

A

Transcription factor	Control					Wasp					Response				
	<i>Trans diverged D. simulans</i> biased		Other			<i>Trans diverged D. simulans</i> biased		Other			<i>Trans diverged D. simulans</i> biased		Other		
	Motif / +	Motif / -	Motif / +	Motif / -	P value	Motif / +	Motif / -	Motif / +	Motif / -	P value	Motif / +	Motif / -	Motif / +	Motif / -	P value
Relish	6	30	63	432	0.67	6	63	44	418	1	0	69	9	453	0.50
Dif	12	149	24	346	0.83	21	29	140	341	0.08	1	8	160	362	0.37
STAT	8	28	76	419	0.39	11	39	73	408	0.29	1	8	83	439	1
Serpent	6	30	94	401	0.90	7	43	93	388	0.47	0	9	100	422	0.30
Dorsal	1	35	37	458	0.47	2	48	36	445	0.53	1	8	37	485	1
CrebA	5	31	93	402	0.61	8	42	90	391	0.78	2	7	96	426	1

B

Transcription factor	Control					Wasp					Response				
	<i>Trans diverged D. sechellia</i> biased		Other			<i>Trans diverged D. sechellia</i> biased		Other			<i>Trans diverged D. sechellia</i> biased		Other		
	Motif / +	Motif / -	Motif / +	Motif / -	P value	Motif / +	Motif / -	Motif / +	Motif / -	P value	Motif / +	Motif / -	Motif / +	Motif / -	P value
Relish	2	45	67	417	0.10	15	120	54	342	0.54	3	14	66	448	0.83
Dif	17	30	144	340	0.45	38	97	123	273	0.60	6	11	155	359	0.85
STAT	9	38	75	409	0.66	21	114	63	333	1	4	13	80	434	0.59
Serpent	12	35	88	396	0.30	23	112	77	319	0.62	3	14	97	417	1
Dorsal	4	43	34	450	0.94	11	124	27	369	0.75	2	15	36	478	0.79
CrebA	8	39	90	394	0.95	21	114	77	319	0.38	2	15	96	418	0.69

C

Transcription factor	Control					Wasp				
	<i>Cis</i> diverged		No <i>cis</i> divergence			<i>cis</i> diverged		No <i>cis</i> divergence		
	Motif / +	Motif / -	Motif / +	Motif / -	P value	Motif / +	Motif / -	Motif / +	Motif / -	P value
Relish	14	230	19	268	0.8108	16	238	17	260	1
Dif	18	226	20	267	0.9896	16	238	22	255	0.5719
STAT	13	231	15	272	1	13	241	15	262	1
Serpent	19	225	26	261	0.7126	19	235	26	251	0.5275
Dorsal	6	238	16	271	0.1148	8	246	14	263	0.3777
CrebA	8	236	21	266	0.0644	8	246	21	256	0.03999

**Table S2. Number of *cis* or *trans* diverged immune responsive genes in the fat body, with or without transcription factor binding site motifs.** Immune-responsive genes in the fat body were classified as to whether they had motifs associated with the binding of six transcription factors. In (A) and (B) Motif/+ genes had the motif in both *D. sechellia* and *D. simulans*. In (C) Motif+ genes had the motif in just one of these species but not the other. These genes in turn were classified as to whether they had diverged in *cis* or *trans* in control or immune-challenged conditions, of if the response to infection had been changed in *trans*. The *P* value is from a chi-squared test.