Su	plemnStary	plemnStatFin	gntnatFin	uSlnsmSet1FAn1	BleCD	1eBEGHI2y.p32a4kB	EGHI2y.p32aEJ IFmSeaseC)tn
- N	LKLKE5675E5KE5KE OKLK Z	pleBlFiinuasn11aunFmāDlemnSta nlemnF eina CACtSmaF1DFamDnFōlemnF eina CACtSmaF1DFamDn	pyHy□ n2TF	HI2y.p32 pyHy HI2y.p32 p2EF	temaset Sunlnu.AFsBleCtu temaset Sunlnu AFsBleCtu	K445EE	KAARF	K F
	LKLK36262E62EKLK4EKLK46p	plennir ena cActoniar i Dean Dieplennir ena cActoniar i Dean Di	D.	HI2y.p32 D	temaset Sunlnu.AFsBleCtu	KHOLL -	KK4K5	4
	LKLK3gUKL47pp44EUKLKIDKI	ymleDSt	yy⊡ x and	HI2y.p32 yy	temaset Sunlnu.AFsBleCtu			ĸ
		E LIESUAAnm Falsabinsel etablem nstAstustbaris Labami Ania4 5 ISelnueStalnuesm F na4 äsme D1F iSs	Lp334 Iv4	H12y.p32 Lp354 H12y.p32 I¥4	temaset Suninu.AFsBleCtu	KKE4K	KKED	4 E
C	p5pBLKLKErEpBKLKBEEKE	g1CmFmSetnaDnleSuF npcDc1SDSuaüleDnleSunaB1CmFmSetnaDnleSuF näiSmesetulS	Fglp 5	HI2y.p32 gp5	temaset Sunlnu.AFsBleCtu	KK54□	KE4E54	Е
44 4E	LKLK \$25 B1B5 B1CKLK 1667 667 667 6 GELKLK B1CKLK 165 65 E65 FELK	7 mAIFtnE-esSFmnuaBCFt1FmnaStF nääFtuapy7aueiFStšetmFStStBaDlemnSta4 11 stsentinal amSen mnlF na41 stsentinal amSen mnlF naF üSmesetulSF1	Egr4 1 HIATHIE	HI2y.p32 Egr4	temaset Sunlnu.AFsBleCtu temaset Sunlnu AFsBleCtu	KEK54 🗆	K4450 KAAKEKA	E 4
45	6HKLKELKLK344KLKLKIL46H	HetmFsmStE esSFmnuaDlemnStISña5	HILp5	HI2y.p32 HIEp5	temaset Sunlnu.AFsBleCtu		Runne	ĸ
40	E4LKLKEKLKEEEEE	LyäunüleBntF naCASCStetnäSletiC1ClaDlemnStaäiSmesētulSF1	ÿ2□	HI2y.p32 y2	temaset Sunlnu.AFsBleCtu		KE S	4
40] 64E4E64E4EKLKIKEKLK⊟⊡] I KI K7EKI KI K256EEKI KER65E	nsnDmelmDnamle StnDlemnStaD@DEmF nanmFplemnStmle StnD@DEmF n F@tuFAaCmanialaRa4	plp□ tat	HI2y.p32 pIp []	temaset Sunlnu.AFsBleCtu temaset Sunlnu AFsBleCtu			ĸ
E	K6E5EKLKIPEKrE6E6	ElaCD mlnFianl nintmAStuStBaDlemnStaE	2p	HI2y.p32 2p	temaset Sunlnu.AFsBleCtu			ĸ
E4	4 65 EKLK IGE 6Er 6Er 6Er ELKL3r15	FyalnDnFmsētmFStStBaDlemnSta□	ÿ□	HI2y.p32 y□	temaset Sunlnu.AFsBleCtu		4KE 54 🗆	4
E E	□ p445K6p445KEDKLKD4BCEB35 □ setset#STKLK#set#DKLKLKD5s	t pel Funt IFmnAStuStBaDlemnSta4pel Funt IFmnAStuStBaDlemnSt	pL3pH4	HI2y.p32 pL3pH4	temaset Sunlnu.AFsBleCtu	KK 54	KE5EEK45	E
E	LKLKgE2DKLKE63ELKLKEy	ZalSAe eiFlaDlemnSta5äiSmesētulSF1	p5	HI2y.p32 p5	temaset Sunlnu.AFsBleCtu			ĸ
E	64KKEKEKE HprpK4p3KK5	2SBtF1aDnDmSuF naseiD1naCACtSmaE	2pH2E	HI2y.p32 2pH2E	temaset Sunlnu.AFsBleCtu	KK 🗆	KKK4E□	E
4	4545EEKLKE45 ADDEKLKE45v2nKEKLKEAADE	nmnleBntneC atCs1nFlalSAetCs1neDlemnStayISn LimlemlEt De etürlSniteDlemnStan#4K	ply⊡ notK	HI2y.p32 pay□ HI2y.p32 patk	temaset Sunlnu.AFsBleCtu			ĸ
	6DKLK 6D6pKyppyEp6g	E2aDlemnF einatetElpF nalnBC1FmelaCACtSma4	p2ÿ4□	HI2y.p32 p2y4	temaset Sunlnu.AFsBleCtu	KKE5E		4
5	pKEKEKLK6EKLES	g1CmFmSetnaDnleSuF na4g1CmFmSetnaDnleSuF n	gp4	HI2y.p32 gp4	temaset Sunlnu.AFsBleCtu	KEK5E		4
	BEKIKEE#FEREN4EHEH6E43	I K2alSAe eiF1aDlemnSta4K U leiFmSeFiSteFeSuanneUAetF n	p4K	HI2y.p32 p4K	temaset Sunlnu.AFsBleCtu	VESE	K4EK4⊟ KKAN⊡	4 F
	LKLK 56337KE63E65K65E6E6	eteCtsmSetF1aH4mnmlFülee1Fmna tmEnäiSmesetulSF1	Iç4 □	HI2y.p32 I¥4	temaset Sunlnu.AFsBleCtu	KEBE	KR4K L	4
	6KK4KE6KK4KEKLK6224126E6E	H1FmIStañFãsESta4H1FmIStañFãsESt	н́тн	HI2y.p32 HIH	temaset Sunlnu.AFsBleCtu	KK44#	KEKKKK 🗆	Е
56	LKLKE6pE6pE2EKLKEg⊡ Anometrikteen	SAe ein#StuStBaDlemnSta4	3p4	HI2y.p32 3p4	temaset Sunlnu.AFsBleCtu	KK II	KK4EE	E
50	35ygKEDKLKKKEKEELKL7LDK	ILIpAStuStBasF nmmna CAEIS1a3ainiAnlaaiSmesētuISF1	L3H3	HI2y.p32 L3H3	temaset Suninu.AFsBleCtu		K4K3E4E	4
55	65KE65KLKLK 10 87 11 5	E2aDlemnF einatetElpF nalnBC1FmelaCACtSma4	p2y4	HI2y.p32 p2y4	temaset Sunlnu.AFsBleCtu	K4545	KEK	Е
50		iimE1GEJaBSFtma1FIEnaDlemnStafie1eBa4	g4 UmpH	HI2y.p32 g4	temaset Sunlnu.AFsBleCtu temaset Sunlnu AFsBleCtu		KKKAKD	K 4
50		4 gnalnuCsmF ngnalnuCsmF naE	mppr	HI2y.p32 HDpH	temaset Suninu.AFsBleCtu		KKK+K L	ĸ
5	6H5ELKLK 6H56H5R 85gH5H50	eCtuFAeCmaeie1eBaE	3E	HI2y.p32 3E	temaset Sunlnu.AFsBleCtu	4550		4
K	H6746745674674674E674E674EKI	LüleinnleSua44AnmFünüleBntF na41SnaDlemnSt	2y4434	HI2y.p32 2y4434	temaset Sunlnu.AFsBleCtu			K
E	6445DKLKEDKLKEUKLKEKee	ZninteDiemnSta□ EILävD#StuStBaDlemnSta5□	2pa ILv3nIvn5	H12y.p32 2p4 H12y.p32 ILv3p	temaset Suninu.AFsBleCtu	KEKE	K5K 🗆	E
