

## Supplementary Table 2

**Unique insert sequences from the prey isolated using the B1H system with each bait.** Bold letters indicate an overrepresented motif identified by MEME analysis. Italic letters indicate the inclusion of constant bases neighboring the randomized region to complete a binding site consistent with an overrepresented motif identified in the other sequences by MEME analysis. The E-value calculated by MEME for each identified motif is indicated. An asterisk following the sequence indicates that it is the reverse complement of the insert sequence. The absence of bold letters indicates a sequence that does not contain a binding site motif based on computational analysis. Numbers in parenthesis indicate how many times a sequence occurred among the analyzed clones.

### Zif268 sites (Two step selection; E-value=3.4e-020)

TTGAGCGCGTGGGTGGAT  
CAACGTTGTGTGGCGTG (2)  
TTGCCCGCGTGGCGTC  
  **CATA**CGGAAGCGGGGTGC\*  
TTGTTGTGCGTAGGCGGC  
  AATCG**AGCGT**GGGAGGGG  
ATCGAGGGCGTGGGTGGC  
TGTAGTTGC**GT**GGGAGGG (2)  
CTAACCC**CG**GTGGCGTA  
CGAGGATGC**GGGGTGTG**  
TTGCC**CCACGT**GGGCGTG  
CGATAATCCGGGGCGTA  
  GAATGTGTGTGGGCGGCT  
ATTG**TAGCGT**GGGCGAT  
GAGCGAAGCGTAGGC**GT**C

**PLAG1 sites** (Motif score = 2.278 calculated by Bioprospector). The Sequence logo for PLAG1 in Figure 2 is the reverse complement of the overrepresented motifs in these sequences. 18 of 21 sequences contain the identified PLAG1 motif.

CCCCCGAAAGGGCCCC (6)  
CCCCCGATTGGGGCCTC (3)  
TTCCCCATTGGGGCCCCT (4)  
AACACCCCTGTGGGCCCC  
  CCCCCTAATGCACCCCAT  
  CCCCCGTGGAGGCCAC (2)  
  CTACCTTTCTGGCCCC (2)  
AACACCCCTAAGGAGCCCC (2)  
CCCCCCTCCTAGGCC (2)  
  CCCCCTACCAGGGCCTCA (2)  
  CCCCCTTGGAGGCCACTA  
CCCCCCCCGTGGCCCCCT  
CCCCACCCCTGGGCC (2)  
CCCCCTAACGCGCCTCCG

**ACCCCCCTTAGTGCCCCC**  
**ACCCCCATTTACCCCCC**  
**GTTCCCCCTTCCGGCCCC**  
**CCCCCTTCACGGGCCCC**  
**GTATCCCAGGTATGCGA**  
**GCAGAACCGGCTAGGAAT**  
**GGGCCAGAATTCCGG**

**Zif268 sites** (One-step selection; E-value=3.0e-06)

**TGTACTGCGTGGGTGTGG\*** (2)  
**ATGCTTGTGCGTACGTGGGCGG** (2)  
**CAGCCTGCGTGGGAGGTT\***  
**TGGACATGCGTAGGCAGG\***  
**CCGCGTGTGTGGGCGGAA\***  
**TTGCCCCACGTGGGCGTG\***  
**CTCGCACGTGTGGGCGTC\***  
**ACTATCACGTGGGCGTG\***

**ZnFp53 sites** (E-value=1.7e-045). The Sequence logo for Znfp53 in Figure 2 is the reverse complement of the overrepresented motifs in these sequences

**ACGTGTCCCGTCAGACGGG**  
**AGCGTGTCCCTCAGTCAG**  
**TGACGTGGCCTGTCACTC**  
**GACGTGGCCGGATGTCCG**  
**AGCTCGTGTCCCATGATCA**  
**GACGTGCCCGGCTATCCC**  
**GACATGTCCACGCCCTAG**  
**TAAGCGTGTCCGTAGCGG**  
**CCCCCGACACGTGTCCCTA\***  
**GGCGTGTCCCGCTGTTA**  
**ACCCGTGTCCAGCTTAG**  
**CCCGCGTGTCCGTAAAGGG**  
**GTCGTGTCCCCATGGTCG**  
**GTCGTGTCCCTGCACGGG**  
**AGTGCCTGTCCCTGGTGCT**  
**AGCGTGTCCCTGGTCTGCT**  
**GACGTGTCCGGATTTCCC**  
**GGTGACGTGTCCCTGGGCA**  
**GCGTGTCCCGCAGCCTGT**  
**ATCGTGTCCGCTCACCTG**

**LAG-1 sites** (E-value=1.4e-023). The Sequence logo for LAG-1 in Figure 3 is the reverse complement of the overrepresented motifs in these sequences. 12 of 17 sequences contain the identified LAG-1 motif.

