

Supplementary figures and legends

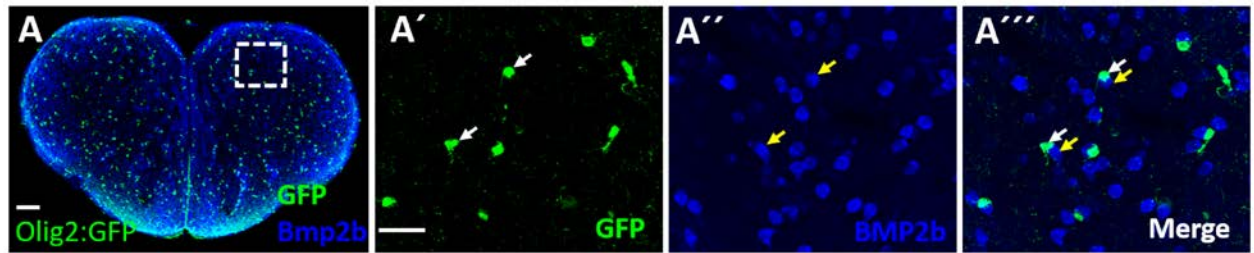


Figure S1. BMP2b is not detectably expressed in oligodendrocytes of the telencephalon.

Cross-sections through the telencephalon of *Tg(olig2:gfp)* transgenic fish marking oligodendrocytes (green) were immunostained with a BMP2b antibody (blue). The boxed-in area in A is magnified in A' to A''' showing individual channels (A', A'') and the merged image (A'''). White arrows show 2 oligodendrocytes, yellow arrows show BMP2b+ neurons.

Scale bars: 100 μ m (A), 20 μ m (A'-A''').

