

Supplementary Table 1 – Full equations for incident colorectal cancer risk models

Colon cancer
Colditz 2000 simple point score model
The risk score was calculated as X; Where $X = 10$ (if family history of colorectal cancer, else 0) + 10 (if BMI ≥ 27 /km/m ² , else 0) + 5 (if alcohol intake ≥ 8 servings per week, else 0) + 5 (if vegetable portions < 3 per week, else 0) + 5 (if height ≥ 5 ft 7 in, else 0) + 10 (if total leisure physical activity time < 3 hours per week, else 0) + 10 (if red meat consumption ≥ 3 per week, else 0) + 5 (if use of birth control pills < 5 years, else 0) + 5 (if use of postmenopausal hormones < 5 years, else 0) + 5 (if currently use aspirin on a regular basis, else 0) + 10 (if not regularly using multivitamins, else 0) + 10 (if has history of inflammatory bowel disease for ≥ 10 years, else 0) + 5 (if consume milk or dairy products (saturated fats) ≥ 3 servings per day, else 0) + 10 (if do not take calcium supplement every day, else 0) + 10 (if do not take vitamin D supplement every day, else 0)
Colorectal cancer
Driver 2007 simple point score model
The risk score was calculated as X; Where $X = \text{age score} + \text{BMI score} + \text{smoking status score} + \text{alcohol intake score}$ Where Age score = 2 if 50-59 years; 4 if 60-69 years; 6 if ≥ 70 BMI score = 1 if 25.0-29.9; 2 if ≥ 30 Smoking status score = 1 if smoker; 0 if else Alcohol intake score = 1 if \geq once/week; 0 if else
Freedman 2009 logistic model
The probability of developing colorectal cancer was calculated using the Gauss Program available at https://dceg.cancer.gov/tools/risk-assessment/ccratgauss
Guesmi 2010 logistic model
The risk score was calculated as X; Where $X = (0.082 * \text{age}) + 1.631$ (if consume processed meat frequently, else 0) - 1.952 (if consume milk frequently, else 0)
Johnson 2013 restricted cubic spine model
Assuming multiplicative joint effects between all risk factors included in the meta-analysis, selecting the “Overall” estimate where multiple estimates are provided, the risk score was calculated as the baseline risk among people with the lowest risk levels of all risk factors multiplied by X; where exp denotes the exponential function and where $X = \exp(0.032 \times (\text{BMI}-22)/8, \text{ if male, else } 0) \times \exp(0.017 \times (\text{BMI}-22)/8, \text{ if female, else } 0) \times \exp(-0.061 \times \text{standardised physical activity}) \times \exp(0.011 \times \text{pack-years} - 0.017 \times \text{rcs}) \times \exp(0.011 \times \text{drinks/week}) \times \exp(-0.120 \times \text{years of current HT use} + 0.415 \times \text{rcs, if current HRT user, else } 0) \times \exp(-0.008 \times \text{years of former HRT use, if former HRT user}) \times \exp(-0.069 \times \text{years of aspirin use} + 0.146 \times \text{rcs}) \times \exp(0.017 \times \text{servings of processed meat per week}) \times \exp(0.024 \times \text{servings of red meat per week}) \times \exp(-0.104 \times \text{servings/day} + 0.128 \times \text{rcs}) \times 1.80, \text{ if CRC family history, else } 1 \times 2.93 \text{ if IBD, else } 1$ $\text{Where: } rcs = (\text{dose} - \text{knot}_1)_+^3 - \frac{(\text{knot}_3 - \text{knot}_1)(\text{dose} - \text{knot}_2)_+^3 - (\text{knot}_2 - \text{knot}_1)(\text{dose} - \text{knot}_3)_+^3}{\text{knot}_3 - \text{knot}_2} \times \frac{1}{(\text{knot}_3 - \text{knot}_1)^2}$ For smoking knots = (0, 14.2, 57.7), for current HRT use knots = (0, 1.7, 14.4), for fruit consumption knots = (0, 1.08, 3.95), for years of aspirin use (0, 2.0, 13.2) and “dose” = the risk factor unit in the model (pack years for smoking, years of HRT use and serving of fruit per day)
Ma 2010 simple point score model
The risk score was calculated as X; Where $X = \text{age score} + \text{BMI score} + \text{smoking status score} + \text{alcohol intake score} + \text{physical activity score}$ Where Age score = 0 if 40-44 years; 1 if 45-49 years; 3 if 50-54 years; 4 if 50-59 years; 5 if 60-64 years; 6 if 65-69 years BMI score = 0 if <25kg/m ² ; 1 if ≥ 25 kg/m ²

