



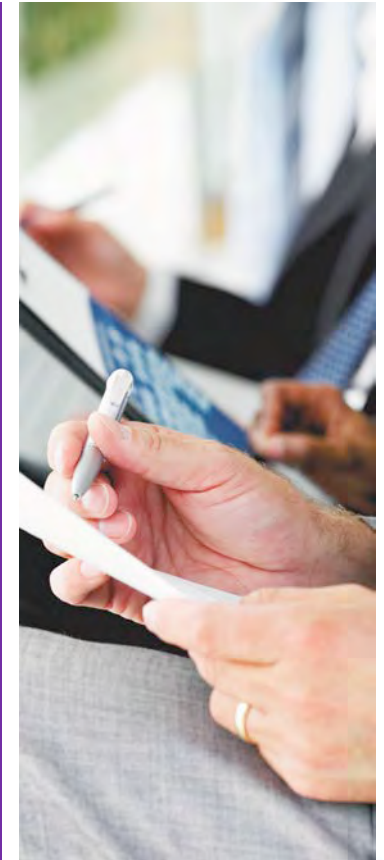
World
Cancer
Research
Fund International

Diet, Nutrition, Physical Activity and Cancer: update from WCRF

6 February 2017

Martin Wiseman

World Cancer Research Fund International & University
of Southampton



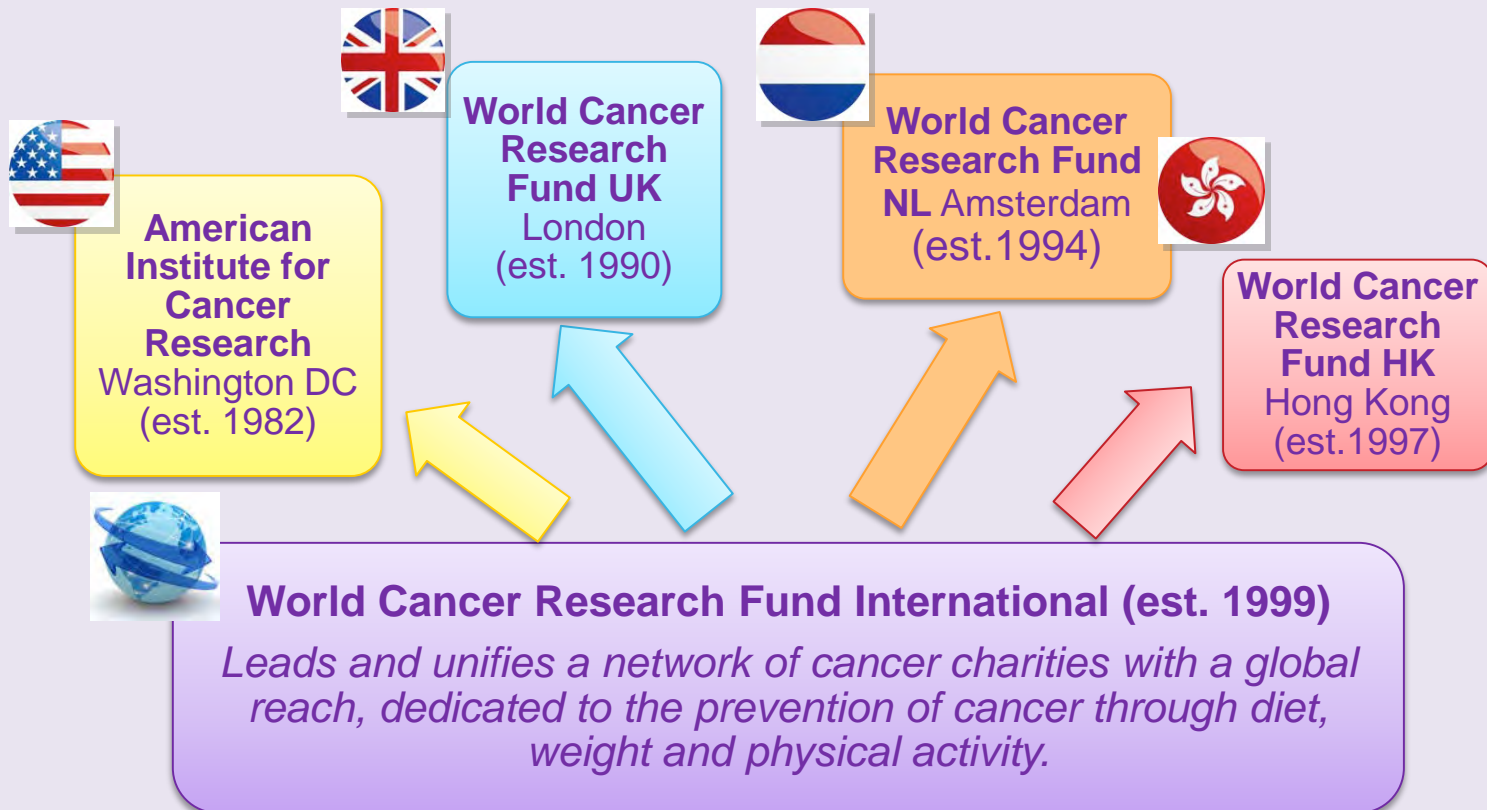
Outline

- Intro to WCRF
- Background
- Continuous Update Project
- Impact of adherence to recommendations
- Changes in emphasis
- Conclusion

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The World Cancer Research Fund Network



Who we are