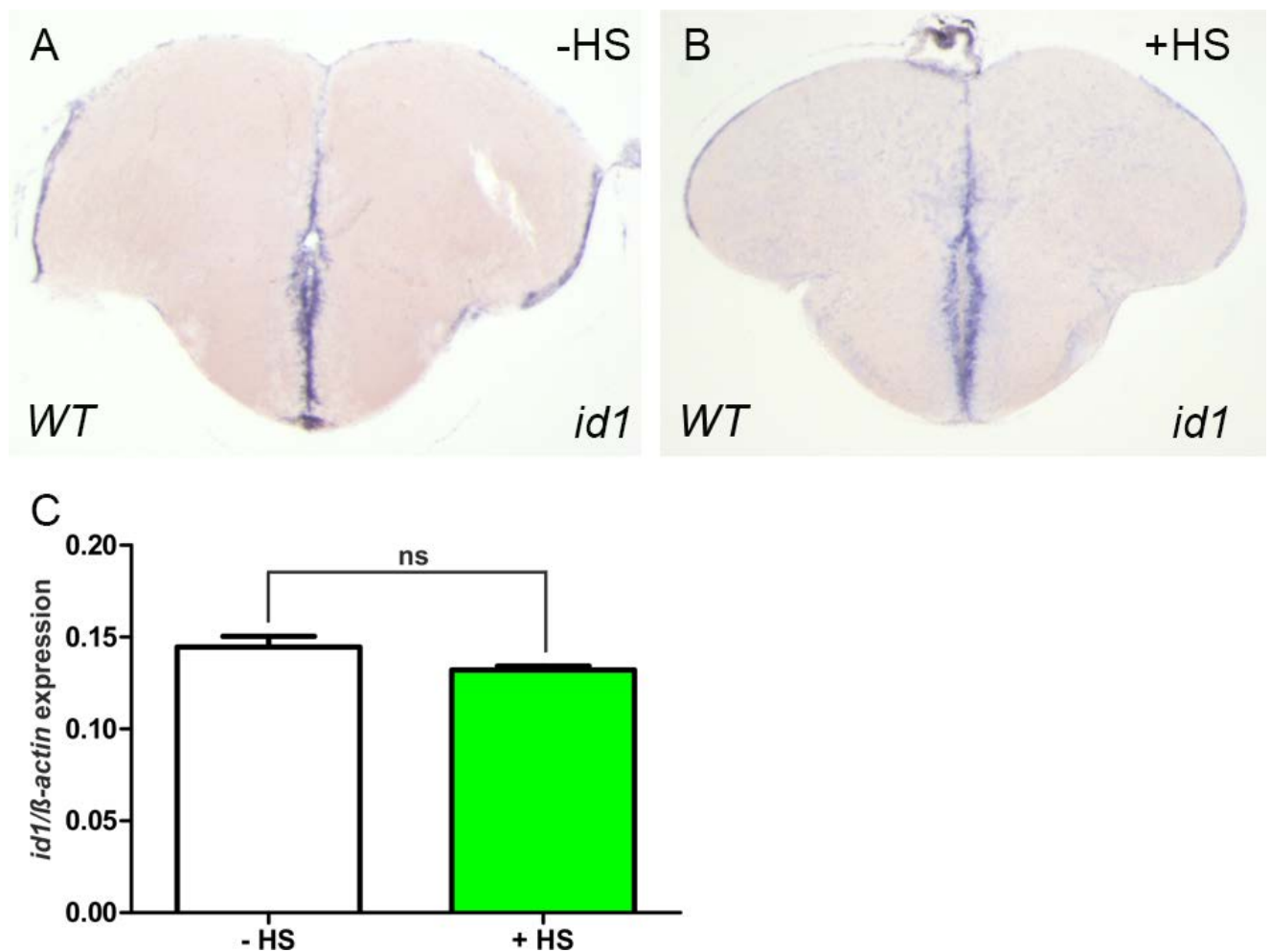


Figure S2. Heat-shock has no influence on the endogenous expression of the *id1* gene.

(A-B) ISH against *id1* on cross- sections of WT telencephala. There is no difference in the expression level for *id1* between non heat-shocked (A) and heat-shocked (B) WT brains. (C) RT-qPCR analysis of *id1* mRNA