

**Supplementary Table 1 Correlation of DLX genes with clinical characteristics of COAD patients**

**DLX1**

Characteristic	Low expression of	High expression of	p	statistic	method
	DLX1	DLX1			
n	239	239			
T stage, n (%)			0.09	6.5	Chisq.test
T1	7 (1.5%)	4 (0.8%)			
T2	39 (8.2%)	44 (9.2%)			
T3	171 (35.8%)	152 (31.9%)			
T4	22 (4.6%)	38 (8%)			
N stage, n (%)			0.1	4.61	Chisq.test
N0	148 (31%)	136 (28.5%)			
N1	57 (11.9%)	51 (10.7%)			
N2	34 (7.1%)	52 (10.9%)			
M stage, n (%)			0.703	0.15	Chisq.test
M0	176 (42.4%)	173 (41.7%)			
M1	31 (7.5%)	35 (8.4%)			
Pathologic stage, n (%)			0.519	2.27	Chisq.test
Stage I	38 (8.1%)	43 (9.2%)			
Stage II	102 (21.8%)	85 (18.2%)			
Stage III	64 (13.7%)	69 (14.8%)			
Stage IV	31 (6.6%)	35 (7.5%)			
Primary therapy					
outcome, n (%)			0.698		Fisher.test
PD	14 (5.6%)	11 (4.4%)			
SD	3 (1.2%)	1 (0.4%)			
PR	6 (2.4%)	7 (2.8%)			
CR	102 (40.8%)	106 (42.4%)			

Gender, n (%)			0.314	1.02	Chisq.test
Female	107 (22.4%)	119 (24.9%)			
Male	132 (27.6%)	120 (25.1%)			
Race, n (%)			0.589	1.06	Chisq.test
Asian	4 (1.3%)	7 (2.3%)			
Black or African American					
American	31 (10.1%)	32 (10.5%)			
White	120 (39.2%)	112 (36.6%)			
Age, n (%)			0.226	1.47	Chisq.test
<=65	104 (21.8%)	90 (18.8%)			
>65	135 (28.2%)	149 (31.2%)			
Weight, n (%)			0.461	0.54	Chisq.test
<=90	95 (34.8%)	94 (34.4%)			
>90	47 (17.2%)	37 (13.6%)			
Height, n (%)			0.997	0	Chisq.test
<170	66 (25.8%)	61 (23.8%)			
>=170	66 (25.8%)	63 (24.6%)			
BMI, n (%)			0.533	0.39	Chisq.test
<25	42 (16.4%)	45 (17.6%)			
>=25	90 (35.2%)	79 (30.9%)			
Residual tumor, n (%)			0.179		Fisher.test
R0	171 (45.7%)	175 (46.8%)			
R1	0 (0%)	4 (1.1%)			
R2	11 (2.9%)	13 (3.5%)			
CEA level, n (%)			0.621	0.25	Chisq.test
<=5	97 (32%)	99 (32.7%)			
>5	49 (16.2%)	58 (19.1%)			
Perineural invasion, n (%)			0.763	0.09	Chisq.test

NO	65 (35.9%)	70 (38.7%)			
YES	24 (13.3%)	22 (12.2%)			
Lymphatic invasion, n (%)			0.758	0.1	Chisq.test
NO	134 (30.9%)	132 (30.4%)			
YES	88 (20.3%)	80 (18.4%)			
History of colon polyps, n (%)			0.302	1.07	Chisq.test
NO	132 (32.4%)	130 (31.9%)			
YES	65 (15.9%)	81 (19.9%)			
Neoplasm type, n (%)			1		Fisher.test
Colon adenocarcinoma	239 (50%)	239 (50%)			
Rectum adenocarcinoma	0 (0%)	0 (0%)			
Colon polyps present, n (%)			0.142	2.16	Chisq.test
NO	74 (29.7%)	88 (35.3%)			
YES	49 (19.7%)	38 (15.3%)			
Age, median (IQR)	68 (57.5, 75)	70 (60, 79)	0.021	25089.5	Wilcoxon

### DLX2

Characteristic	Low expression of DLX2	High expression of DLX2	p	statistic	method
	DLX2	DLX2			
n	239	239			
T stage, n (%)			0.375	3.11	Chisq.test
T1	6 (1.3%)	5 (1%)			
T2	45 (9.4%)	38 (8%)			
T3	163 (34.2%)	160 (33.5%)			
T4	24 (5%)	36 (7.5%)			
N stage, n (%)			0.2	3.22	Chisq.test
N0	150 (31.4%)	134 (28%)			

N1	53 (11.1%)	55 (11.5%)			
N2	36 (7.5%)	50 (10.5%)			
M stage, n (%)			0.005	7.9	Chisq.test
M0	180 (43.4%)	169 (40.7%)			
M1	21 (5.1%)	45 (10.8%)			
Pathologic stage, n (%)			0.014	10.64	Chisq.test
Stage I	45 (9.6%)	36 (7.7%)			
Stage II	100 (21.4%)	87 (18.6%)			
Stage III	67 (14.3%)	66 (14.1%)			
Stage IV	21 (4.5%)	45 (9.6%)			
Primary therapy					
outcome, n (%)			0.036		Fisher.test
PD	12 (4.8%)	13 (5.2%)			
SD	3 (1.2%)	1 (0.4%)			
PR	2 (0.8%)	11 (4.4%)			
CR	110 (44%)	98 (39.2%)			
Gender, n (%)			0.927	0.01	Chisq.test
Female	112 (23.4%)	114 (23.8%)			
Male	127 (26.6%)	125 (26.2%)			
Race, n (%)			0.802	0.44	Chisq.test
Asian	5 (1.6%)	6 (2%)			
Black or African					
American	33 (10.8%)	30 (9.8%)			
White	127 (41.5%)	105 (34.3%)			
Age, n (%)			0.226	1.47	Chisq.test
<=65	104 (21.8%)	90 (18.8%)			
>65	135 (28.2%)	149 (31.2%)			
Weight, n (%)			0.094	2.8	Chisq.test
<=90	97 (35.5%)	92 (33.7%)			

