

Supplementary Figure 1

zebrafish -----MANGAEDVVFCRGTGQSDDSDIWDDTALIKAYDKAVA**S**FKNALKGEDGA
 frog -----MAGLEDGGEVLFRRGAGQSDDSDIWDDTALIKAYDKAVSSFKRALKNEDCT
 mouse ---MAMGSGGAGSEQEDTVLFRRGAGQSDDSDIWDDTALIKAYDKAVASFKHALKNGDIC
 rat ---MAMGSGGAGSEQEDTVLFRRGAGQSDDSDIWDDTALIKAYDKAVASFKHALKNGDMC
 human MAMSSGGSGGGVPEQEDSVLFRRGAGQSDDSDIWDDTALIKAYDKAVASFKHALKNGDIC
 macaque MAMSSGGSGGGVPEQEDSVLFRRGAGQSDDSDIWDDTALIKAYDKAVASFKHALKNGDIC
 cow ----MGGGGGGFPEPEDSVLFRRGAGQSDDSDIWDDTALIKAYDKAVASFKHALKNGDIS
 dog ----MGGGGGLPEPEDSVLFRRGAGQSDDSDIWDDTALIKAYDKAVASFKHALKNGDIS
 cat ---MAMGGGSGVPEQEDSVLFRRGAGQSDDSDIWDDTALIKAYDKAVASFKHALKNGDIS
 chicken -----MAGRVLFRRGAGQSDDSDMWDDTALIKAYDKAVASFKNALKNGDCS
Drosophila -----MSDETNAAVWDDSLLVKTYDESVGLAREALARRAD
C.elegans -----MEVDDVWDDTELKMYDESLQEISKNE-----

zebrafish TPQEND--NPGKKRKNKKNNKSRKRCNAAPDKEWQVGDSCYAFWSEDG--NLYTATITSV
 frog IGAETEENPRTKRKNKKNRKKCNAAPLKKWRIGDTCNAVWSEDG--NIYSATISSI
 mouse ETPDKPKGTA--RRKPAKKNKSQKKNATTPLKQWKVGDKCSAVWSEDG--CIYPATITSI
 rat ETSDKPKGTA--RRKPAKKNKNQKKNATAPLQWKAGDKCSAVWSEDG--CVYPATITSV
 human ETSGKPPTP--KRKPAKKNKSQKKNTAAASLQQWKVGDKCSAIWSEDG--CIYPATIASI
 macaque ETSGKPPTP--KRKPAKKNKSQKKNTAAAPLQWKVGDKCSAIWSEDG--CTYPATIASV
 cow EASEKPKGTP--KRKSAK--NKSQRKNTTSPSKQWKVGDNCAIWSEDG--CIYPATIASI
 dog EASDKPKSTP--KRKPAKKNKSQKKNTAALKQWKVGDKCSAVWSEDG--CIYPATIASI
 cat EASDKPKGTP--KRKPAKKNKSQKKNTAAALKQWKVGDKCSAIWSEDG--CVYPATIASI
 chicken EPSDKQEQRAGVKRKNKKRNRSNAVPLQWKVGDSNAVWSEDG--NVYPATIASI
Drosophila STNKREEENAAAAEEEAGEISATGGATSPEPVSFVKVGDYARATYVDG--VDYEGAVSI
C.elegans -----TSAKITSRKFKGEDGKMYTWKVGKCMAPYEENGEVTDYPATIDTI

zebrafish D-QEKGTCVVFYTDY**G**NEEEQNLSDLLTEPPD**M**DEDALKTAN--VKE**T**ESSSTEESDRSFT
 frog D-AKRRTCIVVYSGYGNSEEHSLADLRFPDTSEAESDQRDQE**E**INGDEHSTDESDRSSR
 mouse D-FKRETCVVVYTGYGNREEQNLSDLLSPTCEVANSTEQNTQENE--SQVSTDDEHSSR
 rat D-LKRETCVVVYTGYGNKEEQNLSDLLSPTCEVANNTEQNTQENE--SQVSTDDEHSSR
 human D-FKRETCVVVYTGYGNREEQNLSDLLSPICEVANNIEQNAQENENESQVSTDESENS-R
 macaque D-FKRETCVVVYTGYGNREEQNLSDLLSPISEVANNIEQNAQENENESQVSTDESENS-R
 cow D-FKRETCVVVYTGYGNREEQNLSDLLSPTSEVAN-IEQNAQENENESQISTDESENSSR
 dog D-FKRETCVVVYTGYGNREEQNVS DLLSPACEVANNVEQDTQENENESQISTDESENSSR
 cat D-FKRETCVVVYTGYGNREEQNVS DLLSPASAVDNNVEQNAQENENESQISTDESENSCR
 chicken N-LKRGTCVVVTGYGNKEEQNLADLLPPASDET-----NENETPYSTDESEKSSQ
Drosophila N-EEKGTCVRLYLGYNEQEVLVLVLLPSWGKVRREQFLIAKKDEDEQLSRPKASAGSH
C.elegans GGADNLEVGVTFIYGGQAVVQMKDLWLNEEAIAADAVKAENDLQKTKKTSTVNSVAHSNS