expression levels in WT with and without heat-shock reveals that the level of *id1* mRNA does not change in response to heat-shock in the WT brain. n=3 brains. Significance is indicated by ns, not significant.

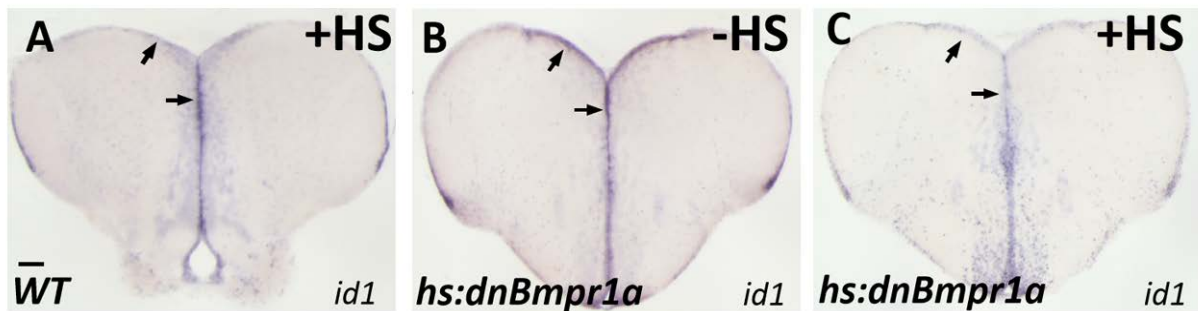


Figure S3. Heat-shock does not influence the expression pattern of *id1* in WT fish.

