

Supplementary Table 3

**Search results using a PWM for Odd built in Target Explorer.** Segments in the *D. melanogaster* genome were required to contain two Odd sites within a window of 300 bp and a PWM score threshold of  $\geq 7.5$  for each site. This search identified a list of 1789 segments throughout the genome. This list was refined by requiring the presence of two Odd sites above the same PWM threshold in corresponding segments of the *D. pseudoobscura* genome. Finally binding sites within exons or more than 20 kb from the nearest gene were removed from the list. 130 segments met these criteria. The score of the Odd sites is the sum of the score of the best pair of sites in each segment divided by the maximum possible score (10.16). Position corresponds to the *Drosophila melanogaster* genome sequence release 3.

*odd* is a primary pair-rule gene that is required for segmentation, head development and central nervous system (CNS) development during embryogenesis<sup>1-4</sup>. This list includes genes required for segmentation (*nkd*, *gsb*, *slp1*, *prd*, *h*), head development (*Gsc*, *ems*, *lab*, *oc*), and CNS development (*sc*, *ac*, *SoxN*). Based on genetic analysis, at least five of these genes are likely to be directly regulated by Odd<sup>3-5</sup>: three pair rule genes (*h*, *prd*, and *slp1*) and two proneural genes (*sc* and *ac*). Some previously identified genetic targets of Odd<sup>3,6</sup>, including *engrailed* and *wingless*, were not recovered in our genomic search using the Odd PWM. These omissions may reflect our stringent search parameters. Both *engrailed* and *wingless* have individual Odd binding sites within 5 kb of their transcriptional start sites that are conserved between the *D. melanogaster* and *D. pseudoobscura* (*data not shown*). More accurate CRM predictions should be possible using a combination of PWMs for multiple factors involved in a common regulatory network, such as the pair-rule genes<sup>7,8</sup>.

Score of Odd sites	Chr.	Position of first Odd site	Gene A	Segment position relative to A	Distance to Transcription Start Site (bp)	Gene B	Segment position relative to B	Distance to Transcription Start Site (bp)
1.97	X	17073340	<i>B-H2</i>	3'	27800			
1.97	2R	20109481	<i>gsb</i>	3'	3763	<i>gol</i>	3'	18409/11762
1.97	3R	4157251	<i>CG7443</i>	3'	7179	<i>CG9603</i>	3'	9527
1.97	3R	12227143	<i>ss</i>	intron	2280			
1.96	3R	2493214	<i>lab</i>	intron	11099			
1.96	X	8530931	<i>Lim1</i>	intron	13843			
1.96	3L	18187778	<i>CheA75a</i>	5'	13198			
1.95	2R	4127672	<i>CG8197</i>	5'	5001	<i>ana</i>	5'	2176
1.95	3L	18970954	<i>nkd</i>	intron	23256			
1.95	X	13643881	<i>CG11072</i>	5'	11599	<i>CG32606</i>	3'	1882
1.95	X	18282867	<i>Bx</i>	intron	17243/14429			
1.95	3L	11327535	<i>CG6163</i>	5'	5118			
1.95	3R	6480770	<i>hth</i>	5'	16366			
1.95	3R	3152555	<i>CG10029</i>	5'	9645	<i>CG31496</i>	5'	18399
1.95	2L	14418526	<i>CG3474</i>	5'	11243	<i>CG15283</i>	3'	14442
1.95	X	7287275	<i>CG32720</i>	5'	10009			

