

Supplementary Figure 1

	$\alpha 1$ (A)	$\beta 1$ (A)	$\alpha 2$ (B)	$\beta 2$ (B)	$\alpha 3$ (C)
	HHHHH	EEEEEEE	HHHHHHHHHHHH	EEEE	HHHHH
			•••	♦♦	
LDHA_HUMAN	MATLKDQLIYNLLKKEEQTPQNKITVVGAVGMACAISILMKDLADELALVDVIEDKCLKG				
LDHA_BOVIN	MATLKDQLIQNLLKKEHVQNKITVVGAVGMACAISILMKDLADEVALVDVIMEDKCLKG				
LDHA_PIG	MATLKDQLIHNLLKKEEHPHNKITVVGAVGMACAISILMKELADELALVDVIMEDKCLKG				
LDHA_RABIT	MAALKDQLIHNLLKKEEHPQNKITVVGAVGMACAISILMKDLADELALVDVIMEDKCLKG				
LDHA_MOUSE	MATLKDQLIVNLLKKEEQAPQNKITVVGAVGMACAISILMKDLADELALVDVIMEDKCLKG				
LDHA_RAT	MAALKDQLIVNLLKKEEQVQNKITVVGAVGMACAISILMKDLADELALVDVIEDKCLKG				
LDHA_MONDO	MGTVQDQLILNLLKKEQTPHNKITVVGAVGMACAISILMKDLADELALVDVIEDKCLKG				
	*.:***** *:***** :.:*****:*****:*****:*****:*****:*****:*****				
		$\beta 3$ (C)	$\beta 4$ (D)	$\alpha 4$ (D)	
	HHHHHHH GGG	EEEE	GGGG	EEEE TT	HHHHHHHHHHHHHH
		♦	♦♦♦•	♦	♦
LDHA_HUMAN	EMMDLQHGSLFLRTPKIVSGKDYNVTANSKLVITITAGARQQEGESRLNLVQRNVNIFKFI				
LDHA_BOVIN	EMMDLQHGSLFLRTPKIVSGKDYNVTANSRLVITITAGARQQEGESRLNLVQRNVNIFKFI				
LDHA_PIG	EMMDLQHGSLFLRTPKIVSGKDYNVTANSRLVVITITAGARQQEGESRLNLVQRNVNIFKFI				
LDHA_RABIT	EMMDLQHGSLFLRTPKIVSGKDYSVTANSKLVITITAGARQQEGESRLNLVQRNVNIFKFI				
LDHA_MOUSE	EMMDLQHGSLFLKTPKIVSKDYCVTANSKLVITITAGARQQEGESRLNLVQRNVNIFKFI				
LDHA_RAT	EMMDLQHGSLFLKTPKIVSKDYSVTANSKLVITITAGARQQEGESRLNLVQRNVNIFKFI				
LDHA_MONDO	EMMDLQHGSLFLKTPKIVSKDYAVTANSKLVITITAGARQQEGESRLNLVQRNVNIFKFI				
	*****:*****:*** *****:*.*****:*****:*****:*****:*****:*****				
	$\alpha 4$ (E)	$\beta 5$ (E)	$\alpha 5$ (1F)	$\beta 6$ (F)	$\alpha 6$ (2F)
	HHHHHHH	EEEE	HHHHHHHHHHHH	GGEEEE	HHHHHHHHHHHHHH
		•••		•	•
LDHA_HUMAN	IPNVVKYSPNCKLLIVSNPVDILTIVAWKISGFPKNRVIGSGCNLDSARFRYLMGERLGV				
LDHA_BOVIN	IPNIVKYSPNCKLLVVSNPVDILTIVAWKISGFPKNRVIGSGCNLDSARFRYLMGERLGV				
LDHA_PIG	IPNIVKYSPNCKLLVVSNPVDILTIVAWKISGFPKNRVIGSGCNLDSARFRYLMGERLGV				
LDHA_RABIT	IPNVVKYSPHCKLLVVSNPVDILTIVAWKISGFPKNRVIGSGCNLDSARFRYLMGERLGV				
LDHA_MOUSE	IPNIVKYSPHCKLLIVSNPVDILTIVAWKISGFPKNRVIGSGCNLDSARFRYLMGERLGV				
LDHA_RAT	IPNVVKYSPQCKLLIVSNPVDILTIVAWKISGFPKNRVIGSGCNLDSARFRYLMGERLGV				
LDHA_MONDO	IPNIVKYSPNCKLLVVSNPVDILTIVAWKLSGFPKNRVIGSGCNLDSARFRYLMGEKLGVI				
	:**:*****:*****:*****:*****:*****:*****:*****:***				
	$\beta 7$ (G)	$\beta 8$ (H)	$\beta 9$ (J)	$\beta 10$	$\alpha 7$
	GGG	EE	TT EE	GGG	EETTEEHHHH
		•			
LDHA_HUMAN	HPLSCHGWVLGEHGDSSVPVWSGMNVAGVSLKTLHPDLGTDKDKEQWKQVHKQVVDSEAYE				
LDHA_BOVIN	HPLSCHGWILGEHGDSSVPVWSGVNVAGVSLKNLHPDLGTDADKEQWKAVHKQVVDSEAYE				
LDHA_PIG	HPLSCHGWILGEHGDSSVPVWSGVNVAGVSLKNLHPDLGTDADKEHWKAVHKQVVDSEAYE				