	ECKLK H27 BEHE H4 HE 5643 H K	LyäunüleBntF naCASCStetnä4aF1DEa CAseiD1nāCACtSma4KaiSmesētulSF1	yE4K	HI2y.p32 yE4K	temaset Sunlnu.AFsBleCtu	KKE	KKE4K4	E
	6IEKLK54	2euSCi.üleBntansFtBnlaZeuSCi.üleBntansFtBnl		HI2y.p32 2HL	temaset Sunlnu.AFsBleCtu	WW2224	V PPP 4-	K
	LKLK BK B	3F F1asn11aFunSetaie1nsC1n	раранаен ЗНГП	H12y.p32 p4	temaset Suninu.AFsBieCtu	K455KK4		E
ĸ	LKLKOp	I224ISnaDlemnSt	I224 🗆	HI2y.p32 I224	temaset Sunlnu.AFsBleCtu		K4454	4
4	62ELKLK BLKLE5562	nCSInuaēlainSemSsatCs1nFlauSS/SetaDlemnSta4aēie1eB	ça.	HI2y.p32 🖗	temaset Sunlnu.AFsBleCtu		K4KEE□	4
E	LKL5EBF4EKgEEKgEKLKEKgE 65x4EKLKEKgEF4a565x4EB676E	K mntunua (FDmemFBiStE GetEmSotFlainStFl	ZIE	HI2y.p32 ZIE	temaset Sunlnu.AFsBleCtu	K5#4□ KE#44	K4K5	E
5	6p7EKLKK	IlēiAe DetuStamDnfaueiFStsētmFStStBaDlemnStaL	12yE	HI2y.p32 I2yE	temaset Sunlnu.AFsBleCtu	KK45	K III	4
	LKLK EpiEpiE6lgK	EsCe1FlaDlemnSta elmStBE esSFmnuaDlemnStaIL4aēie1eB	П.4	HI2y.p32 II.4	temaset Sunlnu.AFsBleCtu	K454	KE54E 🗆	Е
	684ECKLK#6884p858p8158p8	LpEaseiD1naCACtSmaiC	LpE4	HI2y.p32 LpE4	temaset Sunlnu.AFsBleCtu	KE55K	KKE444	E
	6v6vELKLK 6v6v6v53E3K6634	32nSCInaISiaDlemnStaE	27E	HI2y.p32 27E	temaset Suninu.AFsBleCtu	K4E44	KKK45E	E
4	51255515151CKLK5151CKLK74	ICielaDlemnStay5	IpyEE	HI2y.p32 IpyEE	temaset Sunlnu.AFsBleCtu	K 🕮 4	KEK #4	Е
E	644G445644EKLKB5yyH644E	ple mFB1FtuStaātmEna	pIg2⊡ II 4=14⊡	HI2y.p32 pIg2	temaset Sunlnu.AFsBleCtu	KK45K 🗆	K4EKK5	E
5	n4KEKLK4EKLKEKLKELE	2fFDmeFtStEAStuStBaDlemnSt	2F3n2F3nH4	H12y.p32 H4el4	temaset Suninu.AFsBieCtu		KK4K5	к 4
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	HEIg5LKLK740E32□	CElēmSsamlFt 1FmSetaStSmSFmSetaEsmelaāCACtSma□	i 🗆	HI2y.p32 r□	temaset Sunlnu.AFsBleCtu	KKKKK45	KK450	E
	SUKLKVALKLKESESUKLKy2g□ v355d5E5d5LKLK7K6pd6pd6pd6	r esSmlFmnaunuleBntF naLya.CACtSmaAnmFatSmesetulSF1r.esSmlFmnaunuleBntF naLy r rte Sme1 añFS De DEmnaFtuau SDe De Ste Sme1DntmFS De DEmnaStF naF rte Sme1 añFS	ppro Eporo #	HI2y.p32 ry3 HI2y.p32 pprpE	temaset Suninu.AFsBleCtu temaset Suninu AFsBleCtu	KEE5E 🗆	KEE44 □	E K
ĸ	65IPEKLKK44p25HE	Stn StISnaDlemnStarEStn StISnaDlemnSt	rD	HI2y.p32 EL	temaset Sunlnu.AFsBleCtu			к
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5	35yr36pkEEkLKEr6pkEE6pkE 64kkE64kkEkLKER6pkEE6pkE	l 2nlStn.mlñetStnDlemnStaStF naLæ (SmesētulSF1 aSttnlaini AlFtnaDlemnStaF4 □	DEL TATI	HI2y.p32 D4 HI2y.p32 D4	temaset Sunlnu.AFsBleCtu temaset Sunlnu AFsBleCtu	KKARD	KSKET	K F
	LKLKEKKEKEHKK	nClF1asn11aFuniSetaie1nsC1na4ISnaDlemnStplesn nuatnClF1asn11aFuniSetaie1nsC1na4	H4	HI2y.p32 H4	temaset Sunlnu.AFsBleCtu	KSK5	KK4K55K5	E
	LKLKK6467EKLKKH256EEEKI	$. I \ddot{a} u e i FSt \ddot{s} e tm FSt St Ba E i S1 \\ \ddot{a} D lem n St a E \ddot{a} u e i FSt \ddot{s} e tm FSt St Ba E i S1 \\ \ddot{a} D lem n St a E E \ddot{a} u e i FSt \\ \ddot{s} e tm FSt St Ba E i S1 \\ \ddot{a} D lem n St a E E \\ \ddot{a} u e i FSt \\ \ddot{s} e tm FSt \\ \dot{s} e tm \\ s$	IyIyEIy4	HI2y.p32 Iÿ□	temaset Sunlnu.AFsBleCtu			к
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К	K66KKKK4KEIKPEEEEPEEEEK	ämnleBntneC atCs1nFlalSAetCs1neDlemnSta⊡	p⊡_	HI2y.p32 p⊡	temaset Sunlnu.AFsBleCtu			к
4	644DKLKgEEDKLK#4DKLKgEpi	EUF1sieuC1Fmels4	4 211-4	HI2y.p32 4	temaset Sunlnu.AFsBleCtu	KK5E IE =	KK54EEE	E
ь П		veCA1ninlFtunuaDAStuStBaDlemnSta2mFCiitacie1eBa4	2114	HI2y.p32 2IL4	temaset Suninu.AFsBleCtu	KEKK4	K5KEK4	4
5	pEpELKLK#5p□	LIEDmStE	LepE	HI2y.p32 LEpE	temaset Sunlnu.AFsBleCtu		KK5554 🗆	4
	LKLK #g43yK64#E64#E2He7EKL	l EyäunüleBntF naCASCStetnä4aF1DEa CAseiD1nā/CACtSma□	y∐⊡ 2=1=	HI2y.p32 yII	temaset Sunlnu.AFsBleCtu	KEII	KK5E	E
	LKLK #gK # EK 35 yn 55E	s 4 ayra ni nteDiemnSt I stDiemnStam Sen malF 194	2p4	HI2y.p32 2p4	temaset Suninu.AFsBleCtu temaset Suninu AFsBleCtu	K455	K45F F4	K F
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4KI	KKE65EKLK#7EE	plemnF einaFsmSEmelaseiD1naCACtSmaE	p2E	HI2y.p32 p2E	temaset SunInu.AFsBleCtu	K444	KE4E5	E
4K4	4 41K3EKLK Eg46E56E56E5E Exemplify K Reference and the second	LIpünDntuntmaEañISsF nayy4 F2nDlemnF nelnBC1EmelñCACtSme4K3	yy4	HI2y.p32 yy4	temaset Sunlnu.AFsBleCtu	KK5KEE Kaeekko	KE4K4E	E
4K	LKLKEKL43KgIgEKL43KgEKEI	Luntle CssStFmna1En	Ly2	HI2y.p32 Ly2	temaset Sunlnu.AFsBleCtu	RHEEKK	KHE L	ĸ