>90	53 (19.4%)	31 (11.4%)			
Height, n (%)			0.812	0.06	Chisq.test
<170	69 (27%)	58 (22.7%)			
>=170	73 (28.5%)	56 (21.9%)			
BMI, n (%)			0.207	1.6	Chisq.test
<25	43 (16.8%)	44 (17.2%)			
>=25	99 (38.7%)	70 (27.3%)			
Residual tumor, n (%)			0.002		Fisher.test
R0	174 (46.5%)	172 (46%)			
R1	0 (0%)	4 (1.1%)			
R2	5 (1.3%)	19 (5.1%)			
CEA level, n (%)			0.252	1.31	Chisq.test
<=5	101 (33.3%)	95 (31.4%)			
>5	47 (15.5%)	60 (19.8%)			
Perineural invasion, n (%)			0.959	0	Chisq.test
NO	73 (40.3%)	62 (34.3%)			
YES	24 (13.3%)	22 (12.2%)			
Lymphatic invasion, n (%)			0.013	6.23	Chisq.test
NO	148 (34.1%)	118 (27.2%)			
YES	72 (16.6%)	96 (22.1%)			
History of colon polyps, n (%)			0.565	0.33	Chisq.test
NO	133 (32.6%)	129 (31.6%)			
YES	69 (16.9%)	77 (18.9%)			
Neoplasm type, n (%)			1		Fisher.test
Colon adenocarcinoma	239 (50%)	239 (50%)			
Rectum adenocarcinoma	0 (0%)	0 (0%)			

Colon polyps present, n (%)		1	0	Chisq.test
NO	87 (34.9%)	75 (30.1%)		
YES	47 (18.9%)	40 (16.1%)		
Age, median (IQR)	69 (57, 76)	69 (60, 78)	0.069	25817
DLX3				Wilcoxon
	<b>Low expression of DLX3</b>	<b>High expression of DLX3</b>		
Characteristic	DLX3	DLX3	p	statistic
n	239	239		
T stage, n (%)			0.618	1.79
T1	5 (1%)	6 (1.3%)		
T2	40 (8.4%)	43 (9%)		
T3	168 (35.2%)	155 (32.5%)		
T4	26 (5.5%)	34 (7.1%)		
N stage, n (%)			< 0.001	14.27
N0	160 (33.5%)	124 (25.9%)		
N1	50 (10.5%)	58 (12.1%)		
N2	29 (6.1%)	57 (11.9%)		
M stage, n (%)			< 0.001	14.77
M0	183 (44.1%)	166 (40%)		
M1	17 (4.1%)	49 (11.8%)		
Pathologic stage, n (%)			< 0.001	26.91
Stage I	41 (8.8%)	40 (8.6%)		
Stage II	116 (24.8%)	71 (15.2%)		
Stage III	62 (13.3%)	71 (15.2%)		
Stage IV	17 (3.6%)	49 (10.5%)		
Primary therapy outcome, n (%)			0.088	Fisher.test
PD	9 (3.6%)	16 (6.4%)		

SD	0 (0%)	4 (1.6%)			
PR	6 (2.4%)	7 (2.8%)			
CR	109 (43.6%)	99 (39.6%)			
Gender, n (%)			0.054	3.7	Chisq.test
Female	124 (25.9%)	102 (21.3%)			
Male	115 (24.1%)	137 (28.7%)			
Race, n (%)			0.971	0.06	Chisq.test
Asian	6 (2%)	5 (1.6%)			
Black or African American					
American	32 (10.5%)	31 (10.1%)			
White	118 (38.6%)	114 (37.3%)			
Age, n (%)			0.05	3.83	Chisq.test
<=65	86 (18%)	108 (22.6%)			
>65	153 (32%)	131 (27.4%)			
Weight, n (%)			0.29	1.12	Chisq.test
<=90	91 (33.3%)	98 (35.9%)			
>90	47 (17.2%)	37 (13.6%)			
Height, n (%)			0.045	4.01	Chisq.test
<170	73 (28.5%)	54 (21.1%)			
>=170	57 (22.3%)	72 (28.1%)			
BMI, n (%)			0.658	0.2	Chisq.test
<25	42 (16.4%)	45 (17.6%)			
>=25	88 (34.4%)	81 (31.6%)			
Residual tumor, n (%)			< 0.001		Fisher.test
R0	182 (48.7%)	164 (43.9%)			
R1	2 (0.5%)	2 (0.5%)			
R2	3 (0.8%)	21 (5.6%)			
CEA level, n (%)			1	0	Chisq.test
<=5	94 (31%)	102 (33.7%)			

