

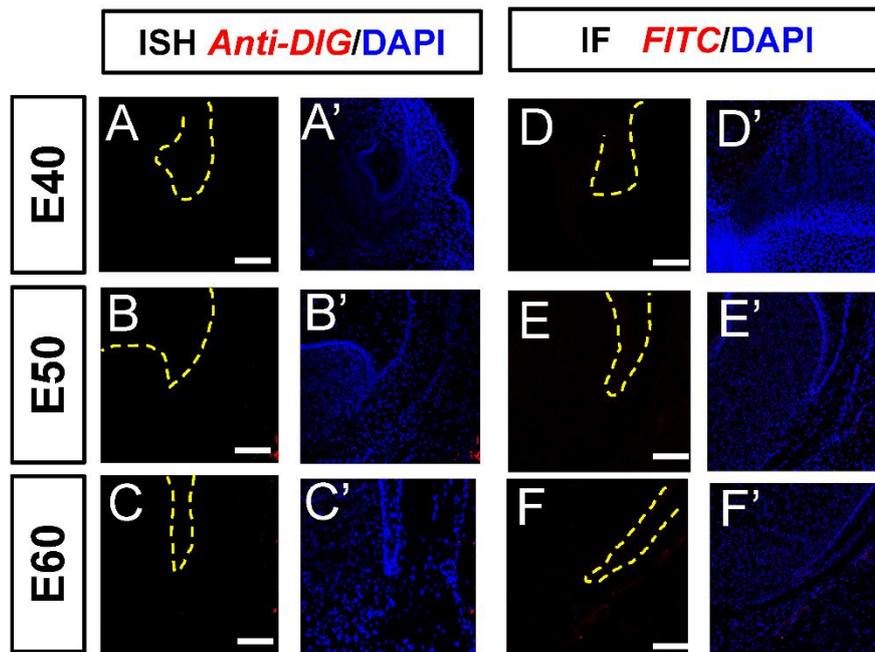
## Supplementary files

**Table S1 Primers of genes for *in situ* hybridization**

Gene name	Accession no.	Sense primer	Antisense primer	Size of PCR product (bp)
<i>FGF3</i>	XM_003122421.2	tattctccggacgctacctg	atctcgtggctccttggtgc	298
<i>FGF4</i>	XM_003122418.2	cacctaaccgcacactgga	aaagtgggtgacctcatgg	621
<i>FGF7</i>	XM_021073771.1	ggatcctgccaagtttgctc	cataggaagaaaatgggctgtt	551
<i>FGF9</i>	NM_213801.1	ggacagcccggttttgtaa	gtactttgtccgggtccact	512
<i>FGFR1</i>	XM_005671767.3	agcgacaccactactctc	accgccaagcatgtatact	708
<i>FGFR2</i>	NM_001099924.2	ggtgggaatcgacaaaagaga	agaggctgactgaggcca	828
<i>FGFR3</i>	XM_021100906.1	ccaccgacaaggagctagag	caccgacaggtccaggctact	1349
<i>SPRY2</i>	XM_001927097.5	atcagagccatccgaaacac	cacagttgtcctcgtcatcg	577
<i>SPRY4</i>	XM_003124009.4	cccttctggacagccgtat	cagaaaggcttgcgggtct	841