Smoking status score = 0 if non-smoker or former smoker; 1 if current smoker Alcohol intake score = 0 if never or occasional; 1 if regular <300g/week; 2 if regular ≥300g/week Physical activity score = 0 if <24.7 MET-h/day; -1 if ≥24.7 MET-h/day
Ma 2010 cox proportional hazards model
The probability of developing colorectal cancer was calculated as $1 - S(t)^{\exp(f(x,M))}$; Where $S(t) = 0.9882$ $f(x,M) = ((0.080 * (\text{individual's age} - 52.9)) + ((0.04 * (\text{individual's BMI} - 23.4)) + ((-0.019 * (\text{individual's physical activity} - 28.7)) + (0 * (1 \text{ if never smoker, else } 0)) + (0.071 * (1 \text{ if ex-smoker, else } 0) - 0.239)) + (0.239 * (1 \text{ if current smoker, else } 0) - 0.525)) + (-0.163 * (1 \text{ if never drinks alcohol, else } 0) - 0.235)) + (0 * (1 \text{ if occasionally drinks alcohol, else } 0)) + (0.358 * 1 \text{ if regular alcohol consumption } <300\text{g/w, else } 0) - 0.481)) + (0.659 * (1 \text{ if regular alcohol consumption } \geq 300\text{g/w, else } 0) - 0.208))$
Tao 2014 simple point score model
The risk score was calculated as X; Where $X = (6 * \text{individual's age}) + (104 * (1 \text{ if male, else } 0)) + (35 * \text{number of first degree relatives with bowel cancer}) + (1 * \text{pack years of smoking}) + (1 * \text{alcohol use g/day}) + (-31 * (1 \text{ if ever regular use of NSAID, else } 0)) + (-147 * (1 \text{ if previous colonoscopy, else } 0)) + (187 * (1 \text{ if history of colonic polyp, else } 0)) + (47 * (1 \text{ if red meat consumption } > 1 \text{ time/day, else } 0))$
Wei 2009 logistic model
The risk score was calculated as X; Where $X = -0.24846 \text{ (if a former smoker)} + 0.00995 \text{ (if a current smoker)} + 0.832909 \text{ (if former alcohol intake)} + 2.15924 \text{ (if current alcohol intake)} + 0.482426 \text{ (if has family history of colorectal cancer)} + 0.329304 \text{ (if BMI } > 24)$
Wells 2015 cox proportional hazards model (male)
The probability of developing colorectal cancer was calculated as $1 - S(t)^{\exp(f(x,M))}$; Where: $S(t) = 0.9846654$ $X = (-6.6419738 ($ + 0.091669179 * (age in years) - 3.7611814e-05 * (the maximum of age - 47 or 0) ^ 3) + 7.794128e-05 * (the maximum of age - 60 or 0) ^ 3) - 4.0529466e-05 * (the maximum of age - 72 or 0) ^ 3) - 0.16728044 * (1 if ethnicity is white, else 0) + 0.00022581331 * (pack years of smoking) - 1.1341047e-05 * (the maximum of pack years of smoking or 0) ^ 3) - 1.3522018e-05 * (the maximum of pack years of smoking - 6.375 or 0) ^ 3) + 2.1809706e-06 * (the maximum of pack years of smoking - 39.525 or 0) ^ 3) + 0.28379769 * (alcohol intake in drinks per day) - 0.21424251 * (the maximum of alcohol intake or 0) ^ 3) + 0.22570057 * (the maximum of alcohol intake - 0.14457189 or 0) ^ 3) - 0.011458065 * (the maximum of alcohol intake - 2.8477722 or 0) ^ 3) + 0.018020786 * (BMI in kg/m ²) + 9.4715899e-05 * (the maximum of BMI - 22.047175 or 0) ^ 3) - 0.00015791645 * (the maximum of alcohol intake - 25.941735 or 0) ^ 3) + 6.3200548e-05 * (the maximum of alcohol intake - 31.778341 or 0) ^ 3) + 0.072052428 * (number of years of education) - 0.00060634342 * (the maximum of the number of years of education - 7 or 0) ^ 3) + 0.0016674444 * the maximum of the number of years of education - 14 or 0) ^ 3) - 0.001061101 * (the maximum of the number of years of education - 18 or 0) ^ 3) - 0.20960315 * (current aspirin use=1, else 0) + 0.24250922 * (family history of colorectal cancer=1, else 0) - 0.19175375 * (multivitamin use=1, else 0) + 0.073141733 * (red meat intake per day) - 0.0043503755 * (the maximum of the amount of red meat intake - 0.59081962 or 0) ^ 3) + 0.0065250851 * (the maximum of the amount of red meat intake - 2.0822052 or 0) ^ 3))