>5	51 (16.8%)	56 (18.5%)			
Perineural invasion, n (%)			0.968	0	Chisq.test
NO	67 (37%)	68 (37.6%)			
YES	22 (12.2%)	24 (13.3%)			
Lymphatic invasion, n (%)			0.365	0.82	Chisq.test
NO	143 (32.9%)	123 (28.3%)			
YES	82 (18.9%)	86 (19.8%)			
History of colon polyps, n (%)			0.616	0.25	Chisq.test
NO	130 (31.9%)	132 (32.4%)			
YES	77 (18.9%)	69 (16.9%)			
Neoplasm type, n (%)			1		Fisher.test
Colon adenocarcinoma	239 (50%)	239 (50%)			
Rectum adenocarcinoma	0 (0%)	0 (0%)			
Colon polyps present, n (%)			1	0	Chisq.test
NO	81 (32.5%)	81 (32.5%)			
YES	44 (17.7%)	43 (17.3%)			
Age, median (IQR)	69 (61, 78.5)	68 (55.5, 76)	0.012	32364	Wilcoxon

#### DLX4

Characteristic	Low expression of DLX4		High expression of DLX4		
	DLX4	DLX4	p	statistic	method
n	239	239			
T stage, n (%)			0.309	3.59	Chisq.test
T1	4 (0.8%)	7 (1.5%)			
T2	38 (8%)	45 (9.4%)			
T3	171 (35.8%)	152 (31.9%)			

T4	26 (5.5%)	34 (7.1%)			
N stage, n (%)			0.336	2.18	Chisq.test
N0	148 (31%)	136 (28.5%)			
N1	54 (11.3%)	54 (11.3%)			
N2	37 (7.7%)	49 (10.3%)			
M stage, n (%)			0.944	0	Chisq.test
M0	175 (42.2%)	174 (41.9%)			
M1	34 (8.2%)	32 (7.7%)			
Pathologic stage, n (%)			0.063	7.3	Chisq.test
Stage I	34 (7.3%)	47 (10.1%)			
Stage II	107 (22.9%)	80 (17.1%)			
Stage III	60 (12.8%)	73 (15.6%)			
Stage IV	34 (7.3%)	32 (6.9%)			
Primary therapy					
outcome, n (%)			0.618		Fisher.test
PD	12 (4.8%)	13 (5.2%)			
SD	1 (0.4%)	3 (1.2%)			
PR	5 (2%)	8 (3.2%)			
CR	107 (42.8%)	101 (40.4%)			
Gender, n (%)			0.647	0.21	Chisq.test
Female	110 (23%)	116 (24.3%)			
Male	129 (27%)	123 (25.7%)			
Race, n (%)			0.512	1.34	Chisq.test
Asian	4 (1.3%)	7 (2.3%)			
Black or African					
American	30 (9.8%)	33 (10.8%)			
White	121 (39.5%)	111 (36.3%)			
Age, n (%)			0.926	0.01	Chisq.test
<=65	98 (20.5%)	96 (20.1%)			

>65	141 (29.5%)	143 (29.9%)			
Weight, n (%)			0.77	0.09	Chisq.test
<=90	96 (35.2%)	93 (34.1%)			
>90	45 (16.5%)	39 (14.3%)			
Height, n (%)			0.315	1.01	Chisq.test
<170	70 (27.3%)	57 (22.3%)			
>=170	62 (24.2%)	67 (26.2%)			
BMI, n (%)			0.25	1.32	Chisq.test
<25	40 (15.6%)	47 (18.4%)			
>=25	92 (35.9%)	77 (30.1%)			
Residual tumor, n (%)			0.243		Fisher.test
R0	181 (48.4%)	165 (44.1%)			
R1	1 (0.3%)	3 (0.8%)			
R2	9 (2.4%)	15 (4%)			
CEA level, n (%)			0.55	0.36	Chisq.test
<=5	85 (28.1%)	111 (36.6%)			
>5	51 (16.8%)	56 (18.5%)			
Perineural invasion, n (%)			0.903	0.01	Chisq.test
NO	65 (35.9%)	70 (38.7%)			
YES	21 (11.6%)	25 (13.8%)			
Lymphatic invasion, n (%)			0.498	0.46	Chisq.test
NO	140 (32.3%)	126 (29%)			
YES	82 (18.9%)	86 (19.8%)			
History of colon polyps, n (%)			0.417	0.66	Chisq.test
NO	124 (30.4%)	138 (33.8%)			
YES	76 (18.6%)	70 (17.2%)			

Neoplasm type, n (%)			1		Fisher.test
Colon adenocarcinoma	239 (50%)	239 (50%)			
Rectum adenocarcinoma	0 (0%)	0 (0%)			
Colon polyps present, n (%)			0.963	0	Chisq.test
NO	80 (32.1%)	82 (32.9%)			
YES	44 (17.7%)	43 (17.3%)			
Age, median (IQR)	69 (58, 76)	68 (58.5, 78)	0.484	27503.5	Wilcoxon
<hr/>					
DLX5					
	Low expression of DLX5	High expression of DLX5			
Characteristic	DLX5	DLX5	p	statistic	method
n	239	239			
T stage, n (%)			< 0.001	17.26	Chisq.test
T1	6 (1.3%)	5 (1%)			
T2	56 (11.7%)	27 (5.7%)			
T3	156 (32.7%)	167 (35%)			
T4	20 (4.2%)	40 (8.4%)			
N stage, n (%)			< 0.001	21.17	Chisq.test
N0	165 (34.5%)	119 (24.9%)			
N1	47 (9.8%)	61 (12.8%)			
N2	27 (5.6%)	59 (12.3%)			
M stage, n (%)			0.005	7.83	Chisq.test
M0	185 (44.6%)	164 (39.5%)			
M1	22 (5.3%)	44 (10.6%)			
Pathologic stage, n (%)			< 0.001	24.08	Chisq.test
Stage I	56 (12%)	25 (5.4%)			
Stage II	100 (21.4%)	87 (18.6%)			
Stage III	55 (11.8%)	78 (16.7%)			
Stage IV	22 (4.7%)	44 (9.4%)			