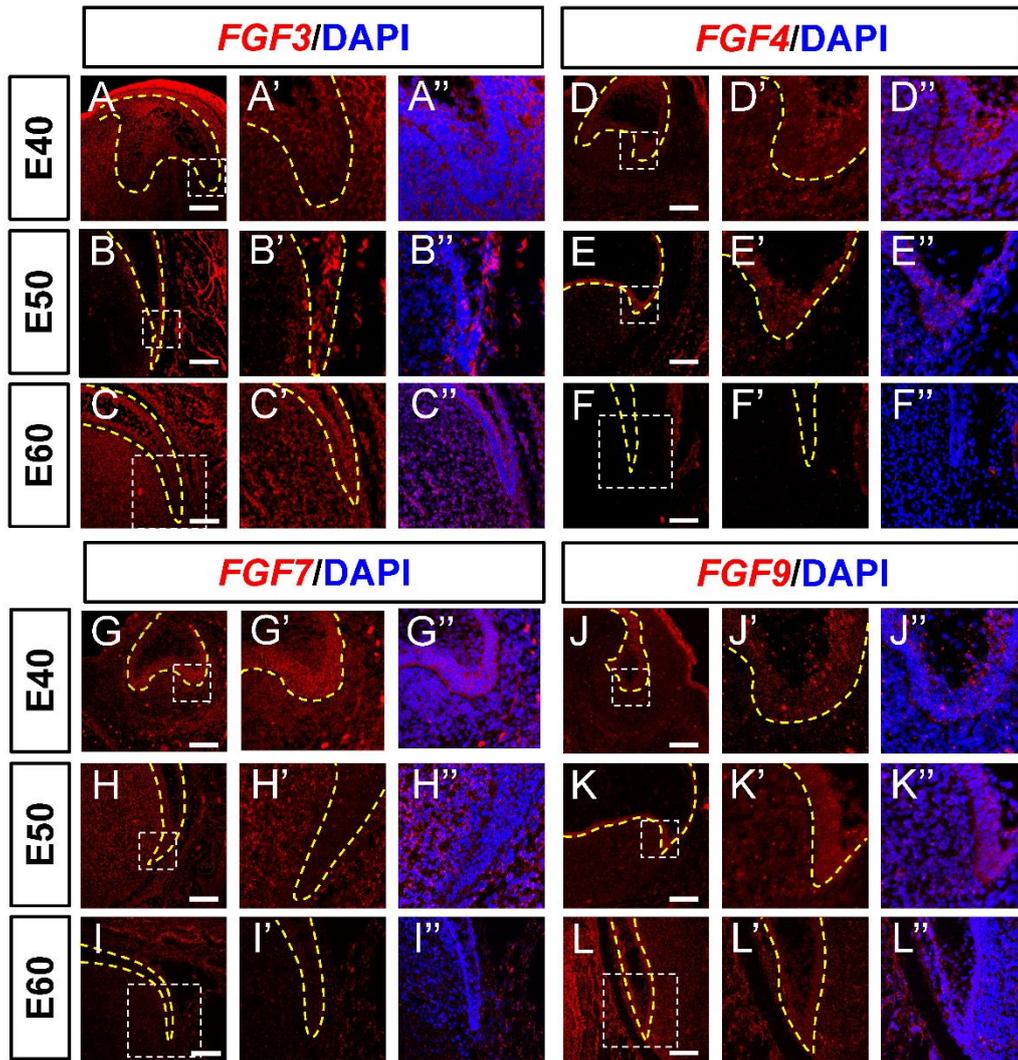
**Table S2 Primers of genes for real-time RT-PCR**

Gene name	Forward primer	Reverse primer
<i>GAPDH</i>	tcttgggtactactgaggac	ccctgttgcctgtagccaaat
<i>FGF3</i>	tattctccggacgctacctg	caccgacaggtaccacagtc
<i>FGF4</i>	gctctatggctcggctttct	tcttgcataagctgtccgt
<i>FGF7</i>	gcacaaggcagacaacagac	tcactgacctctctctattgttct
<i>FGF9</i>	acggatccgaaaaactaacca	tgccgttagtctcgttcc
<i>FGFR1</i>	gactcctaaccacacttgc	aggtgtagttgcccttgcg
<i>FGFR2</i>	aacgattacgggtccatcaa	ccgttctttccacgtgttt
<i>FGFR3</i>	gacgtgcacaacctcgacta	ccgaaggaccagacatcatc
<i>SPRY2</i>	gctcagcacaacacagagag	aagtgcaggagtgctcgtc
<i>SPRY4</i>	aagtgcaggagtgctcgtc	ctcgttgggtcagtggtaga