AICR



WCRF UK



WCRF Netherlands



WCRF Hong Kong

WCRF International



Analysing research on cancer
prevention and survival



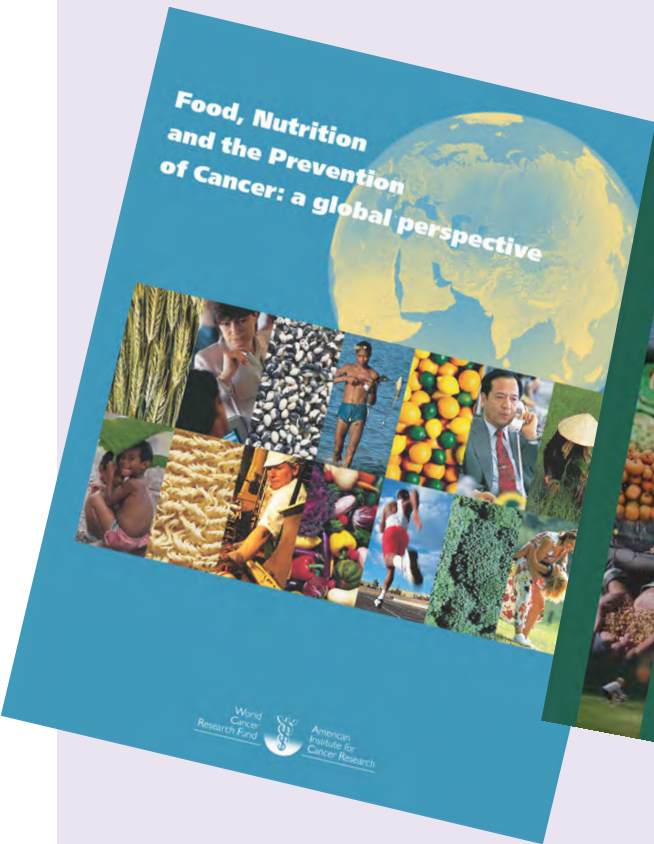
What we do

Fund research on the relationship of diet, nutrition, physical activity and body weight to cancer risk

Interpret the accumulated scientific literature to derive Cancer Prevention Recommendations

Educate people through our national Health Information programmes

Advocate effective policies to help people and populations to reduce their chances of developing cancer



