041 | *Gordonia_polyisoprenivorans*_VH2
 312010261 | *Iscospaera_pallida*_ATCC_43644
 39995168 | *Geobacter_sulfurreducens*_PCA
 83951519 | *Rhodospirillum_ruberum*_ATCC_11170
 108757432 | *Myxococcus_xanthus*_DR_1622
 46252568 | *thermus_thermophilic*_HB27
 159898754 | *Herpetosiphon_aurantiacus*_DSM_785
 38505683 | *Synechocystis_sp*-PCC_6803
 122720959 | *Nostoc_sp*-PCP_7120
 30329160 | *Acetobalochlor_arabaticum*_DSM_5501
 48477076 | *Picrophilus_torridus*_DSM_9790
 154174706 | *Campylobacter_circvus*_525-92
 111224946 | *Frankia_alni* ACN14a
 389844446 | *Mesotoga_prima* MesG1-Ag-4-2
 628317193 | *Rhodothermus_marinus*_DSM_4252
 28211140 | *Clostridium_tetani*_E88
 41614818 | *Nanoarchaeum_equitans*Kin4-M
 289596695 | *Aciduliprofundum_boonei*T469
 18977491 | *Pyrococcus_furius*_DSM_3638
 14590111 | *Pyrococcus_hiroshii*_OT3
 55376277 | *Haloarcula_marismortui*_ATCC_43049
 484492305 | *Methanospaera_stadtmanae*_DSM_3091
 292492937 | *Neuroscoccuss_halophilus*_NC_4
 385805117 | *Fervidicoccus_fontis*_Kam940
 347523047 | *Pyrobolus_fumarii*_1A
 159828285 | *Sulfolobus_solfataricus*_P2
 15922977 | *Sulfolobus_tokodaii*_str_-7
 30234780 | *Acidilobus_saccharovorans*_345-15
 218884091 | *Desulfovibococcus_kamchatkensis*_122ln
 389861366 | *Thermogladius_cellulolyticus*_1633
 156937923 | *Ignicoccus_hospitalis*KIN4-I
 15896973 | *Sulfolobus_solfataricus*_P2
 31831697 | *Pyrobaculum_aerophilum*_str_-IM2
 170290049 | *Candidatus_Korarchaeum_cryptofillum*_OPF8
 126465229 | *Staphylothermus_marinus*_F1
 257125861 | *Lepotrichia_buccalis*_C1013-1B
 124027531 | *Hyperthermus_butylicus*_DSM_5456
 18311770 | *Pyrobaculum_aerophilum*_str_-IM2
 28404053 | *Spirosoma_lingulale*_DSM_74
 122266559 | *Methanoscarcina_mazei*_Gol
 18961575 | *Candidatus_Cloacamonas_acidaminovorans*_str_-Evry
 150401499 | *Methanococcus_aeolicus*_Nankai-3
 1849735 | *Francisella_novicida*_U12
 34557935 | *Wolinella_succinogenes*_DSM_1740
 54296141 | *Legionella_pneumophila*_str_-Paris
 167628644 | *Helio bacterium_modesticaldum*_Ice1
 386854795 | *Deinococcus_gobiensis*_I-0
 184656009 | *Roseiflexus_sp*-RS-1
 118431378 | *Aeropyrum_pernix*_K1
 307595805 | *Vulcanisaeta_distributa*_DSM_14429
 305663328 | *Ignisphaera_aggregans*_DSM_17230
 48403000 | *Candidatus_Nitrosophaera_gargensis*_Ga9-2
 330509015 | *Methanosaeta_concilius*_GP6
 11494961 | *Archaeoglobus_fulgidus*_DSM_4304
 38331961 | *Methanocella_conradii*_H2254
 33359542 | *Pyrococcus_furius*_DSM_3638
 -IESPHYTHSEK--LEVSFGK--VDR1QKRQGD--WIP1YEHKKGRAR1NGPNE-AWE--SDQQCVTVYVALLLE--EATGRINISEKGK--RYHHSK--DLV--KIEI
 FVRSVLMSPA--LGLLJAR--VPPDFTESNGL--VYPVYDKKGSPESD--AWL--TDVKVQLCAQGLL--ENGRYCERGGV--FVTGTR--QRV--EVF
 RVTSLRIESHA--LGLLVAG--LDRVEPDGGDGT--WPS--ADVVYOLQAQALLR--DAGVEYDVRGL--YYAEVH--KRV--VLP1
 HARSVSLADH--HGLLJK--LDLLECADCAG--MIPDVYDKKKPRLADQPVQVWE--PERVOMAVQALVQL--ANCYCRGVRG--YYHQTR--QRV--DVP1
 HARSVSLSSER--LGITAK--IDLVEGBGAY--VSPDVYKRGRPHVAGG--AYE--PERVOLCQAGLLLR--EHCPASDGAL--YFVASR--ERV--EVPA
 KTRRLSLSSE--LGLTAE--LDLLEVEGM--VPIVIEKVGRSPVSEBG--AYL--FERAQVCAQALLR--EAGTYTCLLEGAL--WFAESR--TDL
 -ERIAVLLGSQR--LVRSGR--ADMVEYQFRDAKAGWR--BPFVEVKRGRKAGE--ADRVQVLCQACNLIC--EHGVQVEPEGAL--YGVHR--KRA--AVRP
 -ESGLWNSWSDR--LGLLGV--VDRLEWRKAGE--ACPVYBLGKAGEBAYL--SDAVQLAQAAQCLCR--ESRGKARAFGV--YYHKSH--TTR--EVF
 -ERSVYWSWSDR--WRITGI--VDLLESDRGE--FCPVYEKGLQAR--NDHVQLCAQACLLE--EMPGFVCNQGFI--FSWANR--RRE--QVAF
 -HRQONWWSNDR--LWKVKG1--IDAVEESDGL--LIPVYEKKGKMGKHL--NDHPQVLCQAAQCLCE--EKTGK5ITYGEI--FYYGNR--RRQ--PVE
 -VRA1WLKSND--YQLIGK--SDLIESADQGQ--FIPYVEYKGRKGEND--NDHQLCQACLLE--EMTGQTQVYGYI--YYAHSH--QRQ--LVE
 -30239160 | *Acetobalochlor_arabaticum*_DSM_5501
 -TTQVFVKSSEQ--YRLVKG--IDVVEEKEGE--VPPVYKGRSWE--NDHQLCQACLLE--EQLGVTKITGYL--WFYFSSR--DRE--EV
 -RKGLP1ISTRG--SR--MDNVQ1HKDF--IMVYHEKGRYMK--ADKMLHLYHMIIK--LKENLDI-KGKL--HFLGSK--KVV--DIEL
 -LQDFAFSNLK--FDLKLSKQRGH--YCTITEKKSLSNP--ISGK--TGLN1LVEKGRV--HPLGSK--KVI--AVDD
 -RRDVDLGKA--IDWVTTG--AVVHEKKSRSAP--AHEAOVHYRCYLLE--RRGINRSVGV--HYFLIR--RTV--TVAM
 -DKDVIDEIS--LDLILRNT--REIVEKYKGRVRE--GHIYQVQVYLYL--HKLGLHEFTPL--TYPRAK--KMI--MLSL
 -HVNVEADGPAG--LRLAGR--IDWAEHLRD--GVLHETKRSPSGLE--AHWRLQFLWYLWKLRLGVTRPDQCPF-EQL--NPFPR--RTE--TVL
 -QTE1S1DNIK--IDKUTK--DYLTEVKKSDSDE--AAKWLQLLYLWKL--DGE--EKGKL--EP1EKN--KSKST1I1EL
 -ERKV1IN1ES--DPV1KYYKDF--VLPV1EKKSDKFK--A1WLQVLYLWYL--RL--GLN-VKGKL--VIPENY--KEE--MVE
 -IKNLFGEGV--IDFTRKEGE--VWIYEVKRSSKHLN--AARMQLLFYLWNLK--DR--GVE-VKGKI--AVPKE--YTE--MVEL
 -EKEV1LIGPIR--IDF1SRGDV--IEVHEVKLGRMR--AHEMQALYFYLYL--KH-GIK-AKAIL--HYPKLN--EKR--EVTL
 -WKEV1LIEDK--IDALKTKG--LKV1EIKKSSKLEE--PAWQKLQYLYL--KA-GID-AIGV--SYPKEG--KRE--IIBL
 -RNVS1DGITA--IDVLDL--GRVMVEPKPSSLLK--FAKLQQLYLYLW--QVGVQE--REGVL--AHPTR--KRE--TVAL
 -KKB1RVDNLV--FDFTQTKDK--LTIVFVEKKSSKTI--GALQYLYLYLW--NA-DIE-AKGML--VFPKEK--KRE--SIEL
 -RSTQGLFCFLA--PDRIDWKH--NIVYESHTQGAVE--AVSDQTAFYALL--IATGK--WRATV--HVLSTR--KRR--EVL
 -QFNKKLCHCP--IPVCGS--PDIISGEER--K1IIVEYKHGFSRERGDFQ--DFAQRLTYGWLVQ--EVEGRWKLVL--GREVWQYQ
 -RLLPPLASLR--WP1VIA--PDAYSPTSR--GALLEVVKVARIDS--LYRVAQQLYGVVAR--EN-GLRVERVL--VDAHRS--RVL--WPE
 -YYEVHLRDKD--SGLNGI--DVLVTE--DSV1E1KAPYRFYFN--HFRQLQYLAYALAE--RN-GFVRNTRVL--LMMKE--RKI--NIE
 -PKVYEVCLSY--NGLGCL--IDILAGDRK--YTIVVCFKGRNF--HFRQLQYLAYALV--KRLGI--VERAIL--VMEGK--KVL--EVE
 -LEEPVFLDRD--LGLAAGV--DLVLSGGR--LRLVLEVKFAPRGERL--PFAONQLKFYALVAN--RLLG-BVEREALY--YHGG--DLL--ALVE
 -RG0V1R1STS--GTV1I--IDEV1EKEKG--K11IEKKAFSHNFSR--YVSQAITSYVIARE--KLPGVKVRVIA--KGS--KIV--ELEV
 -GGYSAL--DVEBREBEGGG--VWVVERAKPARDQ--RYMAQLVQAGVIAL--LKGVRVRLRVL--EMCGVR--R--ELEF
 -RYOVPFLRK--LG-SGR--ADAVALCCEGR--LEVVEAKLGGGFKR--QQLAQAAFYCLLAE--EQFGLPCBALWL--CSPE--GCE--RRPP
 -TLEIREVNGKV--Y1KIKGR--ADAI1IRNDNGK--S1VIE1TTSRSKDKGLP--IHHKMQLQ1YLWLF--AEGKGL--VYITPD--RIBA
 -PKPV1YIDAG--WA-HGV--DVIILKVRVA--ASPF1E1KGLP--EHWKQLYAYAELV--AAGCAVYTGYI--TGE--K1Y--RLQI
 -GKEVELRSDR--LGVGGV--DVF1VGRGRE--IMPLVEFKSGRLR--WPKY5ISLYYAMMLE--DSIGRPFVGSVG--LVTGRM--RPI--EV
 -GKQVFLFESKK--YSLAG1--DYV1YVYKRTG--YTP1E1KYSRKLK--WPKY5ITLYALV--ETYYKPKVNAIL--IMPGRK--TTI--INI
 -REFNKELSSK--YHFGYK--LDD1L1EDKT--IIPVYVYNNAKYTAIREEN--RKYV1AMPMLLIE--ETK1QSKYQGYI--LNLK--SLK--KSY
 -LLWEHPFTS1R--LGLLAGR--VDLAAVLSSDT--LWWAEKLSPGKRRSL--HIRVQAAYGAQM--ETLGYPLRMALI--YEAEEN--RIV--EV
 -EHSPLREG--GGLGVG--VDAV1WLRLR--PAP1LEV1KPLAKP1K1PV--HIIKQAAYAAVQO--HTYHGKVASAYI--YYTETG--EVK--
 -KHADQYLTNTEL--IRGV--DVEVLFGFDGT--MAP1DLYKFKAEDKFDYD--HIIKQAAYAAVQO--HTYHGKVASAYI--YYTETG--EVK--
 -ESDVFIASKK--NH1IKGR--DVEVL1PFLEDGT--AAP1LEYKFKAEDKFDYD--TXYKPLQLVQALLR--ENYFOV1NRYAL--CFTRSN--
 -LTDVYLSKSE--LKLVKG--DVEVL1PLADGS--AAP1DLYKFKAEDKFDYD--TXYKPLQLVQALLR--ENYFOV1NRYAL--CFTRSN--
 -YIDVYLSKSE--YR1VIGV--DVE1EVLERDG--YSP1DLYKFKAEDKFDYD--TXYKPLQLVQALLR--ENYFOV1NRYAL--CFTRSN--
 -RPNFLSIKSO--YKLHGI--DWDV1IETDNT--VYVBEK7NDNPNSL--THKVSQVLYLW--EAFVAKY1ALVQ--DNEFVWVKGVY--VV
 -LSTNVMFLPHEH--FPNFGI--CD1L1SSES--VYV2E1KLHGEK1TK--GQKMQLTAYGMLSE--LIFEKPCTRGYI--ITGEKA--KAK--
 -LRQNTMLEHS--YPFYGI--CDG1IEDELE--LIP1EKLPSFASK1SL--NHKYQ1YLATYAMLE--ELIQKPV1RQGFI--KLVNPLADGARPEKSE1ITL
 -RKFHTYIHSAR--LGLLEGK--LDLHLVWHRGE--IIPFVEKTSRTRPGL--GHLKQ1CAYALV--AELGLGCPHGVY--HSSRTR--QTH--MVLF
 -RRYEVRLTNHA--LGLVGT--LDL1IVDGR--AFFPVEKTSRTRPGL--GHLKQ1CAYALV--AELGLGCPHGVY--HSSRTR--QTH--MVLF
 -RFFDVLFLRSET--LGLRKG--DVLVIA1PDRKRQATE--IIVVYESTKZEGKAF--HFLVQLTAYALM1--EANG1PVRGFI--YHMPER--
 -EWAQFLRSKR--LGLQGV--ADGVVVEGSLG--IIVVAKLRSVNRSLWHLTRGR--HFLVQLTAYALM1--EANG1PVRGFI--YHMPER--
 -KLEAVPKSTS--LGLVGR--TDAVINGNR--AVIETKLNTGSKKKLYTKYN--HILQATAYA1ALV--ETMKV7DVKIII--INQEKQ--TTI--
 -YMEYVPKRS--LGLYGM--IDIV1DNGK--VSIAE1KLETSRSLARWLRKSY--HHL1QLTAYTATE--ETFHFK1DRTFI--VTCQIG--EII--EINI
 -KLFN1HIMYES--LRLVEGV--VDCMV1NGNE--RIPVYDKRPMKSGKGEAD--DHRFLQYLAYALLE--EQYST1VRGFI--YLYPDE--
 -KMFREMESET--LGLVGR--LDL1VTRNTE--FIPVYDKRQGSHKGRGSIHL--NHKYQ1YLATYAMLE--ESKFT1VRGFI--HYSDRQ--
 -LTSVYLESKR--YLGKVG--LDALV1VDDG--AFFPVLKTRPQSITY--AWMKQLTAYALV--ENYAVEVKRQYGL--FLLNGQ--KFB--EV
 -VIHDLAIENKT--IGLYGM--VDTVLIMKNGE--VLPVUDVYKSVSVK1--AYFGETVRRGII--YLPQAHQ--KQI--QVEI

Continued

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11499463|Archaeoglobus_fulgidus_DSM_4304
383319613|Methanocella_conradii_HZ254
330509014|Methanosaetae_candida_group_GP6
408402999|Candidatus_Nitrosospaera_gargensis_Ga9-2
167628645|Helicobacterium_modesticaldum_Icel
170290047|Candidatus_Korarchaeum_cryptofilum_OPF8
126465227|Staphylothermus_marinus_F1
156937920|Coccoides_hospitalis_KIN4-I
124027533|Hyperthermus_butyllicus_DSM_5456
118431381|Aeropyrum_pernix_K1
307595809|Vulcanisaeta_distributa_DSM_14429
305663331|Ignisphaera_aggregans_DSM_17230
18311773|Pyrococcus_aerophilum_str_-IM2
18311700|Pyrococcus_aerophilum_str_-IM2
385805114|Fervidicoccus_fontis_Kam940
15898287|Sulfolobus_solfataricus_P2
15922974|Sulfolobus_tokodaii_str_-7
347523054|Pyrolobus_fumarii_LA
18884098|Desulfurococcus_kamchatkensis_122ln
389861364|Thermogladius_cellulosilyticus_1633
332296539|Thermosulfobium_narugense_DSM_14796
167627879|Francisella_philomiragiaATCC_25017
56965355|Bacillus_clausii_SKM-K16
76802275|Natronomonas_pharaonis_DSM_2160
134101617|uncultured_Termite_group_l_bacterium_Rs-D17
134299482|Saccharopolyspora_erythraea_NRRL_2338
433631664|Mycobacterium_cannetti_CIPT_140070010
867419861|Frankia_sp._CcI3
344969861|Candidatuscellulosiruptor_lactoaceticus_6A
-DANLRSWWIIEERKRAIEVNPADPGVAKS-----CSNSFLNFAP--CL-
-NDDLRSSWWIIEERDKKSEMVVEEKPGVGV-----CERKCMYANI---CKE-
-SDEVRQWFIEERDSKQDYLIFYERDPGRCPN-----PGSCRYREF--CF-
-DETLRREFEPLARDLRRMMYIDRDRPGLPRK-----CYAKCQYLRH--CGVGPEQIEVSA
-DDELQRWPVFEARDEKARMVYEEDIPGLAKD-----CPQSCPFFANK--CS-
-TDDLRSLRFLERRDQLQMMLIREDEPKRAER-----CPKYCLFLFSDM--CGVVT-----
-SDDDLRSRFLERKRDQEIQMLLLKGKEPRIPDKE-----CPKRCLFLFRNI--CLGE-
-HEGLRRLRFVLDERAVILGEKEDGPAAPEE-----CPKAQCPFAEV--CKR-
-DDALRSLRFLERDEVAMVNKPVGPGAE-----GCPSPCPFFRGVECPGPG-
-DDHLRTRFLERDVARIAIIEYEDPGLPEA-----CPFGCVFREV--CGV-
-DDT'LRRAFLMDRNVAIRIEHCDDPGPASS-----CPRTCPFVGV--CHG-
-DDSLRRLRFIARDDVARIAIIEYEDPGLPSQ-----CLKMCPFPREV--CGV-
-GDSARTKFELRDEAIDVIAGRDGPGVAPD-----CPKTCPFVH1--CHADRSR-
-GEGLRREFLEERNRNLMLVASGPGLSPK-----CDDKCPCLWRH--CHGSSG-
-SSLYKKFKFLNRNDRDEAIEVLMRATPPRSQN-----CYQVCPFFKA--CNGGRS-
-VSPDMRKREFLARDEVIDEYQSKDGPMPV-----CSPSCPFLNY--CNNNKR-
-SPDMRRELIDKRBEDIEMVLNRDGPVKPV-----CPETCPFYEV--CWK-
-GNEERAEPITARDEAIIIVUSV3DVPGRPPR-----CPDQCPMPRLW--CAP-
-NSYLRRYPIEERDEIIDLMLNREPRPDSA-----CNPSCPFVKV--CNK-
-SNELRNPFLERDEIIDLMLNREPRPKDN-----CPQTCPFYRV--CHO-
-SSEMKEKTMILNENKLLENKIFPLPNGLVH-----PYACK-YCSVSNF--CYFVGNCIS-
-PTSALKLEFEHQITQYKQNPMDNSFRQNI-----EKCK-FCIYANL--CDKTDL-
-TDSLAEKVWEMAKEMUHVYDNYTTPVKVTK-----FPCK-NCSLQH4--CLPKLMNKRSVK
-TEDHRQAVTKIVDQIQSMSATNIPPLTDNP-----DKCD-ACSAEY--CMACTAKLPE
-NKSRLREETFALAQEFHSFLVDSPPEYS-----KCKY-NCSFKEL--CLPEIIPDKQRSV
-SATAEEAERLHLFRARHVASQGQPPLRHK-----SPKCP-RCALLPI--OMDIENELSER
-DEDLRLRMENAVGLARIALNSEIIPPPLAD-----NRCE-GCALVNN--CLPEFVEVGKIKEN
-GNEELRAAVHTAASGVRVARADQSLPRPAND-----ARCP-ACSLNET--CLPRLVDDRRHQ
-TDTLALTRITAERMRDILTTAALPEAVAD-----RRCR-RCSLHHB--CLPELANAAAAGT
-EENKITKLAEIILDNMNRKIAEGLKLPAWEFS-----NKCP-KCSELLN--CFFYEIKQTNIF

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219683006 | *Bifidobacterium_animalis_subsp_-lactis_Ad011*
 347756303 | *Candidatus Chloracidobacterium_thermophilum_B*
 294828260 | *Leptospira_interrogans_serovar_Lai_str_-56601*
 268318441 | *Rhothermus_marinus_DSM_4252*
 378719041 | *Gordonia_polyisoprenivorans_VH2*
 320102061 | *Isospaera_pallida_ATCC_43644*
 39995168 | *Geobacter_sulfurireducens_PCA*
 83591519 | *Rhodospirillum_rubrum_ATCC_11170*
 108757432 | *Mycococcus_xanthus_DK_1622*
 46255268 | *Thermus_thermophilus_HB27*
 159898754 | *Herpetosiphon_auranitacus_DSM_785*
 38505683 | *Synechocystis_sp_-PCC_6803*
 17229059 | *Nostoc_sp_-PCC_7120*
 302391601 | *Acetobacterium_arabaticum_DSM_5501*
 48477046 | *Picrophilus_torridus_DSM_9790*
 154174706 | *Campylobacter_curvus_525-92*
 111224946 | *Frankia_alni_ACN14a*
 389844446 | *Mesotoga_prime_MesG1-Ag-4-2*
 268317193 | *Rhodothermus_marinus_DSM_4252*
 28211140 | *Clostridium_tetani_E88*
 41614818 | *Nanoparchaeum_equitans_Kin4-M*
 289596695 | *Acidilipofundum_boonei_T469*
 18977491 | *Pyrococcus_furius_DSM_3638*
 14590111 | *Pyrococcus_horikoshii_O73*
 55376277 | *Halocarcula_marismortui_ATCC_43049*
 84489235 | *Methanospaera_stadtmanae_DSM_3091*
 292492937 | *Nitrosococcus_philiphius_Nc_4*
 385805117 | *Fervidococcus_fontis_Kam940*
 347523047 | *Pyrolobus_fumarii_1A*
 15898285 | *Sulfolobus_solfataricus_P2*
 15922977 | *Sulfolobus_tokodaii_str_-7*
 302347806 | *Acidilobus_saccharovorans_345-15*
 218884091 | *Desulfurococcus_kanchatkensis_l221n*
 389861366 | *Thermogladius_cellulolyticus_1633*
 156937923 | *Ignicoccus_hospitalis_KIN4-I*
 15896973 | *Sulfolobus_solfataricus_P2*
 18311697 | *Pyrobaculum_aerophilum_str_-IM2*
 170290049 | *Candidatus_Korarchaeum_cryptofilum_OPF8*
 126465229 | *Staphylococcus_marinus_F1*
 257125860 | *Leptothrix_buccalis_C-1013-b*
 124027531 | *Hyperthermus_butylicus_DSM_5456*
 18311770 | *Pyrobaculum_aerophilum_str_-IM2*
 284040853 | *Spirosooma_linguae_DSM_74*
 212266593 | *Methanoscarcina_mazei_Gol*
 218961575 | *Candidatus_Cloacamonas_acidaminovorans_str_-Evry*
 150401499 | *Methanococcus_aeolicus_Nankai-3*
 118497355 | *Francisella_novicida_U112*
 34557935 | *Wolinella_succinogenes_DSM_1740*
 54296141 | *Legionella_pneumophila_str_-Paris*
 167628644 | *Helio bacterium_modesticaldum_Icel*
 386854795 | *Deinococcus_gobiensis_I-0*
 148656006 | *Roseiflexus_sp_-RS-1*
 118431378 | *Aeropyrum_perfringens_K1*
 307595805 | *Vulcanisaeta_distributa_DSM_14429*
 305663328 | *Ignisphaera_aggregans_DSM_17230*
 408403002 | *Candidatus_Nitrosospaera_gargensis_Ga9-2*
 330509015 | *Methanosaeta_conciliif_gfp*
 11499461 | *Archaeoglobus_fulgidus_DSM_4304*
 383319611 | *Methanocella_conradii_H2254*

-RKSODYEAYGSVQEVRKLIKECATAPLPFED---DPRCM-RCSHAGI---CLPEERAHNIP
 -DDAARQELRAAVARAELRHIERPPVTSO---VRLCR-HCALAPV---CLPEEDRLAVIT
 -DEELRSKALKTIDRANLSTNSTPPNQVQNN---ENLCK-NCSLAPV---CLPEETRVTEN
 -DEELITFTRESLEQLRQVRQVPVPPPLED---SPKCP-RCSLVGI---CLPDETHLLQQE
 -GNTVEAAAEVIADEAREVASRENAPLVLGV---SKRCP-RCSLGLL---CMPDTEALLRR
 -DDSLIATTLRALDEARQLSGTTIPPPPLVD---SPKCP-RCSLVGI---CLPDETNLLRNS
 -DDELIGRTLAIDEMGTRALSTMPPPPLDHD---SPKCP-RCSLVGI---CLPDEVFRFLSH
 -TEALVATLVALSRLTVASGRLPPPLDH---SAKCP-RCSLLP1---CLPDEIAWFRKKG
 -DAELQRTEADAJARMHEVLLRQEVPVVARA---PRCD-KCSLEPL---CLPGVTAQGQSA
 -TPELFRAVEAAVVRMALLQSPPRPRVEVP---PSKCP-GCSVQGA---CQPELMWKVGAG
 -DPELRQKTI1AEQAHAIQCQGELPAPTS---KGKCR-DCSLKPL---CLPDEVAYLTH
 -TQALRNMTQEAI-IAAHACNGPMPPRIAN---TKKCA-DCSLERL---CLPKELKKLTK
 -DNLKRLKSALIATIQAVTNLLTTGIMTPNYS---KRCQ-GCSLNQ---CLPKAIDRKVTY
 -DLKLDRKT1TGTESVLLKIMEGEVIPENEYS---RRCR-ACSEIEI---CLPKEVSIKLKG
 -NDENLNELYKSSENIDKLDMAKPPEPVRN---YFCVHGCSYSDF---CWC---
 -SEENFNALETILREITALVNLAKPKPTPQ---KICK-NCAVNGY---CA---
 -DDDARRSALETTQARAVISAPAVPLER---RCCR-GCSYTDY---CWC---
 -TDEDEOQIRRDLDEIKTIPNLNDAPLPLFK---EGCK-GCSYFDL---CFS---
 -TADDEARLRTLVOELRQTAARPPTPSRLRN---RRCFCK-KCAYEEL---CFG---
 -DENNLSELEDVKNIENLIIQENPPEVINE---SKCK-KCAYFEY---CYI---
 -TEKDERIKEILNEIKRIISMDKPREFVK---DYCK-ACAYYPL---CWEVG
 -NEENKKIEDLKLMEMEVEIGKERDPKPKWVR---KICE-KCSYEL---CWA---
 -EGRERREVEEAIVEVEIKSSSKPBPKRS---KICR-KCAYKEL---CWG---
 -KEENIKVIEEAKIGIEEUVUSSPKPKPKV---PYCR-KCSYRDL---CWV---
 -TDENTRVEEAIRGYIDVNTSDSPPAEKE---PFCE-SCAHYDF---CWSC---
 -TED1IKEIDD1IKEIKKISNETKPHFKQK---PYCQ-GCSFREL---CLI---
 -DEARMQQLWESSERLERLAQQTQVPSAKR---PLCA-SCSLAEP---CGYD---
 -SYELHISARHILVERLLKILDSTPPSASAYG---GKCS-YCFFKSV---CSSSNQPAEIL
 -DWGSF1VERLVERLLETVSRLPPPVTPQD---PEKCL-ACRWRW---CPERSV---
 -RKEHVEYIYGKVVNNELESLEDSPSPVWNPS---PLLCK-ACQYRVR---CSVSVLL---
 -TEEHYRYIENVVKKIEEIVNSEKLPNVNE---KCD-FCQYRI---CPVSWNS---
 -DGHLRLEAEELVARVRDVLESEEEPPRAYRS---ERCS-SCWFRV---CPAWD---
 -NDOLVRDVSEAS1SRVERMISESSSLPKPQEO---GKCS-SCWYRPF---CPYA---
 -TGEVLEEARRYLGELERLASSDRPQOPTP---GRCS-SCWYRPF---CPYTF---
 -SY-SLREHVKGLEALERLRDLDVLPRGRPS---RYCS-YCKYSL---CPWRDP---
 -YEINSPDLEATIVLRAEITDILQNSPRSPRN---WECK-YCIFS1---CPAKLT---
 -SKAELKAETLLVKAABEVIEGP PPPVKTG---KCA-YCQYRAI---CPTP---
 -GDSERRFVLOAIEKRRIMB---GELIPRARS---RSCE-NCDFKNR---CLEI---
 -TSNDRNYI1KLTENNNKKTLEQUTTPKPTPS---KSCI-NCDYKN1---CPYQTQKQ---
 -TNNKLEKVKSEAAEALIIETERYPIELEIG---GTKCR-DCYYK1---CGR---
 -TPEMLRKVMEDAARELHMLQRGEPLQHHTVS---PRRCKR-VCSYRGV---CTYART---
 -TRDLVELVKHTAAQIRKIONGWTPTPAVHD---AKCP-SCWYRKY---CGFTSPHVEKI---
 -AETDKVAVKRAAEAIFTIIRERNRFYPKATKA---KARCV-TCTYRN1---C1K---
 -TSSDFKAKEKIIIEEILDIVQXQGLYPKTSRS---SRKCV-DCCYRN1---CV---
 -TPKMLDKAKLILDEIFMFINLEYKPKPAKA---KRKCL-DCTYRN1---CSV---
 -KKKDFFDAVAFVDLIFDIIENEEFPDG1KVN---KRKC1-DCCYRN1---CCK---
 -TDELINKLKITISDILSTLDGSKNPDSAS---DHQCI-QCEYNNP---CNDR---
 -SDGDKAIENNFVKLIEMIELGNLPHSSAD---GSKCT-QCEFLNY---CNDRP---
 -DEKSKPKVIEIACKKKIPEGTLPYSTAN---GLQCI-QCEFLNH---CNDRP---
 -TVMRMRDFFKDVLRIRAMVQKQSLSPKRP---GNRCP-DCEYRVV---CLLDGDVVR---
 -DTRLNRNTRRAIYQVVRGFLIAEQCPSPSTAQ---TVKCL-ECELRFN---CGDTI---
 -SAALRTRVKETVAAIRATVDRHMRPPIPAS---RRQCA-VCEFRRF---CNDVV---
 -SPSLRDRVILRADDMNKTLDGGLDEPPPKPG---RKCVC-ACRFRV---COPWVAERGSE
 -TPNLRTWIKRITIQDMKKHIQNQKEPPPKTIN---KAKCK-ACYFKD1---CPP---
 -SPNLRRTTINIIVEFKKYIQQQKPPPKTPM---KTRCN-HCFYRK1---CV---
 -TESMKTYLKKTTVKA1QNVNIESGEPPVARP---KSRCTGGCGYLWI---CGGIWNRR---
 -GDEIRRTLKMIEKIAQIIEDEIPAGTRN---SARCI-DCEYRKY---CVPNILPKPHT
 -SPDPRKALLRIIKEYIERLIEEEKYPRTSKS---KKCG-YCEVREP---CV---
 -TCEGKEMVKQDIRKIKDLIKSEKMPVNKG---LQCN-YCEMAKF---CS---

Supplementary Figure S2: Cas1, Cas2, Cas4 phylogenetic trees (please see Supplementary Materials and Methods for details and also see Makarova KS, Wolf YI, Koonin EV. **The basic building blocks and evolution of CRISPR-Cas systems**. Biochem Soc Trans. 2013 Dec 1;41(6):1392-400); Sequences are denoted by Genbank identifiers and organism genus and abbreviated species name; automatically identified CRISPR-Cas system subtype and gene in the Cas1 neighborhood are provided). The trees are provided in Newick format, could be open in any tree-viewing software.