- 0.0021747085	* (the maximum of the amount of red meat intake - 5.0656345 or 0) ^ 3)
+ 0.11020556	* (diabetes=1, else 0)
- 0.090669913	* (moderate or strenuous physical activity in hours per day)
+ 0.0093816671	* (the maximum of the amount of PA activity - 0.10714286 or 0) ^ 3)
- 0.011850527	* (the maximum of the amount of PA activity - 0.82142857 or 0) ^ 3)
+ 0.0024688598	* (the maximum of the amount of PA activity - 3.5357143 or 0) ^ 3)))

Wells 2015 cox proportional hazards model (female)

The probability of developing colorectal cancer was calculated as $1 - S(t)^{\exp(t[X,M])}$; Where:

$S(t)=0.9901043$;

$X = (- 5.9026635 ($

+ (0.090012542	* age in years)
- 4.4217156e-05	* (the maximum of age – 47 or 0) ^ 3)
+ 9.2119076e-05	* (the maximum of age – 60 or 0) ^ 3)
- 4.7901919e-05	* (the maximum of age – 72 or 0) ^ 3)
- 0.34056094	* (1 if ethnicity is white, else 0)
+ 0.07443905	* (number of years of education)
- 0.00062546554	* (the maximum of the number of years of education – 7 or 0) ^ 3)
+ 0.0017200302	* (the maximum of the number of years of education – 14 or 0) ^ 3)
- 0.0010945647	* (the maximum of the number of years of education – 18 or 0) ^ 3)
- 0.2450076	* (current oestrogen use=1, else 0)
- 0.044320489	* (past oestrogen use=1, else 0)
+ 0.23328937	* (diabetes=1, else 0)
+ 0.062703176	* (pack years of smoking)
- 0.002446026	* (the maximum of pack years of smoking or 0)^3)
+ 0.003038396	* (the maximum of pack years of smoking - 1.25 or 0) ^ 3)
- 0.00059134632	* (the maximum of pack years of smoking - 6.375 or 0) ^ 3)
- 1.023614e-06	* (the maximum of pack years of smoking - 27.5125 or 0) ^ 3)
+ 0.31589053	* (family history of colorectal cancer=1, else 0)
- 0.1665365	* (multivitamin use=1, else 0)
+ 0.0075233925	* (BMI in kg/m ²)
+ 6.7918662e-05	* (the maximum of BMI - 20.371336 or 0) ^ 3)
- 0.00011039091	* (the maximum of BMI - 25.508027 or 0) ^ 3)
+ 4.2472244e-05	* (the maximum of BMI - 33.722266 or 0) ^ 3)
- 0.236997	* (current NSAID use=1, else 0)
- 0.08856241	* (alcohol intake in drinks per day)
+ 0.62375456	* (the maximum of alcohol intake or 0) ^ 3)
- 0.7191129	* (the maximum of alcohol intake - 0.10740682 or 0) ^ 3)
+ 0.095358349	* (the maximum of alcohol intake - 0.8099724 or 0) ^ 3)))