Primary therapy				
outcome, n (%)			0.005	Fisher.test
PD	5 (2%)	20 (8%)		
SD	3 (1.2%)	1 (0.4%)		
PR	7 (2.8%)	6 (2.4%)		
CR	114 (45.6%)	94 (37.6%)		
Gender, n (%)			0.314	1.02
Female	107 (22.4%)	119 (24.9%)		
Male	132 (27.6%)	120 (25.1%)		
Race, n (%)			0.3	Fisher.test
Asian	7 (2.3%)	4 (1.3%)		
Black or African				
American	24 (7.8%)	39 (12.7%)		
White	101 (33%)	131 (42.8%)		
Age, n (%)			< 0.001	13.2
<=65	77 (16.1%)	117 (24.5%)		
>65	162 (33.9%)	122 (25.5%)		
Weight, n (%)			0.354	0.86
<=90	85 (31.1%)	104 (38.1%)		
>90	32 (11.7%)	52 (19%)		
Height, n (%)			0.321	0.99
<170	57 (22.3%)	70 (27.3%)		
>=170	49 (19.1%)	80 (31.2%)		
BMI, n (%)			0.898	0.02
<25	37 (14.5%)	50 (19.5%)		
>=25	69 (27%)	100 (39.1%)		
Residual tumor, n (%)			0.533	Fisher.test
R0	177 (47.3%)	169 (45.2%)		
R1	1 (0.3%)	3 (0.8%)		

R2	11 (2.9%)	13 (3.5%)			
CEA level, n (%)			0.133	2.26	Chisq.test
<=5	96 (31.7%)	100 (33%)			
>5	42 (13.9%)	65 (21.5%)			
Perineural invasion, n (%)			0.023	5.17	Chisq.test
NO	63 (34.8%)	72 (39.8%)			
YES	12 (6.6%)	34 (18.8%)			
Lymphatic invasion, n (%)			< 0.001	14.88	Chisq.test
NO	150 (34.6%)	116 (26.7%)			
YES	62 (14.3%)	106 (24.4%)			
History of colon polyps, n (%)			0.009	6.9	Chisq.test
NO	121 (29.7%)	141 (34.6%)			
YES	88 (21.6%)	58 (14.2%)			
Neoplasm type, n (%)			1		Fisher.test
Colon adenocarcinoma	239 (50%)	239 (50%)			
Rectum adenocarcinoma	0 (0%)	0 (0%)			
Colon polyps present, n (%)			0.054	3.73	Chisq.test
NO	60 (24.1%)	102 (41%)			
YES	44 (17.7%)	43 (17.3%)			
Age, median (IQR)	71 (62.5, 78)	66 (54.5, 75.5)	< 0.001	34317.5	Wilcoxon

### DLX6

Characteristic	Low expression of DLX6		High expression of DLX6		statistic	method
	DLX6	p	DLX6	p		
n	239		239			
T stage, n (%)				0.119	5.86	Chisq.test

T1	2 (0.4%)	9 (1.9%)			
T2	40 (8.4%)	43 (9%)			
T3	169 (35.4%)	154 (32.3%)			
T4	27 (5.7%)	33 (6.9%)			
N stage, n (%)			0.567	1.13	Chisq.test
N0	147 (30.8%)	137 (28.7%)			
N1	53 (11.1%)	55 (11.5%)			
N2	39 (8.2%)	47 (9.8%)			
M stage, n (%)			1	0	Chisq.test
M0	179 (43.1%)	170 (41%)			
M1	34 (8.2%)	32 (7.7%)			
Pathologic stage, n (%)			0.479	2.48	Chisq.test
Stage I	38 (8.1%)	43 (9.2%)			
Stage II	101 (21.6%)	86 (18.4%)			
Stage III	61 (13.1%)	72 (15.4%)			
Stage IV	34 (7.3%)	32 (6.9%)			
Primary therapy					
outcome, n (%)			0.892		Fisher.test
PD	12 (4.8%)	13 (5.2%)			
SD	2 (0.8%)	2 (0.8%)			
PR	7 (2.8%)	6 (2.4%)			
CR	92 (36.8%)	116 (46.4%)			
Gender, n (%)			0.647	0.21	Chisq.test
Female	110 (23%)	116 (24.3%)			
Male	129 (27%)	123 (25.7%)			
Race, n (%)			0.054	5.85	Chisq.test
Asian	8 (2.6%)	3 (1%)			
Black or African					
American	25 (8.2%)	38 (12.4%)			

White	124 (40.5%)	108 (35.3%)			
Age, n (%)			0.78	0.08	Chisq.test
<=65	95 (19.9%)	99 (20.7%)			
>65	144 (30.1%)	140 (29.3%)			
Weight, n (%)			0.299	1.08	Chisq.test
<=90	89 (32.6%)	100 (36.6%)			
>90	46 (16.8%)	38 (13.9%)			
Height, n (%)			0.104	2.64	Chisq.test
<170	70 (27.3%)	57 (22.3%)			
>=170	57 (22.3%)	72 (28.1%)			
BMI, n (%)			1	0	Chisq.test
<25	43 (16.8%)	44 (17.2%)			
>=25	84 (32.8%)	85 (33.2%)			
Residual tumor, n (%)			1		Fisher.test
R0	168 (44.9%)	178 (47.6%)			
R1	2 (0.5%)	2 (0.5%)			
R2	12 (3.2%)	12 (3.2%)			
CEA level, n (%)			0.412	0.67	Chisq.test
<=5	99 (32.7%)	97 (32%)			
>5	48 (15.8%)	59 (19.5%)			
Perineural invasion, n (%)			0.372	0.8	Chisq.test
NO	67 (37%)	68 (37.6%)			
YES	27 (14.9%)	19 (10.5%)			
Lymphatic invasion, n (%)			0.826	0.05	Chisq.test
NO	134 (30.9%)	132 (30.4%)			
YES	82 (18.9%)	86 (19.8%)			