Cas1 Tree:

```
((347756303|CAS-I-C|Chloracidobacterium_the__6-3-8c-7-5-1+4-2:0.38914,(294828260|CAS-I-C|Leptospira_int__1+4-2-x-x-6-3-8c-7-5:0.54801,162457462|CAS-I-C|Sorangium_cel__6-3-8c-7-5-1-2:0.50926)0.895:0.13172)1.000:0.61947,((15612904|CAS-I-C|Bacillus_hal__cmr5SS-cmr6g7-x-x-x-3-5-8c-7-4-1-2:0.64423,108763792|CAS-I-C|Myxococcus_xan__2-1-4-7-8c-5-3HD:0.49951)0.765:0.13333,(86742032|CAS-I-C|Frankia_Cci__3-5-8c-7-4-1-2:0.71514,433631663|CAS-I-C|Mycobacterium_can__2-1-4-7-8c-5-3:0.60677)0.734:0.20305)0.990:0.32273)0.717:0.14616,(219683006|CAS-I-U|Bifidobacterium_ani__3-csb1-1+4:1.07926,((378719041|CAS-I-U|Gordonia_pol__2-1+4-csb2-csb1-8-3:0.51986,134101617|CAS-III-A|Saccharopolyspora_ery__1+4-2-x-x-x-csm3g7-cmr3g5-csm3g7-10:0.62705)0.819:0.11364,(268318441|unknown|Rhodothermus_mar__2-1+4:0.43464,312114617|CAS-III-A|Rhodomicrobium_van__RT-2-1-csm3g7-csx19-csm3g7-csm3g7-10-x-csm6-csx1-x-csx16:1.15334)0.817:0.13822)0.378:0.07072,(320102061|CAS-I-U|Isosphaera_pal__1+4-2-csb2:0.57635,(39995168|CAS-I-U|Geobacter_sul__3-8-csb1-csb2-x-x-1+4-2:0.33269,83591519|CAS-I-E|Rhodospirillum_rub__2-1-6e-5-7-cse2SS-8e-3-x-x-2-1+4-csb2-x-x-csm3g7-csm3g7-csm3g7-10:0.61850)0.935:0.13614)0.960:0.17395)0.813:0.13734)0.850:0.11671,((38505684|CAS-ID|Synechocystis_PCC__3-10d-csc2g7-csc1g5-6-4-1-2:0.83136,(344996859|CAS-I-D|Caldicellulosiruptor_lac__2-1-4-6-3-csc1g5-csc2g7-10d:0.90289,46255267|CAS-I-A|Thermus_the__2-1-4-x-8a-7-5-6:0.82022)0.822:0.10403)0.844:0.08986,(((302391602|CAS-ID|Acetohalobium_ara__10d-csc2g7-csc1g5-3-4-1-2-6:0.44986,17229060|CAS-I-D|Nostoc_PCC__10d-csc2g7-csc1g5-3-2OG-6-4-1-x-2:0.80140)0.816:0.13790,159898755|CAS-I-D|Herpetosiphon_aur__10d-csc2g7-csc1g5-3-6-4-1-2:0.73922)0.853:0.07489,134299482|CAS-III-A|Desulfotomaculum_red__1+4-2-10-csm2SS-csm3g7-csm4g5-csx1-6:0.60078)0.965:0.13680)0.386:0.06285,((268318378|CAS-III-U|Rhodothermus_mar__csx1-csx20-x-x-csx1-csx1-6-x-2-1+4:0.71035,(336116789|CAS-III-A|Microlunatus_pho__10-csm3g7-cmr3g5-csm3g7-x-x-x-6-1+RT-2-x-10-csm3g7-cmr3g5-csm3g7:1.15764,(55978328|CAS-III-A|Thermus_the__csx1-1-x-10-csm2SS-csm3g7-csm4g5-csm5g7-csx1-x-x-csx1:0.32915,297622675|CAS-III-U|Truepera_rad__2-1-csx1-csx1-6:0.65702)0.999:0.43981)0.614:0.14144)0.987:0.24355,((76802276|CAS-I-D|Natronomonas_phae__R-6-10d-csc2g7-csc1g5-3-4-1-2:0.07764,(390962116|CAS-III-A|Thermococcus_CL1__6-10-csm2SS-csm3g7-csm4g5-csm5g7-R2-csx3-2-1:1.08282,(154174048|CAS-ID|Campylobacter_cur__1-2-4-3-5-7-8-6:1.20432,(((28211139|CAS-I-B|Clostridium_tet__1-4-3-5-7-8b-6:0.80992,(18977490|CAS-III-B|Pyrococcus_fur__2-1-4-3-5-7-8b-cmr6g7-cmr5SS-cmr4g7-csm6-cmr3g5:0.60617,((84489236|CAS-I-B|Methanospaera_stab__6-8-7-5-3-4-1-2:0.38030,(289596694|CAS-III-A|Aciduliprofundum_boo__2-1-4-6-csx1-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10:0.63757,13540937|CAS-III-A|Thermoplasma_vol__2-1-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10-6-csm6:0.86508)0.905:0.15392)0.906:0.14613,14590110|CAS-III-A|Pyrococcus_hor__10-csm2SS-csm3g7-csm4g5-csm5g7-csm6-8b-7-5-x-1-2-4-3-csx1:0.63787)0.819:0.15012,41614813|CAS-I-B|Nanoarchaeum_equ__2-1-x-7-5-3-4:0.77636)0.073:0.03836)0.175:0.14006)0.656:0.10760,268317185|CAS-III-A|Rhodothermus_mar__2-1-x-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10-4-3-5-6:0.62102)0.325:0.06368,(385809790|CAS-III-U|Ignavibacterium_alb__csm5g7-x-csx1-csx20-csx1-csx1-x-x-x-x-1-2:0.72273,150021526|CAS-III-B|Thermosiphon_mel__2-1-2-x-x-x-csm6-cmr1g7-10-cmr3g5-cmr4g7-cmr5SS:0.71602)0.932:0.15479)0.396:0.05380,(389844445|CAS-III-B|Mesotoga_pri__cmr6g7-cmr5SS-cmr4g7-cmr3g5-10-cmr1g7-csm6-2-x-x-1-4-3-5-7:0.68628,48477075|CAS-I-D|Picrophilus_tor__2-1-4-3-csc1g5-csc2g7-10d-6:0.97627)0.470:0.06969)0.718:0.10249,(55376278|CAS-I-B|Haloarcula_mar__6-8-7-5-3-4-1-2:0.65921,(292492938|CAS-ID|Nitrosococcus_hal__6-10d-csc2g7-x-3-4-1-2:1.06977,111224947|CAS-I-B|Frankia_aln__6-x-7-3+5-4-1-2:1.01974)0.382:0.12937)0.495:0.16546)0.929:0.19711)0.540:0.12245)0.997:0.40897)0.849:0.17499,((((317051217|CAS-III-A|Desulfurispirillum_ind__10-csm2SS-csm3g7-csm4g5-csm5g7-x-x-csx1-csx20-x-1-2-x-2:0.86471,17227877|CAS-III-B|Nostoc_PCC__csx18-1-2:0.92025)0.768:0.14478,((85858452|CAS-III-B|Syntrophus_aci__cmr5SS-cmr6g7-csm6-csx1-x-x-x-x-x-1-2-2:0.74830,(328949009|CAS-III-A|Treponema_suc__csx1-x-x-2-2-1-x-x-x-csx20-csx1-x-csm5g7-csm4g5-csm3g7-csm2SS:0.38670,333996012|CAS-III-B|Treponema_azo__csx20-csx1-cmr6g7-cmr5SS-cmr4g7-cmr3g5-10-cmr1g7-6-x-1-2-x-csx1:0.43621)1.000:0.46620)0.996:0.33602,(328953422|CAS-III-A|Desulfobacca_ace__csm5g7-csm6-x-csx20-csx1-x-x-x-Prim2+x-x-1-2:0.58824,(313673559|CAS-III-A|Calditerrivibrio_nit__2-Prim2-1-x-x-x-x-x-x-csx20-csm5g7:0.86969,218961167|CAS-III-A|Cloacamonas_aci__csm5g7-csa3-csx1+x-x-x-x-x-Prim2-x-1-2:0.68621)0.862:0.17223)0.840:0.15794,(154250301|unknown|Fervidobacterium_nod__1:0.90611,116748794|CAS-III-A|Syntrophobacter_fum__csm5g7-csm4g5-csm3g7-csm2SS-10-6-csx20-csx1-x-Prim2-1-2:0.93083)0.434:0.06593)0.080:0.06603)0.699:0.08120,(332296542|CAS-III-A|Thermodesulfobium_nar__4-x-2-1-Prim2-csm5g7-csm4g5-csm3g7-csm2SS-10-csx1-csx1-6-0.88721,337286709|CAS-III-B|Thermodesulfatator_ind__cmr5SS-cmr4g7-x-x-cmr3g5-10-cmr1g7-x-2-x-1-2-6:0.72067)0.539:0.11721)0.713:0.06506)0.646:0.10342,((148657121|CAS-III-U|Roseiflexus_RS__2-x-1-RT-x-x-x-x-csx15-csx1-x-csx1:0.61021,159898907|unknown|Herpetosiphon_aur__RT-1-2:0.55145)1.000:0.38100,(320161859|CAS-III-A|Anaerolinea_the__csx1-10-csm2SS-csm3g7-csm4g5-csm5g7-csx1-csx1-csx15-6-1-2:0.68967,(156741961|CAS-III-A|Roseiflexus_cas__10-csm2SS-csm3g7-csm4g5-csm5g7-csx1-6-1-2:0.47789,383762167|CAS-III-A|Caldilinea_aer__2-1-6-csx15-x-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10-csx1:0.57444)0.084:0.08868)0.989:0.24410)0.762:0.09771,37677204|CAS-III-Vibrio_vul__6-csm3g7-x-csm3g7-5g10-cmr4g7-10-x-x-1+RT-2-2-csx1-csx16-csx16-csm6:1.23121)0.614:0.07387)0.702:0.04517,(((296133514|CAS-III-A|Thermimincola_pot__2-1-csm6-6-csm5g7-csm4g5-csm3g7-csm2SS-10:0.55968,(348026605|CAS-III-A|Megasphaera_els__1-2-10-csm2SS-csm3g7-csm4g5-csm5g7-6-
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csm6:0.72836,114567264|CAS-III-A|Syntrophomonas_wol__csm6-x-2-1-x-6-csm5g7-csm4g5-csm3g7-csm2SS-
 10:0.55211)0.097:0.09427)0.924:0.14860,(116627764|CAS-III-A|Streptococcus_the__1-2-6-csm2SS-csm3g7-csm4g5-csm5g7-x-
 csm6:1.15855,15609954|CAS-III-A|Mycobacterium_tub__2-1-csm6-csm5g7-csm4g5-csm3g7-csm2SS-10-
 6:0.80127)0.948:0.34739)0.991:0.30794,(218442809|CAS-III-A|Cyanothece_PCC__csm6-x-x-RT-1-2:0.82366,(38505760|CAS-III-
 B|Synechocystis_PCC__cmr6g7-cmr5SS-cmr4g7-x-cmr3g5-10-csx18-1-2:1.05148,((119357853|CAS-III|Chlorobium_pha__2-1-
 6:0.57832,119357846|CAS-III-A|Chlorobium_pha__6-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10-x-csx15-csx1-1+RT-
 2:0.71133)0.908:0.16639,(427708213|CAS-III|Nostoc_PCC__csx19-csm3g7-x-x-6-x-x-2-1-x-x-1+RT:1.09536,17228961|CAS-III-
 A|Nostoc_PCC__1+RT-x-x-csm3g7-csx19-csm3g7-csm3g7-x-csm3g7-
 csm3g7:0.37340)0.986:0.34526)0.475:0.09413)0.277:0.05347)0.828:0.10795)0.964:0.17610,(328953000|CAS-III|Desulfobacca_ace__6-x-x-
 x-x-x-RT-x-x-1-2:0.79732,(261415948|CAS-III-A|Fibrobacter_suc__1-2-x-x-csx1-10-csm3g7-x-csm3g7-csx20:0.76621,269798874|CAS-
 III-A|Veillonella_par__6-csm3g7-csx19-csm3g7-5g10-csm3g7-10-csx20-1-2-x-csx1-
 csx1:1.17138)0.843:0.22745)0.760:0.16672)0.679:0.01909)0.789:0.04869,(((326793969|CAS-III-B|Marinomonas_med__2-6-x-x-cmr6g7-
 cmr5SS-cmr4g7-cmr3g5-10-cmr1g7:0.97163,220934970|CAS-III-B|Thioalkalivibrio_sul__cmr1g7-10-cmr3g5-cmr4g7-cmr5SS-cmr6g7-
 csx1-x-csx1-x-1-2-x-csx1:0.60767)0.999:0.58432,(436839745|CAS-III-B|Desulfovibrio_hyd__1+RT-2-x-x-cmr6g7-cmr5SS-cmr4g7-
 cmr3g5-10-cmr1g7:1.37288,(283778924|CAS-III-A|Pirellula_sta__1+RT-2-x-10-csm3g7-
 csm3g7:1.45956,118497969|unknown|Francisella_nov__2OG-x-x-x-x-csa3-2-1-
 4:1.55280)0.320:0.28782)0.806:0.12534,(319790504|unknown|Thermovibrio_amm__2-2-2-1:1.30366,337288662|CAS-III-
 A|Thermodesulfobacterium_OPB__10-csm2SS-csm3g7-csm4g5-csm5g7-6-csx1-x-2-1-
 2:1.11781)0.980:0.53351)0.906:0.16162)0.600:0.09143,(((325980970|unknown|Nitrosomonas_AL2__1:1.73273,(407462288|unknown|Nitr-
 osopumilus_kor__1:1.50397,(21229351|unknown|Methanosaerina_maz__1:0.67446,147921327|unknown|Methanocella_arv__1:0.9953
 7)0.976:0.45554)0.831:0.19198)0.904:0.19310,((150398872|CAS-III-B|Methanococcus_van__10-cmr3g5-cmr4g7-cmr5SS-cmr6g7-x-x-6-
 1-2:0.84562,((((332668016|unknown|Haliscomenobacter_hyd__RT-1-PDEDxK-2:1.20570,(327314603|CAS-III|Prevotella_den__10-
 cmr6g7-5g10-csm3g7-x-csm3g7-csx1-1-Prim-2:0.73847,(379728404|unknown|Saprosira_gra__2-x-
 1:0.87455,392395717|unknown|Flexibacter_lit__2-x-1:0.68194)0.836:0.12742)0.831:0.12483)0.909:0.21557,284040851|CAS-I-
 B|Spirosoma_lin__8-7-5-3-6-1-2-4:0.73413)0.213:0.10519,218961577|CAS-I-B|Cloacamonas_aci__4-2-1-3HD-5-7-8-
 6:0.58565)0.358:0.09219,257125862|CAS-III|Leptotrichia_buc__csm3g7-6-x-x-x-x-4-2-1-x-x-csx1-
 csx20:0.61093)0.324:0.10225,(21226661|CAS-I-B|Methanosaerina_maz__4-2-1-6-3-5-7-8-csa3:0.33047,150401497|CAS-III-
 A|Methanococcus_aeo__6-3-5-7-8-1-2-4-x-csx1-csm5g7-csm4g5-csm3g7-csm2SS-
 10:0.92232)0.484:0.09616)0.933:0.21764)1.000:0.55180,(((11499462|CAS-I-A|Archaeoglobus_ful__cmr1g7-csa3-csa5SS-7-5-csa8a2-3-
 3HD-2-4-1-4:0.41184,383319612|CAS-I-A|Methanocella_con__4-1-4-2-csa3-x-x-x-x-7:0.53988)0.920:0.13193,(288560430|CAS-III-
 A|Methanobrevibacter_rum__1-x-2-6-csx1-csm5g7-csm4g5-csm3g7-csm2SS-
 10:0.47372,148642083|unknown|Methanobrevibacter_smi__1-2:0.88751)0.962:0.26391)0.999:0.31122,((((((156937921|CAS-I-
 A|Ignicoccus_hos__4-1-2-4-3HD-3-x-5-7-csa5SS-R:0.93227,124027534|CAS-I-A|Hyperthermus_but__csaXg8-5-7-csa5SS-R-x-x-4-2-4-
 1-x-x-x-R:0.65197)0.613:0.22777,(305663330|CAS-I-A|Ignisphaera_agg__R-x-7-5-3-3HD-x-6-4-2-1-4:0.36006,307595807|CAS-I-
 A|Vulcanisaeta_dis__3HD-3-x-5-7-x-R-x-4-2-1-4:0.64930)0.000:0.08743)0.820:0.16306,118431380|CAS-I-A|Aeropyrum_per__6-8a2-
 3HD-3-5-7-csa5SS-R-4-2-1-4:0.62305)0.991:0.27322,(347523053|CAS-I-A|Pyrolobus_fum__4-x-x-x-x-2-1-4-R-x-x-3HD-3-8a2-5-7-
 csa5SS:0.78895,((218884093|CAS-I-A|Desulfurococcus_kam__6-3HD-3-csaXg8-5-7-csa5SS-R-4-2-1-4-R-x-cmr3g5-10-cmr6g7-cmr1g7-
 cmr5SS-cmr4g7-csm6:0.52821,389861365|CAS-I-A|Thermogladius_163__6-x-x-4-1-4-R:0.39368)0.988:0.22330,(302347808|CAS-III-
 B|Acidilobus_sac__6-x-4-2-1-4-cmr3g5-10-cmr6g7-csm3g7-cmr4g7-csm6-x-csm6:0.75489,(385805115|CAS-I-
 A|Fervidicoccus_fon__6-R-3HD-3-8a2-5-7-x-x-4-1-2-4-R-R-x-x-csm6-cmr6g7-cmr1g7-cmr5SS:1.07762,(15898286|CAS-I-
 A|Sulfolobus_sol__csaXg8-3HD-3-5-7-csa5SS-R-R-x-4-1-4:0.66104,15922975|CAS-I-A|Sulfolobus_tok__R-4-1-2-4-x-x-csa5SS-7-5-3-
 3HD-csaXg8:0.46000)0.982:0.27868)0.852:0.19905)0.274:0.09946)0.000:0.06113)0.643:0.12690)0.943:0.18315,((18311772|CAS-I-
 A|Pyrobaculum_aer__6-4-2-1-4-csa3-x-8a2-3HD-3-5-7-csa5SS-6:0.90508,18311699|CAS-I-A|Pyrobaculum_aer__3HD-3-x-x-6-csa3-x-
 4-2-1-4:0.82264)0.635:0.11599,15898245|CAS-I-A|Sulfolobus_sol__csx1-csa5SS-7-5-csaXg8-3-3HD-2-1-
 6:1.32874)0.273:0.10934)0.979:0.21717,((34541577|CAS-III-B|Porphyromonas_gin__2-1+RT-cmr6g7-x-cmr4g7-cmr3g5-
 10:1.17610,(386750121|CAS-III|Helicobacter_cet__csm5g7-csx1-x-x-x-x-x-x-2-1:1.23102,328954440|CAS-III-
 A|Desulfobacca_ace__1+RT-x-x-csm3g7-x-csm6-csm3g7-x-csm3g7-csm3g7:0.77196)0.683:0.11126)0.716:0.14667,(54296139|CAS-II-
 B|Legionella_pne__9-1-2-4:0.74021,(34557933|CAS-II-B|Wolinella_suc__9-1-2-4:0.46386,118497353|CAS-II-B|Francisella_nov__9-1-
 2-4:0.52534)0.502:0.15130)1.000:0.52829)0.629:0.07561,(148656004|CAS-III-U|Roseiflexus_RS__csm3g7-10-csx1-csx1-cs3-8b-7-5-3-6-1-
 2-4:0.77295,(386854797|CAS-I-A|Deinococcus_gob__4-2-1-5-7-8a2-3-6:0.75119,167628642|CAS-I-A|Heliobacterium_mod__4-4-2-1-x-6-
 5-7-x-3:0.54240)0.076:0.07286)0.605:0.12022)0.964:0.15886)0.878:0.08588,((126465226|CAS-I-A|Staphylothermus_mar__csm3g7-
 csm4g5-csm5g7-10-x-x-1-4-2-4-6-R-csa5SS-7-5-csa8a2-3:0.60120,170290046|CAS-I-A|Korarchaeum_cry__1-4-2-4-R-csa5SS-7-5-csa8a2-
 x-3:0.38512)1.000:0.74406,(330509017|CAS-I-A|Methanosaeta_con__4-4-2-1-3-x-x-x-5:0.72091,408403000|CAS-
 I|Nitrososphaera_gar__4-1-2-4-x-x-3-5-7-x-
 6:0.85041)0.834:0.11037)0.627:0.05146)0.645:0.08920)0.700:0.09695)0.851:0.07534,(119720226|CAS-III-B|Thermofilum_pen__cmr3g5-
 10-cmr6g7-cmr1g7-cmr5SS-cmr4g7-1-2-3-csc1g5-csc2g7-x-10d:1.21220,(156937861|CAS-III|Ignicoccus_hos__6-x-x-
 1:0.82794,70607609|CAS-II-D|Sulfolobus_aci__3-csc1g5-csc2g7-10d-R-6-x-2-4-1-x-x-x-x-x-x-x-
 csm3g7:0.96376)0.956:0.32568)1.000:0.59773)0.890:0.10459)0.784:0.05200,((((187250661|CAS-II|Elusimicrobium_min__9-1-
 2:0.61891,325972002|CAS-II|Spirochaeta_Bud__2-1-9:0.74940)0.971:0.28775,(((386318631|CAS-II|Staphylococcus_pse__9-1-
 2:0.79215,(15675042|CAS-II-A|Streptococcus_pyo__9-1-2-csn2:0.79682,(352684362|CAS-II-A|Acidaminococcus_int__9-1-2-
 csn2:0.51095,(169823756|CAS-II-A|Finegoldia_mag__csa3-x-x-x-9-1-2-csn2-PDEDxK:0.61846,42525844|CAS-II-

A|Treponema_den___9-1-2-csn2:0.40539)0.759:0.12239)0.928:0.19604)0.863:0.16984)0.000:0.05417,(302336019|CAS-II-
 A|Olsenella_uli___csn2-2-1-9:0.70415,(90961086|CAS-II-A|Lactobacillus_sal___9-x-x-1-2-csn2:0.80559,(310286727|CAS-II-
 A|Bifidobacterium_bif___csn2-1-9:0.67816,406600270|CAS-II-A|Leuconostoc_gel___csn2-2-1-
 9:0.48323)0.903:0.16893)0.956:0.22166)0.823:0.11869)0.980:0.22310,((269123825|CAS-II|Streptobacillus_mon___2-1-
 9:1.01742,(47458867|CAS-II|Mycoplasma_mob___1-9:0.80008,(71894593|CAS-II-A|Mycoplasma_syn___9-1-2-csn2-
 1:0.77706,193216853|CAS-II|Mycoplasma_art___1-2:0.64612)0.976:0.30122)0.975:0.30654)0.473:0.11491,(238924076|CAS-
 II|Eubacterium_rec___9-1-2:0.79548,116627543|CAS-II|Streptococcus_the___9-1-2:0.73103)0.704:0.20755)0.000:0.04759,57865886|CAS-
 III-A|Staphylococcus_epicarditis___6-csm6-csm5g7-csm4g5-csm3g7-csm2SS-10-2-1:1.23206)0.549:0.13652)0.933:0.18660,((187736488|CAS-II-
 II|Akkermansia_muc___2-1-9:1.39758,319957207|CAS-II|Nitratirfractor_sal___9-1-2:0.78599)0.825:0.24500,((189485226|CAS-I-
 C|uncultured_Ter___3-5-8c-7-4-1-2-9-1-2:0.72350,34557789|CAS-II|Wolinella_suc___2-1-9:0.70129)0.951:0.22060,(15602991|CAS-II-
 II|Pasteurella_mul___2-1-9:0.67060,(222109284|CAS-II|Acidovorax_ebr___2-1-9:0.79135,(310780383|CAS-II|Ilyobacter_pol___2-1-
 9:0.66637,220930481|CAS-II|Clostridium_cel___2-1-9:0.45815)0.495:0.12122)0.235:0.10067)0.527:0.07214,(57238545|CAS-II|
 Campylobacter_jej___2-1:0.68707,291276264|CAS-II|Helicobacter_mus___2-1-
 9:0.53967)0.991:0.31535)0.696:0.07398)0.467:0.12246,((90425960|CAS-II|Rhodopseudomonas_pal___2-1-9:0.59972,209542526|CAS-II|
 Gluconacetobacter_dia___9-x-1-2:0.34389)0.999:0.30076,((60683388|CAS-II|Bacteroides_fra___2-1-9:0.67044,154250554|CAS-II|
 Parvibaculum_lav___2-1-9:0.59613)0.616:0.18474,(294086112|CAS-II|Puniceispirillum_mar___9-1-2:0.75736,121608212|CAS-II|
 Verminephrobacter_eis___9-1-
 2:0.66201)0.970:0.28280)0.927:0.14692)0.886:0.15928)0.523:0.07411)0.916:0.17687)0.434:0.13234)0.871:0.16414,(117929157|CAS-II|
 Acidothermus_cel___2-1-9:0.43620,(283456134|CAS-II|Bifidobacterium_den___2-1-9:0.77563,38232679|CAS-II|
 Corynebacterium_dip___9-1-2:0.38942)0.984:0.34995)0.999:0.58714)1.000:0.76830,((50085564|CAS-I-F|Acinetobacter_ADP___3f-x-
 8f-5f-7f-6f-1:0.87259,(77166205|CAS-I-F|Nitrosococcus_oce___3f-1-8f-5f-7f-6f:0.70997,15602176|CAS-I-F|Pasteurella_mul___8f-5f-7f-6f-
 x-x-1-3f:0.74246)0.432:0.16689)0.999:0.75717,(38234755|CAS-I-E|Corynebacterium_dip___5-7-cse2SS-8e-6e-3-1-
 2:1.23531,((53804737|CAS-I-E|Methylococcus_cap___2-1-6e-5-7-cse2SS-8e-3:0.57557,209966592|CAS-I-E|Rhodospirillum_cen___2-1-6e-
 5-7-cse2SS-8e-3:0.74976)0.998:0.72881,((68535718|CAS-I-E|Corynebacterium_jei___1-6e-5-7-cse2SS-8e-3:0.84511,86738746|CAS-I-
 E|Frankia_CclI___3-8e-cse2SS-7-5-6e-1-2:0.59793)0.909:0.30308,(89885834|CAS-I-E|Rhodoferax_fer___1-6e-csf3g5-csf2g7-x-csf1g8-
 Ding:1.24990,16130662|CAS-I-E|Escherichia_col___2-1-6e-5-7-cse2SS-8e-3:0.63487)0.991:0.50550,(51891801|CAS-I-
 E|Symbiobacterium_the___1-3-8e-cse2SS-7-5-6e:0.60107,21674788|CAS-I-E|Chlorobium_tep___3-8e-cse2SS-6e-7-5-1-
 2:1.27314)0.707:0.10574)0.656:0.10724)0.799:0.10534,29834079|CAS-I-E|Streptomyces_ave___1-6e-5-7-cse2SS-8e-
 3:0.75713)0.915:0.20711)0.936:0.38681)0.855:0.25733)0.999:0.98797,(124485266|unknown|Methanocorpusculum_lab___1:0.82627,((15415
 0254|unknown|Methanoregula_boo___1:0.58226,((126178701|unknown|Methanoculleus_mar___1:0.07443,397780289|unknown|Methano-
 culleus_bou___1:0.16232)0.999:0.36213,307352568|unknown|Methanoplanus_pet___1:0.81980)0.633:0.12886)0.049:0.09156,219852754|un-
 known|Methanospaerula_pal___1:0.54240)0.915:0.20557)1.000:0.95417)0.185:0.13326)0.547:0.10037,332661941|CAS-III-
 B|Haliscomenobacter_hyd___2-1-x-RT-cmr6g7-x-cmr4g7-cmr3g5-10:2.03433)0.792:0.07389)0.993:0.24809,((288947693|CAS-III-
 U|Allochromatium_vin___csx1-x-csx16-x-2-2-1:0.63642,431929780|CAS-III-B|Thioflavicoccus_mob___cmr4g7-cmr3g5-10-cmr1g7-csx1-
 x-x-x-2-2-1-RT:0.71938)0.990:0.42353,((292492856|CAS-III|Nitrosococcus_hal___csm4g5-csm5g7-x-csx1-x-x-csx1-csx1-x-2-1-x-
 2:0.61028,((206890539|CAS-III-A|Thermodesulfovibrio_yel___csm2SS-csm3g7-csm4g5-csm5g7-6-x-x-x-2-1-
 2:0.94103,(52425690|CAS-III-U|Mannheimia_suc___2-1-2-x-csx1-csx1-csx16-x-6:0.83608,30248142|CAS-III|Nitrosomonas_eur___2-1-2-
 csx1-x-csx1-csx16-x-csm5g7-csm4g5-csm3g7:0.51689)0.754:0.15286)0.875:0.16834,430760780|CAS-III-
 B|Thioalkalivibrio_nit___cmr4g7-cmr5SS-cmr6g7-csx1-x-csx1-x-6-2-1-2-1-2:0.57693)0.171:0.10187)0.479:0.12125,121997419|CAS-III-
 B|Halorhodospira_hal___1-2-1-2-csx1-csx16-csx1-cmr1g7-10-cmr3g5-cmr4g7-cmr5SS-
 cmr6g7:0.80291)0.973:0.28121)0.994:0.39809)0.748:0.06442)0.802:0.09835,(88602029|CAS-III-A|Methanospirillum_hun___csx1-x-x-x-
 csx1-csx1-x-x-2-1-x-x-x-csm3g7-csm3g7-csm3g7-10:1.27757,83592490|CAS-III-A|Rhodospirillum_rub___csm3g7-csm3g7-
 csm3g7-10-6-2-1:0.90172)0.839:0.25935,(294676823|CAS-III-A|Rhodobacter_cap___RT-x-x-x-x-x-10-csm3g7-csm3g7-2-1+RT-RT-
 6:0.58881,(374998939|CAS-III-B|Azospirillum_lip___2-1+RT-x-csm3g7-x-x-x-cmr1g7-10-cmr3g5-cmr4g7-
 cmr5SS:0.39128,(389875622|CAS-III-A|Tistrella_mob___10-cmr4g7-csm3g7-csm3g7-6-x-1+RT-2:0.63303,(302343124|CAS-III-
 A|Desulfarculus_baa___csm3g7-csm3g7-csm3g7-10-1+RT-2-6:0.64546,288957883|CAS-III-A|Azospirillum_B51___csm3g7-csm3g7-
 csm3g7-6-2-1+RT:0.24663)0.814:0.09915)0.807:0.11976)0.877:0.14374)1.000:0.61254,20094748|CAS-III-A|Methanopyrus_kan___2-x-1-x-
 10-csm2SS-csm3g7-csm4g5-csx1-
 csm5g7:1.51613)0.666:0.10913)0.922:0.15239)0.946:0.10901)0.045:0.02893)0.725:0.07257)0.132:0.03771)0.879:0.12499);

Cas2 Tree:

((((((269123824_Streptobacillus_mon___CAS-II_2-1-9:0.65435000,(90425959_Rhodopseudomonas_pal___CAS-II_2-1-
 9:0.22101000,209542527_Gluconacetobacter_dia___CAS-II_9-x-1-
 2:0.17402000)0.94:0.30411000)0.77:0.09933000,((222109283_Acidovorax_ebr___CAS-II_2-1-
 9:0.42855000,310780382_Ilyobacter_pol___CAS-II_2-1-9:0.13912000)0.66:0.12085000,15602990_Pasteurella_mul___CAS-II_2-1-
 9:0.37916000)0.90:0.20861000,(291276263_Helicobacter_mus___CAS-II_2-1-9:0.31167000,57238544_Campylobacter_jei___CAS-II_2-1-
 9:0.28830000)0.97:0.34796000,189485227_uncultured_Ter___CAS-I-C_3-5-8c-7-4-1-2-9-1-
 2:0.13815000)0.81:0.07699000)0.63:0.13694000)0.85:0.07727000,(15675043_Streptococcus_pyo___CAS-II-A_9-1-2-

csn2:0.46351000,((374307736_Filifactor_alo_CAS-II-A_csn2-2-1-9:0.28271000,302336018_Olsenella_uli_CAS-II-A_csn2-2-1-9:0.43920000)0.76:0.09571000,(42525845_Treponema_den_CAS-II-A_9-1-2-csn2:0.39033000,169823757_Finegoldia_mag_CAS-II-A_csa3-x-x-x-9-1-2-csn2-PDEDxK:0.34299000)0.77:0.11939000)0.84:0.10746000)0.96:0.38688000)0.79:0.08033000,220930480_Clostridium_cel_CAS-II-2-1-9:0.29802000)0.88:0.09183000,((319957208_Nitratifractor_sal_CAS-II_9-1-2:0.70234000,187736487_Akkermansia_muc_CAS-II-2-1-9:0.70126000)0.52:0.04877000,(34557788_Wolinella_suc_CAS-II_2-1-9:0.64671000,((116627544_Streptococcus_the_CAS-II_9-1-2:0.56326000,238924077_Eubacterium_rec_CAS-II_9-1-2:0.13222000)0.94:0.21511000,(90961087_Lactobacillus_sal_CAS-II-A_9-x-x-1-2-csn2:0.11325000,(406600269_Leuconostoc_gel_CAS-II-A_csn2-2-1-9:0.31623000,331702226_Lactobacillus_buc_CAS-II_2-1-9:0.07283000)0.66:0.08097000)0.97:0.21453000)0.73:0.09770000)0.09:0.07068000)0.16:0.13698000)0.85:0.12415000,(60683387_Bacteroides_fra_CAS-II_2-1-9:0.32820000,154250553_Parvibaculum_lav_CAS-II_2-1-9:0.27515000)0.72:0.07487000)0.81:0.13684000,((294086113_Puniceispirillum_mar_CAS-II_9-1-2:0.53000000,(189440766_Bifidobacterium_lon_CAS-II_9-1-2:0.31549000,(117929156_Acidothermus_cel_CAS-II_2-1-9:1.13966000,38232680_Corynebacterium_dip_CAS-II_9-1-2:0.36119000)0.34:0.23863000)0.96:0.35689000)0.60:0.03762000,(121608213_Verminephrobacter_eis_CAS-II_9-1-2:0.53910000,(57865885_Staphylococcus_epii_CAS-III-A_6-csm6-csm5g7-csm4g5-csm3g7-csm2SS-10-2-1:1.29537000,(325972001_Spirochaeta_Bud_CAS-II_2-1-9:0.39221000,187250662_Elusimicrobium_min_CAS-II_9-1-2:0.87228000)0.61:0.25072000)0.68:0.09900000)0.79:0.12183000)0.88:0.15899000)0.82:0.22250000,(193216854_Mycoplasma_art_CAS-II_1-2:0.60540000,71894594_Mycoplasma_syn_CAS-II-A_9-1-2-csn2-1:0.22111000)0.93:0.55087000)0.97:0.66890000,((148642084_Methanobrevibacter_smi_CAS-II_A_9-1-2-csn2-1-2:1.27532000,((((283778925_Pirellula_sta_CAS-III-A_1+RT-2-x-10-csm3g7-csm3g7:0.89402000,319790502_Thermovibrio_amm_CAS-II_2-2-2-1:0.93403000)0.80:0.21342000,((118497968_Francisella_nov_CAS-II_2-1-csm6-csm5g7-1-2:1.15971000,337288663_Thermodesulfobacterium_OPB_CAS-III-A_10-csm2SS-csm3g7-csm4g5-csm5g7-6-csx1-x-2-1-2:0.79480000)0.77:0.26073000,(319790501_Thermovibrio_amm_CAS-II_2-2-2-1:0.48746000,319790503_Thermovibrio_amm_CAS-II_2-2-2-1:0.90092000)0.74:0.15026000)0.72:0.06215000)0.82:0.16445000,((15609953_Mycobacterium_tub_CAS-III-A_2-1-csm6-csm5g7-csm4g5-csm3g7-csm2SS-10-6:0.51178000,((296133513_Thermincola_pot_CAS-III-A_2-1-csm6-6-csm5g7-csm4g5-csm3g7-csm2SS-10:0.29366000,114567263_Syntrophomonas_wol_CAS-III-A_csm6-x-2-1-x-6-csm5g7-csm4g5-csm3g7-csm2SS-10:0.23322000)0.89:0.35153000,348026606_Megasphaera_els_CAS-III-A_1-2-10-csm2SS-csm3g7-csm4g5-csm5g7-6-csm6:0.67555000)0.78:0.13760000)0.90:0.31648000,116627765_Streptococcus_the_CAS-III-A_1-2-6-csm2SS-csm3g7-csm4g5-csm5g7-x-csm6:0.60642000)0.88:0.30205000)0.75:0.03587000)0.86:0.11603000,((((269798875_Veillonella_par_CAS-III-A_6-csm3g7-csm2SS-10-csm3g7-5g10-csm3g7-10-cs20-1-2-x-cs1-cs1:0.77156000,88602028_Methanospirillum_hun_CAS-III-A_csm1-x-x-x-cs1-cs1-x-2-1-x-x-x-cs3g7-csm3g7-x-x-csm3g7:0.85308000)0.87:0.52835000,20094746_Methanopyrus_kan_CAS-III-A_2-x-1-x-10-csm2SS-csm3g7-csm4g5-cs1-csm5g7:1.62011000)0.51:0.19062000,(((37677205_Vibrio_vul_CAS-III_6-csm3g7-x-csm3g7-5g10-cmr4g7-10-x-x-1+RT-2-2-cs1-cs16-cs16-csm6:0.72100000,83592489_Rhodospirillum_rub_CAS-III-A_csm3g7-csm3g7-csm3g7-10-6-2-1:0.71732000)0.78:0.22771000,436839746_Desulfovibrio_hyd_CAS-III-B_1+RT-2-x-x-cmr6g7-cmr5SS-cmr4g7-cmr3g5-10-cmr1g7:0.74051000)0.17:0.14583000,317051220_Desulfurispirillum_ind_CAS-III-A_csm2SS-csm3g7-csm4g5-csm5g7-x-x-cs1-cs20-x-1-2-x-2:1.75303000:0.05677000,((386750120_Helicobacter_cet_CAS-III_csm5g7-cs1-x-x-x-x-x-x-2-1:1.01565000,((((116748795_Syntrophobacter_fum_CAS-III-A_csm4g5-csm3g7-csm2SS-10-6-cs20-cs1-x-Prim2-1-2:0.31037000,218961168_Cloacamonas_aci_CAS-III_csm6-x-cs20-cs1-x-x-x-Prim2-1-2:0.54685000)0.85:0.15969000,(328953423_Desulfobacca_ace_CAS-III_csm6-x-cs20-cs1-x-x-x-Prim2+x-x-1-2:0.41723000,(313673557_Calditerrivibrio_nit_CAS-III_csm5g7-csm3g7-csm2SS-10-cs1-cs1-6:0.43566000)0.40:0.12454000)0.15:0.04882000)0.92:0.18403000,(383762166_Caldilinea_aer_CAS-III-A_2-1-6-cs15-x-cs1-csm5g7-csm4g5-csm3g7-csm2SS-10:0.51565000,((156741962_Roseiflexus_cas_CAS-III-A_10-csm2SS-csm3g7-csm4g5-csm5g7-cs1-6-1-2:0.24695000,320161860_Anaerolinea_the_CAS-III-A_10-csm2SS-csm3g7-csm4g5-csm5g7-cs1-cs1-cs15-6-1-2:0.53996000)0.80:0.22964000,328953001_Desulfobacca_ace_CAS-III_6-x-x-x-x-x-RT-x-x-1-2:0.37791000)0.38:0.04439000)0.63:0.03747000)0.84:0.09147000,(159898756_Herpetosiphon_aur_CAS-I-D_10d-csc2g7-csc1g5-3-6-4-1-2:0.79678000,((134299483_Desulfotomaculum_red_CAS-III-A_1+4-2-10-csm2SS-csm3g7-csm4g5-cs1-6-1-2:0.63110000,119357847_Chlorobium_phae_CAS-III-A_csm1-csm5g7-csm4g5-csm3g7-csm2SS-10-x-cs15-cs1-1+RT-2-x-x-x-x-2-1-6:0.38435000)0.57:0.12431000,(76802277_Natromonas_phae_CAS-I-D_R-6-10d-csc2g7-csc1g5-3-4-1-2:1.07211000,302391603_Acetohalobium_ara_CAS-I-D_10d-csc2g7-csc1g5-3-4-1-2-6:0.31311000)0.70:0.11597000)0.69:0.09059000,75910375_Anabaena_var_CAS-I-D_10d-csc2g7-csc1g5-2OG-6-4-1-2:0.30845000)0.71:0.06011000)0.92:0.23922000)0.82:0.08356000)0.60:0.01366000,(337286707_Thermodesulfatator_ind_CAS-III-B_csm1-cmr6g7-cmr5SS-cmr4g7-x-x-cmr3g5-10-cmr1g7-x-2-x-1-2-6:0.54544000,(119357852_Chlorobium_phae_CAS-III-A_csm1-csm5g7-csm4g5-csm3g7-csm2SS-10-x-cs15-cs1-1+RT-2-x-x-x-2-1-6:0.29854000,(((317051218_Desulfurispirillum_ind_CAS-III-A_csm2SS-csm3g7-csm4g5-csm5g7-x-x-cs1-cs20-x-1-2-x-2:0.47733000,85858453_Syntrophus_aci_CAS-III_cmr6g7-csm6-cs1-x-x-x-x-x-1-2-2:0.90326000)0.42:0.1806000,(328949008_Treponema_suc_CAS-III-A_csm1-x-x-2-2-1-x-x-x-cs20-cs1-x-csm5g7-csm4g5-

csm3g7:0.26185000,333996013_Treponema_azo____CAS-III-B_csx1-cmr6g7-cmr5SS-cmr4g7-cmr3g5-10-cmr1g7-6-x-1-2-x-csx1:0.75344000)0.10:0.23727000)0.72:0.19590000,((34541576_Porphyromonas_gin____CAS-III-B_2-1+RT-cmr6g7-x-cmr4g7-cmr3g5-10:0.69930000,336116790_Microlunatus_ph____CAS-III-A_10-csm3g7-cmr3g5-csm3g7-x-x-x-6-1+RT-2-x-10-csm3g7-cmr3g5-csm3g7:0.98282000)0.85:0.43712000,268318377_Rhodothermus_mar____CAS-III-U_csx1-csx20-x-x-csx1-csx1-6-x-2-1+4:0.44664000)0.92:0.50562000)0.36:0.17225000,((((30248141_Nitrosomonas_eur____CAS-III-A_2-1-2-csx1-x-csx1-csx16-6-x-csm5g7-csm4g5-csm3g7-csm2SS:0.44781000,52425689_Mannheimia_suc____CAS-III-U_2-1-2-x-csx1-csx1-csx16-x-6:0.59489000)0.90:0.32569000,430760782_Thioalkalivibrio_nit____CAS-III-B_cmr3g5-cmr4g7-cmr5SS-cmr6g7-csx1-x-csx1-x-x-6-2-1-2:0.51091000)0.80:0.19099000,(431929779_Thioflavicoccus_mob____CAS-III-B_cmr6g7-cmr5SS-cmr4g7-cmr3g5-10-cmr1g7-csx1-x-x-2-2-1-RT:0.47668000,288947692_Allochromatium_vin____CAS-III-U_cmr6g7-x-x-x-x-csx1-x-csx16-x-2-2-1:0.83407000)0.89:0.32391000)0.83:0.20682000,292492858_Nitrosococcus_hal____CAS-III-A_csm3g7-csm4g5-csm5g7-x-csx1-x-x-csx1-csx1-x-2-1-x-2:0.56529000)0.85:0.21514000,(206890293_Thermodesulfovibrio_yel____CAS-III-A_10-csm2SS-csm3g7-csm4g5-csm5g7-6-x-x-x-2-1-2-1:0.54995000,121997420_Halarhodospira_hal____CAS-III-B_1-2-1-2-csx1-csx16-csx1-cmr1g7-10-cmr3g5-cmr4g7-cmr5SS-cmr6g7-6:0.80448000)0.82:0.18626000)0.92:0.35773000,((268318440_Rhodothermus_mar____CAS-I-C_2-1+4:0.65320000,((320102062_Isosphaera_pal____CAS-I-Cu_1+4-2-csb2:0.62829000,(134101618_Saccharopolyspora_ery____CAS-III-A_1+4-2-x-x-x-csm3g7-cmr3g5-csm3g7-10:0.50543000,(83591518_Rhodospirillum_rub____CAS-I-E_2-1-6e-5-7-cse2SS-8e-3-x-x-2-1+4-csb2-x-x-csm3g7-csm3g7-csm3g7-10:0.49671000,39995169_Geobacter_sul____CAS-I-Cu_3-8-csb1-csb2-x-x-1+4-2:0.23073000)0.92:0.28853000,312114616_Rhodomicrobium_van____CAS-III-A_RT-2-1-csm3g7-csx19-csm3g7-csm3g7-10-x-csm6-csx1:0.80211000)0.90:0.26751000)0.34:0.06159000)0.41:0.13074000,378719040_Gordonia_pol____CAS-I-Cu_2-1+4-csb2-csb1-8-3:0.70293000)0.27:0.09076000)0.90:0.28490000,((162457463_Sorangium_cel____CAS-I-C_6-3-8c-7-5-1-2:0.58650000,347756304_Chloracidobacterium_the____CAS-I-C_6-3-8c-7-5-1+4-2:0.47759000)0.36:0.08796000,24215881_Leptospira_int____CAS-I-C_1+4-2-x-x-6-3-8c-7-5:0.48595000)0.92:0.30625000)0.86:0.18671000)0.74:0.07975000,337286710_Thermodesulfatator_ind____CAS-III-B_csx1-cmr6g7-cmr5SS-cmr4g7-x-x-cmr3g5-10-cmr1g7-x-2-x-1-2-6:0.72111000)0.83:0.20250000)0.69:0.06791000)0.69:0.05198000)0.91:0.17860000)0.78:0.06280000,(((427708212_Nostoc_PCC____CAS-III-A_csm3g7-csx19-csm3g7-x-x-x-6-x-x-2-1-x-x-1+RT:0.48505000,((433631662_Mycobacterium_can____CAS-I-C_2-1-4-7-8c-5-3:0.46431000,86742033_Frankia_CcI____CAS-I-C_3-5-8c-7-4-1-2:0.59968000)0.25:0.10391000,((189485224_uncultured_Ter____CAS-I-C_3-5-8c-7-4-1-2-9-1-2:0.23678000,108759816_Myxococcus_xan____CAS-I-C_2-1-4-7-8c-5-3HD:0.66151000)0.76:0.28677000,15612905_Bacillus_hal____CAS-I-C_cmr5SS-cmr6g7-x-x-x-3-5-8c-7-4-1-2:0.12115000)0.98:0.52307000)0.86:0.20392000,261415949_Fibrobacter_suc____CAS-III-A_1-2-x-x-csx1-10-csm3g7-x-csm3g7-csx20:0.98191000)0.83:0.13763000)0.78:0.05780000,((297622674_Truepera_rad____CAS-III-U_2-1-csx1-csx1-6:0.43915000,344996858_Caldicellulosiruptor_lac____CAS-I-D_2-1-4-6-3-csc1g5-csc2g7-10d:0.72375000)0.71:0.35642000,(17227878_Nostoc_PCC____CAS-III-B_csx18-1-2:0.64378000,326793968_Marinomonas_med____CAS-III-B_2-6-x-x-cmr6g7-cmr5SS-cmr4g7-cmr3g5-10-cmr1g7:0.64153000)0.70:0.19291000)0.62:0.01818000)0.85:0.06567000,((38505761_Synechocystis_PCC____CAS-III-B_cmr6g7-cmr5SS-cmr4g7-x-cmr3g5-10-csx18-1-2:0.33189000,((389875623_Tistrella_mob____CAS-III-A_10-cmr4g7-csm3g7-csm3g7-6-x-1+RT:2:0.59765000,((302343125_Desulfarculus_baa____CAS-III-A_csm3g7-csm3g7-csm3g7-10-1+RT-2-6:0.43962000,294676822_Rhodobacter_cap____CAS-III-A_RT-x-x-x-x-10-csm3g7-csm3g7-2-1+RT-RT-6:0.18051000)0.66:0.10955000,288957882_Azospirillum_B51____CAS-III-A_csm3g7-csm3g7-csm3g7-6-2-1+RT:0.07997000)0.13:0.11882000)0.87:0.18367000,374998938_Azospirillum_lip____CAS-III-B_2-1+RT-x-csm3g7-x-x-x-cmr1g7-10-cmr3g5-cmr4g7-10:0.10149000)0.95:0.45546000,220934971_Thioalkalivibrio_sul____CAS-III-B_10-cmr3g5-cmr4g7-cmr5SS-cmr6g7-csx1-x-csx1-x-1-2-x-csx1:0.86008000)0.88:0.31193000)0.84:0.21401000,((148657119_Roseiflexus_RS____CAS-III-U_2-x-1-RT-x-x-x-csx15-csx1:0.64747000,159898908_Herpetosiphon_aur____unknown_RT-1-2:0.45464000)0.67:0.06560000,((38505685_Synechocystis_PCC____CAS-I-D_3-10d-csc2g7-csc1g5-6-4-1-2:0.70582000,46255266_Thermus_the____CAS-I-A_2-1-4-x-8a-7-5-6:0.80808000)0.34:0.16455000,218442810_Cyanothece_PCC____CAS-III-A_csm6-x-x-RT-1-2:0.60968000)0.55:0.12524000)0.88:0.21577000)0.69:0.03635000)0.73:0.03037000,37677206_Vibrio_vul____CAS-III_6-csm3g7-x-csm3g7-5g10-cmr4g7-10-x-x-1+RT-2-2-csx1-csx16-csx6-csm6:0.81769000)0.32:0.05358000)0.72:0.05232000)0.82:0.11565000)0.15:0.12021000,(((30248143_Nitrosomonas_eur____CAS-III-A_2-1-2-csx1-x-csx1-csx16-6-x-csm5g7-csm4g5-csm3g7-csm2SS:0.47226000,206889468_Thermodesulfovibrio_yel____CAS-III-A_10-csm2SS-csm3g7-csm4g5-csm5g7-6-x-x-x-2-1-2:0.63151000)0.20:0.09160000,(430760779_Thioalkalivibrio_nit____CAS-III-B_cmr3g5-cmr4g7-cmr5SS-cmr6g7-csx1-x-csx1-x-x-6-x-x-2-1-2:0.42728000,121997422_Halarhodospira_hal____CAS-III-B_1-2-1-2-csx1-csx16-csx1-cmr1g7-10-cmr3g5-cmr4g7-cmr5SS-cmr6g7-6:0.48149000)0.23:0.15776000,292492855_Nitrosococcus_hal____CAS-III-A_csm3g7-csm4g5-csm5g7-x-csx1-x-csx1-x-x-csx1-csx1-x-2-1-x-2:0.23880000)0.94:0.26331000)0.81:0.16973000,52425691_Mannheimia_suc____CAS-III-U_2-1-2-x-csx1-csx1-csx16-x-6:1.01735000)0.74:0.20008000,(288947691_Allochromatium_vin____CAS-III-U_cmr6g7-x-x-x-x-csx1-x-csx16-x-2-2-1-RT:0.34180000)0.97:0.82237000)0.89:0.58176000)0.50:0.14925000,328949007_Treponema_suc____CAS-III-A_csx1-x-x-2-2-1-x-x-csx20-csx1-x-csm5g7-csm4g5-csm3g7:0.94210000)0.64:0.06485000)0.83:0.09917000,((((385809791_Ignavibacterium_alb____CAS-III-U_csx1-csx20-csx1-csx1-x-x-x-1-2:0.40745000,150021525_Thermosiphon_mel____CAS-III-B_2-1-2-x-x-csm6-cmr1g7-10-cmr3g5-cmr4g7-cmr5SS-cmr6g7-0.72705000)0.35:0.23605000,268317184_Rhodothermus_mar____CAS-III-A_2-1-x-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10-4-3:0.47741000)0.76:0.10019000,(((289596693_Aciduliprofundum_boo____CAS-III-A_2-1-4-6-csx1-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10:0.27366000,33359262_Pyrococcus_hor____CAS-I-B_csm2SS-csm3g7-csm4g5-csm5g7-csm6-8b-7-5-x-1-2-4-3-

csx1:0.59521000)0.88:0.14079000,(33359542_Pyrococcus_fur____CAS-I-B_2-1-4-3-5-7-8b-cmr6g7-cmr5SS-cmr4g7-csm6:0.30594000,((41614812_Nanoarchaeum_equ____CAS-I-B_2-1-x-7-5-3-4:0.61326000,(48477074_Picrophilus_tor____CAS-I-D_2-1-4-3-csc1g5-csc2g7-10d-6:0.64411000,154174506_Campylobacter_cur____CAS-I-B_1-2-4-3-5-7-8-6:0.68988000)0.77:0.19975000)0.87:0.23894000,13540936_Thermoplasma_vol____CAS-III-A_2-1-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10-6-csm6:0.34232000)0.84:0.15228000,(84489237_Methanospaera_sta____CAS-I-B_6-8-7-5-3-4-1-2:0.35892000,150021527_Thermosiphon_mel____CAS-III-B_2-1-2-x-x-x-csm6-cmr1g7-10-cmr3g5-cmr4g7-cmr5SS-cmr6g7:0.44479000)0.80:0.08895000)0.93:0.15361000)0.11:0.00015000)0.85:0.11140000,(111224948_Frankia_aln____CAS-I-B_6-x-7-3+5-4-1-2:0.86083000,55376279_Haloarcula_mar____CAS-I-B_6-8-7-5-3-4-1-2:0.50539000)0.83:0.21709000)0.72:0.03878000,389844442_Mesotoga_pri____CAS-I-B_csm6-csm6+x-x-cmr6g7-cmr5SS-cmr4g7-cmr3g5-10-cmr1g7-csm6-2-x-x-1-4-3-5-7:0.45847000)0.82:0.17195000)0.68:0.19571000,390962115_Thermococcus_CL1____CAS-III-A_csx1-x-6-10-csm2SS-csm3g7-csm4g5-csm5g7-R2-csx3-2-1:0.80322000)0.79:0.16255000,292492939_Nitrosococcus_hal____CAS-I-D_6-10d-csc2g7-x-3-4-1-2-1:1.14074000)0.86:0.32832000)0.90:0.17979000,((408403001_Nitrosospaera_gar____CAS-I_4-1-2-4-x-x-x-3-5-7-x-6:0.58554000,(18311771_Pyrobaculum_aer____CAS-I-A_6-4-2-1-4-csa3-x-8a2-3HD-3-5-7-csa5SS:1.14033000,((126465228_Staphylothermus_mar____CAS-I-A_csm3g7-csm4g5-csm5g7-10-x-x-1-4-2-4-6-R-csa5SS-7-5-csa8a2-3:0.26154000,170290048_Korarchaeum_cry____CAS-I-A_1-4-2-4-R-csa5SS-7-5-csa8a2-x-3-3HD:0.40242000)0.86:0.17885000,(330509016_Methanosaeta_con____CAS-I-A_4-4-2-1-3-x-x-x-x-5:0.80370000,(148656005_Roseiflexus_RS____CAS-III-U_10-csx1-csx1-csx3-8b-7-5-3-6-1-2-4:1.02105000,386854796_Deinococcus_gob____CAS-I-A_4-2-1-5-7-8a2-3-6:1.37160000)0.38:0.04011000,167628643_Heliobacterium_mod____CAS-I-A_4-4-2-1-x-6-5-7-x-3:0.15876000)0.88:0.14551000)0.87:0.17843000)0.61:0.10408000)0.32:0.17182000)0.69:0.16217000,(((392395715_Flexibacter_lit____unkn_own_2-x-1:0.44069000,379728402_Saprospira_gra____unknown_2-x-1:0.77932000)0.43:0.26216000,327314605_Prevotella_den____CAS-III_10-cmr6g7-5g10-csm3g7-x-csm3g7-csx1-csx1-1-Prim-2:0.60272000)0.91:0.33715000,15898244_Sulfolobus_sol____CAS-I-A_csx1-csa5SS-7-5-csaXg8-3-3HD-2-1-6:0.58386000)0.60:0.05277000,332668018_Haliscomenobacter_hyd____unknown_RT-1-PDEDxK-2:0.97255000)0.80:0.16408000)0.66:0.05231000,(((284040852_Spirosoma_lin____CAS-I-B_8-7-5-3-6-1-2-4:0.51641000,218961576_Cloacamonas_aci____CAS-I-B_4-2-1-3HD-5-7-8-6:0.33190000)0.91:0.29536000,21226660_Methanosarcina_maz____CAS-I-B_4-2-1-6-3-5-7-8-csa3:0.27037000)0.00:0.11790000,150401498_Methanococcus_aeo____CAS-III-A_6-3-5-7-8-1-2-4-x-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10:0.70927000)0.64:0.07037000,257125861_Leptotrichia_buc____CAS-III_csm3g7-6-x-x-x-x-x-4-2-1-x-x-csx1-csx20:0.64294000)0.94:0.33287000,((34557934_Wolinella_suc____CAS-II-B_9-1-2-4:0.68476000,54296140 Legionella_pne____CAS-II-B_9-1-2-4:0.49844000)0.85:0.41634000,118497354_Francisella_nov____CAS-II-B_9-1-2-4:0.44155000)0.97:0.63521000,332661940_Haliscomenobacter_hyd____CAS-III-B_2-1-x-RT-cmr6g7-x-cmr4g7-cmr3g5-10:0.87623000)0.60:0.22102000,150398873_Methanococcus_van____CAS-III-B_10-cmr3g5-cmr4g7-cmr5SS-cmr6g7-x-x-6-1-2:0.74887000)0.46:0.29206000)0.64:0.14281000)0.68:0.06174000)0.81:0.17631000,((385805116_Fervidicoccus_fon____CAS-I-A_R-3HD-3-8a2-5-7-x-x-4-1-2-4-R-x-x-csm6-cmr6g7-cmr1g7-cmr5SS-cmr4g7:0.87975000,(((119720227_Thermofilum_pen____CAS-III-B_cmr3g5-10-cmr6g7-cmr1g7-cmr5SS-cmr4g7-1-2-3-csc1g5-csc2g7-x-10d:0.29880000,70607607_Sulfolobus_aci____CAS-I-D_3-csc1g5-csc2g7-10d-R-6-x-2-4-1:0.72201000)0.98:0.47456000,(347523052_Pyrolobus_fum____CAS-I-A_4-x-x-x-x-2-1-4-R-x-x-3HD-3-8a2-5-7:0.53973000,(305663329_Ignisphaera_agg____CAS-I-A_R-x-7-5-3-3HD-x-6-4-2-1-4:0.18600000,307595806_Vulcanisaeta_dis____CAS-I-A_6-3HD-3-x-5-7-x-R-x-4-2-1-4:0.43362000)0.50:0.11061000,(118431379_Aeropyrum_per____CAS-I-A_6-8a2-3HD-3-5-7-csa5SS-R-4-2-1-4:0.27746000,156937922_Ignicoccus_hos____CAS-I-A_4-1-2-4-3HD-3-x-5-7-csa5SS-R:0.70425000)0.69:0.22932000)0.44:0.18671000)0.91:0.29882000)0.58:0.04873000,124027532_Hyperthermus_but____CAS-I-A_3-3-csaXg8-5-7-csa5SS-R-x-x-4-2-4-1-x-x-x-R:0.62624000)0.91:0.28271000,302347807_Acidilobus_sac____CAS-I-A_6-x-4-2-1-4-cmr3g5-10-cmr6g7-csm3g7-cmr5SS-cmr4g7-csm6-x-csm6:0.54554000)0.84:0.27918000)0.20:0.13179000,15922976_Sulfolobus_tok____CAS-I-A_R-4-1-2-4-x-x-csa5SS-7-5-3-3HD-csaXg8:0.41222000)0.25:0.18822000,218884092_Desulfurococcus_kam____CAS-I-A_6-3HD-3-csaXg8-5-7-csa5SS-R-4-2-1-4-R-x-cmr3g5-10-cmr6g7-cmr1g7-cmr5SS-cmr4g7:0.81926000)0.92:0.64207000)0.74:0.09009000,(288560432_Methanobrevibacter_rum____CAS-III-A_1-x-2-6-csx1-csm5g7-csm4g5-csm3g7-csm2SS-10:0.92672000,(11499460_Archaeoglobus_ful____CAS-I-A_cmr1g7-csa3-csa5SS-7-5-csa8a2-3-3HD-2-4-1-4:0.35880000,383319614_Methanocella_con____CAS-I-A_4-1-4-2-csa3-x-x-x-x-7-x-x-3:0.24133000)0.96:0.43829000)0.62:0.05416000)0.84:0.21065000)0.78:0.21994000,18311698_Pyrobaculum_aer____CAS-I-A_3HD-3-x-x-6-csa3-x-4-2-1-4:1.11375000)0.67:0.07248000,((50085558_Acinetobacter_AD____CAS-I-F_3f-x-8f-5f-7f-6f-x-x-1-3f:0.71605000)0.71:0.25381000)1.00:2.48456000,((90111482_Escherichia_col____CAS-I-E_2-1-6e-5-7-cse2SS-8e-3:0.19505000,77128445_Methylococcus_cap____CAS-I-E_2-1-6e-5-7-cse2SS-8e-3:1.08019000)0.71:0.19962000,21674789_Chlorobium_tep____CAS-I-E_3-8e-cse2SS-6e-7-5-1-2:1.33635000)0.81:0.25236000,(209966591_Rhodospirillum_cen____CAS-I-E_2-1-6e-5-7-cse2SS-8e-3:1.03342000,(86738747_Frankia_CcI____CAS-I-E_3-8e-cse2SS-7-5-6e-1-2:0.29430000,38234756_Corynebacterium_dip____CAS-I-E_5-7-cse2SS-8e-6e-3-1-2:1.16325000)0.85:0.31309000)0.72:0.13876000)0.81:1.00213000)0.76:0.43903000);

Cas4 Tree:

((170290047_Candidatus_Kor__CAS-I-A:0.32712000,126465227_Staphylothermus_mar__CAS-I-A:0.33457000)0.99:0.59491000,((408402999_Candidatus_Nit__CAS-I-A:0.46381000,(330509014_Methanosaeta_con__CAS-I-A:0.57219000,(11499463_Archaeoglobus_ful__CAS-I-A:0.37486000,383319613_Methanocella_con__CAS-I-A:0.37265000)0.99:0.34884000)0.65:0.14787000)0.92:0.15885000,167628645_Heliobacterium_mod__CAS-I-A:0.38142000)0.98:0.30697000,((385805114_Fervidicoccus_fon__CAS-I-A:0.87288000,(218884094_Desulfurococcus_kam__CAS-I-A:0.32294000,389861364_Thermogladius_cel__CAS-I-A:0.20434000)0.51:0.12318000)0.92:0.21199000,((15898287_Sulfolobus_sol__CAS-I-A:0.39623000,15922974_Sulfolobus_tok__CAS-I-A:0.37770000)0.96:0.21913000,((18311700_Pyrobaculum_aer__CAS-I-A:0.81016000,347523054_Pyrolobus_fum__CAS-I-A:0.43936000)0.20:0.06803000,18311773_Pyrobaculum_aer__CAS-I-A:0.72574000)0.77:0.12556000)0.60:0.03377000)0.71:0.16243000,((307595808_Vulcanisaeta_dis__CAS-I-A:0.21634000,124027533_Hyperthermus_but__CAS-I-A:0.57735000)0.86:0.10869000,(118431381_Aeropyrum_per__CAS-I-A:0.41512000,305663331_Ignisphaera_agg__CAS-I-A:0.21018000)0.90:0.10812000)0.92:0.19835000,156937920_Ignicoccus_hos__CAS-I-A:0.23716000)1.00:0.68066000)0.23:0.03755000)0.78:0.09126000,(((111224946_Frankia_aln__CAS-I-B:1.13081000,((((84489235_Methanospaera_sta__CAS-I-B:0.56652000,(14590111_Pyrococcus_hor__CAS-I-B:0.27203000,55376277_Haloarcula_mar__CAS-I-B:0.96864000)0.88:0.20644000)0.80:0.17828000,(389844446_Mesotoga_pri__CAS-I-B:0.83994000,41614818_Nanoarchaeum_equ__CAS-I-B:0.35613000)0.97:0.39504000)0.09:0.06437000,(289596695_Aciduliprofundum_boo__CAS-III-A:0.79798000,(268317193_Rhodothermus_mar__CAS-I-A:1.07613000,(154174706_Campylobacter_cur__CAS-I-B:0.73180000,28211140_Clostridium_tet__CAS-I-B:0.51649000)0.92:0.28583000)0.36:0.15318000)0.71:0.11810000)0.92:0.30744000,18977491_Pyrococcus_fur__CAS-I-B:0.40489000)0.57:0.16692000,(48477076_Picrophilus_tor__CAS-I-D:0.57967000,292492937_Nitrosococcus_hal__CAS-I-D:2.06459000)0.80:0.65148000)0.91:0.44422000)0.99:0.77832000,((((((159898754_Herpetosiphon_aur__CAS-I-D:0.42938000,(38505683_Synechocystis_sp__CAS-I-D:0.60781000,46255268_Thermus_the__CAS-I-A:0.79859000)0.87:0.17858000)0.86:0.16668000,((86742031_Frankia_CcI__CAS-I-C:0.80886000,433631664_Mycobacterium_can__CAS-I-C:0.72421000)0.95:0.40152000,((189485222_uncultured_Ter__CAS-I-C:0.69171000,56965355_Bacillus_hal__check:0.50337000)0.86:0.25583000,108757432_Myxococcus_xan__CAS-I-C:0.55447000)0.02:0.08514000)0.98:0.37358000,(302391601_Acetohalobium_ara__CAS-I-D:0.50266000,17229059_Nostoc_sp__CAS-I-D:0.66094000)0.17:0.14424000)0.77:0.10147000)0.49:0.10657000,344996860_Caldicellulosiruptor_lac__CAS-I-D:1.17966000)0.54:0.03556000,134299482_Desulfotomaculum_red__CAS-III-A:0.72503000)0.83:0.09076000,((((39995168_Geobacter_sul__CAS-I-Cu:0.18797000,83591519_Rhodospirillum_rub__CAS-I-Cu:0.67734000)0.91:0.14642000,320102061_Isosphaera_pal__CAS-I-Cu:0.29118000)0.59:0.09049000,378719041_Gordonia_pol__CAS-I-Cu:0.76296000)0.86:0.12732000,268318441_Rhodothermus_mar__CAS-I-C:0.35064000)0.86:0.14406000,134101617_Saccharopolyspora_ery__CAS-III-A:0.92088000)0.97:0.30155000)0.76:0.10722000,(294828260_Leptospira_int__CAS-I-C:0.34485000,347756303_Candidatus_ChI__CAS-I-C:0.62198000)1.00:0.82703000)0.79:0.16940000,76802275_Natronomonas_phu__CAS-I-D:1.19261000)0.84:0.23501000,167627879_Francisella_nov__check:2.53097000,219683006_Bifidobacterium_ani__CAS-I-Cu:1.03586000:0.60263000)0.12:0.15433000)0.19:0.04486000,(332296539_Thermodesulfobium_nar__CAS-III-A:2.10154000,(18311697_Pyrobaculum_aer__CAS-I-A:1.34349000,15896973_Sulfolobus_sol__check:2.11406000)0.32:0.26671000)0.02:0.20274000)0.43:0.05955000,((18311770_Pyrobaculum_aer__CAS-I-A:1.04920000,(124027531_Hyperthermus_but__CAS-I-A:1.08384000,(156937923_Ignicoccus_hos__CAS-I-A:1.64041000,118431378_Aeropyrum_per__CAS-I-A:0.62763000)0.73:0.12480000,(305663328_Ignisphaera_agg__CAS-I-A:0.64154000,307595805_Vulcanisaeta_dis__CAS-I-A:0.34920000)0.96:0.35574000)0.14:0.05870000)0.24:0.21397000)0.80:0.16355000,(385805117_Fervidicoccus_fon__CAS-I-A:1.48413000,((218884091_Desulfurococcus_kam__CAS-I-A:0.89263000,389861366_Thermogladius_cel__CAS-I-A:0.52075000)0.92:0.41883000,347523047_Pyrolobus_fum__CAS-I-A:1.69446000)0.60:0.04774000,(302347806_Acidilobus_sac__CAS-I-A:1.01638000,(15898285_Sulfolobus_sol__CAS-I-A:0.78934000,15922977_Sulfolobus_tok__CAS-I-A:0.61606000)0.94:0.36412000)0.85:0.25630000)0.84:0.12943000)0.90:0.22262000)0.75:0.19330000,((((284040853_Spirosoma_lin__CAS-I-B:0.63299000,21226659_Methanosparsina_maz__CAS-I-B:0.36828000)0.01:0.10159000,(218961575_Candidatus_Clo__CAS-I-B:0.52444000,150401499_Methanococcus_aeo__CAS-I-B:0.46347000)0.16:0.01868000)0.89:0.24404000,257125860_Leptotrichia_buc__CAS-III-A:1.09081000)0.97:0.41375000,(383319611_Methanocella_con__CAS-I-A:0.66135000,11499461_Archaeoglobus_ful__CAS-I-A:0.51463000)0.97:0.34472000)0.89:0.14938000,(330509015_Methanosaeta_con__CAS-I-A:0.50473000,408403002_Candidatus_Nit__CAS-I-B:0.92921000)0.89:0.21520000)0.72:0.06975000,(((148656006_Roseiflexus_sp__CAS-I-B:0.57761000,386854795_Deinococcus_gob__CAS-I-A:0.79274000)0.84:0.17086000,(54296141 Legionella_pne__CAS-II-B:0.54352000,34557935_Wolinella_suc__CAS-II-B:0.66238000)0.97:0.37315000,118497355_Francisella_nov__CAS-II-B:0.54478000)0.98:0.50460000)0.41:0.18627000,167628644_Heliobacterium_mod__CAS-I-A:0.76117000)0.84:0.18847000)0.85:0.14480000,(170290049_Candidatus_Kor__CAS-I-A:0.72124000,126465229_Staphylothermus_mar__CAS-I-A:0.38343000)1.00:0.75378000)0.80:0.11271000)0.89:0.21765000)1.00:0.90558000);

Supplementary Figure S3. Multiple sequence alignments of type I and type III CRISPR repeats. See Figure 3. The alignments of short (24-26 nt) type I-A (**A**) and long (37 nt) type I and type III (**B**) CRISPR repeats used to create the logplots of consensus repeat sequences are shown.

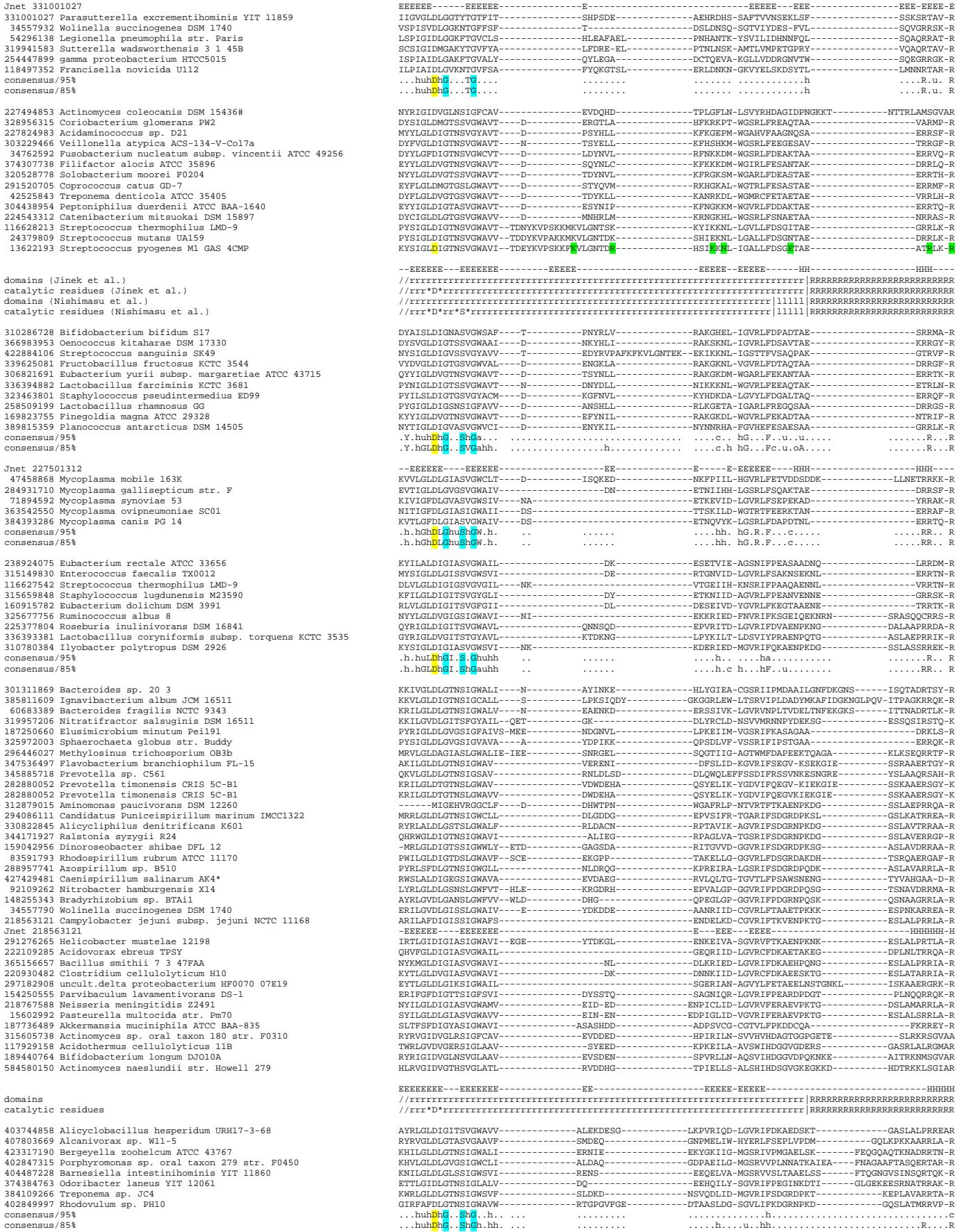
A

<i>Pyrolobus fumarii</i> 1A	GAAACAAC-CAAGAATGAATTGAAAG
<i>Ignisphaera aggregans</i> DSM 17230	GAAAC-TCCAAAAGGGAGTAGAAAG
<i>Hyperthermus butylicus</i> DSM 5456	GAACA-ACTCAAAGAGAATTGCAAG
<i>Thermogladius cellulolyticus</i> 1633	GAAGCAA-TCAAGAA-GAATTGAAAG
<i>Desulfurococcus kamchatkensis</i> 1221n	GAATA--CAATAGAAAGAATTGAAAG
<i>Pyrobaculum aerophilum</i> str. IM2	GAA-TCT-CAAAAAGAGGATTGAAAG
<i>Fervidicoccus fontis</i> Kam940	GAA-TCTGTT-AGATAGAATTGAAAG
<i>Sulfolobus tokodaii</i> str. 7	GATAA-TCCTTAATG-GAATTGAAAG
<i>Sulfolobus solfataricus</i> P2	GATAA-TCTCTTAT-AGAATTGAAAG
<i>Vulcanisaeta distributa</i> DSM 14429	GATAT-TCTCTAAAGAGAATAGAAGG
<i>Sulfolobus solfataricus</i> P2	GATTAATCCAAAAG-GAATTGAAAG
<i>Aeropyrum pernix</i> K1	GCATA-TCCCTAAAGGGAATAGAAG
<i>Ignicoccus hospitalis</i> KIN4/I	GTGAAGTCTCAAAGAGGATAGAAAG
<i>Acidilobus saccharovorans</i> 345-15	GTTGCAACACCATCTCGTGGTTCC-
<i>Pyrobaculum aerophilum</i> str. IM2	GTTTCAACTATTTGATTCTG-G

B

<i>Desulfobacca acetoxidans</i> DSM 11109	GTCGCAATCCCTCGTAATCAGGTCAATGAAGCAGAG
<i>Porphyromonas gingivalis</i> W83	GTCTTAATAGCCTTACGGACTGTGTATGTAGTGAG
<i>Helicobacter cetorum</i> MIT 00-7128	GTTTCAATCCCCTTGAACGGGTCAATCTGTAATAAT
<i>Helio bacterium modesticaldum</i> Ice1	GTTTCACTGGATGACCGCTATGTAGGGATTGAAAG
<i>Roseiflexus</i> sp. RS-1	GTTTCAGTACTCTTCAACGAGTCGAAATTGCCGAAAT
<i>Deinococcus gobiensis</i> I-0	GTTTCAGTCCCCTATGCCCGGGAGTACGTTGCGGC

Supplementary Figure S4. Multiple alignment of Cas9 sequences divided into 5 groups according to tree topology and operon organization (Please see Supplementary Materials and Methods for details); Sequences are denoted by Genbank identifiers and complete organism name on the first page and further by abbreviated organism name. Color bars on the left group sequences according to the Cas9 tree topology (compare with Figure 4). *S. pyogenes* and *A. naeslundii* Cas9 strucutres with domain architecture are mapped on the alignment. r - RuvC motifs, I - loops, R - Arginine-rich motif, N - HNH domain, h - helical lobe, 1 - REC1 recognition lobe domain, 2 - REC2 recognition lobe domain, P - PAM recognition motifs. Functional and conserved amino acids are highlighted: yellow - catalytic, green - RNA-binding, red - target DNA binding, blue - conserved or semi-conserved (see Figure 2). Consensus is predicted using the "Consensus" online tool (<http://coot.embl.de/Alignment/consensus.html>). Map of conserved positions used for the tree reconstruction is shown below each alignment block.



#map

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#map

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#map ----- ***

#map *****

#map -----

#map ----- *----- *----- *----- *----- *----- *----- *-----

#map *-----****-----****-----

#map

www.english-test.net

Jnet 331001027
P. excrementiphilominis
W. succinogenes
L. pneumophila
HTCC5015
F. novicida
consensus/95%
consensus/85%

A. coleocanis
C. glomerans
Acidaminococcus D21
V. atypica
F. nucleatum
F. alocis
S. moorei
C. catus
T. denticola
P. duerdenii
C. mitsuokai
S. thermophilus
S. mutans
S. pyogenes

domains
catalytic residues
domains
catalytic residues
.
B. bifidum
O. kitaharae
S. sanguinis
F. fructosus
E. yurii
L. farcininis
S. pseudintermedius
L. rhamnosus
F. magna
P. antarcticus
consensus/95%
consensus/85%
.
Jnet 227501312
M. mobile
M. gallisepticum
M. synoviae
M. ovipneumoniae
M. canis
consensus/95%
consensus/85%
.
E. rectale
E. faecalis
S. thermophilus
S. lugdunensis
E. dolichum
R. albus
R. inulinivorans
L. coryniformis
I. polystropus
consensus/95%
consensus/85%
Bacteroides sp. 20
I. album
B. fragilis
N. salugrinis
E. minutum
S. globus
M. trichosporium
F. branchiophilum
Prevotella C561
P. timonensis
P. timonensis
A. paucivorans
P. marinum
A. denitrificans
R. syzygii
D. shibae
R. rubrum
Azospirillum B510
C. salinarum
N. hamburgensis
Bradyrhizobium BTAl1
W. succinogenes
C. jejuni
Jnet 218563121
H. mustelae
A. ebreus
B. smithii
C. cellulolyticum
HF0070_07E19
P. lavamentivorans
N. meningitidis
P. multocida
A. muciniphila
Actinomyces F0310
A. cellulolyticus
B. longum
A. naeslundii

domains
catalytic residues
.
A. hesperidum
Alcanivorax W11-5
zoohelcum
Porphyromonas F0450
B. intestinalisomnis
O. laevis
Treponema JC4
Rhodovulum PH10
consensus/95%
consensus/85%

#map

Jnet 33100102
P. excrementum
W. succinogenes
L. pneumophila
S. wadsworthensis
HTCC5015
F. novicida
consensus/95%
consensus/85%

A. coleocanis
C. glomerans
Acidaminococcus D21
V. atypica
F. nucleatum
F. alocis
S. moorei
C. catus
T. denticola
P. duerdenii
C. mitsuokai
S. thermophilus
S. mutans
S. pyogenes

domains
catalytic residues
domains
catalytic residues
.
B. bifidum
O. kitaharae
S. sanguinis
F. fructosus
E. yurii
L. farciminis
S. pseudintermedius
L. rhamnosus
F. magna
P. antarcticus
consensus/95%
consensus/85%
.
Jnet 227501312
M. mobile
M. gallisepticum
M. synoviae
M. ovipneumoniae
M. canis
consensus/95%
consensus/85%

E. rectale
E. faecalis
S. thermophilus
S. lugdunensis
E. dolichum
R. albus
R. inulinivorans
L. coryniformis
I. polytropus
consensus/95%
consensus/85%

Bacteroides sp.20
I. album
B. fragilis
N. salugrinis
E. minutum
S. globus
M. trichosporium
P. branchiophilum
Prevotella C561
P. timonensis
P. timonensis
A. paucivorans
P. marinum
A. denitrificans
R. syzygii
D. shibae
R. rubrum
Azospirillum B510
C. salinarum
N. hamburgensis
Bradyrhizobium BTa1
W. succinogenes
C. jejuni
Jnet 218563121
H. mustelae
A. ebreus
B. smithii
C. cellulolyticum
HF0070_07E19
P. lavamentivorans
N. meningitidis
P. multocida
A. muciniphila
Actinomyces F010
A. cellulolyticus
B. longum
A. naeslundii

domains
catalytic residues
.
A. hesperidum
Alcanivorax W11-5
Zoohelcum
Porphyromonas F0450
B. intestinalis
O. laevis
Treponema JC4
Rhodovulum PH10
consensus/95%
consensus/85%

#map

#map -----*****-----

#map

#map

