

Table S4: Details of HMM profiles used in this work

Profile	Structs.	Seqs	Sequence pattern of identical amino acids in alignment with highlighted active site residues [#]
ALKB	4	1	N _{x{10,25}} H _{x{1}} D _{x{12,14}} S _{x{25,45}} G _{x{13,14}} H
ARGI	2	1	T _{x{7,8}} ALAx _{9,10} D _{x{3}} FL _{x{2}} A _{x{3}} A _{x{2}} LP _{x{2}} L _{x{15,16}} L _{x{2}} RG _{x{1}} PV _{x{1,2}} D _{x{2}} L _{x{2}} TP _{x{19}} L _{x{4}} LG _{x{3}} G _{x{6}} G _{x{1}} V _{x{2}} DV _{x{1}} P _{x{2}} G _{x{1,2}} H _{x{1}} L _{x{1}} S _{x{2}} S _{x{3}} L _{x{2}} HTE _{x{1}} A _{x{7}} YV _{x{1}} L _{x{3}} R _{x{6,8}} T _{x{4}} V _{x{1,3}} L _{x{2}} LDE _{x{1}} T _{x{3}} L _{x{9}} D _{x{15,30}} V _{x{2}} L _{x{1}} G _{x{4}} P _{x{1}} L _{x{2}} D _{x{4}} AP _{x{1} ,3}D _{x{5}} A _{x{3}} L _{x{4}} D _{x{7}} L _{x{2}} GD _{x{4}} DN _{x{1}} R _{x{2}} H _{x{1}} R _{x{2}} F _{x{1}} PR _{x{1}} DG _{x{1}} DRWL _{x{1}} R _{x{2}} I x{2}D
ASPA	3	4	H
CHLO	2	1	F _{x{9}} F _{x{3}} G _{x{1}} IGP _{x{5}} PEEM _{x{7}} R _{x{2}} L _{x{2}} R _{x{2}} A _{x{1}} Y _{x{9,10}} NYDRHL D _{x{1}} D _{x{1}} L _{x{2}} H _{x{2}} R _{x{1}} {1}E _I _{x{2}} R _V _{x{1}} S _{x{2}} G _{x{4}} CWR _{x{1}} E F _{x{1}} PKY _{x{1}} G _{x{1}} EGTD WHQA _{x{1}} TF A _{x{1}} A _{x{1}} G _{x{1}} PQ _{x{2}} 2}WP _{x{4,5}} F _{x{1}} G _{x{1}} I _{x{1}} VW T _{x{1}} FT _{x{5}} NGC _{x{1}} Q _{x{2}} PG _{x{4,52}} A _{x{1}} P _{x{3}} K _{x{1}} G _{x{3}} IF _{x{1}} S _{x{2}} 2} MH _{x{1}} S _{x{1}} P _{x{7,8}} R _{x{1}} G _{x{3}} RYVP _{x{2}} V _{x{1}} VYP _{x{13}} L _{x{3}} G _{x{1}} V _{x{3}} G _{x{6}} NR _{x{7}} G
CLAS	1	1	M _{x{1}} S _{x{0,2}} VDCT _{x{1}} Y _{x{2}} EL _{x{1}} ALAx _{2} LP _{x{2}} PRADL _{x{2}} FLD _{x{1}} A _{x{1}} T _{x{1}} AA _{x{1}} LP _{x{2}} LA x{1}ALDTFN _A _{x{1}} GSEDG _{x{1}} LLLRGLPV _{x{1,2}} D _{x{2}} LP _{x{1}} TP _{x{1}} STPAP _{x{1}} DR _{x{2}} L _{x{1}} MEAM _{x{2}} }L _{x{1}} GRRLGLHTGY _{x{1}} ELRSGTVYHDVYPSGAH _{x{1}} LS SETSETLLEFHTE MA Y H _{x{1}} LQPNYVMLA CSRADHE _{x{2}} A _{x{1}} TLV _{x{1}} SVRKALPLLDE _{x{1}} TRARL _{x{1}} DR _{x{2}} PCCV DVA FRGGVDDPGAIA _{x{1}} V KPLYGD _A _{x{1}} DPFLGYDRELLAPEDPADKEAV _A _{x{1}} LS _{x{1}} ALD _{x{1}} VT _{x{2}} V _{x{1}} L _{x{1}} PGD _{x{1}} LI _{x{1}} DNFR TT HARTPFSPRWGKDR WLHRVY IRTDRNG _{x{1}} LSGGERAGD _{x{3}} F _{x{1}} PR
COLY	1*	13	R _{x{130,172}} W
CP3H	1*	9	H _{x{27}} F _{x{65,73}} E _{x{17}} P _{x{52,115}} P _{x{2}} P _{x{3,4}} F _{x{27,48}} S _{x{2}} H _{x{17}} S _{x{19,77}} V _{x{1}} A _{x{5}} R _{x{5}} W

