<html><head><style type="text/css"></style></head><body>CLUSTAL 2.1 multiple sequence alignment

----MTARGLALG-LLLLLLCPAQVFS-QSCVWYGECG---IAYGDKRYNCEYSGPPK 49 humanNPC1 ----MGAHHPALGLLLLLLCPAQVFS-QSCVWYGECG---IATGDKRYNCKYSGPPK 50 mouseNPC1 zebrafishNPC1 ----MLLLGRNHIFRLLLATVLLSHWVHG-QHCIWYGECGNS-PNVPEKKLNCNYTGPAV 54 drosophilaNPC1 MSPRSPLRISPFGVHILIAAVLFTLIQSSKQDCVWYGVCN---TNDFSHSQNCPYNGTAK 57 yeastNPC1 -----MNVLWIIALVGQLMRLVQGTATCAMYGNCGK--KSVFGNELPCPVPRSFE 48 celegansNPC1 -----MKQLLIFCLLFGSIFHHGDAGCIMRGLCQKHTENAYGPCVTNDTNVEPT 49 humanNPC1 P--LPKDGYDLVQELCPGFFFGN-VSLCCDVRQLQTLKDNLQLPLQFLSRCPSCFYNLLN 106 mouseNPC1 P--LPKDGYDLVQELCPGLFFDN-VSLCCDIQQLQTLKSNLQLPLQFLSRCPSCFYNLMT 107 zebrafishNPC1 P--LPNEGOELLOELCPRLVYAD-NRVCCDTOQLNTLKSNIQIPLQYLSRCPACFFNFMT 111 drosophilaNPC1 E--MATDGLELLKKRCGFLLENSENKFCCDKNQVELLNKNVELAGNILDRCPSCMENLVR 115 yeastNPC1 PPVLSDETSKLLVEVCG-EEWKEVRYACCTKDQVVALRDNLQKAQPLISSCPACLKNFNN 107 celegansNPC1 AFDKTHPAYEKMVEFCPHLLTGD-NKLCCTPSQAEGLTKQIAQARHILGRCPSCFDNFAK 108 :. **:*: *: * .:: . LFCELTCSPRQSQFLNVTATEDY-----VDPVTNQTKTNVKELQYYVGQSFANAMYNACR 161 humanNPC1 mouseNPC1 LFCELTCSPHQSQFLNVTATEDY----FDPKTPENKTNVKELEYYVGQSFANAMYNACR 162 LFCELTCSPRQSQFISVK-----DFLKEKNLTSVGNVTYYITQTFADAMFNACR 160 zebrafishNPC1 drosophilaNPC1 HICQFTCSPKQAEFMHVVATQ-----KNKKGDEYISSVDLHISTEYINKTYKSCS 165 yeastNPC1 LFCHFTCAADQGRFVNITKVE-----KSKEDKDIVAELDVFMNSSWASEFYDSCK 157 celegansNPC1 LWCEFTCSPNQQDFVSISEMKPIEKKEGFTPEYQPAEAYVNTVEYRLSTDFAEGMFSSCK 168 *.:**:. * *:: humanNPC1 DVEAPSSNDKALGLLCGK-DADACNATNWIEYMFNKD--NGQAPFTITPVFSDFPVRG-- 216 DVEAPSSNEKALGLLCGR-DARACNATNWIEYMFNKD--NGQAPFTIIPVFSDLSILG-- 217 mouseNPC1 zebrafishNPC1 DVQAPSSNIKALGLLCGR-DASVCTPQIWIQYMFSIS--NGQVPFGIEPIFTDVPVQG-- 215 drosophilaNPC1 QVSVPQTGQLAFDLMCGAYSASRCNPTKWFNFMGDAT--NPYVPFQITYIQHEPKSNSNN 223 yeastNPC1 NIKFSATNGYAMDLIGGG----AKNYSQFLKFLGDAKPMLGGSPFQINYKYDLANEEKE- 