

Supplemental Table 1: Statistical Analysis of GFP-Positive Cells Counting Data

Figure Number	AVC or Vent	Condition	N	Total Number of GFP-positive cells	Number of Epithelial Cells (%)	Number of Activated Cells (%)	Number of Transformed Cells (%)
Figure 1F	Vent	GFP, 200pM TGFβ2, control siRNA	24	956	737 (77±1.4%)	111 (12±0.4%)	108 (11±0.9%)
	Vent	TGFβR3, TGFβ2, control siRNA	23	957	431 (45±2.0%)	107 (11±1.5%)	419 (44±0.6%)
	Vent	GFP, TGFβ2, Smad4A siRNA	24	960	753 (78±1.6%)	105 (11±1.0%)	102 (11±0.6%)
	Vent	GFP, TGFβ2, Smad4B siRNA	24	867	696 (80±0.8%)	83 (10±0.6%)	88 (10±0.4%)
	Vent	TGFβR3, TGFβ2, Smad4A siRNA	24	765	581 (76±1.3%)	90 (12±1.0%)	94 (12±1.2%)
	Vent	TGFβR3, TGFβ2, Smad4B siRNA	24	765	603 (79±1.4%)	83 (11±0.7%)	79 (11±1.3%)
Figure 1G	Vent	GFP, 5 nM BMP-2, control siRNA	23	763	609 (80±1.2%)	71 (9±0.4%)	83 (11±1.7%)
	Vent	TGFβR3, BMP-2, control siRNA	23	914	437 (48±0.7%)	89 (10±0.6%)	388 (42±0.3%)
	Vent	GFP, BMP-2, Smad4A siRNA	24	863	708 (82±1.6%)	72 (8±0.7%)	83 (10±1.1%)
	Vent	GFP, BMP-2,	24	846	690 (81±1.7%)	76 (9±1.1%)	80 (10±0.6%)

		Smad4B siRNA					
	Vent	TGF β R3, BMP-2, Smad4A siRNA	24	556	419 (75 \pm 2.3%)	65 (12 \pm 1.3%)	72 (13 \pm 1.8%)
	Vent	TGF β R3, BMP-2, Smad4B siRNA	24	514	404 (79 \pm 1.3%)	50 (10 \pm 1.5%)	60 (12 \pm 1.9%)
Figure 2C	Vent	GFP	39	1643	1096 (67 \pm 2.6%)	182 (11 \pm 1.4%)	365 (22 \pm 1.8%)
	Vent	Smad1	39	1292	755 (57 \pm 4.3%)	170 (14 \pm 1.3%)	367 (29 \pm 3.6%)
Figure 2D	Vent	GFP	34	1159	770 (66 \pm 0.5%)	155 (13 \pm 0.4%)	234 (20 \pm 0.6%)
	Vent	Smad3	36	971	643 (66 \pm 1.6%)	121 (12 \pm 0.1%)	207 (21 \pm 1.7%)
Figure 3A	Vent	GFP, 200pM TGF β 2, control siRNA	25	1103	869 (79 \pm 2.3%)	116 (11 \pm 1.5%)	118 (11 \pm 0.9%)
	Vent	TGF β R3, TGF β 2, control siRNA	23	716	332 (46 \pm 1.1%)***	76 (11 \pm 1.9%)	308 (43 \pm 1.2%)***
	Vent	GFP, TGF β 2, Par6cA siRNA	25	993	748 (75 \pm 3.5%)	133 (14 \pm 2.0%)	112 (12 \pm 1.7%)
	Vent	GFP, TGF β 2, Par6cB siRNA	24	910	706 (78 \pm 0.9%)	101 (11 \pm 0.1%)	103 (11 \pm 0.8%)
	Vent	TGF β R3, TGF β 2, Par6cA siRNA	24	687	541 (79 \pm 1.4%)***	75 (11 \pm 1.1%)	71 (10 \pm 0.4%)***
	Vent	TGF β R3, TGF β 2, Par6cB siRNA	24	744	572 (77 \pm 2.0%)***	91 (12 \pm 2.0%)	81 (11 \pm 0.6%)***
Figure 3B	Vent	GFP, 200pM TGF β 2, control	23	942	749 (80 \pm 3.1%)	100 (11 \pm 1.5%)	93 (10 \pm 2.1%)

