



UNIVERSIDAD DE BARCELONA
FACULTAD DE FARMACIA
DEPARTAMENTO DE BIOQUÍMICA Y BIOLOGÍA MOLECULAR

**ESTUDIO DE LOS MECANISMOS DE INHIBICIÓN
DE LA ACTIVIDAD CARNITINA
PALMITOILTRANSFERASA 1**

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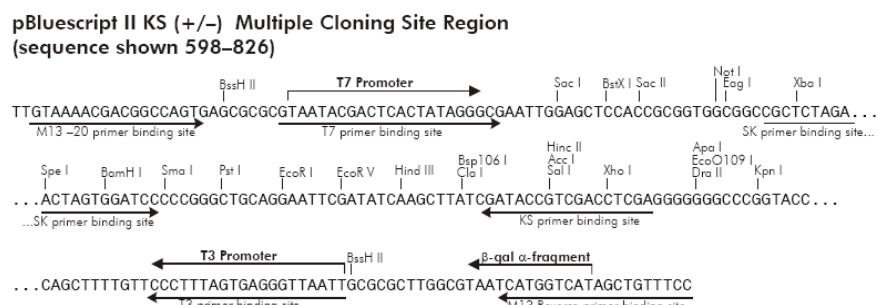
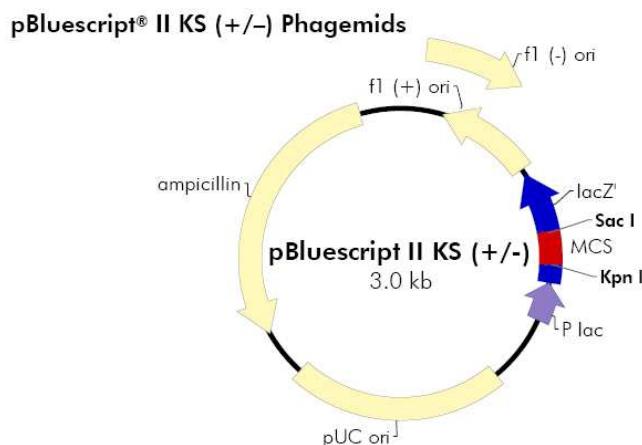
Z

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APÉNDICES

1. MAPA DE RESTRICCIÓN DE LOS VECTORES PLASMÍDICOS

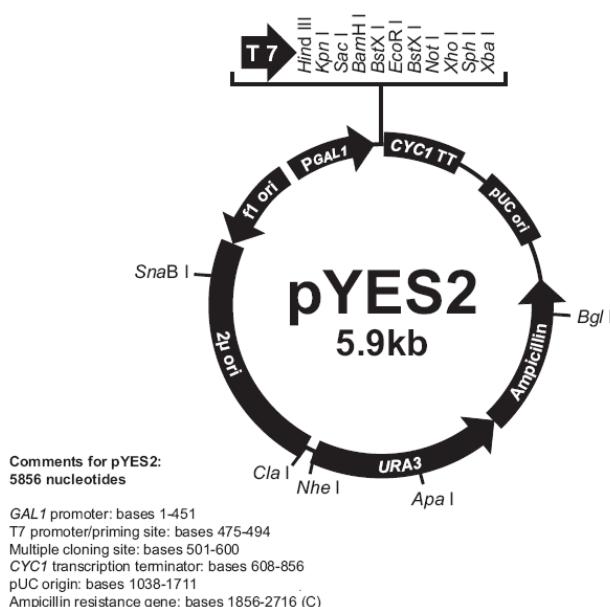
- ❖ Vector *Bluescript* (*pBS*) (Stratagene, ref. 212205) para subclonaje



- ❖ Vector *pYES 2.0* (Invitrogen, ref. V825-20) de expresión en *S. cerevisiae*

Map of pYES2

The figure below summarizes the features of the pYES2 vector. The sequence for pYES2 is available for downloading from our Web site (www.invitrogen.com) or from Technical Service (see page 17).



2. OLIGONUCLEÓTIDOS

OLIGONUCLEÓTIDOS CPT1A	SECUENCIA (5' → 3')
CPT1A.C608A.for	5' GAG ACT GTA CGC TCC <u>GCC</u> ACT TAT GGA GTC CTG C 3'
CPT1A.C608A.rev	5' G CAG GAC TCC ATA AGT GGC GGA GCG TAC AGT CTC 3'
CPT1A.R243T.for	5' GAA TAT ATC TAC CTG <u>ACG</u> GGC CGA GGG CCG CTC 3'
CPT1A.R243T.rev	5' GAG CGG CCC TCG GCC CGT CAG GTA GAT ATA TTC 3'
CPT1A.W682A.for	5' GAG GTA TTG TCT GAG CCA <u>GCG</u> AGG TTG TCT ACG AGC 3'
CPT1A.W682A.rev	5' GCT CGT AGA CAA CCT CGC TGG CTC AGA CAA TAC CTC 3'
CPT1A.A478G.for	5' CAC TCC TGG GCG GAG <u>GGG</u> CCC ATC GTG GGC CAT TTG 3'
CPT1A.A478G.rev	5' CAA ATG GCC CAC GAT GGG CCC CTC CGC CCA GGA GTG3'
CPT1A.M593S.for	5' CTC ACA TAT GAG GCC TCC <u>AGT</u> ACC CGG CTC TTC CGA GAA GG 3'
CPT1A.M593S.rev	5' CC TTC TCG GAA GAG CCG GGT CCT GGA GGC CAC ATA TGT GAG 3'
CPT1A.M593R.for	5' CTC ACA TAT GAG GCC TCC <u>AGG</u> ACC CGG CTC TTC CGA GAA GG 3'
CPT1A.M593R.rev	5' CC TTC TCG GAA GAG CCG GGT CCT GGA GGC CAC ATA TGT GAG 3'
CPT1A.M593K.for	5' CTC ACA TAT GAG GCC TCC <u>AAG</u> ACC CGG CTC TTC CGA GAA GG 3'
CPT1A.M593K.rev	5' CC TTC TCG GAA GAG CCG GGT CCT GGA GGC CAC ATA TGT GAG 3'

OLIGONUCLEÓTIDOS CPT1B	SECUENCIA (5' → 3')
CPT1.HindIII.ATG.for	5' TCG ATA AGC TTA TAA AAT GGC GGA AGC ACA CCA GGC AG 3'
CPT1B.HindIII.rev	5' GGG ACA GGA AGC TTG GGC 3'
CPT1B.AAAA.for	5' CGA CAG GCA TCT TCC TCT TCC GAC AAA C 3'
CPT1B.AAAA.rev	5' GTT TGA CGG AAG AGG AAG ATG CCT GTC G 3'
CPT1B.MseI.rev	5' CTT GTT GGC TCG TGT TCT TAA TAA GC 3'
CPT1B.H473A.for	5' CAG CTG GGC CTC AAC ACA GAA <u>GCC</u> TCA TGG GCA GAT GCT CCC 3'
CPT1B.H473A.rev	5' GGG AGC ATC TGC CCA TGA GGC TTC TGT GTT GAG GCC CAG CTG 3'
CPT1B.H477A.for	5' AAC ACA GAA CAC TCA TGG GCA <u>GCT</u> GCT CCC ATC ATC GGT CAC CTC 3'
CPT1B.H477A.rev	5' GAGGTGACCGATGATGGGAGCAGCTGCCATGAGTGTCTGTGTT 3'
CPT1B.R655N.for	5' GCC ATG ACA GGG GCT GGG ATC GAC <u>AAC</u> CAC CTC TTC TGC CTC 3'
CPT1B.R655N.rev	5' GAG GCA GAA GAG GTG GTT GTC GAT CCC AGC CCC TGT CAT GGC 3'
CPT1B.TET602-604VDN.for	5' ATG TTC CGA GAG GGG CGG <u>GTA GAC AAT</u> GTG CGT TCC TGT ACT AGC 3'
CPT1B.TET602-604VDN.rev	5' GCT AGT ACA GGA ACG CAC ATT GTC TAC CCG CCC CTC TCG GAA CAT 3'
CPT1B.H279A.for	5' G GGA AAC ACC GTT <u>GCC</u> GCC ATG AT 3'
CPT1B.H279A.rev	5' AT CAT GGC GGC AAC GGT GTT TCC C 3'
CPT1B.H483A.for	5' GCT CCC ATC ATC GGT <u>GCC</u> CTC TGG GAG TTC G 3'
CPT1B.H483A.rev	5' C GAA CTC CCA GAG GGC ACC GAT GAT GGG AGC 3'
CPT1B.M593S.for	5' CTG ACC TAT GAG GCC TCC <u>AGC</u> ACA AGA ATG TTC CGA GAG 3'
CPT1B.M593S.rev	5' CTC TCG GAA CAT TCT TGT GCT GGA GGC CTC ATA GGT CAG 3'
CPT1B.COT GGG709-711QGV.for	5' AATCATCTGGGTGCT <u>CAAGGTGTCTTGGCCTGTGGCC</u> 3'
CPT1B.COT GGG709-711QGV.rev	5' GGCCACAGGACCAAAAGACACCTTGAGCACCCAGATGATT 3'
CPT1B.CrAT GGG709-711VMF.for	5' AATCATCTGGGTGCT <u>GTAATGTTCTTGGCCTGTGGCC</u> 3'
CPT1B.CrAT GGG709-711VMF.rev	5' GGCCACAGGACCAAAAGAACATTACAGCACCCAGATGATT 3'

Los nucleótidos marcados en negrita corresponden a la secuencia para una eficiente traducción de las proteínas en levaduras.