4K	5 H564KDKK64KDKLKp64KDKL?	HpaSBtF1e einaseiD1naCACtSma4	gp24	HI2y.p32 gp24	temaset Sunlnu.AFsBleCtu	KK		4
4K	□pKyprEEKLK35EyEKLK prEKLk		HE4el□ 2n4	HI2y.p32 HE4el□	temaset Sunlnu.AFsBleCtu	KK555E	KKE4KEEE5	E
4K	□LKL766K5EKL3F2EKLKLK6□	2nlStn.mlnetStnDlemnStaStF nal	ар4 Ш	HI2y.p32 5p4	temaset Suninu.AFsBleCtu	K 404 🗆	K+BE	ĸ
44 H	CE7p5p5bELKLKLK6pHp55pp5□	ED4agIpF nEsmSEmStBaDlemnSta4	Ep4gLp	HI2y.p32 Ep4gLp	temaset Sunlnu.AFsBleCtu	KEEKEE□		4
445	656DKLKLK25LKLKLKH56E6DK	x gDlemnSt SBtF1StBaieuC1Fmela4	gp24	HI2y.p32 gp24	temaset Sunlnu.AFsBleCtu			K
44	GEGEKLKLKEEg□	2elmStBatnStZelmStBatnStamnliStF11aDlesn nu	2	HI2y.p32 H	temaset Suninu.AFsBleCtu	K4EKKKE 🗆	KEED	E
44	KG161616151KLKLK61E6161	nCletatFSBFmela4	D4	HI2y.p32 L4	temaset Sunlnu.AFsBleCtu		K444K4	4
4EF	X PEEK APEEK AECKLEOKLKLKU E 6e7 n KAK 6e7 n KAS6e7 n K6e7 n K6e	EplemnStSleF DFlmFmnGyE DFlmFmnJaiim EmlFt iilF nplemnStSleF DFlmFmnaiim EmlFt 71 A 168 tmplFemole4	1 pHG 1 3=4	HI2y.p32 pHI4	temaset Sunlnu.AFsBleCtu	K4KKK5K	KEEEE	E
4E	□LKLKH5yşy5yy5EKLKy2EKLKI	LgaDaCAEIS1ainiAnlaE	ppE	HI2y.p32 ppE	temaset Sunlnu.AFsBleCtu	KEEK	K4K4□	Ē
4E5	LKLKLK464EE64EEEH34645pE	j EESñaDlemnStEESñaDlemnStaEEESñaDlemnStaEESñaDlemnSta5	DEDDS	HI2y.p32 EE	temaset Sunlnu.AFsBleCtu	KEK45EK		4
4E0 4E0	□LKLKLKE6E6⊡ □LKLKLK2LFKLKH5va5m444m44	yygäneiFStsetmFStStBaDlemnSta4 Ine1SeSIC alnsnDmel	yyga p□	HI2y.p32 yyg4 HI2y.p32 p□	temaset Sunlnu.AFsBleCtu temaset Sunlnu AFsBleCtu	KEKKED KK55455	KKE# KKEKK□	E
4E	□ 64 EKLKLK2ELKLKy2pE64 64 Ep	üleEsIsentinaLaunüleBntF naiSmesētulSF1	Èy□	HI2y.p32 Ly□	temaset Sunlnu.AFsBleCtu	KEEEK	KEEKE	E
4E	BKK54EKLKLK2r	ēs īnaseiD1 nāseiDetntma⊡	H = 2=	HI2y.p32 H	temaset Sunlnu.AFsBleCtu	K55E5		4
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44	6K 5GK 5GK 5GK 5ELKLKLK2EKU	DSmiil SF1 auS seSuStaueiFStsetmFStStBalnsnDmela4	yy4	HI2y.p32 yy4	temaset Sunlnu.AFsBleCtu	K54K4□		4
4E	LKLK2E75pE55KEEKLKLKIpE5 64EE64EEKI KI KIP4	s r eEiniHeLaunüleBntFnäiSmesētulSF1 yn i mEGE5 Lündel alnu CamEn	ry: vHE5	HI2y.p32 ry	temaset Sunlnu.AFsBleCtu	K4EK45□	KK4K□	E K
45	pK4KpK4K87y4pK4KEpK4K5L	kLsIHeL&StuStBaDlemnSt	y3r	HI2y.p32 y3r	temaset Sunlnu.AFsBleCtu	K45E5□	K44KE	E
4□	6prE6prEELKLKLKI6prEHK5	nCmlF1 as E1 n mnle1 an mnlaule1F na4	H4	HI2y.p32 H4	temaset Sunlnu.AFsBleCtu	KE55	KE555E	E
4⊡ 4⊡	61E364#EKLKLKIK64#E6IE3	HC118tE nlemn\$ta1\$nBCFluaE	HE DrF	HI2y.p32 HE HI2y.p32 DeF	temaset Sunlnu AFsBleCtu temaset Sunlnu AFsBleCtu	KEK5K	KKK4KE	E F
4□	6363635LKLKLK165 @Eg5HE 1070	nDC1 SnaBCSuFtsnaie1nsC1naL	 gD	HI2y.p32 gD	temaset Sunlnu.AFsBleCtu	KEEK5KK5	K4KK	Ē
4□	pEKLKLK16gpE	ayFaiSmesētulSF1aDlemne1SDSu	pH45elE	HI2y.p32 p	temaset Sunlnu.AFsBleCtu			ĸ
451	SONELKLKLEGE⊡ SEIKIKIEGE	leteInBC1884 nKEaDlemnStaHelnCmFmSnaDXStaStDaDlemnStaCetSm-F	HZLIYA Həlfif	H12y.p32 M± H12y.p32 H±1□	temaset Sunlnu.AFsBleCtu	KEK4EE	K4KK5□	E K
45E	p DKLKL56yrK EDKr4p4r4p En Sr	i HeletSt⊞eletStSmesëtulSF1aSiDelmaSttnlainiAlFtnamlFt 1esF na CACtSmaIr4⊡	HBp14p14	HI2y.p32 H	temaset Suninu.AFsBleCtu	K54 K4	KKK	E
455	p5EEKLK35EL5E	timeFs1HeLamSe1F naiSmesetulSF1	LHLLE	HI2y.p32 LHLLE	temaset Sunlnu.AFsBleCtu			ĸ
45 : 45 :	D P4 GK 5DK LK 35 EH FigE Fif4 EKHK	1 IIFT 1 FmSetF1 1 Setmle 1 1 nuamCielaDlemnSt 1 Ete mnle 1 a45F1 D Fauninm TF n	1p14 Hodi 4	HI2y.p32 IpI4	temaset Sunlnu.AFsBleCtu	KK4EED KKF4KD	KK 5440 44 maa	E
45	LKLKH5y46E55	IIFt iniAlFtnaaCDnlEiS1ainiAnla5	125	HI2y.p32 125	temaset Sunlnu.AFsBleCtu	KK4EE		4
45	64EELKLK5EKLK264EEKLKEI					KKEE 🗆	KEIE	E
45	I VI VIIS SECTOR	yS alFlBnačieleBa4	yg4	HI2y.p32 yg4	temaset Suninu.AFsBleCtu	and an an an and an		E
414	LKLKH5y7Ep4Ep4Ep HnKfiKLKH5ya4si	{yS älFlBnačieleBa4 LIECIEmF naLLIECIEmF naLaseiDetntma3LIECIEmF naLaseiDetntmaH 5X/alSAe eiFaDlemnSta21.	yg4 L2L n2L	HI2y.p32 yg4 HI2y.p32 L2L HI2y.p32 p21	temaset Sunhu.AFsBleCtu temaset Sunhu.AFsBleCtu temaset Sunhu AFsBleCtu	K4K5E□ K41044	KKK45	F
4K 44	□LKLKH5y7Ep4Ep4ED HpREKLKH5yg4p⊐ D68EDKLyH68DKLKH5ygE468555	§yS a1FIBnatie1eBa4 LUC(1EmF naLLUC(1EmF naLaseiDetntma3LUC(1EmF naLaseiDetntmaH 5K2alSAc eiF1aDlemnSta2L LA1aStmmlFmeIaE	yg4 1.21. p21. 1.3rE	HI2y.p32 yg4 HI2y.p32 L2L HI2y.p32 p2L HI2y.p32 L3rE	temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu	K4K5E□ K412044	KKK45□ K4KE□	E 4