```
#map -----
```

Philip Morris Inc. / Philip Morris Products S.A. / Philip Morris Inc. / Philip Morris Inc. / Philip Morris Inc.

Jnet 331001027
P. excrementiphominis --EE-----EEEEE----HHHHHHHHHHHHH---
W. succinogenes --GTI--FFELVG--PRVIFNYIVGGAASSLKEIFSBACKER
L. pneumophila --VKI--QLRTDGSISNIVURKNAADPTTLSFSEHIQKLK-
S. wadsworthensis --CRM--KIVSTG--KIVTEFEFSDDSTPQWIQTLYVTQLKK-
 HTCC5015 --SEI--FIENVGNAKHIFRFWYIVVSSNKKMNESYNNVSKS-
F. novicida --SNL--FILETG--QKLTIEFANGANAEVKAYSLRKA--
consensus/95% --KVL--EILK---QSTIIEFESSGFNKTIKEMLGMKLAGI
consensus/85% .h.....h.....h.....h.....h.....h.....

A. coleocanis AMPEVIRRDGLGR-KRKFPSYSGLPVSWQG-----
C. glomerans NVHLI-DQSVTGM-FERRTKIGL-----
Acidaminococcus D21 DIRLV-EQSASGL-FVSKSQNLLEYL-----
V. atypica EFKII-NESITGL-YSNEVITIV-----
F. nucleatum NNNKELEESVTGL-FVKKIKL-----
F. alocis KILLV-NQSVTGL-YENRREL-----
S. moorei SVFKVIHQSIITGL-FEKEMDLLK-----
C. catus KQISVINQSPSTGI-YKEEIDLKL-----
T. denticola DCNLLHQSIITGL-FEKRIDLLKV-----
P. duerdenii KECKLINSQSIITGL-FENEVDLLNL-----
C. mitsukai NNNVPIQSOPSTGI-YTKVYKL-----
S. thermophilus KDATLIHQSVTGL-YETRIDLAKLGEG-----
S. mutans LNATLHQSIITGL-YETRIDLNLKGGD-----
S. pyogenes LDATLHQSIITGL-BEIRDLSQLGGD-----

 domains -----EEEE-----
 catalytic residues ccccccccccccccccccccccccccccccccccccc|
 domains ccc|
 catalytic residues PPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPP|
 PPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPPP|

B. bifidum DEDEPIFOSPSGL-FEKRTVTGELKKAKKEVNSKYRTNEK
O. kitaharae DTDEP IPQSVTGL-FSTQKTVQALYQETK-----
E. sanguinis LNATLHQSIITGL-YETRIDLNLKGED-----
F. fructosus DNSEMIYQSATGI-FETRKVKISDL-----
E. yurii KEFKLHQSIITGL-YSCEDKDLMTI-----
L. farciminis NNDFIIPQSVTGL-YETRIHIE-----
S. pseudintermedius KDWKIAUTSISGLKTTPKSLFKLAESRNEL-----
L. rhamnosus KDTQIVVQSPSGL-FKRRIPLAIDL-----
F. magna DNTYLVYKSITGI-YEKRRTRIK-----
P. antarcticus EEAIVGYESITGLKYRKPRSVVGTKR-----
consensus/95% ...h...S...oh.....
consensus/85% ...h...S...oh.a...h.....

.Jnet 227501312 --EEEEE-----EEEEE-----
M. mobile -WKV-NLTYMGF-IFKK-----
M. gallisepticum LLDADKDNFDILGL-SKNRIDEILNSKLGLDKIVK-----
M. synoviae EYKIIIEHVNPDLGI-NLNWTKKLENNN-----
M. ovipneumoniae EYIPI-HLDELGN-EYFVKIKEHTDDEKLMTIK-----
M. canis NYEQV-DISPGFI-NKKIFE-----
consensus/95% .h....h..h...
consensus/85% .h....h..h...

E. rectale FIKKY-RYDILGN-KYSCSEEKFTSFC-----
E. faecalis KLKP-FNDILGK-KHVLFYKEKEPKNIK-----
S. thermophilus SIVK-YRTDVGLN-QHIKNEGDKPKLDF-----
S. lugdunensis SIEKL-TTDVGLN-VFTNTQYTKEPQLLFKRGN-----
E. dolichum RIDKY-ATDVGN-LYKVVKNTLKFEFD-----
R. albus NGKGISGEPLSL-LKEKN-----
R. inulinivorans IFEKC-QVDILGN-ISVVRHENRQEFP-----
L. coryniformis KFKY-QVDVGLN-YHKVHEKKRQLFVKRDE-----
I. polytropus NIKY-QVDVGLN-YVKVRERQTF-----
consensus/95%hu.....
consensus/85% .hKh...Dh#

Bacteroides sp. 20 ICFFPI-KVDRGLN-IIQVGSCILTN-----
I. album VCIKI-RVDRGLN-VRPL-----
B. fragilis TIVKV-RVNHIGQ-IVSVEGY-----
N. salugrinis RSEVFGTBLBEGS-IEWFKEESGYGRVEDDPH-----
E. minutus NPKRL-ILSSCGF-IKYRSPILKNGE-----
S. globus DVRKV-QLSSAGL-VRVSPPLLVDKIEDEVALCGE-----
M. trichosporium EFKPF-NVDRFGR-KHEVERELRTWRGETWRKGAYI-----
F. branchiophilum KMHNH-QFTAFVEGIDFKVLPSPGK--FEKI-----
Prevotella C561 SVKSL-NFLIMGENRDFVIKN-GKIIFNHR-----
P. timonensis SINNI-NILVDGI--DFTIDILGKVTLKE-----
P. timonensis SINNI-NILVDGI--DFTIDILGKVTLKE-----
A. paucivorans SMSKY-QVDPLGE-IRLVGSEKPPFVL-----
P. marinum SARVK-HISPTGL-IREG-----
A. denitrificans KTRRV-TIISPIGE-VRDPGFKG-----
R. syzygi GARPI-FVDSLIGR-VLDPGIGKGD-----
D. shibae GIRRV-SVDEIGR-LRDGGTRPI-----
R. rubrum KIRT-TCTALGR-LRLSKKAT-----
Azospirillum B510 GARPV-TVDPGLR-VRVHAPGARVGIGGDAGRTPAEDI
C. salinarum VAKGD-TVKKDGKTYKVGV--ITQGIFGMPVDAAGSAPRTP
N. hamburgensis AAVPV-RVDELGR-WVWMPN-----
Bradyrhizobium BTail NAEVL-RIDTLQG-PWRRKRGLETGSEDATRIGWTRPKWP
M. succinogenes DVKRY-QIDPLGY-YYEVQKEKRLGTIPQMRMSAKLKVKK--
C. jejuni VFKEY-IVSALGE-VTKAEFRQRREDPKK--
 Jnet 218563121 EEEE-----EEE-----
H. mustelae VFQKV-KLSVLGL-ELEHKPDRNRQNTALKTTPKHV-----
A. ebreus SVEPKV-NVDVLGR-IYLAAPPETRSGLA-----
B. smithii RFKEY-QVDVGLN-IHKVKGKERVGLAAPTQOKKGTVDSDL
C. cellulolyticum SIEKY-QNDLGN-KSIVKGEPRRGMEKYNSFKSN-----
 HFPAKV-QVNLGK-VIK-----
 HFPAKV-QVNLGK-VIK-----
 HFPAKV-QVNLGK-VIK-----
P. lavantlivorans DAKKV-SIDFLIGR-VREPSND-----
N. meningitidis SFQKY-QIDELGK-EIRPCRLKRPBPVR-----
P. multocida SPEKY-QVDELGK-NRQICRPQQRPQVR-----
A. muciniphila NWREVDLISLLKEK-YQMKRYPSTSQTGPR-----
Actinomycetes F0310 GLIRVIRRNALGE-VRTSPKSGLPISLNLR-----
A. cellulolyticus GLTVI-RRTALQG-PRWRRGHLPIWSWRPWSDAPWSGTP-----
B. longum AVRIV-RRNACGE-PRLSSAHHMPCSWQRHIE-----
A. naeslundii HPTVVV-RRDALGR-PRYSSRSRNLPSTSWTIE-----

 domains -----EE-----
 catalytic residues ccccccccccccccccccccccccccccccccc|
 ccccccccccccccccccccccccccccccccccccc|

A. hesperidum VFEKY-VVGPLGD-THPVYKERRMPFVERKMN-----
Alcanivorax W11-5 KPADL-LSAKWGR-LKVGGRNIHLLRLCAE-----
B. zoothelcum NCFLK-EIDRLGN-IVKVKIR-----
Porphyromonas F0450 NIRKV-RVDDLGLR-ISL-----
B. intestiniphominis NPHKV-HISVLGE-IKEIS-----
O. laneus NPVKV-QIDELGK-ITFLNGPLC-----
Treponema JC4 KARLV-TVSPIGR-VFVK-----
Rhodovulum PH10 GCVAV-RDPIGV-VTLRRRSNV-----
consensus/95%
consensus/85% ...h...h...

#map -----*****-----

Supplementary Figure S5. Multiple alignment of Cas1 sequences associated with type II CRISPR-Cas systems (Please see Supplementary Materials and Methods for details); Sequences are denoted by Genbank identifiers and complete organism name.

16130662 *Escherichia coli* str_K-12 substr_MG1655
 11500001_Aerobacterium_fulgidus_DSM_4304
 18977490 *Pyrococcus furiosus* DSM_3638
 46447796 *Desulfovibrio vulgaris* str_Hildenborough
 54296139 *Legionella pneumophila* str_Paris
 331001028 *Parasutterella exrementimominis* YIT_11859
 319941582 *Sutterella wadsworthensis* 3 1 45B
 118497353 *Francisella novicida* U112
 34557933 *Wolinella succinogenes* DSM_1740
 187736488 *Akkermansia muciniphila* ATCC_BAA-835
 407803668 *Alcanivorax* sp_W1-5
 325972002 *Sphaerocheta globus* str_Buddy
 187250661 *Elusimicrobium minutum* Peil91
 47458867 *Mycoplasma mobile*_163K
 363542551 *Mycoplasma ovipneumoniae* SC01
 71894593 *Mycoplasma synoviae* 53
 384393287 *Mycoplasma canis* PG_14
 284931711 *Mycoplasma gallisepticum* str_F
 117929157 *Acidothermus cellulolyticus* 11B
 317482065 *Bifidobacterium* sp_12_1_47BFAA
 315605739 *Actinomyces* sp_ oral taxon_180 str_F0310
 294086112 *Candidatus Puniceispirillum marinum* IMCC1322
 282880053 *Prevotella timonensis* CRIS_5C-B1
 374384762 *Odoribacter laneus* YIT_12061
 423317188 *Bergeyella zoohelcum* ATCC_43767
 385181160 *Ignivibacterium* album JCM_16511
 402847305 *Porphyromonas* sp_ oral taxon_279 str_F0450
 345885719 *Prevotella* sp_C561
 404487227 *Barnesiella intestinihominis* YIT_11860
 301311870 *Bacteroides* sp_20_3
 60683380 *Bacteroides fragilis* NCTC_9343
 42849996 *Rhodovulum* sp_PH10
 154250554 *Parvibaculum* lavamentivorans DS-1
 159042957 *Dinoroseobacter shibae* DFL_12
 288957740 *Azospirillum* sp_B510
 330822846 *Alicyphilobacter denitrificans* K601
 34171926 *Ralstonia syzygii* R24
 319957207 *Nitratiruptor saluginis* DSM_16511
 315149831 *Enterococcus faecalis* TX0012
 116627543 *Streptococcus thermophilus* LMD-9
 238924076 *Eubacterium rectale* ATCC_33656
 1218563120 *Campylobacter jejuni* subsp_jejuni NCTC_11168 - ATCC_700819
 291276264 *Helicobacter mustelae* 12198
 326577757 *Ruminococcus* albus 8
 296446028 *Methylosinus trichosporium* OB3b
 34557789 *Wolinella succinogenes* DSM_1740
 222109284 *Acidovorax ebreus* TPSY
 36339380 *Lactobacillus coryniformis* subsp_ torquens KCTC_3535
 218767587 *Neisseria meningitidis* 22491
 15602991 *Pasteurella multocida* subsp_ multocida str_Pm70
 31078033 *llyobacter polytopus* DSM_2926
 312879014 *Amimonomas paucivorus* DSM_12260
 403744859 *Alicyphilobacter hesperidium* URH17-3-68
 421874296 *Brevibacterium laterosporous* GI-9
 220930481 *Clostridium cellulolyticum* H10
 225377803 *Roseburia inulinivorans* DSM_16841
 315659847 *Staphylococcus lugdunensis* M23590
 160915783 *Eubacterium dolichum* DSM_3991
 324363802 *Staphylococcus pseudintermedius* ED99
 389815358 *Planococcus antarcticus* DSM_14505
 328956316 *Coriobacterium glomerans* PW2
 422884107 *Streptococcus sanguinis* SK49
 116628212 *Streptococcus thermophilus* LMD-9
 24379808 *Streptococcus mutans* UA159
 13622194 *Streptococcus pyogenes* M1_GAS
 227824982 *Acidaminococcus* sp_D21
 169823756 *Finegoldia magna* ATCC_29328
 305282779 *Solobacterium moorei* F0204
 303229394 *Veillonella atypica* ACS-134-V-Co17a
 304438953 *Peptoniphilus duerdenii* ATCC BAA-1640
 374307737 *Fillipaccharia alocis* ATCC_35896
 256845020 *Fusobacterium* sp_3_1_36A2
 306821690 *Eubacterium yurii* subsp_ margaretiae ATCC_43715
 42525844 *Treponema denticola* ATCC_35405
 291520706 *Coprococcus catus* GD-7
 258509198 *Lactobacillus rhamnosus* GG
 336394883 *Lactobacillus farciniminis* KCTC_3681
 339625080 *Fructobacillus fructosus* KCTC_3544
 310286727 *Bifidobacterium bifidum* S17
 366983954 *Oenococcus* kitaharae DSM_17330
 #map of conserved positions used for tree reconstruction

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#map of conserved positions used for tree reconstruction
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Continued

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 34557933 *Wolinella succinogenes* DSM_1740
 187736488 *Akkermansia muciniphila* ATCC_BAA-835
 407803668 *Alcanivorax* sp_WII-5
 325926002 *Sphaerochaeta globus* str_Buddy
 187250661 *Elumimicrobium minutum* Pei191
 47458867 *Mycoplasma mobile* 163K
 363542551 *Mycoplasma ovipneumoniae* SC01
 71894593 *Mycoplasma synoviae* 53
 384393278 *Mycoplasma canis* PG_14
 284931711 *Mycoplasma gallisepticum* str_F
 117929157 *Acidothermus cellulolyticus* 11B
 317482065 *Bifidobacterium* sp_12_1_47BFA
 315605739 *Actinomyces* sp_cral_taxon_180_str_F0310
 294086112 *Candidatus Puniceispirillum marinum* IMCC1322
 282880053 *Prevotella timonensis* CRIS_5C-B1
 374384762 *Odoribacter* laneus YIT_12061
 423317188 *Bergeyella zoohelcum* ATCC_43767
 385816001 *Ignavigacterium album* JCM_16511
 402847305 *Porphyromonas* sp_oral_taxon_279_str_F0450
 345885719 *Prevotella* sp_C561
 404487227 *Barnesiella intestinihominis* YIT_11860
 301311870 *Bacteroides* sp_20_3
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 336393380 *Lactobacillus coryniformis* subsp _torquens_KTC_3535
 218767597 *Neisseria meningitidis* Z2491
 156022991 *Pasteurella multocida* subsp _multocida str_Pm70
 310780383 *Ilyobacter polytropus* DSM_2926
 312819404 *Aminomonas paucivorans* DSM_12260
 403744859 *Alicyclobacillus hesperidum* URH17-3-68
 421874296 *Beijerinckia laterosporus* GI-9
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 328956316 *Coriobacterium glomerans* PW2
 422884107 *Streptococcus sanguinis* SK49
 116628212 *Streptococcus thermophilus* LMD-9
 24379808 *Streptococcus mutans* UA159
 13622194 *Streptococcus pyogenes* M GAS
 227824982 *Acidaminococcus* sp_D21
 169824656 *Fingolida magna* ATCC_29328
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 304438953 *Peptoniphilus duerdenii* ATCC_BAA-1640
 374379817 *Filicibacter alocis* ATCC_35896
 256845020 *Bubobacterium* sp_3_1_36A2
 306821190 *Eubacterium yuii* subsp _margaretaiae ATCC_43715
 425258442 *Treponema dentale* ATCC_35405
 291520706 *Coprococcus catus* GD-7
 258509198 *Lactobacillus rhamnosus* GG
 336394883 *Lactobacillus farciniminis* KCTC_3681
 339625080 *Fructobacillus fructosus* KCTC_3544
 310286727 *Bifidobacterium bifidum* S17
 366983954 *Oenococcus kitaharae* DSM_17330
 #map of conserved positions used for tree reconstruction

Continued

16130662 *Escherichia coli* str_K-12 substr_MG1655
 11500011 *Archaeoglobus fulgidus* DSM 4304
 18977490 *Pyrococcus furiosus* DSM_3638
 46447796 *Desulfovibrio vulgaris* str_Hildenborough
 54296139 *Legionella pneumophila* str_Paris
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 -----SD---VIASLQHLLSAKS-----IV-ELMG-----AEAARNAYYT-KDEILKNFEGKRSRMPPE-----
 -----S-----DF---SNHLKELQGAKK-----V-T-EIMN-----VEGRIRQEYRARWDESLEPGFGRIGKTRRPRPK-----
 -----H-GDKVDVGVAEEVLRLRACMRLLQPGAG-----LD-AVRG-----IEGEAAKGYFSVFDN----LILTREAFRFEGRSRRPP-----
 -----K---NRPNEKILHAASQKMAGIVANIT-----ET-NWIQLQDWRTLLGFEGSAARIYWEVTKEVNLVCSDFEG-----RRPRVNK-----

331001028_Parasutterella_excrementihominis_YIT_11859
 319941582_Sutterella_wadsworthensis_DSM_3_1_45B
 118497353_Francisella_novicida_U112
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#map of conserved positions used for tree reconstruction

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16130662_Escherichia_coli_str_K-12_substr_MG1655
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 46447796_Porfobacter_vulgaris_str_Hildenborough
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 407803668_Alcanivorax_sp_W11-5
 325972002_Sphaerochaeta_globus_str_Buddy
 187250661_Elusimicrobium_minutum_Pei191

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 -DINTSMLYNAYSISSLWSVVAITNAGLEYAGILTSKPGPKPSLVDMEERFWCDRIFTKLHARAQKQN-1---TPALKK---TLIGEMANLYK
 -EITNAALNNGYAIOSYIQLNAGNAGLEYAGILFLHQRGKPMVLDOMEERYAWWDNRNIVLRSRCKLGSEQNFL--D---T---ALKKQIVLELIDK
 -EIVNAALNNGYAIOSYIQLNAGNAGLEYAGILFLHQRGKPMVLDOMEERYAWWDNRNIVLRSRCKLGSEQNFL--D---T---EIKSAIVNAIDE
 QELANSALNNGYAIOSYIQLNAGNAGLEYAGILFLHQRGKPMVLDOMEERYAWWDNRNIVLRSRCKLGSEQNFL--D---T---KLSILISEIQA
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 -DIINSALNNGYAIOSYIQLNAGNAGLEYAGILFLHQRGKPMVLDOMEERYAWWDNRNIVLRSRCKLGSEQNFL--D---T---EIKSAIVNAIDE
 -P--SAKKYMASVIES

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 301311870 *Bacteroides* sp_ 20_3
 66863388 *Bacteroides fragilis* NCTC_9343
 402849998 *Rhodovulvus* sp_PH10
 154250554 *Parvibaculum lavamentivorans* DS-1
 159042957 *Dinoroseobacter shibae* DFL_12
 288957740 *Azospirillum* sp_B510
 330822846 *Alicyclophilus denitrificans* K601
 443171926 *Ralstonia syzygii* R24
 319597207 *Nitratiruptor salsuginosus* DSM_16511
 315149831 *Enterococcus faecalis* TX0012
 116627543 *Streptococcus thermophilus* LMD-9
 238924076 *Uabacterium rectale* ATCC_33656
 18563120 *Campylobacter jejuni* subsp_ *jejuni* NCTC_11168_- ATCC_700819
 291276264 *Helicobacter mustelae* 12198
 325677757 *Ruminococcus albus* 8
 296464602 *Methyllosinus trichosporum* OB3b
 34557789 *Wolinella succinogenes* DSM_1740
 222109284 *Acidovorax ebreus* TPSY
 33639380 *Lactobacillus coryniformis* subsp_ *torquens* KCTC_3535
 218767587 *Neisseria meningitidis* 22491
 15602991 *Pasteurella multocida* subsp_ *multocida* str_ Pm70
 310780383 *Ilyobacter polytropus* DSM_2926
 312879014 *Ammononas paucivirans* DSM_12260
 403744859 *Alicyclobacillus heisnerdum* URH17-3-68
 421874296 *Brevibacillus laterosporus* Gi_9
 220930481 *Clostridium cellulolyticum* H10
 225377803 *Roseburia inulinivorans* DSM_16841
 31659847 *Staphylococcus lugdunensis* M23590
 160915783 *Uabacterium dolichus* DSM_3991
 323463002 *Staphylococcus pseudointermedius* ED99
 389815358 *Planoococcus antarcticus* DSM_14505
 328956316 *Coriobacterium glomerans* PW2
 422884107 *Streptococcus sanguinis* SK49
 116628212 *Streptococcus thermophilus* LMD-9
 34739808 *Streptococcus mutans* UA159
 13622194 *Streptococcus pyogenes* M1 GAS
 227824982 *Acidaminococcus* sp_D21
 169823758 *Finegoldia magna* ATCC_29328
 320528779 *Solobacterium mcorei* F0204
 30322939 *Veillonella atypica* ACS-134-V-Co17a
 304438953 *Peptoniphilus duerdenii* ATCC_BAA-1640
 374307737 *Filifactor alocis* ATCC_35896
 256845020 *Fusobacterium* sp_ 3_1_36A
 306821690 *Uabacterium yurii* subsp_ *margaretaiae* ATCC_43715
 42525844 *Treponema denticola* ATCC_35405
 291520706 *Coprococcus catcus* GD_7
 20278198 *Lactobacillus rhamnosus* GG
 336394883 *Lactobacillus farcinimis* KCTC_3681
 339625080 *Fructobacillus fructosus* KCTC_3544
 310286727 *Bifidobacterium bifidum* S17
 366983954 *Oenococcus kitaharae* DSM_17330
 #map of conserved positions used for tree reconstruction

Continued

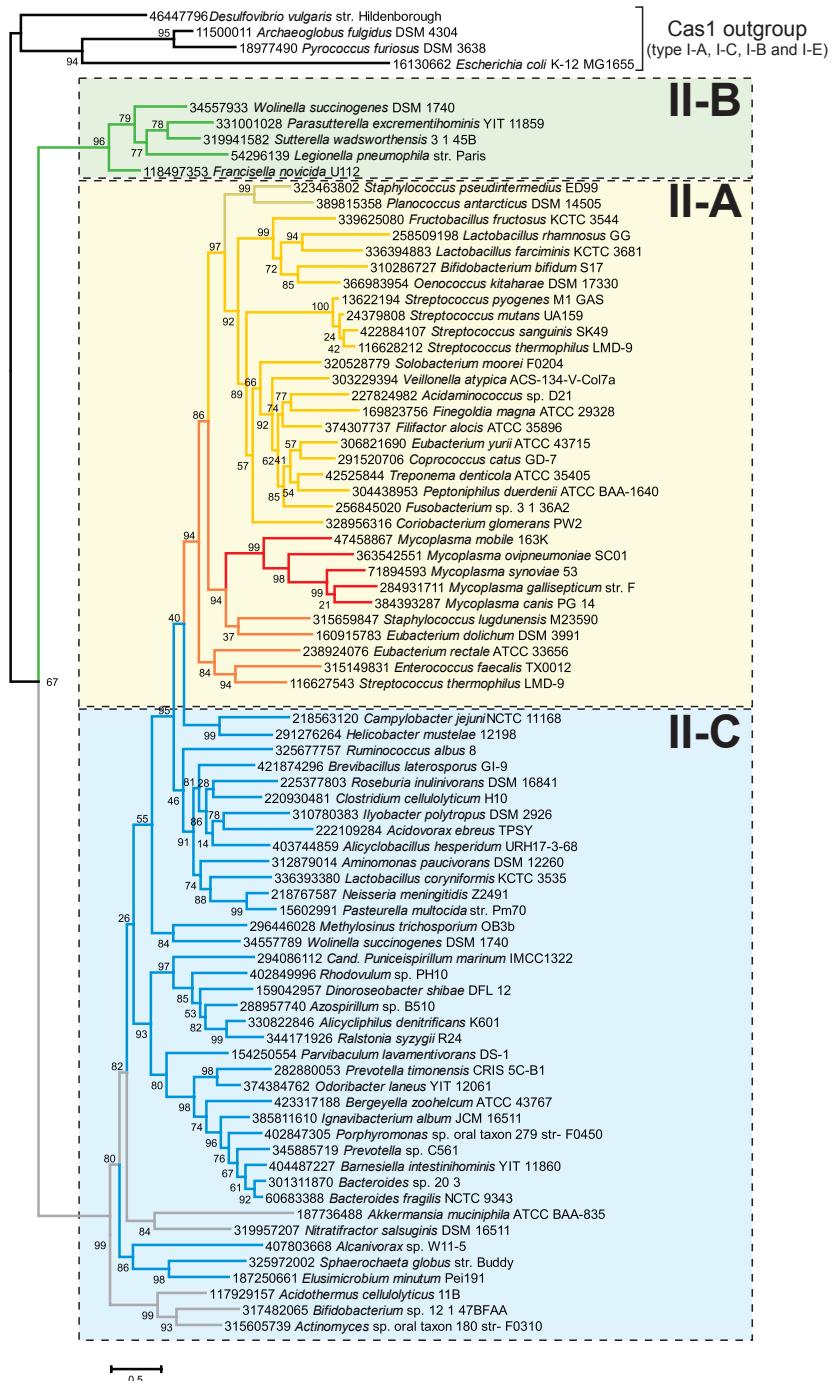
16130662 *Escherichia coli* str_ K-12 substr_MG1655
115000111 *Archaeoglobus fulgidus* DSM_4304
18977490 *Pyrococcus furiosus* DSM_3638
46447796 *Desulfobivibrio vulgaris* str_Hildenborough
54296139 *Legionella pneumophila* str_Paris
331001028 *Parasutterella* *excrementumominis* YIT_11859
31994158 *Sutterella wadsworthensis* 3_1_45B
118497353 *Francisella novicida* UI12
34557933 *Wolinella succinogenes* DSM_1740
117736486 *Akkermansia muciniphila* ATCC_BAA-835
407803668 *Alcanivorax* sp_WL-5
325972002 *Sphaerotilus globus* str_Buddy
187250661 *Elusimicrobium minutum* Peil191
47548867 *Mycoplasma mobile* 163K
363542551 *Mycoplasma ovipneumoniae* SC01
71894593 *Mycoplasma synoviae* 53
384393287 *Mycoplasma canis* PG_14
284931711 *Mycoplasma gallisepticum* str_F
117929157 *Acidothermus cellulolyticus* 1IB
317482063 *Bifidobacterium* sp_12_1_47BFAA
315605739 *Actinomyces* sp_ oral taxon 180 str_F0310

S-----KTLAKIPLIEDVLAAGEIQPPAPPEDAQ-----PVAIPLPVSLGDAGHRSS-----
RLEKTVKHKLRLNRKVSYQLRILECYLKVKHFTAAVEK-----YSPFVMMW-----
ELTEKTVKHPKLGLGVTRRLRILEAKYIKKHLVGWE-----YSPLVAFW-----
RKQDEIVHPFLRPLGQVHQAQMILMARWLKGDLD-----GYF-PFWVK-----
PL-----PYHKRISLESIIQRCVQVHLSAVFSGEKG-----YKGYLFKW-----
TMST-----KINYKQCKLLENALQRCVQVLAGAIVEKGK-----YRGJRFK-----
TMAN-----AVSQGKSVLRNENALQRCVQVRAILGTVVDGKR-----YKSIRFRW-----
TIAKKY-----IYNGKULKLEHHIIRQRVYRLSGEFAEGHN-----YKPVIFKW-----
TMNK-----KLYNGKULSLSQVQVQVRLVGAQMQQNR-----YKSYIFKW-----
SV-----MYDRQKLPKAAVEAVCSFRKAVLAQGS-----EPYEPWLMTTIKWA-----
TV-----CLNGQDFPRLPAIRTKVASTYSLRILEAGGG-----GRRRKLLEPCP-----
EV-----VFEQEVDFESYALRLYVLSYLFSLRSNRD-----AIQFPKYRYAWAV-----
NL-----YFEDGAFNLNTAGIQKYIQSYIAFLEEREN-----RIIFPAILK-----
KV-----IYKGKFYVNDTFDQADNUVNRSWEWY-----ELWD-----
HI-----EYKGKFYKHSPEMEIMIDILNKENYKE-----ITTEDSOS-----
KI-----LLNKNPSISVNEYICKLIENTKDFNPES-----LEIDWNKK-----
KI-----AIKNTRLSVNEYIDKIVKSVINNNLICED-----PIIEWS-----
FI-----FPNGKNU---LNQVMDILMKAIINNNKINERD-----FNIDWS-----
QF-----DASGATVG-----TAVERFAQVGRYVEGEIR-----SLRPAMELSHA-----
Q-----FNGSGLTIPNSLNDFAQPGFLYCEGKID-----RLQVPEVYGES-----
-----RFDCDGHDGIPAVAEALASQFGRYVEGDDID-----RLCVALWQGPSSVAVED-----

294086112_Candidatus_Puniceispirillum_marinum_IMCC1322
 282880053_Prevotella_timonensis_CRS_B-1
 374384762_Odoribacter_laneus_YIT_12061
 42331788_Bergeyella_zoochelum_ATCC_43767
 385811610_Ignavabacterium_album_JCM_16511
 402847305_Porphyromonas_sp._oral_taxon_279_str_F0450
 345885719_Prevotella_sp_C561
 404487227_Barnesiella_intestinohominis_YIT_11860
 301311870_Bacteroides_sp_20_3
 60683388_Bacteroides_fragilis_NCTC_9343
 402849996_Rhodovulum_sp_PH10
 154250554_Parvibaculum_lavamentivorans_DS-1
 1590427597_Dinoroseobacter_shibae_DFL_12
 288957740_Azospirillum_sp_BS10
 330822846_Alicycliphilus_denitrificans_K601
 344171926_Ralstonia_syzygii_R24
 319957207_Nitratirfractor_salsuginis DSM_16511
 315149831_Enteroctococcus_faecalis_TX0012
 116627543_Streptococcus_thermophilus_LMD-9
 238924076_Eubacterium_rectale_ATCC_33656
 218563120_Campylobacter_jejuni_subsp._jejuni_NCTC_11168_-ATCC_700819
 291276264_Helicobacter_mustelae_12198
 325677757_Ruminococcus_albus_8
 296446028_Methylosinus_trichosporium_OB3b
 34557789_Wolinella_succinogenes_DSM_1740
 222109284_Acidovorax_ebreus_TPSY
 336393380_Lactobacillus_coryniformis_subssp._torquens_KCTC_3535
 218767587_Neisseria_meningitidis_Z2491
 15602991_Fastereulla_multocida_subsp._multocida_str_Pm70
 310780383_Ilyobacter_polytropus_DSM_2926
 312879014_Aminimonas_paucivorans_DSM_12260
 403744859_Alicyclobacillus_hesperidum_URH17-3-68
 421874296_Brevibacterium_laterosporus_GI-9
 220930481_Clostridium_cellulolyticum_H10
 225377803_Roseburia_inulinivorans_DSM_16841
 315659847_Staphylococcus_lugdenensis_M23590
 160915783_Eubacterium_dolichum_DSM_3991
 323463802_Staphylococcus_pseudointermedius_ED99
 389815358_Plancoccus_antarcticus_DSM_14505
 328956316_Coriobacterium_glomerans_PW2
 422884107_Streptococcus_sanguinis_SK49
 1166282122_Streptococcus_thermophilus_LMD-9
 24379800_Streptococcus_mutans_UA159
 13622194_Streptococcus_pyogenes_M1_GAS
 227824982_Acidaminococcus_sp_D21
 169823756_Finegoldia_magna_ATCC_29328
 320528779_Solobacterium_moorei_F0204
 303229394_Uvillonella_atypica_ACS-134-V-Co17a
 304438953_Peptoniphilus_duerdenii_ATCC_BAA-1640
 374307737_Filifactor_alocis_ATCC_35896
 256845020_Fusobacterium_sp_3_1_36A2
 306821690_Eubacterium_yurii_subsp._margaretiae_ATCC_43715
 42525844_Treponema_denticola_ATCC_35405
 291520705_Coprococcus_catus_GD-7
 258509198_Lactobacillus_rhamnosus_GG
 336394883_Lactobacillus_farciminis_KCTC_3681
 339625080_Fructobacterium_fructosus_KCTC_3544
 310286727_Bifidobacterium_bifidum_S17
 366983954_Oenococcus_kitaharae_DSM_17330
 #map of conserved positions used for tree reconstruction

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Cas1



Supplementary Figure S6. Phylogenetic tree of representative Cas1 sequences. The tree was reconstructed from selected, informative positions of the Cas1 multiple sequence alignment (see Supplementary Figure S5 and Supplementary Materials and Methods), and was rooted to selected Cas1 orthologs of the type I CRISPR-Cas outgroup. Shaded boxes group Cas1 sequences associated with three major type II CRISPR-Cas subtypes. Colored branches indicate the CRISPR-Cas loci with similar architecture. Compare with Figure 4. Note the similar topology of Cas1 and Cas9 trees (Figure 4).

Supplementary Figure S7. Analysis of Csn2 orthologs. (A) Complete multiple sequence alignment of selected Csn2 orthologs and selected ATPases. See Figure 5. Underlined, regions of homology with short Csn2 ortholog according to HHPRED. (B) HHPRED hits for selected Csn2 orthologs representative of five families. See Figure 5. (C) Selected proteins with structures similar to long and short Csn2 structures (3zth and 3toc) according to PDBe and their structural alignment with long and short Csn2 representatives (3toc and 3zth). Walker A, Walker B and signature motifs are highlighted in boxes. See Figure 5 and Supplementary Materials and Methods.

A

3ZTH_SS E---EEE---HH---HH---HHHHHHHHH---H---HH---EEE---HHH---HE---HHHHHHHHHHHHHHHHH---EEE---HHHHHHHHH---EEE---HHHHHHHHH---HHHHHHHHH
3ZTH IENKNLNSFE---WVS-NI---DKLSSLFLEM---L-DH---LLSQTTKEKYLIVLKNIDGFISEE---SYTIFYRQICHLVKVYKPNTLFILFP---SDQGYL---KIDEENSRFVNILSDQ---VEHLYDVEFMYERVMKYPSNDFPTREGFRMSLET---VTPYLLTKMLRQPSSLVDSVLINLNQLHFHSY---RIRCSQTDPKDELLQKFLE---
365905381 DSNRTFPLE---FME-TE---SLVDELLNL---I-ES---SLKKANNDTWIVLYNLDNSFVSRS---AKKTILKLKEFTDR---ELVKVYLG---NSLTD---SITAETEKVVAADE---FHQLPPIEINLSKS1KMRYPNDPFPHSTEQLVLSLQI---VIPYVGKN---ENVSLSNRLDVLLK1VND1IGET---SLDYSDQ1LNSAETKFLLD---
406671115 YDGQEVPLD---TFT-LN---LAYQDMLEL---I-KF---RVEEEGRVLLVILL---DLEPFRTINFTEVAEFTIQLIEIANNTHLKLIIFD---DGDLNQ---K1---PLSISKT1IYTDD---IQOMPDEIFYQSISIKRNPDKITCSDSNLDRDFYC---ICKYLGHQIYKNIISLSSKDVLVLLYVVKDLAADNT---YIETSVDSLSEAEEKFYLASKNTRY---
422729713 ENQQLSPLM---MDM-TG---ELLAEIFYL---L-QR---SISRREQPWLWITNPEFSELE---SIHQLFQKLNQKLAQETKQLKFFIFS---NRSRLP---PYTTEDEVKTILLYDQ---YQQLPVFVQFSIERHYPDQLSWTNQQLIDAFYR---VCHFVGQD---YTKNYLLPKDMILLKLKELLDONS---ECVETSMEKLTVLEQYFRLRKE
406838713 ENSQVFLPQ---MLD-VN---KTIDEFCRCL---L-RN---FLISESKIPWLWIKPNPSFLAKG---SLKFKTIAVLKTTNETKLRLRVFVS---DDFDL---AYEPEDMVENTVLYINE---IQOLPPYDIFLNSIKNNYQFLDKNNMNLLLKLVELDGENCQFLQKGNRNELETCPEENFLYKG---
238924078 DTE---LS---DKS-NA---ELILILLNI---V-DN---VLKNNPKTTLIFENLDLHVLSLK---EYVEIIHAADTISKY---DLYFIFSS---SLNKVY---CCDIFLEGISVFGDV---YFQMPEYNRLY---LFLKENYPIYEKEFNSHEVKK1ICN1IQNIGQSNFLNTVENVCKLINKTL---LLDDKIPDEKMPMELKA
381184142 YEERTYATE---YVD-GL---TKYMSLLDL---L-EI---NLESSSEMILLMIKNIIDTLYQ---EYEEVMNSMKRLTTHKPNFVCMVFP---SQIGYV---YI SEEHEIESI LVVGE---AHSLADGRNLTRVCAVYPDQTIPSYNEFLGRKM---IAPYLFESTSKKQSIQMNVKELVLIKINELFQYD---LIPVAVLQHQSLEYDFYES---
116100822 IENKNLNSFE---WVS-NI---DKLSSLFLEM---L-DH---LLLQTTEKYLIVLKNIDGFISEE---SYTIFYRQICHLVKVYKPNTLFILFP---SDQGYL---KIDEENSRFVNILSDQ---VEHLYDVEFMYERVMKYPSNDFPTREGFRMSLET---VTPYLLTKMLRQPSSLVDSVLINLNQLHFHSY---RIRYQKSDKEKKLQKFLESKD---

4199_SS -----HHH---HHH---HHHHHHHHH---H-HH---EEE---HHH---HE---HHHHHHHHHHHHHHHHH---EEE---HHHHHHHHH---HHH---HHHHHHHHH---HHHHHHHHH
4199 AGKDVKRKE---AXS-CG---AKTALTAFL---V-FA---IQK-FKPAFPFLFDEI DAHLLDA---NVKRVADLAIKESS---ESQFIVIT
20NK AHLLDRKPA---RLS-GG---ERQRVALAR---A-LV---I---QPRLLLDEPLSAVDLK---TKGVMEELRFVQREFD---VPLHVT
205V DFPASDYAS---RGE-GR---TVALALRRA---EEL---LREKFGEDPVPLLLDFTAELDPH---RRQYLLDAAV---PQAIVTG

3TOC_SS -----HHH---HHHHHHHHH---H---EEE---HHH---HHH---HHHHHHHHH---EEE---HHH---HHHHHHHHH---EEE---HHH---HHHHHHHHH
3TOC -----TQ---SDT-IF---EKCLEIIQI---F-KY---LT---KKKL1LIVFNSGAFLTKD---EVASLQYEISLTN---LTVLFL---PRELYD---FP---QYILD---LITKNM
13622196 -----TQ---SDT-IF---EKCLEIIQI---F-KY---LT---KKKL1LIVFNSGAFLTKD---EVASLQYEISLTN---LTVLFL---PRELYD---FP---QYILEDYDFFLITKNMV
339625078 -----KG---NNSTF---GKIEVEHTV---GM---LD---ESRYLFLTNASLYCQLS---DLNQLHCELLAEG---VNLSISD---LSLEKPNFSDHYC---HFH1DQDFVFLYS
333398276 -----TL---LTSNSF---GKIEDVVSV---A-GE---FS---EQRLIVFTNLYNLLSVD---QIDYLNLLAKTMN---LCLISLN---LTQNPVMTKQGLPP---B1F1DEDFFVQFGAD
372325148 -----MP---ILNTPY---DKIRSTIDL---A-AE---LS---DPLHVFNTNAFQYLTMG---QTNQLAESRSME---RNVLFL---RFDSKIDLTAEAN---SYFFDKDVFQF---
423349697 -----VP---EYENFY---DIAQAVIDI---A-GA---LS---EKRMVLFLHISEYCTND---QLEVLVSDIKRQE---LQVLYLE---RTDHLLRGFD---ER---SYVVEDDFVQFS---
310286726 -----TS---SLGSY---DKISTAVDT---A-GA---LA---ESRMLVTLHTTYQCDND---QLLYLHRTLLRYQ---LQLLDL---CCSKVVMLTE---GM---SHVDEDYVQFS---
336394885 -----FS---SWANYC---DKIMDVISF---Y-SD---FT---NKKML1FNNIGRLNNVN---QLINEIHYAHLKSD---LKLVSLESYPMIFKEEKLN---AK---VYISIDNDHVRFDY---
408790125 -----TS---SMDDN---PKIVKIL---Y-HK---FY---PKMLMVLNLGFNYLNSS---QINDLQIMKALD---LXVLLD---FSADFRKELADK---CR---YRVDSDFVRFNY
258509196 -----EE---VLTDPF---SKVEAAIKT---L-AK---LE---EKRLVIFTNISHYLDV---SLNKLDDQVKETS---LELLLIE---FSDVERRKFFER---CQ---YIYIDKDFMDSRELID
206662217 -----TQ---VIRQPY---GIIETDQL---H-LE---LK---DCVGLYCNVANVYLP---QIEQFATLVEK---LAVLVL---FSEKRETQLQQLQD---IYR1DRDFWDWHE
354808133 -----TV---VQHDPE---GIESIVQT---A-VE---LN---ESKI1LINVSHYLSIN---QFNLNEVRLVATLN---VKLFLIE---FSEEVKSQDQYQK---CY---YHIDNDYVWEWRYE
392947439 -----TD---LTHNPS---AVLEALVOT---H-VE---LG---LQKCPVMTNVSHYLNPE---QWAFFEQVADLG---TTVIIIE---YSEMRNMRMFKD---CC---YYVDEDLWDWRDMN
333394443 -----TS---LSTNCY---GIIETIILT---A-KE---LN---EQRILGFMMNISDYLTLQ---ELQDFFLIELR---LTVLIIK---FSENQRSVEFNN---CR---YYIJDQYVWDWRYE
323340065 -----SS---IAYDPC---AIIETDIKI---H-AE---CA---LKKAI1GVSGLATYLP---EFSY1DQVVKEMD---VPLLMID---YNGSGWKSYFDN---CN---VMIYIDBEDFIDWH
308344195 -----AL---LWKKNAY---DRISSVVKL---H-VE---LN---RKIRK1LNTVNAHYLTLEG---EFQELVSLVKMTN---SSMFIE---FTEKNGQRFFEN---CD---NYIJDYEDVWY---
328956318 -----PN---POETFL---DNLIDFLSF---V-LD---IQ---CSKVMFVFLNKTFLTE1---ELEKLYEHWVFSSK---TRVLLIE---NKAOPNRHEN---EI---KTIIDLDLFIK---
374307735 -----EE---EGR-FV---ERLIDYSKT---M-FE---LL---HKN1F1VFGCSGLTK1---DFEYLRQHQFSQK---I1LFLVFE---SHQIDLESP---KK---QTIIDBEDLCEI---
224543315 -----RK---AEEKKY---DLRMLVYD---V-TE---LV---KKPVFLVLYNCFDYLDBE---ELMELIKYKKNYK---LHLLFIE---NKNREVDSTAFR---KTIIDBEDLCEYD KIEEVQSD
227824980 -----LD---YNG-LG---EKIFVLMMLD---I-SR---FE---GDK1F1VNLRSYLSSE---EIQ1LFWDTALAH---LKFLLID---NRRYPLFHQ---EK---RVLIDNDLCV
169823758 -----VD---YF---DKFVFLFDI---Y-QN---VL---K1DVFYTINLQ1QYTFEE---ELKELKSY1IKINN---IC1IYND---NILVDSNYISK---KLLFDNDLRCV
303229343 -----FN---RKG-EV---MDLMSYDVE---V-DT---LL---SPM1YIM1NLDLWNDD---E1NAFYQNMLSRO---LRLVCLT---TGSFDRESLDKNLIN---GY1LDNDFCII
315926106 -----EQ---SLS-LE---EKLAEYFKI---L-SL---VL---RKPI1LCLLHAIYFSEK---QIRELIQLSKYQK---LQLLFI---AREPKK1FDEDED---IY1LDQDRCVIK
306821688 -----EN---DZC-LL---EKLINVYKI---I-SQ---VS---FTKFLVFLVNHDFLFSK---E1DMLYB1ALY---S1LVI1E---SHQREKSSY---EH---TY1LDNNCLIEF
320528781 -----HN---YSS-LL---E1IVDY1DI---F-SQ---II---TKTLFVFI1SLRSYLTGB---E1NNFK1MDYKG---IR1L1IE---RY1LSLNSIVDN---QV1IDKDLCVI
342218249 -----WG---EME-LL---ERLVTYLDI---I-CD---LL---KSKL1LVLNVNRKYMTEE---ELKLVLYEH1CIRKK---I1FICIE---SGYTCHSVDELK---VY1IDKDLCFIY
227501315 -----TK---FTG-LM---ANIIEYDII---S-HK---LL---NKK1F1FVGIDDYFEER---ELYDLNEYLYCND---I1IVICLQ---NRLDRDLLP---EN---LRIIDEDLGEI
304438951 -----YE---KNS-VL---ERLIEYIKV---T-SE---LL---KTKVY1IVNLDLSFLSEE---ELVELEKFLLYNLD---IKVLAQ---NAIRREVIPS---EN---LRIVDKDLCEI
34762595 -----RS---ED---LL---LNLFEWLKI---L-NE---IL---GY1FFFINIENFLSED---ELLEFSK1IVYKNK---YKVFL---N1FYNRLKSDN---DN---LIV1DNDLCIEF
241889886 -----VE---NDL-ID---ENFVKFLKV---L-TE---LC---CFK1F1VFLGHTVFTQ---E1E1IYKEVCLNK---INI1VNE---YQQFNNLSAKN1KEI---VY1IDKDNCEI
291520707 -----EM---EDD-FF---ERI1RYVKI---A-VE---VL---STKVFVFN1RSYLTDE---Q1QELKE1R1QE---AK1LFLM---SQERACLEG---GM---VY1IDRDGCEIY
42525846 -----TE---YTN-LL---E1KIDY1VA---S-VE---LN---Q1C1FF1FLN1QFLHFK---DLSELY1F1PACY1K---T1NLFLIE---G-SYST1KEDC---EK---SY1IDKDLCEIY
299144349 -----FD---YGN-TI---DRLISYMV---L-RK---YI---G1KL1FL1NIK1E1L1TC---EMKEFFY1C1MNE---FKV1L1E5GKADYEN1DL1K1Y---IK---RVIIDBEDLCDL
116101487 -----TK---SCT-VF---EK1FD1QI---F-KY---LV---KKR1L1VFN1SLSYFSKD---E1YQ1LEY1TKLSQ---ADVLFL---PRQ1E6---IQ---QF1LDKDY1LMPYNN

MYCO -----H---HHHHHHHHH---H-H---EEE---HHHHHHH---EEE---HHHHHHHHH---EEE---HHHHHHHHH---EEE---HHHHHHHHH---HHHHHHHHH
350546883 -----DL---IN1-NK1---E1K1---N---LFD---ERMTFVFC1D1WNN---IND1SEY1---NS---HN1F1VTDN---R1LKN1V1T1Q1E1C1V1C1N1D---LVE1YDYNK1D1WNN---K1V1E1N1L1K1N1L1N1---
193216855 -----ET---SLL-TR---DLFFKWL---N-Y---DEV---GQKV1K1V1K1N1PWVK---1EDLVQY1---D1EDV1LTD---FFDNC1R1F1EATCAF1WEGKR---F1WV1H1V1W1E1K1N1L1Q1E1K1N1F1STFD---GEK1LFFN1L1SEVFS
47458866 -----KD---LL1-NE---IN1L1N1K1---I-DK---YSN---ET1S1IY1G1K1LN---FSEF1---F-NK---L1N1Y1L1D1K---L1N1L1S1F1E1N1Q1L1E1K1N1---
363526557 -----ED---Y1-DH---E1F1Y1S1E1L---H---WSK---EVK1V1L1F1D1F1D1N1D---L1K1R1S1N1T---Q1---T1N1L1M1D1Y1L1E1N1K1M1D1N1P1S1F1E1D1F1V1M1D1---
490549417 -----E1K---LL1-SK---NN1L1N1L1D1---L-S---SSN---F1K1P1L1V1K1D1Y1T---I1E1L1K1Y---S1N---L1N1L1T1S1D---FTK1Y1Q1SYN1L1E1V1F1Y1N1G1---
363542553 -----ND---ILI-NE---ENFENLAEI---I-F---STM---K1T1V1L1K1D1Y1---FEK1L1F1Y---NN---V1F1L1T1D---FTK1Y1K1S1F1E1L1V1F1Y1---
71894595 -----ND---KPI-DN---KSL1K1WMEN---Q-K---YNN---Q1K1N1L1K1N1F1D1V---IN1L1K1F1S---F1N1V1L1D---F1N1S1R1N1E1K1F1D1F1S1N1F1---
419703977 -----TK---EFI-TS---SLLYSFLRN---I---SQA---E1K1N1I1K1MD1D1S---L1S1L1T1F1---F1N1V1M1T1N1---F1N1V1T1N1---S1F1N1K1S1S1E1V1L1F1N1D1E1S1N1F1E1V1F1---

STAPH_S -----H---HHHHHHHHH---HHH---HHHHHHHHH---EEE---HHH---HHHHHHHHH---EEE---HHHHHHHHH---HHHHHHHHH---HHHHHHHHH
389815356 -----TV---SDE---PMGAYQYQ1FL1K1S1W1L1---V-NK---ATNVCFYDFPENELNKS---EL1K1F1Q1F1S1Q1---CTM1C1---TSPQ1V1N1Q1V1G1S1N1V1H1L1K1F1E1L1V1F1---
386318633 -----NQ---LKD---L1E1L1N1K1F1A1F1D---L-IH---PT---K1E1N1L1F1P1E1S1L1G1---D1K1F1H1L1K1Y---F1T1F1F1---N1H1P1S1I---
414160480 -----ND---LDD---L1N1A1E1Q1R1K1F1D1---L-TS---TE---K1E1V1L1V1F1P1E1A1G1---M1F1S1I1T1---SHPY1F1---I1E1S1D1L1A1K1T1N1G1---

STAPH_L -----HHHHHHHHH---HHH---HHHHHHHHH---EEE---HHH---HHHHHHHHH---EEE---HHH---HHHHHHHHH---EEE---HHH---HHHHHHHHH
315659845 -----LKD1Y---L1F1R1Q1V1P1L1---V-IE1L1K1Q1H1---N1Q1L1I1Y1P1E1A1N1S1P1---E1Q1R1K1Q1L1S1D1---V1I1V1L1---
403411239 -----Y1S1D1Y---T1R1Q1V1P1L1---V-V1D1M1Y1Q1F1V1---E1K1F1Y1L1P1S1K1---E1P1F1R1F1V1D1---P1V1F1---
425737239 -----L1A1D1Y---L1V1R1K1Q1L1---V-IE1M1N1S1N1---N1G1L1I1Y1P1E1A1N1S1P1---E1R1F1R1F1V1D1---T1K1V1L1---
-----OS1R1Y1F1---D1G1T1D1S1G1L1N1R1S1N1K1M1T1V1F1---F1E1N1C1Y1F1P1E1S1L1N1F1---OS1R1Y1F1---D1G1T1D1K1S1L1A1S1Y1L1S1F1---

B

HHPRED hits for Csn2 long

No	Hit	Prob	E-value	P-value	Score	SS	Cols	Query	HMM	Template	HMM
1	4i99_A Chromosome partition pr	95.0	2.5	1.7E-05	43.4	14.0	230	2-245	1-305	(354)	
2	glxeww.l_c.37.1.12 (X;Y); Smc	94.6	3.1	2.1E-05	42.3	13.3	219	4-236	6-278	(329)	
3	3auy_A DNA double-strand break	81.5	8.8	5.9E-05	37.8	4.9	66	1-67	4-72	(371)	
4	4ad8_A DNA repair protein RECN	76.4	18	0.00012	40.1	5.9	47	1-48	39-88	(517)	
5	4aby_A DNA repair protein RECN	75.0	21	0.00014	38.5	5.7	46	1-47	39-87	(415)	
6	3ktba_B Chromosome segregation	73.2	71	0.00048	32.3	8.4	115	113-242	9-123	(173)	
7	gf1f2t.1_c.37.1.12 (A;;B); Rad5	72.6	1.1E+02	0.00074	30.8	9.4	216	13-244	15-263	(292)	
8	d2za4b_c.9.1.1 (B:); automated	72.3	70	0.00047	29.0	7.3	62	175-243	26-89	(90)	
9	2fnfaA01	71.5	60	0.0004	31.6	7.3	87	161-249	95-181	(208)	
10	d1e69a_c.37.1.12 (A:); Smc hea	70.9	13	8.7E-05	37.8	3.1	47	4-51	5-55	(308)	
11	PF02463_SMC_N; Recf/Recn/SMC	69.4	23	0.00016	35.2	4.4	46	1-47	1-51	(220)	
12	d2hxxa_c.9.1.1 (A:); automated	69.3	1.1E+02	0.00073	27.8	7.8	52	191-243	34-88	(89)	
13	PF13304_AAA_21; AAA domain; P	68.1	1.6E+02	0.0011	27.9	9.0	215	22-244	1-295	(303)	
14	lw1w_A Structural maintenance	67.3	1.3E+02	0.00086	31.5	8.9	124	113-244	269-393	(430)	
15	3zgx_A SMC, chromosome parti	66.5	20	0.00014	36.5	3.4	44	4-48	5-52	(426)	

Probab=95.02 E-value=2.5 Score=43.40 Aligned_cols=230 Identities=11% Similarity=0.085 Sum_probs=0.0

>glixine_1.c.37.1.12 (X,Y): Smc head domain (*Pyrococcus furiosus* [TaxId: 2261])

Q ss_pred	EEEEEEccccce---EEEccccc-eEEEecCcHHHHHHHHHHHHHHHHCCCCcc-----	
Q gi 365905381 r	2 RLKIEIIDQSF---FELDFGD-IVYISGYNQNWMWKIYRSLYYFNKSPM-----	47 (340)
Q Consensus	4 ~l~i~h~~~~~---ie~~~g~~tq~G~n~qlk~y~l~q~L~y~F~gkys~ ++ + . +. + ++ + ++ . + . -- = = ++++ .++ +... ~l~f~~~lnvi~G~NGsGKS~il~AI~~~L~~~~~r~~~~~ ~N~F~k~s~~~~~	50 (365)
T Consensus	6 ~i~i~-NFks~~~~~l~f~~~lnvi~G~NGsGKS~il~AI~~~L~~~~~r~~~~~ ~N~F~k~s~~~~~	84 (329)
T g1xew.1	6 KLELK-GFKSYGNKVKV~PFPSKGFTAIVGANGSKNSNIGDAHILGLGSKAMRASRISDLIFAGSKNEPAPAKYAEVAI	84 (329)
T ss_dssp	E~~~~~SBTTBCSSCEEECSSEEE~~~~EECTSSHHHHHHHHHHHTTSSCCCC-----	CSEEEEEE
T ss_pred	EEEEEE-CeEcccCCEEEeCCCcEEEEEECCCCCCCCHHHHHHHHHHHHCCCCchhhccccchhhccccccccchhhhhh	

Q	ss_pred		HHHHHHHHHHhcCCCCEEEEeChHHHHHhcHHHHHHHHHHHHhcC		
Q	gi 365905381 r	179	LLNLIESSLKKANNDTWIVLYNLDLSFVSRS-AKKTILNLKLEFTDRF	224	(340)
Q	Consensus	190	-L~l~~l~~~~~l~lv~l~Knid~fis~~~~~s~s~s~i~l~~~l~t~k~ .+.~~~~~+...+.~+++.~+ ~...~.~+.~+.~.++++++	236	(365)
T	Consensus	234	~l~~~~~s~~~~~s~~~~~ide~e~l~~~~~s~~~~~l~~~~~	278	(329)
T	g1xew.1	234	ALAFVFAlQKFKP-PAPFYLFDEIDAHLLDA-NVKRVAIDLKESSKES	278	(329)
T	ss_dssp		HHHHHHHHHHHS-CCSEEEESTTTSCH-HHHHHHHHHHHHHTTS		
T	ss_pred		HHHHHHHHHHhc-ccchhhhhhRRhCcHH-HHHHHHHHHHHHccC		

No 8
>d2za4b_c.9.1.1 (B:) automated matches {Bacillus amyloliquefaciens [TaxId: 1390]}
Probab=72.27 E-value=70 Score=29.00 Aligned_cols=62 Identities=10% Similarity=0.091 Sum_probs=0.0