(A-C) Cross-sections through telencephala of heat-shocked WT, (A) and *hs:dnBMPR1a* transgenic animals without (B) and with heat-shock (C). Heat-shock of WT animals did not influence the expression of *id1*. In contrast, decrease of *id1* expression was noted after inhibition of the BMP pathway by heat-shock of *Tg(hs: dnBmpr1a)* animals (C) relative to transgenics without heat-shock (B). Black arrows show the expression of *id1* in the ventricular zone of WT (A) and *Tg(hs:dnBmpr1a)* animals without heat-shock (B) and after heat-shock (C). n=3 animals (A-C). Scale bar: 20 μ m. HS, heat-shock.

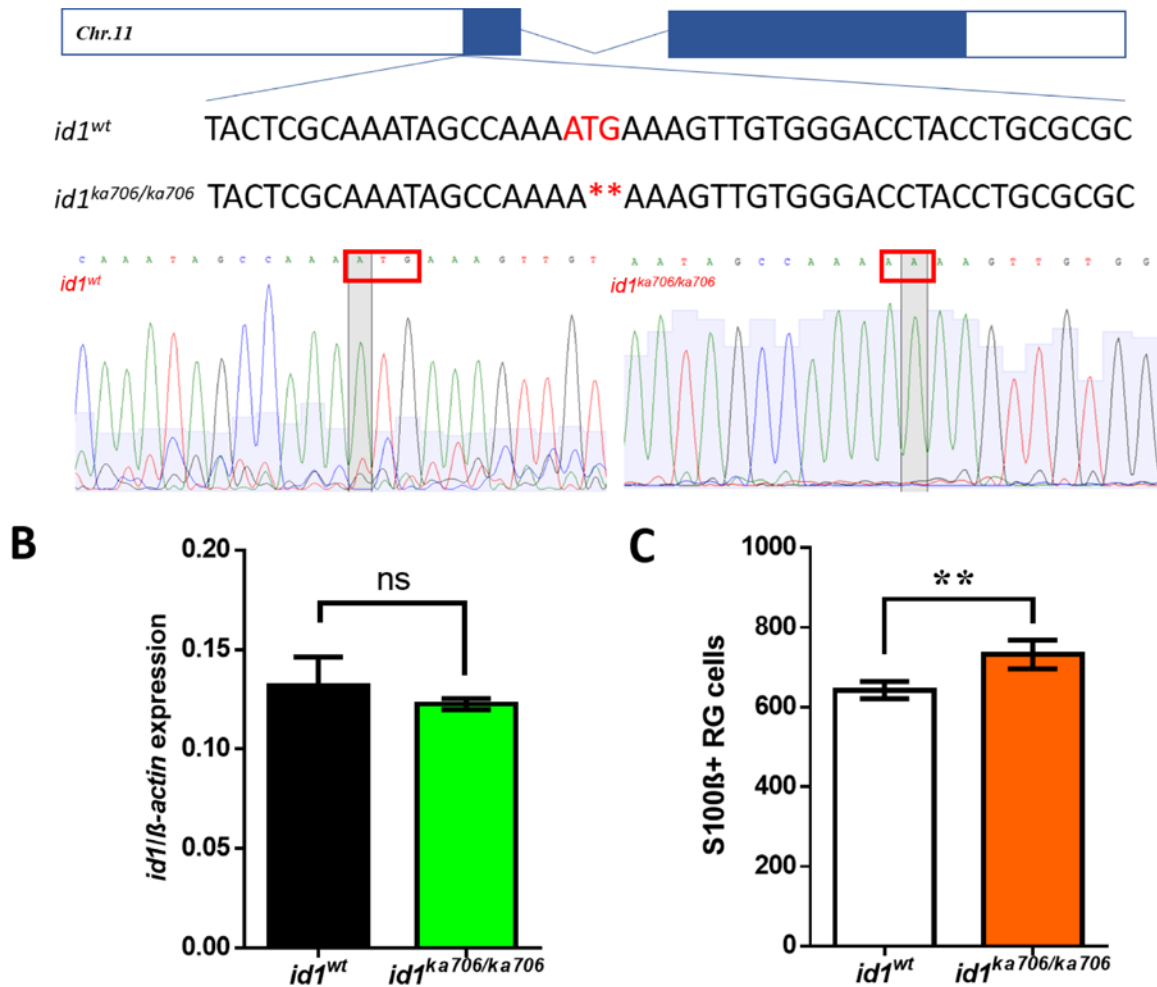


Figure S4. Generation of a CRISPR/Cas 9-mediated *Id1* loss- of- function mutant *id1*^{ka706/ka706}. (A) Schematic representation of the *id1* locus on chromosome 11. The gene consists of 2 exons. Red letters represent the start codon of the *id1* coding sequence. The Sanger sequencing results of WT (*id1*^{WT}) and homozygous (*id1*^{ka706/ka706}) sequences are displayed underneath. In the homozygous sequences T and G of the start codon are deleted. (B) RT-qPCR analysis of *id1* mRNA expression levels in WT and *id1*^{ka706/ka706} telencephala reveals that the level of *id1* mRNA is not changed in the mutant. (C) Quantification of the S100β+ RGCs/NSCs in WT and *id1*^{ka706/ka706} telencephala without injury. n=3 brains (B, C). Significance is indicated by asterisks: ns, not significant; **, *p* < 0.01.