CP4H	3	6	----
CYCL	2	1	Sx{3,4}LLLRGx{7,27}Ex{41,48}Hx{4}Yx{4}Px{6}Cx{2}Ax{7,8}Tx{3}Sx{3}ALx{1}Lx{1}Dx{5,8}Rx{9,12}Ax{41,52}Ex{2}Ax{1}Lx{3}Lx{12,14}Dx{2}Ix{1}DNx{4}Hx{1}Rx{8,11}Rx{1}Lx{1}RVx{1}Ix{12}R
DACS	1	4	VPx{1}Fx{4}Lx{2}Gx{12,13}Gx{1}FYLx{3}Gx{14,15}DFFx{2}Gx{3}EKx{1}AVx{7}RRGx{3}LEx{1}ESTAx{2}Tx{1}TGx{1}Yx{1}DYSx{3}SMGx{3}Nx{1}FPx{3}FEx{2}Wx{2}YFx{3}Yx{1}Ax{4}ARx{1}VLx{3}Gx{14,17}Px{1}Lx{1}Ry{1}RyFPx{1}VPEx{1}Rx{1}AEx{2}Px{1}RMx{1}PHYDLSx{2}Tx{3}QTx{1}CANGFVSLQx{1}Ex{2}Gx{5}Px{4}Ax{2}Vx{1}CGAx{3}Lx{3}Gx{3}APx{1}Hx{1}Vx{2}Px{6}GSx{1}RTSSVFLRPx{2}DFx{1}Fx{1}Vx{3}Rx{2}Gx{2}Vx{4}Ex{2}TFx{2}Wx{1}Gx{1}NY
DSAT	3	18	----
ECTO	1	3	Dx{1}YPx{1}Rx{8,11}Rx{1}Dx{1}Vx{8,11}Px{5}Lx{6}GFLx{8}DEVx{9}Lx{12}Ix{1}Ex{2}Sx{3}RSx{1}Fx{1}VHx{5}Fx{5}Dx{1}Rx{5}Rx{3}Gx{1}DVYVHQSRINx{1}KPGFx{4}Fx{1}WHSDFETWHx{1}EDGx{1}Px{1}Mx{3}Sx{1}SIx{1}Lx{2}Nx{3}NGx{1}LMx{2}PGSHx{2}Fx{2}Cx{1}Gx{1}TPx{4}Kx{1}SLx{2}Qx{2}Gx{1}Px{2}Ex{1}Lx{7}Gx{0,1}Ix{3}TGx{1}AGx{4}Fx{2}Nx{2}HGSx{2}Nx{2}Px{2}Rx{1}Nx{1}Fx{1}Vx{1}Nx{3}Nx{4}Px{6}RP
FLAV	1	17	Rx{3}Ex{13,19}Px{17,24}Ax{3}WGx{6}Hx{16}FFx{1}Lx{3}EKx{26,32}Wx{9}Px{8}WPx{2}Px{9}Yx{14}LSx{3}Gx{5}Lx{13,16}Nx{1}YPx{1}CPx{1}Px{1}Lx{1}LGx{3}HTDx{4}Tx{6}Vx{1}GLQx{6,8}Wx{14}GDx{5}SNGx{6}Hx{3}Vx{5}Rx{1}Sx{3}Fx{2}Px{7,8}P
GBBH	1*	3	Ax{5}Ax{6}Wx{1}Dx{3}Sx{5}WLRDNCx{1}Cx{1}DCx{10}Lx{23,24}Wx{3}Hx{1}Sx{5}WLx{7}Qx{5,7}Ex{3}Px{4}Wx{5}LPx{6}Vx{3}Dx{5}WLx{5}VGx{3}Lx{1}Gx{5}Gx{4}Lx{1}KRx{2}Fx{6}Gx{4}Vx{2}Kx{1}DAx{3}AYTx{3}Lx{2}HTDx{1}Px{5}PGx{5}C
GIAC	0	81	Gx{3}Hx{1}Dx{13}Gx{8,12}Wx{15,16}Dx{6}N