LDHA_RABIT	HALSCHGWILGEHGDSSVPVWSGMNVAGVSLKTLHPDLGTDADKEQWKQVHKQVVDSEAYE				
LDHA_MOUSE	HALSCHGWVLGEHGDSSVPVWSGVNVAGVSLKSLNPELGTADKEQWKQVHKQVVDSEAYE				
LDHA_RAT	HPLSCHGWVLGEHGDSSVPVWSGVNVAGVSLKSLNPELGTADKEQWKQVHKQVVDSEAYE				
LDHA_MONDO	HSSSCHGWILGEHGDSSVPVWSGVNVAGVSLKSLHPALGTDSDSEQWKQVHKQVVDSEAYE				
	*.*****:*****:*****:*****:*****:*****:*****:*****:*****				
	$\alpha 9$ (3G)	$\beta 11$ (K)	$\beta 12$ (L)		
	HHHHH	HHHHHHHHHHHHHH	EEEEEEEE TT TT	EEEEEEEEETTEE	
	•	•			
LDHA_HUMAN	VIKLKGYTSSWAIGLSVADLAESIMKNLRRVHPVSTMIKGLYGIKDDVFLSVPCILGQNGI				
LDHA_BOVIN	VIKLKGYTSSWAIGLSVADLAESIMKNLRRVHPVSTMIKGLYGIKEDVFLSVPCILGQNGI				
LDHA_PIG	VIKLKGYTSSWAIGLSVADLAESIMKNLRRVHPVSTMIKGLYGIKEDVFLSVPCILGQNGI				
LDHA_RABIT	VIKLKGYTTWAIGLSVADLAESIMKNLRRVHPVSTMIKGLYGIKEDVFLSVPCVILGQNGI				
LDHA_MOUSE	VIKLKGYTSSWAIGLSVADLAESIMKNLRRVHPVSTMIKGLYGINEDVFLSVPCILGQNGI				
LDHA_RAT	VIKLKGYTSSWAIGLSVADLAESIMKNLRRVHPVSTMIKGLYGIKEDVFLSVPCILGQNGI				
LDHA_MONDO	VIKLKGYTSSWAIGLSVADLAESIMKNLRRVHPVSTMIKGLYGINEDVFLSVPCILGQNGI				
	*****:*****:*****:*****:*****:*****:*****:*****:*****				
	$\beta 13$ (M)	$\alpha 10$ (H)			
	EEEE	HHHHHHHHHHHHHHHH			
LDHA_HUMAN	SDLVKVTLTSEEEARLKKASADTLWGIQKELQF				
LDHA_BOVIN	SDVVKVTLTHEEEACLKKSADTLWGIQKELQF				
LDHA_PIG	SDVVKVTLTPEEEAHLKKSADTLWGIQKELQF				
LDHA_RABIT	SDVVKVTLTSEEEAHLKKSADTLWGIQKELQF				
LDHA_MOUSE	SDVVKVTLTPEEEARLKKASADTLWGIQKELQF				
LDHA_RAT	SDVVKVTLTPEEEARLKKASADTLWGIQKELQF				
LDHA_MONDO	SDVVKVTLTTEEESRLKQASADTLWGIQKELQF				
	:**:***:***:*****:*****				

Multiple sequence alignment of human (UniProtKB accession code: P00338), bovine (UniProtKB accession code: P19858; 93.7% sequence identity), porcine (UniProtKB accession code: P00339; 92.8% sequence identity), rabbit (UniProtKB accession code: P13491; 93.7% sequence identity), murine (UniProtKB accession code: P06151, 93.9% sequence identity), rat (UniProtKB accession code: P04642; 94.3% sequence identity) and opossum (UniProtKB accession code: Q9XT87; 90.9% sequence identity) LDH-A. The active site loop is highlighted in yellow and the catalytic residue His193 is highlighted in orange. Other active site pocket residues are marked with a rhombus or dot depending on their proximity to the adenine or oxamate/nicotinamide moieties, respectively. Secondary structure elements are shown above the sequences whereby E denotes for β -strand, H for α -helix, G for 3_{10} helix and T for β -turn. A sequential and the historical (in brackets) nomenclature are shown for secondary structure elements.