```
No 9  
>2fnnaA01  
Probab=71.53 E-value=60 Score=31.56 Aligned cols=87 Identities=24% Similarity=0.332 Sum probs=0.0
```

Q ss_pred		HhhcccccccCcceeceeechHHHHHHHHHHHHHHcCCcEEEEeChHHHhcchHHHHHHHHHHHHHHcCCCEE		
Q g1 365905381 r	150	LKYFLKVSYEDSNRTFPLEFMETESLVDELLNLIELSSLLKANNNDTIVLYNLDLSFSVRSR-AKKTLINLKKEFTDRF-ELK	227	(340)
Q Consensus	161	1~k~~~~~l~le~v~~~~~L~l~~~l~~~~~1~l~v~l~K~d~f~s~~~~~s~i~l~lt~k~l~l~t~k~l~	240	(365)
		+.+.-.----+.-.----+.+++.+++.+++++!+-+.-.----+.-.----+.+++.+++.+++.++.		
T Consensus	95	~~~~~DE~~~~~l~~~~~l~~~~~l~~~~~l~~~~~l~~~~~l~~~~~l~~~~~l~~~~~l~~~~~l~~~~~l~~~~~	172	(208)
T 2fnaA01	95	LKAKLKK1E1GIE1IRGLSLE1LKLKKS1E1FELLE1KLE1KKGK1V1L1D1E1Q1K1L1K1B-G-KD1LS1ALK1LD1NL1K1V1L	172	(208)
T ss_pred		HHHHcCCcCCCCC1CcCcCCCCC1HHHHHHHHHHHHHHcCCCEEEECChHHhccc-c-hHHHHHHHHHHHHCCCEE		

```

Q ss_pred EEEEcCCCC
Q g1|365905381|r 228 VIYLGNSLT 236 (340)
Q Consensus 241 -iif-s~~~ 249 (365)
                ++++.+|..+
T Consensus 173 ~~~~gs~~~ 181 (208)
T 2fnaa01 173 FIATGSSVG 181 (208)
T ss pred EEEeeCcRh

```

