

Motif	I	II	III	IV	V	VI	Source	ref.
uvrD	26	VLGAGSGKTRVLV_174	NILVDEFQNTN_16	VMIVGDDDDQSIY_26	QNYRSTSI_267	IMTLHS-AGKL-EFPQVFIWG_23	LAYVGVTRA	(20)
rep	19	VLGAGSGKTRVIT_175	YLLVDEYQDTN_16	FTVVGDDDDQSIY_26	QNYRSSGI_271	IMTLHA-SKGL-EFPYVVMVG_22	LAYGVITRA	(2)
recB	20	IFASAGTKTFTTA_345	VAMIDEFQDTN_18	LLILGDPKQAIY_24	TWRSAPGM_286	IVTHK-SKGL-EFPLVWLF_44	LLYVALTRS	(4)
recD	164	ISGPGGTGKTTTVA_82	VLVWDEASMD_16	VIFLGRDQLAS_24	QLSRLGTGH_198	AMTVHK-SQGS-EFDHAALIL_11	LVYTAVTRA	(21)
EBV	69	ITGTAGAGKSTSVS_113	VIVVDEAGTLS_26	IVCVGSPTQIDA_44	NNKRCITDVQ_426	AMTIK-AQGL-SLNKVAICF_9	HVYVALSRA	(22)
HCMV	117	VTGTAGAGKSTSIQ_125	IIVIDECEGLM_26	IICVGSPTQTEA_44	NNKRCITDLD_513	AMTIK-SQGL-SLEKVADEF_10	HVYVAMSRV	(6)
HSV	94	ITGNAGSGKSTCVQ_136	VIVIDEAGLLG_26	LVCVGSPTQTAS_44	NNKRCVEHE_442	AMTIR-SQGL-SLDKVAICF_8	SAYVAMSRT	(6)
VZV	87	ISGNAGSGKSTCIQ_135	VIVIDEAGLLG_26	IVCVGSPTQIDS_44	NNKRCQEDD_447	AMTIAR-SQGL-SLEKVAICF_8	SVYVAMSRT	(23)
PIF	255	YTGSAGTGKSLILR_46	ALVWDEISMLD_25	LIFCGDFFQLFP_29	KVFRQRGDV_219	MQTIHQNSAGKRLRLVIFRKA_33	QAYVALSRA	(10)
AIMV	821	VDVGAGCGCKTTNIK_55	RLIFDECEFLQH_15	VIGFGDTEQIIPF_22	ITWRSPADA_66	IFTTHE-AQCK-TFDNVYFCR_19	NGLVALSRH	(1)
BMV	687	VDVGAGCGCKTTAIK_54	RLVDEAGLLH_15	VLAGDTEQISF_22	KTYRCEQDV_78	IKTVHE-AQGI-SVDNVTLVR_13	YCLVALTRH	(1)
CMV	709	VDVGAGCGCKTTAIK_54	RLVDEAVLLH_15	ALCGDSEQIAF_22	TTFRSPQDV_79	IKTVHE-SQGI-SEDHVTLVR_13	YCLVAVTRH	(1)
TMV	829	VDVFGCGCKTKELL_57	RLFIDEGLMLH_15	AYVGTQOIPY_24	TLRCPADV_62	VHTVHE-VQGE-TYSDVSLVR_14	HVLVALSRH	(1)
ToMV	829	VDVFGCGCKTKELL_57	RLFIDEGLMLH_15	AYVGTQOIPY_24	TLRCPADV_62	VHTVHE-VQGE-TYADVSLVR_14	HVLVALSRH	(24)
TRV	903	VDVFGCGCKSTMIY_56	VLHDEALMAH_15	CICQGDQOIPY_24	EYRSPADV_64	VSTVHE-SQGE-TFKDVSLVR_13	YLVALSRH	(1)
SFV	181	VDFGPGSGSKAIK_50	ILVYDEAFACH_16	VVLGDPKQCGF_18	ISRRCTRPV_58	VMTAAA-SQGL-TRGVYAVR_14	HVNVLTRT	(1)
SV	183	VIGTFPGSGSKAIK_50	VLVYDEAFACH_16	VVLGDPKQCGF_24	ISRRCTRPV_58	VMTAAA-SQGL-TRGVYAVR_14	HVNVLTRT	(1)
IBV	1209	VQGGPFGSGKSHFAI_54	ILLVDEVSMILT_15	VVYVGDPAQLPA_30	KCYRCPEKI_82	VQTVDS-SQGS-EYDVIKFCV_11	RFNVALTRA	(25)
BNYV1	893	VKGPGGTGKSLFLIR_48	IIFVDEFTAYD_11	IYLVGDEQGTGI_25	MNFRNPVHD_72	KITVRA-NQGS-TYDNVFLV_12	LNLVALSRH	(26)
BNYV2	121	VLGAGFGGKSTSIK_49	ITLVDEVTIRVH_11	VICFGDPAQLN_18	ASRREGKAT_67	SILYSD-AHQGT-YDVITLIL_13	VRVALTRA	(26)
BSMV2	267	ISGFGGSGKSTIVR_41	LLIIDEVTIAE_11	VLLVGVDAQGLA_28	TTYRLQET_62	CALAID-VQGE-EFDSVTLFL_12	LRLVALSRH	(27)
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	V G A G K S	V D E	G D Q	R	T	S Q G E	V	VALSR
	I A P T	I	S			L A K S A		TLVT
	Y	F				V H T		GM
						N A R		