HILY	11	12	-----
HP4H	3	5	-----
HYOS	0	6	ATx{5}Sx{7,28}Fx{11}Vx{2}Gx{3,6}Px{1}Ix{1}Lx{7,13}Vx{2}Ix{2}Ax{4}Gx{1}FQx{2}NHGx{1}Px{3}Mx{9}Fx{18,26}Kx{2}LYx{13}Wx{1}DTx{13,17}Ex{1}Px{3}Rx{4}KYSx{3}Rx{1}Lx{9}Ex{1}LGLKx{14,15}Lx{2}YYPx{1}CPx{1}Px{2}Tx{1}Gx{3}Hx{1}Dx{4}Tx{1}Lx{1}Qx{3}Gx{2}Qx{6,7}Wlx{1}Vx{2}IPx{1}Ax{1}Vx{1}Nx{4}Lx{4}Nx{1}Kfx{4}HRVx{7}Rx{1}Sx{5}Gx{8,15}Px{1}KELx{4}NPPx{1}YKx{6}Fx{2}Ix{13}Px{3}N
NUHY	0	7	G
OGFD	0	2	Gx{6}Vx{3}Fx{21}Px{2}Hx{1}Vlx{8,21}Lx{4}KELMx{3,4}Fx{1}EKx{2}Dx{1}Yx{3}Qx{1,7}Dx{1}Lx{1}Kx{8}LRx{4,8}Ex{1}FRSx{4}Ix{6,8}Sx{2}Dx{1}Sx{3}Yx{5}Lx{2}HDDx{4}Rx{1}Ix{2}ILYLVx{1}Px{0,3}Wx{4}GGx{1}Lx{1}Lx{6,7}Fx{7}Sx{2}Px{1}WNx{1}Lx{1}FFx{1}Vx{1}Px{1,2}SFHx{1}Vx{1}EVx{3,4}Kx{1}Rx{2}ISGWfHx{1}Px{4}Px{1,4}NYx{17}Lx{7}Yx{16}Sx{7}Lx{1}Px{4}Kx{9}Vx{31,33}SEx{2}Fx{9}Kx{3}Lx{4}Ex{4,12}Dlx{9}Px{9}Sx{5}Sx{2}Qx{8,13}Sx{18}Lx{6}Ax{16}Px{13}Dx{2}LLx{6}Nx{3}LVYRDx{3}Lx{1}FVKx{10,19}WD
PHYT	2	3	Lx{3}Qx{7}Gx{1}LVx{51,82}Kx{36,38}Mx{1}Ix{1}Kx{1}Px{8,11}HQDx{7}Px{6,7}Wx{1}Ax{1}Ex{5}NGCLx{3}PGx{1}Hx{29,33}Vx{6}Gx{5}Hx{4}Hx{1}Sx{2}Nx{5}Rx{1}Ax{3}H
SULF	2	1	AAx{6}Px{3}Lx{47}Hx{3}Gx{13}Ix{7}Sx{11}Tx{18}Dx{7}Ax{49}Lx{9}Gx{2}Ix{1}Gx{2}Lx{3}Lx{16}Vx{3}Rx{1}Qx{17}Gx{4}Hx{36}WHTDVx{6}Px{10}Px{2}GGDTx{7}Ax{3}Lx{2}Px{7}Lx{3}Hx{19}Rx{13}HPx{1}VRx{1}HPx{6}Lx{5}Fx{3}Ix{7}Sx{2}Lx{3}Lx{12}Rx{2}Wx{6}IWDNRx{2}QHx{1}Ax{2}Dx{6}Rx{3}Rx{1}Tx{5}P
TDLP	1	1	Lx{6}Px{2}Gx{2}Ix{1}Gx{4,5}LSDx{5}Lx{3}Vx{4}VVx{2}Rx{1}Qx{3,6}Px{9}FGx{1}LHIHx{4}Hx{5}Ex{8,17}Dx{7}WHTDVx{5}PPx{8}Ex{1}Px{2}GGDTLx{4}Ix{1}Ax{3}LSx{1}Px{4}LSx{1}Lx{3}Hx{6,14}Ex{1}7}HPx{1}VRx{1}HPVx{2}Kx{2}Lx{1}VNx{2}Fx{3}IVx{5}ESEx{1}LLx{1}FLx{9,10}Qx{1}Rx{2}Wx{1}Px{4}IWDNRx{2}QHx{1}Ax{2}Dx{5,17}Rx{7}Lx{1}Dx{3}Y