**TGTGAGAATTCCCCACGGT** (2)  
**GAGTTCCCACGGTCGACA\*** (3)  
**GGTCACTTTCCCCACGGTG** (3)  
**TCAACTTTTCCCCACTGA\*** (3)

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CCTTTCCCACGGTTACCC*( 3 )
AGTTGTTTTCTCACGGT( 2 )
    CAGTTCCCACGATTCCGC*
    GATTTCCCACGTGTGGC*
    CATTCCCACGCTTCAGA*( 2 )
    TCATTCCCACGGTAGCAG*
    CTCATTCCCACGGTCCAG*
TCAACTTTTCCCACCTG*
CTTCACGGCTCTGACCGG
GGTATGCAATCGATAGAG
GTCTGACCATGTCCTTCC
ATCCGATGTGTTGATCGC
CCCGCGGACACGAACGGT

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**Dorsal sites** (E-value = 5.5e-058). The Sequence logo for Dorsal in Figure 2 is the reverse complement of the overrepresented motifs in these sequences. 32 of 35 sequences contain the identified Dorsal motif.

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CCACCACGGGAATTCCC
AGGCACAGAGGGAAATTCCC
CCCACCACGGGAATTCCC( 3 )
CTGTGGACAGGGATTCCC
ATAGCCAGAGGGATTCCC
CCGGATTCTGGGATTCCC
TTTCGCGGGGGGTTTCCC
    ATCCGGCGGGAAAACCC
    CGCGTACCAGGGAAAACCC
    GGTGCGTCTGGGAAAACCC
    TAATCACGGGGAAAACCC
GGTGCGTCTGGGAAAACCC( 2 )
GGTGTGGCGTGGAAAACCC
    CCACCGGACGGAATTCCC
GCCATGTACCGGAATTCCC
TCGTATACTGGGAAAAACC
CGCCTGCGCTGGAAAAACC
ACCTCGCCTGGAAAACCC
    CGGTGCCTGGGATTCCC
ACCTGCTCTGGGAATTCC( 2 )
CCTACCTCGGAGAATTCCC
    CAGTGAACTGGGAAAACCC
CCTGGTAGCCGGAAAACC
    CCCTAGCGGGTTTCCCC
TGTTCACCCGGGTTTCCC
AACCTGGAGCGGAAAACC( 2 )
ACTGCATCACGGGATTCCC( 2 )
    TGGTAGAATTCCCATGGT*
TCAATCGGAGGGAAAACC
ACCCAGCTTGGGAAATCC
TCGTACGGTGGTTTCCC

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CATCGGTTT**GGGGTTTCCC**  
GCCATGCTATCGGCCGCG  
CACCCCCATCAGAATCCTC  
CGAACCGTTAGCAGCTTAC

**Paired site** (E-value = 3.0e-028). 22 of 32 sequences contain the identified Paired motif.

**AATTCGTCACGGTTGATG**  
**GATTCTCGTCACGCTTGCTT**  
**TGATTTCTGTTACGGTCTG**  
**CGATTAGTCACAGTGCCTA**  
**CTAATTCTCCGCTCCC(4)**  
**CCGATTCTCCACTACC**  
**TTAATTCTTCCGCTCCT**  
**CCGATTCTCCCTCTACC**  
**ACAATTCTACGGTGGT**  
**CCGATTCAACCCAGTCAC**  
**CCGATTAGTCTCGGTTCA**  
**CCGCTTCGTACGCTATG**  
**CTGATTGATACGCTTCG**  
**CATATCCGTACGGTGT**  
**AAGAACGTCACGGTTCAC**  
**CTCGCTCGTCACGGTTGG**  
**ACAATTCTACTGTGGT**  
**CACTAGTTACTGTACCGC\***  
**CGTTTCTATAAGGCTGCGT\***  
**AAGAACGTCACGGTTCAC**  
**ATAAATCTGACCGGTGGG\***  
**TCATACGGCCCCCTTCCTC**  
GCGGTACAGTATGTCCCA(2)  
TGAGTTATTATGGCGGGG  
AGTGATTGTATGGTATAC  
AACCTGTGCTAGAGTACC  
AGTAAAGACCGTGCTGCC  
CAAGCTGAGCCAAAATG  
TCTTATTTGTCAGTAGGA  
CGGGCGGCTCGGATTAAT(2)  
GGAATACTCTGAAAACAC  
GCCCGCTGCAGAATTAA

**Runt/Bgb sites** (Two step selection; E-value = 1.1e-067) The Sequence logo for Runt in Figure 2 is the reverse complement of the overrepresented motifs in these sequences.