212 celegansNPC1 DVTFG--GQPALRVMCTS---TPCTLTNWLEFIGTQN-LDLNIPIHTKFLLYDPIKTPPS 222 . *: :: ::::: humanNPC1 -- MEPMNNATKGCDESVDEVTAPCSCQDCS-IVCGPKPQPPPPPAPWTILGLDAMYVIMW 273 --MEPMRNATKGCNESVDEVTGPCSCQDCS-IVCGPKPQPPPPPMPWRIWGLDAMYVIMW 274 mouseNPC1 zebrafishNPC1 --MTPMNNRTFNCSQSLDDGSEPCSCQDCS-EVCGPTPVPPPIPPPWIILGLDAMSFIMW 272 drosophilaNPC1 --FTPLNVTTVPCNQAVSSKLPACSCSDCD-LSCPQGPPEPPRPEPFKIVGLDAYFVIMA 280 yeastNPC1 --WQEFNDEVYACDDAQYK----CACSDCQ-ESCPHLKPLKDGVCKVGPLPCFSLSVLIF 265 celegansNPC1 DRSTYMNVNFTGCDKSARVGWPACSTSECNKEEYANLIDLDDGKTSGQTCNVHGIACLNI 282 *: .:*. humanNPC1 ITYMAFLLVFFGAFFAVWCYRKRYFVSEYTP-IDSNIAFSVNASDK---GEASCCDPVSA 329 mouseNPC1 VTYVAFLFVFFGALLAVWCHRRRYFVSEYTP-IDSNIAFSVNSSDK---GEASCCDPLGA 330 zebrafishNPC1 CSYIAFLLIFFGVVLGAWCYRRSVVTXEYGPILDSNQPHSLNSDGTDLIDEASCCETVGE 332 drosophilaNPC1 AVFLVGVLVFL---MGSFLFTQGSSMDDNFQVDGNDVSDEMPYSEN----DSYFEKLGA 332 YTICALFAFMWYYLCKRKKNGAMIVDDDIVPESGSLDESETNVFES----FNNETN 317 yeastNPC1 FVMLAFIGSLAVLLCVGFVFTS--YDEDYTNLRQTQSGEESPKRNR-----IKRTGA 332 celegansNPC1 AFEGCLRRLFTRWGSFCVRNPGCVIFFSLVFITACSSG-LVFVRVTTNPVDLWSAPSSQA 388 humanNPC1 AFDDCLRRMFTKWGAFCVRNPTCIIFFSLAFITVCSSG-LVFVQVTTNPVELWSAPHSQA 389 mouseNPC1 zebrafishNPC1 RFENSLRLVFSRWGSLCVROPLTIILSSLVLICICSAG-LSYMRITTNPVELWSAPSSRA 391 drosophilaNPC1 HTETFLETFFTKWGTYFASNPGLTLIAGASLVVILGYG-INFIEITTDPVKLWASPNSKS 391 yeastNPC1 FFNGKLANLFTKVGQFSVENPYKILITTVFSIFVFSFIIFQYATLETDPINLWVSKNSEK 377 celegansNPC1 WIHNFMENNARDIGMMAGRNPKSHFFIGCAVLIFCLPG-MIYHKESTNVVDMWSSPRSRA 391

humanNPC1	RLEKEYFDQHFGPFFRTEQLIIRAPLTDKHIYQPYPSGADVPFGPPLDIQILHQVLDLQI	448
mouseNPC1		449
zebrafishNPC1	RQEKNCFDQHFGPFFRTEQLIITTPWTEEGGFS-TITGDIIPFSPILNLSLLHQVLDLQL	450
drosophilaNPC1	RLEREFFDTKFSPFYRLEQIIIKAVNLPQIVHNTSNGPYTFGPVFDREFLTKVLDLQE	449
yeastNPC1	FKEKEYFDDNFGPFYRTEQIFVVNETGPVLSYETLHWWFDVEN	420
