Appendix 1. PAML summary results for TRIM5a

A. PAML (1) Analysis of Entire Data Set (hominids+OWM+NWM)

1. Log Likelihood Scores and Parameter Estimates for Four Models of Variable ω's Among Sites Assuming the F3x4 Model of Codon Frequencies.

Model	Parameter Estimates	Sites* with $\omega > 1$	l
M0: one ratio	Average dN/dS for each branch = 1.160		-5772.38
	S = 1.55260		
Site models			
M1: neutral	$(\omega_0 = 0) f_0 = 0.354$	Not allowed	-5,725.00
	$(\omega_1 = 1) (f_1 = 0.646)$		
	Average dN/dS for each branch = 0.646		
	S = 1.56259		
M2: selection	$(\omega_0 = 0) f_0 = 0.281$	(47–51: gaps)	-5,667.01
	$(\omega_1 = 1) f_1 = 0.566$	175 Q 0.9958**	
	$\omega_2 = 4.624 \ (f_2 = 0.153)$	213 S 0.9635*	
	Average dN/dS for each branch = 1.273	215 T 0.9955**	
	S = 1.62074	228 L 0.9942**	
		310 C 0.9924**	
		311 A 0.9603*	
		317 K 0.9757*	
		324 K 0.9995**	
		325 P 0.9933**	
		(326-338: gaps)	
		339 F 0.9994**	
		340 V 0.9999**	
		(385-386: gaps)	
		389 K 0.9953**	
		407 G 0.9974**	
		418 F 0.9997**	
		421 P 0.9769*	
		423 V 0.9633*	
		471 Q 0.9535*	

Model	Parameter Estimates	Sites* with $\omega > 1$	l
		483 G 0.9936**	
M3:discrete	$\omega_0 = 0.342 f_0 = 0.662$	Too many to list.	-5,664.25
	$\omega_1 = 2.575 \ (f_1 = 0.305)$	Note: M3 vs. M0 is a	
	$\omega_2 = 7.922 \ (f_2 = 0.033)$	test of heterogeneity	
	Average dN/dS for each branch = 1.273	among sites and not an	
	S = 1.63377	explicit test of positive	
		selection.	
Μ7:β	$p = 0.02830 \ q = 0.01638$	Not allowed	-5,724.62
	Average dN/dS for each branch = 0.6160		
	S = 1.55777		
	0.0000 0.14001		
M8: β and ω	$p = 0.2299 \ q = 0.14801$	(47-51: gaps)	-5,666.71
	$f_0 = 0.829$	139 Q 0.9534*	
	$\omega_1 = 4.308 \ (f_1 = 0.171)$	175 Q 0.9967**	
	Average dN/dS for each branch = 1.2426	213 S 0.9739*	
	S = 1.62038	215 T 0.9966**	
		228 L 0.9958**	
		257 1 0.9511*	
		310 C 0.9943**	
		311 A 0.9/1/*	
		31 / K 0.9830*	
		324 K 0.99996**	
		325 P 0.9946**	
		(320-338: gaps)	
		339 F 0.9993**	
		340 0.9999	
		(385-386: gaps)	
		389 K 0 9964**	
		390 N 0 9611*	
		407 G 0 9979**	
		418 F 0 9998**	
		421 P 0 9830*	
		423 V 0 9737*	
		125 1 0.7757	

Model	Parameter Estimates	Sites* with $\omega > 1$	l
		471 Q 0.9670*	
		483 G 0.9952**	

p and *q* are parameters of the β distribution. *f* is the proportion of sites assigned to an individual ω (= dN/dS) category or to a β distribution with shape parameters *p* and *q*. The proportion $f_{1,2}$ in parentheses is not a free parameter. Sites assigned to $\omega_{1,2}$ are those with posterior probabilities (*P*) > 0.95, and those with *P* > 0.99 are in bold. Analyses were conducted using κ as a free parameter. S = tree length. *Sites assigned to ω > 1 are those with posterior probabilities *P* > 0.95 (*) or *P* > 0.99

(**).

2. Log Likelihood Scores and Parameter Estimates for Four Models of Variable ω's Among Sites Assuming the F61 Model of Codon Frequencies.