```
No 10  
>de1e69a_c.37.1.12 (A:) Smc head domain {Thermotoga maritima [TaxId: 2336] }  
Probab=70.93 E-value=13 Score=37.79 Aligned cols=47 Identities=21% Similarity=0.330 Sum probs=0.0
```

Q ss_pred	EEEEEEccccce---EEEccccc-eEEEecCcHHHHHHHHHHHHHHHHhCCCCccH	
Q gi 365905381 r	2 RKLIEIDDSQSF---FELDFGD-IVYISGYNHQNMWKIYRSLYYFFNKSMP--	47 (340)
Q Consensus	4 ~l~h~h~~~~~l~e~~~~g~~~tq~i=G~n~qlk~y~i=q~L~y~f~gkyse	51 (365)
	++ + .+ . .++ + + .+ .+- =+= + . .++	
T Consensus	5 ~i~I~-NFksf~~~~~i~f~~~lnvlvG~NgsGKS~iL~Ai~~~lg~~~~~	55 (308)
T dle69a_	5 KLYLK~GFKSFRGRPSLIGFSDRVTIAVPGNGSKNSIIDAIKWPVEQEKSKE	55 (308)
T ss_dssp	EEEEEE-SBTBCSCE~~~~~ECCCC~~~~~CTTCSTHHHHHHHHHTSCC-----	
T ss_pred	EEEEEE-CeEEECCCEEeeCCCCee=EECCCCCCCCHHHHHHHHHHHHHCCCCChhh	

No 11
 >PF02463 SMC_N: RecF/RecN/SMC_N terminal domain; InterPro: IPR003395 This domain is found at the N terminus of structural maintenance of chromosomes (SMC) proteins, which function together with other proteins in a range of chromosomal transactions, including chromosome condensation, sister-chromatid cohesion, recombination, DNA repair and epigenetic silencing of gene expression []. The domain is also found in RecF and RecN proteins, which are involved in DNA metabolism and recombination.; PDB: 3HTK_A 1WIW_C 2WD5_A 3L51_A 1XEW_Y 3KTA_B 3NW_C_B 1XEX_A 1GXL_C 1GXK_A
 Prob=69.41 E-value=23 Scores=35.22 Aligned cols=46 Identities=24% Similarity=0.339 Sim prob=0.0

Q ss_pred	CceEEEEEc(ccc)---EEEccc-eEEEeCcHHHHHHHHHHHHHHHHhCCC	
Q gi 365905381 r	1 --MRLKIEIDDQSF---FELDFGD-IVYISGYNHQNWMK1YRSLYYFNKS	45 (340)
Q Consensus	1 ~mM~l-i-h~~~~~i-e~~~~g~t-q~n-qlk~y-i-q~L~y~F~gk .++ + + ++.+.++ ++ . . -. =+.. =+.+. .++	47 (365)
T Consensus	1 ~I~~l~i~nFr~~~~~i~~~~~i~G~NgSgKS~ileAi~~l~~~	51 (220)
T PF02463_consens	1 MIKSLEIE-NFRNFKGKNAELSFSPLGNVINGPNGSGKSNLIEAIEFVLGGR	51 (220)
T ss_dssp	EEEEEEEEE-SBTCT-SCEEEETTSSEEEEEEESTSSHHHHHHHHHHHTSS	
T ss_pred	CccEEEE-CceEEEeCcEEEEeccccCCCCEEEccccCCCCHHHHHHHHHHHHHC	

No 12
>d2hxxa_c.9.1.1 (A:) automated matches {Bacillus amyloliquefaciens [TaxId: 1390] }
Probab=69.32 E-value=1.1e+02 Score=27.82 Aligned cols=52 Identities=10% Similarity=0.179 Sum probs=0.0

Q	ss_pred	HHHHHHHHHcCCC-cEEEEEEeChHHHHHhc--cHHHHHHHHHHHHHHHHcCCCEEEE		
Q	gi 365905381 r	180 LNLNIELSSKKANN-DTWIVLYNLDSFVSR--S-AKKTILNLKLEFTDRF-ELKVII	230	(340)
Q	Consensus	191 L-ll~~~l-~~~~~l-lvlvKNId~fls~~~-s~~~i~~~l-llt-k~~~l-ii + - -.. .. + ++ +.++.. .+.. ++.. ++ ++	243	(365)
T	Consensus	34 1DALD~D~L-~~~~~p~ ~~~~~l~~~~~l~~~~~l~~~~~l~~~~~l~~~~~l~~~~~	88	(89)
T	d2hxxa_	34 DLDALXDALTGXVEYFLVLXRQFESQSKQLTEN-GAESVLQVFREAKAEGADITIIL	88	(89)
T	ss_dssp	HHHHHHHHHHHTCCSSEEEEEEHHHHHHHHHHHHHHHHHHHHHHCTCCEEEE		
T	ss_pred	HHHHHHHHHCCCCCCC-EEEECCHHHHHHHHHHHHHHHHHHHHHHHCCCCeEc		

No 13
>PF13304 AAA_21: AAA domain; PDB: 3QKS_B 1USS_B 1F2U_B 1F2T_B 3QKT_A 1II8_B 3QKR_B 3QKU_A.
Probab=68.12_E_val=1.6e102_Score=27.89_aligned_col=215_Identities=165_Similarity=0.181_Sum_probab=0.0

No 14
 >1wlw_A Structural maintenance of chromosome 1; cohesin, chromosome segregation, cell adhesion, kleisin, MIT cell cycle; HET: AGS; 2.90A
 {Saccharomyces cerevisiae} SCOP: c.37.1.12
 Probab=67.30 E-value=1.3e+02 Score=31.47 Aligned_cols=124 Identities=11% Similarity=0.092 Sum_probs=0.0

No 15
>3zgx_A SMC, chromosome partition protein SMC; cell cycle; 3.40A {Bacillus subtilis}
Probab=66.45 E-value=20 Score=36.46 Aligned cols=44 Identities=20% Similarity=0.338 Sum probs=0.0

Q	ss_pred	EEEEEEccccce---EEEccc-eEEEEEccCccHHHHHHHHHHHHHHHHhCCCC		
Q	gi_316905381 r	2 RLKIEIDQDFG---FELDFGD IVTIVSINGHNQNNMKIYRSLYYVNFKNSP	46	(340)
Q	Consensus	4 ~l-i~h~~~~~i~~e~~g~~t~g~i~G~n~q~l~y~i~q~l~Y~y~g~k~k~ ++ + .+ + .+ ++ ++ +. - = ++ . .+ ++ .	48	(365)
T	Consensus	5 ~l-i~nFks~~~~~i~f~~~v~i~v~G~NGSGKS~ileAi~~lg~~	52	(426)
T	3zgx_A	5 RLDV1-GFKFSFAERISVDFVKGVTAVGPNGSGKSNTDAIRWVLGEQS	52	(426)
T	ss_dssp	C~~~~SSSSBSCCCBC~~~~SEEEEEECC---CTTHHHHHHHHHHTCCCC		
T	ss_pred	EEEEEE-C~~~~EECCCCC~~~~EECCCCC~~~~HHHHHHHHHHHHCCCC		

HHPRED hits for Csn2 Mycoplasma

No Hit	Prob	E-value	P-value	Score	SS	Cols	Query	HMM	Template	HMM
1 PIRSF037215 Peptidase_T	21.0	2.9E+02	0.002	31.2	3.1	33	90-122	293-325	(419)	
2 PF07693 KAP_NTPase:	KAP famil	17.6	1.8E+03	0.012	22.8	7.1	81	143-237	157-246	(325)

No 9
>PIRSF037215 Peptidase_T
Probab=20.96 E-value=2.9e+02 Score=31.18 Aligned_cols=33 Identities=15% Similarity=0.278 Sum_probs=0.0

Q ss_pred	cchHhhhhhHHHHHHHHHHHHHHHHhhccchheccc
Q gi 350546883 r	83 -IETNKILGNDKI-HNKEYINSKIGFDVIEEK 113 (250)
Q Consensus	90 ~~~~dki~~~~~i~~~I~~~iN-kiGfdfv~~~e 122 (265)
T Consensus	+ +.+-.-+++.++++ ++ - .. .++
T PIRSF037215	293 DFD~~~F~~~K~~~~~v~~~~n~~~g~~~~~e 325 (419)
T ss_pred	293 DFDRDGFEARKEELIKEIVDKLNAYGRARIKLE 325 (419)
	CCChHHHHHHHHHHHHHHHHhhccccEEE

No 10
>PF07693 KAP_NTPase: KAP family P-loop domain; InterPro: IPR011646 The KAP (after Kidins220/ARMS and PifA) family of predicted NTPases are sporadically distributed across a wide phylogenetic range in bacteria and in animals. Many of the prokaryotic KAP NTPases are encoded in plasmids and tend to undergo disruption to form pseudogenes. A unique feature of all eukaryotic and certain bacterial KAP NTPases is the presence of two or four transmembrane helices inserted into the P-loop NTPase domain. These transmembrane helices anchor KAP NTPases in the membrane such that the P-loop domain is located on the intracellular side [].

Probab=17.57 E-value=1.8e+03 Score=22.80 Aligned_cols=81 Identities=16% Similarity=0.138 Sum_probs=0.0

Q ss_pred	HHHHHHHHHHhhccccEEEecCCCCcHHHHHHHhc-----ceeEEEeCcHHHHHhHHHHHHHHhhccCCCC
Q gi 350546883 r	134 KNIFISLIKNNLFD-ERMIFVFCDIWLNLNIDSEYIN-----SHNFIFVTND-LRKNVNITQIESCVICND--- 199 (250)
Q Consensus	143 ~~~~s~len~~~~~e~~~v1Kd~d~I~i~~L~e~~n-----n~IiltNd~~~~~i~~f~~iE~~~f~~~~~ 213 (265)
T Consensus	++,.++.+.+.+=...++ ++ - + -.-+...+.+.
T PF07693_consens	157 ~~~~~~1~~~~~iViIDoLDR~~~~~i~~~l~~ik~~~~~i~~~l~~D~~~~~l~~i~~~~~ 228 (325)
T ss_pred	157 EELISKIKKLKESKKRIVIIIDDLRCSPEETVELLEAIKLLLDFPNIFIILAFDP--EILEKAIKNYGE GF----- 228 (325)
	HHHHHHHHHHhhccCceEEEEEccchhcCCCCeEEEEeEcH--HHHHHHHHhhCcCc----

Q ss_pred	EEEEEccHHHHHHHHHHhhCCCC
Q gi 350546883 r	200 FLVEIYDYNKLISYLELKMNTLIN 223 (250)
Q Consensus	214 ~iv-I~~~y~1E~~~~~i~ 237 (265)
T Consensus	+...-..! +. !. +.
T PF07693_consens	229 -----yLeKiq~~~ 246 (325)
T ss_pred	229 -----DEIDGREYLEKIIQVPFS 246 (325)
	-----ccccHHHHHHHHhhCeEEE

HHPRED hits for Csn2 Staphylococcus long

No	Hit	Prob	E-value	P-value	Score	SS	Cols	Query	HMM	Template	HMM
1	PF13304 AAA_21: AAA domain; P	91.0	4.4	3E-05	33.6	7.1	59	175-234	241-302	(303)	
2	COG3593 Predicted ATP-dependen	89.8	1.9	1.3E-05	48.4	5.6	91	132-236	255-347	(581)	
3	3zgx A SMC, chromosome parti	87.6	9.7	6.5E-05	35.0	7.4	60	176-235	334-395	(426)	
4	COG3910 Predicted ATPase [Gene	87.5	3.5	2.3E-05	42.5	5.2	54	182-235	136-192	(233)	
5	g1ii8.1 c.37.1.12 (A,:B:) Rad5	86.8	2.9	1.9E-05	37.0	3.7	35	201-235	311-348	(369)	
6	PIRSF034888 P-loop_UCPO34888	86.4	5.8	3.9E-05	40.2	6.0	61	175-236	308-372	(444)	
7	4i99 A Chromosome partition pr	84.0	4.2	2.8E-05	38.5	3.7	35	201-235	276-312	(354)	
8	g1f2t.1 c.37.1.12 (A,:B:) Rad5	82.0	6.5	4.4E-05	35.4	3.9	35	201-235	234-271	(292)	
9	SUPFAM0046693 c.37.1 P-loop co	80.5	8.3	5.6E-05	30.1	3.5	34	201-234	235-271	(293)	
10	d1e69a c.37.1.12 (A:) Smc hea	79.8	33	0.00022	32.3	7.5	60	176-235	225-286	(308)	
11	PF13175 AAA_15: AAA ATPase do	78.9	21	0.00014	35.2	6.3	57	178-234	351-415	(415)	
12	PF11398 DUF2813: Protein of u	71.3	17	0.00011	39.3	4.1	36	201-236	293-328	(373)	
13	SUPFAM0050337 c.37.1 P-loop co	69.9	24	0.00016	30.2	3.8	36	201-236	363-401	(427)	
14	SUPFAM0050943 c.37.1 P-loop co	66.1	31	0.00021	28.1	3.6	34	201-234	256-291	(330)	
15	SUPFAM0036790 c.37.1 P-loop co	65.9	34	0.00023	27.5	3.7	35	201-235	250-286	(308)	

```
No 1
>PDB13304 AAA 21: AAA domain; PDB: 3QKS_B 1US8_B 1F2U_B 1F2T_B 3QKT_A III8_B 3QKR_B 3QKU_A.
Probab=90.97 E-value=4.4 Score=33.65 Aligned cols=59 Identities=29% Similarity=0.388 Sum probs=0.0
```

No 2
>COG3593 Predicted ATP-dependent endonuclease of the OLD family [DNA replication, recombination, and repair]
Probab=89.82 E-value=1.9 Score=48.39 Aligned cols=91 Identities=29% Similarity=0.317 Sum probs=0.0

Q ss_pred	HHHHHHhcCC--CcEEEEcCcHHHHH		
Q gi 315659845 r	200 RLKQLLISLD--VNIVLTGSMHFLAQ	224	(333)
Q Consensus	212 r~r~l~L~~i~i~iVLT~S~~FLae	236	(345)
	.+#+ + .+ .+ +. .		
T Consensus	321 ~~~l~l~~l~~l~~Q~IvTThS~~~s~	347	(581)
T COG3593	321 VLWDLLNNPLQLQRIVTTSHPILLSL	347	(581)
T ss pred	HHHHHHhcCcccEEEEcCcCCCCC		

No 3
>3zgx_A_SMC, chromosome partition protein SMC; cell cycle; 3.40A (Bacillus subtilis)
Probab=87.60 E-value=9.7 Score=35.02 Aligned cols=60 Identities=17% Similarity=0.113 Sum probs=0.0

Q ss_pred	HHHHHHHHHHHHHHhhCeEEEEEEccCCCCHHHHHHHHHHhhC-CcEEEecCChhhh	
Q gi 315659845 r	164 EQVPLPVIEELKKQHNNQLLLIYLYPEANLSPKEQIRLKQLLISL--DVNIIVLTGSMHFLA	223 (333)
Q Consensus	176 e@vlpLV~leem~n~q~k~l11YLYPEANL~K~e~g~r~r~l~L~L~i~i~V~L~S~F~L~A	235 (345)
T Consensus	334 k~~~lal~a~~~~~ilDPe~Lhp~~~~~l~l~s~s~QiitTHsp~~~~	395 (426)
T 3zgx_A	334 RALTAIALLFSILKVRPVPFCVLDEVEAALDEANVRFAQYLKKYSSDTQFIVITHRKGTME	395 (426)
T ss_dssp	HHHHHHHHHHHHHHHHCSCEEEESTTSCTHHHHHHHHHHHHHHCSSEEEECSCHHHH	
T ss_pred	HHHHHHHHHHHHHHhCCCCEEEECCCCCCCCHHHHHHHHHHHHHHHHHhCCCCEEEEeeCHHHH	

No 4
>COG3910 Predicted ATPase [General function prediction only]
Probab=87.49 E-value=3.5 Score=42.49 Aligned cols=54 Identities=35% Similarity=0.431 Sum probs=0.0

No 5
>gll18.1 c.37.1.12 (A,B:) Rad50 {Archaeon Pyrococcus furiosus [TaxId: 2261]}
Prob-abh-86.76 E-value=2.9 Score=36.99 Aligned cols=35 Identities=11% Similarity=0.218 Sum probs=0.0

Q ss_pred	ccCCCCCHHHHHHHHHHHcC---CcEEEeCcChH		
Q g1 315659845 r	189 PEANLSPKEQIQLRQLLTL---DVNIVILTGSMHFLA	223	(333)
Q Consensus	201 PEAnLs~Keqir~r~l~L---i~iIVLT~S~Fla	235	(345)
T	+ +. . +- ++ ++ ..+ ..-+ + ++.		
T Consensus	311 pe~l~h~~~~~l~~l~~~~~QvitiTHs~~~~~	348	(369)
T glii8.1	311 PTPYLDEERRKLITIMERYLKKIPQVILVSHDEELKD	348	(369)
T ss_dssp	CSSSSSCSHHHHHHHHHHHHHHHTGGGSEEEEEECSGGGG		
T ss_pred	CCCCCCCCHHHHHHHHHHHHHHHHHHHHHHHHHH		

```
No 6
>PIRSF034888 P-loop_UCP034888
Probab=86.44 E-value=5.8 Score=40.15 Aligned_cols=61 Identities=36% Similarity=0.480 Sum_probs=0.0

Q ss pred HHHHHHHHHHHHHHHhhcCeEEEEEEcccCCCCHHHHHHHHHHHHhC---C-CcEEEEEcCChHHHHH
Q g1|315659845|r 163 IEQVLPVLVIEELKKHQHNQLLIYLYPEANLSPKEQIRLKQLLISL---D-VNIIVLTGSMHFLAQ 224 (333)
Q Consensus 175 IEQVLPVLVeeem~q~ll~YIYLYPEANL~KEGir~r~l~v~L---i~iIVLT~S~FLae 236 (345)
    ++++=+====+...+.+=|=||.|=|+..+..+=+ +.+=|=|.+=|+..+
T Consensus 308 ~~~~LPi~La~~~K~~~vIIEPEAHLHP~Q~~~l~~~gr~~~v~~~THSd~~~ 372 (444)
T PIRSF034888 308 VAYVLPVLLALR~LSSREGSLVIIPEEPAHLFKPAQSVLADLLADLVSGRVQQVVTTTHSDHLNR 372 (444)
T ss pred HHHHHHHHHHHHHhhcCcCCCCEEEcccCCCCHHHHHHHHHHHHHHHHHHHHCcEEECCCCHHHHHHHHHHHH
```


No 15

>SUPFAM0036790 c.37.1 P-loop containing nucleoside triphosphate hydrolases (52540) SCOP seed sequence: d1e69a_.
Probab=65.91 E-value=34 Score=27.51 Aligned_cols=35 Identities=34% Similarity=0.552 Sum_probs=0.0

Q ss_pred	ccCCCCHHHHHHHHHHHhcC--CCcEEEECcCChhHHH
Q gi 315659845 r	189 PEANLSPKEQIRLKQLLISL--DVNIIVLTGSMHFLA 223 (333)
Q Consensus	201 PEAnLS~KEqir~r~l~l~i~i~iIVLT~S~FLa 235 (345) .++ .++..+.++..++ .++ .+-+ +. ..+.++.
T Consensus	250 p~~~Ld~~~~~j~~~~~h~~~~~ 286 (308)
T SUPFAM0036790	250 PTSGLDPETREALLLRELAKGRTVIVVTHDLELLA 286 (308)
T ss_pred	ccccCCHHHHHHHHHHHHHHhcCCEEEEccHHHHh

HHPRED hits for Csn2 Staphylococcus short

No	Hit	Prob	E-value	P-value	Score	SS	Cols	Query	HMM	Template	HMM
1	gllii1.8 c_37.1.12 (A:,B:) Rad5	96.9	0.017	1.1E-07	49.4	7.8	79	158-237	282-363 (369)		
2	g1f2t.1 c_37.1.12 (A:,B:) Rad5	96.3	0.045	3E-07	47.6	6.8	59	177-237	224-286 (292)		
3	cd03275 ABC_SMCl_euk Eukaryoti	96.2	0.32	2.2E-06	45.2	11.9	123	110-237	111-237 (247)		
4	3gkt_A DNA double-strand break	95.6	0.34	2.3E-06	43.3	8.8	76	158-235	252-330 (339)		
5	PRK11147 ABC transporter ATPas	95.5	0.31	2.1E-06	51.7	9.8	93	138-236	420-515 (635)		
6	3auy_A DNA double-strand break	95.5	1.2	8E-06	38.9	11.5	153	77-237	209-365 (371)		
7	4i99_A Chromosome partition pr	95.5	0.33	2.2E-06	44.8	8.7	82	155-237	244-327 (354)		
8	SUPFAM0046693 c_37.1 P-loop co	95.5	0.58	3.9E-06	35.8	8.4	73	161-235	207-284 (293)		
9	cd03293 ABC_NrtD_SsuB transpor	95.3	0.75	5E-06	41.5	10.1	93	138-236	112-211 (220)		
10	d1le69a_c_37.1.12 (A:) Sma hea	95.2	0.36	2.4E-06	43.8	8.0	76	161-237	224-301 (308)		
11	TIGR02211 LoloD_lipo_ex lipoprote	95.2	0.82	5.5E-06	41.1	10.0	94	138-239	122-221 (221)		
12	SUPFAM0050337 c_37.1 P-loop co	95.2	0.74	5E-06	38.1	8.7	74	161-235	337-413 (427)		
13	3zgx_A SMC, chromosome partiti	95.0	0.42	2.8E-06	42.7	7.7	77	161-239	333-411 (426)		
14	1w1w_A Structural maintenance	95.0	0.79	5.3E-06	41.5	9.2	77	161-239	338-418 (430)		
15	PRK10247 putative ABC transpor	94.9	1	7E-06	41.2	10.0	94	138-238	117-217 (225)		

```
No 1
>gllii8.1.c.37.1.12 (A;,B:) Rad50 {Archaeon Pyrococcus furiosus [TaxId: 2261]}
Probs=96.90 E-value=0.017 Score=49.40 Aligned cols=79 Identities=10% Similarity=0.151 Sum probs=0.0
```

Q	ss_pred		Ccc		
Q	g1 389815356 r	233	GER	235	(288)
	Consensus	235	ge-	237	(291)
			++		
T	Consensus	361	g-s	363	(369)
T	glib8.1	361	GSS	363	(369)
T	ss_dssp		SSE		
T	ss_pred		CEE		

```
No 2
>g1f2t.1 c.37.1.12 (A:,B:) Rad50 {Archaeon Pyrococcus furiosus [TaxId: 2261]}
Probab=96.35 E-value=0.045 Score=47.57 Aligned cols=59 Identities=14% Similarity=0.197 Sum probs=0.0
```

No 3
 >cd03275 ABC_SMC1_euk Eukaryotic SMC1 proteins. SMC proteins are large (approximately 110 to 170 kDa), and each is arranged into five recognizable domains. Amino-acid sequence homology of SMC proteins between species is largely confined to the amino- and carboxy-terminal globular domains. The amino-terminal domain contains a 'Walker A' nucleotide-binding domain (GxxGxGKS/T, in the single-letter amino-acid code), which by mutational studies has been shown to be essential in several proteins. The carboxy-terminal domain contains a sequence (the DA-box) that resembles a 'Walker B' motif, and a motif with homology to the signature sequence of the ATP-binding cassette (ABC) family of ATPases. The sequence homology within the carboxy-terminal domain is relatively high within the SMC1-SMC4 group, whereas SMC5 and SMC6 show some divergence in both of these sequences. In eukaryotic cells, the proteins are found as heterodimers of SMC1 paired with SMC3, SMC2 with SMC4, and SMC5 with SMC6 (for Probab=96.25 E-value=0.32 Score=45.21 Aligned cols=123 Identities=11% Similarity=0.083 Sum probes=0.0

No 4
>3qkt_A DNA double-strand break repair RAD50 ATPase; RECA-like fold, coiled-coils, ATP binding, DNA bindi MRE11, replication; HET: DNA ANP; 1.90A
{Pyrococcus furiosus} PDB: 3qku_A* lii8_A 3qks_B* 3qkr_B* lii8_B
Probab=95.57 E-value=0.34 Score=43.27 Aligned cols=76 Identities=12% Similarity=0.135 Sum probs=0.0

Q	ss_pred	C
Q	gi 389815356 r	233 G 233 (288)
Q	Consensus	235 g 235 (291)
		+
T	Consensus	330 ~ 330 (339)
T	3qkt_A	330 N 330 (339)
T	ss_dssp	T
T	ss_pred	C

No 5
>PRK11147 ABC transporter ATPase component; Reviewed
Probab=95.53 E-value=0.31 Score=51.69 Aligned_cols=93 Identities=16% Similarity=0.246 Sum_probs=0.0

Q ss_pred	CCHHHHHHrcCceeeeeeCCCc	
Q gi 389815356 r	213 TSPQVNVGVLNSNHLIKRTGE	234 (288)
Q Consensus	215 ~hP~I~i~v~n~LiK~nge +++++.++ .+..++.+ +	236 (291)
T Consensus	495 Hd~~~~~d~i~~l~~g~~~	515 (635)
T PRK11147	495 HDROFVDNT~VTECWLIFEGNGE	515 (635)
T ss_pred	CCHHHHHHr-CCEEEEEECCe	

No 6
>3auy_A DNA double-strand break repair RAD50 ATPase; DNA repair, ABC transporter ATPase domain-like; HET: DNA ADP; 2.70A (Methanocaldococcus jannaschii) PDB: 3auy_A* 3av0_B*
Probab=95.51 E-value=1.2 Score=38.93 Aligned_cols=153 Identities=13% Similarity=0.067 Sum_probs=0.0

```

Q ss_pred          CCCc
Q g1_389815356|r 231 RTGER 235 (288)
Q Consensus       233 ~nge~ 237 (291)
                           .+|.+.
T Consensus        361 ~~~~s 365 (371)
T 3auv_A          361 DGNVS 365 (371)
T ss_dssp         SSSCE
T ss_pred         CCCCC

```

No 7
>4i99_A Chromosome partition protein SMC; winged-helix domain and SMC head domain, chromosome condensin SCPB, DNA binding protein; 2.30A {Pyrococcus furiosus}
Probab=95.47 E-value=0.33 Score=44.85 Aligned_cols=82 Identities=13% Similarity=0.182 Sum_probs=0.0

Q ss_pred		CCCC		
Q gi 389815356 r	231	RTGER	235	(288)
T Consensus	233	~nge~	237	(291)
		.+ .+		
T Consensus	323	~~~s	327	(354)
T 4199_A	323	RDGV	327	(354)
T ss_dssp		SSSC		
T ss_pred		ECCce		

No 8
 >SUPFAM0046693 c.37.1 P-loop containing nucleoside triphosphate hydrolases (52540) SCOP seed sequence: d1f2t.1.
 Probab=95.46 E-value=0.58 Score=35.81 Aligned cols=73 Identities=16% Similarity=0.195 Sum probs=0.0

No 9
 >cd03293 ABC_NrtD_SsuB_transporters NrtD and SsuB are the ATP-binding subunits of the bacterial ABC-type nitrate and sulfonate transport systems, respectively. ABC transporters are a large family of proteins involved in the transport of a wide variety of different compounds, like sugars, ions, peptides, and more complex organic molecules. The nucleotide binding domain shows the highest similarity between all members of the family. ABC transporters are a subset of nucleotide hydrolases that contain a signature motif, Q-loop, and H-loop/switch region, in addition to, the Walker A motif/P-loop and Walker B motif commonly found in a number of ATP- and GTP-binding and hydrolyzing proteins.
 Probabs=95.32 F-value=0.75 Score=41.53 Aligned cols=93 Identities=14% Similarity=0.162 Sum probss=0.0

Probab=95.32 E-value=0.75 Score=41.53 Aligned_cols=93 Identities=14% Similarity=0.162 Sum_probs=0.0

Q	s_pred		EEccccHHHHHHCceeeeeeccc	
Q	g1_389815356 r	210	CLTSPSVINQVGLSNVHLKR-TGE	234 (288)
Q	Consensus	212	i-T~hP~i~t~i~~~nv~LiK~nge	236 (291)
T	Consensus	187	+ + + +.++ +.++ +.++ +. +	
T	cd03293	187	i~sH~~~~~d~i~l~~~G~	211 (220)
T	s_pred		LVTHD1DEAVFLDRVVLSRAPGR	211 (220)
T			EEccccHHHHHHCcEEEEEECCCC	

```
No 10  
>d1e69a_c.37.1.12 (A:) Smc head domain (Thermotoga maritima [TaxId: 2336])  
Probab=95.25 E-value=0.36 Score=43.76 Aligned_cols=76 Identities=13% Similarity=0.133 Sum_probs=0.0
```

No 11
>TIGR02211 *LolD_lipo_ex* lipoprotein releasing system, ATP-binding protein. This model represents *LolD*, a member of the ABC transporter family (pfam00005). *LolD* is involved in localization of lipoproteins in some bacteria. It works with a transmembrane protein *LolC*, which in some species is a paralogous pair *LolC* and *LolE*. Depending on whether the residue immediately following the new, modified N-terminal Cys residue, the nascent lipoprotein may be carried further by *LolA* and *LolB* to the outer membrane, or remain at the inner membrane. The top scoring proteins excluded by this model include homologs from the archaeal genus *Methanosa*cina.
P-value=0.22, E-value=0.83, Score=41.08, Aligned_col=94, Identities=13%, Similarity=0.382, Sum_pscore=0.0

Probab=95.22 E-value=0.82 Score=41.08 Aligned_cols=94 Identities=13% Similarity=0.282 Sum_probs=0.0

Q	ss_pred	EEeCCCCHHHHRRCcCeeeeeeeCCccc	
Q	gi 389815356 r	210 CLTTSPQVINQVGLSNVWHLIKRTGERYS	237 (288)
Q	Consensus	212 i~T~h~P~i~L~I~i~v~n~V~K~ng~e~ ++ +.++ +.++ +.++ ++ ++	239 (291)
T	Consensus	197 ~T~h~m~m~m~m~d~v~v~l~G~i~ TIGR02211 197 VVTTHDLEELAKKL~DRVLEMKG~DQLFNM	221 (221) 221 (221)
T	ss_pred	EEeCCCCHHHHRRhbcc~CCEEEEEE~CCEcc	

No 12
>SUFCAF0050337 c.37.1 P-loop containing nucleoside triphosphate hydrolases (52540) SCOP seed sequence: dlwiwa_.
Probab=95.16 E-value=0.74 Score=38.14 Aligned cols=74 Identities=18% Similarity=0.277 Sum probs=0.0

Q ss_pred		HHHHHHhhhhhccCccccEEEEEEChhcCHHHHHHHHHHHHh---	ceeEEEeCCHHHHHhcCceeeeeecc	
Q gi 389815356 r	159	QYQSFLLKSWLHLVHNKATNVCFYDPENELENKSELKELFQFATSQ--	-ACTMCLTTSQPQVINQVGLSNVHLIKRTG	233 (288)
Q Consensus	161	qrk-Fid~~~L-h~~keNvl~~lFPE-hlg~~sd~K~f~~~~~t~Li~t~h~b~i~n~v~Li~ng	+++.+++.+++.+++.++ = +.+++.+++.+++.++ .+++.+++.+++.+++.+++.++.	235 (291)
T Consensus	337	e~~~~la~~~~~1~~~~~1~~~~~1~~~~~1~~~~~1~~~~~h~~~~~g~~~~~		413 (427)
T SUPFAM0050337	337	EKQLRLALALALALIAELLLPPPLLDEPTAGLDPEENRERLLELRELASEGGQVIVTVTHDPELAARL-ADRLVLIKDDG		413 (427)

```

Q ss pred          c
Q gi|389815356|r 237 S 237 (288)
Q Consensus       239 ~ 239 (291)
                           +
T Consensus        411 s 411 (426)
T 3zgx_A          411 S 411 (426)
T ss_dssp          B
T ss pred          E

```

No 14
>1wlw_A Structural maintenance of chromosome 1; cohesin, chromosome segregation, cell adhesion, kleisin, MIT cell cycle; HET: AGS; 2.90A
{Saccharomyces cerevisiae} SCOP: c.37.1.12
Probab=94.95 E-value=0.79 Score=41.46 Aligned cols=77 Identities=8% Similarity=0.119 Sum probs=0.0

No 15

>PRK10247 putative ABC transporter ATP-binding protein YbbL; Provisional
Probab=94.94 E-value=1 Score=41.20 Aligned_cols=94 Identities=13% Similarity=0.127 Sum_probs=0.0

Q ss_pred CCC
Q gi|389815356|r 235 RYS 237 (288)
Q Consensus 237 ~~~ 239 (291)
 .++
T Consensus 416 ~~~ 418 (430)
T 1wlw_A 416 NSS 418 (430)
T ss_dssp TEE
T ss_pred Cce

Q ss_pred HHHHHhhcCccc--hhccccchHHHHHHHHHHHHHHccCCcccEEEEECchhcCHHHHHHHHHHHHHHh----ceeeE
Q gi|389815356|r 136 FEQLIKIAPIE---VRTVSDEPMGAYQYQSFLLKSWLHLVHNKATNVCFYDFPENELNKSELKELFQFATSQ----ACTM 208 (288)
Q Consensus 138 l~klik~f~~d---fhn~lddin~~qrk~Fid~~~l~h~~keNvl~~lFPE~hLg~sd~Kf~~~t~ 210 (291)
 .+++++.+++ .+....++-++|+=-..++ ..|++|=+-|...-..+++.+.|+
T Consensus 117 ~~~~l~~~l~~~~~LS~G~~qrv~laral~~~~~p~111DEPT~LD~~~l~~~l~~~~~tv 191 (225)
T PRK10247 117 FLDDLERFALPDTILTAKNIAELSGGEKQRISLIRNLQF----MPKVLLDEITSALDESNKHNVNEIIHRYVREQNIAV 191 (225)
T ss_pred HHHHHHHhCCChHHhCCcccCCHHHHHHHHHHHHHhC----CCCEEEEeCCcccCCHHHHHHHHHHHHHhCCCE

Q ss_pred EEEeCCHHHHhCcceeecCCCC
Q gi|389815356|r 209 ICLTSPQVINQVGLSNVHLIKRTGERY 236 (288)
Q Consensus 211 Ii~T~hp~~i~~~nv~LiK~nge~~ 238 (291)
 ||+|.++..+. . .+..+..++..+
T Consensus 192 ii~sh~~~~~d~i~~l~~~~~ 217 (225)
T PRK10247 192 LWVTHDKDEIN-H-ADKVITLQPHAGEM 217 (225)
T ss_pred EEEECChHHH-h-CCEEEEeccccH

HHPRED hits for Csn2 short

No	Hit	Prob	E-value	P-value	Score	SS	Cols	Query	HMM	Template	HMM
1	2y15_A Beta-N-acetylhexosamini	68.5	45	0.0003	36.0	6.5	70	141-210	3-112	(442)	
2	d1t5ia_c.37.1.19 (A): Spliceo	66.9	64	0.00043	29.8	6.3	45	163-212	13-57	(168)	
3	d1s2ma2_c.37.1.19 (A:252-422)	64.2	67	0.00045	29.6	5.9	45	163-212	18-62	(171)	
4	d2gjx1a_c.1.8.6 (A:167-528) be	51.3	81	0.00054	33.3	4.7	61	150-210	5-88	(362)	
5	d2rb4a1_c.37.1.19 (A:307-474)	51.2	1.3E+02	0.00085	28.1	5.4	48	160-212	15-62	(168)	
6	c0d06564 GH20_DspB_LnbB-like G1	50.5	1.2E+02	0.00079	31.6	5.6	61	150-210	4-99	(326)	
7	d1oywa3_c.37.1.19 (A:207-406)	49.2	2.2E+02	0.0015	26.8	6.7	44	164-212	17-60	(200)	
8	d1fukfa_c.37.1.19 (A): Initiat	48.9	2.7E+02	0.0018	25.9	7.1	44	164-212	14-57	(162)	
9	2hzjv_A ATP-dependent RNA helic	48.3	2.4E+02	0.0016	25.6	6.6	45	163-212	21-65	(163)	
10	PF14258 DUF4350: Domain of un	47.5	2.1E+02	0.0014	22.7	5.4	59	142-212	10-69	(70)	
11	c0d02742 GH20_hexosaminidase Be	47.1	1.9E+02	0.0013	29.9	6.4	61	150-210	3-89	(303)	
12	SUPFM00408920_c.1.8 (Trans)gly	46.0	2.4E+02	0.0016	26.4	6.4	61	150-210	5-107	(443)	
13	c0d06565 GH20_GcnA-like Glycosy	45.7	2.2E+02	0.0015	29.6	6.6	61	150-210	3-77	(301)	
14	1t51_A C_terminal domain of A	45.2	2.5E+02	0.0017	25.9	6.3	45	163-212	17-61	(172)	
15	PF09664 DUF2399: Protein of u	44.3	1.1E+02	0.00077	28.6	4.1	31	22-64	20-50	(152)	

No 1
 >2y15_A Beta-N-acetylhexosaminidase; hydrolase; 2.15A (Streptococcus pneumoniae) PDB: 2yla A* 2y19_A*
 Probab=68.49 E-value=45 Score=36.01 Aligned_cols=70 Identities=16% Similarity=0.270 Sum_probs=0.0

```

Q ss_pred      ----ChHrHCCCCHHHHHHHHHHHHHHCCeEE
Q g1_339625078|r 170 ---NASLYCKLSDLNLHECLLAEGVNLI 195 (224)
Q Consensus    185 ---Nl~~Ylt~~~el~~L~~~i~~~l~~~v~l 210 (251)
                .-+.|..+|..++|..++|..++|..+|
T Consensus    83 ~~~~~~YT~~~di~~ElV~YA~~rgI~VI 112 (442)
T 2y15_A       83 YYDDPFNGTALTQAEVTELIEYAKSKDIGH 112 (442)
T ss_dssp      HCCCCCTCSCBHHHHHHHHHHHHHHHTTCCEE
T ss_pred      cccCCCCCcHHHHHHHHHHHHHHHCeEE

```

```
No 2
>d1t5ia_c.37.1.19 (A) Spliceosome RNA helicase BAT1 (UAP56) {Human (Homo sapiens) [TaxId: 9606]}
Probab=66.89 E-value=64 Score=29.83 Aligned cols=45 Identities=13% Similarity=0.269 Sum_probs=0.0
```

No 3
>d1s2ma2 c.37.1.19 (A:252-422) Putative ATP-dependent RNA helicase DHH1 (Baker's yeast (*Saccharomyces cerevisiae*) [TaxId: 4932])
Probab=64.23 E-value=67 Score=29.63 Aligned cols=45 Identities=11% Similarity=0.121 Sum probs=0.0

Q	ss_pred	HHHHHHHHHHHHhhCcCEEEEEECHhCCHHHHHHHHHHHHHhCCeEEEE		
Q	gi 339625078 r	418 GKIIEVIIHVMGLMDESRYFLTNASLYCKLSDLNQLHECLLAEGVNLI SI	197	(224)
	Consensus	163 eki~~~iki~~~kll~fvNl~~Ylt~el~L~~i~~~l~vl~	212	(251)
		+ +....+.... + + +....+++.++.		
T	Consensus	18 K~~~L~~~l~~~k~~~i~vF~~~~~~l~~~L~~~g~~~	62	(171)
T	dis2ma2	18 QKLHCLNTLFSKLQIINQAIIFCN--STNRVELLAKKITLDGYSYY S	62	(171)
T	ss_dssp	GHHHHHHHHHHHHCCCCCCCCCCCC		
T	ss_pred	HhhHHHHHHHHHHCCCCCCCCCCCC		

No 4
>2d7jxal c.1.8.6 (A:167-528) beta-hexosaminidase A (Human (Homo sapiens) [TaxId: 9606])
Probab=51.25 E-value=81 Score=33.35 Aligned cols=61 Identities=13% Similarity=0.296 Sum probs=0.0