celegansNPC1	RQEEMVFNANFGRPQRYQQIMLLSHRDFQSSGKLYGPVFHKDIFEELFDILN	443
	*. *: :*. * :*::	
humanNPC1	AIENITASYD-NETVTLQDICLAPLSPYNTNCTILSVLNYFQNSHSVLDHKKGDD	502
mouseNPC1	AIESITASYN-NETVTLQDICVAPLSPYNKNCTIMSVLNYFQNSHAVLDSQVGDD	503
zebrafishNPC1	EIENLIAEYK-GENVTLKDICVSPLSPYNDNCTILSVLNYFQNSHEVLDHEFQDE	504
drosophilaNPC1	GIKEINANGTQLKDICYAPLSDDGSEIDVSQCVVQSIWGYFGDDRERLDDHDEDN	504
yeastNPC1	FITEELQSSENIGYQDLCFRPTEDSTCVIESFTQYFQGALPNKDS	465
celegansNPC1	AIKNISTQDSDGRTITLDDVCYRPMGPGYDCLIMSPTNYFQGNKEHLDMKSNKE *	497
	*	
humanNPC1	FFVYADYHTHFLYCVRAPASLNDTSLLHDPCLGTFGGPVFPWLVL	547
mouseNPC1	FYIYADYHTHFLYCVRAPASLNDTSLLHGPCLGTFGGPVFPWLVL	548
zebrafishNPC1	FFLYNDYHTHLLYCASSPTSLDDTSRLHDPCMGTFGGPVFPWLVL	549
drosophilaNPC1	GFNLCLAPYGGPVDPAIAL	
yeastNPC1	CLPTFQQPLKTNLLF	494
celegansNPC1	ETVSEDDDAFDYFSSEATTDEWMNHMAACIDQPMSQKTKSGLSCMGTYGGPSAPNMVF : * * * * .: * .: :	555
	: : * * * * : : * : :	
humanNPC1	GGYDDQNYNNATALVITFPVNNYYNDTEKLQRAQAWEKEFINFVKNYKNPNL	599
mouseNPC1	GGYDDQNYNNATALVITFPVNNYYNDTERLQRAWAWEKEFISFVKNYKNPNL	600
zebrafishNPC1	GGYEDSAYNNATALFFTFPVTNCLNDTEKLGKALAWEKEFIRFMKNYENPNL	601
drosophilaNPC1	GGFLPPGDQLTGSTKFELANAIILTFLVKNHHNKTD-LENALTWEKKFVEFMTNYTKNNM	
yeastNPC1		
celegansNPC1	GKNSTNHQAANSIMMTILVTQRTEPEIQKAELWEKEFLKFCKEYREKSP * ::*: : **::: : : : : : : : : : : : :	604
	. * ::*: : . **:.: :	
humanNPC1	TISFTAERSIEDELNRESDSDVFTVVISYAIMFLYISLALGHIKSCRRLLVD	651
mouseNPC1	TISFTAERSIEDELNRESNSDVFTVIISYVVMFLYISLALGHIQSCSRLLVD	652
zebrafishNPC1	TVSFSSERSIEDEIDRESNSDVSTIVISYIIMFVYISVALGRINSCRTLLVD	653
drosophilaNPC1	SQYMDIAFTSERSIEDELTRESQSDVLTILVSYLIMFMYIAISLGHVKEFKRVFID	653
yeastNPC1 celegansNPC1	LRISFNTEISLEKELNNNNDISTVAISYLMMFLYATWALRRKDGK	582 662
CelegalishPC1	KVIFSFMAERSITDEIENDAKDEIVTVVIALAFLIGYVTFSLGRYFVCENQLWSILVH .:* :* *: .*: .:: *: :: *: :*	002
humanNPC1	${\tt SKVSLGIAGILIVLSSVACSLGVFSYIGLPLTLIVIEVIPFLVLAVGVDNIFILVQAYQR}$	711
