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**Supporting Information**

**for**

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**Ubiquitination of HLA-DO by MARCH family E3 ligases**

Supplemental Figure 1. **Lysine residues are conserved in the cytoplasmic tail of mammalian DO $\beta$  molecules.**

Amino acid sequences of the cytoplasmic tails of the homologues of HLA-DO $\alpha$  and  $\beta$  from 24 mammals are shown. The ubiquitination site at the residue corresponding to human DO $\beta$  lysine 225 is conserved across most mammalian species. The C-terminal parts of the putative transmembrane domains are highlighted in grey, lysine residues are shown in red, and Yxx $\Phi$  and dileucine motifs are in bold print. Sequences were obtained from the Ensemble database, Release 62. All species for which both  $\alpha$  and  $\beta$  chains were available are shown. In orangutan, mouse lemur, horse, dog and elephant, the DO $\beta$  STOP codon was not found in the database (...).

Species	Species name	DO $\alpha$	DO $\beta$
Human	<i>Homo sapiens</i>	...LVGTVLIIIMGTIVSSVPR	...LVGIVIQLR <b>AQ</b> K <b>GYVRT</b> QMSGNEVSRAV <b>LL</b> PQSC
Chimpanzee	<i>Pan troglodytes</i>	...LVGTVLIIIVGTIVSSVPR	...LVGIVIQLR <b>AQ</b> K <b>GYVRT</b> QMSGNEVSRAV <b>LL</b> PQSC
Orang Utan	<i>Pongo pygmaeus</i>	...LVGTVLIIIMGTIVSSARR	...LVGIVIQLR <b>AQ</b> K <b>GYVRT</b> QMSGNEVSRAV <b>LL</b> PQ [...]
Macaque	<i>Macaca mulatta</i>	...LVGTILIIIMGTIVSSAPR	...LMGIIIQLR <b>AQ</b> K <b>GYVRT</b> QMSGDEVSRAV <b>LL</b> PQPC
Marmoset	<i>Callithrix jacchus</i>	...LVGTILIIIMGICVSSAPR	...LVGIVIQLR <b>AQ</b> KGHVGTMSGDEVSRAV <b>LL</b> PQSY
Galago	<i>Otolemur garnettii</i>	...LVGTILII <b>K</b> GT <b>CQ</b> SSAPR	...LVGIVIH <b>LR</b> AQ <b>KGYMEIQ</b> KSGDEVSRAV <b>LF</b> PQPH
Mouse Lemur	<i>Microcebus murinus</i>	...LVGTALVITGTRLSSAPGQ	...LLLVGVL <b>LR</b> RAQR [...]
Tarsier	<i>Tarsius syrichta</i>	...LVGTILIIIRGT <b>C</b> ESHAPR	...LVGIVIH <b>LR</b> AR <b>KGYMDT</b> Q <b>Q</b> SGDEVPRTVMPWQPC
Large Flying Fox	<i>Pteropus vampyrus</i>	...LVGTVLIIITSTCLHSAPR	...LVGI <b>AV</b> HLRAQ <b>KGYAET</b> QLSGD <b>KV</b> SR <b>SV</b> LPPLTY
Pika	<i>Ochotona princeps</i>	...LQGTVLII <b>I</b> ATRMSRAPRQ	...LVGII <b>TY</b> LR <b>AW</b> KGCMETQ <b>RS</b> AAEVSRAGF <b>PS</b> QPY
Mouse	<i>Mus musculus</i>	...LLGTVLMITGTRRPSIRR	...LVGVV <b>IH</b> L <b>KAQ</b> KASVETQ-PGNEASRESLHSQP.
Rat	<i>Rattus norvegicus</i>	...LLGTLLMVTATRMPSTRR	...LVG <b>AV</b> IH <b>LKAQ</b> KASVETQ-PGNEAS <b>RALL</b> PPHPY
Kangaroo Rat	<i>Dipodomys ordii</i>	...LMGTILIIILATTMADAHR	...LAG <b>LI</b> IHL <b>RAW</b> KGS <b>AE</b> AQ-AGHEVSR <b>ALL</b> PSQPY
Ground Squirrel	<i>Spermophilus tridecemlineatus</i>	...LMGTILIIITGTCMSSCPR	...LVGII <b>IHL</b> RAR <b>KGS</b> VETQ-SGD <b>KISRAIL</b> Q <b>K</b> PN <b>S</b>
Horse	<i>Equus caballus</i>	...LVGTILLIIITSTCLSGAPR	...LVGIF <b>IHL</b> RAR <b>KG</b> [...]
Pig	<i>Sus scrofa</i>	...LVGVTLIIITGRRRSSSPR	...LVGII <b>I</b> HVRAR <b>KGR</b> VETQLSGDEPH
Alpaca	<i>Vicugna pacos</i>	...LVGIVLIIITGSRLSRAPR	...LVGITVHVRAR <b>KGYVET</b> QLSGD <b>KVL</b> GAVCP <b>EQ</b> PH
Bottlenose Dolphin	<i>Tursiops truncatus</i>	...LVGVILIIITGTCLSSTPR	...LVGIVIHTRAR <b>KM</b> -----VTVPRAVLR <b>PQ</b> PH
Dog	<i>Canis lupus</i>	...LVGTIFIIIRGTCLSSGPRYRGPQ	...LVGTVIC <b>LR</b> AQ <b>KGYVET</b> Q <b>F</b> SGDEVM [...]
Cat	<i>Felis catus</i>	...LVGFLVGTILIIIRGTCLSSAPR	...LVGTVIC <b>LR</b> AQ <b>KGYMET</b> QLSGDEVSRAV-P <b>SP</b> PH
Nine-Banded Armadillo	<i>Dasypus novemcinctus</i>	...LAGLVVGTILIFAGTRLSSAPR	...LVGII <b>I</b> CLRAR <b>KGYMET</b> QLSN <b>DG</b> VSRA <b>LL</b> PPQLYQ
African Elephant	<i>Loxodonta africana</i>	...LVGTIFIIISGTCLSSTPRCRGPR	...LVGIVIHFRG <b>QK</b> [...]
Hyrax	<i>Procavia capensis</i>	...LVGTIFIIISGACLS <b>SDPK</b>	...LVGIF <b>IY</b> FRAQ <b>KGYVKT</b> QL <b>SV</b> D <b>KV</b> SR <b>SV</b> LP <b>PQ</b> PYE
Lesser Hedgehog Tenrec	<i>Echinops telfairi</i>	...LLGTVLIIIHSTCCSSVLR	...LVGIC <b>IRL</b> L <b>KAQ</b> RGHVET <b>PQ</b> LHDEGSRSV-PI <b>Q</b> SC