Alignment of conserved motifs from six protein families (single-letter code). Hyphens, gaps introduced during alignment. The number of residues between each motif is shown. At the base of alignment, characteristic residues have been shown where there are less than five alternatives. The main RNA virus family includes: AIMV, alfalfa mosaic virus 1a; BMV, brome mosaic virus 1a; CMV, cucumber mosaic virus 1a; TMV, tobacco mosaic virus (common strain) p126; ToMV, tobacco mosaic virus (tomato strain) p126; TRV, tobacco rattle virus p134; SFV, Semliki Forest virus nsP2; SV, Sindbis virus nsP2. The last two are alphaviruses infecting animals. Conserved regions in this protein family were used to construct motifs for the program TPT<sup>14</sup>, which extracted uvrD from the Protein Information Resource database. A general database search using FASTP<sup>15</sup> then matched a region from recD with Epstein-Barr virus (EBV) BBLF4. Related genes in other herpesvirus: human cytomegalovirus (HCMV) PS3 (J.A. Martignetti, personal communication), herpes simplex virus (HSV) UL5, and varicella-zoster virus (VZV) gene 55, are strongly conserved. Similarities between the yeast PIF sequence<sup>6</sup> and uvrD, and between infectious bronchitis virus (IBV) F2, beet necrotic yellow vein virus (BNYV1) 237K protein and the alphavirus nsP2 family were noted by the original authors. The relationship between the (BNYV2) 42K and barley stripe mosaic virus (BSMV2) 58K proteins was also noted, but their similarity to all the other proteins was not. Asterisks, positions used for statistical analysis<sup>16</sup>. The probability of each motif occurring by chance was  $\sim 10^{-6}$ , and for the six- and seven-member patterns  $1.9 \times 10^{-33}$  and  $1.6 \times 10^{-39}$ , respectively.

DNA-dependent ATPases sharing 37% amino-acid identity<sup>2</sup>. The recB and recD are subunits of ExoV. The former probably carries out the helicase activity of the ExoV holoenzyme because it has been shown to bind ATP<sup>3</sup> and single-stranded DNA<sup>4</sup>; also, it has 20% amino-acid identity with the above helicases. Its extra sequences presumably carry out ExoV-specific functions. The recD does not have a demonstrated helicase function, binds ATP only in the presence of the other subunits<sup>3</sup>, and has diverged further than the others. The structural resemblances between these proteins strongly suggest that they belong to a DNA helicase family that is distinct in evolutionary terms from other *E. coli* helicases, although they are related to PIF, a yeast nuclear gene product involved in mitochondrial DNA repair and recombination<sup>5</sup>.

Genetic data in herpes simplex virus<sup>6</sup> show that UL5 is essential for DNA replication, a finding that is consistent with its stringent conservation among other herpesviruses. UL5 could also be a helicase<sup>6</sup>, a function possibly common to all groups within the superfamily, although the level of similarity between the groups is too low to allow a direct functional comparison. The available evi-

dence shows that the RNA viral protein domains containing these conserved motifs are involved in replication<sup>7</sup>, but are distinct from the suspected polymerase domain<sup>8</sup>.

The very broad occurrence of these motifs in otherwise unrelated proteins strongly suggests that they all carry out crucial nucleoside triphosphate-dependent steps in nucleic acid replication. Motif I is well-known<sup>9</sup>, and is found almost exclusively in ATP and GTP binding proteins. Where the crystal structure is known<sup>10-12</sup>, the motif forms a loop which binds one of the phosphates, another being bound by an aspartate via a magnesium ion. There is a suitably located, strictly conserved aspartate in motif II. Motif III was previously noted<sup>13</sup> to resemble a conserved region from viral DNA polymerases. This study shows that the proteins have no overall relationship, though it remains possible that they have similar local folds. Note also that only the presumed DNA binding proteins have tyrosine at position 3 of motif VI and another motif between motifs I and II.

These motifs probably represent common secondary structures that make up functional sites for pyrophosphate, magnesium or nucleic acid binding, while the rest of the sequence provides the struc-

tural framework, the specificity and any supplementary functions. The resolution of the exact structure, function, and evolutionary significance of these motifs is still awaiting further crystallographic analysis.

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## Lifting heavy loads not just by the Egyptians

SIR—The astute suggestion made by John Cunningham (*Nature* **332**, 22-23; 1988) that certain old Egyptian art work illustrates a method of raising heavy weights prompted me to look into that compendium of primitive clever invention, Francis Galton's *Art of Travel* (5th edn, David and Charles, Newton Abbot, 1971).

Sure enough, two related systems, which make use of this principle of successive small forces being applied to a series of springy supports, are mentioned under the heading 'Accumulation of Efforts'.

To lift and swing forward a heavy log in a tropical forest, 'the labourer gets hold of one of these creepers that runs from the top of the boughs of a tree in the direction in which he wants to move his log, and pulling this creeper home with all his force, bending down the bough, he attaches it to the log; then he goes to another creeper and does the same with that; and so on until he has accumulated strain of many bent boughs, urging the log forward and of sufficient power to move it'. The other example concerns a commercially available 'accumulator', consisting of cords of india-rubber each hooked to a fixed ring at one end, the other ends being then hooked one by one to the object to be moved.

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