TFDA	0	6	Px{6}Ax{6}Lx{2,3}Lx{19}Fx{2}Qx{5}Qx{1}Ix{1}Fx{3}FGx{8,23}Ix{16,25}Gx{3,6}WHx{1}DSx{13}Ax{2}VPx{2}GGx{1}Tx{1}Fx{2}Mx{2}Ax{3}Lx{14}Hx{8}Lx{18,27}PLVx{2}Hx{3}GRx{2}Lx{10,11}Gx{3}AEx{4}Lx{2}Lx{3}Ax{10}Wx{7}WDNx{1}Cx{2}Hx{10}R
THYD	0	2	Kx{3}Ex{1}Ax{20}Fx{12}Ax{1}Ex{3}Kx{3}Ex{11}Dx{4}Fx{5,18}Fx{4}Gx{4,7}Vx{1}IPx{3}Px{6}AAx{1}VLEx{1}Ax{5}LRx{4}GGx{2}Px{2}Gx{2}GYx{1}DYx{3}Px{3}KCRx{1}Tx{1}Fx{12}Px{8}Yx{4}Px{1}Hx{3}Qx{2}Ax{1}Px{6}Gx{2}FSTx{1}TVNx{2}FRTAx{1}HTDx{1}GDFx{2}Gx{1}GVLx{1}Vx{2}GEx{2}Gx{2}Lx{7}Fx{5}DVLLFDTx{1}Lx{1}Hx{1}NTEx{7,9}Wx{1}Rx{1}Sx{1}Vx{1}Yx{1}Rx{2}Lx{4}Cx{2}Ex{1}Rx{2}Lx{20}NGx{5}Px{4}Px{2}Lx{1,5}Px{3}AAx{6,12}GCx{2}AMx{3}Lx{0,32}Hx{5}Nx{3}Mx{2}Lx{8}DGLx{15,16}Nx{5}LGx{1,2}Fx{3}Gx{2}Lx{10,17}Dx{8}Gx{10}Rx{2,3}Lx{2}Vx{6}Qx{11}Wx{1}Nx{7,11}Fx{2}Lx{5}Qx{3}Gx{2}Gx{25}Nx{4}Lx{15}Yx{10}Mx{8}Rx{3}Lx{4}Ax{3}Ax{1}Ix{1}DSx{12,13}Qx{13}Ix{12}Ex{11}Gx{3}VLx{9}Rx{14,16}Ix{5}Ax{1}Rx{24}Nx{17}DFx{4}Hx{22}Rx{5}Vx{1}Vx{2}Lx{19,22}Fx{9}Rx{9}Cx{5}Dx{10}Kx{5,11}LNx{8}RE
THYE	0	2	Vx{5,7}Px{1}Ix{1}Fx{2}FLx{1}Gx{4}KLx{1}TAx{3}Lx{6}GFx{2}Lx{1}Nx{5}Dx{5,7}Fx{5}FFx{1}LPx{1}Ex{1}Kx{4}Wx{3}Ex{1}NRGYx{1}Ax{1}GREx{1}VTQx{1}TDPx{2}Ix{1}Kx{1}Rx{2}APDx{1}KEx{2}EIGx{5}Gx{0,1}Px{3}Px{1}Ex{6,10}FKx{2}MNX{1}FFx{5}LHx{1}Ex{1}MRAx{1}Ax{2}MGLx{4}FDx{2}Vx{6}LRLLx{1}YPx{2,4}Ex{2}Klx{2}Gx{2}RAGx{1}Hx{1}DYGSx{1}TLLx{1}QDx{2}GGLx{1}Vx{4}Gx{1}Fx{2}Ax{1}Px{3}TVVvNx{1}GDLLx{1}RWSNDx{2}KSTx{1}HRVVx{1}Px{7,9}Px{1}Rx{1}SIAx{1}Fx{1}NPx{5}Ix{1}Ax{1}PGTx{1,5}Ex{1}KYEx{6}YLVx{1}RLx{2}TY
XANT	0	3	Mx{3}Tx{6,10}Px{1}Gx{10,12}Dx{3}Lx{1}Dx{1}DFx{5}ALx{1}Tx{2}Vx{1}Vx{3}Qx{4}Px{2}QYx{2}Tx{1}Rx{1}DPx{4,6}YGHx{8,11}Sx{1}Lx{2}Dx{4}PHQPQVQx{2}Gx{1}Gx{5}EGx{5}Lx{1}HPx{1}Hx{1}TFHx{12,13}TRFYRWHIDx{1}ALYGx{2,5}PPx{2}TTLLx{3}VPx{4,5}Qx{3}Yx{1}Dx{1,2}Gx{9}Tx{1}Fx{1}SGx{6}LSx{6}Ax{4}Vx{1}YAPHPYx{3}Sx{1}Ax{5}GLx{3}Sx{3}Ex{5}LPx{3}Ex{5}LPMx{1}Wx{1}NPx{1}TGx{3}LQx{1}Hx{3}Vx{7}Gx{1}Vx{2,3}Lx{3}Rx{3}Yx{3}RPx{1}Ix{1}Px{2}VYAHx{1}Wx{2}GDLVx{1}FHNx{1}Gx{2}HSVx{1}Gx{1}Fx{3}Ex{1}RLx{2}QCNx{1}Ax{2,25}V

(#) Active site residues with side chains oriented towards HATM,

(*) PDB IDs in absence of HATM

HATM: atoms of Fe (II), 2-OG, and/or substrate