Los nucleótidos subrayados son los mutados con el fin de cambiar un amino ácido original por otro de interés (tal y como se indica en el nombre del oligonucleótido). Los otros oligonucleótidos, se utilizaron para la secuenciación del cDNA.

3. SECUENCIAS

3.1 Secuencia de CPT1A de rata

Secuencia de la carnitina palmitoiltransferasa I de hígado de rata (CPT1A), publicada por Esser (Esser, 1993), con número de acceso en GeneBank Data Libraries L07736.

DEFINICIÓN: mRNA de carnitina palmitoiltransferasa I, cDNAs completos.

NÚMERO DE ACCESO: L07736

FUENTE: cDNA correspondientes a mRNA de hígado de macho adulto de *Rattus norvegicus*.

AUTORES: Esser,V., Britton,C.H., Weis,B.C., Foster,D.W. and McGarry,J.D.

TÍTULO: Cloning, sequencing and expression of a cDNA encoding rat liver carnitine palmitoyltransferase I. Direct evidence that a single polypeptide is involved in inhibitor interaction and catalytic function.

PUBLICACIÓN: *J. Biol. Chem.* **268**, 5817-5822 (1993).

ZONA CODIFICANTE: 103 – 2424 (773 aminoácidos).

"MAEAHQAVAFQFTVTPDGIDLRLSHEALKQICLSQLHSWKKFIRFKNGIITGVFPA
NPSSWLIVVVGVISSMHAKVDP SLGMIAKISRTLDTGRMSSQTKNIVSGVLFGTGLW
VAVIMTMRYSLKVLLSYHGWMFAEHGKMSRSTKIWMAMVKVLSGRKPMLYSFQTSLPR
LPVPAVKDTVSRYLESVRPLMKEEDFQRMTALAQDFAVNLGPKLQWYLKLKSWWATNY
VSDWWEYIYLGRGRGPLMVNSYYAMEMLYITPTHIQAARAGNTIHAILLYRRTLDRE
ELKPIRLLGSTIPLCSAQWERLFNTSRIPGEETDTIQHIKDSRHIVVYHRGRYFKVWL
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EKHQHLYRLAMTGAGIDRHLFCLYVVSKYLAVDSPFLKEVLSEPWRLLSTSQTPOQQVE
LFDFEKNPDYVSCGGGFGPVADDGYGVSYIIVGENFIHFHISSKFSSPETDSHRFKGKH
LRQAMMDIITLFGLTINSKK"

RECUENTO DE BASES: 1109 a; 1115 c; 1121 g; 1032 t

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61 tgcttgctcc ggggagtgcgca gagcaatagg tccccactca agatggcaga ggctcaccaa
121 gctgtggcct tccagttcac cgtcacccccc gatggcattt acctccgcct gagccacgaa
181 gccctcaaac agatctgcct gtcggggctg cactcctgga agaagaagtt catccggttc
241 aagaatggca tcatacactgg tgtgttcccc gcgaatccgt ccagctggct tatcgtggtg
301 gtgggtgtga tttcatccat gcatgccaaa gtggaccctt ccctggccat gatcgcaaag
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421 ggcgtcctct ttggcacagg gctctgggtg gcagtcatca tgaccatgcg ctactcgctg
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1021 cagtgggagc gactttcaa tacttccgg atccctgggg aggagacaga caccatccaa
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1141 ctctaccacg atgggaggt gctgaggccc cgagagctgg agcagcagat gcagcagatc
1201 ctggatgatc cctcagagcc acagctggg gaggccaagc tggccgcctt cactgctgca
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2701 gaattcctgg attctggggg ttgtttcttc acatgtgtt gaggtgacag acttcctcag
2761 tggtgaccct gtgaatactt gggcgtctga ctccacccag gcagtgtag gagtcaccc
2821 tggaaagaga aagtgtcttc agagccagca gaggcaacag ctgtagctaa cacatctgta
2881 acacactaat ggaatggta ggcctgggg ttaagggttct gctatgagtg acagccactg
2941 tcccttgga agttcacatt tccaggaagc agagtaccac ctccccagtg ccacccctt
3001 cacacatctt caaaaacccag tgccttaaag aagggggcca ttggcaagcg ggaccataga
3061 gaagacttag catctgtgaa gcctttgggt ggatatgtga ggatgtgtct tcccccttact
3121 gtttcctgca taaagatgtc cctaagtaag cacttcccc accccctagaa aatgaggtcc
3181 ttggtaagg cagggatgct ggagtctcat tgcctgccag ttccattaag ccacaaaata
3241 gcagacatgt gtccacagag ggaggggctt ggttagtcaaa ggctgcatag ctggacaaca
3301 gcgggagagt gtggcttgct gtatttgaca gctgtggca agaggagtga gaccctggc
3361 accaagttag acatactgac acaggcagcc aaagctcactc gagccaggag atatagatag
3421 atactggctt gtattctggc agatacaccc ctgggcttat cttctaattcc cacccttgtca
3481 gattccaacc agagtcaaat tccatagaag gctaggtcat tttggcgaca gactcaggga
3541 tctcaagtaa tgggtgcctt tacccatcg ccattccctca gtgggagtgc ctttctgaa
3601 agcatccaaat gggctaaagc agctctacca agtctgttgc gtatttaatg taaacattag
3661 catataatgga gtggctcttc ctacctgtca ccattctgtc ctgacaagct tagctctccg
3721 agtttacat catgtattta tttccagtg ccccttggc cttgtttgat tcctgccc
3781 gtgccagaag tggcccaagaa gtgaggggtg gggtgaccag cagtgcagag aggtgtggc
3841 tgaacagttc atgtgtgtct tatgggtata catgtataaa tttgtaatg taaaaaaaaaa
3901 aatcatacct aaaaggccca aagtttttt ttttttact aaaaccaaga aaacaaaaga
3961 caacataaaag acataaggcag aaacaaaactg ttgtaagtca gagcggcctg actctcgctg
4021 ctgtgaccac tcaccaacct gtgttactca gagtagcccg gctagtgcgg gagtgggaca
4081 ttccctcttc aggtttccag tgtccttgct gctcctgagc agttaccaat gcaatttcgc
4141 attccttaca aggcaagaaga gtgggtcttc actgtatgtt ttcaaaggag gaggttaagac
4201 tattgtgtat ttaatttaat gtggaaacaaa atatagtctt accgcagcca aggttcaaat
4261 ttgggtttct aatctgtcca ttgcattgtaa ataccatatac tgtttggata taaaatcttag
4321 aagtgcattgt gtgagcgaat gtagctggcc attaataaaa catataact gtctact

3.2 Secuencia de CPT1B de rata

Secuencia de la carnitina palmitoiltransferasa I de músculo de rata (CPT1A), publicada por Yamazaki (Yamazaki, 1995), con número de acceso en GeneBank Data Libraries D43623.

DEFINICIÓN: mRNA de carnitina palmitoiltransferasa 1, cDNAs completos.

NÚMERO DE ACCESO: D43623

FUENTE: cDNA correspondientes a mRNA de músculo de macho adulto de Rattus norvegicus.

AUTORES: Yamazaki,N., Shinohara,Y., Shima,A. and Terada,H.

TÍTULO: High expression of a novel carnitine almitoyltransferase I like protein in rat brown adipose tissue and heart: isolation and characterization of its cDNA clone.

PUBLICACIÓN: *FEBS Lett.* **363** (1-2), 41-45 (1995).

ZONA CODIFICANTE: 27 – 2345 (772 aminoácidos).

