

Table S1. The time-line of the pull-down method

Day	Task	time
1	Harvest lymphoblastoid cells or blood drawing	30 min
	↓	
	Extraction of total RNA	45 min
	↓	
	cDNA synthesis by reverse transcription	110 min
	↓	
	The process of the pull-down method (details in Methods)	125 min
	01. Preparation of the annealing mixture containing cDNA (100 ng) and biotinylated RNA probe (100 pmol)	5 min
	↓	
	02. Heat-denaturation at 95 °C for 5 min	5 min
	↓	
	03. Annealing at room temperature	30 min
	↓	
04. Add streptavidin-resin to the mixture	5 min	
↓		
05. Incubation at 37 °C with gentle agitation	60 min	
↓		
06. Wash the resin with wash buffer followed by centrifugation 3 times	10 min	
↓		
07. Elution and collection of cDNA fragments from the resin by 5 mM biotin solution	10 min	
↓		
SNP typing by the TaqMan SNP Genotyping assay	120 min	
Examination of trinucleotide repeat sequences by PCR and gel electrophoresis		



Table S3: Genotype of the cSNPs in the *HTT* gene in HD patients and healthy individuals

SNP ID	Exon	Alleles	Heterozygosity <sup>a)</sup>	HD patients							Healthy individuals	
				C0142	C0160	C0166	C0221	C0222	HD995	N0001	N0002	
rs363099	29	C/T	0.333	C/T	T/T	C/C	C/T	T/T	T/T	C/C	C/T	
rs362331	50	C/T	0.494	C/T	C/C	C/T	C/C	C/C	C/C	T/T	C/T	
rs362273	57	A/G	0.370	A/G	G/G	A/A	A/G	G/G	G/G	A/A	A/G	
rs362272	61	A/G	0.374	A/G	A/A	G/G	A/G	A/A	A/A	G/G	A/G	

<sup>a)</sup> The heterozygosities are derived from the NCBI Entrez dbSNP database: <http://www.ncbi.nlm.nih.gov/snp>.

Table S4. Synthetic siRNA sequences targeting cSNP sites (1/2)

SNP ID (alleles)	siRNA name		5' - sequence - 3' <sup>(b)</sup>	
rs363099 (C/T)	siRNAs targeting the C-allele			
	siRs099_C9	ss <sup>a)</sup>	GGGUUUCU <u>C</u> CGCUCAGCCUUU	
		as	AGGCUGAGCGGAGAAACCCUU	
	siRs099_C10	ss	AGGGUUUCU <u>C</u> CGCUCAGCCUU	
		as	GGCUGAGCGGAGAAACCCUUU	
	siRs099_C11	ss	GAGGGUUUCU <u>C</u> CGCUCAGCUU	
		as	GCUGAGCGGAGAAACCCUCUU	
	siRs099_C12	ss	GGAGGGUUUCU <u>C</u> CGCUCAGUU	
		as	CUGAGCGGAGAAACCCUCCUU	
	siRNAs targeting the T-allele			
	siRs099_T9	ss	GGGUUUCU <u>U</u> CGCUCAGCCUUU	
		as	AGGCUGAGCGAAGAAACCCUU	
	siRs099_T10	ss	AGGGUUUCU <u>U</u> CGCUCAGCCUU	
		as	GGCUGAGCGAAGAAACCCUUU	
	siRs099_T11	ss	GAGGGUUUCU <u>U</u> CGCUCAGCUU	
		as	GCUGAGCGAAGAAACCCUCUU	
	siRs099_T12	ss	GGAGGGUUUCU <u>U</u> CGCUCAGUU	
		as	CUGAGCGAAGAAACCCUCCUU	
	rs362331 (C/T)	siRNAs targeting the C-allele		
		siRs331_C8	ss	CCCUCAU <u>C</u> CACUGUGUGCAUU
			as	UGCACACAGUGGAUGAGGGUU
		siRs331_C9	ss	CCCUCAU <u>C</u> CACUGUGUGCAUU
			as	UGCACACAGUGGAUGAGGGUU
		siRs331_C10	ss	UCCCUCAU <u>C</u> CACUGUGUGCUU
as			GCACACAGUGGAUGAGGGAUU	
siRs331_C11		ss	CUCCCUCAU <u>C</u> CACUGUGUGUU	
		as	CACACAGUGGAUGAGGGAGUU	
siRs331_C12		ss	GCUCCCUCAU <u>C</u> CACUGUGUUU	
		as	ACACAGUGGAUGAGGGAGCUU	
siRNAs targeting the T-allele				
siRs331_T8		ss	CCCUCAU <u>U</u> CACUGUGUGCAUU	
		as	UGCACACAGUGAUGAGGGUU	
siRs331_T9		ss	CCCUCAU <u>U</u> CACUGUGUGCAUU	
		as	UGCACACAGUAGAUGAGGGUU	
siRs331_T10		ss	UCCCUCAU <u>U</u> CACUGUGUGCUU	
		as	GCACACAGUAGAUGAGGGAUU	
siRs331_T11		ss	CUCCCUCAU <u>U</u> CACUGUGUGUU	
		as	CACACAGUAGAUGAGGGAGUU	
siRs331_T12		ss	GCUCCCUCAU <u>U</u> CACUGUGUUU	
		as	ACACAGUAGAUGAGGGAGCUU	