4K 44 4E	□LKLKH5y7Ep4Ep4Ep H prK tikLKH5yg4p□ D68EEKLy168EKLKH5ygE4685p LKLKH5yg46pK6pKE6pK	sys är Fillmariel cBa4 LIC (TEM Fall LTC) Em F nal asciDetntma3ULC (TEM F nal asciDetntmaH 5K2afsAc eiff inDiemaSta2L LA laStmmFisnelatz HF IsSC (StrunDatuntmaDE DEI SDF nal EBFiif	yg4 1.21. p21. 1.3rE pp11.	H12y.p32 yg4 H12y.p32 L2L H12y.p32 p2L H12y.p32 L3rE H12y.p32 ppE	temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu	K4K5E□ K4⊞44 KK5554	K45E K4KE⊡ K4KE⊡	E 4 E
4K 4₫ 4E 4⊡	□LKLKH5y7Ep4Ep4Ep HpiKUKLKH5yg4g⊐ DobeELKLyF1GELKLKH5ygE4685p LKLKH5yg46pK6pKE6pK LKLKH5yg63D505655KH54K63□ P46FLKLKH5yg68D50555KH54K63□	sys är FBBanzieleBa4 LHC UEmF naLLEC (EmF naLaseüDetntmabUEC (EmF naLaseüDetntmaH SK2alSAc ei FlaDlemaSta2L LLAIaStmmFsmelaE HF1scS (SULDNItutmaDEDEI SDF naLEBFiif Zy4TSaaDlemaSt EHDel and LESS eindE me füSm====1821	уğ4 1.21. j21. 1.3гЕ ррПП 2y4 HrF	H12y.p32 yg4 H12y.p32 L2L H12y.p32 p2L H12y.p32 L3rE H12y.p32 ppD⊡ H12y.p32 2y4□ H12y.p32 H=F	temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu	K4K5E□ K41044 KK5554 KE5E□ KE52□	K43E KKK45□ K4KE□ KK45□ KK45□	E 4 E F
4K 44 4E 40 45 40	LKLKHSyTE#EB#ED HF#KUKLKHSyg49D DEBELKL#BGELKLKHSygE4635F LKLKHSyg66FE6665KKH5K63D B4EEKLKHSygE8EB6365KH5K63D B4EEKLKHSygE82F636516415670+F	yS ai FiBaaciel eBa4 LIC (TEM Fail LIC (TEM Fail asceiDetatma3ULC (TEMF nal.asciDetatmaH SK2alKAc eiFi JaDlemaSta2L LI AlaStimulFwiela& HF1sG (SRtunDatontmaDe Del SDF nal.EBFiiF 224 VSaaDlemaSt tetHE1auri n#SS einH Fail äiSmesetulSF1 Hys färlmSBM	уд4 1.21. 13-Е ррПП 23-4 HrE HyEП	H12y.p32 yg4 H12y.p32 L2L H12y.p32 p2L H12y.p32 p2L H12y.p32 p3 H12y.p32 2y4□ H12y.p32 2y4□ H12y.p32 HrE H12y.p32 HvE□	temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu	K4KSE= K42844 KESE= KESE= K42E5KKE	K4KE K4KE K4KE K4K5 K5K K455 K455 K455 K	E 4 E E E E
4K 44 4E 40 45 40 4K	LIKLKH57Ep4Ep4EP HERUEKLKH57ef4E DGSELKLYHGSLKLKH5ygE4GSSF LKLKH5yg46pKGpKE6pK LKLKH5yg6BEGGGKKH5KG LKLKH5ygKGBEGGEKH5KG LKLKH5ygKGP5G7pG7pG4KG LKLKH5ygKGEGEEEEG6KGG	sys är FiBanciel cBa4 LIC (TEF m Fall LTC (TEF Fall aseiDetntmäd LIE C1EmF nal.aseiDetntmäH SK2alSAc eiF laDlemaSiraL LI AlaSimalFønenlaE HF1 sSCSStunDutuntmaDEDeI SDF nal.EBFiiF 2y4TSimDemaSt EHTLe Jaun I mFaS einfF mE äiSmeseitulSF1 HyE äFimSbat Lägipf miSms Stms/BaDlemaStagr14	уğ4 L2L D2L L3FE D5D 224 H7E HyE grl4	HI2y,p32 yg4 HI2y,p32 L2L HI2y,p32 L2L HI2y,p32 L3FE HI2y,p32 L3FE HI2y,p32 2y4□ HI2y,p32 HyE□ HI2y,p32 HyE□ HI2y,p32 HyE□	temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu temaset Sunlnu.AFsBleCtu	K4K5ED K4E44 KK5554 KE5ED KK5ED K4EE5KKE	KKK450 K4KE0 K4KE0 K4450 KK4450 K45E4E0 K45E4E0	E 4 E E E K
4K 44 4E 40 45 40 4K 44 44	LKLKH5yZEP4Ep4EE HFKLKHK15yZ4GZ LG3ELKL¥H5yZ4GZ LKLKH5yZ4GPKGFKEGFK LKLKH5yZ6GPEG5GKH5KGS LKLKH5yZ6GPEG5GZ LKLKH5yZ6GPEG5G2G LKLKH5yZ6GEGGE12GrKEGG GEIKLKH5yZ6GGC GEIKLKH5yZ6GGC	sys är Filbanätel cBa4 LIC (TEm Faul LIC (TEm Faul aseciDetatumä3UE C1EmF nal.aseiDetatumäH 5K2alSAe ciF1 aDlemnSta2L LA Li StmmFsmelaE HF1sc (Stunbuttumtan3DeDe1SDF nal.EBFiiF 2y4tSmaDtennSt CHITeLaun InFaS cinF mä äiSmeseituISF1 Hy5 äF1mSBat Lägipt näSmsEms(BaDlemnStagr14 HineOIF i SsaDe DEmSG18te Sne1 amFit ilaDlemnSta4 UEDDStrie i III co 4 CcE mod	yğ4 121. p21. L3rE ppD□ 2340 HrE HyE□ grl4 pripH4 U=24	HB2yp32 yg4 HI2yp32 L2L HI2yp32 L2L HI2yp32 p2L HI2yp32 l3rE HI2yp32 2y4□ HI2yp32 2y4□ HI2yp32 HrE HI2yp32 gr14 HI2yp32 gr14 HI2yp32 gr14 HI2yp32 pr1pH4	temaset Sunlm. AF BilleCtu temaset Sunlm. AF BilleCtu	K41K5ED K41944 KK5554 KE5ED K41E5KKE K41K5D K41K5D	KKK45 KKK45 K4K0 KK445 KK445 KK45E K45E4E	E E E E K 4 F