Model	Parameter estimates	Sites* with $\omega > 1$	l
M0: one ratio	Average dN/dS for each branch = 1.144		-5,748.06
	S = 1.55005		
Site models			
M1: neutral	$(\omega_0 = 0) f_0 = 0.359$	Not allowed	-5,696.78
	$(\omega_1 = 1) (f_1 = 0.641)$		
	Average dN/dS for each branch = 0.641		
	S = 1.56922		
M2: selection	$(\omega_0 = 0) f_0 = 0.291$	(47-51: gaps)	-5,641.84
	$(\omega_1 = 1) f_1 = 0.551$	175 Q 0.9954**	
	$\omega_2 = 4.451 \ (f_2 = 0.158)$	213 S 0.9544*	
	Average dN/dS for each branch = 1.2545	215 T 0.9970**	
	S = 1.62398	228 L 0.9957**	
		310 C 0.9930**	
		311 A 0.9669*	
		317 K 0.9736*	
		324 K 0.9994**	
		325 P 0.9903**	

Model	Parameter estimates	Sites* with $\omega > 1$	l
		(326-338: gaps)	
		339 F 0.9987**	
		340 V 0.9999**	
		(385-386: gaps)	
		389 K 0.9935**	
		407 G 0.9958**	
		411 S 0.9536*	
		418 F 0.9994**	
		421 P 0.9784*	
		423 V 0.9680*	
		483 G 0.9909**	
M3:discrete	$\omega_0 = 0.301 f_0 = 0.631$	Too many to list.	-5,639.09
	$\omega_1 = 2.379 \ (f_1 = 0.328)$	Note: M3 vs. M0 is a	
	$\omega_2 = 7.224 \ (f_2 = 0.041)$	test of heterogeneity	
	Average dN/dS for each branch = 1.2692	among sites and not an	
	S = 1.63693	explicit test of positive	
		selection.	
Μ7:β	$p = 0.029 \ q = 0.0167$	Not allowed	-5,696.39
	Average dN/dS for each branch = 0.6144		
	S = 1.56601		
M8: β and ω	$p = 0.16877 \ q = 0.11056$	(47-51: gaps)	-5,641.64
	$f_0 = 0.825$	139 Q 0.9532*	
	$\omega_1 = 4.190 \ (f_1 = 0.175)$	175 Q 0.9963**	
	Average dN/dS for each branch = 1.2331	213 S 0.9659*	
	S = 1.62300	215 T 0.9975**	
		228 L 0.9967**	
		257 T 0.9607*	
		310 C 0.9945**	
		311 A 0.9749*	
		317 K 0.9805*	
		324 K 0.9995**	
		325 P 0.9920**	
		(326-338: gaps)	
		339 F 0.9989**	
		340 V 0.9999**	
		(385-386: gaps)	

Model	Parameter estimates	Sites* with $\omega > 1$	l
		389 K 0.9949**	
		390 N 0.9587*	
		407 G 0.9966**	
		411 S 0.9653*	
		418 F 0.9995**	
		421 P 0.9832*	
		423 V 0.9756*	
		471 Q 0.9570*	
		483 G 0.9929**	

p and *q* are parameters of the β distribution. *f* is the proportion of sites assigned to an individual ω (= dN/dS) category or to a β distribution with shape parameters *p* and *q*. The proportion $f_{1,2}$ in parentheses is not a free parameter. Sites assigned to $\omega_{1,2}$ are those with posterior probabilities *P* > 0.95; those with *P* > 0.99 are in bold. Analyses were conducted by using κ as a free parameter. S = tree length.

Sites assigned to $\omega > 1$ are those with posterior probabilities P > 0.95 () or P > 0.99 (**).