Table S4. Synthetic siRNA sequences targeting cSNP sites (2/2)

SNP ID (alleles)	siRNA name		5' - sequence - 3' <sup>b)</sup>	
rs362273 (A/G)	siRNAs targeting the A-allele			
	siRs273_A8	ss	UGCUGCU <u>A</u> CAGAUCAACCCUU	
		as	GGGUUGAUCUG <u>U</u> AGCAGCAUU	
	siRs273_A9	ss	CUGCUGCU <u>A</u> CAGAUCAACCCUU	
		as	GGUUGAUCUG <u>U</u> AGCAGCAGUU	
	siRs273_A10	ss	GCUGCUGCU <u>A</u> CAGAUCAACUU	
		as	GUUGAUCUG <u>U</u> AGCAGCAGCUU	
	siRs273_A11	ss	AGCUGCUGCU <u>A</u> CAGAUCAAUU	
		as	UUGAUCUG <u>U</u> AGCAGCAGCUUU	
	siRs273_A12	ss	AAGCUGCUGCU <u>A</u> CAGAUCAUU	
		as	UGAUCUG <u>U</u> AGCAGCAGCUUUU	
	siRNAs targeting the G-allele			
	siRs273_G8	ss	UGCUGCU <u>G</u> CAGAUCAACCCUU	
		as	GGGUUGAUCUG <u>C</u> AGCAGCAUU	
	siRs273_G9	ss	CUGCUGCU <u>G</u> CAGAUCAACCCUU	
		as	GGUUGAUCUG <u>C</u> AGCAGCAGUU	
	siRs273_G10	ss	GCUGCUGCU <u>G</u> CAGAUCAACUU	
		as	GUUGAUCUG <u>C</u> AGCAGCAGCUU	
	siRs273_G11	ss	AGCUGCUGCU <u>G</u> CAGAUCAAUU	
		as	UUGAUCUG <u>C</u> AGCAGCAGCUUU	
	siRs273_G12	ss	AAGCUGCUGCU <u>G</u> CAGAUCAUU	
		as	UGAUCUG <u>C</u> AGCAGCAGCUUUU	
	rs362272 (A/G)	siRNAs targeting the A-allele		
		siRs272_A8	ss	GCACGGC <u>A</u> UCCUCUAUGUGUU
		as	CACAUAGAGGA <u>U</u> GCCGUGCUU	
siRs272_A9		ss	UGCACGGC <u>A</u> UCCUCUAUGUUU	
		as	ACAUAGAGGA <u>U</u> GCCGUGCAUU	
siRs272_A10		ss	CUGCACGGC <u>A</u> UCCUCUAUGUU	
		as	CAUAGAGGA <u>U</u> GCCGUGCAGUU	
siRs272_A11		ss	CCUGCACGGC <u>A</u> UCCUCUAUUU	
		as	AUAGAGGA <u>U</u> GCCGUGCAGGUU	
siRs272_A12		ss	CCCUGCACGGC <u>A</u> UCCUCUAUU	
		as	UAGAGGA <u>U</u> GCCGUGCAGGGUU	
siRNAs targeting the G-allele				
siRs272_G8		ss	GCACGGC <u>G</u> UCCUCUAUGUGUU	
		as	CACAUAGAGGA <u>C</u> GCCGUGCUU	
siRs272_G9		ss	UGCACGGC <u>G</u> UCCUCUAUGUUU	
		as	ACAUAGAGGA <u>C</u> GCCGUGCAUU	
siRs272_G10		ss	CUGCACGGC <u>G</u> UCCUCUAUGUU	
		as	CAUAGAGGA <u>C</u> GCCGUGCAGUU	
siRs272_G11		ss	CCUGCACGGC <u>G</u> UCCUCUAUUU	
		as	AUAGAGGA <u>C</u> GCCGUGCAGGUU	
siRs272_G12		ss	CCCUGCACGGC <u>G</u> UCCUCUAUU	
		as	UAGAGGA <u>C</u> GCCGUGCAGGGUU	

<sup>a)</sup> ss: sense-strand, as: antisense-strand.