4K 44 40 45 40 4K 44 44 42 40	LIKLKISYZEPIEDEE HARIKKIKISYZEPIEDE HARIKKIKISYEL LIKLKISYSEDEGGGKKIKISKE LIKLKISYSEDEGGGKKIKISKE LIKLKISYSECDEGGEGKKIST LIKLKISYSECDEGEGEGEGGKKIGGI GELKKIKISYSEGKECE DEEKKIKISYSEG	yS ai FiBanciel eBa4 LIC (TEP na LLE) CTE ni na LasceiDetntmaJLIE CTEmF na LasciDetntmaH Sk2afAck eiFlaDlemsKraL LA LAStmniFwrelaE HF1sGCSRunDhotntmaDEDEJSDF naLEBFiiF 2y4ISanDbensKt fetHeLanol ni Se einiF mär äismesetuISF1 Hy5 äirMSBnt Läglpf näsmsSms/BBdDemsKragri4 HHEDFF SisaDEDmsSLtSE sen ei amlift ilaDlemsKra4 HHEDFF SisaDEDmsSLtSE sen ei amlift ilaDlemsKra4 HHEDFF SisaDEDmsSLtSE sen ei amlift ilaDlemsKra4	у§4 121. р21. L3rE ррС 2343 HrE HyE grl4 prfpH4 H1р24 E	HI2y.p32 yg4 HI2y.p32 L2L HI2y.p32 L2L HI2y.p32 J3F HI2y.p32 pfL HI2y.p32 gr4□ HI2y.p32 HFE HI2y.p32 HFE HI2y.p32 gr14 HI2y.p32 pr1pH4 HI2y.p32 FL	temaset Sunina_AFsBicCtu temaset Sunina_AFsBicCtu temaset Sunina_AFsBicCtu temaset Sunina_AFsBicCtu temaset Sunina_AFsBicCtu temaset Sunina_AFsBicCtu temaset Sunina_AFsBicCtu temaset Sunina_AFsBicCtu temaset Sunina_AFsBicCtu temaset Sunina_AFsBicCtu	K41K5ED K41944 KK5554 KE5ED K45ED K4EE5KKE K41KKSD K15D	KK54 KK545 KK545 KE5K K55K K55K K55K	E E E E K 4 E K
4K 44 4C 45 40 44 44 45 44 45 45	LKLKHSYZEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIE	sys är IPBanciel cBa4 LLT (LTP m Fall LLT CTP m Fall asseiDetatuma JLT CTEm Fall asseiDetatuma H SK2al SA4 eif FalDremsfrad L LLAtsfrum Fibrenela E HF Is SC StrumDatontuman DE Del SDF mal EBFiiF 2 yA1 SmaDlemsft Eif Helaman In FaS einf F mä äis meseitul SF1 Eilly Earlinn Stan Läglp fränsnör Läglp fränsnör Strast BaDlemsftagr 14 Hine Olt F IS sabe DEm Sul Ste Smel am IFF niaDlemsfta4 HTH POFStar II-La CAC Sma4 Steenniasi Stem CAC In Ge SSF munaDlemsft Stafie	y64 121. p21. L3rE p10- 2340 HFE HyE gr414 H1p24 E E y23	$\begin{array}{l} H2_{2}, p32 & yg4 \\ H12_{2}, p32 & [2L \\ H12_{3}, p32 & [2L \\ H12_{3}, p32 & [2L \\ H12_{3}, p32 & [p5D \\ H12_{3}, p32 & [p5D \\ H12_{3}, p32 & 2y4 \\ H12_{3}, p32 & HrE \\ H12_{3}, p32 & HrE \\ H12_{3}, p32 & grlpH1 \\ H12_{3}, p32 & grlpH1 \\ H12_{3}, p32 & [rlpH2 \\ H12_{3}, p32 & Hlp2 \\ H12_{3}, H12_{3}$	temaset Sunina AFsBicCtu temaset Sunina AFsBicCtu	K4KSE K4884 KESS4 KESE K4SE K4EESKKE K4EESKKE K4KKS K50	KKK 450 KKK 450 KAUK0 KK450 KK450 KK450 KA5E4E0 KE 50 KE 50 KE 544K0	E 4 E E E K 4 E K 4
4K 44 45 40 45 40 4K 44 45 40 45 40 40	LKLKHSyZEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIE	sys air Filemaich c Bea Lit C (The Final LIT) C Ther Final asceiDetintma&LIE C (TEm Final asceiDetintmaH \$K2al5Ac eif FinDfermaSta2L LA fastimulFsmelaß HYS & SixunDatuntmaDe Del SDF naLEBFiiF 2y4fSmBlemaSt EtHelaun InFaS einiF mE äiSmesetulSF1 HyE äFimSbat Lägipf nämsFms/BaDlemaStagrI4 HineDI F SsapE DEmsulSte SmelamFit änDlemaSta4 HIFIDFStai Fit La CAC(Sma4 sSteunläiSislem CAC(Tär & esSFmuaDlemaSt1StänE ynilEbBF Julia La CAC(Sma4 SteunläiSislem CAC(Tär & esSFmuaDlemaSt1StänE ynilEbBF Julia Cac(Sma4 SteunläiSislem CAC(Tär & esSFmuaDlemaSt2B) = fast1 SteunläiSislem CAC(Tär & esSFmuaDlemaSt2B) = fast2 SteunläiSislem CAC(Tär & esSFmuaDlemaSt2B) = fast2 SteunläiSislem CAC(Tär & esSFmuaDlemaSt2B) = fast2 SteunläiSislem CAC(Tär	y¥4 L2L. p2L. L3-rE ppE□ 2y40 H+FE HyE□ gr14 pr0pH4 HLp24 E y23 y23 y21	$\begin{array}{c} H12_{7}p_{32} & y_{33} \\ H12_{7}p_{32} & g_{21} \\ H12_{7}p_{32} & g_{21} \\ H12_{7}p_{32} & g_{32} \\ H12_{7}p_{32} & g_{32} \\ H12_{7}p_{32} & g_{32} \\ H12_{7}p_{32} & g_{41} \\ H12_{7}p_{32} & HrE \\ H12_{7}p_{32} & HrE \\ H12_{7}p_{32} & g_{71} \\ H12_{7}p_{32} & g_{7$	temaset Sunina-AFsBicCtu temaset Sunina-AFsBicCtu	KARSED KARSA KESSA KESED KASED KAKEN KSD KREKES KSP	NULLE KKK 450 KKK 450 KK4450 KK4450 KK450 KK55K0 KK55K0 KK55K0 KK55K0	E 4 E E E E K 4 E K 4 E F
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4K 44 45 40 45 44 45 40 45 40 40 40 40	LKLKHSYZEPIEPIEPIEP HØKEKLKHSYZEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIE	sys är Fillmaciet elBa4 LIC (TEIm Faul LEIT CFER Final asseiDetntmak LEIT CTEIm Final asseiDetntmak H Sk2alske eiFlaDlemstrad. LIAIstinnelFørnelafe HF1ssCtStrunDetnutmanDEDetSDF maLEBFiiF 24/StanDlemst FilleLannt nF2s einf Finzle äiSmeseitulSF1 Hyf FärfmSBmt Hip FärfmSBmt HillPDFStar Fill a CAC CIS ms4 HineOD FilssaDet DenSutIste SmellamfFi ilaDlemsSta4 HIPDFStar Fill a CAC CIS ms4 SteennlaisStem CAC Lind esSFmmanDlemsStIStäa yt lSsLBDE DeciSBe FisSHSmanDlemsStIStäa yt lSsLBDE DeciSBe FisSHStaDlemstata4 BDE DEISDSuaFsLIDT fill mp51LLS üClcabFSSL	y64 121. p21. L3-rE p010 23-40 HrE Hy50 gr14 pr1p144 H11p24 