		siRNA					
	Vent	TGF β R3, TGF β 2, control siRNA	22	727	354 (49 \pm 2.5%)***	72 (10 \pm 1.0%)	301 (41 \pm 3.1%)***
	Vent	GFP, TGF β 2, Smurf1A siRNA	24	945	702 (75 \pm 2.3%)	124 (13 \pm 1.9%)	119 (13 \pm 0.6%)
	Vent	GFP, TGF β 2, Smurf1B siRNA	22	832	645 (78 \pm 2.0%)	93 (11 \pm 0.9%)	94 (11 \pm 1.1%)
	Vent	TGF β R3, TGF β 2, Smurf1A siRNA	23	673	523 (78 \pm 3.5%)**	76 (11 \pm 1.6%)	74 (11 \pm 2.0%)**
	Vent	TGF β R3, TGF β 2, Smurf1B siRNA	23	758	586 (77 \pm 0.5%)**	86 (11 \pm 1.2%)	86 (11 \pm 1.3%)**
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Figure 4A	Vent	GFP, 5 nM BMP-2, vehicle (DMSO)	30	1126	849 (75 \pm 1.1%)	142 (13 \pm 1.4%)	135 (12 \pm 0.6%)
	Vent	GFP, BMP-2, 2.5 μ M SB431542	31	921	768 (83 \pm 0.7%)	72 (8 \pm 0.4%)	81 (9 \pm 0.7%)
	Vent	TGF β R3, BMP-2, vehicle	28	564	356 (63 \pm 2.0%)**	52 (9 \pm 1.6%)	156 (28 \pm 2.2%)**
	Vent	TGF β R3, BMP-2, SB431542	28	474	425 (90 \pm 0.7%)**	25 (5 \pm 0.9%)	24 (5 \pm 0.6%)**
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Figure 4C	Vent	GFP, TGF β 2, control siRNA	25	803	631 (79 \pm 1.7%)	77 (9 \pm 1.1%)	93 (12 \pm 0.6%)
	Vent	TGF β R3, TGF β 2, control siRNA	25	1027	497 (48 \pm 1.0%)***	85 (8 \pm 0.2%)	445 (43 \pm 0.8%)***
	Vent	GFP, TGF β 2, ALK5A siRNA	25	876	720 (82 \pm 0.4%)	72 (8 \pm 0.7%)	84 (10 \pm 1.1%)

	Vent	GFP, TGF β 2, ALK5B siRNA	24	831	720 (79 \pm 2.1%)	72 (9 \pm 0.4%)	84 (12 \pm 1.7%)
	Vent	TGF β R3, TGF β 2, ALK5A siRNA	24	668	529 (81 \pm 1.8%)***	60 (9 \pm 0.5%)	79 (10 \pm 1.3%)***
	Vent	TGF β R3, TGF β 2, ALK5B siRNA	23	642	523 (77 \pm 1.7%)***	56 (12 \pm 1.3%)	63 (11 \pm 1.0%)***
Figure 4D	Vent	GFP, BMP-2, control siRNA	23	763	609 (80 \pm 1.2%)	71 (9 \pm 0.4%)	83 (11 \pm 1.7%)
	Vent	TGF β R3, BMP-2, control siRNA	23	914	437 (48 \pm 0.7%)***	89 (10 \pm 0.6%)	388 (42 \pm 0.3%)***
	Vent	GFP, BMP-2, ALK5A siRNA	24	948	764 (81 \pm 0.7%)	91 (10 \pm 0.5%)	93 (10 \pm 0.2%)
	Vent	GFP, BMP-2, ALK5B siRNA	24	904	747 (83 \pm 0.7%)	70 (8 \pm 0.5%)	87 (10 \pm 0.3%)
	Vent	TGF β R3, BMP-2, ALK5A siRNA	24	470	361 (77 \pm 0.6%)***	53 (11 \pm 0.7%)	56 (11 \pm 0.9%)***
	Vent	TGF β R3, BMP-2, ALK5B siRNA	24	524	403 (77 \pm 1.7%)**	61 (12 \pm 1.3%)	60 (11 \pm 1.0%)***
Figure 4E	Vent	GFP, BMP-2, control siRNA	24	778	603 (77 \pm 1.2%)	83 (11 \pm 1.3%)	92 (12 \pm 0.4%)
	Vent	GFP, BMP-2, Par6cA siRNA	24	968	779 (80 \pm 0.6%)	82 (8 \pm 0.4%)	107 (11 \pm 1.1%)
	Vent	GFP, BMP-2, Par6cB siRNA	24	906	743 (82 \pm 0.2%)	76 (8 \pm 0.2%)	87 (10 \pm 0.3%)