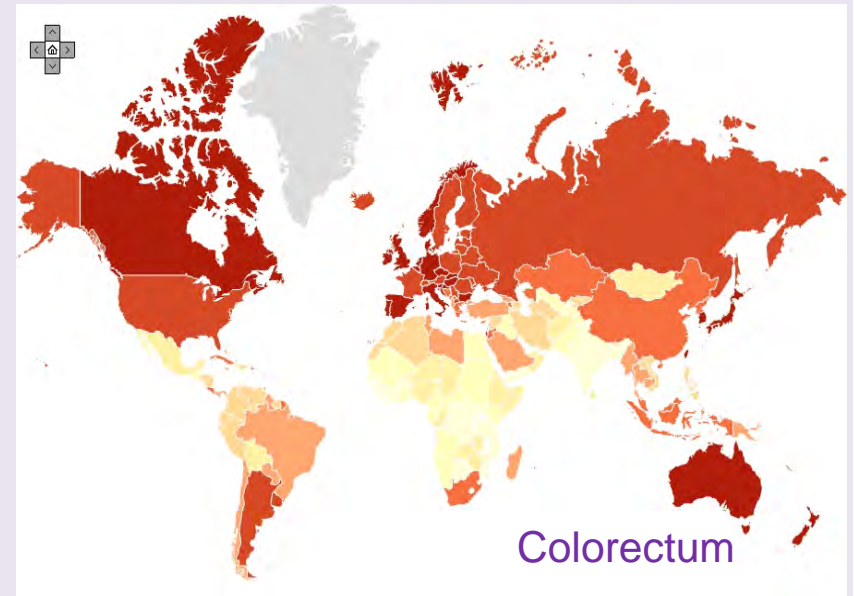
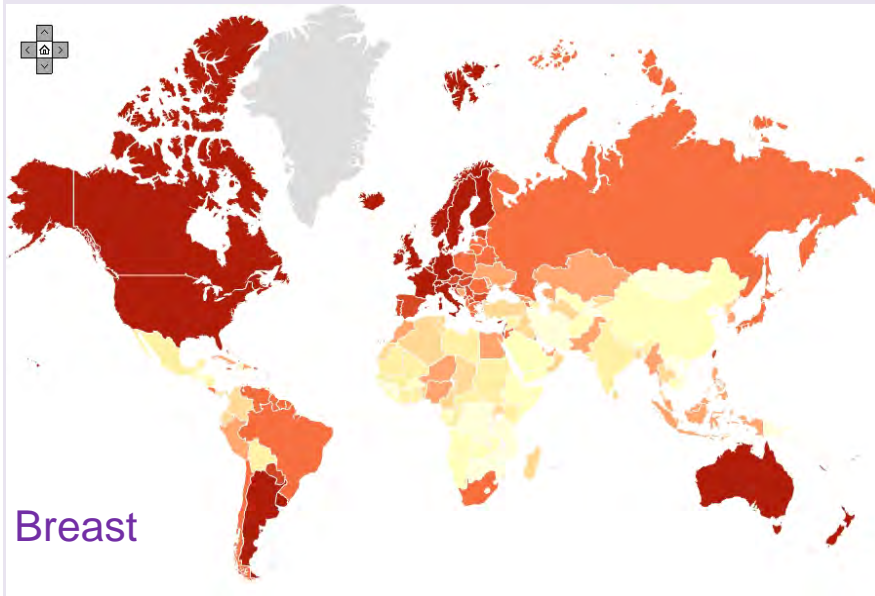
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Global variation in cancer incidence