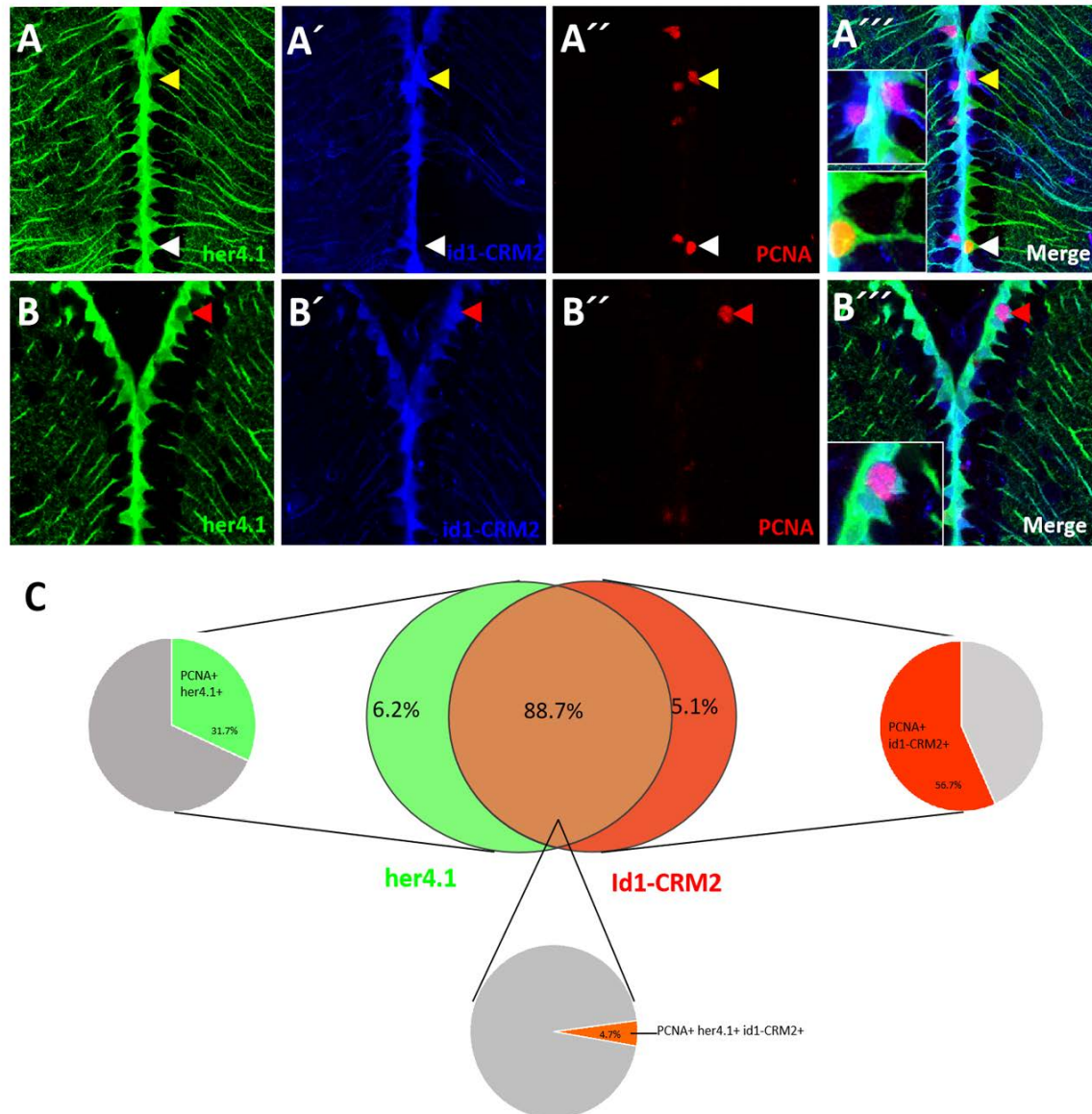


Figure S5. Both *notch3* and *her4.1* are expressed together with *id1* in the RGCs of the telencephalon. (A-B''') Immunostaining on cross sections of the telencephalon of *Tg(her4.1:gfp;id1-CRM2:mCherry)* double transgenics with antibodies against GFP (green), mCherry (blue) and the proliferation marker PCNA (red). GFP and mCherry signals co-localize indicating that *her4.1* and *id1* are co-expressed. A''' inset: magnified image of PCNA+/id1-CRM2:mCherry+ cells (yellow arrowhead, upper image) and a PCNA+/her4.1+ cell (white arrowhead, lower image). B''' inset: a magnified view of a PCNA+/id1-CRM2:mCherry+/her4.1:gfp+ cell (red arrowhead). (C) Summary of co-expression analysis of (*Tg(id1-CRM2:mCherry)*, red) and (*Tg(her4.1:gfp)*, green). Pie charts represent total number

of *Tg(her4.1:gfp)* or *Tg(id1-CRM2:mCherry)* or both *Tg(her4.1:gfp)* and *Tg(id1-CRM2:mCherry)* expressing cells and the fraction of cells co-expressing PCNA. Cells expressing either *Tg(id1-CRM2:mCherry)* alone or *Tg(her4.1:gfp)* alone show a higher proportion of PCNA+ cells than cells expressing both markers. Scale bar = 20 μ m. n=3 brains.