```

Q ss_pred          CeEE
Q gi|339625078|r 192 VNLI  195 (224)
Q Consensus       207 l-vL  210 (251)
               +_+_
T Consensus        85 I~VI   88 (362)
T dgjxal          85 IRVL   88 (362)
T ss_dssp          CEEE
T ss_pred          CeEE

```

No 5
>d2rb4a1 c.37.1.19 (A:307-474) ATP-dependent RNA helicase DDX25 (Human (*Homo sapiens*) [TaxId: 9606])
Probab=51.19 E-value=1.3e+02 Score=28.07 Aligned cols=48 Identities=8% Similarity=0.167 Sum probs=0.0

No 6
 >cd06564 GH20_DspB_LnbB-like Glycosyl hydrolase family 20 (GH20) catalytic domain of dispersin B (DspB), lacto-N-biosidase (LnbB) and related proteins. Dispersin B is a soluble beta-N-acetylglucosaminidase found in bacteria that hydrolyzes the beta-1,6-linkages of PGA (poly-beta-(1,6)-N-acetylglucosamine), a major component of the extracellular polysaccharide matrix. Lacto-N-biosidase hydrolyzes lacto-N-biose (LNB) type I oligosaccharides at the nonreducing terminus to produce lacto-N-biose as part of the GNB/LNB (galacto-N-biose/lacto-N-biose I) degradation pathway. The lacto-N-biosidase from *Bifidobacterium bifidum* has this GH20 domain, a carbohydrate binding module 32, and a bacterial immunoglobulin-like domain 2, as well as a YSIRK signal peptide and a G5 membrane anchor at the N and C termini, respectively. The GH20 hexosaminidases are thought to act via a catalytic mechanism in which the catalytic nucleophile is not provided by solvent or the enzyme, but by the substrate itself.
 Probab=50.45 E-value=1.2e+02 Score=31.57 Aligned_cols=61 Identities=18% Similarity=0.315 Sum_probs=0.0

Q	ss_pred	HHHHHHHHHHHCCeEE		
Q	g1 339625078 r	180 LNQLCLLAEAGVNL	195 (224)	
Q	Consensus	195 1~L~i~~~i~~~l~v~L	210 (251)	
		+.+ .+ ++..+. +		
T	Consensus	84 i~eiv~A~y~g~r~i~V~I	99 (326)	
T	cd06564	84 FELIYAKDRGVNII	99 (326)	
T	ss_pred	HHHHHHHHHHCCeEe		

No 7
>d1loywa3 c.37.1.19 (A:207-406) RecQ helicase domain {Escherichia coli [TaxId: 562]}
Probab=49.19 E-value=2.2e+02 Score=26.83 Aligned cols=44 Identities=7% Similarity=0.167 Sum_probs=0.0

No 8
>difuka_c.37.1.19 (A:) Initiation factor 4a (Baker's yeast (*Saccharomyces cerevisiae*) [TaxId: 4932])
Probab=48.91 E-value=2.7e+02 Score=25.86 Aligned cols=44 Identities=14% Similarity=0.210 Sum probs=0.0

Q ss_pred	H~~~~~HHHHHHHHhhCCCEEEEChHHhCCHHHHHHHHHHHHHhCCeEEE	
Q gi 339625078 r	149 KIEEVIHVMGMLDESRYLFLTNASLYCKLSDLNQLHECLLAEGVNLSI	197 (224)
Q Consensus	164 ki~~~ki~~~l~~kkllyfvnl~Ylt~~~el~~l~~~i~~~l~v~i	212 (251)
	+....++...+ ++ + ..+....++....++..+	
T Consensus	14 K~~~L~~~l~~~k~iF~~~~~-~~~~~l~~~L~~~~~	57 (162)
T difuka_	14 KYECLTBLDYSLVTQAVIFN----TRRKVEELTTKLNRDKFVSAI	57 (162)
T ss_dssp	HHHHHHHHHHHHHTTCSCEEEEEE-----SHHHHHHHHHHHTTCEEEE	
T ss_pred	HHHHHHHHHHHHCCCCCEEEEEEE-----EeChHHHHHHHHhCCeEEE	

No 9
 >2hvj_A ATP-dependent RNA helicase DBPA; parallel alpha-beta, hydrolase; 1.95A (Bacillus subtilis)
 Probable=48.29 F-value=2.4e+02 Score=25.60 aligned cols=45 Identities=18% Similarity=0.144 Sum probs=0.0

No 10
>PF14258 DUF4350: Domain of unknown function (DUF4350)
Probab=47.4% E-value=2.1e02 Score=22.67 Aligned cols=59 Identities=22% Similarity=0.207 Sum proba=0.0

Q ss_pred		HHHHHHhCcEeccCCCCHHHHHHHHHHHHHHhCCCEEEEChHHhCCH-HHHHHHHHHHHHhCcEEE	
Q gi 339625078 r	127	LERLRLHSQNIVVEKGNNSTIFGKIEEVIVHGMGLDESRYLFLTNASLYCKL-SDLNQLHECLLAEGVNLSI	197 (224)
Q Consensus	142	~~~llk~~~kiae~~~~~ekia~~iki~~l~~kkllvfvnl~~~lyt+~~el~~l~~i~~i~~l~~vl~i	212 (251)
		+.+++++.+ --..+++.++..+....+ --...+++.+ ++ .+++.++# ++.	
T Consensus	10	~~~L~~g~~v~~~~~l~~~t~~~tll~i~~~~~l~~~v~~~G~~l~v~l	69 (70)
T PF14258_consen	10	YQVLLER~PQGVKEERKK~PYAELEA~DDGTLLVIGPDRLSPEFEA~AGNTLVIA	69 (70)
T ss_pred		HHHHHHHCCCCeEEccc~~CHHHhCCH-~~~~CCEEEECCCCCCCCHHHHHHHHHHHHCCCCeEE	

No 11
 >cd02742 GH20_hexosaminidase Beta-N-acetylhexosaminidases of glycosyl hydrolase family 20 (GH20) catalyze the removal of beta-1,4-linked N-acetyl-D-hexosamine residues from the non-reducing ends of N-acetyl-beta-D-hexosaminides including N-acetylglucosides and N-acetylgalactosides. These enzymes are broadly distributed in microorganisms, plants and animals, and play roles in various key physiological and pathological processes. These processes include cell structural integrity, energy storage, cellular signaling, fertilization, pathogen defense, viral penetration, the development of carcinomas, inflammatory events and lysosomal storage disorders. The GH20 enzymes include the eukaryotic beta-N-acetylhexosaminidases A and B, the bacterial chitobiases, dispersin B, and lacto-N-biosidase. The GH20 hexosaminidases are thought to act via a catalytic mechanism in which the catalytic nucleophile is not provided by the solvent or the enzyme, but by the substrate itself.
 Probab=47.13 E-value=1.9e+02 Score=29.93 Aligned cols=61 Identities=18% Similarity=0.322 Sum_probs=0.0

Q ss_pred	CcEeccCCCcHHHHHHHHHHHHHHhhCcCCE-EEEE--	-ChHHhCCHHHHHHHHHHHH	
Q g1 339625078 r	135 NIVVEKGNNSTKIGEVIHVGMLDESSR-LFLT-----	-NASLYCCLLSDLNQLHECLL	188 (224)
Q Consensus	150 +kie~~~~~ +ki~~~~l~~~kkl~lfv~-----	-N~+YLT~el~l~~~~i~	203 (251)
	+.+ +...=-+.-+ ..+ +.+....-+ +==+-----	--.+. + +.+ .+ ++	
T Consensus	3 G~mld@Ar~~~~~lk~id~ma~K~N~1hl~D~~~~~le~~~p~l~~~g~~~~~yT~d~el~iv~yA	82 (303)	
T cd02742	3 GIMLDVSRHLFSVSEIKRTDVLARYKINTFHWHLTDQAWRIESKFFPELAEGKGQINPRSPGGFYTAQKLDIYEAYA	82 (303)	
T ss_pred	ccccccccCcHHHHHHHHHHHHHHhhCcEEEEeeccCccEeeCccchhhhcccccCCCCCcEeCHHHHHHHHHHHHH		

Q ss_pred	HhCCeEE	
Q gtl339625078 r	189 AEGVNLL	195 (224)
Q Consensus	204 ~~~l~vL	210 (251)
	..++. +	
T Consensus	83 ~rgl~vi	89 (303)
T cdo2742	83 ARGIEVI	89 (303)
T ss_pred	HccccEE	

No 12
>SUPFAM0048920 c.1.8 (Trans)glycosidases (51445) SCOP seed sequence: dlqbaa3; InterPro: IPR017853.
Probab=46.02 E-value=2.4e+02 Score=26.43 Aligned_cols=61 Identities=10% Similarity=0.224 Sum_probs=0.0

Q ss_pred	CcEeccCCCcHHHHHHHHHHHHHHHHcCCEEEE--	-ChH		
Q gi 339625078 r	135 NIVVEGGNNSTIFGKIEEVIVHVMGLMDESRYLFIT--	-NAS	172	(224)
Q Consensus	150 ~kicke~~~~~ekia~~iki~~~kklivfv~	-Nl~	187	(251)
	+.+ +...-.-.+ ... +.+....+.-.	..-		
T Consensus	5 G~~D~~R~~~~~k-N~~l~~h D~~~~~P~l~~~~~		84	(443)
T SUPFM0048920	5 GLMLDVARHFFSVEITIKRLIDAMALYKLNLVHLTDQGWRLEIKSYPELTEVGAYRGSDLVETALLPQLSLPDGG		84	(443)
T ss_pred	ccccccCCCcHHHHHHHHHHHHHHHHcCCEEEE--ecccCCc~~ecccCcHHhhcccccchheccccccccCCCC			

No 13
 >cd06565 GH20_GcnA-like Glycosyl hydrolase family 20 (GH20) catalytic domain of N-acetyl-beta-D-glucosaminidase (GcnA, also known as BhsA) and related proteins. GcnA is an exoglucosidase which cleaves N-acetyl-beta-D-galactosamine (NAG) and N-acetyl-beta-D-galactosamine residues from 4-methylumbelliferylated (4MU) substrates, as well as cleaving NAG from chito-oligosaccharides (i.e. NAG polymers). In contrast, sulfated forms of the substrate are unable to be cleaved and act instead as mild competitive inhibitors. Additionally, the enzyme is known to be poisoned by several first-row transition metals as well as by mercury. GcnA forms a homodimer with subunits comprised of three domains, an N-terminal zincin-like domain, this central catalytic GH20 domain, and a C-terminal alpha helical domain. The GH20 hexosaminidases are thought to act via a catalytic mechanism in which the catalytic nucleophile is not provided by solvent or the enzyme, but by the substrate itself.
 Probab=45.73 E-value=2.2e+02 Score=29.59 Aligned cols=61 Identities=7% Similarity=0.221 Sum probs=0.0

No 14
>1t51_A_C-terminal domain of A probable ATP-dependent RNA helicase; RECA-like fold, PRE-mRNA processing protein; 1.90A (Homo sapiens) SCOP: c.37.1.19
Brohabas45 25 E-value=2.5e+02 Score=25.94 Aligned cols=45 Identities=13% Similarity=0.269 Sum prob=0.0

No 15
>PFO9664 DUF2399: Protein of unknown function C-terminus (DUF2399); InterPro: IPR024465 This domain is found in archaeal, bacterial and eukaryotic proteins. Its function is unknown.
Probab=44.26 E-value=1.1e+02 Score=28.56 Aligned cols=31 Identities=13% Similarity=0.228 Sum probs=0.0

C

Proteins with structure most similar to *S. thermophilus* Csn2 structure (3zth)

	Target	Q-score	P-score	Z-score	Seq%
1w1w:A	<i>S. cerevisiae</i> Smc1	0.1447	-0	6.767	11.73
4i99:A	<i>P. furiosus</i> Smc	0.1404	0.6379	8.12	11.9
1vpl:A	<i>T. maritima</i> ABC transporter ATP-binding protein	0.1374	-0	5.211	13.38
2onk:A	<i>A. fulgidus</i> Molybdate/tungstate ABC transporter, ATP-binding protein	0.1346	-0	5.001	7.273
4glu:D	<i>Y. pestis</i> Hemin import ATP-binding protein HmuV	0.1308	1.04e-09	5.957	13.69
4hlu:C	<i>T. maritima</i> Energy-coupling factor transporter ATP-binding protein EcfA	0.1288	-0	5.253	11.52
2olk:A	<i>G. stearothermophilus</i> Amino acid ABC transporter	0.1271	-0	5.316	10.19
1g6h:A	<i>M. jannaschii</i> high-affinity branched-chain amino acid transport ATP-binding protein	0.1266	-0	5.862	10.69
4fi3:C	<i>E. coli</i> Vitamin B12 import ATP-binding protein BtuD	0.1253	-0	5.085	10.78
2fgk:B	<i>E. coli</i> Alpha-hemolysin translocation ATP-binding protein hlyB	0.1231	-0	4.874	12.26
2yz2:A	<i>T. maritima</i> Putative ABC transporter ATP-binding protein TM_0222	0.1219	-0	5.316	0.1562
1l2t:A	<i>M. jannaschii</i> Hypothetical ABC transporter ATP-binding protein MJ0796	0.1216	0.001364	5.357	0.1241

Proteins with structure most similar to *S. pyogenes* Csn2 structure (3toc)

	Target	Q-score	P-score	Z-score	Seq%
2onk:A	<i>A. fulgidus</i> Molybdate/tungstate ABC transporter, ATP-binding protein	0.1986	0.05449	4.77	9.028
2pcj:A	<i>A. aeolicus</i> Lipoprotein-releasing system ATP-binding protein loloD	0.1959	0.00532	4.49	7.692
3gfo:A	<i>C. perfringens</i> Cobalt import ATP-binding protein cbiO 1	0.1766	0.1979	5.312	14.39
1ji0:A	<i>T. maritima</i> ABC transporter TM_1139	0.1728	0.859	5.647	15.15
3n70:D	<i>E. coli</i> P-loop NTPase domain of the Sigma-54 transport activator	0.172	0.09933	3.876	6.522
2fgj:A	<i>E. coli</i> Alpha-hemolysin translocation ATP-binding protein hlyB	0.1675	0.3295	5.372	8.759
2olk:B	<i>G. stearothermophilus</i> Amino acid ABC transporter ArtP	0.1624	-0	4.349	4.511
3vx4:A	<i>S. mutans</i> Putative ABC transporter, ATP-binding protein ComA	0.1614	0.493	5.853	6.923
2ihy:A	<i>S. aureus</i> ATPase subunit of ABC transporter	0.1602	-0	4.327	11.03
1g9x:C	<i>M. jannaschii</i> High-affinity branched-chain amino acid transport ATP-binding protein	0.1597	-0	4.683	10.08
1l7v:C	<i>E. coli</i> Vitamin B12 transport system permease protein Btuc	0.1591	8.58e-10	4.305	8.209

3zth:Ass	-----EE-EE-----EE-EE-----EEEEEE--HHHHHHHHHHHHHHHHH--HH-----HHHHHH
3zth:A (1)	--X--KF-FV-QH--P-----Y--K-----ERIE-LNI-G--AITQIVGQNNELKYYTWOIISWYF--GG--KK-----YSSEDLS-----IFDYEEP-----TILDEAREIVKRSSY-----
1w1w:Ass	--EE--EE-EEE-----EE-E-EE-----EEEEEE-----HHHHHHHHHHH--HH-----HH-----HHH-----EEEEEEEEE-----EEE-----EEEEE
1w1w:A (2)	GRLV--GL-ELSNF--K-----SY-RG-----VT-K-VGFGES-NFTSIIGP-NGS--GKSNNMMDAIS--FVLGV-----LK-----DLIYRGPQSAYVKAF-----YQKGKNLVELMRI1SRNGDT
4i99:A (2)	--PYI-EKLE--LKGFK-----S-YGNKKV--VIPFS-K--GFTAIVGA-NGS--GKSNIGDAILFVLG--GLSAKAXRASRISDLIFAGSKNEPPAKYAE-----VAIY-----FNNEDRGFP1IDEDEVVIRR
1vp1:A (2)	--GAVVV-KDLRK--RIG-----KKEILK-----GI-S-FEI--EEGEIIFGLGP-NGA--GKTTTLRIIS--T-----L-----I-----KPS-----SGIVT-----VFGKNV
2onk:A (1)	--MF--LK-VRAEK--RLG-----NFR--L-----NV-D-FEM-GR-DYCVLIGP-TGA--GKSVFLELIA--G-----I-----V-----KPD-----RGEVR-----LNGADIT
4glu:D (10)	--ALL-EA-SHLHY-HVQ-----QQALIN-----DV-S-LHI-ASGEMVAIIGP-NGA--GKSTLLRLLT--G-----Y-----L-----SPS-----HGECH-----LLGQNLN
4hlu:C (999)	GSGRI-EL-NSVSF--RYN-----GDYVLK-----DV-N-AEF-ETGKIYVVVGK-NGS--GKTTLLKILA--G-----L-----L-----AA-----AGEIF-----LDGSPAD
2olk:A (1)	---MDIV-HQLKK--SFG-----SLEVLK-----GI-N-VHI--REGEVVVVVIGP-SGS--GKSTFLRCLN-----L-----L-----E-----DFD-----EGRII-----IDGINLKA
1g6h:A (4)	-TMEILRT-ENIVK--YFG-----EFKALD-----GV-S-ISV-NKGDVTLLIIGP-NGS--GKSTLLINVIT--G-----F-----L-----KAD-----EGRVY-----FENKDIT
4fi3:C (2)	--SIVMLQ-QDVAE--S-----TRLG-----PL-S-GEV--RAGEILHLVGP-NGA--GKSTLLARMA--G-----M-----T-----SG-----KGSIQ-----FAGQP
2fgk:B (467)	--DI-TF-RNRF--RYKP-----DSPVILD-----NI-N-LST-KQGEVIGIVGR-SGS--GKSTLLTKLIQ--R-----F-----Y-----IPE-----NGQVL-----IDGHDLA
2yz2:A (1)	--MRI-EV-VNVSH--IFHRGTPLEKKALE-----NV-S-LVI-NEGECLLVAGN-TGS--GKSTLLQIVA--G-----L-----I-----EPT-----SGDVL-----YDGERK
11t2:A (1)	--MI-KL-KNVTK--TYKMG-EEIYIYALK-----NV-N-LNI-KEGEFVSIIMP-SGS--GKSTMLNIIG--C-----L-----D-----KPT-----EGEVY-----IDNIKTNDL

3toc:Ass	HHHH-----HHHHHHHHHH-----H-----HHHHHHHHHHHH-----EEEEEE-H-HHH-----HHHHHHHHHHHHHH-----EEEEEE-----EEEEEE-EEE-----
3toc:A	ECLENELDLEYDEITILELIKSLGVKVEQSDF-T--IFEKCLEILQIFKYLTK--KKLLIFVNS-GAFL--TKDEVASLQEYISLTN--L-TVLFLEFREL--Y-D---F--P--QYILDLITKNX-
2pcj:Ass	-----HHHHHHHHHH-----HHHHHH-HHHHHHHHHHHHH-----EEEEEE-----HHHHHHHHHHHHHHHHHH-----EEEEEE-----HHHH-H-----H-----EEEEEE-EeeeeEE-----
2pcj:A	-----KERGEYLLSELGLGDKLSKPYELS-GGEQQRVAIARALA--NE----PILLFADEP-TGNLDSANTKRVMDFILKINEG-G---T-SIVMVT--HEREL-A----E-LTH---RTLEMKDGVKVGEITRV-----
Zonk:A	-----RRVREMAEKLGIHAHLLDKPARLS-GGERQRVALARALV--IQ----PRLLLDEP-LSAVDLKTKGVLMEEFLRFVQREFD--V-PILHVT--HDLIE-AA----M-LAD---EVAVMLNDRIVEKGKLKEF-----
3gfo:A	-----RKRVDNALKRTGTIEHLKDPTHCLS-FGOKKRVAAIGVLV--ME----PKVLLIDEP-TAGLDPMGVSEIMKLLEVQMKELG--I-TIIIA----DIDIV-P----L-YCD---NVFVMKEGRVILQGNPKEVFA-----
1ji0:A	-----RDLEWIFSLSFPRLKERLKQLGGTLS-GGEQQXLAIGRALX--SR----PKLXXDEP-SLGLAPILVSEVFVEIQKINQE-G---T-TILLVE--QNALGAL----K-VAH---YGYVLETGQIVLEGKASELL-----
3n70:D	-----PDNAP-QLNDFIALA--Q----G-GTLVLSH-PEHL---TREQQYHVLQQLSQSEH--RPFRIGI--GDTSL-VEIAELYCFA----XTQ-----
2fgj:A	-----KLAGAHDFISELREGYNTIVGEQGAGLSSGQ-RQRIAIARALV--NN----PKILIFDEATSALD--YESEHVIMRMNHICKG--R-TVIIIA--A--R-L----S-TVKN-ADRIIVMEKGKIVEQGKHKELL-----
2olk:B	-----AEAKAMELLDKVGLKDKAHYPDLSL-GGOAQQRVAIARALA--ME----PKIMLFDEP-TSALDPEMVGVLVSMVKQLANE-G---M-TMVVVT--H--E-M----G-FAREVGDRVLFDGGYIIEEGKPEDLFL-----
3vx4:A	-----CEIAEIRSDIEQMPQGYQTELSDGAGISGGQ-KQRIALARALL--TQ----APVLLDAA-TSSL--DILTEKKIISNLNQM-TE--K-TIIFVA--H--R-L----S-ISQR-TDEVIVMDQGKIVEQGTHKELLA-----
2ihy:A	-----NEAHQLLKLVGXSAKAQQYIGYLS-TGEKVRXIAARALX--GQ----PQVLILDEP-AAGLDFIARESLLSILSDSDSYT--L-AXIYVTH--FI-EE-I----T-ANF--SKILLKDGQSITQQGAVEDEF-----
19gx:C	IPKEEMVEAKFALKFLKSHLDKAGELES-GGQMKLVEIGRALM--TN----PKMIVMDP-IAGV--APGLAHDIFNHVL-EL-KAKGI-TFLIIEH--R--L-D----I-VLYN-IDLHLYVMFNGQIJAEGRGEET-----
117v:C	-----LLNDVAGALALDDKLGHSTNQLSGGE-WQRVRLAAVVL--QITPQANPAQOLLNIDEP-XNSL--DVAQOSALDKILSALCQOGL-AIVXSS--H--D-L----N-HTLRHAHRAWLLKGGKXLASGRREEVLT-----

3zth:Ass	HHH-----HHHHHHHHH-----EEE--EEHHH-----HH-----HHHHHHHHHHHHHHHHH-----
3zth:A	XNRHINLNCQVEYHLENPLTLTLDQLLTKNFSPFFAIENKNLSFE-----WV-----SNIDKLSLFLEXLDHLLSQE-----TTE
1w1w:Ass	-----HHHHHHHHH-HHHHHHHH-----HHHHHHH--EEEE--EEEE-----HHHHHHHHHHHHHHHHHHH-----
1w1w:A	-----QSPVELSRMFTFDYVSDH-----LDAIYRELTGNASLTKYHATPPLKRFK-DMEYLSGGEKTVAAALLFAINSYQP-----S
4i99:A	-----SPLERRLLIDISGIKEKKNVFXRTFEAISRNFSEIFAKLSPGGSGARLILENPEDPFSGGLEIAKPKGDKVKRIEAXSGGEKALTALAFVFAIQKF-KP-----A
1vpl:A	-----RNMQGIEYLRFVAGFYASSSSE-----IEEMVERATEIAGLGEKIKDRVST-----YSKGMVRKLLIARALMVN-----P
2onk:A	-----PHLSVYRNIAYGLRNVERVER-----DRRVREMAEKLGIAHLLDRKPA-----RLSGGERQRVALARALVIQ-----P
4glu:D	-----RQALQQVMAQTDCCLALAQRDYL-----VL-----SGEQQRVQLARVLAQL-----W
4hlu:C	-----MRKRIKKVLELVGLSGLAAADPL-----NL-----SGGQKQRLAIAASMLARD-----T
2olk:A	-----PHMTVLNNITLAPMKVRKP-----KAEAKAMELLDKVGLKDKAHAYPD-----SLSGGQAQRVAIARALAME-----P
1g6h:A	-----G-----EL-----SGGQMKLVEIGRALMTN-----P
4fi3:C	-----N-----QL-----SGGEWQRVRLAAVVLQITPQANPAG-----P
2fgk:B	-----AKLAGAHDFISELREGYNTIVGEQGA-----GL-----SGGQRQRIATARALVNN-----P
2yz2:A	-----FAERVFDEVAFAVKNFYPRDRP-----VPLVKKAMEFVGGLDFDSFKDRVPF-----FLSGGEKRRVIAIASVIVHE-----P
1l2t:A	-----PLLTALENVELPLIFKYRGAMSG-----EERRKRRALECLKMAELEERFANHKPN-----QLSGGQQQRVAIARALANN-----P

A

312126587 | *Caldicellulosiruptor hydrothermalis* 108
 43408285; *Stanieria cyanosphaera* PCC 7437
 414076241; *Anabaena* sp. 90
 428399778 | *Calothrix* sp. PCC 6303
 17233043 | *Nostoc* sp. PCC 7120
 166363864 | *Microcystis aeruginosa* NIES-843
 166367659 | *Microcystis aeruginosa* NIES-843
 307150343 | *Cyanotheca* sp. PCC 7822
 307153352 | *Cyanotheca* sp. PCC 7822
 428306661 | *Crinalium epiphamnum* PCC 9333
 307150912 | *Cyanotheca* sp. PCC 7822
 428775289 | *Halotheca* sp. PCC 7418
 428779516 | *Dactylococcopsis salina* PCC 8305
 428778599 | *Dactylococcopsis salina* PCC 8305
 428781696 | *Dactylococcopsis salina* PCC 8305
 428215045 | *Oscillatoria acuminata* PCC 6304
 428313425 | *Microcoleus* sp. PCC 7113
 158340109 | *Acarachloris marina* MBIC11017
 428775002 | *Halotheca* sp. PCC 7418
 307150700 | *Cyanotheca* sp. PCC 7822

312126663 | *Caldicellulosiruptor hydrothermalis* 108
 312621424 | *Caldicellulosiruptor krotonskvensis* 2002
 222530402 | *Caldicellulosiruptor bescii* DSM 6725
 222530431 | *Caldicellulosiruptor bescii* DSM 6725

307591386 | *Cyanotheca* sp. PCC 7822
 302391131 | *Acetohalobium arabaticum* DSM 5501
 435854579 | *Bacteroides halobius* DSM 5150
 435852864 | *Bacteroides halobius* DSM 5150
 435854272 | *Bacteroides halobius* DSM 5150
 313123202 | *Lactobacillus delbrueckii* ND02
 773620250 | *Pseudoaalteromonas haloplanktis* TAC125
 298675374 | *Methanohalobium evestigatum* Z-7303
 298674707 | *Methanohalobium evestigatum* Z-7303
 298675090 | *Methanohalobium evestigatum* Z-7303
 169834784 | *Clostridium botulinum* A3 str. Loch Maree
 229887454 | *Clostridium botulinum* B4 str. 657

297585104 | *Bacillus selenitireducens* MLS10
 188587532 | *Natronarrobacter thermophilus* JW-NM-WN-LF
 312793946 | *Caldicellulosiruptor kristjanssonii* 177R1B
 344996431 | *Caldicellulosiruptor lactoaceticus* 6A
 217077332 | *Thermosiphon africanus* TCF52B
 375009168 | *Geobacillus thermolevoevarans* CCB US3 UF5
 407477159 | *Exiguobacterium antarcticum* B7
 172057416 | *Exiguobacterium sibiricum* 255-15
 169823684 | *Finegoldia magna* ATCC 29328
 169092252 | *Petrotoga mobilis* SJ95
 169092616 | *Petrotoga mobilis* SJ95

333978211 | *Desulfotomaculum kuznetsovii* DSM 6115
 260892456 | *Ammonifex degensii* KC4
 260893631 | *Ammonifex degensii* KC4
 302391857 | *Acetohalobium arabaticum* DSM 5501
 428778891 | *Dactylococcopsis salina* PCC 8305
 260892735 | *Ammonifex degensii* KC4
 260893993 | *Ammonifex degensii* KC4
 433655673 | *Thermoaerobacterium M0795*
 332982963 | *Mahella australiensis* 50-1 BON
 332983053 | *Mahella australiensis* 50-1 BON
 222530385 | *Caldicellulosiruptor bescii* DSM 6725
 298674346 | *Methanohalobium evestigatum* Z-7303
 298675173 | *Methanohalobium evestigatum* Z-7303
 292490518 | *Nitrosococcus halophilus* NC 4
 428778759 | *Dactylococcopsis salina* PCC 8305
 300114477 | *Nitrosococcus watsonii* C-113
 292491781 | *Nitrosococcus halophilus* NC 4
 292491869 | *Nitrosococcus halophilus* NC 4
 292491893 | *Nitrosococcus halophilus* NC 4
 292491036 | *Nitrosococcus halophilus* NC 4
 292490884 | *Nitrosococcus halophilus* NC 4
 292491492 | *Nitrosococcus halophilus* NC 4
 297563003 | *Noardiopsis dassonvillei* DSM 43111
 374989417 | *Streptomyces bingchenggensis* BCW-1
 374989654 | *Streptomyces bingchenggensis* BCW-1
 271963428 | *Streptosporangium roseum* DSM 43021
 271964894 | *Streptosporangium roseum* DSM 43021
 258511478 | *Alicyclobacillus acidocaldarius* DSM 446
 169092264 | *Petrotoga mobilis* SJ95
 121604978 | *Polaromonas naphthalenivorans* CJ2
 347754630 | *Chloracidobacterium thermophilum* B
 218667603 | *Acidithiobacillus ferrooxidans* ATCC 23270
 218667415 | *Acidithiobacillus ferrooxidans* ATCC 23270
 344201059 | *Acidithiobacillus ferrivorans* SS3
 344201127 | *Acidithiobacillus ferrivorans* SS3
 288940385 | *Allochromatium vinosum* DSM 180
 288941921 | *Allochromatium vinosum* DSM 180
 288940786 | *Allochromatium vinosum* DSM 180
 288941171 | *Allochromatium vinosum* DSM 180
 288941306 | *Allochromatium vinosum* DSM 180
 121582913 | *Polaromonas naphthalenivorans* CJ2
 121604604 | *Polaromonas naphthalenivorans* CJ2
 91791204 | *Polaromonas* sp. JS666
 91791283 | *Polaromonas naphthalenivorans* CJ2
 121582916 | *Polaromonas naphthalenivorans* CJ2
 121582983 | *Polaromonas naphthalenivorans* CJ2
 428304158 | *Crinalium epiphamnum* PCC 9333
 428204921 | *Chroococcidiopsis thermalis* PCC 7203
 307153539 | *Cyanotheca* sp. PCC 7822
 428316193 | *Oscillatoria acuminata* PCC 6304
 414079584 | *Anabaena* sp. 90
 172055147 | *Cyanotheca* sp. ATCC 51142
 428316510 | *Oscillatoria nigro-viridis* PCC 7112
 428314951 | *Oscillatoria nigro-viridis* PCC 7112
 307591386 | *Cyanotheca* sp. PCC 7822
 75812341 | *Anabaena variabilis* ATCC 29413
 17233293 | *Nostoc* sp. PCC 7120
 75812782 | *Anabaena variabilis* ATCC 29413
 434047609 | *Cylindrospermum stagnale* PCC 7417
 414078405 | *Anabaena* sp. 90
 440685156 | *Anabaena cylindrica* PCC 7122
 428315656 | *Oscillatoria nigro-viridis* PCC 7112
 428204926 | *Chroococcidiopsis thermalis* PCC 7203
 218440244 | *Cyanotheca* sp. PCC 7424
 428775092 | *Halotheca* sp. PCC 7418
 414075475 | *Anabaena* sp. 90
 428305120 | *Crinalium epiphamnum* PCC 9333
 427713881 | *Synechococcus* sp. PCC 6312

S. pyogenes Cas9
 S. pyogenes Cas9 domains
 A. naeslundii Cas9
 A. naeslundii Cas9 domains

(7) KKYSIGL
 RRTTYTT

(12) LRGVI
 RTTYY

Supplementary Figure S8. Distant Cas9 homologs. Compare with Figure 2. (A) Multiple sequence alignment of Cas9 distant homologs (See Supplementary Material and Methods). Sequences are denoted by Genbank identifiers and the complete organism name. Mapped sequences and domains of *A. naeslundii* and *S. pyogenes* Cas9 are shown in the last rows of the alignment with catalytic residues highlighted in yellow. For Cas9 domain description, see Supplementary Figure S4. Mapping was done manually according to HHPRED hits for Cas9 homologs shown in (B). (B) HHPRED hits for Cas9 distant homologs. Top ten hits and selected hits to Cas9, RuvC and transposase structures are shown.

312165687 Caldicellulosiruptor hydrothermalis 108	DPGSKMEGVSKS--	-KA	-HTVNLQI-AYA	-VTHVKSAVET	-RRNMRARRRY--	-RK-	-TRK
34408285 Stianieria cyanosphaera PCC 7437	DPGSKREGVSKS--	-KA	-HQVNLQI-ATT	-VDWNEHHLKT	-RREMRARRRGG-	-RK-	-TPC
414076241 Anabaena sp. 90	DPGKLFGIAVQSK--	-KK	-FTLQLMLHVLVP	-FKTVKDRMDQ	-RSMMRNRKRG--	-RRINRK-	-LSP--NK
428299778 Calothrix sp. PCC 6303	DPGKMFMSGVAVQSK--	-KK	-YTLQMLHVLVP	-FKTVKRNMRQ	-RSIMRGRGKG--	-RRINRK-	-LSP--KK
17233043 Nostoc sp. PCC 7120	DPGKFFGIGVQSK--	-AK	-FTLYTAHILP	-FQTVRDMDA	-RRLMRGRGKG-	-RRIDRK-	-VNF--LK
166367659 Microcystis aeruginosa NIES-843	DPGKLFGIGVQSK--	-SL	-FTLWKAHLELP	-FKRVKERLDN	-RRLMRGRGKG-	-RRINRQ-	-LSP--NL
166367659 Microcystis aeruginosa NIES-843	DPGKLFGIGVQSK--	-SL	-FTLWKAHLELP	-FKRVKERDMN	-RRLMRGRGKG-	-RRINRQ-	-LSP--NL
307150343 Cyanothec sp. PCC 7822	DPGKLFGIGVQSK--	-SS	-YSLWTHGLELP	-FKTVKRMED	-RMMRMRARRG-	-RRINRN-	-LPP--EL
307153352 Cyanothec sp. PCC 7822	DPGKLFGIGVQSK--	-SS	-YSLWAGHLELP	-FTKIKRMED	-RMMRMRTRRG-	-RRINRQ-	-LPP--EL
428306661 Crinalium epiphamnum PCC 9333	DPGKLFGIGVQSK--	-SR	-FTLWTAHLELP	-FKRVKERDMN	-RRLMRGRGKG-	-RRINRQ-	-LPP--NC
310510912 Cyanothec sp. PCC 7822	DPGKLFGIGVQSK--	-SH	-FSLWLKAHLELP	-FKRVKERDMN	-RRLMRGRGKG-	-RRINRQ-	-ISF--ES
428775289 Halotheca sp. PCC 7418	QAEFHNSGIGVQSK--	-SK	-TTLFATAHLELP	-FQVQRKDMET	-RAMMRNRGKG-	-RRINRK-	-LPP--DQ
428779516 Dactylococcopsis salina PCC 8305	DPGKHYSGIGVQSK--	-SF	-MTLFATAHLELP	-FQVQRKDMET	-RAMMRNRGKG-	-RRIDRK-	-LPP--NO
428778599 Dactylococcopsis salina PCC 8305	DPGKHYSGIGVQSK--	-SK	-ATLFATAHLELP	-FQVQRKDMET	-RAMMRNRGKG-	-RRINRK-	-LPP--DQ
428781696 Dactylococcopsis salina PCC 8305	DPGKHYSGIGVQSK--	-SQ	-ATLFATAHLELP	-FQVQRKDMET	-RAMMRNRGKG-	-RRINRK-	-LPP--DQ
428215045 Oscillatoria acuminata PCC 6304	DPGKLGYGIGVQSK--	-AK	-ATLFWMAHILP	-FQTVKDRMEO	-RMMRMRNRKG-	-RRINRQ-	-LEY--SK
428313425 Miccoleus sp. PCC 7113	DPGKHYSGIGVQSK--	-SK	-VTLFMAHILP	-FQS1KDRMEO	-RMMRMRNRKG-	-RRINRQ-	-APY--NO
158340109 Acarychloris marina MBLIC11017	DPGKSYSVIGIAVOS--	-AH	-FTLYTAHILP	-FQR1KERLAAGAVIKNGKV1KVNVRGALRVRVKG-	-RINRK-	-VPF--HL	
428775002 Halotheca sp. PCC 7418	DPGKSYSVIGVQSK--	-AK	-CTLLQLHILP	-FGVRVRSRMDQ	-RLLRRLRSRS-	-RRINRD-	-VPF--KL
310510700 Cyanothec sp. PCC 7822	DPGKFFGIGVQSK--	-SL	-FTLWKAHLELP	-FQVQRERIDN	-RGLMRGRGKG-	-RRINRK-	-ISF--ES
312126663 Caldicellulosiruptor hydrothermalis 108	DGGYHICGFVVC-EL--	-KGKGL	-IVYCTGILVSR	-IPEKIELITE	-RKAHRRNRRYYSSRCKKRLSARONRVLTKF		
312621424 Caldicellulosiruptor kromotskyensis 2002	DGGYHHGPAV-EL--	-RWGVL	-IVCYCIGVLETR	-IPEIKDLMTK	-RGYRNRNRRYYHSRCRKRMKSQSVLTKF		
322530402 Caldicellulosiruptor bescii DSM 6725	DGGYHHGPAV-EL--	-RWGVL	-IVCYCIGVLETR	-IPEIKDLMTK	-RGYRNRNRRYYHSRCRKRMKSQSVLTKF		
222530431 Caldicellulosiruptor bescii DSM 6725	DGGYHHGPAV-EL--	-RWGVL	-IVCYCIGVLETR	-IPEIKDLMTK	-RGYRNRNRRYYHSRCRKRMKSQSVLTKF		
307591386 Cyanothec sp. PCC 7822	DPGSKTTGIALVQDD-	-	-KLINGAELIHR	-GQQ1KDNLIT	-RRQIRRSRRN-	-RK-	-TRY
302391131 Aceto halobium arabaticum DSM 5501	DGDGAKVYGVVALQDC-	-KTKN	-KVLFKAKMVKQR	-QDVSAAKME	-RNDYGRYRGG-	-RK-	-TRY
435854579 Halobacteroides halobius DSM 5150	DGDGKVGFLVQIQLKC-	-QSKN	-KVLFKATMKMR	-QEVSKLMLTQ	-RREYRRYRQQ-	-HK-	-TRY
435852864 Halobacteroides halobius DSM 5150	DGDGKVGFLVQIQLKC-	-QTKN	-KVLFKAVMEQR	-QDVSKMKEE	-RGYRGRYRRS-	-HK-	-TRY
313123202 Lactobacillus delbrueckii ND2	DGTQRH1GFAIVSNS-	-QTKN	-KVLHQSVEVELR	-QDVHKNLYT	-RK1YRRSRKN-	-RK-	-TRY
77362205 Pseudoaletromonas haloplanktis TAC125	DGTSKHKVGAACALAN-	-	-KIVYQSEVELR	-TDVSKMMQO	-RAMYRTRRNN-	-RK-	-LRY
288675374 Methanohalobium evestigatum z-7303	DGSGSKY1GCAAVAND-	-	-EVMYQSEVELR	-DDVSKMMKD	-RADHMRMRRY-	-RK-	-TRY
288674707 Methanohalobium evestigatum z-7303	DGSGSKH1GCAAVAND-	-	-EVLYQSEVELR	-DDVSKVQVO	-RASYRNRNRS-	-RK-	-TRY
288675090 Methanohalobium evestigatum z-7303	DGSGSKH1GCAAVAND-	-	-EVLYQSEVELR	-DDVSKRMKD	-RADYRNRNRS-	-RK-	-TRY
169384784 Clostridium botulinum A3 str. Loch Maree	DSCYLN1GFSAITKE-	-	-NEL1SGEVKLL	-KGMERLKE	-KAMYRQRQRS-	-R-	-LRY
229587454 Clostridium botulinum Ba4 str. 657	DSCYLN1GFSAITKE-	-	-KEL1SGEVKLL	-KCMERLKE	-KAMYRQRQRS-	-R-	-LRY
297585104 Bacillus selenitireducens MLS10	DAGTKHIGFASAPTEK-	-	-DVLFEQDVQLR	-TD1QDKLAT	-RRQYRRSRRS-	-RK-	-TRY
188587532 Natranaerobius thermophilus JW-NM-WN-LF	DAGTKHVGGLSATTK-	-	-EVLFGEAQLR	-TN1QELLST	-RREARRAARN-	-RK-	-TRY
312739496 Caldicellulosiruptor kristjanssonii 177RIB	DIGSKVGVGSIA1TEK-	-	-QELFSSEVELR	-QDIRKLILLE	-RREHRRFRRY-	-RK-	-TRY
344996431 Caldicellulosiruptor lactoaceticus 6A	DIGSKVGVGSIA1TEK-	-	-QELFSSEVELR	-QDIRKLILLE	-RREHRRFRRY-	-RK-	-TRY
210770372 Thermosiphon africanus TCF52B	DIGSKVGVGSIA1TEK-	-	-QELLSMEVELR	-QDIRKLILLE	-RREYRNRNRRY-	-RK-	-TRY
375009168 Geobacillus thermeleovorans CCB US3 UF5	DGDTCHV1GSA1TCK-	-	-DVLFEAAEOLR	-TD1VELLA1	-RQFQRRSRRN-	-RK-	-TRY
407747159 Exiguobacterium antarcticum B7	DAGTKHIGFASPTATT-	-	-KVLLEGEVQLR	-TD1QDLILLAT	-RRAAMRSARRS-	-RK-	-TRY
172057416 Exiguobacterium sibiricum 255-15	DAGTKHIGFASPTATT-	-	-KVLLEGEVQLR	-TD1QDLILLAT	-RRAAMRSARRS-	-RK-	-TRY
169823684 Finegoldia magna ATCC 29328	DGSGSKY1GSLASTEN-	-	-KET1SAEYELR	-ND1VKLSS-	-REECRTRRN-	-R-	-LRY
160902522 Petrotoga mobilis S95	DAGSKS1GSLASTEN-	-	-KEL1FAAEVEIR	-DD1PKL1SQ	-KQYRKGRRF-	-RK-	-TRY
160902616 Petrotoga mobilis S95	DAGSKTGT1GSLASTEN-	-	-KEL1FASEVEIR	-DD1PKL1SQ	-KQYRDRDRL-	-RK-	-TRY
333978211 Desulfotomaculum kuznetsovii DSM 6115	DGDSKWG1ALVNH--	-TG	-EVVFRGTLIQR	-GDVVR-LLT	-REYTRNRRYY-	-RL-	-VRH
620892456 Ammonifex degensis KC4	DGDSRYG1ALVNH--	-TG	-EVVFRGLVLR	-GDVVR-LLT	-REYRGRYRYY-	-R1-	-VRH
620893631 Ammonifex degensis KC4	DGDSRYG1ALVNH--	-TG	-EVVFRGLVLR	-GDVVR-LLT	-REYRGRYRYY-	-R1-	-VRH
320391857 Aceto halobium arabaticum DSM 5501	DGDSKTG1ALVNING-	-	-QVAFANLHHR	-ADT1KQNLQT	-RREHRRFRRS-	-Q-	-LKY
428778891 Dactylococcopsis salina PCC 8305	DGDTATGW1QTCGCE-	-	-EII1FMEMLKR	-GGL1KQLSLEQ	-RADFRIGKRRS-	-RN-	-LRY
260893993 Ammonifex degensis KC4	DGDSYGT1GSLASTEN-	-GGKE	-VVVYFAEVHVR	-TDVPGKLLS	-RQARQRRSRR-	-RK-	-TRY
433655673 Thermomoanaerobacterium thermosphaerolyticum M0795	DGDSYGT1GSLASTEN-	-GGKE	-VVVYFAEVHVR	-TDVPGKLLS	-RQARQRRSRR-	-RK-	-TRY
332982963 Mahella austriensis 50-1 BON	DGDSYGT1GSLASTEN-	-TG	-EVVFRGTLIQR	-GDVVR-LLT	-REYTRNRRYY-	-RL-	-VRH
332983053 Mahella austriensis 50-1 BON	DGDSYGT1GSLASTEN-	-TG	-EVVFRGLVLR	-GDVVR-LLT	-REYRGRYRYY-	-R1-	-VRH
222530385 Caldicellulosiruptor bescii DSM 6725	DGDSKTG1ALVNING-	-	-QVAFANLHHR	-ADT1KQNLQT	-RREHRRFRRS-	-Q-	-LKY
288674346 Methanohalobium evestigatum z-7303	DGDSYGT1GSLASTEN-	-K	-DVIWLDQVHHR	-TD1KKLDD	-RNPFRRRRKS-	-QN-	-LRY
288675173 Methanohalobium evestigatum z-7303	DGDSYGT1GSLASTEN-	-K	-DVIWLAQ1HHR	-TD1KKLDD	-RSPFRRRRKS-	-KN-	-LRY
229405198 Nitrosococcus halophilus NC 4	DGDSYGT1GSLAQF-	-NSGT	-VCI1FAMOLEHR	-GRQ1ECSLDS	-RGVGRSRRSRR-	-RN-	-TRY
428778759 Dactylococcopsis salina PCC 8305	DGDSYGT1GSLAQF-	-G	-EVVWRMOLERE	-GG1IKRKLREQ	-RALRRLRKKR-	-R-	-LRY
300114477 Nitrosococcus watsonii C-113	DGDSYGT1GSLAQF-	-PQQR	-VVLWAGNHLTHR	-GG1QARDLER	-RALRRLRKKR-	-R-	-LRY
229419781 Nitrosococcus halophilus NC 4	DGDSYGT1GSLAQF-	-PQQR	-VVLWAANLTHR	-GG1YARERL	-RSLRRLRGG-	-RK-	-TRY
22941869 Nitrosococcus halophilus NC 4	DGDSYGT1GSLAQF-	-PQQR	-VVLWAANLTHR	-GG1YARERL	-RSLRRLRGG-	-RK-	-TRY
229419893 Nitrosococcus halophilus NC 4	DGDSYGT1GSLAQF-	-PQQR	-VVLWAANLTHR	-GG1YARERL	-RSLRRLRGG-	-RK-	-TRY
22941036 Nitrosococcus halophilus NC 4	DGDSYGT1GSLAQF-	-SVQR	-VVLWAANLHR	-GQS1VDRDLER	-RSLRRLRGG-	-RK-	-TRY
229419492 Nitrosococcus halophilus NC 4	DGDSYGT1GSLAQF-	-PQQR	-VVLWAANLHR	-GQS1VDRDLER	-RSLRRLRGG-	-RK-	-TRY
297563003 Nocardiopsis dassonvillei DSM 43111	DGDSYGT1GSLAQF-	-PQQR	-VVLWAANLHR	-GQS1VDRDLER	-RSLRRLRGG-	-RK-	-TRY
374989417 Streptomyces bingchenggensis BCW-1	DGDSYGT1GSLAQF-	-AGER	-RGRYAVLHDHR	-GAV1KRRQ	-RAYRGRGRR-	-SN-	-LRY
374989654 Streptomyces bingchenggensis BCW-1	DGDSYGT1GSLAQF-	-AGER	-RGRYAVLQDH	-GAV1KRRQ	-RAYRGRRRT-	-SH-	-LRY
271963428 Streptosporangium roseum DSM 43021	DGDSYGT1GSLAQF-	-GGSR	-TGRYALQ1LHDHR	-GAI1KDLKAS	-RALRRLRGRS-	-RN-	-LRY
271964894 Streptosporangium roseum DSM 43021	DGDSYGT1GSLAQF-	-GGSR	-IGLYALQ1LHDHR	-GQ1Q1DLKAS	-RALRRLRGRS-	-RN-	-LRY
288511478 Alcylobacillus acidocaldarius DSM 446	DAGSKTGT1GVALVADF-	-QRKX	-TWWVAEE1QHR	-GQ1Q1RALLQ	-RMLRRLRGG-	-RK-	-TRY
160902264 Petrotoga mobilis S95	DAGSKTGT1GVALVDF-	-KNGK	-K1W1GAE1HHR	-GQ1Q1KALDT	-RGVGRSRRSRR-	-RK-	-TRY
121604978 Polaromonas naphthalenivorans CJ2	DAGSKTGT1GVALVDF-	-	-EVGADTGV1QPTM-	-GAT1K1SLGA	-RSAMRRLRGG-	-R-	-LRY
374754630 Chloracidobacterium thermophilum B	DAGSKTGT1GVALVRES-	-	-QEVGADTGV1QPTM-	-GAT1K1SLGA	-RSLRRLRGG-	-R-	-LRY
218667603 Acidithiobacillus ferrooxidans ATCC 23270	DAGSKTGT1GVALVREK-	-	-TIVDVTG1VLR1A-	-GQ1Q1KALDT	-RGAFRPERRR-	-Q-	-LRY
226674145 Acidithiobacillus ferrooxidans ATCC 23270	DAGSKTGT1GVALVREO-	-	-ESVDED1G1QIRKA-	-I1V1MLLEQHR	-RRAFRPERRR-	-N-	-LRY
344201059 Acidithiobacillus ferrivorans SS3	DAGSKTGT1GVALVRE-	-	-APGSKTGT1GVALVRA-	-TIVD1ATGV1TPTV-	-RRAFRPERRR-	-K-	-LRY
442011217 Acidithiobacillus ferrivorans SS3	DAGSKTGT1GVALVRE-	-	-APGSKTGT1GVALVRA-	-VW1MLLEQHR	-RGHRLRARR-	-K-	-TRY
288940385 Allochromatium vinosum DSM 180	DAGSKTGT1GVALVRE-	-	-DAGSKTGT1GVALVRE-	-HGLN1GELAHR	-RGHRLRARR-	-R-	-TRY
288941921 Allochromatium vinosum DSM 180	DAGSKTGT1GVALVRE-	-	-DAGSKTGT1GVALVRE-	-HGLW1GELVHR	-RGHRLRARR-	-R-	-TRY
288940786 Allochromatium vinosum DSM 180	DAGSKTGT1GVALVRE-	-	-DAGSKTGT1GVALVRE-	-HGLW1GELVHR	-RGHRLRARR-	-R-	-TRY
288941171 Allochromatium vinosum DSM 180	DAGSKTGT1GVALVRE-	-	-DAGSKTGT1GVALVRE-	-HGLW1GELVHR	-RGHRLRARR-	-R-	-TRY
288941306 Allochromatium vinosum DSM 180	DAGSKTGT1GVALVRE-	-	-DAGSKTGT1GVALVRE-	-HGLW1GELVHR	-RGHRLRARR-	-R-	-TRY
121582913 Polaromonas naphthalenivorans CJ2	DAGSKTGT1GVALVRE-	-	-ESVDASTGI1RHH-	-GRO1SEALTA	-RGQMRRRR-	-N-	-LRC
121582983 Polaromonas naphthalenivorans CJ2	DAGSKTGT1GVALVRE-	-	-ESVDASTGI1RHH-	-GRO1SEALTA	-RGQMRRRR-	-N-	-LRC
428304158 Crinalium epiphamnum PCC 9333	DAGSKTGT1GVALVRE-	-	-ERGAV-	-AVLN1LFEMLHR	-RGH1RLLRGG-	-K-	-LRY
428204921 Chroococcidiopsis thermalis PCC 7203	DAGSKTGT1GVALVRE-	-	-DAGSKTGT1GVALVRE-	-ERGAV-	-RGH1RLLRGG-	-K-	-LRY
307153539 Cyanothec sp. PCC 7822	DAGSKTGT1GVALVRE-	-TG	-EV1WAGELTHR	-GQ1KISEALTA	-RGQMRRRR-	-N-	-LRY
428126193 Oscillatoria acuminata PCC 6304	DAGSKTGT1GVALVRE-	-TG	-EV1WAAELVHR	-GQ1KISEALTA	-RGQMRRRR-	-N-	-LRY
414079584 Anabaena sp. 90	DAGSKTGT1GVALVQNN-	-	-DAGSKTGT1GVALVQNN-	-GSA1KELES	-RGQMRRRR-	-R-	-TRY
4282316510 Oscillatoria nigro-viridis PCC 7112	DAGSKTGT1GVALVQNN-	-TG	-RMI1GAELEHR	-GFR1KDSL5	-RGQMRRRR-	-R-	-TRY
4282314951 Oscillatoria nigro-viridis PCC 7112	DAGSKTGT1GVALVQNN-	-TG	-DAGSKTGT1GVALVQNN-	-GQ01KNDLES	-RGQMRRRR-	-R-	-TRY
30751386 Cyanothec sp. PCC 7822	DAGSKTGT1GVALVQNN-	-TG	-DAGSKTGT1GVALVQNN-	-GQ01KNDLES	-RGQMRRRR-	-R-	-TRY
758123341 Anabaena variabilis ATCC 29413	DAGSKTGT1GVALVQNN-	-ND	-DAGSKTGT1GVALVQNN-	-GQ01KNDLES	-RGQMRRRR-	-R-	-TRY
172332933 Nostoc sp. PCC 7120	DAGSKTGT1GVALVQNN-	-NG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
578127822 Anabaena variabilis ATCC 29413	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
434407609 Cyanothece stagnale PCC 7417	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
414078405 Anabaena sp. 90	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
404685156 Anabaena cylindrica PCC 7122	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
288215656 Oscillatoria nigro-viridis PCC 7112	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
282049244 Cyanothec sp. PCC 7424	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
428775092 Halotheca sp. PCC 7418	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
414075475 Anabaena sp. 90	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
428305120 Crinalium epiphamnum PCC 9333	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY
427713881 Synechococcus sp. 6312	DAGSKTGT1GVALVQNN-	-TG	-EVVPAEELKHR	-GFAIRDAITS	-RGQMRRRR-	-R-	-TRY

S. pyogenes Cas9

A. naeslundii Cas9

A. *naeslundii* Cas9

312126587 | *Caldicellulosiruptor hydrothermalis* 108
 434408285 | *Stanieria cyanosphaera* PCC 7437
 414076241 | *Anabaena* sp. 90
 428299778 | *Calothrix* sp. PCC 6303
 17233043 | *Nostoc* sp. PCC 7120
 166363864 | *Microcystis aeruginosa* NIES-843
 166367659 | *Microcystis aeruginosa* NIES-843
 3071510343 | *Cyanothece* sp. PCC 7822
 3071513352 | *Cyanothece* sp. PCC 7822
 428306661 | *Crinalium epiphamnum* PCC 9333
 307150912 | *Cyanothece* sp. PCC 7822
 428775289 | *Halotheca* sp. PCC 7418
 428779516 | *Dactyloccoccopsis salina* PCC 8305
 428778599 | *Dactyloccoccopsis salina* PCC 8305
 428781696 | *Dactyloccoccopsis salina* PCC 8305
 428215045 | *Oscillatoria acuminata* PCC 6304
 428313425 | *Microcoleus* sp. PCC 7113
 158340109 | *Acarychloris marina* MBIC11017
 428775002 | *Halotheca* sp. PCC 7418
 307150700 | *Cyanothece* sp. PCC 7822

 312126663 | *Caldicellulosiruptor hydrothermalis* 108
 312621424 | *Caldicellulosiruptor kronotskyensis* 2002
 222530402 | *Caldicellulosiruptor bescii* DSM 6725
 222530431 | *Caldicellulosiruptor bescii* DSM 6725

 307951386 | *Cyanothece* sp. PCC 7822
 302391131 | *Acetohalobium arabaticum* DSM 5501
 435654579 | *Halobacteroides halobius* DSM 5150
 435652864 | *Halobacteroides halobius* DSM 5150
 435654272 | *Halobacteroides halobius* DSM 5150
 313123202 | *Lactobacillus delbrueckii* ND02
 773620520 | *Pseudoaeromonas haloplanktis* TAC125
 298675374 | *Methanohalobium evestigatum* Z-7303
 298674707 | *Methanohalobium evestigatum* Z-7303
 298675090 | *Methanohalobium evestigatum* Z-7303
 169834784 | *Clostridium botulinum* A3 str. Loch Maree
 229587454 | *Clostridium botulinum* Ba4 str. 657

 297585104 | *Bacillus selenitireducens* MLS10
 188507532 | *Natronaerobium thermophilus* JW-NM-NN-LF
 312793946 | *Caldicellulosiruptor kristjanssonii* 177RIB
 344996431 | *Caldicellulosiruptor lactoaceticus* 6A
 217077332 | *Thermosiphon africanus* TCF52B
 375009168 | *Geobacillus thermeleovorans* CCB US3 UFS
 407477159 | *Exiguobacterium antarcticum* B7
 172057416 | *Exiguobacterium sibiricum* 255-15
 169823684 | *Finegoldia magna* ATCC 29328
 160902522 | *Petrotoga mobilis* S795
 160902616 | *Petrotoga mobilis* S795

 333978211 | *Desulfotomaculum kuznetsovii* DSM 6115
 260892456 | *Ammoxenix degensii* K4
 260893631 | *Ammoxenix degensii* K4
 302391857 | *Acetohalobium arabaticum* DSM 5501
 428778891 | *Dactyloccoccopsis salina* PCC 8305
 260892735 | *Ammoxenix degensii* K4
 260893993 | *Ammoxenix degensii* K4
 433655673 | *Thermaanaerobacterium*
thermosacccharolyticum M0795
 332982963 | *Mahella australiensis* 50-1 BON
 332983053 | *Mahella australiensis* 50-1 BON
 222530385 | *Caldicellulosiruptor bescii* DSM 6725
 298675436 | *Methanohalobium evestigatum* Z-7303
 298675173 | *Methanohalobium evestigatum* Z-7303
 292490518 | *Nitrosococcus halophilus* Nc 4
 428778759 | *Dactyloccoccopsis salina* PCC 8305
 300114477 | *Nitrosococcus watsonii* C-113
 292491781 | *Nitrosococcus halophilus* Nc 4
 292491869 | *Nitrosococcus halophilus* Nc 4
 292491893 | *Nitrosococcus halophilus* Nc 4
 292491036 | *Nitrosococcus halophilus* Nc 4
 292490884 | *Nitrosococcus halophilus* Nc 4
 292491492 | *Nitrosococcus halophilus* Nc 4
 297563003 | *Nocardiosis dasonvilliei* DSM 41111
 374889417 | *Streptomyces bingchengensis* BCW-1
 374889654 | *Streptomyces bingchengensis* BCW-1
 271963428 | *Stereosporangium roseum* DSM 43021
 271964894 | *Stereosporangium roseum* DSM 43021
 258511478 | *Alicyclobacillus acidocaldarius* DSM 446
 160902264 | *Petrotoga mobilis* S795
 121604978 | *Palromonas naphthalenivorans* CJ2
 347754630 | *Chlorobacterium thermophilum* B
 218667603 | *Acidithiobacillus ferrooxidans* ATCC 23270
 218667415 | *Acidithiobacillus ferrooxidans* ATCC 23270
 344201059 | *Acidithiobacillus ferrivorans* SS3
 344201127 | *Acidithiobacillus ferrivorans* SS3
 288940385 | *Allochromatium vinosum* DSM 180
 288941921 | *Allochromatium vinosum* DSM 180
 288940786 | *Allochromatium vinosum* DSM 180
 288941171 | *Allochromatium vinosum* DSM 180
 288941306 | *Allochromatium vinosum* DSM 180
 121582913 | *Palromonas naphthalenivorans* CJ2
 121604604 | *Palromonas naphthalenivorans* CJ2
 91791204 | *Palromonas* sp. JS666
 91791283 | *Palromonas* sp. JS666
 121582916 | *Palromonas naphthalenivorans* CJ2
 121582983 | *Palromonas naphthalenivorans* CJ2
 428304158 | *Crinalium epiphamnum* PCC 9333
 428204921 | *Chroococcidiopsis thermalis* PCC 7203
 307513539 | *Cyanothece* sp. PCC 7822
 428216193 | *Oscillatoria acuminata* PCC 6304
 414079584 | *Anabaena* sp. 90
 172055147 | *Cyanothece* sp. ATCC 51142
 428316510 | *Oscillatoria nigro-viridis* PCC 7112
 428314951 | *Oscillatoria nigro-viridis* PCC 7112
 307591386 | *Cyanothece* sp. PCC 7822
 75812341 | *Anabaena variabilis* ATCC 29413
 17233293 | *Nostoc* sp. PCC 7120
 75812782 | *Anabaena variabilis* ATCC 29413
 434407609 | *Cylindrospermum stagnale* PCC 7417
 414078405 | *Anabaena* sp. 90
 440685156 | *Anabaena cylindrica* PCC 7122
 428315656 | *Oscillatoria nigro-viridis* PCC 7112
 428204926 | *Chroococcidiopsis thermalis* PCC 7203
 218440244 | *Cyanothece* sp. PCC 7424
 428775092 | *Halotheca* sp. PCC 7418
 414075475 | *Anabaena* sp. 90
 428305120 | *Crinalium epiphamnum* PCC 9333
 427713881 | *Synechococcus* sp. PCC 6312

S. pyogenes Cas9
 S. pyogenes Cas9 domains
 A. naeslundii Cas9
 A. naeslundii Cas9 domains

31216587	Caldicellulosiruptor hydrothermalis	108		NW	FYA-ELSLR1-G--KVHVKIGRE
34408285	Staineria cyanophanaera	PCC 7437		DW	FYS-ELAKL1-A--EVHTLQGME
414076241	Anabaena	sp. 90		RY	QIS-RLSKL1-T--NVLKLGKWE
428297778	Calchtrix	sp. PCC 6303		RF	QIN-RLSEI-A--SILKKKGWE
17233043	Nostoc	sp. PCC 7120		KW	MLK-QLEQP-A--PVAKVEGYQ
166363864	Microcystis aeruginosa	NIES-843		KW	AIE-QLSQL-A--TVHTRFGWQ
166367659	Microcystis aeruginosa	NIES-843		KW	AIE-QLSQL-A--TVHTRFGWQ
307150343	Cyanothece	sp. PCC 7822		KW	MLE-QLRQL-A--NVHTRGYWQ
307153352	Cyanothece	sp. PCC 7822		KW	MLE-QLRQL-G--NVDVTRYGWQ
428306661	Crimatium epipsammum	PCC 9333		KW	AIE-QLSKP-A--RVHTRFGWQ
3107150912	Cyanothece	sp. PCC 7822		KW	MLE-QLRQL-G--NVDVTRYGWQ
428775289	Halothecia	sp. PCC 7418		KV	MTEQWLNP1-A--PVTAKYGE
428779516	Dactylococcopsis	salina PCC 8305		KV	MTEQWLPO1-A--PVTAKYGE
428778599	Dactylococcopsis	salina PCC 8305		KV	ITEQWLPS1-A--PVTAKHGWG
428781696	Dactylococcopsis	salina PCC 8305		KV	MTEQWLPS1-T--PVTAKFGWQ
428215045	Oscillatoria acuminata	PCC 6304		KV	MLG-WLSKL1-A--PIETOLGYQ
428313425	Microcoleus	sp. PCC 7113		KW	MLE-HLSKL1-A--PTSTLYGWE
158340109	Acaryochloris marina	MBIC11017		YW	AMC-QLEAI-A--PVIKRHGWQ
428775002	Halothecia	sp. PCC 7418		QF	CIS-ELEKI-A--PVYVERGWQ
307150700	Cyanothece	sp. PCC 7822		KW	MLK-QLKQF-G--NVHTRLGWQ
312126663	Caldicellulosiruptor hydrothermalis	108			
312621424	Caldicellulosiruptor kronotskyensis	2002			-GRVYIP-VEGVRQWRA-LGTVNA--IIGRLREI-
222530402	Caldicellulosiruptor bescii	DSM 6725			-GRVYIP-VDGVQRQWRA-IIGRLREI-
222530431	Caldicellulosiruptor bescii	DSM 6725			-GRVYIP-VEGVRQWRA-LGTMNA--IIGQLREI-
307591386	Cyanothece	sp. PCC 7822			-GRVYIP-VEGVRQWRA-LGTMNA--IIGQLREI-
302391131	Acetobacterium arabaticum	DSM 5501			
435854579	Halobacteroides halobius	DSM 5150			NOP-S-LLEPKIL-KQSQSKLKN-VAAVNTRW--A--LFLNLKETG
435852864	Halobacteroides halobius	DSM 5150			KDQ-LITK1-DGKELNLPK-AMHVMQGKTS-W--LRELSEK
435854272	Halobacteroides halobius	DSM 5150			KDQ-EFLNII-DGKELNPK-AMHVMQGKT-W--LRELSKIV
313123202	Lactobacillus delbrueckii	ND02			KDQ-FLAID-SKELSDLKS-ASHVMQGKT-W--LRLDKLSKIA
77362205	Pseudoalteromonas haloplanktis	TAC125			KW-MKAK-CVKTQKLKG-ETPMFLIRK-R--IMTPPEAS
298675374	Methanohalobium	investigatum Z-7303			--Q-GKPELVRK-SKTKHATEI-G--ILKLAQKQC
288674707	Methanohalobium	investigatum Z-7303			NKR-P-SNTKH-PTEMGIVRS-Q--IRKLN--
288675090	Methanohalobium	investigatum Z-7303			NKR-P-SNTKH-PTEMGIVRS-Q--IRKLN--
169834784	Clostridium botulinum	A3 str. Loch Maree			GW-QPKIKSFKE-ATFMSMIRW-R--LVNTLGCKH
229587454	Clostridium botulinum	Ba4 str. 657			GW-HPKVKFSKE-ATFMSMIRW--R--LVNTLGCKH
297585104	Bacillus selenitireducens	MLS10			
188857532	Natraneraerobius	thermophilus Z-NW-WN-LP			-EGLEH--LF-QRKKKPFRD-ASQAMVRW-F--IYNGLKARY
312793946	Caldicellulosiruptor	kristjanssonii 177R1B			-ESLEN--YF-HPKDGFKFD-ESQFTLIRW-F--IYNGLKEKY
344996431	Caldicellulosiruptor	lactoaceticus 6A			-IKIPR--SK-LEKVK1FKD-ASHVST1GW-Y--IVNRLLKQY
217077332	Thermosiphon	africanus TCF52B			-IKIPR--SK-LEKVK1FKD-ASHVST1GW-Y--IVNRLLKQY
375009168	Geobacillus thermaeleovorans	CCB US3 UP5			-IKIPR--SK-LEKVRILKLD-ASHVST1GW-H--IVNRLLKQY
407477159	Exiguobacterium	antarcticum B7			-KGLEH--IF-QRKPPMRD-ASQMTAMRW-A--MFSPREKBY
172057416	Exiguobacterium	sibiricum 255-15			-EKLEH--IF-QRKSRFSRD-ASQMTVMRW-F--IYNAVKETY
169823684	Finegoldia	magna ATCC 29328			-ENLEH--LF-QRKSRFSRD-ASQMTVMRW-F--IYNAVKETY
160902522	Petrotoga	mobilis SJ95			-GKIK--LN-IKRGKSLKD-SAFMGIMRW-A--LYNRKLEIY
160902616	Petrotoga	mobilis SJ95			-GKIK--LD-FKRGQKFKD-ATSMNIMRW-T--LYNRKLDLY
160902616	Petrotoga	mobilis SJ95			-GKIK--LN-FKRGQKFKD-ATFMGIMRW-T--LYNRKLDLY
333978211	Desulfotomaculum	kuznetsovii DSM 6115			
260892456	Ammonifex	degensii KC4			-GEVS--LD-REGWTFAWV-AHAVLGKAY--LLALLSRPG
260893631	Ammonifex	degensii KC4			-GGAT--LD-RGKTFAWV-AHAVLGKAY--LLALLSRPG
302391857	Acetobacterium	arabaticum DSM 5501			-GEAA--LD-RGKTFAWV-AHAVLGKAY--LLALLSRPG
427788981	Dactylococcopsis	salina PCC 8305			DLK-EQDFRK--KLPKVYRNQVQLESVKVDISLDK-AGRVNSVRY-K--LDDQLKSCIS
260892735	Ammonifex	degensii KC4			ELK-D--VI-AEAPKPLDS-GSANVTRY-A--IVBMAKNLC
260893993	Ammonifex	degensii KC4			ELK-DSGRAVRDKRAENPK-AACMNATRY-V--LVERLRLALG
433656573	Thermomoanerobacterium				ELK-NSGRAVRDKRAENPK-AACMNATRY-V--LVERLRLALG
3730298263	Thiomicrosphaera	charolyticum M7095			
332983053	Maheila	australiensis 50-1 BON			-GYPD--IQ-QAKQPLKD-AAMINATRW-K--TYNLLKEMY
222530385	Caldicellulosiruptor	bescii DSM 6725			-GYPE--IQ-QAKQPLKD-AAMINATRW-A--LYNNLKKFG
298674466	Methanohalobium	investigatum Z-7303			-GYPE--IQ-QAKQPLKD-AAMINATRW-A--LYNNLKKFG
298675173	Methanohalobium	investigatum Z-7303			-GYPE--IQ-KLVKAPLD-KALVNTRW-R--IYEVLKNTG
292490518	Nitrosococcus	halophilus Nc 4			-GHPB--VY-EKAKRSLKD-AVINATKRD-K--VLDVLKNTG
428778759	Dactylococcopsis	salina PCC 8305			-GHPB--VY-EKAKRSLKD-AVINATKRD-K--VLDVLKNTG
300114477	Nitrosococcus	watsonii C-113			-KRK-N-KLEQQRSLMRAIL-KGHRTPLKD-AAVANTRY-A--VGRV1KOLI
292491781	Nitrosococcus	halophilus Nc 4			-KPK-T-ILES-VL-QKAKPLKD-ASAANVTRY-K--IVEV1SOHS
292491869	Nitrosococcus	halophilus Nc 4			-GYPK--IQ-AKAKPLLD-A-AVNTRY-A--IGRA1QSLIS
292491893	Nitrosococcus	halophilus Nc 4			-GYPK--IQ-SKSKPLKD-AAVANTRY-A--IGCT1QSVG
292491036	Nitrosococcus	halophilus Nc 4			-GYPK--IQ-TKANQPLKD-AAVANTRY-A--IGRA1QSVG
292490942	Nitrosococcus	halophilus Nc 4			-GYSQ--IQ-SKANQPLKD-AAVANTRY-A--VGR1QSVG
292491492	Nitrosococcus	halophilus Nc 4			-GYSQ--IQ-SKANQPLKD-AAVANTRY-A--VGR1QSVG
297563030	Nocardiosis	dassonvillei DSM 43111			-NP-R-ALAG--IS-AQTAKPLRD-AAVADTRW-A--LWRSLDCB
374989417	Streptomyces	bingchengensis BCW-1			-DK-S-KLAK--AL-AQAKPLRD-AAVQSTRW-A--LWRALDQM
374989654	Streptomyces	bingchengensis BCW-1			-DK-T-ML-SRAKPLRD-AAVQSTRW-A--LWRALDQM
271963428	Streptosporangium	roseum DSM 43021			-AKP-T-LLAK--IL-QKAKPLRD-AAVANTRW-A--LWRALAEATO
271964894	Streptosporangium	roseum DSM 43021			-DRP-V-VLVK--IL-QOSKPLRD-AAVANTRW-A--LWRALATGAT
258511478	Alicyclobacillus	acidocaldarius DSM 446			-HDO-E-RLHR--IE-SQRKAPLKD-TAAVNSTRW-A--LWRQLVATG
160902264	Petrotoga	mobilis SJ95			-NNP-E-RLKQ--IS-SESKRPLKD-TAAVNSRRW-A--LWRQLLEBTG
121604978	Polaromonas	naphthalenivorans CJ2			-KDP-K-RLAK--RL--V-IPNQLRGNS-AAVNTTRW-T--LFGALQKTDG
347754630	Chloracidobacterium	thermophilum B			-KDP-K-RLAR--IE-AQAKPLRD-AAVNTTRW-E--LNRLQRTA
218667603	Acidiibacter	ferrooxidans ATCC 23270			-KDP-E-RIRK--IL-AQAKPLRD-AAVNTTRW-E--LNRLQRTA
218667415	Acidiibacter	ferrooxidans ATCC 23270			-KDP-HDPR--IL-AQAKPLRD-AAVNTTRW-A--LWRQLVATG
344201059	Acidiibacter	ferrivorans SS3			-HDO-K-RLTR--IE-SQRKAPLKD-TAAVNSRRW-A--LWRQLLEBTG
344201127	Acidiibacter	ferrivorans SS3			-HDT-K-RLAH--IE-AVAKPLRD-TAAVNSTRW-A--LWRQLVDTG
288940385	Allochromatium	vinosum DSM 180			-RQP-A-LLRQ--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
288941921	Allochromatium	vinosum DSM 180			-RQP-V-RLHQ--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
288940786	Allochromatium	vinosum DSM 180			-RQP-A-LLH--IQ-AQAKPLQD-AAVANTRW-A--LFLAFAKQ
288941171	Allochromatium	vinosum DSM 180			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
288941306	Allochromatium	vinosum DSM 180			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
121582913	Polaromonas	naphthalenivorans CJ2			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
121604604	Polaromonas	naphthalenivorans CJ2			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
91791204	Polaromonas	sp. JS666			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
91791283	Polaromonas	sp. JS666			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
121582916	Polaromonas	naphthalenivorans CJ2			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
121582983	Polaromonas	naphthalenivorans CJ2			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
282304158	Crimatium	epipsammum PCC 9333			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
428204921	Chrococcidiopsis	thermalis PCC 7203			-RQP-A-LLH--IQ-AQAAQPLQD-AAVANTRW-A--LFLAFAKQ
307153539	Cyanothece	sp. PCC 7822			-AKS-D-LAKK--IL-QSKPLKD-TAAVNSTRW-A--LNQLQKPF
282126193	Oscillatoria	acuminata PCC 6304			-KRP-E-IL-QSKPLKD-TAAVNSTRW-A--LNQLQKPF
414079584	Anabaena	sp. 90			-KRP-E-IL-SQAKPLKD-ASAANVSTRW-A--LNRLKETG
17205147	Cyanothece	sp. ATCC 51142			-KRP-E-IL-SQAKPLKD-ASAANVSTRW-A--LNRLKETG
428316510	Oscillatoria	nigro-viridis PCC 7112			-KRP-E-IL-SQAKPLKD-ASAANVSTRW-A--LNRLKETG
282314951	Oscillatoria	nigro-viridis PCC 7112			-KRP-E-IL-SQAKPLKD-ASAANVSTRW-A--LNRLKETG
307591386	Cyanothece	sp. PCC 7822			-QOP-E-LFDR--IA-KQAKPLKD-AAVANSTRW-Q--LFNRLKETBS
578123411	Anabaena	variabilis ATCC 29413			-NQP-S-LLEK--IL-QSKPLKD-AAVANSTRW-A--LNRLKETG
17233293	Nostoc	sp. PCC 7120			-KDK-P-RLEK--IL-ALTKKPLD-AAVANSTRW-A--LLEVLKSTG
57812782	Anabaena	variabilis ATCC 29413			-KDK-S-RLQK--IL-QAKPLD-AAVANSTRW-A--LLEVLKSTG
434407609	Cylindrospermum	stagnale PCC 7417			-KDK-T-RLEK--IL-QVQKPLD-AAVANSTRW-A--LLEVLKSTG
414078405	Anabaena	sp. 90			-KDK-T-RLQK--IL-QAKPLD-AAVANSTRW-A--LLEVLKSTG
440685156	Anabaena	cylindrica PCC 7122			-KDK-S-RLEK--IL-QAKPLD-AAVANSTRW-A--LLEVLKSTG
282315656	Oscillatoria	nigro-viridis PCC 7112			-KDK-D-VLKK--IL-SQAKPLKD-AAVANSTRW-A--LFLNLGMTC
28204926	Chrococcidiopsis	thermalis PCC 7203			-NKP-D-VLKK--IL-SQAKPLKD-AAVANSTRW-A--LFLNLGMTC
218440244	Cyanothece	sp. PCC 7424			-SKS-D-LINH--VL-QKAKPLD-AAVANSTRW-Q--LFNRLKETG
282775092	Halothecia	sp. PCC 7418			-SKP-D-VLNR--IL-QSKPLD-AAVANSTRW-Q--LFNRLKETG
414075475	Anabaena	sp. 90			-KRP-D-LLKC--IL-QAKPLD-AAVANSTRW-A--LFLNLQRTG
282035120	Crimatium	epipsammum PCC 9333			-KRP-D-VLKR--IL-QAKPLD-AAVANSTRW-A--LMNRLKOTG
427713881	Synechococcus	sp. PCC 6312			-KRP-E-LLQR--IL-RKAKPLD-AAVANSTRW-A--LMNRLKOTG

S. pyogenes Cas9
S. pyogenes Cas8

S. pyogenes Cas9 domains
A. naeslundii Cas9

A. *naeslundii* Cas9

312126587 | *Caldicellulosiruptor hydrothermalis* 108
 434408285 | *Stanieria cyanosphaera* PCC 7437
 414076241 | *Anabaena* sp. 90
 428299778 | *Calothrix* sp. PCC 6303
 172330433 | *Nostoc* sp. PCC 7120
 166363864 | *Microcystis aeruginosa* NIES-843
 166367659 | *Microcystis aeruginosa* NIES-843
 307150343 | *Cyanothec* sp. PCC 7822
 307153352 | *Cyanothec* sp. PCC 7822
 428306661 | *Crinalium epiphamnum* PCC 9333
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 428779516 | *Dactyloccopsis salina* PCC 8305
 428778599 | *Dactyloccopsis salina* PCC 8305
 428781696 | *Dactyloccopsis salina* PCC 8305
 428215045 | *Oscillatoria acuminata* PCC 6304
 428313425 | *Microcoleus* sp. PCC 7113
 158340109 | *Acarychloris marina* MBIC11017
 428775002 | *Halothec* sp. PCC 7418
 307150700 | *Cyanothec* sp. PCC 7822

 312126663 | *Caldicellulosiruptor hydrothermalis* 108
 312621424 | *Caldicellulosiruptor kronotskyensis* 2002
 222530402 | *Caldicellulosiruptor bescii* DSM 6725
 222530431 | *Caldicellulosiruptor bescii* DSM 6725

 307951386 | *Cyanothec* sp. PCC 7822
 302391131 | *Acetohalobium arabaticum* DSM 5501
 435654579 | *Halobacteroides halobius* DSM 5150
 435652864 | *Halobacteroides halobius* DSM 5150
 435654272 | *Halobacteroides halobius* DSM 5150
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 773620250 | *Pseudoaeromonas haloplanktis* TAC125
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 298674707 | *Methanohalobium evestigatum* Z-7303
 298675090 | *Methanohalobium evestigatum* Z-7303
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 297585104 | *Bacillus selenitireducens* MLS10
 188507532 | *Natronaerobium thermophilus* JW-NM-NF-LF
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 160902616 | *Petrotoga mobilis* S795

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 260894245 | *Ammonifex degensii* K4
 260893631 | *Ammonifex degensii* K4
 302391857 | *Acetohalobium arabaticum* DSM 5501
 428778891 | *Dactyloccopsis salina* PCC 8305
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thermosaccharolyticum M7095
 332982963 | *Mahella austrialiensis* 50-1 BON
 332983053 | *Mahella austrialiensis* 50-1 BON
 222530385 | *Caldicellulosiruptor bescii* DSM 6725
 298675436 | *Methanohalobium evestigatum* Z-7303
 298675173 | *Methanohalobium evestigatum* Z-7303
 292490518 | *Nitrosococcus halophilus* Nc 4
 428778759 | *Dactyloccopsis salina* PCC 8305
 300114477 | *Nitrosococcus watsonii* C-113
 292491781 | *Nitrosococcus halophilus* Nc 4
 292491869 | *Nitrosococcus halophilus* Nc 4
 292491893 | *Nitrosococcus halophilus* Nc 4
 292491036 | *Nitrosococcus halophilus* Nc 4
 292490884 | *Nitrosococcus halophilus* Nc 4
 292491492 | *Nitrosococcus halophilus* Nc 4
 297563003 | *Nocardiopsis dassonvillei* DSM 41111
 374889417 | *Streptomyces bingchenggensis* BCW-1
 374896654 | *Streptomyces bingchenggensis* BCW-1
 271963428 | *Streptosporangium roseum* DSM 43021
 271964894 | *Streptosporangium roseum* DSM 43021
 258511478 | *Alicyclobacillus acidocaldarius* DSM 446
 160902264 | *Petrotoga mobilis* S795
 121604978 | *Polaromonas naphthalenivorans* CJ2
 347754630 | *Chlorobacterium thermophilum* B
 218667603 | *Acidithiobacillus ferrooxidans* ATCC 23270
 218667415 | *Acidithiobacillus ferrooxidans* ATCC 23270
 344201059 | *Acidithiobacillus ferrivorans* SS3
 344201127 | *Acidithiobacillus ferrivorans* SS3
 288940385 | *Allochromatium vinosum* DSM 180
 288941921 | *Allochromatium vinosum* DSM 180
 288940786 | *Allochromatium vinosum* DSM 180
 288941171 | *Allochromatium vinosum* DSM 180
 288941306 | *Allochromatium vinosum* DSM 180
 121582913 | *Polaromonas naphthalenivorans* CJ2
 121604604 | *Polaromonas naphthalenivorans* CJ2
 91791204 | *Polaromonas* sp. JS666
 91791283 | *Polaromonas* sp. JS666
 121582916 | *Polaromonas naphthalenivorans* CJ2
 121582983 | *Polaromonas naphthalenivorans* CJ2
 428304158 | *Crinalium epiphamnum* PCC 9333
 428204921 | *Chroococcidiopsis thermalis* PCC 7203
 307153539 | *Cyanothec* sp. PCC 7822
 428316193 | *Oscillatoria acuminata* PCC 6304
 414079584 | *Anabaena* sp. 90
 172055147 | *Cyanothec* sp. ATCC 51142
 428316510 | *Oscillatoria nigro-viridis* PCC 7112
 428314951 | *Oscillatoria nigro-viridis* PCC 7112
 307513866 | *Cyanothec* sp. PCC 7822
 75812341 | *Anabaena variabilis* ATCC 29413
 17233293 | *Nostoc* sp. PCC 7120
 75812782 | *Anabaena variabilis* ATCC 29413
 434407609 | *Cylindrospermum stagnale* PCC 7417
 414078405 | *Anabaena* sp. 90
 440685156 | *Anabaena cylindrica* PCC 7122
 428315656 | *Oscillatoria nigro-viridis* PCC 7112
 428204926 | *Chroococcidiopsis thermalis* PCC 7203
 218440244 | *Cyanothec* sp. PCC 7424
 428775092 | *Halothec* sp. PCC 7418
 414075475 | *Anabaena* sp. 90
 428305120 | *Crinalium epiphamnum* PCC 9333
 427713881 | *Synechococcus* sp. PCC 6312

TKQLRE-----SYGLKTKS---KKLETFE-----SHCVDW-CLAN-----WTCGGHVQP-----DNTELLVIIPLRF
 TKQIRD----ELGLKKT-T-KNKAETPFE-----AHCVDSF-ALAY-----SVVGGNNSLP-----KNTNLIVPPRLF
 TSNLRK----YLGLKHEK---SDKSLSQIPE-----THAVDAV-ALASSEFPVKYQTWEAGK-----NHGASWIGGVBDITEQS-----TIVRPP
 TSNLRK----HLGMHKKEK---TDKSLQIPE-----THAVDAI-TLASESFVKYKSFEKGK-----TRGASWGNVSITESQF-----TIVRPP
 TASTRK----YLGLTKNS---SDKSLSQIPE-----THAVDGV-AIAATAFVEVYQHTVK-----VNGGDWNFGNVVITFALP-----KLIRPP
 TSNLRK----YLGLEKS---KNKAQSP-----SHANDGI-ALACCPFLYWPMPHNYN-----GHGYDWNKGSKVVTNAPF-----AVIKRPP
 TSNLRQ----HLGLLEKS---KDKAQSKS-----SHANDGI-ALACCPFLYWPMPHNYN-----GHGYDWNKGSKVVTNAPF-----AVIKRPP
 TSNLRK----HLGLLEKS---KDKAQSKS-----SHANDGI-ALACCPFLYWPMPHNYN-----GHGYDWNKGSKVVTNAPF-----AVIKRPP
 TSNLRK----HLGLLEKS---KDKAQSKS-----SHANDGI-ALACCPFLYWPMPHNYN-----GHGYDWNKGSKVVTNAPF-----AVIKRPP
 TSKREB----HLGLEKS---KDKAQSKT-----SHANDGI-ALACCPFLYWPMPHNYN-----GHGYDWNKGSKVVTNAPF-----AVIKRPP
 TSAREE----MLGLEKD---REKSNSQTPA-----THAVDGV-ALASYQFLKVKYCYKGW-----FEGSVETINAPP-----STIKRPP
 TAKRDR----TLGLQED---RNRSQTPA-----THAVDGV-ALASYQFLKVKYKLKGW-----FEGSVETINAPP-----SVIKRPP
 TARRD-----SLGLERD---REKSNSQNP-----THAVDGV-ALASHDFLKVKYKLKGW-----FEGSVETINAPP-----SVIKRPP
 TASLRT----QLGLQED---RKDSKSKT-----SHAVDG-VLACSEF1IYKPKPQTCA-----THGHTWGSVITITNCF-----VVVR
 TSKREB----HLGLEKS---KDKAQSKT-----SHAVDG-VLACSEF1IYKPKPQTCA-----THGHTWGSVITITNCF-----VVVR
 TSAREE----MLGLEKD---REKSNSQTPA-----THAVDGV-ALASLYQFLKVKYKLKGW-----FEGSVETINAPP-----STIKRPP
 TAKRDR----TLGLQED---RNRSQTPA-----THAVDGV-ALASLYQFLKVKYKLKGW-----FEGSVETINAPP-----SVIKRPP
 TARRD-----SLGLERD---REKSNSQNP-----THAVDGV-ALASHDFLKVKYKLKG

B

No	Hit		Prob	E-value	P-value	Score	SS	Cols	Query	HMM	Template	HMM
1	PF14239	RRXRR: RRXRR protein	100.0	1.6E-79	2.7E-84	564.5	20.9	174	5-178	1-176	(176)	
2	4oge_A	HNH endonuclease domain	99.7	6.1E-19	1.1E-23	202.5	4.6	189	128-321	476-748	(1101)	
3	4cmp_A	CAS9, crisper-associated	99.7	6.7E-18	1.2E-22	197.5	6.4	196	125-321	732-999	(1372)	
4	COG3513	Predicted CRISPR-assoc	99.5	3.6E-14	6.2E-19	154.1	6.0	138	184-321	561-748	(1088)	
5	2qgp_A	HNH endonuclease; Q39X4	99.4	1.8E-13	3.2E-18	119.1	4.5	60	186-245	27-88	(112)	
6	PF13395	HNH_4: HNH endonuclea	99.2	4.6E-12	7.9E-17	96.8	2.4	46	197-242	1-53	(54)	
7	PF01844	HNH endonuclease	99.1	2.7E-11	4.7E-16	88.5	1.9	44	197-240	1-47	(47)	
8	COG1403	McrA Restriction endon	98.8	5.4E-09	9.4E-14	91.5	3.3	61	184-245	59-120	(146)	
9	PF14279	HNH_5: HNH endonuclea	97.1	0.00038	6.7E-09	56.6	2.0	39	197-238	1-41	(71)	
10	PF13391	HNH_2: HNH endonuclea	97.0	0.0006	1E-08	53.1	2.5	44	197-240	1-61	(66)	

13 le71_A GP49, recombination end 91.2 0.29 5E-06 46.0 3.7 53 184-238 10-66 (157)

16	lhj _A	Holliday junction resol	87.7	1.5	2.7E-05	41.0	5.7	41	56-96	2-46	(158)
17	PF02075	RuvC: Crossover junct	87.4	1.4	2.4E-05	40.7	5.3	40	57-96	1-45	(149)
18	4ep4_A	Crossover junction endo	87.0	1.3	2.3E-05	41.8	5.0	41	56-96	2-50	(166)

22 **4oge_A** HNH endonuclease domain 80.5 1.3 2.3E-05 52.9 2.6 74 56-129 12-99 (1101)

37 **4cmp_A** CAS9, crispr-associated 58.7 5.3 9.2E-05 49.2 1.0 53 54-110 7-79 (1372)

44 COG0675 Transposase and inacti 50.6 75 0.0013 31.6 7.5 152 40-233 161-328 (364)

37 6665915 Predicted CRISPR spacer 36.7 36 6.66632 16.7 2.1 65 35 117 1 38 (168)

>**PF14239** RRXRR: RRXRR protein
Probab=100.00 E-value=1.6e-79 Score=564.55 Aligned_cols=174 Identities=61% Similarity=1.015 Sum_probs=0.0

T PF14239_consens 1 VVFLDKNGRPMPCEPARAKRDLRQGRAKVVRRYFTIQLQRYEPGSGYTQFIRIGIDPGSKITGIAVSEKKVLUWAEELE 80 (176)
T ss_pred CeBECCCCCCCCCCCCHHHHHHHHCCCCEEEecCCCCeCCCCccccCC