"MAEAHQAVAFQFTVTPGVDFRLSREALRHIYLSGINSWKKRLIRIKNGILRGVYPGS
PTSWLVVVMATVGSNYCKVDISMGLVHICQRCLPTRYGSYGTQTETLMSMVIFSTGVW
ATGIFLFRQTLKLLSYHGWMFEMHSKTSHATKIWAICVRLLSSRRPMLYSFQTSPLKL
PVPSVPATIHYRLDSVRPLLDEAYFRMESLAKEFQDKIAPRLQKYLVLKSWATNYVS
DWWEELYVYLRGRSPIMVDSNYYAMDFVLIKNTSQQAARLGNTVHAMIMYRRKLDREEIK
PVMALGMVPMCSYQMERMFNTTRIPGKETDLLQHLSESRHVAVYHKGRFFKVWLVEGSC
LLKPRDLEMQFQRILDDTSPPQPGEEKLAALTAGGRVEWAEARQKFFSSGKNKMSLDTI
ERAAFFVALDEDSHCYNPDDEASLSLYGKSLLHGNCYNRWFDKSFTLISCKNGQLGLNT
EHSWADAPIIGHLWEFVLATDTFHLYTETGHCVGEPNTKLPPPQRMQWDIPEQCQTAI
ENSYQVAKALADDVELYCFQFLPGKGLIKKCRTSPDAFVQIALQLAHFRDKGFCLTY
EASMTRMFREGRTEVRSCTSESTAFVRAMMTGSHKEQDLQDLFRKASEKHQNMYRLAM
TGAGIDRHLFCLYIVSKYLGVRSPFLDEVLSEPWSLSTSQIPQFQICMFDPKQYPNHLG
AGGGFGPVADHGYGVSYMIAgentMFFHVSSKLSSSETNALRFGNHIRQALLDIADLFK
ISKTDS"

RECUENTO DE BASES: 656 a; 788 c; 727 g; 642 t

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1 ctgagctgtg ctgactaaac cccaggatgg cggaagcaca ccaggcagta gctttccagt
61 tcactgtgac cccagacggg gtcgacttcc ggcttagtcg ggaggctctg agacacatct
121 acctgtctgg aatcaactcc tggaaagaaaac gccttattcg aatcaagaat ggtatccta
181 ggggtgtgta ccctggcagc cctaccagct ggctggttgt tgtcatggca acagttggtt
241 ccaactactg caaaagtggac atctccatgg ggctggtcca ttgcattccag agatgcctcc
301 cgacaaggta tggctcctac gggaccccac agaccgagac acttctcagt atggcatct
361 tctccaccgg agtctggcg acaggcattt ttttattccg acaaaccctg aagctgctgc
421 tttccttatca tgggtggatg ttcgagatgc acagcaagac cagccatgcc accaagatct
481 gggctatctg tggtcgctc ctgtccagcc ggcggccat gctctatgc ttccaaacat
541 cactgcccaa gcttcctgtc cccagtgtgc cagccacaat tcaccggta ttggattctg
601 tgcggccctt gctggatgac gaagcctatt tccgcatgga gtcgttggcc aaagaattcc
661 aggacaagat tgccccaga ctgcagaaat acctgggtct gaagtcatgg tgggcaacca
721 actatgtaaag tgactggtg gaagagtacg tctacctccg aggcaggagc cccatcatgg
781 tgaacagcaa ctattacgccc atggattttg tgcttattaa gaacacgagc caacaagcag
841 cacgttggg aaacaccgtt caccatgtc tcatgtatcg ccgcaaaactg gaccgagaag
901 agatcaagcc ggtaatggca ctgggtatgg taccatgtc ctcctaccag atggagagga
961 tggtcaacac tacacgcatac ccaggcaaag agacagactt gctacagcac ctctcagaga
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1021 gcaggcacgt ggctgtctac cacaaggc gcttctcaa gtttggtc tatgagggct
1081 cgtgcctgca caagccccga gacctcgaga tgcagttcca gagaatcc tcgacacact
1141 ccccgctca gcctggagag gaaaagctgg cagccctcac cgccaggagga agggtagagt
1201 gggcagaagc acgtcagaag ttcttttagt ctggcaagaa caagatgtcc ctggatacc
1261 tcgaacgtgc tgcttccttt gtggccctgg acgaagactc tcactgttac aaccctgatg
1321 acgaggccag tctcagcc tcacggcaaat ccctgctgca cggcaactgc tataacagg
1381 gttcgacaa atctttcaact ctcatctcct gcaagaatgg ccagctggc ctcaacacag
1441 aacactcatg ggcagatgct cccatcatcg gtcacccctg ggagttcgtc ctggccactg
1501 ataccttca cctgggctac acggagacag gacactgtgt gggtaaccc aacaccaagt
1561 tgccggccgc tcagcggatg cagtggaca ttcccgagca gtgcagaca gccatcgaga
1621 attcgatcca agtagccaag gcctggctg atgatgtgga gttatactgc ttccagttct
1681 tacccttcgg caaaggcctg atcaagaagt gtcggaccag ccctgatgcc tttgtgcaga
1741 ttgccctgca gctggctcat ttccggaca aaggcaagtt ctgcctgacc tatgaggcct
1801 ccatgacaag aatgttccga gaggggcgga cagagactgt gcgttcctgt actagcgagt
1861 ccacggcctt tgtgcggcc atgatgacgg ggtcccataa gaaacaagac ctccaagacc
1921 tcttccggaa agcctccgaa aaacacccaaa acatgtaccg cctagccatg acagggcgt
1981 ggatcgacag gcacctttc tgcctctaca tgcgtccaa gtacttaggg gttagatctc
2041 ctttcctgca cgaggtgctt tcggaaccct ggagccctc caccagccag atccccagt
2101 tccagatctg catgttgac ccaaaggcgt accccaatca tctgggtgct ggaggtggct
2161 ttggtcctgt ggccgaccac ggatacgggg tttcctacat gatgcgaggc gaaaacacaa
2221 tgttcttcca tgttccagc aagttatcga gttcagaaac gaacgcctg cgttcggga
2281 accacatccg tcaagcactg ctggatatcg ccgaccttt caaaatttcc aagactgaca
2341 gctgagacca ggagacacac cagtcgcctt ttggccccca cctgggtggag gaaggggtct
2401 gtggccagtt cacaggcata aggggtggca tgcacacgtg cccagttctg agaccagctc
2461 cagcgcaggg gctccccagg cagacactgc tcctccaggc ccggcagg tgggattgga
2521 gtggtgaggg aactttgate ttttttttc ccccggtctt ggtagatgct aataaaaaata
2581 aggctgtata attctctctc agcccttagg tgcctatgtt tggtagaga actagaaggc
2641 ctttcccctg cccctgctca ggttagggtg gtggcgactg aaggccggg tgaatgttca
2701 taatggcttt ttacctgctt tgaaatgtgt gctttccctg aataatgcgg acttcgagag
2761 tgctgtccaa cctctcatgt gcacttggaa taaattctta ctttagaacc ttt

4. DEFICIENCIAS HUMANAS EN CPT1A y CPT2

Posición	Mutación	Observación	Referencia
Arg ¹²³ (exón 4)	R123C	Mobilidad aberrante.	Brown, 2001
Ala ²⁷⁵ (exón 8)	A275T	↓ eficiencia catalítica y estabilidad de la proteína.	Gobin, 2002
Cys ³⁰⁴ (exón 9)	C304W	↓ estabilidad de la proteína y no hay actividad CPT1	Gobin, 2003
Arg ³¹⁶ (exón 9)	R316G	Actividad CPT1 no detectable.	Bennett, 2004
Phe ³⁴³ (exón 10)	F343V		
Arg ³⁵⁷ (exón 10)	R357W	↓ estabilidad y actividad de la proteína.	Brown, 2001
Glu ³⁶⁰ (exón 10)	E360G	↓ estabilidad de la proteína y no hay actividad CPT1.	Ogawa, 2002
Del 1183_1185 (exón 11)	delR395	↓ estabilidad de la proteína, no hay actividad CPT1 y se deleciona la Arg ³⁹⁵ .	Brown, 2001 Bonnefont, 2004
Ala ⁴¹⁴ (exón 11), altamente conservado	A414V	Residuo posiblemente del centro catalítico del enzima y de unión del palmitoil-CoA. ↓ eficiencia catalítica y estabilidad de la proteína.	Gobin, 2002 Gobin, 2003
Asp ⁴⁵⁴ (exón 12)	D454G	↓ estabilidad, actividad y nivel (Western Blot) de la proteína	LIJ, 1998
Gly ⁴⁶⁵ (exón 12)	G465W		
Pro ⁴⁷⁹ (exón 12)	P479L	↓ actividad de CPT1. Resistencia parcial al malonil-CoA.	Bennett, 2004
Leu ⁴⁸⁴ (exón 12)	L484P	↓ estabilidad de la proteína .	Brown, 2001
Tyr ⁴⁹⁸ (exón 13), altamente conservado	Y498C	Residuo del centro catalítico del enzima y de unión del palmitoil-CoA. ↓ estabilidad de la proteína y eficiencia catalítica para ambos sustratos.	Gobin, 2002 Gobin, 2003
Thr ⁵⁶⁸	T568A	Presenta un polimorfismo.	Prip-Buus, 2001
Gly ⁷⁰⁹ (exón 17)	G709E	Afectan al sitio activo y elimina la actividad CPT1. Residuos	Gobin, 2003
Gly ⁷¹⁰ (exón 17)	G710E	implicados en la unión de carnitina en CPT2 de rata (Brown, 1994).	Prip-Buus, 2001
Tyr ³² (exón 2)	Y32X	Mutación sin sentido, causa una terminación prematura que resulta en una proteína CPT1 inmadura.	Ogawa, 2002
Gln ¹⁰⁰ (exón 4)	Q100X	Mutación sin sentido, trunca la proteína a 671 aminoácidos antes del final del TM2 del dominio N-terminal.	Gobin, 2002
Arg ¹⁶⁰ (exón 5)	R160X	Mutación sin sentido, produciría una proteína severamente truncada que carece de la mayor parte del centro catalítico y del dominio de unión de la carnitina.	Bennett, 2004
Trp ⁴⁷⁵ (exón 12)	W475X	Mutación sin sentido, ↓ la actividad CPT1 y no se detecta proteína en el análisis por Western Blot.	Ogawa, 2002
Tyr ⁴⁹⁸ (exón 13)	Y498X		
1876-2028del (exón 15)		Mutación por <i>splicing</i> , deleciona 51 residuos desde 626 hasta 676. CPT1 es inmadura e inestable.	Gobin, 2002 Ogawa, 2002
2027-2028+2delAAGT (exón 16)		Mutación por <i>splicing</i> , provoca tres transcritos aberrantes:	
• 1876-2028del		Inserción de 6 residuos desde la posición 676 a 677 y resulta en ↓ de actividad CPT.	Ogawa, 2002
• 2027-2028insGTCTCTT			
CCCTTCTTCC			
• 2026-2028del		Delección de 3 nucleótidos delección del E676del.	
Eliminación de 8 kb comprendido entre el intrón 14 hasta el exón 17 del gene. Resulta una proteína truncada desde el residuo 581 hasta 702. La delección provoca un mARN inestable.			
La inserción de 113-bp predice al addición de 8 residuos (FLPYHELS) que se terminen con un codón de stop, truncando así la proteína por 240 aminoácidos. Se inserta un residuo E525.			