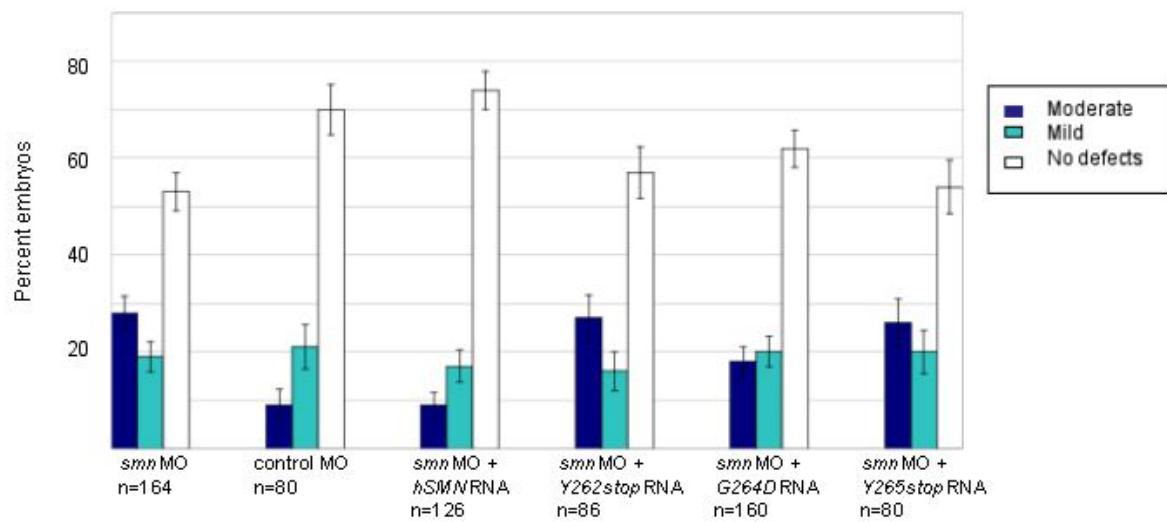
zebrafish PQKSGHAKHKSNSFPMPGPPSWF**P**SFPPGPPPPPHFKKMDGRGEGPGP-----S
 frog SHQSKDPQNRETPKSSQ----WNGQFPVPPPFMPGFRHG-EKLDQAHP-----F
 mouse SLRSKAHSK-SKAAP-----WTSFLPPPPPMPGSGLPGPKGLKFNGPPPPPLPPPF
 rat SLRSKAHSK-SKAAP-----WTSFLPPPPPVPAGLGPGLRFSGPPPPPPPPPPF
 human SPGNKSDNIKPKSAP-----WNSFLPPPPPMPGPRLPGPKGLKFNGPPPPPPPPPHL
 macaque SPGSKSDNIKSKSAP-----WNSFLPPPPPMPGPRLPGPKGLKFNGPPPPPPPPPHL
 cow SPLNKPNNIRSRAAP-----WNSFLPPPPHMPRSGLPGKSGLNFSGPPPPPPPP-PHF
 dog SPGNKPNNIKSKAAP-----WNSFLPPPPPMGSGSLGPGLPGVKFSGPPPPPPPP-HF
 cat SPGNKSNNVSKATS-----WNSFLPPPPPMGAGLGPAPKGLKFNGPPPPPPPP-PHF
 chicken SHHNENNCTKARFSP-----KNLRFPIPPTPPG---LGRHGSKFRTLPP-----F
Drosophila SKTPKSSRRSRSRISGG-----
C.elegans KSTSSAPNTS-----

zebrafish FPGWPPMIPLGPPMIPPPPMSPDFGEDDEALGSM LISWYMSGYHTGY**Y****MGL**RQGRKEAA
 frog LSGWPPPFPLGPPMIPPPPMSPDACEDEALGSM LISWYMSGYHTGYYLGLKQGRMESS
 mouse LPCWMPPFPSPGPIIIPPPPISPDCLDDTDALGSM LISWYMSGYHTGYYMFRQNKEGK
 rat LPCWMPPFPSPGPIIIPPPPISPDCLDDTDALGSM LISWYMSGYHTGYYMFRQNKEGK
 human LSCWLPPFPSPGPIIIPPPPICPDSSLDDADALGSM LISWYMSGYHTGYYMFRQNKEGR
 macaque LSCWMPPFPSPGPIIIPPPPICPDSSLDDADALGSM LISWYMSGYHTGYYMFRQNKEGR
 cow LSRWLPPFPAGPPIIIPPPPICPDSSLDDADALGSM LISWYMSGYHTGYYMFRQNKEGR
 dog LSCWLPPFPSPGPIIIPPPPICPDSSLDDADALGSM LISWYMSGYHTGYYMFRQNKEGR
 cat LSCWLPPFPSPGPIIIPPPPICPDSSLDDADALGSM LISWYMSGYHTGYYMFRQNKEGR
 chicken LSCWPPPFAGPPLIIPPPPMGPSPEDDEALGSM LIAWYMSGYHTGYYLGLKQSRMEAA
Drosophila -----LVMPMPMPVPPMIVGQGDGAEQDFVAML TAWYMSGYYTGLYQGKKEASTTSG
C.elegans -----MPFPSFAPPVPPN-IIAMAPVNQKEAMNSMLMSWYMSGYHTGYYQALADQKNVQN

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zebrafish ASKKSHRK
    frog IKGPPHQK
    mouse -CSHTN--
    rat KCSHTN--
    human CSHSLN--
    macaque CSHSLN--
    cow YSHFN---
    dog CSHFN---
    cat CSHFN---
    chicken LEREAYLK
    Drosophila KKTPKK
    C. elegans -----
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Supplementary Figure 1: zebrafish *smn* mutations identified by TILLING. Protein sequence alignments of zebrafish Smn with other species showing the eight point mutations identified by TILLING (red). The identified *smn* mutations were distributed throughout the Smn protein; S38A, G120E, M137V, T150I, P186A, Y262stop, G264D, and L265stop.

Supplementary Figure 2



Supplementary Figure 3



Supplementary Figure 4