3. Likelihood Ratio Test Statistics (2 δ) for Models of Variable Selective Pressure Among Sites.

	2δ	df	<i>P</i> value
Model 0 (one dN/dS ratio for all branches)			
F3x4			
One ratio vs. M3 $(k = 3)$	216.26	4	<i>P</i> < 0.0001
M1 vs. M2	115.98	2	<i>P</i> < 0.0001
M7 vs. M8	115.82	2	<i>P</i> < 0.0001
F61			
One ratio vs. M3 $(k = 3)$	217.94	4	<i>P</i> < 0.0001
M1 vs. M2	109.88	2	<i>P</i> < 0.0001
M7 vs. M8	109.50	2	<i>P</i> < 0.0001

4. Likelihood Ratio Test Statistics (2 δ) for Models of Variable Selective Pressure Along Branches.

	l	2δ	df*	P value
TRIM5				
Model 0 (same dN/dS ratio for all branches)	-5,784.326			
Model 1 (different dN/dS ratio for each branch)	-5,755.534	57.584	37	<i>P</i> < 0.02
Apobec3G [†]				
Model 0 (same dN/dS ratio for all branches)	-4,209.048			
Model 1 (different dN/dS ratio for each branch)	-4,201.337	15.422	20	P > 0.75 (N.S.)

*df denotes degrees of freedom that is equal to one less than the total number of branches in the phylogeny. For a 20-taxa tree like in TRIM5, the total number of branches is 38, so df = 37. N.S., not significant.

[†]Sawyer, S. L., Emerman, M. & Malik, H. S. (2004) *PLoS Biol.* **2**, e275.

5. Likelihood Ratio Test Statistics for Models of Selection Along the Human and Gibbon Lineages (OWM + hominids only).

	ω (dN/dS)					
	(human or	ω (dN/dS)				
	gibbon	(remainder				
	branch)	of tree)	l	2δ	df*	P value
Human						
Model 0 (two-ratio, human ω	1.00	1.132	-5,748.233			
=1)						
Model 1 [two-ratio, human $\omega > 1$	2.985	1.133	-5,747.543	1.38	1	0.24 (N.S)
(estimated by PAML; ref. 1)]						
Gibbon						
Model 0 (two-ratio, gibbon ω	1.00	1.087	-5,748.718			
=1)						
Model 1 [two-ratio, gibbon $\omega > 1$	5.594	1.091	-5,744.345	8.746	1	0.0031
(estimated by PAML; ref. 1)]						

df denotes degrees of freedom; here, df = 1.

6. *P* Value Statistics for Positive Selection Along the Human and Gibbon Lineages, Based on Monte Carlo Simulation.

	Confidence ancestral		
	sequence (0-1) per site	ω (dN/dS)	P value
Human vs. Ancestral Node	0.99971	3.6	$P < 10^{-4}$
Gibbon vs. Ancestral Node	0.99633	6.5	$P < 10^{-4}$

B. PAML (1) Analysis of Partial Data Set (hominids+OWM)

1. Log Likelihood Scores and Parameter Estimates for Four Models of Variable ω's Among Sites Assuming the F3x4 Model of Codon Frequencies.

Model	Parameter estimates	Sites* with $\omega > 1$	l
M0: one ratio	Average dN/dS for each branch = 1.1923		-3,519.244
Site models			
M1: neutral	$(\omega_0 = 0) f_0 = 0.307$	Not allowed	-3,513.813
	$(\omega_1 = 1) (f_1 = 0.693)$		
	Average dN/dS for each branch = 0.693		
M2: selection	$(\omega_0 = 0) f_0 = 0.185$	330 G 0.9960**	-3,492.320
	$(\omega_1 = 1) f_1 = 0.783$	332 R 0.9929**	
	$\omega_2 = 11.283 \ (f_2 = 0.032)$	335 R 0.9890*	
	Average dN/dS for each branch = 1.1429	337 Q 0.9981**	
		340 V 0.9993**	
M3:discrete	$\omega_0 \ 0.00001 \ f_0 = 0.380$	Too many to list.	-3,490.526
	$\omega_1 = 1.667 \ (f_1 = 0.601)$	Note: M3 vs. M0 is a	
	$\omega_2 = 16.866 \ (f_2 = 0.019)$	test of heterogeneity	
	average dN/dS for each branch = 1.3219	among sites and not	
		an explicit test of	
		positive selection.	
Μ7:β	$p = 0.00592 \ q = 0.00273$	Not allowed	-3,513.818
	Average dN/dS for each branch = 0.7		