mouseNPC1	SKISLGIAGILIVLSSVACSLGIFSYMGMPLTLIVIEVIPFLVLAVGVDNIFILVQTYQR	712
zebrafishNPC1	·	713
drosophilaNPC1	SKITLGIGGVIIVLASVVSSVGVFGYIGLPATLIIVEVIPFLVLAVGVDNIFILVQTHQR	713
yeastNPC1 celegansNPC1	TRLLLGISGLLIVLASIVCAAGFLTLFGLKSTLIIAEVIPFLILAIGIDNIFLITHEYDR SRICLGMLSVIINLLSSFCSWGIFSMFGIHPVKNALVVQFFVVTLLGVCRTFMVVKYYAQ	042 722
celegalishrci	::: **: .::* * * .: *:: : * * *:: : : :	122
humanNPC1	DERLQGETLDQQLGRVLGEVAPSMFLSSFSETVAFFLGALSVMPAVHTFSLFAGLA	
mouseNPC1	DERLQEETLDQQLGRILGEVAPTMFLSSFSETSAFFFGALSSMPAVHTFSLFAGMA	768
zebrafishNPC1	DERMPEEELHQQIGRILGDVAPSMFLSSFSETVAFFLGALSTMPAVRTFSLFAGLA	769
drosophilaNPC1	DQRKPNETLEQQVGRILGKVGPSMLLTSLSESFCFFLGGLSDMPAVRAFALYAGVA	769
yeastNPC1	NCEQKPEYSIDQKIISAIGRMSPSILMSLLCQTGCFLIAAFVTMPAVHNFAIYSTVS ORVSMPYMSPDOCPEIVGMVMAGTMPAMFSSSLGCAFSFFIGGFTDLPAIRTFCLYAGLA	699 782
celegansNPC1	QRVSMPYMSPDQCPEIVGMVMAGTMPAMFSSSLGCAFSFFIGGFTDLPAIRTFCLYAGLA : : : *::::::::::::::::::::::::::::::	102
humanNPC1	VFIDFLLQITCFVSLLGLDIKRQEKNRLDIFCCVRGAEDGTSVQASESCLFRFFK	
mouseNPC1	VLIDFLLQITCFVSLLGLDIKRQEKNHLDILCCVRGADDGQGSHASESYLFRFFK	
zebrafishNPC1	IFIDFLLQISCFVSLLGLDIKRQEANRMDILCCVKLS-DGQE-EKSEGWLFRFFK	
drosophilaNPC1	LIIDFLLQITCFVSLFTLDTKRREENRMDICCFIKGK-KPDSITSNEGLLYKFFS	
yeastNPC1	VIFNGVLQLTAYVSILSLYEKRSNYKQITGNEETKESFLK	139

celegansNPC1	VLIDVVLHCTIFLALFVWDTQRELNGKPEFFFPYQIKDLLGAYLIGRQRATDTFMTQFFH	842
	:::: :*: : : : : : : : : : : : : : : :	
humanNPC1	NSYSPLLLKDWMRPIVIAIFVGVLSFSIAVLNKVDVGLDQSLSMPDDSYMVDYFKSISQY	882
mouseNPC1	NYFAPLLLKDWLRPIVVAVFVGVLSFSVAVVNKVDIGLDQSLSMPNDSYVIANFKSLAQY	883
zebrafishNPC1	KIYAPFILKDWVRPLVVAVFVGMLSFSIAVVNKVEIGLEQTLSMPDDSYVLNYFGNLSKY	
drosophilaNPC1	SVYVPFLMKKIVRASVMVIFFAWLCFSIAIAPRIDIGLDOELAMPODSFVLHYFOSLNEN	883
yeastNPC1	TFYFKMLTQKRLIIIIFSAWFFTSLVFLPEIQFGLDQTLAVPQDSYLVDYFKDVYSF	796
celegansNPC1	FQVAPFLMHRMTRIITGIIFIASFITTVILSSKISVGFDQSMAFTEKSYISTHFRYLDKF	902