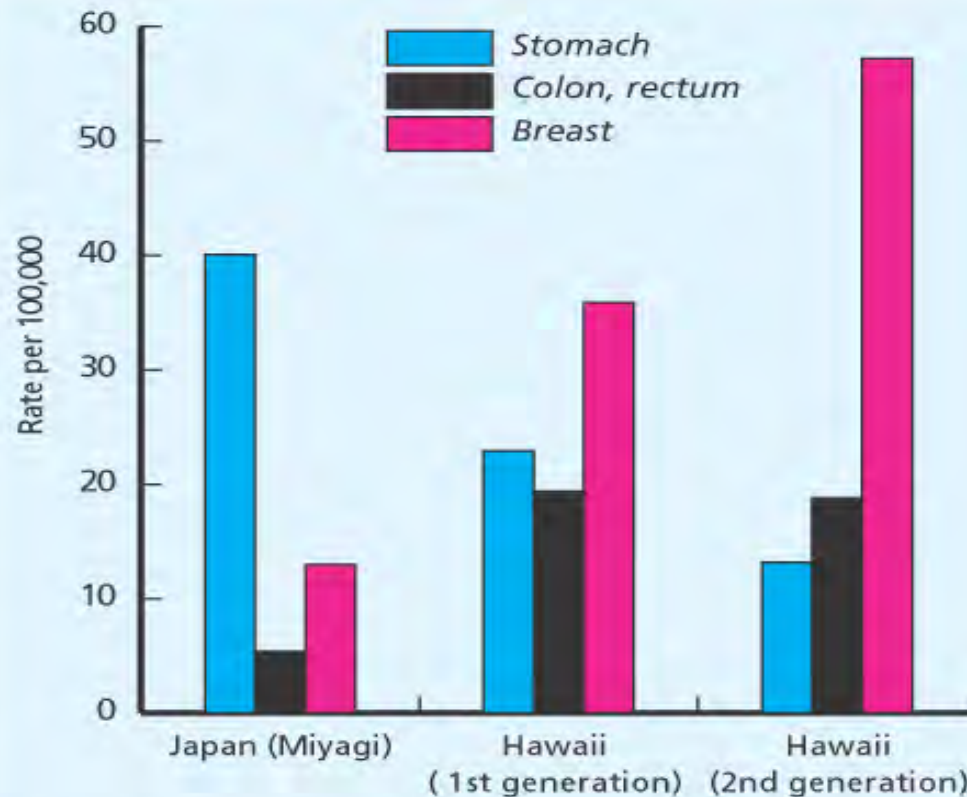


Globocan, WHO



Migration data

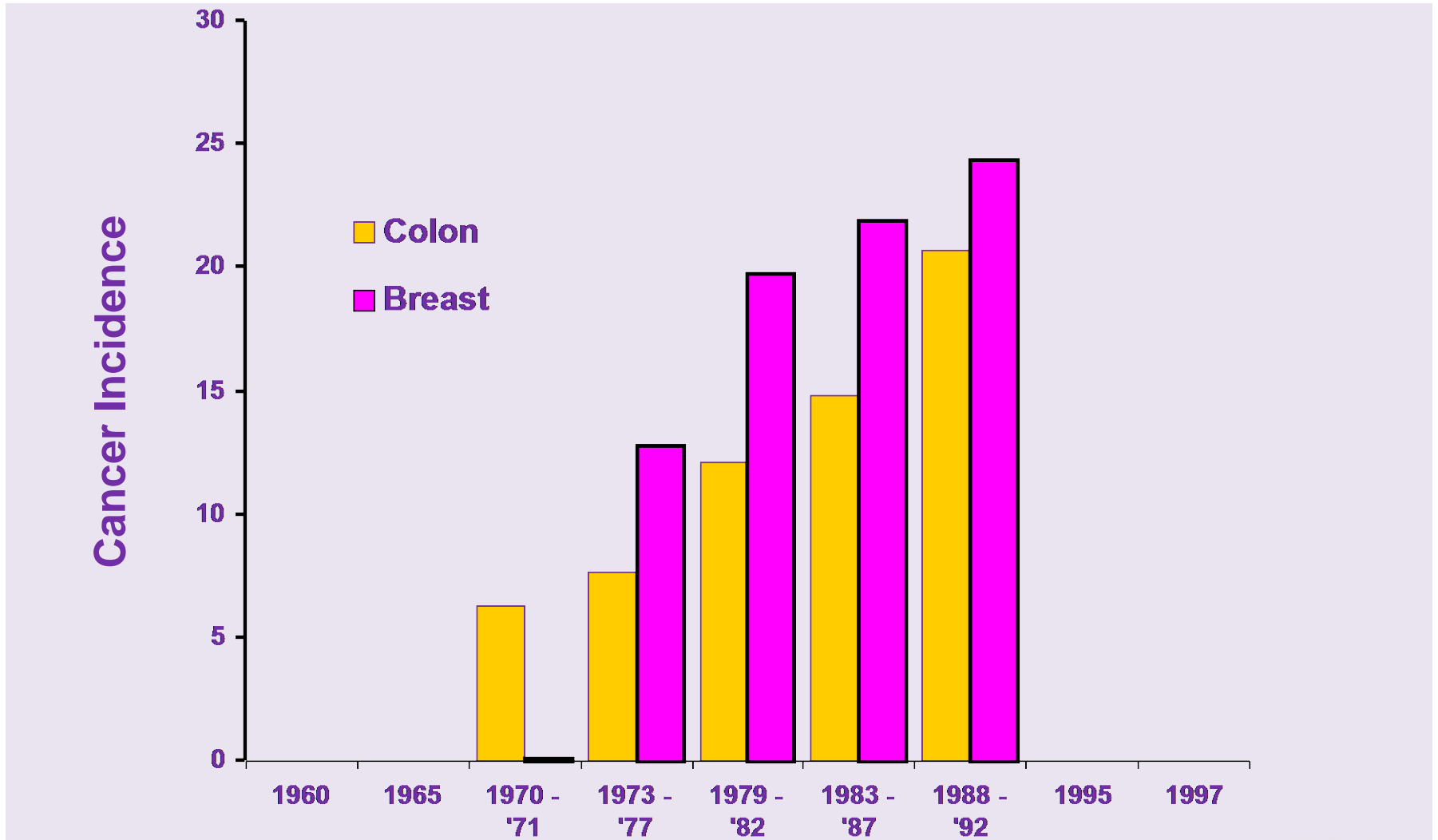
Figure 1.2.20 Cancer incidence for selected cancers in Japanese women by generation in Hawaii and Japan, 1968–1977