E E 23 y23 y23 y23 y23 y23 y23 y23 y23 y24 y23 y23 y23 y24 y23 y23 y23 y24 y23 y23 y24 y23 y23 y24 y23 y23 y24 y23 y24 y23 y24 y23 y23 y24 y24 y23 y24 y23 y23 y24 y23 y23 y24 y23 y24 y23 y23 y24 y23 y24 y23 y24 y24 y24 y23 y24 y23 y24 y23 y24 y23 y24 y23 y24 y23 y24 y23 y24 y24 y24 y24 y24 y24 y24 y24 y24 y24	$\begin{array}{c} H12_{7}p_{32} > y_{34} \\ H12_{7}p_{32} & p_{21} \\ H12_{7}p_{32} & p_{21} \\ H12_{7}p_{32} & p_{21} \\ H12_{7}p_{32} & p_{7} \\ H12_{7}p_{32} & p_{7} \\ H12_{7}p_{32} & p_{7} \\ H12_{7}p_{32} & g_{7} \\ H12_{7}p_{3} \\ H12_{7}$	temaset Sunina AFsBICCU temaset Sunina AFsBICCU	K4KSE K4KS K4KSS54 KKSS50 K4EESKKE K4KKSC K50 KKEKKES K45ED	KK K K KK K K KK K K KK K KE KE KE KE KE KE KE KE KE K	E 4 E E E E K 4 E E K K 7
4K 44 4E 45 40 4K 44 4E 45 40 40 40 40 40 40 40	LKLKHSyZEPIEPIEPIEP HØKKKLKHSYZEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIEPIE	sys är Fillmaide (EBa4 LIC (Eller final LIC (Eller final aseiDetntmak LIE C1EmF naLaseiDetntmak Sk2alsAc eiFlaDtemastraL LA LastimatFismelaE HF1scSCStanDatuntmaDEDe1SDF naLEBFiiF 2y4TSinDtemast eiHelaant mFaS einfF mE äiSmeseitulSF1 Liggfor fin3msEmstBaDlemastragr44 HmeD1F FssanDeDfmSul3te SmelamiFr tiaDtemaSta4 HF1DFSts inf1 La CAC (Sma4 SteennäisSlem CAC (Eller SeSFmunaDlemaStISñaE yet1StaD5De De1SBE FesSFbanDlemastBis E ImFr tilf maSägFa CAC (Sm 4) eriaemSafuzzianei Staet mStBablemastrag yet2StaD5De De1SBE FesSFbablemastBis E ImFr tilf maSägFa CAC (Sm 4) eriaemSafuzzianei Staet mStStBablemastraf 2DE DE1SDSuaFstImFr tilf majHL15 aC (CAnFSSt	y64 121. p21. 123.e p15 23.4 13.5 13.5 14.5 14.5 14.5 14.5 14.5 15 16	H12x p32 y64 H12x p32 k2L H12x p32 k2L H12x p32 k2L H12x p33 p5D H12x p33 p5D H12x p33 p5D H12x p33 p5D H12x p33 k2K H12x p32 k4K H12x p32 k4K H12x p32 k4K H12x p32 k2K H12x p32 k2K H12x p32 k2K H12x p32 k2K H12x p32 k2K H12x p32 k1L H12x	temaset Sunina-AFsBicCtu temaset Sunina-AFsBicCtu	KKSEC KHII4 KKSSS KESEC KSSC KSEC KAELSKE KSC KSC KSEC KASEC	KESC KKK5C KARC KARC KESC KESC KESC KESC KESSAC KESSAC KESSAC KESSAC	E 4 E E E E E K 4 E K 4 E E K K E E K 4 E E K 4 E E K 4 E E K 4 E E E K 4 E E E K 4 E E E E
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5	weights (kDa). The inactive proform of the CTSD enzyme (~52-kDa) contains a pro-peptide
6	(orange) that is cleaved off during maturation, yielding in an active single chain form (~48-kDa).
7	Additionally, mannose residues are attached at two asparagine residues, at position N134, and
8	N263, respectively, through N-glycosylation. The active mature form of CTSD consists of a light
9	chain (~14-kDa) and a heavy chain (~34-kDa). (B) Representative Coomassie Brilliant Blue
10	(CBB)-stained SDS-PAGE-gel of recombinant proCTSD after purification and size exclusion
11	chromatography in indicated elution fractions. (C) CTSD activity of recombinant proCTSD after
12	combining all fractions positive for proCTSD. To measure enzymatic activity assay, the pH was
13	adjusted to pH 4.5 and aspartyl protease inhibitor pepstatin A (PepA) was added to examine
14	specificity of assay and enzymatic activity of recombinant enzyme ($n = 3$ independently purified
15	rHsCTSD). (D) Representative immunoblot of CTSD deficient H4 cells treated with 20 µg/ml
16	rHsCTSD from 30 min to 72 h showing pro- (~ 52-kDa), single chain (~ 48-kDa) and mature (~ 34-
17	kDa) CTSD forms. Additionally, to show all bands that appear as mature CTSD, a higher exposed
18	image is shown. An unspecific band detected by CTSD antibody was present in all samples and
19	labeled with an asterisk (*). (E) Quantification of single chain and mature CTSD forms analyzed
20	together as active CTSD in H4 cells deficient in CTSD (CTSD KO) treated with 20 μ g/ml
21	rHsCTSD for 24, 48 and 72 h. Active CTSD was normalized to TUBB3 and expressed as fold
22	change $(n = 3)$. (F) Representative western blot analysis of lysate and lysosomal-enriched fraction
23	of H4 CTSD KO cells treated with 20 μ g/mL rHsCTSD for 72 h. CTSD pro- and single chain forms
24	(~52-kDa/48-kDa) as well as the mature CTSD (~34-kDa) are enriched in the lysosomal fraction.