History of colon polyps, n (%)		0.724	0.12	Chisq.test
NO	130 (31.9%)	132 (32.4%)		
YES	69 (16.9%)	77 (18.9%)		
Neoplasm type, n (%)		1		Fisher.test
Colon adenocarcinoma	239 (50%)	239 (50%)		
Rectum adenocarcinoma	0 (0%)	0 (0%)		
Colon polyps present, n (%)		0.963	0	Chisq.test
NO	80 (32.1%)	82 (32.9%)		
YES	44 (17.7%)	43 (17.3%)		
Age, median (IQR)	69 (59, 77)	68 (57, 77.5)	0.547	29470.5
				Wilcoxon

**Supplementary Table 2 The list of genes associated with DLX genes**

Gene names	Gene names	Gene names
ABCC2	FGF19	PIP4K2B
ADAMTSL2	FGF20	PRDM11
AKAP7	FOXC2	PRR9
AL035425.2	FXYD1	PRSS56
AMBN	FZD7	PTP4A3
AMER3	GAL	QSOX2
APCDD1	HAVCR1	RASL11B
ARHGAP36	HCN4	RIMS4
ASPSCR1	HEY1	RTL5
ATOH8	HS3ST4	SATB2
ATXN7L2	ID4	SBK2
AXIN2	IGDCC3	SERPINI1
BCAM	IGF2BP1	SEZ6
BCORL1	ISM1	SH3GL2
BSX	IVL	SHISA7
C10orf82	KCNN1	SLC22A11
C1QL4	KLF14	SLC30A2
CAB39L	KRT23	SLC38A3
CACNG7	KRT31	SLC38A4
CDK5R1	KRT35	SLC38A8
CELSR2	KRT37	SLC46A1
CER1	KRTAP4-6	SLITRK1
CHRND	LHX3	SMAD7
CHST13	LMX1A	SP7
CILP2	LRRC43	SPATA31A5
CKB	LY6G6D	SRCIN1
CLDN19	MAFA	STK36

CNPY1	MAGEF1	STRIT1
COCH	METAP1D	SYT6
COMP	MFAP2	SYT7
CWH43	MGP	TCP11X1
DBF4B	MTARC1	TESMIN
DEFB106B	MYH7B	TFAP2B
DLX1	MYL3	TLCD3B
DLX2	NCAN	TNFAIP8L3
DLX3	NELL2	TNFRSF19
DLX4	NEUROD1	TRIM71
DLX5	NEUROD4	TTLL4
DLX6	NKD1	TTPA
DMRT2	NOTUM	UNK
DNMT3A	NXF2B	VENTX
DNMT3B	NXPH3	VWA5B2
DYNC1I1	OGDHL	WNT3
EDC3	OLFML3	WNT6
ELavl2	OR12D2	ZDHHC22
ETNK2	PABPC1L	ZNF385C
F10	PCSK2	ZNF620
FBXO17	PGBD5	
FGF1	PHF12	
FGF18	PHYHIPL	

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### Supplementary Table 3 GO and KEGG analysis of genes associated with DLX genes

ONT	ID	Descript	Gen	BgR	pval	p.adj	qval	geneID	Co
OLO		ion	eRat	atio	ue	ust	ue		un
GY		io							t
BP	GO:0007389	pattern specification	15/1	446/35	8.352	0.001	0.001	DLX1/DLX2/CELSR2/FGF1/FOXC2/NEUROD1/WNT3A/WNT6/LHX3/AXIN2/CER1/DMRT2/SATB2/HEY1/NKD1	15
BP	GO:0030022	regionalization	13/1	351/35	1.632	0.001	0.001	DLX1/DLX2/CELSR2/FGF1/FOXC2/NEUROD1/WNT3A/LHX3/AXIN2/CER1/DMRT2/HEY1/NKD1	13
BP	GO:0048705	skeletal system morphogenesis	10/1	239/35	9.526	0.006	0.005	COMP/COCH/DLX2/DLX5/FOXC2/MGP/AXIN2/FGF18/CER1/SATB2	10

BP	GO:0	endocho	6/13	73/1	1.462	0.007	0.006	COMP/COCH/DLX5/AXIN2/FGF18/CER1	6
	06035	ndral	5	8670	73E-	17836	51686		
	0	bone			05	9	7		
			morpho						
			genesis						
BP	GO:0	hippoca	6/13	81/1	2.659	0.007	0.006	DLX1/DLX2/ID4/LMX1A/NEUROD1/CDK5R1	6
	02176	mpus	5	8670	95E-	62249	92006		
	6	develop			05	9	9		
			ment						
BP	GO:0	cell fate	10/1	270/	2.729	0.007	0.006	DLX1/DLX2/FOXC2/NEUROD1/WNT3/WNT6/LHX3/	10
	04516	commit	35	1867	95E-	62249	92006	FZD7/SATB2/NEUROD4	
	5	ment		0	05	9	9		
BP	GO:0	connecti	10/1	273/	2.999	0.007	0.006	COMP/COCH/DLX2/FOXC2/ID4/MGP/AXIN2/FGF18	10
	06144	ve tissue	35	1867	3E-05	62249	92006	/CER1/SATB2	
	8	develop		0		9	9		
			ment						