```

T P14259_consensus 61 QPDIQKRLRQKAIAKGAKIATKAKRPLDNGKFRGWIFPSEKIRVDTIHLRNVAKLCKRFSTIAIVVEELRDFI 100 (170)
T ss_pred          ccHHHHHHHHHHHHHHhhccccccccccccccccCCCCCccCccHHHHHHHHHHHHHHHHHhCcccccEEEEEeh

```

```

Q Thu_Feb_20_16: 163 QKLQNPEISGVEYQQG 178 (425)
Q Consensus      163 qk~np~i~G~~YQgg 178 (425)
                  [|+|+|||||+|+|||||]
T Consensus       161 q~l~np~i~G~~YQGG 176 (176)
T PF14239_consen 161 QKLQNPEISGVEYQQG 176 (176)
T ss_pred         hhccCcCcccccccCCCC

```

No 2
>4oge_A HNH endonuclease domain protein; crisper-CAS, CAS9, RUV, RNA-guided DNA endonuclease, cytoplasmic, hydrolyase; HET: SPP; 2.20A (Actinomycetes naeslundii) PDB: 4oge_A

Q ss_pred	hhCCCc-----cccccceeeccc			
Q Thu_Feb_20_16:	301 TRLDLPK-----AHWLDAAVCQGVDA	321	(425)	
Q Consensus	301 ~~~gL-K-----H- DA- cig~~~	321	(425)	
	.. +.+ ++ +.++			
T Consensus	711 ~-wgl-----K-R-d-rHHAIDA-VIA---	748	(1101)	
T 4oge_A	711 KAAGIDSRLNLIGEKGRKDRIDRHHAVDASVVALMEA	748	(1101)	
T ss_dssp	HHHTCTTTSCSSCCCCSSCCBTTHHHHHHHHHHHHCCH			
T ss_pred	HHCCchhhhccccccCCCCCchHHHHHHHHHCCH			

No 3