CelegalishPC1		902
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h		0.4.2
humanNPC1	LHAGPPVYFVLEEGHDYTSSKGQNMVCGGMGCNNDSLVQQIFNAAQLDNYTRIGFAPSSW	
mouseNPC1	LHSGPPVYFVLEEGYNYSSRKGQNMVCGGMGCDNDSLVQQIFNAAELDTYTRVGFAPSSW	
zebrafishNPC1	LHTGPPVYFVVEDGHDYKTFEGQNAVCGGVGCNNDSLVQQIYTASLMSNYTRISNVPSSW	
drosophilaNPC1	LNIGPPVYFVLKGDLAYTNSSDQNLVCAGQYCNDDSVLTQIYLASRHSNQTYIARPASSW	
yeastNPC1	LNVGPPVYMVVKN-LDLTKRQNQQKICGKFTTCERDSLANVLEQERHRSTITEPLANW	
celegansNPC1	FDVGPPVFFTVDGELDWHRPDVQNKFCTFPGCSDTSFGNIMNYAVGHTEQTYLSGEMYNW	962
	:. ****:::: . *: . *: . *: . *	
humanNPC1	IDDYFDWVKPQSS-CCRVDNITDQFCNASVVDPACVRCRPLTPEGKQR	989
mouseNPC1	IDDYFDWVSPQSS-CCRLYNVTHQFCNASVMDPTCVRCRPLTPEGKQR	990
zebrafishNPC1	LDDYFDWVKPQST-CCRYYNSTGAFCNASVVDKSCVHCRPMTSSGKQR	989
drosophilaNPC1	IDDYFDWAAAASS-CCKYRKDSGDFCPHQDTSCLRCN-ITKNSLLR	987
yeastNPC1	LDDYFMFLNPQNDQCCRLKKGTDEVCPPSFPSRRCETCFQQGSWNYNMSGF	904
celegansNPC1	IDNYLEWISRKSPCCKVYVHDPNTFCSTNRNKSALDDKACRTCMDFDYVANSYPKSSIMY	1022
	:*:*:	
humanNPC1	PQGGDFMRFLPMFLSDNPNPKCGKGGHAAYSSAVNILLG-HGTRVGATYFMTYHTVLQ	1046
mouseNPC1	PQGKEFMKFLPMFLSDNPNPKCGKGGHAAYGSAVNIVGDDTYIGATYFMTYHTILK	
zebrafishNPC1	PNGTEFMHFLPMFLSDNPNIKCGKGGHAAYGTAVDLKDNNTDVGATYFMSYHTILK	
drosophilaNPC1	PEEKEFVKYLPFFLKDNPDDTCAKAGHAAYGGAVRYSNSHERLNIEASYFMAYHTILK	
yeastNPC1	PEGKDFMEYLSIWIN-APSDPCPLGGRAPYSTALVYNETSVSASVFRTAHHPLR	
celegansNPC1	HRPSIEVFYRHLRHFLEDTPNSECVFGGRASFKDAISFTSRGRIOASOFMTFHKKLS	
ceregalishrer	*. ** ::. *. * .*:*: *:	1019
humanNPC1	TSADFIDALKKARLIASNVTETMGINGSAYRVFPYSVFYVFYEOYLTI	100/
mouseNPC1	TSADYTDAMKKARLIASNITETMRSKGSDYRVFPYSVFYVFYEQYLTI	
mouseNPC1 zebrafishNPC1		
	NSSDFINAMKMARELTDNITQTLSTHDKSYKVFPYSVFYVFYEQYLTI	
drosophilaNPC1	SSADYFLALESARKISANITQMLQGRLMSNGVPMASALTVEVFPYSVFYVFYEQYLTM	
yeastNPC1	SQKDFIQAYSDGVRISSSFPELDMFAYSPFYIFFVQYQTL	
celegansNPC1	ISNSSDFIKAMDTARMVSRRLERSIDDTAHVFAYSKIFPFYEQYSTI	1126