25	GAPDH and CBB-stained SDS-Page-gel were shown as loading controls. (G) Quantification of
26	CTSD activity of H4 CTSD KO enriched lysosomal fractions incubated with 20 μ g/mL rHsCTSD
27	for 72 h. Data shows CTSD activity in relative fluorescence units (RFU) ($n = 3$ independent
28	lysosome-enrichment fractionations) (H) ToxiLight cytotoxicity assay of H4 CTSD KO cells

treated with 20, 40, 60, 80 and 100 µg/ml rHsCTSD for 72 h, expressed as fold change (measured 29 30 in triplicates for three independent experiments, n = 3). (I) Representative immunofluorescence images of H4 CTSD KO cells treated with 20 and 100 µg/mL rHsCTSD for 72 h and stained for 31 32 cleaved CASP3 (red) as an apoptotic marker. DAPI (blue) was used to stain the nucleus. As a positive control H4 CTSD KO cells were incubated with 500 µM H₂O₂ for 4 h. Scale bar: 20 µm. 33 34 Data represent mean \pm SEM. Statistical analyses were performed by using a two-tailed unpaired 35 Student's t-test for (C and G) and a one-way ANOVA together with a Dunnett's multiple 36 comparison test (E and H). Statistical differences are shown toward the PBS treatment (E) and toward positive control for ToxiLight cytotoxicity assay (H). ****p < 0.0001, ***p < 0.001, *p < 37 38 0.05.



Figure S2. Functional analysis of rHsCTSD *in vitro* and in H4 cells overexpressing WT SNCA. (A) Representative CBB-stained SDS-PAGE-gel showing ability of purified enzyme to process recombinant monomeric SNCA. 20 μ M SNCA was incubated with 0.3 μ M rHsCTSD for 1 h and 24 h at pH 7.4 and pH 4.5 (lysosomal pH). (B) Representative CBB-stained SDS-PAGE gel illustrating the capacity of rHsCTSD to cleave and process monomeric as well as oligomeric forms (partly) after 24 h of digesting by rHsCTSD (0.3 μ M). Fibrillary forms of SNCA were not processed by rHsCTSD after 24 h. (C) Representative negative stain TEM pictures of SNCA fibrils

48	after incubation with and without rHsCTSD after 24 h. Scale bar: 100 nm. (D) LDH cytotoxicity
49	assay and (E) ToxiLight cytotoxicity assay of H4 cells (SNCA overexpression) treated with
50	different concentrations of rHsCTSD (20, 40, 60, 80, 100 μ g/mL), expressed as fold change
51	(measured in triplicates of three independent experiments, $n = 3$). (F) Immunofluorescence images
52	of H4 cells (SNCA overexpression) incubated with 20 and 100 $\mu g/mL$ rHsCTSD for 72 h and
53	stained for cleaved CASP3 (red) as an apoptotic marker. DAPI (blue) was used to stain the nucleus.
54	As a positive control, H4 cells were incubated with 500 μ M H ₂ O ₂ for 4h. Scale bar: 20 μ m. (G)
55	Representative western blot of H4 cells (SNCA overexpression) treated with PBS, 20 $\mu g/mL$
56	rHsCTSD and 20 $\mu g/mL$ rHsCTSD together with 5 μM Leupeptin (LeuP) for 72 h and stained for
57	SNCA (C-20 and syn-1) and SAPC. GAPDH signal and CBB-stained SDS-gel were used as
58	loading controls. (H) Quantification of SNCA (syn-1 and C-20) and SAPC signal were normalized
59	to GAPDH and expressed as fold change (n = 3). All data represent mean \pm SEM. Statistical
60	analyses were performed by one-way ANOVA together with Dunnett's multiple comparison test
61	with statistical differences toward the positive control (D and E) and PBS for (H). **** $p < 0.0001$,
62	**p < 0.01, *p < 0.05.



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Figure S3. Effects of SNCA on endo-lysosomal and autophagic system in H4 cells overexpressing SNCA. Representative western blot of H4 cells incubated with Dox to decrease SNCA expression and PBS as an untreated control for 72 h. Signal for SNCA was detected by syn-1 and as autophagic markers, LAMP1 and SQSTM1 were used. Corresponding quantifications of LAMP1 and SQSTM1 are shown in Figure 3D. GAPDH and CBB stained SDS-PAGE were used as loading controls.