Tabla 1. Resumen de las mutaciones sin sentido y missense descritas en CPT1A humana relacionadas con la actividad catalítica. Se indican las características provocadas por cada mutación en la actividad CPT1A.

Exón	Nucleótido ^a	Aminoácido	Referencia
1	109Ins GC		Martin, 2000
	112-113 Ins GC	S38fs ^c	Martin, 2000
	149C>A	P50H ^b	Verderio, 1995
2	216G>C/T	L72F	Ijlst, 1998
3	338C>T	S113L ^b	Taroni, 1993
	IVS3 + 5G>A ^d	Del179_113	Deschauer, 2003
4	359A>G	Y120C	Martin, 1999; 2000
	370C>T	R124X	Yang, 1998
	371G>A	R124Q	Thuillier, 2003
	437A>C	N146T	Thuillier, 2003
	452G>A	R151Q	Yang, 1998
	481C>T	R161W	Thuillier, 2003
	490A>T	K164X	Ijlst, 1998
	520G>A	E174K ^b	Yamamoto, 1996
	533-534insT; 534-558 del	L178F; N179-I186 del	Yang, 1998
	628T>G	Y210D	Ijlst, 1998
	641T>C	M214T	Wieser, 1997
	680C>T	P227L	Taroni, 1994
	821A>T	K274M	Ijlst, 1998
	890C>A/G	Y290X	Ijlst, 1998
	906C>T	R296X	Ijlst, 1998
	907G>A	R296Q	Ijlst, 1998
	907-918ins	L302fs ^c	Gellera, 1994
	983A>G	D328G	Thuillier, 2003
	1145G>A	R382K	Yang, 1997
	1148T>	F383Y ^b	Yamamoto, 1996
5	A1238-1239delAG	Q413fs ^c	Taggart, 1999
	1342T>C	F448L	Wieser, 1997
	1436A>T	Y479F	Wieser, 1997
	1459G>A	E487K	Bruno, 2000
	1507C>T	R503C	Taggart, 1999
	1543-1546delGCCT	515del4	Deschauer, 2002
	1646G>A	G549D	Taggart, 1999
	1649A>G	Q550R	Yang, 1998
	1657G>A	D553N ^b	Verderio, 1995
	1798G>A	G600R	Ijlst, 1998
	1810C>T	P604S	Yang, 1998
	1823G>C	D608H	Thuillier, 2003
	1883A>C	Y628S ^b	Bonnefont, 1996
	1891C>T	R631C ^b	Taroni, 1992

Tabla 2. Resumen de las mutaciones humanas en CPT2. Actualización de mutaciones en CPT2 que generan deficiencias descritas en la literatura. a = Nucleótido 1: A del codón ATG; b = Mutaciones expresadas en células COS; c fs = marco de lectura cambiado; d = Intrón 3. La tabla fue tomada de la referencia (Bonnefont, 2004).

5. ALINEAMIENTO DE SECUENCIAS CORRESPONDIENTES A CARNITINA/COLINA ACILTRANSFERASAS

Representación de secuencias de aminoácidos de 22 enzimas que catalizan acyl-CoAs de cadena de átomos de carbono corto como sustratos: CrAT (CACP) de levadura, *C. tropicalis*, humano, ratón y paloma; y ChAT (CLAT) de ratón, rata, cerdo, humano, mosca de fruta, y *C. elegans*; y las enzimas que catalizan acyl-CoAs de cadena de átomos de carbono media y larga como sustratos: CPT1A (CPT1) de rata, ratón y humano; CPT1B (CPTM) de rata y ratón; CPT2 (CPT2) de rata, ratón y humano; y COT (OCTC) de humano, rata y bovino; fueron obtenidos del banco de datos SwissProt y alineados utilizando el programa ClustalW. Los residuos han sido coloreados por conservación según el programa Belvu versión 2.8 (disponible en <http://www.sanger.ac.uk/~esr/Belvu.html>).

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CPT2_RAT      1 .....MMPRLLFRAWP,RCPSLVLGAPS... 24
CPT2_MOUSE    1 .....MMPRLLRDWP,RCPSLVLGAPS... 24
CPT2_HUMAN    1 .....MMPRLLLRQWP,RGPAGPGAPSRP 24
CACP_YEAST    1 .....MRICHSRTLSNLKDLPITSRRMHS... 35
CACP_CANTR   1 .....MFNFKLSSQVLKN..STKS 17
CACP_HUMAN    1 .....AFARTVVKP 10
CACP_MOUSE    1 .....MLAFAAARTVVKP 12
CACP_COLLI   1 .....MDRKQKQAEKARP 13
CLAT_MOUSE   1 .....P 1
CLAT_RAT     1 .....P 1
CLAT_PIG     1 .....P 1
CLAT_HUMAN   1 MGLRTAKKRGGLGGGKUKREEGGGTRGRREV... 110
CLAT_DROME   1 ...VASNEASTAAGSGPESAAFLS... 50
CLAT_CAEEL   1 .....MEKEK 5
CPTM_RAT     1 .....MAEAHQAVAFQTFTPDGVDFRLS... 53
CPTM_HUMAN   1 .....MAEAHQAVAFQTFTPDGVDFRLS... 53
CPT1_RAT     1 .....MAEAHQAVAFQTFTPDGVDFRLS... 53
CPT1_MOUSE   1 .....FQFTVTPDGIDLRLSHEALK... 44
CPT1_HUMAN   1 .....MAEAHQAVAFQTFTPDGVDFRLSHEALK... 44

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CPT2_RAT      25 LSAVS.....GPDYDQLQH...SIVPTMHYQD 47
CPT2_MOUSE    25 LSAVS.....GPAEYLQH...SIVPTMHYQD 47
CPT2_HUMAN    25 LSAGS.....GPGQQYLR...SIVPTMHYQD 47
CACP_YEAST    36 QFPVETNNGEHY...WAE.....KPNFKFYQNK... 72
CACP_CANTR   18 IMPILK.....KP...FSTS...HAKGDLFYQD 40
CACP_HUMAN    11 LGFLKPF.....SLMKASS...RFAKAHD 31
CACP_MOUSE    13 LGLLKPS.....SLMKVSG...RFAKAHD 33
CACP_COLLI   14 YGLLKPA.....ALGKIPG...RFGQHQE 34
CLAT_MOUSE   2 ILEKVP.....PKMPVQ...SS.CEEVL 21
CLAT_RAT     2 ILEKAP.....QKMPVKA...SS.WEEEL 20
CLAT_PIG     2 ILEKTP.....PKMAAKS...PSS...EEE... 21
CLAT_HUMAN   111 ILEKVP.....RKMAAKT...PSS...EES... 129
CLAT_DROME   51 LSNITPSDTGWQKDSILS.....IPKKWLS...AESVDEFGFP 85
CLAT_CAEEL   6 VDE.....LPP...NDNNUYET 18
OCTC_HUMAN   1 .....MENOLAKS...TEERTFQYQD 18
OCTC_RAT     1 .....MENOLAKS...IEERTFQYQD 18
OCTC_BOVIN   1 .....MENOLAKS...TEERTFQYQD 18
CPTM_RAT     54 VYPGSFTSULVVVMMATVGSNYCKVDISMGHLVHCICRQLCPTRGYSYGT... 162
CPTM_HUMAN   54 VYPGSFTSULVVVIMATVGSFCNVDISMGHLVSCICRQLCPQCGPYQT... 162
CPT1_RAT     54 VFPANPSSMULIVVVGVISSMHAKVDPDSLGMIAKISRTLDTTG... 160
CPT1_MOUSE   45 VFPASPFSSULIVVVGVISSMHTKVDPSLGMIAKINRTLDTG... 151
CPT1_HUMAN   54 VYPASPFSSMULIVVVGVMTTMYAKIDPSLGI... 160

