

Supplementary Table 1. Pooled multivariate relative risk (RR) and 95% confidence interval (CI) for colorectal cancer according to cumulative alcohol intake, stratified by one-carbon nutrients and family history of colorectal cancer in post-fortification period (1998-2008)

Alcohol (g/d)	0	0.1-4.9	5.0-9.9	10-14.9	15-29.9	30+	<i>P</i> for trend
Vitamin B6							
High (>3.9 for women; >4.3 for men; median)							
No. of cases	133	120	56	41	55	30	
Multivariate RR*	1.00	1.08 (0.83-1.40)	1.19 (0.85-1.66)	0.97 (0.66-1.42)	1.43 (1.00-2.03)	1.33 (0.78-2.25)	0.29
Low (<=3.9 for women; <=4.3 for men; median)							
No. of cases	179	151	72	59	64	38	
Multivariate RR*	1.00	1.05 (0.84-1.32)	1.20 (0.90-1.59)	1.08 (0.78-1.51)	1.25 (0.90-1.72)	1.05 (0.71-1.55)	0.53
<i>P</i> for interaction						0.40	
Vitamin B12							
High (>12.4 for women; >13.7 for men; median)							
No. of cases	134	120	64	39	58	29	
Multivariate RR*	1.00	0.97 (0.75-1.25)	1.22 (0.69-2.12)	0.86 (0.58-1.27)	1.33 (0.79-2.24)	1.07 (0.68-1.67)	0.17
Low (<=12.4 for women; <=13.7 for men; median)							
No. of cases	178	151	64	61	61	39	
Multivariate RR*	1.00	1.09 (0.87-1.36)	1.10 (0.81-1.48)	1.18 (0.82-1.72)	1.17 (0.84-1.64)	1.19 (0.82-1.75)	0.41
<i>P</i> for interaction						0.69	
Betaine							
High (>91.5 for women; >116.6 for men; median)							
No. of cases	147	116	64	52	61	37	
Multivariate RR*	1.00	0.88 (0.67-1.15)	1.10 (0.80-1.50)	0.98 (0.68-1.41)	1.29 (0.91-1.83)	1.27 (0.83-1.94)	0.10
Low (<=91.5 for women; <=116.6 for men; median)							

No. of cases	165	155	64	48	59	31	
Multivariate RR*	1.00	1.18 (0.94-1.49)	1.16 (0.79-1.73)	0.95 (0.67-1.34)	1.16 (0.84-1.61)	1.02 (0.67-1.54)	0.91
<i>P</i> for interaction						0.52	
Choline							
High (>291.4 for women; >360.8 for men; median)							
No. of cases	143	142	67	52	62	26	
Multivariate RR*	1.00	1.15 (0.90-1.47)	1.28 (0.94-1.75)	1.17 (0.82-1.66)	1.43 (1.02-2.03)	1.06 (0.67-1.70)	0.41
Low (<=291.4 for women; <=360.8 for men; median)							
No. of cases	169	129	60	47	57	42	
Multivariate RR*	1.00	1.00 (0.79-1.27)	1.06 (0.69-1.62)	0.92 (0.65-1.31)	1.16 (0.83-1.61)	1.22 (0.84-1.78)	0.20
<i>P</i> for interaction						0.64	
Methionine							
High (>1.5 for women; >1.9 for men; median)							
No. of cases	153	132	60	48	63		
Multivariate RR*	1.00	0.90 (0.70-1.15)	0.96 (0.58-1.59)	0.95 (0.66-1.37)	0.96 (0.68-1.35)		0.96
Low (<=1.5 for women; <=1.9 for men; median)							
No. of cases	159	139	67	52	124		
Multivariate RR*	1.00	1.19 (0.94-1.51)	1.31 (0.97-1.77)	1.04 (0.74-1.46)	1.39 (1.06-1.83)		0.03
<i>P</i> for interaction							0.05
Family history of colorectal cancer							
Yes							
No. of cases	74	51	31	22	31		
Multivariate RR*	1.00	0.93 (0.54-1.61)	1.40 (0.88-2.24)	1.03 (0.45-2.34)	1.09 (0.27-4.49)		0.82
No							
No. of cases	238	220	97	77	156		

Multivariate RR*	1.00	1.11 (0.92-1.34)	1.08 (0.76-1.54)	1.03 (0.78-1.35)	1.28 (0.85-1.91)	0.27
<i>P</i> for interaction					0.86	

*Multivariate RRs are adjusted for age, gender (combined data only), total pack-years of smoking (<10, 10-<30, 30-<50, 50-<70, 70+), physical activity (quintiles of metabolic equivalent tasks/wk), body mass index (BMI) (<23, 23-<25, 25-<30, 30-<35, and 35+ kg/m²), height (continuous), family history of colorectal cancer in parents and siblings (yes, no), history of endoscopy (yes, no), aspirin use (never, past, current use of 1-2, ≥3 tablets/wk), postmenopausal hormone (PMH) use (only in women; premenopausal, never, past, and current), red meat intake (quintiles), calcium intake from foods (continuous), total folate intake (quintiles at baseline), and total energy intake (continuous).