Sequences prior to counterselection

**TAGCCGCAGATTGACTGC**  
GATCTACAGGCGCTATGA

TCTCGGACGCAGAGTACG  
AAACTGTCCCACCCCTAC  
CACGCTATACTGCAACAC  
ACATGTATACTTGAGAG  
AGCTACTCCTGGGAGAAT  
ACTGGTATATTAGTCGG

#### Sequences after counterselection

**CAAACCGCAATCCTCGCC (2)**  
AAGTTCCA**AAACCGCAAGC**  
**GGTAGCCTAACCGCAAGT\***  
**ACTTAACCGCAAAGGATG (2)**  
**AATAACCGCAAACCTACCC (3)**  
**AAAACCACAAGCGCTCCT**  
**ATAAACCGCAGGCTCTAC**  
**CTCGAAAAACCGCAGCCT\***  
**CTGAAACCGCAGTTCGG**  
**TAAACCGCAGGTTCCAGC**  
**AAGCTAACCGCAGAAGTG**  
**CTAACCGCAGCGCTATT**  
**ACCCCAGTAACCACAAGT\***  
**CTAAACCCACAGCTCCCCG**  
**CTCAACCGCAAGGACGAC**  
**ACCCAAACCACAAAAGGTG**  
**ACCGCAA**AAGCCACAATC\*** (2)**  
**AACGACCACAAAGTCCTTG**

#### **Odd sites**

##### 1.5 mM 3-AT (E-value=4.3e-031)

**GCTACTGGTCTTAGGATCT**  
**CTGCTACTGTAAATGGTAAC**  
**GCTACCGGTTCTTACAGA**  
**GCTACCGGAAGTCATCGTA**  
**GGATCTGTTGTTGGTG**  
**GAGCTACTGTCTTGCAGG**  
**GCTACTGCTTAGTCTTGGT**  
**GCTACCGGACTAGTTACCC**  
**AGCTTCTGAAATTCCAGA**  
**GCTACTGGACCCACCTAG**  
**GCTACTGGCGGGCAACTC**  
**GCTACTGGACTGTTGTCGC**  
**GCTACCGATGCTCACTCC**  
**GAGGATCCGGAAGCAACG**  
**GGTCCTTCCGTGCTACTGGC**  
**GCTACTGGTGTCTTAGT**  
**GCTACTTGTCAAGGTCCGA**  
**TGCTACCGAGCATGAGCG**  
**GCTACTGCAAGCGGTGCGTA**

**ACTACCGTTAGTTGGCG**  
**GACTACTGGAAGCCTGGT**  
**CGCTTCCGAAACTACCG**  
**GGGATCCGTTGGAGCGAT**

2.5 mM 3-AT (E-value=9.5e-43)

**GCTACCGTTTACCCGGG**  
**GCTACTGGTCTTAGGATCT ( 2 )**  
**TGCTACTGTGTAATGC**  
**TGCTACTGGAAGTTGGTC**  
**CGCTACTGTTGTGTGG**  
**GCTTCTGTATTGTTGCCA**  
**TGGCTACTGTCTGCCACC**  
**TTGCTACTGTACCCGGGG**  
**GCTACCGTGTGACTCGGA**  
**GCTACTGTAGGCGAACAA**  
**GCTACCGTAGACGCTCGG**  
**GCTACCGTGTGTACCGCA**  
**AGCTACCGGAGCCCTAAG**  
**GCTATCTGGTGGAAAGTTG**  
**GCTACCGGAAGTTCTAG**  
**GCTACCGTTATGTCGTCG**  
**GCTACTGATTCCCCCTTA**  
**GCTACTGATTGCGGAAGCA**  
**GCTTCCGGAAAAGGCGTG**  
**GCTACTGGTGCTGCACCC**  
**GGCTACTGGTAACAGAGT**  
**GGGCTACCGGGGAAATTC**  
**GCTACCGTGAAGTTGTTGT**  
**GCTACCGGACGGTCCCTC**  
GGATCCGTATGAGTGCCTC

5 mM 3-AT (E-value=3.2e-049)

**GCTACCGGTTCCCTTACAGA**  
**CGCTACCGAATTAAAGGG**  
**GCTACCGTATACTTTCCA**  
**GCGCTACTGTTGGGGTG**  
**GCTACTGTTCTGTCACC**  
**GCTACTGGTAAAACCAAGT**  
**GCTACTGTTGCGTGCAAGG**  
**TGCTACTGGACAAAATCG**  
**TGCTACTGTTCACTCGGG ( 3 )**  
**TGCTACTGGAAGTTGGTC**  
**TGCTACTGGAAGGGGTGA**  
**AGCTACTGTATCAGTGGT ( 3 )**  
**GCTACTGTTACGCATGGG**  
**TGCTACCGTAGCGACGAG**  
**GCTACTGGTCTTAGGATCT**

**TGCTACTGTGCAATGCGG**  
**TGCTACTGTGGATGAGCG**  
**TGCTACTGTGTAATGCAGC (3)**  
CGGCTACTGTGCAGCTAG  
    **GCTACTGTGCAGTTGCTG**  
CGGCTACTGTATACTGTCC  
GTGCTACTGTAAGCTCAG  
GTGCTACCGTTGTTCGG  
GGACTAGGCTGTCTACCC  
GACACATCCTCGGCGGCT  
GGCCTAGGGAAAGTCTGT  
GCCCGAGGAGCATAAAG