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CPT2_RAT      48 .....SLPRLPIP KLEDTMKRYLNQKPLLD...SQFRRT... 141
CPT2_MOUSE    48 .....SLPRLPIP KLEDTMKRYLNQKPLLD...SQFRRT... 141
CPT2_HUMAN    48 .....SLPRLPIP KLEDTMKRYLNQKPLLD...SQFRRT... 141
CACP_YEAST    73 DPLSLPVPFLKSTIUDKYL... 166
CACP_CANTR   41 QLPKL... 135
CACP_HUMAN    32 ALPRLPVPPLQQLSDIHYL... 125
CACP_MOUSE    34 ALPRLPVPPLQQLSDIHYL... 128
CACP_COLLI   35 ALP... 128
CLAT_MOUSE   22 DPLKL... 115
CLAT_RAT     21 DPLKL... 114
CLAT_PIG     22 G... 115
CLAT_HUMAN   130 G... 223
CLAT_DROME   86 DTLKL... 180
CLAT_CAEEL   19 ALPKPVPSLEATUDRYLEYAAVVAVGOKASLATTHDAAHKFVRQ... 112
OCTC_HUMAN   19 SLP... 111
OCTC_RAT     19 SLP... 111
OCTC_BOVIN   19 SLP... 111
CPTM_RAT     163 RPMLYSFOTSPLKLPVPSPWATIHYL... 264
CPTM_HUMAN   163 HPMLYSFOTSPLKLPVPSPWATIHYL... 264
CPT1_RAT     163 KPMLYSFOTSPLKLPVPSPWATIHYL... 262
CPT1_MOUSE   152 KPMLYSFOTSPLKLPVPSPWATIHYL... 253
CPT1_HUMAN   161 KPMLYSFOTSPLKLPVPSPWATIHYL... 262

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Apéndices

CPT2_RAT	142	...KSEYNDQ.LTRATNLTVSAWRFKLTKIAGL..LEPEVFHLNPKSDTDAFKRLLIRFVPPSLSWYGAYLVNAPLDMQYFRFLNSTRIPRPNR.DE.....,LFTD	237
CPT2_MOUSE	142	...KSEYNDQ.LTRATNLTVSAWRFKLTKIAGL..LEPEVFHLNPARSDTDAFKRLLIRFVPPSLSWYGAYLVNAPLDMQYFRFLNSTRIPRPNR.DE.....,LFTD	237
CPT2_HUMAN	142	...KSEYNDQ.LTRATNLTVSAWRFKLTKIAGL..LEPEVFHLNPKSDTDAFKRLLIRFVPPSLSWYGAYLVNAPLDMQYFRFLNSTRIPRPNR.DE.....,LFTD	237
CACP_YEAST	167	NHLSKID.NDP.IKATAIIISTVWKFIEAIKDES.PVEIIKG..,PFCMNSFSLMNTSRLPGKFE.DNQ..DTNIFYSV	239
CACP_CANTR	136	NIIG...QDQ.LKATLIAAYTIEFQEKLVDSES.LPDEVIKGN..,PFCMNAFKYMHNNNSRVPAE..GS..,DITQHYNG	202
CACP_HUMAN	126	...DFVBLDQGQ.LRFRAAKLIEGVWDFKVMIDNET.LPVEYLG..,GPQLCMQNYQQQLSSCRVPGPKQ.DT,VSNSFS,KTKK	196
CACP_MOUSE	129	...DFVBLDQGQ.LRFRAAKLIEGVWDFKVMIDNET.LPVEFLG..,GPQLCMQNYQQQLSSCRVPGPKQ.DS,WNFL,KSKR	199
CACP_COLLI	129	...DFQDRQCGQ.LRFRAAKLIEGVWDFKVMIDNET.LPVEYMG..,GPQLCMQNYQQQLSSCRVPGPKQ.DS,IVNYA,KGKK	199
CLAT_MOUSE	116	..HFQDTNDQ.LRFRAASLISGVWSYKALLDSHS.PTDWAKGQLS..,GQPLCMQDYYRFLFSSYRLPGHTQ.DT,LVAQK,SSIM	190
CLAT_RAT	115	..HFQDTNDQ.LRFRAACLISGVWSYKALLDSHS.PTDWAKGQLS..,GQPLCMQDYYRFLFSSYRLPGHTQ.DT,LVAQK,SSIM	189
CLAT_PIG	116	..HFQDTNDQ.LRFRAASLISGVWSYKALLDSHS.PIDCAKPELS..,GQPLCMQDYYRFLFSSYRLPGHTQ.DT,LVAQK,SSIM	190
CLAT_HUMAN	224	...HFPGLTDDQ.LRFRAASLISGVWSYKALLDSHS.PIDCAKPELS..,GQPLCMQDYYRFLFSSYRLPGHTQ.DT,LVAQK,SSIM	298
CLAT_DROME	181	...SLQDPRPR.AHFARLLDGISLHREMLDSGE.UPLERALA.EK..,NQPLCMQDYYRFLFSSYRLPGHTQ.DS,QLPFLPSR,ERLN	255
CLAT_CAEEL	113	...FETKEDH.TKYTTALTRGLLEVKNLIDTQVKCREKSTGAQ..,KLONCMEQDYRVLSCYREPGVGE.DT,...QIR,OKT	183
OCTC_HUMAN	112	H..YWPPKEGTQDQERGSITLWHNINWYJQLLRRKEK..PVHKV..,GNTPLDNNQFRMLFSTCKPGITR.DSIMNYFR,TESE	185
OCTC_RAT	112	H..YWPPKEGTQDQERGSITLWHNINWYJQLLRRKEK..PVHK..,GNTPLDNNQFRMLFSTCKPGITR.DSIMNYFR,TESE	185
OCTC_BOVIN	112	H..YWPPKEGTQDQERGSISLWHNINWYJQLLRRKEK..AVEKV..,GNTPLDNNQFRMLFSTCKPGITR.DSIINYFR,TESE	185
CPTM_RAT	265	...KNT.SQD.AARLGNTVHAMMMYRRKLDRREE.IKPMVALG..,MVPNCYSQMERMFNTTRIPGKE.TD..,LQH..329	
CPTM_HUMAN	265	...KNT.DQD.AAARLGNIHAMMMYRRKLDRREE.IKPMVALG..,IVPNCYSQMERMFNTTRIPGKD.TD..,LQH..329	
CPT1_RAT	263	...TPT.HIQ.AARAGNTIHAIILYRRTLDREE.UKPIRLLGS..,TIPLCASQWERLFNTSRIPIGEE.TD..,IQHV..328	
CPT1_MOUSE	254	...TPT.HIQ.AARAGNTIHAIILYRRTVDREE.UKPIRLLGS..,TIPLCASQWERLFNTSRIPIGEE.TD..,IQHV..319	
CPT1_HUMAN	263	...LPT.HIQ.AARAGNTIHAIILYRRLKDRREE.UKPIRLLGS..,TIPLCASQWERMFNTSRIPIGEE.TD..,IQHM..328	