Hippisley-Cox 2015 cox proportional hazards model (QCancer10)* (male)

The probability of developing colorectal cancer within 10 years among men was calculated as $1 -$

$S_0(10)^{\exp(X)}$ where exp denotes the exponential function and where:

$S_0(10)= 0.997250497341156$

and

$X = X_{\text{categorical}} + X_{\text{continuous}} + X_{\text{Boolean}} + X_{\text{Interaction}}$

$X_{\text{categorical}} = X_{\text{alcohol}} + X_{\text{ethnicity}} + X_{\text{smoking}}$

$X_{\text{alcohol}} = 0$ if alcohol intake = none, else

$X_{\text{alcohol}} = 0.047922717327152427$ if alcohol intake = <1 unit / day, else

$X_{\text{alcohol}} = 0.1306450076527492$ if alcohol intake = 1-2 units/day, else

$X_{\text{alcohol}} = 0.26424169760665095$ if alcohol intake = 3-6 units / day, else

$X_{\text{alcohol}} = 0.48248078623669538$ if alcohol intake = 7-9 units / day, else

$X_{\text{alcohol}} = 0.44338870524869628$ if alcohol intake = >9 units / day

$X_{\text{ethnicity}} = 0$ if ethnicity = White or not stated

$X_{\text{ethnicity}} = -0.58523471759902834$ if ethnicity = Indian

$X_{\text{ethnicity}} = -0.58643377805068653$ if ethnicity = Pakistani

$X_{ethnicity} = -0.88110809361453912$ if ethnicity = Bangladeshi
 $X_{ethnicity} = -0.4825211311076027$ if ethnicity = Other Asian
 $X_{ethnicity} = -0.35029857021954686$ if ethnicity = Black caribbean
 $X_{ethnicity} = -0.28686430423891179$ if ethnicity = Black African
 $X_{ethnicity} = -0.21258459900822296$ if ethnicity = Chinese
 $X_{ethnicity} = -0.5250625721689498$ if ethnicity = Other
 $X_{smoking} = 0$ if smoking status= never smoker
 $X_{smoking} = 0.054436520378164946$ if smoking status=ex-smoker
 $X_{smoking} = 0.066967983943209852$ if smoking status=light smoker (1-9 per day)
 $X_{smoking} = 0.026825493057743323$ if smoking status=moderate smoker (10-19 per day)
 $X_{smoking} = 0.12301347198058288$ if smoking status=heavy smoker (20+ per day)
 $X_{continuous} = age_1 \times 2.6296558591088894 + age_2 \times -0.14957281392190663 + BMI \times 0.015645211942720651 + Town \times 0.0089992920978670451$
 $X_{Boolean} = H/O \text{ blood cancer (1 if present, else 0)} \times 0.42840787527086416 + H/O \text{ colitis (1 if present, else 0)} \times 0.60167262208697581 + H/O \text{ Lung Cancer (1 if present, else 0)} \times 0.62731856184500678 + H/O \text{ Oral cancer (1 if present, else 0)} \times 0.48251770458128385 + H / O \text{ Colorectal polyps (1 if present, else 0)} \times 0.4092716863938517 + H / O \text{ Type 2 diabetes (1 if present, else 0)} \times 0.23892082443419493 + \text{Family History of bowel cancer (1 if present, else 0)} \times 0.78029895261674453$
 $X_{interaction} = age_1 \times \text{Family History of bowel cancer (1 if present, else 0)} \times -1.6277021020802243 + age_2 \times \text{Family History of bowel cancer (1 if present, else 0)} \times 0.12483775307131491$
and
 $age_1 = age/10 - 4.425716876983643$
 $age_2 = (age/10)^2 - 19.586969375610352$
 $BMI = BMI - 26.309040069580078$
 $Town = \text{Townsend Score} - 0.260672301054001$