<sup>b)</sup> The nucleotides corresponding to the SNP sites are underlined.

Table S5. Mismatched siRNA sequences

SNP ID (alleles)	SNP position	Mismatched position	siRNA name	5' - sequence - 3' <sup>b)</sup>
<b>rs363099 (C/T)</b>				
siRNAs targeting the C-allele				
	9	13	siRs099_C9(A13)	SS <sup>a)</sup> GGG <u>UUU</u> CU <u>CC</u> GGcA <u>CA</u> GC <u>UUU</u> AGG <u>CU</u> G <u>u</u> GC <u>GG</u> GA <u>AAAA</u> CC <u>UUU</u>
	9	14	siRs099_C9(G14)	SS GGG <u>UUU</u> CU <u>CC</u> GC <u>U</u> gA <u>GC</u> CU <u>UU</u> AGG <u>CU</u> cA <u>GC</u> GG <u>GA</u> AA <u>AA</u> CC <u>UUU</u>
	9	15	siRs099_C9(U15)	SS GGG <u>UUU</u> CU <u>CC</u> GC <u>U</u> C <u>u</u> GC <u>UUU</u> AGG <u>Ca</u> GA <u>G</u> CG <u>GA</u> AA <u>AA</u> CC <u>UUU</u>
	9	16	siRs099_C9(U16)	SS GGG <u>UUU</u> CU <u>CC</u> GC <u>U</u> CA <u>u</u> CC <u>UUU</u> AGG <u>u</u> GA <u>G</u> CG <u>GA</u> AA <u>AA</u> CC <u>UUU</u>
<b>rs362331 (C/T)</b>				
siRNAs targeting the C-allele				
	9	14	siRs331_C9(A14)	SS CC <u>CU</u> CA <u>U</u> CC <u>A</u> CU <u>Ga</u> GU <u>G</u> CA <u>UU</u> UG <u>CA</u> C <u>u</u> CA <u>G</u> U <u>GG</u> AU <u>G</u> AG <u>GG</u> UU
	9	15	siRs331_C9(C15)	SS CC <u>CU</u> CA <u>U</u> CC <u>A</u> CU <u>G</u> U <u>c</u> UG <u>CA</u> <u>UU</u> UG <u>CA</u> gA <u>CA</u> GU <u>GG</u> AU <u>G</u> AG <u>GG</u> UU
	11	14	siRs331_C11(A14)	SS CU <u>CC</u> CU <u>CA</u> U <u>CC</u> A <u>Ca</u> GU <u>GU</u> UU CA <u>CA</u> C <u>u</u> G <u>U</u> GG <u>AU</u> GAG <u>GG</u> A <u>GU</u> U
	11	15	siRs331_C11(C15)	SS CU <u>CC</u> CU <u>CA</u> U <u>CC</u> A <u>CU</u> c <u>U</u> GU <u>GU</u> CA <u>CA</u> gA <u>GU</u> GG <u>AU</u> GAG <u>GG</u> A <u>GU</u> U

<sup>a)</sup> ss: sense-strand, as: antisense-strand.

<sup>b)</sup> The nucleotides corresponding to the SNP sites and mismatched nucleotides are underlined and represented in small letters, respectively.