CPT2_RAT	238	TK.ARHLVLRKGHFV.FFDVLQDQGN.IVNPLTEQAHLYKILS.DSS.PVLEPF..,VAYLTSENRDVMAELRQLLIFDQGN..EETLKKVDSAVFCCLCDD..	329
CPT2_MOUSE	238	TK.ARHLVLRKGHFV.FFDVLQDQGN.IVNPLTEQAHLYKILS.DSS.PVLEPF..,LAYLTSENRDVMAELRQLLIFDQGN..EETLKKVDSAVFCCLCDD..	329
CPT2_HUMAN	238	DK.ARHLVLRKGNFY.FFDVLQDQGN.IVNPLTEQAHLYKILS.DSS.PVLEPF..,LAYLTSENRDVMAELRQLLIFDQGN..EETLKKVDSAVFCCLCDD..	329
CACP_YEAST	240	YE,NNFTI1AYKKGFKYKLMLTHGDNDKPLSE,NEIWPFLYSVVFQGSQ.SDPLKLG..IGSLTSPLRQDREVHELMKD,PIS.QDSLETI1HKSSFLMLCLLDQSP..	338
CACP_CANTR	203	EE,NQFTI1AYKKGFKYKLMLTHGDNDKPLSE,NEIWPFLYSVVFQGSQ.SDPLKLG..IGSLTSPLRQDREVHELMKD,PIS.QDSLETI1HKSSFLMLCLLDQSP..	296
CACP_HUMAN	197	P..PHTHTVWHYDFFELDVWHSQDT..PLTSDIIFVDEKIJUNSSL..,QTNK..EPVGILTSNHRNTPUAKAYHNLKDKVN..RESVNSIOKSITTFCLDQKQ.VP..	291
CACP_MOUSE	200	P..PHTHTVWHYDFFELDVWHSQDT..PLTSDIIFVDEKIJUNSSL..,QSNK..EPVGILTSNHRNTPUAKAYHNLKDKVN..RESVNSIOKSITTFCLDQKQ.VP..	294
CACP_COLLI	200	Q..SRHTTVWHNFQFFELDVWHSQDT..PLTSDIIFVDEKIJUNSSL..,QSNK..EPVGILTSNHRNTPUAKAYHNLKDKVN..RESVNSIOKSITTFCLDQKQ.VP..	294
CLAT_MOUSE	191	PE,PEHVIVACCCNQFFVLDWVIN.FR.RLSEGDLFTQDRLKIVKMRAS..,NEDERL.PPIGLLTSQDRSEMAKARTVLLKDSTN..RDSLDMIERCICLVCCLDGP.GT..	287
CLAT_RAT	190	PE,PEHVIVACCCNQFFVLDWVIN.FR.RLSEGDLFTQDRLKIVKMRAS..,NEDERL.PPIGLLTSQDRSEMAKARTVLLKDSTN..RDSLDMIERCICLVCCLDGP..	286
CLAT_PIG	191	PE,PEHVIVACCCNQFFVLDWVIN.FR.RLSEGDLFTQDRLKIVKMRAS..,NEDERL.PPIGLLTSQDRSEMAKARTVLLKDSTN..RDSLDMIERCICLVCCLDGP..GG..	287
CLAT_HUMAN	299	PE,PEHVIVACCCNQFFVLDWVIN.FR.RLSEGDLFTQDRLKIVKMRAS..,NEDERL.PPIGLLTSQDRSEMAKARTVLLKDSTN..RDSLDMIERCICLVCCLDGP..GG..	395
CLAT_DROME	256	DE,DRHVVICRNDQMCVVLQDSDP.KLSESEIASDILYVLSDAP..CLIAKP..VPVGILTSQDRSEMAKARTVLLKDSTN..RDSLDMIERCICLVCCLDGP..GG..	357
CLAT_CAEEL	184	NDGNEHVLVMCRNQTFLLHSRIN,GA,LVSYADVEYQLAQIEEISKINQNTAN..,IGASGVGPRDNAALFIQJDMLTVEQN..,SKSYEWVKSAL..,VCLDME.DPIDYG	285
OCTC_HUMAN	186	GRSPNHVLCRGRPFVDFWVHED..,CLITTPPELROLTYIHKKCHS..,EDQGD..GIAALTSEERTWAKAREYHLDGPEN..LALLEKIOSLSSLYVSMEDS.SP..	283
OCTC_RAT	186	GHCPTHIAVLCRGRGFVDFWVHED..,CLITTPPELROLTYIHKKCHS..,EDQGD..GIAALTSEERTWAKAREYHLDGPEN..LALLEKIOSLSSLYVSMEDS.SP..	283
OCTC_BOVIN	186	GHSPHSLAVLCRGRGFVDFWVHED..,CLITTPPELROLTYIHKKCHS..,EDQGD..GVAALTEERL..,RNLTEKIOSLSSLYVSMEDS.SP..	283
CPTM_RAT	330	SE,SRHIVAYHKGRFFFKMLYIYE,GSC,LLKPRDLENQFQRILD,DTSPSPQRGE..,KLAALTAGGRVEMAEARQKFFSSGK..,NKSMLDITIERRAFYVALI..E..DS..	425
CPTM_HUMAN	330	SD,SRHIVAYHKGRFFFKMLYIYE,GAR,LLKPKDQLENQFQRILD,DTSPSPQRGE..,KLAALTAGGRVEMAEARQKFFSSGK..,NKSMLDITIERRAFYVALI..E..ES..	425
CPT1_RAT	329	KD,SRHIVVYHGRGYFKWJLYH,DGR,LLRPRELEQMQMQLD,DTSPSPQRGE..,KLAALTAADRVPMWAKRCOTYFARGK..,NKQSLDAVEKAFFVTLD..E..SE..	424
CPT1_MOUSE	320	KD,SRHIVVYHGRGYFKWJLYH,DGR,LLRPRELEQMQMQLD,DTSPSPQRGE..,KLAALTAADRVPMWAKRCOTYFARGK..,NKQSLDAVEKAFFVTLD..E..SE..	415
CPT1_HUMAN	329	RD,SKHIVVYHGRGYFKWJLYH,DGR,LLKPREMEQMQMQLD,DTSPSPQRGE..,R..,KLAALTAADRVPMWAKRCOTYFARGK..,NKQSLDAVEKAFFVTLD..E..TE..	424

CPT2_RAT	330	...FPMKDLI.....,HLSHTMLHQ..,DGTNRUFDKSFNLFNIVAEQDGTAAVHFHESWQDGVAVLRFNNEVFRDSTQ..,TPAIPATQOS.QPAATNTSSA	410
CPT2_MOUSE	330	...FPMKDLV.....,HLSHTMLHQ..,DGTNRUFDKSFNLFNIVAEQDGTAAVHFHESWQDGVAVLRFNNEVFRDSTQ..,TPAIPATQOS.QPAATNTSSV	410
CPT2_HUMAN	330	...FP1KDLV.....,HLSHTMLHQ..,DGTNRUFDKSFNLFNIVAEQDGTAAVHFHESWQDGVAVLRFNNEVFRDSTQ..,TPAIPATQOS.QPAATNTSSV	410
CACP_YEAST	339	...VTLTE.EK.....,SRNCWHLH..,DGTNRFYDQKSLFQDPLVFTGNGSSGFLAEHKSMQDGTPTFLFLNNVYCQQLNKLDDV..,DDFMRKVITP..,SSTVAM	418
CACP_CANTR	297	...VTLTE.EK.....,SKNCWHLH..,DGNQRFYDQKSLFQDPLVFTGNGSSGFLAEHKSMQDGTPTFLFLNNVYCQQLNKLDDV..,DDFMRKVITP..,SSTVAM	375
CACP_HUMAN	292	...PVSE.DVYPS..,HVGQMLHGGGSPLNSNPUSNPUDPFTLQDIFVIAEDGSCLVYEHAAEEFPVITLDDVWVIEYTKPP..,NDLIVEIGS..,SAPRFG	374
CACP_MOUSE	295	...RVSD.DVYRN..,HVGQMLHGGGSFKFNSGRUJFQDFKTLQDIFVIAEDGSCLVYEHAAEEFPVITLDDVWVIEYTKPP..,ELVRSP..,MVPLP	377
CACP_COLLI	295	...RVSD.DIVYKS..,PVVAGMLHGGGSRLNUSGRUJFQDFKTLQDIFVIAEDGSCLVYEHAAEEFPVITLDDVWVIEYTKPP..,ELVRSP..,MIPLP	377
CLAT_MOUSE	288	..,GDL5.DTHR..,ALQLLHGCGGCSLNLGNGRNUYDKSFLQVWGRBGTGCVCEHSPFDGIVLVQVCTEHLKKHMTGN..,K.KLVRVDS..,VSEL..	370
CLAT_RAT	287	..,GELS.DTHR..,ALQLLHGCGGCSLNLGNGRNUYDKSFLQVWGRBGTGCVCEHSPFDGIVLVQVCTEHLKKHMTGN..,K.KLVRVDS..,VSEL..	369
CLAT_PIG	288	..,MELS.DTHR..,ALQLLHGCGGCSLNLGNGRNUYDKSFLQVWGRBGTGCVCEHSPFDGIVLVQVCTEHLKKHMTGS..,K.KMVRADS..,VSEL..	370
CLAT_HUMAN	396	..,VELS.DTHR..,ALQLLHGCGGCSLNLGNGRNUYDKSFLQVWGRBGTGCVCEHSPFDGIVLVQVCTEHLKKHMTGS..,R.KLIRADS..,VSEL..	478
CLAT_DROME	358	ARGGTAGTP..,TVHLAGRD..,ETMNAHEMIHKGGSEYNSGRUJFQDFKTMQIITCTWVQCLCYEHCSCS..,IAVOLLEKDYKKIEHP..,D.EDNGLP..,DHHE..	453
CLAT_CAEEL	286	KMD,TMSIS,EKEKE....,FVARGYSTLTGHGSKFLGFLNRMUQDITQIQLVMSSSGVNGLCIEHSTAEQVIIINMAETAIRYAQYF..,KS.KMWN..,DVRNV	376
OCTC_HUMAN	284	..,HVT,PEDYS..,EIIAIIILG..,DPTVPRUGDKSYNLSFVQDFVQDFPQDAMIVMNISYYDEKIFQVQ..,EG.RWKGSE..,KVRDIP	363
OCTC_RAT	284	..,HAT,PE NFS..,QVFEMLLGG..,DPTVPRUGDKSYNLSFVQDFVQDFPQDAMIVMNISYYDEKIFQVQ..,EG.RWKGSE..,KVRDIP	363
OCTC_BOVIN	284	..,HVT,PEDYS..,QVSALKING..,DPTVPRUGDKSYNLSFVQDFVQDFPQDAMIVLWVQVYVYDENILEN..,EG.RWKGSE..,KVRDIP	363
CPTM_RAT	426	..,HCYN,PDQ..,EASLSLYGKSSLH..,NCYNRUFDKSFTLISCKNGQLGLNLEHNSWADAPIGHJLWEFLVTDGFLH..,GYTET.GHCVGE..,PNTKLE	513
CPTM_HUMAN	426	..,QSYD,PED..,EASLSLYGKALLH..,NCYNRUFDKSFTLISCKNGQLGLNLEHNSWADAPIGHJLWEFLVTDGFLH..,GYTET.GHCLGK..,PNPALA	513
CPT1_RAT	425	..,QGYR,EDDP..,EASISDYKSSLH..,RCFDRWDFDKSITVFWFKNSK1GINAEHNSWADAPIVGHJLWEYMATDQL..,GYSED.GHCKGD..,TNPNTI	513
CPT1_MOUSE	416	..,QGYR,EDDP..,EASISDYAKSLLH..,RCFDRWDFDKSITVFWFKNSK1GINAEHNSWADAPIVGHJLWEYMATDQL..,GYSED.GHCKGD..,KNPNP	504
CPT1_HUMAN	425	..,EGYR,SED..,DTSMDSYAKSLLH..,RCYDRWDFDKSITVFWFKNSK1GINAEHNSWADAPIVGHJLWEYMATDQL..,GYAED.GHCKGD..,INPNP	513