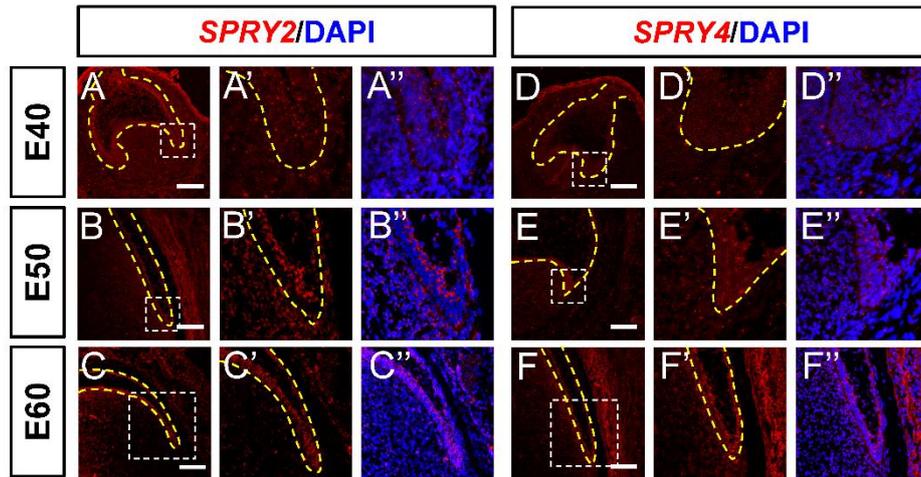


**Figure S1. Negative control for the *in situ* hybridization with TSA and immunofluorescence**

(A–C) Negative control for the *in situ* hybridization with TSA of E40, E50, and E60; Signals were in red; (A'–C') nuclei stained with DAPI (blue) for A-C; (D–F) Negative control for the immunofluorescence of E40, E50, and E60. Signals were in red; (D'–F') nuclei stained with DAPI (blue) for D-F. The yellow dotted line marked the boundary of tooth epithelium and mesenchyme. Scale bar represents 100  $\mu\text{m}$ .

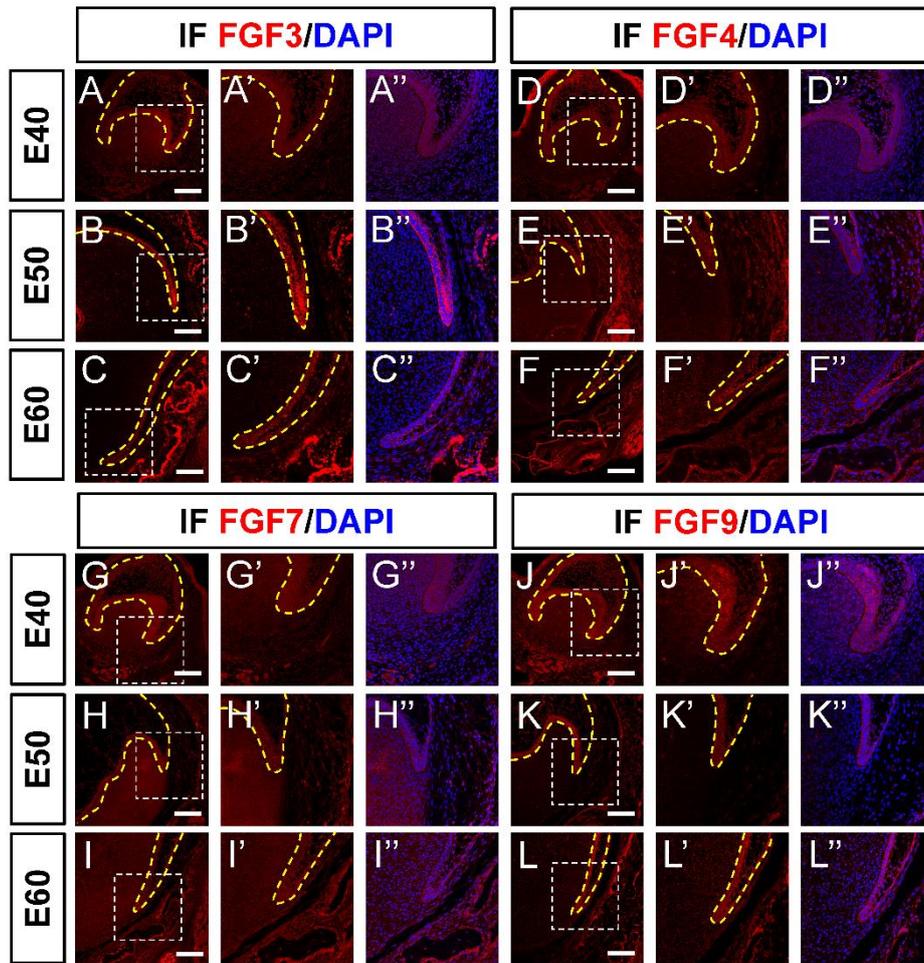


**Figure S2. Dynamic expression of genes encoding FGF ligands of cervical loop during morphogenesis of DM3.** (A–L) *In situ* hybridization (ISH) showing the mRNA expression of FGF ligands (red) and nuclei stained with DAPI (blue) from E40 to E60. White boxed regions in A–L are magnified in A'–L', and DAPI stained were overlaid in A''–L''. Expression of *FGF3* (A–C) and *FGF4* (D–F) mRNA from E40 to E60; (G–I) Expression of *FGF7* mRNA from E40 to E60; (J–L) Expression of *FGF9* mRNA E40 to E60. Yellow dotted line, boundary of tooth epithelium and mesenchyme. Scale bar represents 100  $\mu\text{m}$ .