BP	GO:000995	anterior/posterior pattern specification	9/13	219/5	3.106	0.007	0.006	CELSR2/FOXC2/NEUROD1/WNT3/AXIN2/CER1/DMRT2/HEY1/NKD1	9
2	or		0	05	9	9			
BP	GO:002187	forebrain neuron fate commitment	3/13	10/1	4.275	0.009	0.008	DLX1/DLX2/SATB2	3
7	neuron		5	8670	21E-05	32471	46542		
BP	GO:004358	skin development	12/1	419/35	5.366	0.010	0.009	COMP/IVL/KRT31/KRT35/TFAP2B/KRT37/KRT23/GAL/TNFRSF19/KRTAP4-6/APCDD1/PRR9	12
8	ment		0	05	7	1			

BP	GO:02195	central nervous system	8/13	183/5	5.614	0.010	0.009	DLX1/DLX2/DLX5/ID4/LMX1A/WNT3/LHX3/SATB2	8
		neuron		1867	21E-01881		09556		
		differentiation		0	05	7	1		
BP	GO:00340	neural retina development	5/13	62/15	8.531	0.013	0.012	DLX1/DLX2/NEUROD1/TFAP2B/NEUROD4	5
		ment		8670	53E-95616	67007			
BP	GO:04871	negative regulation of oligodendrocyte differentiation	3/13	13/15	0.000	0.015	0.013	DLX1/DLX2/ID4	3
				8670	10028	14288	74743		
				4	9	9			



BP	GO:0	artery	5/13	76/1	0.000	0.023	0.021	COMP/FOXC2/SMAD7/TFAP2B/HEY1	5
	04884	morpho	5	8670	22466	21132	07234		
	4	genesis			4	3	8		
BP	GO:0	pallium	7/13	172/	0.000	0.025	0.022	DLX1/DLX2/ID4/LMX1A/NEUROD1/CDK5R1/IGF2BP	7
	02154	develop	5	1867	25664	18971	86842	1	
	3	ment		0	5	2	5		
BP	GO:0	regulati	9/13	292/	0.000	0.025	0.023	DLX1/FGF1/SMAD7/TFAP2B/FGF18/CER1/ADAMTS	9
	09028	on of	5	1867	27720	91214	52428	L2/PTP4A3/RASL11B	
	7	cellular		0	6				
		response							
		to							
		growth							
		factor							
		stimulus							
BP	GO:0	eye	10/1	362/	0.000	0.027	0.024	DLX1/DLX2/FOXC2/MFAP2/NEUROD1/TFAP2B/WN	10
	00165	develop	35	1867	30560	26810	75528	T6/NEUROD4/NKD1/PRSS56	
	4	ment		0	3	3	8		

BP	GO:0	visual	10/1	366/	0.000	0.028	0.025	DLX1/DLX2/FOXC2/MFAP2/NEUROD1/TFAP2B/WN	10
	15006	system	35	1867	33336	45195	83004	T6/NEUROD4/NKD1/PRSS56	
	3	develop		0	5	1	2		
		ment							
BP	GO:0	cartilage	4/13	47/1	0.000	0.028	0.026	COMP/COCH/AXIN2/CER1	4
	06035	develop	5	8670	36645	90314	23966		
	1	ment			8	8			
		involved							
		in							
		endocho							
		ndral							
		bone							
		morpho							
		genesis							
BP	GO:0	sensory	10/1	371/	0.000	0.028	0.026	DLX1/DLX2/FOXC2/MFAP2/NEUROD1/TFAP2B/WN	10
	04888	system	35	1867	37100	90314	23966	T6/NEUROD4/NKD1/PRSS56	
	0			0	3	8			



BP	GO:0	odontog	5/13	90/1	0.000	0.032	0.029	AMBN/DLX1/DLX2/DLX3/WNT6	5
	04247	genesis of	5	8670	49302	26031	28745		
	5	dentin-		6		2			
		containi							
		ng tooth							
BP	GO:0	kidney	6/13	140/	0.000	0.034	0.031	FGF1/FOXC2/SMAD7/TFAP2B/WNT6/CER1	6
	07207	epitheli	5	1867	54326	40060	23051		
	3	um		0		2	2		
		develop							
		ment							
BP	GO:0	forebrai	4/13	53/1	0.000	0.035	0.032	DLX1/DLX2/DLX5/SATB2	4
	02187	n	5	8670	58181	69066	40169		
	9	neuron		4	8	5			
		different							
		iation							

BP	GO:0	cerebral	3/13	24/1	0.000	0.039	0.036	DLX1/DLX2/ID4	3
	02189	cortex	5	8670	66956	67221	01633		
	5	neuron		6	9	7			
		different							
		iation							
BP	GO:0	ureteric	5/13	97/1	0.000	0.039	0.036	FGF1/FOXC2/SMAD7/WNT6/CER1	5
	00165	bud	5	8670	69442	67221	01633		
	7	develop		1	9	7			
		ment							
BP	GO:0	mesone	5/13	98/1	0.000	0.039	0.036	FGF1/FOXC2/SMAD7/WNT6/CER1	5
	07216	phric	5	8670	72756	67221	01633		
	3	epitheli		9	7				
		um							
		develop							
		ment							