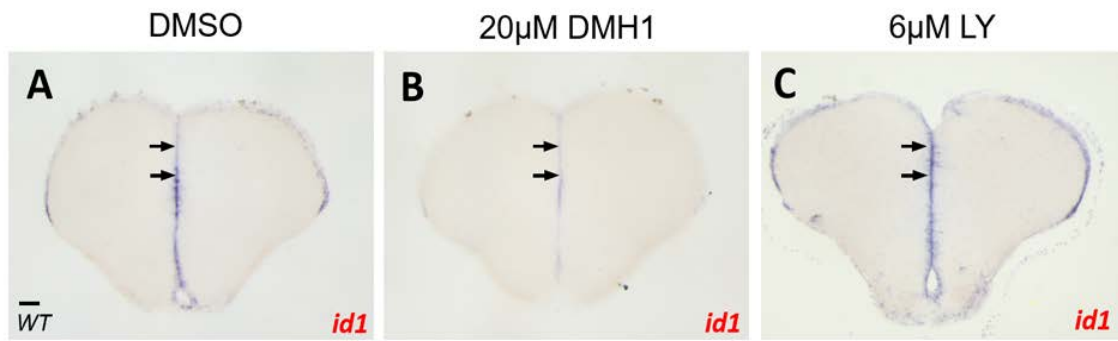


Figure S6. Expression of *id1* is independent of Notch signaling

(A-C) Expression of *id1* revealed by ISH on control brains (DMSO, A) or brains treated with different concentrations of DMH1 (B) or LY (C). (B) Inhibition of BMP signaling by 20 μ M DMH1 lead to the reduction of *id1* expression. (C) Inhibition of Notch signaling by 6 μ M LY411575 did not influence the expression of *id1* but strongly reduced *her4.1* expression (See Fig, 7C). Black arrows show the expression of *id1* in the ventricular zone of untreated (A) and treated (B and C) adult zebrafish telencephala. n=3 brains. Scale bar: 100 μ m.

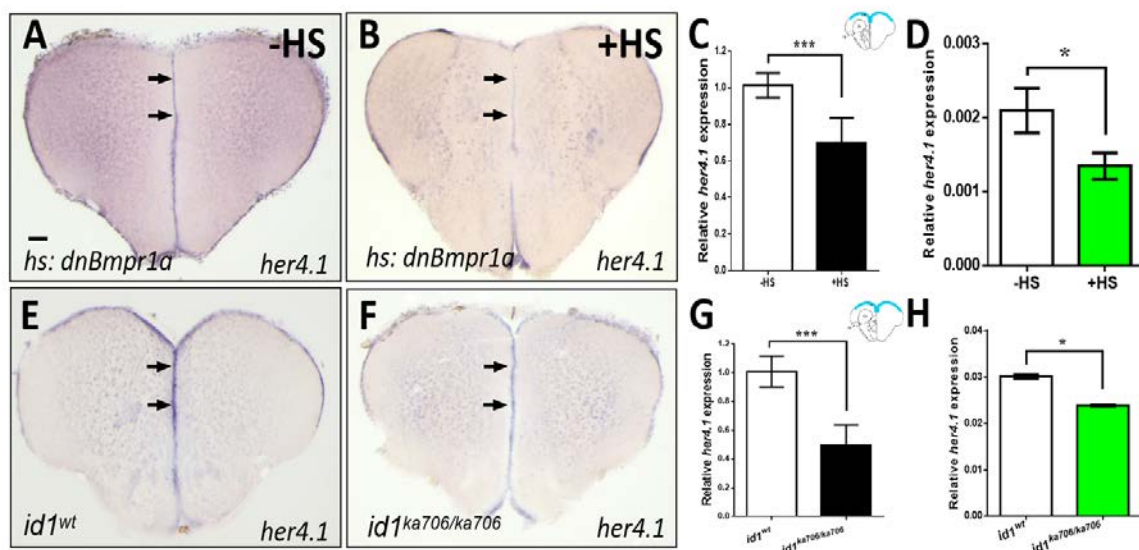


Figure S7. Inhibition of BMP signaling or deletion of *id1* leads to a reduction of *her4.1* expression.

(A-B) ISH reveals decrease of *her4.1* expression after inhibition of the BMP pathway by heat-shock of *Tg(hs:dnBmpr1a)* animals. Black arrows show the expression of the *her4.1* gene in the ventricular zone of *Tg(hs:dnBmpr1a)* animals without (A) and with heat-shock (B). (C) Quantification of *her4.1* expression in panels A and B (scheme in the upper right-hand corner displays the quantified area in blue). (D) RT-qPCR quantification of *her4.1* mRNA expression of brains shown in A-B. (E-F) Expression of *her4.1* mRNA is reduced in *id1^{ka706/ka706}* telencephala. Black arrows show the region of expression of *her4.1* in the ventricular zone of *id1^{ka706/ka706}* mutants and WT siblings. (G) Quantification of *her4.1* expression (scheme in the upper right-hand corner displays the quantified area in blue). (H) RT-qPCR quantification confirms reduction of *her4.1* in *id1^{ka706/ka706}* telencephala. HS, heat-shock. n=3 independent brains (C, G), n=15 sections D, H

Table S1. Genes co-expressed with *id1*+/*ccdn1*- RGCs/NSCs according to single cell sequencing data from (41).