	Vent	GFP, BMP-2, Smurf1A siRNA	24	921	740 (80±0.8%)	87 (9±0.3%)	94 (10±0.6%)
	Vent	GFP, BMP-2, Smurf1B siRNA	24	896	748 (84±0.6%)	69 (8±0.2%)	79 (9±0.5%)
	Vent	TGFβR3, BMP-2, control siRNA	24	843	410 (49±1.9%)***	80 (10±0.4%)	353 (42±2.2%)**
	Vent	TGFβR3, BMP-2, Par6cA siRNA	24	538	420 (78±1.9%)***	54 (10±0.9%)	61 (12±1.1%)**
	Vent	TGFβR3, BMP-2, Par6cB siRNA	24	549	430 (78±0.9%)***	54 (10±0.1%)	65 (12±0.8%)**
	Vent	TGFβR3, BMP-2, Smurf1A siRNA	24	504	387 (77±0.9%)***	63 (12±0.8%)	54 (12±0.2%)**
	Vent	TGFβR3, BMP-2, Smurf1B siRNA	24	515	398 (77±1.2%)***	62 (12±0.6%)*	55 (11±0.8%)**
Figure S1	Vent	GFP, vehicle	24	780	614 (79±1.1%)	80 (10±0.7%)	86 (11±0.6%)
	Vent	GFP, 200 pM TGFβ2	23	812	651 (80±1.6%)	87 (11±0.8%)	74 (9±0.9%)
	Vent	GFP, 5 nM BMP- 2	25	908	718 (79±0.3%)	88 (10±0.5%)	102 (11±1.2%)
	Vent	TGFβR3, vehicle	22	617	504 (82±0.8%)	58 (9±1.0%)	55 (9±1.6%)
	Vent	TGFβR3, TGFβ2	25	737	324 (44±0.9%)***	71 (10±1.1%)	342 (46±0.8%)***
	Vent	TGFβR3, BMP-2	24	743	362 (49±1.1%)***	89 (12±0.7%)	292 (39±0.2%)***
Asterisks denote significance as follows: *P<0.05, **P<0.01, ***P<0.001							

Supplemental Table 2. siRNA target sequences

ALK5A	5'(GCUACGACAUGAAAAUUTT)3'	
ALK5B	5'(GGAUUUUGCUGCCUUUUATT)3'	
Smad1A	5'(GGAGUUCGCUCAGCUCUATT)3'	
Smad1B	5'(GCAGGGAGAUGAAGAAGAATT)3'	
Smad2A	5'(GGAUUGAACUUCAUCUGAATT)3'	
Smad2B	5'(CCACCUCCAGGAUAUAUCATT)3'	
Smad3A	5'(GGAUGUAACUUGAAUUAUUUTT)3'	
Smad3B	5'(GGAUGUGCACGAUUCGGAUTT)3'	
Smad5A	5'(GCAAUACAAUGAUCCCUCATT)3'	
Smad5B	5'(GGUUUGCUCUCAAUGUUATT)3'	
TGF β R3	5'(GGAAGUAAAUCUACUUGAATT)3'	
Scrambled high GC content (62%)	5'(AGACTGTCGCGTGCTCTGTCC)3'	
Scrambled medium GC content (38%)	5'(CTCCTUGTCAATTACCGCTT)3'	
Scrambled low GC content (29%)	5'(AAAAGGAUCUGAUACGUATT)3'	

Supplemental Table 3. Primer pairs

Target	Forward (5'→3')	Reverse (5'→3')
Smad4	GTCGCCACAGACAGATGCAACAACA	TTTGACGAAGCTCATGCGGAGGAT
ALK5	ACCAGAGTGGCGTGTAAAGAAGGT	TGC ACAGAAAGGACCCAAGCAAC
GAPDH	GGGCACGCCATCACTATCTTCC	GAGGGGCCATCCACCGTCTT
Smad1	CTGCTCTCCAATGTTAACCG	GCATTCA CGCGTACACTTCTC
Smad2	TTTGTTAGAGCCCCAAC	ACCTGATTCA CAGCCTGAG
Smad3	TGTAAACCAGGGCTTCGA	AGGTGCAGCTCAATCCA
Smad5	ATGTGCACCATCCGAATG	GGGTGCTTGTGACATCTT
TGF β R3	GCTGAAGGCAGCAGCTTGATTAT	ACATTGGTGATGTGAGGCTCAGGA