```
>4cmp_A CAS9, crispr-associated endonuclease CAS9/CSN1; hydrolase, DNAse, RNA-guided, immunity, crRNA, genome editin; 2.62A [Streptococcus pyogenes] PDB: 4cmp_A
  Probab=99.71 E-value=6.7e-18 Score=197.53 Aligned_cols=196 Identities=20% Similarity=0.121 Sum_probs=0.0
```

Q ss_pred		HhccCChhh-----HHHHHHHHH-----HHHHHHHHhC-----CCeEeechhhhTFN	299 (425)
Q Thu_Feb_20_16:	258	RKA[KAP]LKD-----AAAVNSTRW-----ALFNLNATAKG-----LPLTTGDKTFKN	299 (425)
Q Consensus	258	--~k--lkd-----a--vn~TR-----~l--Lk-----V--G--Tk--	299 (425)
		.+.+.. .+.+ +.+.. .+. +. +. +. .	
T Consensus	891	~-~is--K--L~-----e~-----Fi-RqL-DTrYisK-----L-----Vv-vKG-Ts-1	970 (1372)
T 4cmp_A	891	HANAKDITQRKFDNLTKAERGGLSLEDKAGFICIKRQLVTRQIITKHVAQILDSMRNTKYDENDKLIREVKVITLHSSRMMHCF	970 (1372)
T ss_dssp		HHTTSSSSCHHHHHHHHH-----CCHHHHHHHHHHHHHHHCCCCHHHHHHHHHHCCCCBCTTSCKBCCCCCEEKKHHSSHHHH	
T ss_pred		HhhhhcCHHHHHHHHHhhccccchhhcchhhhccccCHHHHHHHHHHHHHHHHHHHHHcccccccCCCCCEEEEBCChhhhHHH	

Q ss_pred		hhnCCCc---cccccccccccccc	
Q Thu_Feb_20_16:	300	FTRIDLPK-----AHWLDAAVGVDAA	321 (425)
Q Consensus	300	R~~~gl-K~~~~~H-DA:cig~~~ .. + +++.+	321 (425)
T Consensus	971	R-wgl-K-r~~~n-HHA-DA-lia~~~	999 (1372)
T 4cmp_A	971	RKDFOQFYKVRBИННYYHAAHDAYLNAAVGGT	999 (1372)
T ss_dssp		HHHHTCCCCCTTCSHHHHHHHHHHHHHHHHHH	
T ss_pred		HHHcCCCCcCCCCchhhhhhhhhhhchch	

No 4
SOC351

>COG3513 Predicted CRISPR-associated nuclease, contains McrA/HNH-nuclease and RuvC-like nuclease domain [Defense mechanisms]
Probab=99.48 E-value=3.6e-14 Score=154.15 Aligned cols=138 Identities=28% Similarity=0.304 Sum probs=0.0

PF085ab-339.48 E-value=3.0e-14 Score=154.15 Aligned_CDSs=138 Identities=28% Similarity=0.504 Sum_pICoS=0.0

Q ss_pred	hhCCCC-----ccccccccc			
Q Thu_Feb_20_16:	301 TRLDPK-----AHWLDAACVGQVDA	321	(425)	
Q Consensus	301 ~~gL-K-----H~DA~cig~~~	321	(425)	
	... - + ++ +...			
T Consensus	721 ~~wgl~kvr~N~~HHA~DAVVA~a~~	748	(1088)	
T COG3513	721 KTWGLYKVRENNNYHIAHDAAVVVACALN	748	(1088)	
T ss_pred	hhcceeeecccccchhhhhhhheccccH			

No 5

```
>2qgp_A HNH endonuclease; Q39X46, GMR87, X-RAY, NESG, structural genomics, PSI-2, protein structure initiative;
  2.60A [Geobacter metallireducens gs-15] PDB: 4h9d_A
  Probab=99.41 E-value=1.8e-13 Score=119.11 Aligned cols=60 Identities=28% Similarity=0.499 Sum probas=0.0
```

Probab=99.41 E-value=1.8e-13 Score=119.11 Aligned_cols=60 Identities=28% Similarity=0.499 Sum_probs=0.0

No 6

>PF13395 HNH_4: HNH endonuclease
Probab=99.23 E-value=4.6e-12 Score=96.76 Aligned_cols=46 Identities=50% Similarity=0.959 Sum_probs=0.0

No. 7

>PF01844 HNH: HNH endonuclease PDB: 2QGP_C.
Probab=99.12 E-value=2.7e-11 Score=88.50 Aligned_cols=44 Identities=41% Similarity=0.954 Sum_probs=0.0

No 8

>**COG1403** McrA Restriction endonuclease [Defense mechanisms]
 Probab=98.76 E-value=5.4e-09 Score=91.49 Aligned_cols=61 Identities=36% Similarity=0.641 Sum_probs=0.0

Q ss_pred	hHHHHHHHHhCCccccCCC-CcEEeEeeeeCCCCCc	hhhhhhhhhhhhhhccccHHHH
Q Thu_Feb_20_16:	184 EVREYLLKEWKGRKCTYCGTK-DVPLEVEHIIHPRSRGGTD	RVSNLTMACHACNQSKGNQIRD
Q Consensus	184 -vrey~l-k~~~C-yCg~~~~Lev-HI-prs-gGsd~sNl~1-C~Cn~K-n~~~~f	245 (425)
T Consensus	59 .+ ..++. ++ + +. .-++ + ++ . .+..+ .++ ++....+	245 (425)
T COG1403	59 KTRRAVLLRDNGLCQYCGSGT-DLEVHDIVPLSRGGASAWENLET	CERCHNNKGSSRLPGKA
T ss_pred	HHHHHHHHhcccccccCcCC-CcEEeEeeeeCcCc	HHHHHHHHHHHHccccccccccc

No 9

>**PF14279** HNH_5: HNH endonuclease
 Probab=97.05 E-value=0.00038 Score=56.63 Aligned_cols=39 Identities=38% Similarity=0.837 Sum_probs=0.0

Q ss_pred	ccccCCcCCee--EeeeeCCCCCc	hhhhhhhhhhhhhhhhcc
Q Thu_Feb_20_16:	197 CTYCGTKDVPLE--VEHIIHPRSRGGTD	RVSNLTMACHACNQSKGNQ
Q Consensus	197 C-yCg~~~~Lev-HI-prs-gGsd~sNl~1-C~Cn~K-	238 (425)
T Consensus	1 Ci~C-~s-EHIIP-sLGG~VC~CN~g	41 (71)
T PF14279_consens	1 CIYCNKEKSESNFSEEHIIPESLGGKLKINN--VCDKCNKKFG	41 (71)
T ss_pred	CccCCCCCccCCccccCc	hhcCCCCcccc--hhHHHHHH

No 10

>**PF13391** HNH_2: HNH endonuclease
 Probab=96.99 E-value=0.0006 Score=53.06 Aligned_cols=44 Identities=27% Similarity=0.269 Sum_probs=0.0

Q ss_pred	ccccCCc---CcEEeEeeeeCC--Cc-----	chhhhhhhhhhhhhhhcc
Q Thu_Feb_20_16:	197 CTYCGTKD---VPLEVEHIIHPRSRG-GTD-----	RVSNLTMACHACNQSKGNQ
Q Consensus	197 C-yCg~~~~Lev-HI-prs-gGsd-----sNl~1-C~Cn~K-n	240 (425)
T Consensus	1 C-itg~~~~~eaAH-P-s~~~~~Ngl~L~~~lH~~fd~~	61 (66)
T PF13391_consens	1 CVITGIRDPSWPWELAHHIVPFSLGSSWMNWFGEYANDWISPSNGLLRPDHLKLFDRG	61 (66)
T ss_pred	CccCccCccCCCCcEEEccccCc	hhhhhhhccCCCCcEEEccC

No 13

>**le71_A** GP49, recombination endonuclease VII; resolvase, holliday junction, DNase; HET: SO4; 1.32A {Bacteriophage T4} SCOP: **a.140.4.1 d.4.1.5** PDB: **2qnc_A*** **len7_A** **1etd_A** **2qnf_A***
 Probab=91.18 E-value=0.29 Score=45.98 Aligned_cols=53 Identities=19% Similarity=0.421 Sum_probs=0.0

Q ss_pred	hHHHHHHHHhCCccccCCC---CCeEEeEeeeeCCCCCc	hhhhhhhhhhhhhhhhcc
Q Thu_Feb_20_16:	184 EVREYLLKEWKGRKCTYCGTK---DVPLEVEHIIHPRSRGGTD	RVSNLTMACHACNQSKGNQ
Q Consensus	184 -vrey~l-k~~~C-yCg~~~~Lev-HI-prs-gGsd~sNl~1-C~Cn~K-	238 (425)
T Consensus	10 ~~~~l~~~Q-g-C-IC~~~l~~~vDHDh~~~g~tG-VRGLlC~CN~lG	66 (157)
T le71_A	10 EEKQKFYDAQNQKCLICQRELNPDVQANHLDHDHELN-GPKAGKVRLGCNLCDAE	G
T ss_dssp	HHHHHHHHHTTBCTTTCCBCSSGGSEEECCCS-STTTTBEEEEECHHHHHHH	
T ss_pred	HHHHHHHHhCcCcCCCCccccccccCCCCC-CCCCCchhhcHhchhc	

No 16

>**lhjr_A** Holliday junction resolvase (RUV)C; site-specific recombinase; 2.50A {Escherichia coli} SCOP: **c.55.3.6**
 Probab=87.67 E-value=1.5 Score=40.99 Aligned_cols=41 Identities=22% Similarity=0.353 Sum_probs=0.0

Q ss_pred	EEEEEEcCCCCcEeeeeEC---	EEEEEEEEEeCc
Q Thu_Feb_20_16:	56 IELKIDPGSKVGTIAIKQGN---KIIFGAELQHRGQ-QIKDALLS	96 (425)
Q Consensus	56 i-lgiDpG-k~G-av~~~~~l~~el~r~~vk~l~~	96 (425)
T Consensus	2 ~lgidPGL~tG-gvi~~g~~v~G-i~t~~~~~Rl~~	46 (158)
T lhjr_A	2 IILGIDPGSRVTGYGVIRQVGRQLSYLGS CIRTKVDDLPSRLKL	46 (158)
T ss_dssp	EEEEEECCSSEEEEEEETTEEEEEECCCCSC	HHHHHHHH
T ss_pred	EEEEEEccCCCCeEeeeeCcCc	EEEEECCCCHHHHHH

No 17

>**PF02075** RuvC: Crossover junction endodeoxyribonuclease RuvC; PDB: **1HJR_A**.
 Probab=87.43 E-value=1.4 Score=40.65 Aligned_cols=40 Identities=23% Similarity=0.387 Sum_probs=0.0

Q ss_pred	EEEEEcCCCCeEeeeeEC---	EEEEEEEEEeCc
Q Thu_Feb_20_16:	57 ELKIDPGSKVGTIAIKQGN---KIIFGAELQHRGQ-QIKDALLS	96 (425)
Q Consensus	57 i-lgiDpG-k~G-av~~~~~l~~el~r~~vk~l~~	96 (425)
T Consensus	1 ~lgidPGL~tG-avi~~~~~i~G~I~t~~~~~Rl~~	45 (149)
T PF02075_consens	1 IILGIDPGSLNTGYAVIeedggkLRLDYGTIKTSSKDSLPERLKE	45 (149)
T ss_dssp	EEEEEE-SSEEEEEEETTEEEEEE--S-	HHHHHHHH
T ss_pred	CCCCCCCCeEeeeeCcCc	EEEEECCCCHHHHHH

No 18

>**4ep4_A** Crossover junction endodeoxyribonuclease RUVC; resolvase, hydrolase; 1.28A {Thermus thermophilus} PDB: **4ep5_A**
 Probab=86.99 E-value=1.3 Score=41.83 Aligned_cols=41 Identities=17% Similarity=0.267 Sum_probs=0.0

Q ss_pred	EEEEEEcCCCCeEeeeeEC---	EEEEEEEEEeCc
Q Thu_Feb_20_16:	56 IELKIDPGSKVGTIAIKQGNK---IFGAELQHRGQ-QIKDALLS	96 (425)
Q Consensus	56 i-lgiDpG-k~G-av~~~~~l~~el~r~~vk~l~~	96 (425)
T Consensus	2 rILGIDPGl~tG-gvi~~g~~v~G-I~t~~~~~Rl~~	50 (166)
T 4ep4_A	2 VVAGIDPGITHLGLGVVAVEKGALKARLHGEVVKTSPQEPAKERVGR	50 (166)
T ss_dssp	EEEEEECCSSEEEEEECCSSSSCEEEEEECCCTTS	CHHHHHHH
T ss_pred	EEEEEEccccCcEeeeeCcCc	EEEEECCCCHHHHHH

No 22

```
>4oge_A HNH endonuclease domain protein; crisper-CAS, CAS9, RUVCR, RNA-guided DNA endonuclease, cytoplasmic, hydrolase; HET: SPD; 2.20A {Actinomycetes naeslundii} PDB: 4oge_A*  
Probab=80.53 E-value=1.3 Score=52.86 Aligned_cols=74 Identities=19% Similarity=0.209 Sum_probs=0.0
```

No 37

```
>4cmq_A CAS9, crispr-associated endonuclease CAS9/CSN1; hydrolase, DNAse, RNA-guided, immunity, crRNA, genome editin; 2.62A [Streptococcus pyogenes] PDB: 4cmq_A
Probab=58.70 E-value=5.3 Score=49.16 Aligned_cols=53 Identities=28% Similarity=0.396 Sum_probs=0.0
```

No 44

>COG0675 Transposases and inactivated derivatives [DNA replication, recombination, and repair]
Probab=0.60 E-value=75 Score=31.60 Aliqnded cols=152 Identities=18% Similarity=0.323 Sum probs=0.0

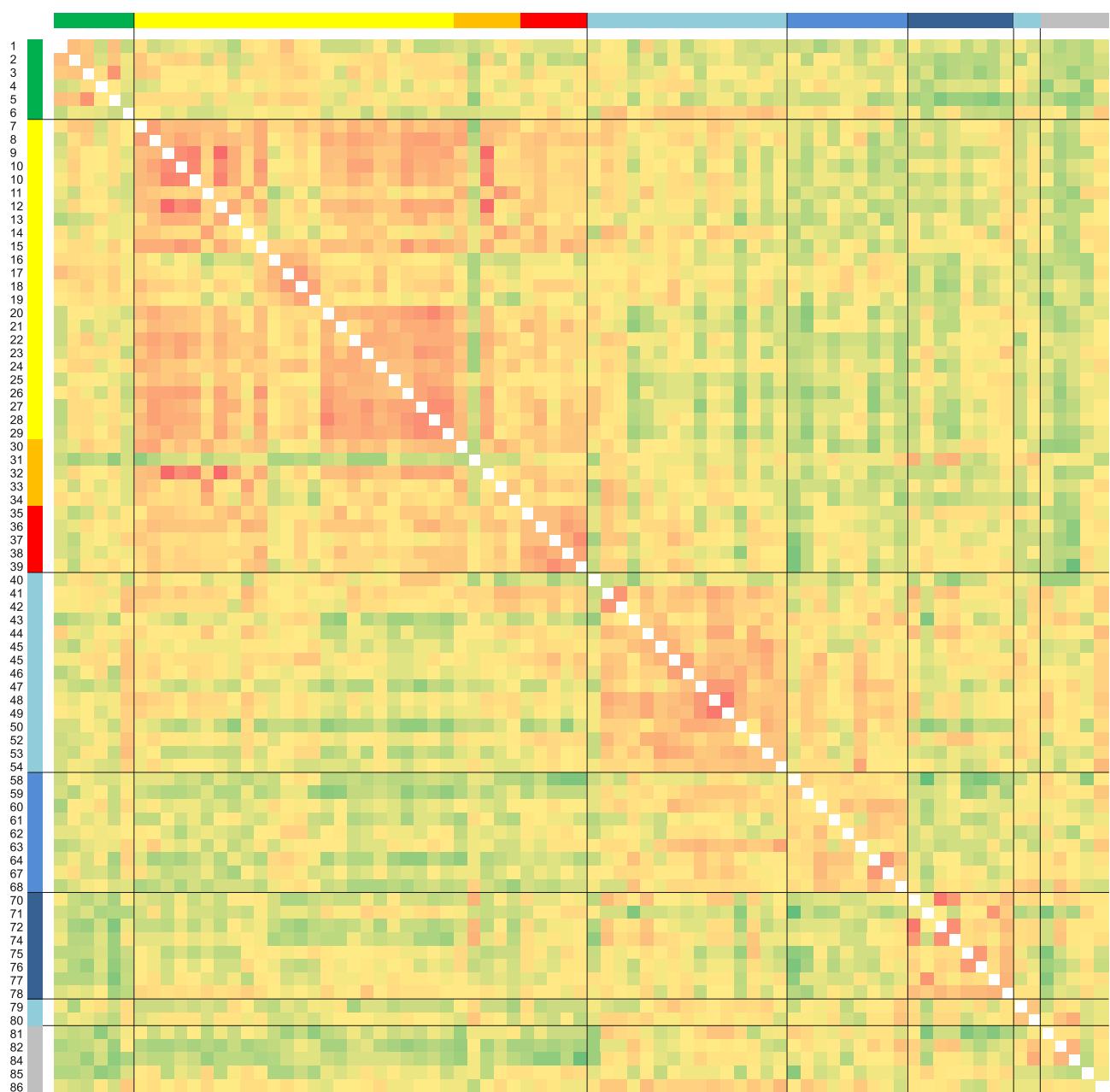
No 57

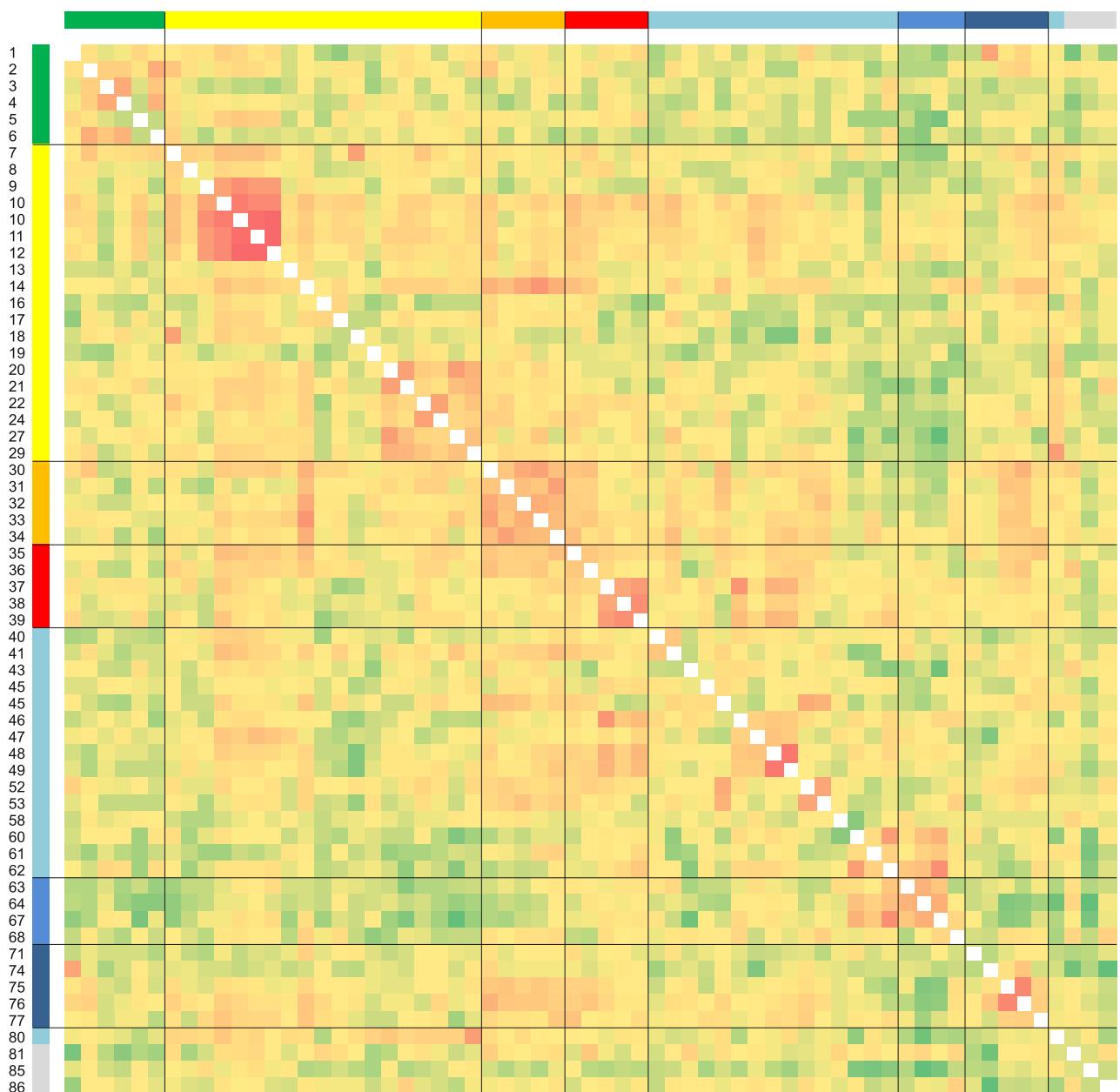
>COG3513 Predicted CRISPR-associated nuclease, contains McrA/HNH-nuclease and RuvC-like nuclease domain
Probab=36.67 E-value=30 Score=40.66 Aligned_cols=63 Identities=32% Similarity=0.385 Sum_probs=0.0

No 75

>PF01385 OrfB_IS605: Probable transposase;
Probab=29.04 E-value=39 Score=31.85 Aligned_cols=110 Identities=16% Similarity=0.156 Sum_probs=0.0

A



B

Supplementary Figure S9. CRISPR repeats and tracrRNA orthologs pairwise identity. **(A)** Sequence identity of each type II CRISPR repeat pair is represented with a color scale from green (lowest) to red (highest identity). The repeat sequences are ordered according to the corresponding Cas9 sequences on the Cas9 phylogenetic tree. The numbers on the left indicate the exact position on the tree. Color bars on the top and left correspond to the colors of Cas9 tree clusters. (see Figure 4 and Supplementary Table S1) Note the high similarity of the repeat sequences belonging to the same, large clusters of the Cas9 tree. **(B)** Sequence identity of every tracrRNA sequence pair is represented with a color scale from green (lowest) to red (highest identity). The tracrRNA sequences are ordered according to the corresponding Cas9 sequences on the Cas9 phylogenetic tree and the numbers on the left indicate the exact position on the tree. Color bars on the top and left correspond to the colors of the Cas9 tree clusters (see Figure 4 and Supplementary Table S1). Note that tracrRNA sequences show higher similarity only in small clusters of closely related loci.

Type II-B

F. novicida 3523

GUUUCAGUUACUAGAUUAUCUGAUUAACUA-UCUUAG
GUUUCAGUAGCUAGAUUAUUUGGUACAUACUAGUGUU
UUUCGUUAUUAGGUAUUUGAUUAACUGCUUCAA
AUUUUAGCGGUUAUUAUCUGAUUAACUUACCCAG
GUUUCGGUAGCUAGGUAUUUGAUUAACUGCUGUAG
UUAAAAGUAGCUAGAUUAUUUGAUUAACUGCUGUAG
GUUUCAGUAGCUAGAUUAUUUGAUUAACUGCUGUAG

F. novicida U112 and Fx1

UUUUUAGUAAACUAGACUAUUAUAAAACUA-UGUUAG
GUUUCAGUUGGUAGAU-UAUUUGGUAGUACUUGUGUU
UUUUUAAAUGGAAU-GCUUUGGUAGUAUGUUGAGAU
GUUUUCGUGAUUAUUAU-UAUUUGAUACAAUGUGUCAA
GUUUCAGUUGCUGAAU-UAUUUGGUAAACUACUGUUAG

Gammaproteobacterium HTCC5015

AUACAAGGGAGUAUUUGCUGGAUAAGCUCUGAAC
GUUUCAGCUGUUGGUUGGUUGGGAUAGCUCUGAAC

Distance from 1st repeat: 55 nt instead of ~33 nt spacer

Wolinella succinogenes DSM1740

UUUGGUAAAGCUGGAUGGGAUUAUUAUAGAGAGUGUUGC
GUUUCACAGGCUAAGCGGAUUGCUAUAAAGUGUUGC

Distance from 1st repeat: 37 nt, like other spacers (~33-37 nt)

Legionella pneumophila str. Paris

AUUUCAUUAGUAAAUUUUUAGAUGAGGGAUUAUUGG
GUUUCAGUGGUUGGAUUUUUAGAUGAGGGAUUAUUGG

Distance from 1st repeat: 38 nt, like other spacers (~34-36 nt)

Sutterella wadsworthensis 3_1_45B

GUUUCAGCGCUUUAGCUUGCAGCGUUAUGAUCUUAU
GUUUCAGUGCUAUAGCUCGUAGCGUUAUGAUCUUCGC

Distance from 1st repeat: 38 nt, like other spacers (~33-39 nt)

Type II-A

Streptococcus pyogenes M1 GAS

GUUUUAGAGCUAUGCUGUUUUGAAUGGUCCAAAAC
GUUUUAGAGCUAUGCUGUUUUGAAUGGUCCAUUC

Streptococcus thermophilus LMD-9

GUUUUUGUACUCUCAAGAUUUAGUAACUGUACAAAC
GUUUUUGUACUCUCAAGAUUUAGUAACUGUACAGU

Treponema denticola ATCC 35405

GUUUGAGAGUUGUGUAUUUAAGAUGGAUCUCAAAC
GUUUGAGAGUUGUGUAUUUAAGAUGACAACCAAGC

Type II-C

Neisseria meningitidis Z2491

GUUGUAGCGCCCAUUCUCAUUUCGCAGUGCUACAAU
GUUGUAGCCUCCUUUCUCAUUUCGCAGUGCUACAAU

Campylobacter jejuni subsp. *jejuni* NCTC 11168

AUUUUAGUCCUUUUAAAUUUCUUUAUGGUAAAAAU
GUUUUAGUCCUUUUAAAUUUCUUUAUGGUAAAAAU

Candidatus Puniceispirillum marinum IMCC1322

UGGGGGGUAGGCUCUCAAUCACCAGAGUGCUAUACU
GUUGCUCUAGGCUCUCAAUCACCAGAGUGCUAUACU

Supplementary Figure S10. Degenerated repeats of type II CRISPR-Cas. Black: consensus repeat sequence; red: degenerated repeat sequence; yellow highlighted: scaRNA repeat. Nucleotides mutated in the degenerated repeats are highlighted in grey. For every strain, the repeats are listed in the direction of 5' to 3' end of the CRISPR array. For the type II-B, the degenerated repeat distance from the first, regular repeat in the array is given. Note that unlike type II-B, type II-A and II-C repeats display a lower amount of mutations and their distribution are typically at one of the repeat ends.

A

<i>Streptococcus pyogenes</i> SF370	GUUUUAG--AGCUAUGCUGUUUUGAAUGGUCCCCAAAC • AAAUGAACGUAACGACAAAACUUACCAAGGUUGUU
<i>Listeria innocua</i> Clip11262	GUUUUAG--AGCUAUGUUAUUUUGAAUGGUACAAAAAC • AAAUGAACGUAACAAAACUUAUAGAUUGUUUA
<i>Streptococcus thermophilus</i> LMD-9	GUUUUAG--AGCUGUGUUGUUUCGAAUGGUCCCCAAAC • AAAUGAGCGACACAACAAAGCUUACCAAAGUUUGG
<i>Streptococcus mutans</i> UA159	GUUUUAG--AGCUGUGUUGUUUCGAAUGGUCCCCAAAC • AAAUGAACGACACAACAAAGCUUACUAAGGUUGG

B

C

D

<i>Eubacterium yurii</i> subsp. <i>margaretiae</i> ATCC 43715	GUUUGAG-A-ACCUUGUAAAUCAAUAAGUAUGUAAAAC • CAAACUUGAGUGGAACAUUUAGUUACAUUAC-UAAA
<i>Solobacterium moorei</i> F0204	GUUUGAG-A-ACUAUGUAAAUAU---GCUGGUA---GCAAAAC • • • CAAACUUGAUAUGAUACAUUUAAAGAAUGACCACAUUCUGUAAAAG

Supplementary Figure S11. Compensatory mutations within CRISPR repeat:tracrRNA duplexes suggest coevolution of type II CRISPR-Cas RNA components. Basepairings: upper sequence, repeat (5' to 3' orientation); lower sequence, tracrRNA antirepeat (3' to 5' orientation). Highlighted are nucleotides different from the corresponding ones in top duplexes for every panel. Color bars indicate clusters of Cas9 tree of selected loci (see Figure 4). **(A, B)** RNAs associated with very closely related Cas9 orthologs (same BLASTclust group, see Supplementary Materials and Methods and Supplementary Table S1). **(C, D)** RNAs associated with Cas9 orthologs from the same Cas9 phylogenetic tree

cluster but from different BLASTCLUST groups. Note the conservation of the cofolding pattern in (**A**, **B**, **C**) and 5' part of (**D**). The conserved structure at the 5' part of the repeat in (**D**) is most likely present in mature crRNA:tracrRNA duplex. Note, that bacterial strains in (**C**) belong to different families (*Bacilli* and *Clostridia*) and in (**D**) to different phyla (*Firmicutes* and *Tenericutes*).

<i>F. novicida</i> U112	CAACGTCTAGTTGCCATCCATTAAAGTGATGTTCACTAATTCTTAATCCAAAAGGAATAATTGGTCT-ATAACTATTATAACAATACTGTATAAAAACAATACTTAAAGGAA
<i>F. novicida</i> Fx1	CAACGTCTAGTTGCCATCCATTAAAGTGATGTTCACTAATTCTTAATCCAAAAGGAATAATTGGTCT-ATAACTATTATAACAATACTGTATAAAAGCTATAAAAACAATACTTAAAGGAA
<i>F. novicida</i> 3523	CAACGTCTGGTTGCCATCCATTAAAGTGATGTTCACTAATTCTTAATCCAAAAGGAATAATTGGTCT-ATAACTATTATAACAATACTGTATAAAAGCTATAAAAACAATACTTAAAGGAA
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	TATTTAAAGACCCTTAGCTGGTAGTTGAGATTTTATTAAACTGATAATCTCAATATTAGTTAGTTACATTGGTAGCTATCGCGTTATAGCAACCCATATGCTTATACACTTTATTAG
<i>F. novicida</i> Fx1	TATTTAAAGACCCTTAGCTGGTAGTTGAGATTTTATTAAACTGATAATCTCAATATTAGTTAGTTACACTTTGGTAGCTGTGCGTTATAGCAACCCATATGCTTATACACTTTATTAG
<i>F. novicida</i> 3523	TATTTAGAAACTCTTTA-TTTGCTAGTTGAGCTTTATTAAACTGATAATCTCAATATTAGTTAGTTACACTTTGGTAGCTATGGTATTGGCAACTTATATTCTATACTACTTTATTAG
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	CTATTGTTAAGCTAATTGAGCGATATATCATTAACTATTCTAAAATGATTCAAAAATAATTCTATAAAATAATTGCTTAAAGTCTATCAAACACTTGTGCTATTCTGAAAGTAGTATTAGATAA-
<i>F. novicida</i> Fx1	CTATTGTTAAGCTAATTGAGCGATATATCATTAACTATTCTAAAATGATTCAAAAATAATTCTATAAAATAATTGCTTAAAGTCTATTAAACACTTGTGCTATTCTGAAAGTAGTATTAGATAA-
<i>F. novicida</i> 3523	CTATTGTTAAGCTAATTGAGCGATATATCATTAACTATTCTAAAATGATTCAAAAATAATTGCTTAAAGTCTCTAAACACTTGTGTTACAGCTCTAGATTAGGTTTATAAGCAAG
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	----AATCTTATGAAAAATTA--TTTTAAATGAGTTTTAGAACTAGACTATTAAACTATGTTAGAATTATCAAAGCAAATG-AAAAAGATTAGCTGTAAGTTTCAGTTGTTAGATTGGTATGACTTGTGTTA-
<i>F. novicida</i> Fx1	----AAGCTTATGAAAAATTA--TTTTAAATGAGTTTTAGAACTAGACTATTAAACTATGTTAGAATTATCAAAGCAAATG-AAAAAGATTAGCTGTAAGTTTCAGTTGTTAGATTGGTATGACTTGTGTTA-
<i>F. novicida</i> 3523	TAATGATCTTACTAAATATTGGTAAAG---TTTCAGT-TACTAGATTCTGATATACTCTTAGGATTAGTCCAGCAAATGAAAAAGATTAGCTGTAAGTTTCAGTAGCTAGATTGGTACATACTAGTGTGTTAAGG
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	GTTTAAAGTAGCT----AGAAAATTCACTTTAGACCTAC-TTATTGGTCT-ATGCTT-GGTAGTATGGAGATCAACATTTGCTGAAACTGTTTCGTGATTATATTGGTATGACAATGTC
<i>F. novicida</i> Fx1	GTTTAAAGTAGCT----AGAAAATTCACTTTAGACCTAC-TTATTGGTCT-ATGCTT-GGTAGTATGGAGATCAACATTTGCTGAAACTGTTTCGTGATTATATTGGTATGACAATGTC
<i>F. novicida</i> 3523	GTTAGGATTAACCTAACATAAAGAACATTAGCTGAAATTTCGTAAATTAGGTTATTGATATACTGCTTCAAATCTGACTTGAATTGATATTGCTATATAACTATTTAGCGTTATATTCTGATATACTTACCCAGA
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	TAGAAACTTTAGGAATAATTACAAAATTAATTAA-----
<i>F. novicida</i> Fx1	TAGAAACTTTAGGAATAATTACAAAATTAATTAA-----
<i>F. novicida</i> 3523	TAGAAACTCTAAATATTTCGTAATTAAAGTTTCGGTAGCTAGGTTATTGATGACAATAAAACTTACCTGTTGATGCTAGGTATTAGTGTATTATAAGTAGCTAGATTGGTATGACA
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	-----GTTTCAGTTGCTGAATTATTGGTAAACTACTGTTAGACAATGGCAAGCTGATTACTTTAGTGTGAT-----GTTTCAGTTGCTGAATTATTGGTAAACTACTGTTAGAAGGATTTAAC
<i>F. novicida</i> Fx1	-----GTTTCAGTTGCTGAATTATTGGTAAACTACTGTTAGCTTAACTCATCAAATCACCTATAAAAGTTTCAGTTGCTGAATTATTGGTAAACTACTGTTAGATCAAATCGACAG
<i>F. novicida</i> 3523	ATGACTATAACCAACACTCGTCCAGAGTTTCAGTAGATTATTGATATACTGCTGTTAGTTCTTAACTCTAGGTATCTAAACAAATTCTGTTAGCTAGATTGGTATGACAATGCA
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----

<i>F. novicida</i> U112	CTTAAAATTATCCAATTTATGTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGTGCTTAATCCTCAGCAGCAACACTTAATTCAAGAATGTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGTTACACATTCAAGCTGGT
<i>F. novicida</i> Fx1	TAAATACATATGAGAACG--GTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGTAACGTTGGAAATAAGTCACATTTCCTTT--GTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGATAACATTGTATTAA
<i>F. novicida</i> 3523	AGATATTTATTTAATTTCAT-GTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGCTAATGTAGAAGCTATATTCTATTCAACATT-GTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGCTGGTGGAGGCAGTCAA
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	TCTTTAGCATTATT--AACGTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGAGTAATGCCAATGCTTTAAACTACTGATATATACTGTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGGCAATGACAGAAAACAGCACA
<i>F. novicida</i> Fx1	TGAGAGAGACAGATCCAACGTTTCAGTTGCTGAATTATTTGGTAACTGTTAGTTACCGTAAAGACTGGTTACTACATCCAGA--
<i>F. novicida</i> 3523	ACTGGCGCCGAGGGTGA--GTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGAGTGCTGAACTCAAGCCTGTTGAAACGG--GTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGATAATATGCTAACATATGT
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	GATAATAAAAAACTAGTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGCTATAGGGTACCTATCTTGAGTGTGCAAATAAGTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGGTAATGGCAGGTTGTATGGTC
<i>F. novicida</i> Fx1	-----
<i>F. novicida</i> 3523	TTTCTAGCGATATT-GTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGTTAACCGAGATAATGAATATCAAGAGTATGTCGA---GTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGCAATTATGGAACGATTATGGGAT
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	ATATAGGAGT--GTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGACTATTAAGATAACTCGACCAATATTGACAAA--GTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGATCTGGTAAGATGTAATCGAAATTGAT
<i>F. novicida</i> Fx1	-----
<i>F. novicida</i> 3523	TGGTTAGATGAAAGTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGCTATATTGGTTATAAAGTCAACTTCGAGGCCTAGTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGTGCTATGAGCATTGCAAGCAAAACAAT
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	CCGCTTAAGTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGCAGTCTCGCAATGACACAACCACAAGCCGATAGTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGAAGAGATATCTAACCTTGGTTTGGTTTAC
<i>F. novicida</i> Fx1	-----
<i>F. novicida</i> 3523	AAAAAAAAAGTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGCTGCTGCCATACGGGTACAATTCAAGGTTATATGGTTCAAGCTAGCTAGATTATTTGATATACTGCTGTTAGTGCCATACGACATATATTCTTTTAAAT
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	CGTTTCAGTTGCTGAATTATTTGGTAAACTACTGTTAGGGGAGTGATTAGCAGTTGATATACCAATCTG-----
<i>F. novicida</i> Fx1	-----
<i>F. novicida</i> 3523	-GTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGACCAAAAGCAACATTGCAATTGCAAGATAGTGCTGGTTTCAGTAGCTAGATTATTTGATATACTGCTGTTAGTGCTAGGCTAAAGGCTGTTAACAGCTGTTACGTTTCAG
<i>F. holarktica</i> LVS	-----
<i>F. tularensis</i> SCHU S4	-----
<i>F. novicida</i> U112	----- GTTTCAAGTTGCTG
<i>F. novicida</i> Fx1	----- GTTTCAAGTTGCTG
<i>F. novicida</i> 3523	TAGCTAGATTATTTGATATACTGCTGTTAGCCTCAACTTAGCCTTAGGCTCAGTCCTTGGCTTCAAGCTAGCTAGATTATTTGATATACTGCTGTTAGTGCTAGGTTACTGGTTTGCTGAAGGTGGTTACTGGCAAGTTTCAGTAGCTA
<i>F. holarktica</i> LVS	----- GTTTCAAGTTGCTG
<i>F. tularensis</i> SCHU S4	----- GTTTCAAGTTGCTG

<i>F. novicida</i> U112	AATTATTTGGTAAACTACTGTTAGAGCCAAAACAAACCTAGTGTTCAGGCCTTGAGAGTTTTAAATGGTATTTTCGCAAATACCCAATTGTAaaaaaaCGTAGACATTGCGAAAAAAT	ATCTAAAATTATAATGTAC
<i>F. novicida</i> Fx1	AATTATTTGGTAAACTACTGTTAGAGCCAAAACAAACCTAGTGTTCAGGCCTTGAGAGTTTTAAATGGTATTTTCGCAAATACCCAATTGTAaaaaaaCGTAGACATTGCGAAAAAAT	ATCTAAAATTATAATGTAC
<i>F. novicida</i> 3523	GATTATTTGATATACTGCTGTTAGAGCCAAGAACAAACCTAGAGTTGCAAGGCTTGAGAGAATTATGGCTGGTTTCGCAAACATCCTACTGTAaaaaaaGTAGACATTGCGAAAAAAT	ATCTACAATTAAAAGCGTAC
<i>F. holarctica</i> LVS	AATTATTTGGTAAACTACTGTTAGAGCCAAAACAAACCTAGTGTTCAGGCCTTGAGAGTTTTAAATGGTATTTTCGCAAATACCCAATTGTAaaaaaaCGTAGACATTGCGAAAAAAT	ATCTAAAATTATAATGTAC
<i>F. tularensis</i> SCHU S4	AATTATTTGGTAAACTACTGTTAGAGCCAAAACAAACCTAGTGTTCAGGCCTTGAGAGTTTTAAATGGTATTTTCGCAAATACCCAATTGTAaaaaaaCGTAGACATTGCGAAAAAAT	ATCTAAAATTATAATGTAC
<i>F. novicida</i> U112	AAATAATTAAATGCTCTGTAATCATTAAAAGTATTTGAACGGACCTCTGTTGACACGCTCTGAATAACTAAAAGCAAAATTGCCACCTAAGTGGCTTTTTGTTGGCAATTATATCTGTCGCTCTAGCTCGAT-TAC	
<i>F. novicida</i> Fx1	AAATAATTAAATGCTCTGTAATCATTAAAAGTATTTGAACGGACCTCTGTTGACACGCTCTGAATAACTAAAAGCAAAATTGCCACCTAAGTGGCTTTTTGTTGGAAATTATATAACTTGTGCTCTAGCTCGAT-TAC	
<i>F. novicida</i> 3523	AAATAATTAAATGCTCTGTAATCATTAAAAGTATTTGAACGGACCTCTGTTGACACGCTCTGAATAACTAAAAGCAAAATTGCCACCTAAGTGGCTTTTTGTTGGAAATTATATAACTTGTGCTCTAGCTCGAT-TAC	
<i>F. holarctica</i> LVS	AAATAATTAAATGCTCTGTAATCATTAAAAGTATTTGAACGGACCTCTGTTGACACGCTCTGAATAACTAAAAGCAAAATTGCCACCTAAGTGGCTTTTTGTTGGCAATTATATCTGTCGCTCTAGCTCGAT-TAC	
<i>F. tularensis</i> SCHU S4	AAATAATTAAATGCTCTGTAATCATTAAAAGTATTTGAACGGACCTCTGTTGACACGCTCTGAATAACTAAAAGCAAAATTGCCACCTAAGTGGCTTTTTGTTGGCAATTATATCTGTCGCTCTAGCTCGAT-TAC	
<i>F. novicida</i> U112	ATCAAGATGGC-TGAGAAATTAA-----ATATGGAGATT-----CTAAATTTG---CGAGATTTCCACCTTCAGTGGCAATTAGA-----	
<i>F. novicida</i> Fx1	ATCAAGATGGC-TGAGAAATTAA-----ATATGGAGATT-----CTAAATTTG---CGAGATTTCCGCCTTCAGTGACAATTAGA-----	
<i>F. novicida</i> 3523	ATAAAAATGTCAAGCTAAATTATTCAGCATCTCATTATAAATATAGATTTATAAAAAAAACATAATTACATAATTGATGACAAAATTACTACTTTATGCCTAGCAGACTATCTTTACTTTATGATATCTTTAGTTAAAA	
<i>F. holarctica</i> LVS	ATCAAGATGGC-TGAGAAATTAA-----ATATGGAGATT-----CTAAATTTG---CGAGATTTCCGCCTTCAGTGGCAATTAGA-----	
<i>F. tularensis</i> SCHU S4	ATCAAGAAGGC-TGAGAAATTAA-----ATATGGAGATT-----CTAAATTTG---CGAGATTTCCGCCTTCAGTGGCAATTAGA-----	
<i>F. novicida</i> U112	-----GGTTTACTA TCAACGATCATTACAAAAGTTGTAATATTCACATTGTATACATTGATCCGATGCAGATGA	
<i>F. novicida</i> Fx1	-----GATTTACTA TCAACGATCATTACAAAAGTTGTAATATTCACATTGTATACATTGATCCGATGCAGATGA	
<i>F. novicida</i> 3523	TTATTCTGACAAGCTCGAAGGAATTGATT TTAGCGATCATTACAGAAAATTAAATCTCACACTGTATACATTGATCCGATGCAGATGA	
<i>F. holarctica</i> LVS	-----GATTTACTA TCAACGATCATTACAAAAGTTGTAATATTCACATTGT-----TCCGATGCAGATGA	
<i>F. tularensis</i> SCHU S4	-----GATTTACTA TCAACGATCATTACAAAAGTTGTAATATTCACATTGT-----TCTGATGCAGATGA	

Supplementary Figure S12. Alignment of *Francisella* sp. CRISPR loci. See Figure 6. *F. novicida* DNA sequence from the 3'-end portion of the radical SAM protein-encoding gene located upstream of the repeat-spacer array to the 5'-end portion of the *cas4* gene was aligned with the *F. holarctica* and *F. tularensis* region between *cas4* and the gene encoding the transposase inserted within the CRISPR locus using MUSCLE with manual corrections. Blue highlighted, 3'-end of the radical SAM protein-encoding gene; yellow highlighted, scaRNA; red highlighted, degenerated repeats; grey highlighted, repeats; red highlighted, tracrRNA; green highlighted, 5'-end of the *cas4* gene. See as well Supplementary Figure S9 for the degenerated repeat alignments.