CPT2_RAT	411	SVETLSNLSGALKAGITAAKEKFDTTWTKLISIDS1QFQRGCKEFLKQKLSPLDAVQLAQAFQMAFLRQYQG..,QTVATYESCSTAAFKHGRTEТИR..,PASIFTKRCSEAFV	517
CPT2_MOUSE	411	SVQKLSMQLSLLALKAGVTAKEKEFDATMKTLLTIDQIAQFQRGCKEFLKQKLSPLDAVQLAQAFQMAFLRQYQG..,QTVATYESCSTAAFKHGRTEТИC..,PASIFTKRCSEAFV	517
CPT2_HUMAN	411	TVQKLNELTDLAKTGTIAKEKEFDATMKTLLTIDQIAQFQRGCKEFLKQKLSPLDAVQLAQAFQMAFLRQYQG..,QTVATYESCSTAAFKHGRTEТИC..,PASIFTKRCSEAFV	517
CACP_YEAST	419	KPMELPFIITPKIHKAIASEQALQKFETIGEHDLRVWVYHKYKGTFTIKRHGMSPDFAITQVQILOAVFKYLK..,RGLPTYEAASTRKFLKGRTETR..,SVTASLEFVSKWQ	525
CACP_CANTR	376	NAEIPEPDINPTTRANIKDIAKDFATIAHDEEIQHFGYKGLIKKFVKSFDAYQMLQLAQYRMYG..,HSCATYEESASLRFMLGRTDTIR..,STSIESHKFVQSD	482
CACP_HUMAN	375	MPKKLRLNIPTEIKNDIEKAQKNLISIMIQLDQIMMLTFHHFGKDFPKSEKLSPLDPAITQVQILOAVFKYLK..,QADATYEESASLRFMLGRTDTIR..,SASIDSLAFVKGM	484
CACP_MOUSE	378	MPKKLRLNIPTEIKNDIEKAQKNLISIMIQLDQIMMLTFHHFGKDFPKSEKLSPLDPAITQVQILOAVFKYLK..,QADATYEESASLRFMLGRTDTIR..,SASIDSLAFVKGM	484
CACP_COLLI	378	MPKKLRLNIPTEIKNDIEKAQKNLISIMIQLDQIMMLTFHHFGKDFPKSEKLSPLDPAITQVQILOAVFKYLK..,QADATYEESASLRFMLGRTDTIR..,SASIDSLAFVKGM	484
CLAT_MOUSE	371	APRRLRLUKCSPEIQQGLLASSAAEKLQRIKVLNDFIVYKFDNYGKTFIKHQKCSPLDPAITQVQILOAVFKYLK..,R.LVPTEYESAISTRFQEGRVDNIR..,SATPEALAFVQAMT	477
CLAT_RAT	370	APRRLRLUKCSPEIQQGLLASSAAEKLQRIKVLNDFIVYKFDNYGKTFIKHQKCSPLDPAITQVQILOAVFKYLK..,R.LVPTEYESAISTRFQEGRVDNIR..,SATPEALAFVQAMT	476
CLAT_PIG	371	APRRLRLUKCSPEIQQGLLASSAAEKLQRIKVLNDFIVYKFDNYGKTFIKHQKCSPLDPAITQVQILOAVFKYLK..,R.LVPTEYESAISTRFQEGRVDNIR..,SATPEALAFVQAMT	477
CLAT_HUMAN	479	APRRLRLUKCSPEIQQGLLASSAAEKLQRIKVLNDFIVYKFDNYGKTFIKHQKCSPLDPAITQVQILOAVFKYLK..,R.LVPTEYESAISTRFQEGRVDNIR..,SATPEALAFVQAMT	585
CLAT_DROME	454	PPERLEHIVGCPOLQLPFDASKVSKCICDLDLFFVYRQYDQYGTFTIKSCQVSPDQVYIQLATATGSLDQVY..,RTSGHLLRKCVHSTISOPPRRLHQSGQHGCIVGQAM	560
CLAT_CAEEL	377	HPKSLTWHFSNSRNILKKQAEVFLDANELELEVLFNEFGKDSIKHWRVSPDGFQIQLIMQLAHYKTHG..,HLVSTYESASVRFQGAVRDNIR..,ANTQALEMVTAM	483
OCTC_HUMAN	364	LPEELIIVDEKVLNIDNOAKAQYREASDLQQAAYAFTSFQGKLLTKNMLHDPDFTIQLQALQLAHYRRL..,HGHPGCCYETAMTRHRYHGRTETR..,SCTEAVRWCQSD	470
OCTC_RAT	364	LPEELIIVDEKVLNIDNOAKAQYREASDLQQAAYAFTSFQGKLLTKNMLHDPDFTIQLQALQLAHYRRL..,HGHPGCCYETAMTRHRYHGRTETR..,SCTEAVRWCQSD	470
OCTC_BOVIN	364	VPEELVTVDEKVLNIDNOAKAQYKQVSDLQLVWYVATSFQGKLLTKNMLHDPDFTIQLQALQLAHYRRL..,HGHPGCCYETAMTRHRYHGRTETR..,PETVEAVNJCOSQ	470
CPTM_RAT	514	PPQRMQDIPEDQOQATIENSYBVAKLADDVYLCFOFLPFGKGLIKKCRSPDPAITQVQILOAVFKYLK..,RGKFDLTYEASMTPMFRGRTETR..,SCTSSESTAFVRAM	620
CPTM_HUMAN	514	PPTRLODIPGECQEVIDQSLSSAFLNNDLHSFFD5FQGKGLIKKCRSPDPAITQVQILOAVFKYLK..,RGKFDLTYEASMTPMFRGRTETR..,SCTSSESTAFVRAM	620
CPT1_RAT	514	KPTRLQDIPGECQEVIDQSLSSAFLNNDLHSFFD5FQGKGLIKKCRSPDPAITQVQILOAVFKYLK..,MKGKFDLTYEASMTPMFRGRTETR..,SCTMESCNFVQAM	620
CPT1_MOUSE	505	KPTRLQDIPGECQEVITSLSSAFLNNDLHSFFD5FQGKGLIKKCRSPDPAITQVQILOAVFKYLK..,MKGKFDLTYEASMTPMFRGRTETR..,SCTECNSCNFVQAM	611
CPT1_HUMAN	514	YPTRLQDIPGECQEVITSLNTANLLANDVHFSEPFVAFQGKGIKKCRSPDFTVQILOAVFKYLK..,MKGKFDLTYEASMTPMFRGRTETR..,SCTECNSCNFVQAM	620