BP	GO:0	mesone	5/13	98/1	0.000	0.039	0.036	FGF1/FOXC2/SMAD7/WNT6/CER1	5
	07216	phric	5	8670	72756	67221	01633		
	4	tubule			9		7		
		develop							
		ment							
BP	GO:0	artery	5/13	100/	0.000	0.042	0.038	COMP/FOXC2/SMAD7/TFAP2B/HEY1	5
	06084	develop	5	1867	79736	30361	40524		
	0	ment		0	8	9	9		
BP	GO:0	monoval	3/13	26/1	0.000	0.042	0.038	CKB/ABCC2/TFAP2B	3
	05508	ent	5	8670	85107	79626	85249		
	3	inorgani		8			1		
		c anion							
		homeost							
		asis							
BP	GO:0	mesone	5/13	102/	0.000	0.042	0.038	FGF1/FOXC2/SMAD7/WNT6/CER1	5
	00182	phros	5	1867	87205	79626	85249		
	3			0	8		1		

		develop									
		ment									
BP	GO:0	cerebell	5/13	102/	0.000	0.042	0.038	CKB/LMX1A/NEUROD1/CDK5R1/SEZ6			5
	02154	um	5	1867	87205	79626	85249				
	9	develop		0	8		1				
		ment									
BP	GO:0	negative	9/13	344/	0.000	0.043	0.039	DLX1/DLX2/ID4/LMX1A/TTPA/WNT3/CDK5R1/HEY			9
	01072	regulati	5	1867	90165	16932	19117	1/GAL			
	1	on of cell		0	2	7	9				
		develop									
		ment									
BP	GO:0	drug	7/13	214/	0.000	0.044	0.040	ABCC2/SYT7/SLC38A3/FGF20/GAL/SLC46A1/SYT6			7
	01589	transpor	5	1867	94540	18630	11444				
	3	t		0	2	8	3				
BP	GO:0	BMP	6/13	157/	0.000	0.044	0.040	COMP/DLX1/DLX5/SMAD7/TFAP2B/CER1			6
	03050	signalin	5	1867	98967	50954	40789				
	9			0	9	2	1				





BP	GO:0	cornifica	5/13	112/	0.001	0.052	0.047	IVL/KRT31/KRT35/KRT37/KRT23	5
	07026	tion	5	1867	32619	29447	47542		
			8		0	9	2	2	
BP	GO:0	forebrai	4/13	66/1	0.001	0.052	0.047	DLX1/DLX2/DLX5/SATB2	4
	02187	n	5	8670	33200	29447	47542		
		generati			4	2	2		
		on	of						
		neurons							
BP	GO:0	fibroblas	5/13	114/	0.001	0.055	0.050	FGF1/FGF18/FGF19/FGF20/TRIM71	5
	00854	t growth	5	1867	43469	15869	07569		
		factor		0	1	2	8		
		receptor							
		signalin							
		g							
		pathway							

BP	GO:0	response	6/13	170/	0.001	0.055	0.050	COMP/DLX1/DLX5/SMAD7/TFAP2B/CER1	6
	07177	to BMP	5	1867	48925	15869	07569		
			2		0	7	2	8	
BP	GO:0	cellular	6/13	170/	0.001	0.055	0.050	COMP/DLX1/DLX5/SMAD7/TFAP2B/CER1	6
	07177	response	5	1867	48925	15869	07569		
		to BMP			0	7	2	8	
			stimulus						
BP	GO:0	positive	9/13	371/	0.001	0.055	0.050	DLX1/DLX2/DNMT3B/NEUROD1/SERPINI1/WNT3/F	9
	04566	regulati	5	1867	52382	39379	28913	GF20/CACNG7/SLITRK1	
		on of			0	3	2	3	
			neuron						
			different						
			iation						
BP	GO:0	develop	7/13	235/	0.001	0.057	0.052	FGF1/SH3GL2/WNT3/CDK5R1/GAL/CACNG7/NKD1	7
	06056	mental	5	1867	62154	87432	54107		
		growth			0	2	2	7	
			involved						

in

## morphogenesis

BP	GO:000175	somitogenesis	4/13	70/1	0.001	0.058	0.052	FOXC2/AXIN2/DMRT2/NKD1	4
			6		7	2	3		
BP	GO:0005189	positive regulation of protein kinase B signaling	6/13	176/1867	0.001	0.061	0.055	F10/FGF1/FGF18/FGF19/FGF20/TNFAIP8L3	6
			7	5	77599	16266	52639		
BP	GO:0000320	cardiac ventricle morphogenesis	4/13	73/1867	0.001	0.065	0.059	FOXC2/SMAD7/MYL3/HEY1	4
			8	5	93485	48474	45018		
				2	7	5			

BP	GO:0	epiderm	10/1	464/	0.002	0.066	0.060	IVL/KRT31/KRT35/KRT37/KRT23/GAL/TNFRSF19/K	10
	00854	is	35	1867	02449	87335	71083	RTAP4-6/APCDD1/PRR9	
	4	develop		0	9	6			
		ment							
BP	GO:0	camera-	8/13	314/	0.002	0.066	0.060	DLX1/DLX2/FOXC2/NEUROD1/TFAP2B/WNT6/NEU	8
	04301	type eye	5	1867	04922	87335	71083	ROD4/PRSS56	
	0	develop		0	8	6			
		ment							
BP	GO:0	growth	3/13	36/1	0.002	0.066	0.060	COMP/COCH/CER1	3
	00341	plate	5	8670	21719	87335	71083		
	7	cartilage		3	6				
		develop							
		ment							
BP	GO:0	embryo	5/13	126/	0.002	0.066	0.060	DLX1/DLX2/FOXC2/DMRT2/SATB2	5
	04870	nic	5	1867	22813	87335	71083		
	6	skeletal		0	4	6			
		system							