Hippisley-Cox 2015 cox proportional hazards model (QCancer10)* (female)

The probability of developing colorectal cancer within 10 years among women was calculated as $1 - S_0(10)^{\exp(X)}$ where exp denotes the exponential function and where:
 $S_0(10) = 0.997171759605408$

and

$X = X_{categorical} + X_{continuous} + X_{Boolean} + X_{Interaction}$

$X_{categorical} = X_{alcohol} + X_{ethnicity} + X_{smoking}$

$X_{alcohol} = 0$ if alcohol intake = none, else

$X_{alcohol} = 0.016932317022484617$ if alcohol intake = <1 unit / day, else

$X_{alcohol} = 0.050471611879332329$ if alcohol intake = 1-2 units/day, else

$X_{alcohol} = 0.081302218350274985$ if alcohol intake = 3-6 units / day, else

$X_{alcohol} = 0.31884276410032175$ if alcohol intake = 7-9 units / day, else

$X_{alcohol} = 0.30711384933305064$ if alcohol intake = >9 units / day

$X_{ethnicity} = 0$ if ethnicity = White or not stated

$X_{ethnicity} = -1.0494971746256023$ if ethnicity = Indian

$X_{ethnicity} = -0.75800858721677611$ if ethnicity = Pakistani

$X_{ethnicity} = -0.16206418242822976$ if ethnicity = Bangladeshi

$X_{ethnicity} = -0.52075515892893942$ if ethnicity = Other Asian

$X_{ethnicity} = -0.3390081073116003$ if ethnicity = Black caribbean

$X_{ethnicity} = -0.36639919625711181$ if ethnicity = Black African

$X_{ethnicity} = -0.49625527321514168$ if ethnicity = Chinese

$X_{ethnicity} = -0.22289770148509797$ if ethnicity = Other

$X_{smoking} = 0$ if smoking status= never smoker

$X_{smoking} = 0.067943065824236953$ if smoking status=ex-smoker

$X_{smoking} = 0.10017056701706135$ if smoking status=light smoker (1-9 per day)

$X_{smoking} = 0.19403251493802176$ if smoking status=moderate smoker (10-19 per day)

$X_{smoking} = 0.16095242485295916$ if smoking status=heavy smoker (20+ per day)

$X_{continuous} = age_1 \times 33.92788234800954 + age_2 \times -83.924050851993556$

$X_{Boolean} = H/O \text{ breast cancer (1 if present, else 0)} \times 0.15096136873617999 + H/O \text{ cervical cancer (1 if present, else 0)} \times 0.55134659369582772 + H/O \text{ colitis (1 if present, else 0)} \times 0.56087545120009641 +$

H/O ovarian cancer (1 if present, else 0) x 0.68400222067217209 + H / O Colorectal polyps (1 if present, else 0) x 0.74904091187947031 + H / O Type 2 diabetes (1 if present, else 0) x 0.14828780874778763 + H/O endometrial cancer (1 if present, else 0) x 0.47923953865647911 + Family History of bowel cancer (1 if present, else 0) x 0.66160452485446319
 $X_{\text{interaction}} = \text{age_1} \times \text{Family History of bowel cancer (1 if present, else 0)} \times 10.985496891972508 + \text{age_2} \times \text{Family History of bowel cancer (1 if present, else 0)} \times -0.094897933584199498$
and
 $\text{age_1} = (\text{age}/10)^{-2} - 0.049712907522917$
 $\text{age_2} = (\text{age}/10)^{-2} \times \log(\text{age}/10) - 0.074606411159039$

* The beta coefficients not included in the original paper were downloaded in September 2016 from the website of the model using a link obtained from the investigators: <http://svn.clinrisk.co.uk/opensource/qcancer/10yr/>