Model	Parameter estimates	Sites* with $\omega > 1$	l
M8: β and ω	$p = 0.01037 \ q = 0.00248$	330 G 0.9959**	-3,492.324
	$f_0 = 0.968$	332 R 0.9927**	
	$\omega_1 = 11.191 \ (f_1 = 0.032)$	335 R 0.9888*	
	Average dN/dS for each branch = 1.1308	337 Q 0.9980**	
		340 V 0.9992**	

p and *q* are parameters of the β distribution. *f* is the proportion of sites assigned to an individual ω (= dN/dS) category or to a β distribution with shape parameters *p* and *q*. The proportion $f_{1,2}$ in parentheses is not a free parameter. Sites assigned to $\omega_{1,2}$ are those with posterior probabilities *P* > 0.95; those with *P* > 0.99 are in bold. Analyses were conducted using κ as a free parameter. S = tree length.

Sites assigned to $\omega > 1$ are those with posterior probabilities P > 0.95 () or P > 0.99 (**).

2. Log Likelihood Scores and Parameter Estimates for Four Models of Variable ω's Among Sites Assuming the F61 Model of Codon Frequencies

Model	Parameter estimates	Sites* with $\omega > 1$	l
M0: one ratio	Average dN/dS for each branch		-3,469.858
	= 1.1375		
Site models			
M1: neutral	$(\omega_0 = 0) f_0 = 0.331$	Not allowed	-3,462.818
	$(\omega_1 = 1) (f_1 = 0.669)$		
	Average dN/dS for each branch = 0.669		
M2: selection	$(\omega_0 = 0) f_0 = 0.215$	330 G 0.9914**	-3,443.237
	$(\omega_1 = 1) f_1 = 0.749$	332 R 0.9934**	
	$\omega_2 = 10.019 \ (f_2 = 0.036)$	335 R 0.9840*	
	Average dN/dS for each branch =	337 Q 0.9984**	
	1.1057	338 T 0.6807	
		340 V 0.9993**	
M3:discrete	$\omega_0 = 0.00001 f_0 = 0.415$	Too many to list.	-3,441.147
	$\omega_1 = 1.761 \ (f_1 = 0.568)$	Note: M3 vs. M0 is a test	

Model	Parameter estimates	Sites* with $\omega > 1$	l
	$\omega_2 = 17.069 \ (f_2 = 0.017)$	of heterogeneity among	
	Average dN/dS for each branch =	sites and not an explicit	
	1.2835	test of positive selection.	
Μ7:β	$p = 0.00484 \ q = 0.00310$	Not allowed	-3,463.372
	Average dN/dS for each branch = 0.6		
M8: β and ω	$p = 0.00553 \ q = 0.00144$	330 G 0.9919**	-3,443.266
	$f_0 = 0.964$	332 R 0.9938**	
	$\omega_1 = 10.263 \ (f_1 = 0.036)$	335 R 0.9849*	
	Average dN/dS for each branch =	337 Q 0.9985**	
	1.2368	340 V 0.9994**	

p and *q* are parameters of the β distribution. *f* is the proportion of sites assigned to an individual ω (= dN/dS) category or to a β distribution with shape parameters *p* and *q*. The proportion $f_{1,2}$ in parentheses is not a free parameter. Sites assigned to $\omega_{1,2}$ are those with posterior probabilities *P* > 0.95; those with *P* > 0.99 are in bold. Analyses were conducted using κ as a free parameter. S = tree length.

* Sites assigned to $\omega > 1$ are those with posterior probabilities P > 0.95 (*) or P > 0.99 (**).

	2δ	df	P value
Model 0 (one ω for all branches)			
F3x4			
One ratio vs. M3 $(k = 3)$	57.436	4	<i>P</i> < 0.0001
M1 vs. M2	42.986	2	<i>P</i> < 0.0001
M7 vs. M8	42.988	2	<i>P</i> < 0.0001
F61			
One ratio vs. M3 $(k = 3)$	57.422	4	<i>P</i> < 0.0001
M1 vs. M2	39.162	2	<i>P</i> < 0.0001
M7 vs. M8	40.212	2	<i>P</i> < 0.0001

3. Likelihood Ratio Test Statistics (2δ) for Models of Variable Selective Pressure Among Sites.

1. Yang, Z. (1997) Comput. Appl. Biosci. 13, 555–556.