	. *: * :: *: ** *: *: *: *: *: *: *: *: *: *:	
1 275.61		1154
humanNPC1	IDDTIFNLGVSLGAIFLVTMVLLGCELWSAVIMCATIAMVLVNMFGVMWLWGISLNAVSL	
mouseNPC1	IDDTIFNLSVSLGSIFLVTLVVLGCELWSAVIMCITIAMILVNMFGVMWLWGISLNAVSL	
zebrafishNPC1	VDDTALNLGVSLSAIFIVTAVLLGFELWSAVLVCFTIAMILINMFGVMWLWSISLNAVSL	
drosophilaNPC1	WSDTLQSMGISVLSIFVVTFVLMGFDVHSALVVVITITMIVVNLGGLMYYWNISLNAVSL	1163
${\tt yeastNPC1}$	GPLTLKLIGSAIILIFFISSVFL-QNIRSSFLLALVVTMIIVDIGALMALLGISLNAVSL	1056
celegansNPC1	MPILTTQLFITVVGVFGIICVTLGIDVKGAACAVICQVSNYFHIVAFMYIFNIPVNALSA	1186
	: :: :* : * : ::	
humanNPC1	VNLVMSCGISVEFCSHITRAFTVSMKGSRVERAEEALAHMGSSVFSGITLTKFGGIV	1211
mouseNPC1	VNLVMSCGISVEFCSHITRAFTMSTKGSRVSRAEEALAHMGSSVFSGITLTKFGGIV	1211
zebrafishNPC1	VNLVMSCGISVEFCSHIVRAFSISTRSSRVERAEEALAHMGSSVFSGITLTKFGGIL	1210
drosophilaNPC1	VNLVMAVGISVEFCSHLVHSFATSKSVSQIDRAADSLSKMGSSIFSGITLTKFAGIL	1220
yeastNPC1	~	1116
celegansNPC1	TNLVMSSGILIEFSVNVLKGYACSLRQRAKDRAESTVGSIGPIILSGPVVTMAGSTM	
· - = - 9 14	.**:. *: :**. :: :::	
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humanNPC1	VLAFAKSQIFQIFYFRMYLAMVLLGATHGLIFLPVLLSYIGPSVNKAKSCATEERYKGTE	1271
mouseNPC1	VLAFAKSQIFEIFYFRMYLAMVLLGATHGLIFLPVLLSYIGPSVNKAKRHTTYERYRGTE	
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zebrafishNPC1 drosophilaNPC1 yeastNPC1 celegansNPC1	ILALSKSQIFQIFYFRMYLAIVLLGAAHGLIFLPVLLSYAGKSLSPRTFMICLEAVVHLK VLAFAKSQIFQVFYFRMYLGIVVIGAAHGLIFLPVLLSYIGAPVSNARLRYHSQAAAEHE VLAFAQSKIFDVFYFRMWFTLIIVAALHALLFLPALLSLFGGESYRDDSIEAED FLSGAHLQIITVYFFKLFLITIVSSAVHALIILPILLAFGGSRGHGSSETSTNDNDEQHD .*: :: :*: :: :: :: :: :: :: :: :: :: ::	1270 1280 1170 1303
humanNPC1 mouseNPC1 zebrafishNPC1 drosophilaNPC1 yeastNPC1 celegansNPC1	RERLLNF	1278 1278 1277 1287 1363
humanNPC1 mouseNPC1 zebrafishNPC1 drosophilaNPC1 yeastNPC1 celegansNPC1	SLPCTSRMPRAHIEPDLRSL 1383	

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