Apéndices

CPT2_RAT	518	R.....DPSKHSGV.....ELQHMMAECSKYHGQLTKEAAMGQGFDRHLIYALARYLATARGNLN.....PE.....LYLDPAYQQMN.....HNILSTSTLN	593
CPT2_MOUSE	518	R.....EPSKHSVG.....ELQHMMAECSKYHGQLTKEAAMGQGFDRHLIYALARYLAARGVTL.....PE.....LYDOPAYQRIN.....HNILSTSTLS	593
CPT2_HUMAN	518	R.....EPSRHSAG.....ELQHMMVECSKYHGQLTKEAAMGQGFDRHLIYALRHLAAKGIL.....PE.....LYDOPAYQGIN.....HNVLSTSTLS	593
CACP_YEAST	526	I G.....DPIRAEK.....IAQLKHSAKEHSTYLKNAANGNSVDRFFGKLKNNLKSNDDQ.....IPP.....LFKDPLFNYSS.....TWLISSTSOL	599
CACP_CANTR	483	P.....NASSDK.....VATFQARAKQAVAYLSAARDGKVDRHLFLGKQMIQPGEP.....IPE.....IFTDPFIFSYSQ.....TWYISSSQV	555
CACP_HUMAN	482	ISSVTEHQK.....VELLRKVAQHARGYTDRAIRGEAFDRHLGLKLQIAEDLWST.....PD.....IFNDTSYAZAM.....HFHLSTSQV	555
CACP_MOUSE	495	ISTVPELQK.....VELLRKVAQHARGYTDRAIRGEAFDRHLGLKLQIAEDLWSM.....PD.....IFNDTSYAZAM.....HFNLSTSQV	558
CACP_COLLI	495	S.....PDKSDQEK.....ADLLRRTPQHKEYTMMAIQNAIDRHLGLKLQIAEDLWSI.....PE.....LFMDTAYAVAM.....HFNLSTSQV	558
CLAT_MOUSE	478	DHK.....AAVLASEK.....LQLLQRIQOQEYTVMAITGMADINHULLALRELARDLCKEP.....PE.....MFMDETYLMSN.....RFILSTSQV	553
CLAT_RAT	477	DHK.....AAMPASEK.....LQLLQTPMQCHQGYTVMAITGMADINHULLALRELARDLCKEP.....PE.....MFMDETYLMSN.....RFVLSTSQV	552
CLAT_PIG	478	DHK.....SAMPSEK.....LLLKDPIRAQTOYTVMAITGMADINHULLGLRELAREVCKEL.....PE.....MFMDETYLMSN.....RFVLSTSQV	553
CLAT_HUMAN	586	DHK.....AVPASEK.....LLLKDPIRAQTOYTVMAITGMADINHULLALRELAREVCKEL.....PE.....MFMDETYLMSN.....RFVLSTSQV	553
CLAT_DROME	561	QEGEGANPVLSEPLEDDEEEESRKVKFSIYSKDHLERLFRCAVARQTEVMVRISWAMASTROWPHRGQYGRHRRDA.....RA.....VQRRVLQQCSQ.....CNLLSTSQV	661
CLAT_CAEEL	484	SK.....KESKERK.....LELFKKAVLKQVKVTLLENISGYGVBNHLCALFCLAREEREETTEDIPS.....LFQDPLWSEVM.....RFPPLSTSQV	561
OCTC_HUMAN	471	D P.....SVNLRE.....QOKMLQFAKHNKMMDCSAGKGFDRHLGLLULIAKEEGLPV.....PE.....LFDTPLFSKSGGG.....GNFWLSTSILV	546
OCTC_RAT	471	D P.....SASLLER.....QOKMLQFAKHNKMMDCSAGKGFDRHLGLLULIAKEEGLPV.....PE.....LFDTPLFSKSGGG.....GNFWLSTSILV	546
OCTC_BOVIN	471	D P.....STSLLER.....KHMMLFQFAKHNKMMDCSTGKGFDRHLGLLULIAKEEGLPV.....PE.....LFDTPLFSKSGGG.....GNFWLSTSILV	546
CPTM_RAT	621	TG.....SHKKQDL.....QDLFRKASEKHONMYRLAMTGAGIDRHLFCYIVSKYLGWS.....PFLDEVLSERPWLSTSQIPOFQTCMFDPKQY	703
CPTM_HUMAN	621	E G.....SHTKADL.....RDLFQVSLPWLSTSQIPOSQIRMFDPEQHP.....PFLAEVLSERPWLSTSQIPOSQIRMFDPEQHP	703
CPT1_RAT	621	D P.....KSTAEQR.....LKLFIKICCEKHQHLYRLAMTGAGIDRHLFCYIVSKYLGWS.....PFLKEVLSERPWLSTSQTPOQQQVELDFEKNP	703
CPT1_MOUSE	612	D P.....TTTAEQR.....LKLFIKICCEKHQHLYRLAMTGAGIDRHLFCYIVSKYLGWS.....PFLKEVLSERPWLSTSQTPOQQQVELDFEKYP	694
CPT1_HUMAN	621	D P.....AQTEQR.....LKLFIKICCEKHQHLYRLAMTGAGIDRHLFCYIVSKYLGWS.....PFLKEVLSERPWLSTSQTPOQQQVELDFELEN	703

CPT2_RAT	594	SPAVS...LGGFAPVVPDGFCIAYAVH,DWJIGCNVSSYSGR...NAREELHCVQKCLEDFDALE.....GKAIKT.....	658
CPT2_MOUSE	594	SPAVS...LGGFAPVVPDGFCIAYAVH,DWJIGCNVSSYSGR...NAREELHCVQKCLEDFDALE.....GKAIKT.....	658
CPT2_HUMAN	594	SPAVN...LGGFAPVVPDGFCVGVAVH,DWJIGCNVSSYSPGR...NAREELQCVKALEDIFDALE.....GKSIKS.....	658
CACP_YEAST	600	SEYFD...GYUSQVNNDNGFLGLAMLN,NEWLJHINIVNPKPNSGASVNRLLHYLSQADEE.....ENKRKAHL.....	670
CACP_CANTR	556	SEFFQ...SUGMISQVNDDGFLGLAMLN,NDWJHVNHHISCKRN, GLOSDHDLQVLYLVDSANEMIKDWLTKGL.....LTDAPKKL	627
CACP_HUMAN	556	AKTDC...VMF6GPVVPDGYGVYNP,EAHINELSLSYNSCAETNAARMALYHEKALLDIDLRLQSHPR.....AKL.....	624
CACP_MOUSE	559	AKTDC...VMF6GPVVPDGYGVYNP,EAHINELSLSYNSCAETNAARMALYHEKALLDIDLRLQSHPR.....AKL.....	627
CACP_COLLI	559	AKTDC...VMC6GPVVPDGYGVYNP,GEHINFADISFNSCAETNAARMALYHEKALLDIDLRLQSHPR.....SKL.....	627
CLAT_MOUSE	554	TTMEM...FCCYGPVVPNGYGACYNP,AEAIFTFCISSFHGQKETTSSVETAEAVGASLVDRDCSSRQP.....ADSKPPTAKERAGPSQAKQS	640
CLAT_RAT	553	TTMEM...FCCYGPVVPNGYGACYNPQ,PEAIFTFCISSFHGQKETTSSVETAEAVGASLVDRDCSSRQP.....ADSKPAPKEKARGPSQAKQS	639
CLAT_PIG	554	TTMEM...FCCYGPVVPNGYGACYNPQ,PESILFCISSFHGQKETTSSVETAEAVGASLVDRDCSSRQP.....ADSKPAPKEKARGPSQAKQS	640
CLAT_HUMAN	662	TTTEM...FCCYGPVVPNGYGACYNPQ,PETILFCISSFHGQKETTSSVETAKAVEEESFIENKGUCSLSQS.....GMGKPLATKEKTRPSQVHOP	748
CLAT_DROME	658	CSTDs...FMGYGPVTPREGYCSVNP,PEQIVFCWSFYSCEDTSASRYVAKSLQDSLIDRDCSLLPP.....TESKPLATEKTRPSQGHOP	720
CLAT_CAEEL	562	TSLDIPDCYLTQAVVVRDGYCPVNIQ,PDRWTFIAFTFPRDPTDQLQHEKKSLAGMPDKELLSN.....	627
OCTC_HUMAN	547	GYLRV...QGVVPMVHNGYFFYHIR,DDRFTVVAWSAWKSCPTDAREKLVQLTFCAFHDIQLMNSTHL.....	612
OCTC_RAT	547	GYLRI...QGVVPMVHNGYFFYHIR,DDRFTVVTCSMJKSCPTDAREKLVQOVFHAFCDIMQMMPMEPHL.....	612
OCTC_BOVIN	547	GYLRV...QGVVPMVHNGYFFYHIR,DDRFTVVSCHMJKSCPTDAREKLVQOVFHAFCDIMQMMPMEPHL.....	612
CPTM_RAT	704	NHLGA...GGGF6GPVADDGYGVSV,MIAGENTMFFHVS5KLSSSETNARLREGNHIRQDALLDIADEFK1SKTDS.	772
CPTM_HUMAN	704	NHLGA...GGGF6GPVADDGYGVSV,MIAGENTMFFHVS5KLSSSETNARLREGNHIRQDALLDIADEFK1SKTDS.	772
CPT1_RAT	704	DYVSC...GGGF6GPVADDGYGVSV,IIVGENFIIFHIS5KFSSPETDSHREGRKHLRQAMMDIT,FGLTANSKK.	773
CPT1_MOUSE	695	DYVSC...GGGF6GPVADDGYGVSV,IIVGENFIIFHIS5KFSSPETDSHREGRKHLRQAMMDIT,FGLTANSKK.	764
CPT1_HUMAN	704	EYVSS...GGGF6GPVADDGYGVSV,ILVGENLNFHIS5KFSSPETDSHREGRHLKEAMTDIT,FGLSSNSKK.	773