

Supplementary Material

Table S1. Spearman's rank correlation coefficient between the rhTPH2 genotype/haplotype and phenotypic data

Haplotype or Locus	Cortisol		ACTH		CORTRES		PERDEX		Self-biting rate	
	AM	PM	AM	PM	15 min	30 min	Night	Day	1999	2000
M2	-0.3901[#]	0.1672	-0.1124	0.1573	-0.0741	0.0519	0.5043[*]	0.3729	0.0119	0.2234
M3	-0.1308	-0.0077	-0.0667	-0.0501	0.5564^{**}	0.2120	0.0935	0.1870	0.0000	0.0133
M4	0.1994	0.1246	-0.1072	-0.1072	-0.0910	-0.0820	-0.5296[*]	-0.3841	-0.2921	-0.2104
M5	0.3309	0.0432	-0.0630	-0.1732	-0.0423	0.0181	-0.5705[*]	-0.3483	-0.0388	-0.0243
M5'	-0.2415	-0.4087	0.1602	0.1602	-0.0637	-0.2641	0.3027	0.2596	0.0000	-0.1098
M9	0.3539	0.4665[*]	-0.0696	-0.0870	-0.2851	-0.1322	0.3116	0.0104	0.1041	0.0699
-1605T>C	-0.3539	-0.4665[*]	0.0696	0.0870	0.2851	0.1322	-0.3116	-0.0104	-0.1041	-0.0699
-1491Tn	-0.3539	-0.4665[*]	0.0696	0.0870	0.2502	-0.0036	-0.3116	-0.0104	-0.1041	-0.1624
-1485(AT)n	-0.0062	-0.3009	0.0015	0.0573	0.2489	-0.0663	-0.6047^{**}	-0.2039	-0.3287	-0.1491
(AT)6	-0.0219	0.5319[*]	-0.0972	-0.0486	-0.2357	-0.0302	0.5608[*]	0.1645	0.1842	0.3764[#]
(AT)7	-0.1983	-0.2192	0.0225	0.0450	0.5007[*]	0.0911	-0.4105[#]	-0.1457	-0.2612	-0.1372
(AT)8	0.1489	-0.2830	0.0667	-0.0501	-0.0855	-0.1588	-0.4400[#]	-0.2348	-0.0401	-0.0982
-1454A>G	0.2415	0.4087[#]	-0.1602	-0.1602	0.0637	0.2641	-0.3027	-0.2596	0.0000	0.1098
-1325In>Del	-0.1014	0.2946	-0.0972	-0.0134	0.1616	0.1951	0.5548[*]	0.3364	0.0683	0.1321
-363T>G	-0.0562	0.1366	-0.0904	0.0000	0.2887	0.2037	-0.0948	-0.2933	0.1244	0.2425
2051A>C	0.6538^{**}	0.0385	-0.1501	-0.3503	0.3824[#]	0.4729[*]	-0.0877	-0.3839	-0.4068[#]	-0.3842[#]
2128S>L	-0.2727	-0.2258	0.0797	0.2347	-0.1068	-0.4552[*]	-0.5012[*]	-0.3009	0.2339	0.1877

* $p<0.05$, ** $p<0.01$, # $0.05 \leq p < 0.10$; CORTRES: cortisol response to ACH challenge; PERDEX: DEX suppression of cortisol excretion (%). For each haplotype or allele, the carrier and non-carrier were designated as "1" and "2", respectively, while for each locus the genotype was designated as "1", "2" and "3" for the wild-type homozygote, wild-mutant heterozygote and mutant-mutant homozygote, respectively. The 2051A>C genotypes were designated as "1" for AA&AC and "2" for CC.

Supplementary Material

Table S2. Distribution of the rhTPH2 5'-FR and 3'-UTR haplotypes in the cohort of 32 monkeys

Group	n	5'-FR haplotype										3'-UTR haplotype			
		M2	M3	M4	M4'	M5	M5'	M7	M9	N1	N2	N3	ACS	AAS	GAL
Overall	64	6 0.09	20 0.31	4 0.06	2 0.03	14 0.22	6 0.09	1 0.02	8 0.13	1 0.02	1 0.02	1 0.02	30 0.47	18 0.28	16 0.25
SW (1)	40	1	10	3	2	10	5	1	6	1	1	-	18	12	10
HFB (1a)	20	1	8	2	-	4	2	-	2	1	-	-	9	5	6
LFB (1b)	10	-	1	1	1	4	1	-	2	-	-	-	5	4	1
NW (2)	22	5*	10	-	-	3	1	-	2	-	-	1	11	5	6
HFB (2a)	4	1	1	-	-	1	-	-	1	-	-	-	-	1	3
LFB (2b)	14	3	7	-	-	2	-	-	1	-	-	1	9 [#]	3	2
HFB (a)	24	2	9	2	-	5	2	-	3	1	-	-	9	6	9 [#]
LFB (b)	24	3	8	1	1	6	1	-	3	-	-	1	14	7	3

Chi-Square test 5': 1, 2: $p=0.0326$; a, b: $p=1.0000$; 1a, 1b: $p=0.1804$; 2a, 2b: $p=0.7397$; 1a, 2a: $p=0.8749$; 1b, 2b: $p=0.0375$
(R×C) 3': 1, 2: $p=0.6248$; a, b: $p=0.1658$; 1a, 1b: $p=0.2218$; 2a, 2b: $p=0.1647$; 1a, 2a: $p=0.3029$; 1b, 2b: $p=0.3862$

SW: self-wounder; NW: non-wounder; HFB: high-frequency biter; LFB: low-frequency biter. * $p<0.05$ for SW vs NW,
$p<0.05$ for HFB vs LFB assessed for the individual haplotype (2×2 table). The incidence of each haplotype was given for the overall group. P values were given for the comparisons of haplotype distribution between SIB groups (R×C table). The Fisher's exact test and Mantel-Haenszel Chi-Square test were used for 2×2 and R×C tables, respectively.