1.95	X	8384268	oc	intron	5328			
1.94	2R	13626166	CG30115	5'	811			
1.93	X	160065	sc	3'	6567	<i>l(1)sc</i>	5'	7096
1.92	3R	21907920	dys	intron	3712			
1.92	2R	4770073	CG10459	3'	1802	<i>dap</i>	5'	2883
1.92	2R	9499021	phyl	5'	2960	CG17388	5'	6944
1.92	2L	15262288	CG15261	5'	9175	CG15263	5'	824
1.92	3L	553243	CG17090	intron	29544/26543			
1.89	3R	9716184	E5	5'	16264	<i>ems</i>	5'	11416
1.82	2L	16388914	CG5888	intron	37635			
1.81	2L	8810494	SoxN	5'	7362			
1.8	2R	17198073	a	intron	17531/17089			
1.8	3L	710829	CG13896	5'	18626	CG13897	3'	8920
1.8	3R	20789396	CG13647	3'	2555	CG31422	3'	25147
1.79	3L	16437084	CG33158	intron	68411			
1.79	X	8406035	oc	5'	16439	CG12772	5'	10025
1.79	3L	20601963	CG32431	5'	4317			
1.79	2L	15326224	esg	3'	14275	CG15258	3'	13449
1.78	3R	12019746	sra	intron	7072			
1.78	2R	2256768	pk	intron	45164/8604			
1.78	3R	21836710	m2	5'	4153	HLHm3	5'	92
1.78	X	17065733	B-H2	3'	20193			
1.78	X	124671	y	3'	10724	ac	5'	2798
1.78	X	1321046	CG14796	5'	6102			
1.78	3R	5114769	CG8454	3'	21999	CG8861	3'	39474/5015
1.78	2L	5450620	CG31647	5'	9203	CG6634	5'	3236
1.77	3L	12387146	toe	intron	3069			
1.77	2L	15077733	CG15270	3'	24003	CG15269	5'	9937
1.77	3L	2522454	Spn	intron	12452			
1.76	3L	22533345	CG6914	3'	9698			
1.76	2L	4350753	CG15429	3'	5209	<i>Traf1</i>	5'	13806
1.76	2R	4126236	CG8197	5'	3565	<i>ana</i>	5'	3612
1.76	2L	12070805	CG14947	3'	5111	<i>prd</i>	3'	4522/3444
1.76	3R	13330528	CG31190	intron	876			
1.76	3R	11056313	mRpS10	3'	906/820	CG6535	3'	11570
1.75	3R	19142388	pnt	intron	20153/11914			
1.75	2L	14432503	CG15283	intron	465			
1.75	2L	17491891	Dhc36C	intron	12341			
1.75	3R	27835883	CG2053	5'	17578			
1.75	3R	24372095	CG14063	3'	2644			
1.75	2L	8907731	CG15867	5'	335			
1.75	3R	17655621	Gr93a	3'	6432	CG13416	5'	1970
1.75	X	8409695	CG12772	5'	6365			
1.75	3R	22717622	T48	intron	20801			
1.74	2R	10946734	sli	intron	7845			
1.74	3R	13983718	CG14316	3'	5861	CG14315	5'	789