BP	GO:0	endothel	2/13	10/1	0.002	0.066	0.060	FGF1/FGF18	2
	03576	ial	cell	5	8670	24841	87335	71083	
	8	chemota			6	6			
		xis	to						
		fibroblas							
		t growth							
		factor							
BP	GO:0	regulati	2/13	10/1	0.002	0.066	0.060	DLX1/DLX2	2
	04653	on	of	5	8670	24841	87335	71083	
	2	photorec			6	6			
		eptor							
		cell							
		different							
		iation							
BP	GO:2	regulati	2/13	10/1	0.002	0.066	0.060	FGF1/FGF18	2
	00054	on	of	5	8670	24841	87335	71083	
	4	endothel			6	6			





BP	GO:0	regulati	3/13	39/1	0.002	0.076	0.069	DLX1/DLX2/ID4	3
	04871	on	of	5	8670	79351	82135	7421	
	3	oligoden			9	6			
		drocyte							
		different							
		iation							
BP	GO:0	Notch	6/13	193/	0.002	0.076	0.069	DLX1/DLX2/FOXC2/PTP4A3/HEY1/NEUROD4	6
	00721	signalin	5	1867	81769	82135	7421		
	9	g		0	6	6			
		pathway							
BP	GO:0	dopami	3/13	40/1	0.003	0.079	0.072	SYT7/FGF20/SYT6	3
	01404	ne	5	8670	00418	69220	34839		
	6	secretio			9	2	1		
		n							
BP	GO:0	regulati	3/13	40/1	0.003	0.079	0.072	SYT7/FGF20/SYT6	3
	01405	on	of	5	8670	00418	69220	34839	
	9	dopami			9	2	1		



BP	GO:0	cellular	2/13	12/1	0.003	0.083	0.075	CKB/ABCC2	2
	03000	anion	5	8670	26657	27643	60232		
	2	homeost			4	3	8		
			asis						
BP	GO:0	cellular	2/13	12/1	0.003	0.083	0.075	CKB/ABCC2	2
	03032	monoval	5	8670	26657	27643	60232		
	0	ent			4	3	8		
			inorgani						
			c anion						
			homeost						
			asis						
BP	GO:0	negative	3/13	43/1	0.003	0.092	0.084	DLX1/DLX2/HEY1	3
	04574	regulati	5	8670	69383	96149	39489		
	6	on	of		4	5			
			Notch						
			signalin						

BP	GO:004434	cellular response to fibroblast growth factor stimulus	5/13	143/5	0.003	0.093	0.085	FGF1/FGF18/FGF19/FGF20/TRIM71
				1867	84364	94128	28438	
			4	0	3	9		
BP	GO:003411	endochondral bone growth	3/13	44/1	0.003	0.093	0.085	COMP/COCH/CER1
			5	8670	94336	94128	28438	
			6	2	3	9		
BP	GO:006004	retina development	5/13	144/5	0.003	0.093	0.085	DLX1/DLX2/NEUROD1/TFAP2B/NEUROD4
			1	1867	95922	94128	28438	
			1	0	8	3	9	



BP	GO:0	regulati	4/13	91/1	0.004	0.099	0.090	DLX1/SMAD7/TFAP2B/CER1	4
	03051	on	of	5	8670	30007	30645	15514	
	0	BMP			6		4		
		signalin							
		g							
		pathway							
CC	GO:0	integral	6/13	152/	0.000	0.093	0.086	CHRND/SYT7/CACNG7/SLITRK1/SYT6/SHISA7	6
	09969	compon	8	1971	70838	80697	40116		
	9	ent	of	7	2	6	2		
		synaptic							
		membra							
		ne							
CC	GO:0	intrinsic	6/13	164/	0.001	0.093	0.086	CHRND/SYT7/CACNG7/SLITRK1/SYT6/SHISA7	6
	09924	compon	8	1971	05167	80697	40116		
	0	ent	of	7	3	6	2		
		synaptic							



CC	GO:0	intrinsic	4/13	77/1	0.002	0.093	0.086	CHRND/CACNG7/SLITRK1/SHISA7	4
	09894	compon	8	9717	09390	80697	40116		
	8	ent	of		6	6	2		
		postsyn							
		aptic							
		specializ							
		ation							
		membra							
		ne							
CC	GO:0	neurona	10/1	497/	0.002	0.098	0.090	CKB/KCNN1/PCSK2/SERPINI1/SH3GL2/CDK5R1/GA	10
	04302	l	cell	38	1971	64135	61047	82544	L/CACNG7/SRCIN1/SEZ6
	5	body		7	2	8			
MF	GO:0	DNA-	13/1	439/	1.796	0.004	0.004	DLX2/DLX3/DLX5/FOXC2/LMX1A/NEUROD1/TFAP2	13
	00122	binding	28	1769	19E-	53608	21853	B/LHX3/SATB2/VENTX/SP7/KLF14/MAFA	
	8	transcri		7	05	6	7		
		ption							
		activator							

activity,

RNA

## **polymer**

ase II-

## specific

MF	GO:000510	fibroblast growth factor receptor binding	4/12	25/1	2.935	0.004	0.004	FGF1/FGF18/FGF19/FGF20	4
			00510	t	7697	98E-05	53608	21853	
			4	growth factor		6	7		
				receptor					
				binding					
KEGG	hsa05224	Breast cancer	9/54	147/8076	4.543	3.771	3.061	FGF1/WNT3/WNT6/AXIN2/FZD7/FGF18/FGF19/HEY1/FGF20	9
					81E-07	36E-05	09E-05		
KEGG	hsa05226	Gastric cancer	8/54	149/8076	5.643	0.000	0.000	FGF1/WNT3/WNT6/AXIN2/FZD7/FGF18/FGF19/FGF20	8
					26E-06	23419	19008	20	
					5	9			