Table S3. The effect of the rhTPH2 5' (-1485(AT)_n) and 3' (2015A>C) genotype and their interaction on the physiology and self-biting behavior in mother- and peer-reared monkeys

Variable	Genotype	Mother-Reared				Peer-Reared			
		Mean1 _(n)	Mean2 _(n)	F	p	Mean1 _(n)	Mean2 _(n)	F	p
HIAA	5'	33.7 ₍₅₎	49.5 ₍₅₎	4.82	0.070	43.4 ₍₂₎	45.6 ₍₆₎	0.03	0.867
	3'	49.9 ₍₃₎	33.3 ₍₇₎	5.31	0.061	46.0 ₍₄₎	42.4 ₍₄₎	0.11	0.755
	5'x3'	-	-	4.25	0.085	-	-	5.35	0.082
CORT-AM	5'	29.8 ₍₆₎	24.1 ₍₅₎	0.19	0.677	32.4 ₍₂₎	33.3 ₍₆₎	0.01	0.918
	3'	32.7 ₍₃₎	21.2 ₍₈₎	4.61	0.069	42.7 ₍₄₎	23.4 ₍₄₎	4.67	0.097
	5'x3'	-	-	0.94	0.364	-	-	0.02	0.895
CORT-PM	5'	12.7 ₍₆₎	21.3 ₍₅₎	7.79	0.027	14.2 ₍₂₎	18.5 ₍₆₎	0.89	0.400
	3'	20.2 ₍₃₎	13.7 ₍₈₎	2.86	0.134	14.6 ₍₄₎	18.0 ₍₄₎	0.54	0.502
	5'x3'	-	-	2.10	0.191	-	-	1.28	0.321
ACTH-AM	5'	71.2 ₍₅₎	66.7 ₍₅₎	0.07	0.805	68.8 ₍₂₎	60.6 ₍₆₎	0.38	0.571
	3'	69.6 ₍₃₎	68.3 ₍₇₎	0.0	0.948	55.6 ₍₄₎	73.9 ₍₄₎	1.88	0.243
	5'x3'	-	-	2.01	0.206	-	-	0.40	0.563
ACTH-PM	5'	65.9 ₍₅₎	63.8 ₍₅₎	0.15	0.711	64.2 ₍₂₎	49.6 ₍₆₎	1.40	0.303
	3'	63.8 ₍₃₎	65.9 ₍₇₎	0.04	0.842	43.6 ₍₄₎	70.2 ₍₄₎	4.64	0.098
	5'x3'	-	-	3.74	0.101	-	-	0.37	0.578
CORTRES15	5'	17.4 ₍₇₎	17.9 ₍₆₎	0	0.964	17.0 ₍₃₎	14.0 ₍₅₎	1.67	0.265
	3'	20.3 ₍₃₎	15.3 ₍₁₀₎	3.32	0.102	17.9 ₍₃₎	13.2 ₍₅₎	4.06	0.114
	5'x3'	-	-	0.28	0.607	-	-	1.66	0.268
CORTRES30	5'	14.1 ₍₇₎	12.5 ₍₆₎	0.32	0.583	12.4 ₍₃₎	11.9 ₍₅₎	0.04	0.852
	3'	15.3 ₍₃₎	11.3 ₍₁₀₎	2.24	0.169	14.2 ₍₃₎	10.2 ₍₅₎	2.16	0.215
	5'x3'	-	-	0.31	0.593	-	-	0.52	0.510
DEX-N	5'	56.4 ₍₇₎	93.8 ₍₄₎	25.2	0.001	30.0 ₍₂₎	64.5 ₍₄₎	1.60	0.333
	3'	46.0 ₍₂₎	75.3 ₍₉₎	3.14	0.114	46.8 ₍₃₎	47.8 ₍₃₎	0	0.974
	5'x3'	-	-	-	-	-	-	2.71	0.241
DEX-D	5'	62.9 ₍₇₎	80.3 ₍₄₎	0.16	0.703	24.0 ₍₂₎	41.0 ₍₄₎	0.91	0.442
	3'	35.2 ₍₂₎	76.8 ₍₉₎	3.92	0.083	23.5 ₍₃₎	41.5 ₍₃₎	1.02	0.420
	5'x3'	-	-	-	-	-	-	1.81	0.311
SB-1999	5'	0.050 ₍₆₎	0.094 ₍₅₎	0.22	0.652	0.060 ₍₂₎	0.088 ₍₄₎	0.22	0.683
	3'	0.020 ₍₂₎	0.081 ₍₉₎	0.72	0.422	0.023 ₍₃₎	0.125 ₍₃₎	3.10	0.221
	5'x3'	-	-	-	-	-	-	0.09	0.792
SB-2000	5'	0.068 ₍₇₎	0.105 ₍₆₎	0.11	0.746	0.018 ₍₃₎	0.125 ₍₅₎	1.53	0.284
	3'	0.025 ₍₃₎	0.148 ₍₁₀₎	1.23	0.296	0.030 ₍₃₎	0.113 ₍₅₎	0.89	0.398
	5'x3'	-	-	0.06	0.813	-	-	0.29	0.618

*Statistics were performed by ANOVA2. Mean1 represents the Tukey-Kramer least-squares mean for -1845(AT)6 carrier (5') or 2051CC (3') genotype, while Mean2 represents the Tukey-Kramer least-squares mean for -1845(AT)6 non-carrier (5') or 2051AA&AC (3'). The sample size (n) for each group was shown in parenthesis.