Figure S4. Characterization and functional analyses of DA-iPSn A53T mutant and isogenic control 74 75 (A53T corr.) lines. (A) Representative immunofluorescence images of DA-iPSn A53T and A53T 76 corr. using SNCA antibody LB509 (red) and dopaminergic marker tyrosine hydroxylase (TH; green) (n = 7 individual neurons per group). Scale bar: $10 \mu m$. (B) LDH cytotoxicity assay of A53T 77 78 corr. and mutant treated with and without rHsCTSD for 25 days (measured in triplicates of five 79 independent treatments, n = 5), shown as fold change compared to the positive control. Positive 80 control was provided by the Kit. (C) Quantification of neurofilament release to determine neuronal 81 viability and early cell death in DA neurons performed by an in-cell western. Signals were 82 normalized to cell stain (Celltag700) and are displayed as fold change compared to A53T PBS 83 (measured in triplicates of four independent experiments, n = 4). (D) Representative 84 immunofluorescence pictures of DA-iPSn A53T corr. and A53T mutant with and without 85 rHsCTSD stained for cleaved CASP3. Scale bar: 100 µm. (E) Quantification of soluble SNCA by 86 syn-1 antibody (corresponding western blot is shown in Figure 4D), normalized to GAPDH and 87 expressed as fold change (n = 3). (F) Full size western blot membrane from T-sol (Triton-soluble) 88 lysates of DA-iPSn A53T PBS and rHsCTSD treated cells. Left, Membrane was stained with the 89 SNCA C-20 antibody, which was detected by the infrared channel 800. TUBB3 and its unspecific 90 protein signal, which is marked with an asterisk (*), is also shown. Right, the same full size western 91 blot stained with SNCA antibody syn-1 and detected by the infrared channel 680. (G) Full western 92 blot membrane from T-insol (Triton-insoluble) lysates of DA-iPSn A53T treated with PBS or 93 rHsCTSD stained with SNCA antibody C-20. (H) Representative immunoblot of T-sol lysates of 94 A53T corr. DA-iPSn, stained for CTSD (pro- and single chain forms and mature forms) and SNCA 95 (C-20 and syn-1 antibody). GAPDH, TUBB3 and CBB-stained SDS-gels were used as loading 96 control. (I) Quantification of pro-/single chain form and mature CTSD signal intensities, 97 normalized to GAPDH and expressed as fold change (n=3) (J) Representative immunoblot of T-

98 insol lysates of A53T corr. DA-iPSn, indicating the absence of SNCA signal in the corrected line. 99 TUBB3 and CBB-stained SDS-gels were used to control for equal protein loading (n = 3). (K) Quantification of active electrodes and (L) weighted mean firing rate (Hz) in A53T corr. and A53T 100 101 before treatment (n = 16, each point represents one active well of a 48-well MEA plate). Green 102 labeled graphs were exposed to rHsCTSD at day 44 of DA neuronal differentiation (Figure 4 J and 103 K). Data represent mean ± SEM. Statistical analyses were performed by one-way ANOVA together 104 with Dunnett's multiple comparison test (A and B). Statistical differences are shown in (B) toward 105 positive control and in (C) towards A53T mutant line with PBS. A two-tailed unpaired Student's 106 t-test was used for (E) and (I) and one-way ANOVA together with a Tukey's comparison test for (K and L). ****p < 0.0001, **p < 0.01; n.s., not significant. 107

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111 Figure S5. Characterization of lysosomes and autophagic vesicles from DA-iPSn harboring SNCA mutation (A53T). (A) Corresponding recombinant controls of first round in vitro SNCA seeding 112 113 assay of DA-iPSn-derived lysosomal enriched fraction shown in Figure 5A. As negative controls 114 purified recombinant SNCA monomers were incubated alone (mono only) and with SNCA fibril 115 alone (fibril only) and as a positive control SNCA monomer was incubated together with SNCA 116 fibril (fibril+mono). Graph shows relative Thio T signal normalized to positive control (fibril+ a-117 syn mono) (n = 3). (**B**) Analyses of two rounds of *in vitro* SNCA seeding assay of DA-iPSn A53T 118 corr. and A43T mutant with rHsCTSD and without (PBS)-derived lysosomal enriched fraction and 119 corresponding controls (n = 3). (C) TEM pictures of ultrastructural analysis of DA iPSn A53T corr. 120 corresponding to Figure 5H. Blue: nucleus, yellow arrows: intracellular vesicles. Exemplary 121 mitochondria is depicted as an orange star. Scale bars: 25 µm. (D) Quantification of lysosomal 122 GBA activity in DA-iPSn A53T corr., A53T mutant and mutant treated with rHsCTSD for 21 days 123 expressed as fold change (n = 3). Data represent mean \pm SEM. Statistical analyses were performed 124 by using two-way ANOVA with a Tukey's multiple comparison test (A and B). To highlight

125 statistical differences asterisks (*) were used for A53T corr. vs. A53T and mono only vs. 126 fibril+mono. Diamonds (#) were used for A53T vs. A53T +rHsCTSD and fibril+mono vs. fibril 127 only. For (D) statistical analysis were performed by one-way ANOVA together with a Dunnett's 128 multiple comparison test with statistical differences toward A53T corr.. ****/####p < 0.0001, 129 **/##p < 0.01, */#p < 0.05; n.s., not significant.



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132 Figure S6. Analysis of SNCA levels within *ctsd* KO animals after treatment with rHsCTSD. (A) 133 Cartoon illustrating the time line of intracranial injection (i.c.) time points of 100 µg rHsCTSD (P1 134 and P19) as well as time point of mice perfusion and sample acquisition (P23). (B) Representative 135 immunoblot of T-sol and (C) T-insol from whole brain lysates extracted from wildtype (WT) and 136 ctsd KO and ctsd KO mice i.c. injected with rHsCTSD (KO +rHsCTSD). (D) Respective 137 quantification of T-sol (left) and T-insol (right) SNCA levels detected by syn-1 antibody, 138 normalized to TUBB3 and expressed as fold change (n = 4 mice per group). (E) Representative dot 139 blot from T-sol whole brain lysates of WT, ctsd KO and ctsd KO mice i.c. injected with rHsCTSD. 140 Total SNCA was detected by syn-1 antibody and total protein stain (DirectBue71) was used to 141 ensure equal protein load. (F) Representative dot blot of T-insol fraction of whole brain lysates of 142 WT, ctsd KO and ctsd KO mice treated with rHsCTSD. SNCA was detected by syn505 antibody 143 and total protein staining was used as loading control. (G) Respective quantification of T-sol SNCA

by syn-1 (left) and pathological T-insol SNCA by syn505 (right). SNCA signals were normalized to total protein and expressed as fold change (n = 4 mice per group), compared to WT. (H) Schematic representation of the mouse brain in a sagittal cut. The region marked by the red square shows where the confocal images were taken. Data represent mean \pm SEM. Statistical analyses were performed by one-way ANOVA together with Tukey's multiple comparison test. *** p < 0.001, **p < 0.01, *p < 0.05, n.s., not significant.





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152 Figure S7. Immunofluorescence images of WT mouse brain and *ctsd* KO mouse brain treated with 153 PBS or rHsCTSD and stained for SNCA. (A) Representative confocal microscopy images of WT 154 mouse brain treated with PBS. Pathology-associated SNCA was detected by the conformation-155 specific antibody MJFR-14-6-4-2 (red) and co-stained for neurofilament (green). The following 156 brain regions are shown: Hypothalamus (HT), fornix (FX), basal forebrain/thalamus (BF/Thal), 157 hippocampus (HC) and midbrain (MB). OT indicate the optical tract and CC the corpus callosum. 158 The white dotted line indicates boundary between regions. The region marked by the white square 159 in 20x is magnified in 40x. Images were taken with the 20x (scale bar: 100μ m) and 40x (scale bar: 160 50 µm) magnifications. (B) Representative confocal microscopy images of mouse brain from *ctsd* 161 KO treated with PBS or 100 µg of rHsCTSD. The corresponding higher magnified (40x) images 162 were marked by the white square and are shown in Figure 6F. Images were taken with the 20x 163 (scale bar: 100 µm). The same antibodies and brain regions were used as shown in Figure 6F and 164 S7A. (C) Representative immunofluorescence images *ctsd* KO mice treated with PBS and (D) with 165 rHsCTSD stained with MJFR-14-6-4-2 (red) and co-stained for syn-1 (green). The images are 20x 166 magnified (scale bar: 100 µm) and the marked white square indicate the 40x magnification (scale 167 bar: 50 µm). Immunofluorescence images show the hypothalamus (HT), fornix (FX) and 168 hippocampus (HC) of the mouse brain. The white dotted line indicates boundary between regions 169 and OT stands for optical tract whereas CC for the corpus callosum. (E) Representative 170 immunofluorescence microscopy images of ctsd KO hippocampus (HC), fornix (FX) and midbrain 171 (MB) mouse brain areas i.c. injected with PBS or rHsCTSD. Brain areas are stained for SNCA by 172 Acris antibody (red) and neurofil. (green). Mouse brain images were taken with a 20x 173 magnification (scale bar: 100 µm). (F) Corresponding immunofluorescence stainings of primary 174 neurons from WT, Het, KO and KO +rHsCTSD animals are shown in Figure 6L (merged). SNCA 175 (syn-1) is shown in red, CTSD in green and nucleus in blue. Scale bar: $10 \,\mu m$.