Name	ID	Pearson_cor	Function	Reference
<i>smad1</i>	ENSDARG000000027199	0.448256850	signal transducer	(81)
<i>acvr2b</i>	ENSDARG000000044422	0.379378967	activin receptor type II	(82)
<i>smad5</i>	ENSDARG000000037238	0.323494119	signal transducer	(81)
<i>bambia</i>	ENSDARG000000055381	0.222851826	target gene and modulator	(83)
<i>bmpr2b</i>	ENSDARG000000020057	0.191787248	Bmp type II receptor	(73)
<i>acvr2aa</i>	ENSDARG000000011188	0.181544807	activin receptor type II	(82)
<i>bmpr1ab</i>	ENSDARG000000105045	0.088841975	Bmp type I receptor	(75)
<i>smad9</i>	ENSDARG000000021938	0.061794402	signal transducer	(39)

Table S2. Number of WT and *idl*^{ka706/ka706} adult zebrafish used in the multiple stab wound experiments.

	Start number	Fish sacrificed at 5dpl	Fish sacrificed after second stab injury	Fish survived after 3 months
<i>idl</i> ^{wt}	25	2	7	16
<i>idl</i> ^{ka706/ka706}	25	2	7	11

Table S3. PCR primer sequences for ISH probes

<i>bmp2a</i> Forward	5'-ACGCAGAGCAGGTTAGCA-3'
<i>bmp2a</i> Reverse	5'- GAAGGGACCGACTTACGC-3'
<i>bmp7a</i> Forward	5'-CAGTTGCTGCTGATTTGTTG-3'
<i>bmp7a</i> Reverse	5'-TGAACATCTGTGAGGGGATT-3'
<i>bmp7b</i> Forward	5'-CAGGTTGTGCGCTCTGGAAACGCAA-3'
<i>bmp7b</i> Reverse	5'-CCTTTTGAATCCTGGCCAATCG-3'
<i>bmpr2b</i> Forward	5'- TATTGTGCGCTGTTCTTTG-3'
<i>bmpr2b</i> Reverse	5'- GCAGATAGGCCAGTCCTCTG-3'

Table S4. Restriction enzymes and RNA polymerases used for DIG probe synthesis:

Gene name	Restriction enzyme	RNA-polymerase
<i>idl</i>	XhoI	T3
<i>her4.1</i>	CeuI	T7
<i>notch3</i>	ApaI	SP6
<i>bmp2a</i>	ApaI	SP6
<i>bmp2b</i>	SpeI	T7
<i>bmp4</i>	BamHI/SpeI	T7
<i>bmp7a</i>	SacII	SP6
<i>bmp7b</i>	NcoI	SP6
<i>bmpr1ab</i>	EcoRI	T7
<i>bmpr2b</i>	PstI	T7

Table S5. PCR primer sequences for RT-qPCR.

<i>β-actin</i> RT Fwd	5'-GCC TGA CGG ACA GGT CAT-3'
<i>β-actin</i> RT Rev	5'-ACC GCA AGA TTC CAT ACCC-3'
<i>ascl1a</i> RT Fwd	5'-ATC TCC CAA AAC TAC TCT AAT GAC ATG AAC TCT AT-3'
<i>ascl1a</i> RT Rev	5'-CAA GCG AGT GCT GAT ATT TTT AAG TTT CCT TTT AC-3'
<i>her4.1</i> RT Fwd	5'-GCT GAT ATC CTG GAG ATG ACG-3'
<i>her4.1</i> RT Rev	5'-GAC TGT GGG CTG GAG TGT GTT-3'
<i>idl</i> RT Fwd	5'-ATG TTG TCC GCT GCC TCT-3'
<i>idl</i> RT Rev	5'-GCT GGC TTT CTT GTT GGT C-3'

References

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