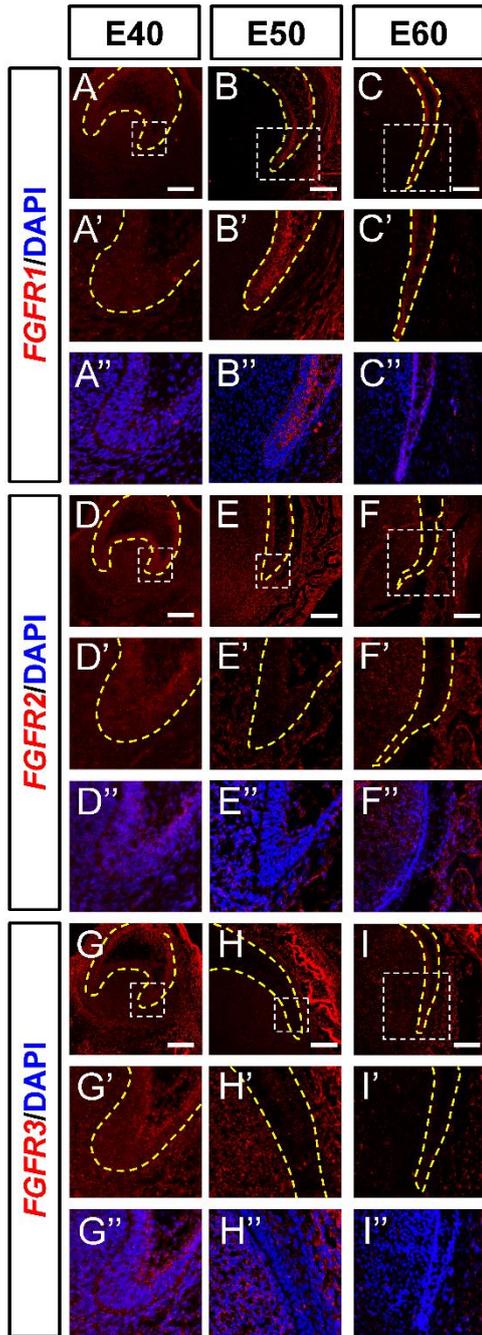
**Figure S3. Dynamic expression of genes encoding FGF antagonists of cervical loop during morphogenesis of DM3.**

(A–F) *In situ* hybridization (ISH) showing the mRNA expression of FGF antagonists (red) and nuclei stained with DAPI (blue) from E40 to E60. The white boxed regions in A–F are magnified in A'–F', and DAPI staining is overlaid in A''–F''. Expression of *SPRY2* (A–C) and *SPRY4* (D–F) mRNA from E40 to E60. Yellow dotted line, boundary of tooth epithelium and mesenchyme. Scale bar, 100  $\mu$ m.



**Figure S4. Protein expression of FGF ligands of cervical loop during morphogenesis of DM3.**

(A–F) Immunofluorescence (IF) staining shows protein expression of FGF ligands (red) and nuclei stained with DAPI (blue) from E40 to E60. White boxed regions in A–L are magnified in A'–L', and overlaid with DAPI staining in A''–L''. Expression of (A–C) FGF3, (D–F) FGF4, (G–I) FGF7 and (J–L) FGF9 from E40 to E60. Yellow dotted line, boundary of tooth epithelium and mesenchyme. Scale bar, 50  $\mu$ m.



**Figure S5. Dynamic expression of genes encoding FGF receptors of cervical loop during morphogenesis of DM3.**

(A–I) *In situ* hybridization (ISH) shows mRNA expression of FGF receptors (red) and nuclei stained with DAPI (blue) from E40 to E60. White boxed regions in A–I are magnified in A'–I', and DAPI staining is overlaid in A''–I''. Expression of (A–C) *FGFR1*, (D–F) *FGFR2*, and (G–I) *FGFR3* mRNA from E40 to E60. Yellow dotted line, boundary of tooth epithelium and mesenchyme. Scale bar, 100  $\mu$ m.