Table S6. IC50 values of the siRNAs examined in Fig. S6

SNP ID	siRNA name	IC50 (nM)		IC50 (nM)	
		Target allele ( <i>Renilla Luc.</i> )	Non-target allele ( <i>Photinus Luc.</i> )	Target allele ( <i>Photinus Luc.</i> )	Non-target allele ( <i>Renilla Luc.</i> )
rs363099	siRs099_T10	0.21 ± 0.081	>20	0.18 ± 0.107	>20
	siRs099_C9	0.20 ± 0.077	0.86 ± 0.171	0.12 ± 0.063	0.26 ± 0.055
	siRs099_C9(G14)	0.29 ± 0.048	>20	0.25 ± 0.043	>20
rs362331	siRs331_T9	0.07 ± 0.015	0.52 ± 0.431	0.05 ± 0.007	0.57 ± 0.125
	siRs331_T10	0.12 ± 0.032	7.63 ± 2.663	0.15 ± 0.074	0.98 ± 0.157
	siRs331_C9	0.48 ± 0.151	0.94 ± 1.685	0.18 ± 0.033	0.38 ± 0.044
	siRs331_C9(A14)	0.28 ± 0.066	8.19 ± 1.131	0.11 ± 0.032	>20
	siRs331_C9(C15)	0.14 ± 0.080	>20	0.08 ± 0.033	>20
rs362273	siRs273_G9	0.27 ± 0.032	>20	0.19 ± 0.048	>20
	siRs273_A9	0.26 ± 0.031	>20	0.38 ± 0.092	>20
rs362272	siRs272_G12	0.43 ± 0.141	>20	0.46 ± 0.148	>20
	siRs272_A9	0.53 ± 0.219	>20	0.71 ± 0.483	>20

Averaged IC50 values (mean ± S.D., n=4) are indicated.

Data fallen short of the half-maximal inhibition at the highest concentration of siRNA (20nM) is indicated by '>20'.

Table S7. Synthetic DNA oligonucleotides used in construction of reporter alleles

SNP ID (alleles, heterozygosity <sup>a)</sup> , exon)	Sequence name <sup>b)</sup>	5' - sequence - 3' <sup>c)</sup>
rs363099 (C/T, 0.333, exon 29)	C-allele	rs363099(C)-ss rs363099(C)-as ct agcatgCGTTTGGAGGGTTTCTCCGCTCAGCCTTGGATa ggcctATCCAAGGCTGAGCGGA <del>AA</del> AACCCCTCCAAA <del>C</del> gcatg
	T-allele	rs363099(T)-ss rs363099(T)-as ct agcatgCGTTTGGAGGGTTTCTTCGCTCAGCCTTGGATa ggcctATCCAAGGCTGAGCGGA <del>AA</del> AACCCCTCCAAA <del>C</del> gcatg
rs362331 (C/T, 0.494, exon 50)	C-allele	rs362331(C)-ss rs362331(C)-as ct agcatgCGCCTGCTCCCTCATCCACTGTGTGCACTTCAa ggcctTGAAGTGCACACAGTGGATGAGGGAAGCAGGCGcatg
	T-allele	rs362331(T)-ss rs362331(T)-as ct agcatgCGCCTGCTCCCTCATCTTACTGTGTGCACTTCAa ggcctTGAAGTGCACACAGT <del>TA</del> GATGAGGGAAGCAGGCGcatg
rs362273 (A/G, 0.370, exon 57)	A-allele	rs362273(A)-ss rs362273(A)-as ct agcatgCCGAGAAGCTGCTGT <del>TA</del> CAGATCAA <del>CC</del> CGAGA ggcctCTCGGGTTGATCTG <del>TA</del> GACAGCCTTCTCGgcatg
	G-allele	rs362273(G)-ss rs362273(G)-as ct agcatgCCGAGAAGCTGCTGTGCAGATCAA <del>CC</del> CGAGA ggcctCTCGGGGTTGATCTG <del>CA</del> GACAGCCTTCTCGgcatg
rs362272 (A/G, 0.374, exon 61)	A-allele	rs362272(A)-ss rs362272(A)-as ct agcatgCGGAGCCCTGCACGGCA <del>T</del> CCTTATGTGCTGga ggcctCCAAGCACATAGAGGA <del>AG</del> CCGTTGCAAGGCTCCgcatg
	G-allele	rs362272(G)-ss rs362272(G)-as ct agcatgCGGAGCCCTGCACGGCGT <del>C</del> CCTTATGTGCTGga ggcctCCAAGCACATAGAGGA <del>AG</del> CCGTTGCAAGGCTCCgcatg

<sup>a)</sup> Heterozygosities are derived from the NCBI SNP database: <http://www.ncbi.nlm.nih.gov/snp>.

<sup>b)</sup> 'ss' and 'as' fixed to the names represent the sense- and anti-sense strand sequences, respectively.

<sup>c)</sup> SNP sites and linker sequences are underlined and represented in small letters, respectively.