1.74	X	15278472	CG32584	3'	28205			
1.74	2R	16008812	<i>hbn</i>	5'	4359	CG15649	3'	8663
1.74	2L	5254918	<i>tkv</i>	intron	9006			
1.74	2L	15299975	<i>nht</i>	5'	1984	<i>esg</i>	5'	11974
1.73	2L	14372370	<i>eIB</i>	intron	17540			
1.73	3R	25321433	<i>Trc8</i>	5'	2548	CG2014	5'	9500
1.73	3L	13907610	CG32137	intron	19478/19477			
1.73	2R	14873689	<i>Obp56h</i>	5'	5601			
1.73	3L	6734508	CG32392	5'	11733	<i>vvl</i>	5'	15181/8148
1.72	3L	8630244	<i>h</i>	5'	4425			
1.72	2R	9989803	CG10249	5'	13860	CG10253	3'	7223
1.72	X	6932312	CG32729	3'	2207	CG9650	5'	1140
1.72	2L	591812	<i>Gsc</i>	intron	4540			
1.72	X	20232340	CG15455	3'	18661	CG15454	5'	5229
1.72	3L	15122159	CG7011	5'	9656			
1.72	2R	848020	<i>ap</i>	intron	17681			
1.72	2L	12624482	<i>nub</i>	3'	17264			
1.72	2R	2470747	CG1701	intron	2235			
1.71	2R	18683619	<i>retn</i>	intron	7439			
1.71	3L	20591496	CG32431	5'	14784			
1.71	3L	7922305	<i>exex</i>	3'	14584	RNaseX25	5'	7819
1.71	X	11513842	<i>dy</i>	5'	4509	<i>Karl</i>	5'	5661
1.71	3L	13265462	<i>Acp70A</i>	3'	6259			
1.7	2L	8848506	CG32986	5'	6532			
1.7	2L	3826483	<i>slp1</i>	3'	8234	<i>slp2</i>	5'	2928
1.69	2R	1690306	<i>Vha16</i>	intron	3724/2977			
1.64	3R	17248719	<i>lbl</i>	intron	2090			
1.62	3L	16713314	<i>Nrt</i>	5'	1640			
1.6	3L	19613099	CG8780	intron	521			
1.6	3L	8981081	<i>Doc2</i>	5'	3131	<i>Doc1</i>	5'	19006
1.6	2L	7356176	<i>Wnt6</i>	3'	13047	<i>Wnt10</i>	5'	10817
1.59	3R	4152688	CG7443	3'	2616	CG9603	3'	14090
1.59	3L	12437330	CG10663	5'	1694	CG10660	3'	10305
1.58	2R	10887316	<i>tun</i>	5'	8678	<i>Poxn</i>	3'	11184
1.58	3L	502848	<i>klar</i>	intron	18370			
1.57	3R	3154983	CG10029	5'	12073	CG31496	5'	15971
1.57	2R	15184841	<i>18w</i>	3'	9618			
1.57	2L	8852440	CG32986	5'	2598			
1.57	X	14689532	CG9030	5'	3418	<i>eag</i>	5'	12738
1.57	3R	16845088	<i>Rlip</i>	intron	1645			
1.57	3R	12562470	<i>Ubx</i>	5'	2104		3'	14659/9656
1.56	3R	24381227	CG14063	3'	11776	<i>fkh</i>	3'	18944
1.56	2L	20346069	<i>Ugt37a1</i>	5'	2360	CG16798	3'	6028
1.55	3L	7897471	CG8268	3'	1567	<i>exex</i>	5'	10250
1.55	2R	873539	<i>ap</i>	5'	7838	<i>l(2)09851</i>	5'	19860/4490
1.55	2L	10795834	CG31869	5'	2340	CG6508	5'	15698

1.55	3R	15490545	<i>ort</i>	3'	5058	CG7432	5'	2934
1.55	3R	17253878	<i>lbl</i>	5'	3069	<i>lbe</i>	3'	9128
1.55	3L	12394551	<i>toe</i>	5'	4336			
1.55	3R	10421211	<i>stumps</i>	intron	3008/1395			
1.54	X	7282988	CG32720	5'	5722			
1.54	3L	16890459	CG9665	intron	60			
1.54	3L	12889583	CG32115	5'	18390			
1.53	X	2154224	CG14045	5'	9788	CG12496	3'	13447/2441
1.52	3R	17775673	<i>Eip93F</i>	5'	16639			
1.52	3R	21607770	CG31093	5'	2212	CG5024	5'	18413
1.52	2L	15999265	<i>beat-1c</i>	intron	20644			
1.52	X	7874749	CG2120	intron	108			
1.51	3R	6871365	CG4695	intron	1653			
1.51	2L	11442500	<i>salm</i>	5'	8026			
1.51	X	20387452	<i>run</i>	3'	20596	CG1324	3'	20923
1.51	X	19988339	CG1504	5'	3868	CG1631	5'	11953
1.51	2L	16479848	<i>Tpr2</i>	intron	10222/4998			
1.5	3L	13450203	CG10741	intron	5831/2992			
1.5	2L	702374	<i>ds</i>	intron	14259			
1.49	3R	13904260	CG14317	5'	2598			
1.49	2R	3091987	CG14762	5'	17332	<i>Optix</i>	5'	393
1.49	2R	13765541	CG15071	3'	5187	CG18604	5'	138

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