Supplemental Table 1. Demographics and case information for the 61 Vanderbilt Medical Center patients (test dataset) represented by 6 adenomas and 55 adenocarcinoma patients. The Vanderbilt test set included 15 patients from the University of Alabama-Birmingham Medical Center (one adenoma and 14 tumors provided by Martin J. Heslin, M.D.). All patients were diagnosed with adenoma or colorectal adenocarcinoma (Stages I-IV) according to current American Joint Commission on Cancer (AJCC) guidelines. *Other* in the VMC medical record implies not otherwise specified or Asian.

Supplementary Table 1  VMC Demographics	VMC-adenomas	VMC-adenocarcinomas	
Sample size	6	55	
Mean Age (s.d.)	61.7 (15.2)	62.3 (14.1)	
Sex (%male)	2 (33.3%)	30 (54.5%)	
Stage I	n/a	4 (7.3%)	
Stage II	n/a	15 (27.3%)	
Stage III	n/a	19 (34.5%)	
Stage IV	n/a	17 (30.9%)	
Median Follow-up Months (Min - Max)	n/a	50.2 (0.4 - 111.3)	
Number of deaths	n/a	20 (36.3%)	
Caucasian (%)	5 (83.3%)	50 (90.9%)	
Black (%)	0 (0%)	4 (7.3%)	
Other (%)	1 (16.7%)	1 (1.8%)	

## Supplemental Table 2. Intestinal polyp development in Min mice with los of Rab25

		No. of polyps/mouse				
		Small Intestine				Colon
Rab25	No. of Mice	Proximal	Middle	Distal	Total	
+/+	7	$7.4 \pm 0.9$	12.6±5.5	25.4±3.7	45.4±5.1	1.9±0.5
+/-	16	$13.9 \pm 2.4$	38.9±5.1**	45.8±4.2*	98.5±9.2**	$3.5 \pm 0.6$
-/-	12	20.7±3.1**	60.1±1.6**##	94.3±7.8**##	175.0±9.4**##	$4.7{\pm}0.8^{**}$

## Supplement Table 3: Primers for Mouse Genotyping

Rab25	NEO-NM32	TCTATCGCCTTCTTGACGAGTTC	
	R25KO-S	TCCCCTCAGTGTCTCTGCCAG	
	R25KO-AS	GGGAACACACGGCAGAGTCTC	
Apc <sup>Min</sup>	oIMR0033	GCCATCCCTTCACGTTAG	
	oIMR0034	TTCCACTTTGGCATAAGGC	
	oIMR0758	TTCTGAGAAAGACAGAAGTTA	
Smad3	489	TGGACTTAGGAGACGGCAGTCC	
	490	CTTCTGAGACCCTCCTGAGTAGG	
	542	CTCTAGAGCGGCCTACGTTTGG	

Supplemental Figure 1. Expression of Rab25 protein in normal C57BL/6 mouse tissues by immunohistochemistry using anti-Rab25 antibodies. A. Skin B. Higher magnification views of A C. Dorsal surface of tongue D. Esophagus E. Kidney (collecting duct) F. Stomach (fundus) Bar=50 fm.

Supplemental Fig. 1