Age-adjusted to the World Standard Population
(From Kolonel et al, 1980)



Cancer Incidence in Japan*

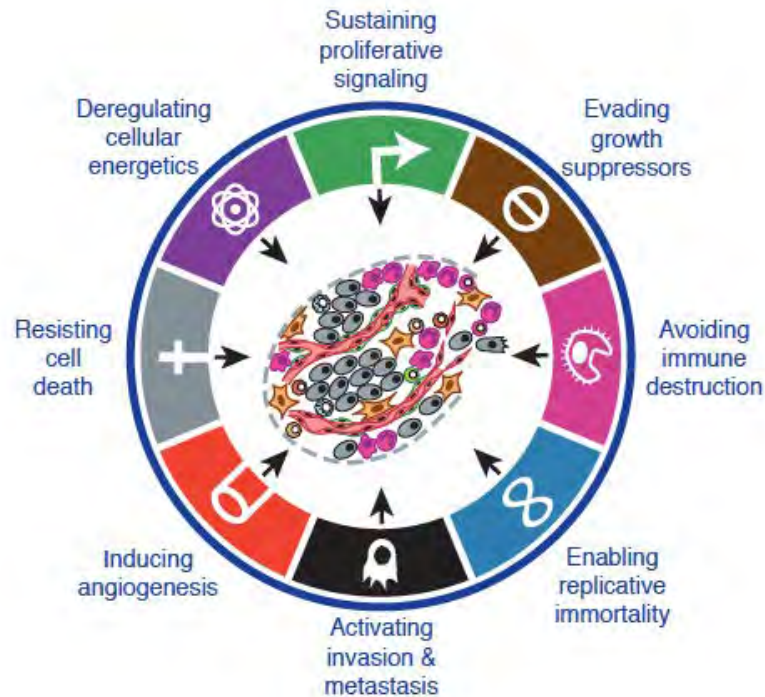


* Per 100,000, world population standard

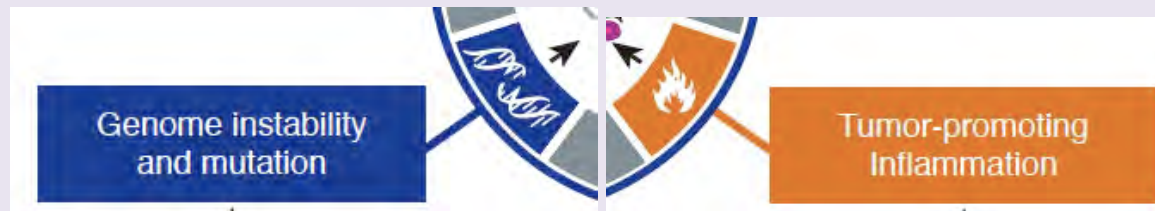
Hallmarks of cancer



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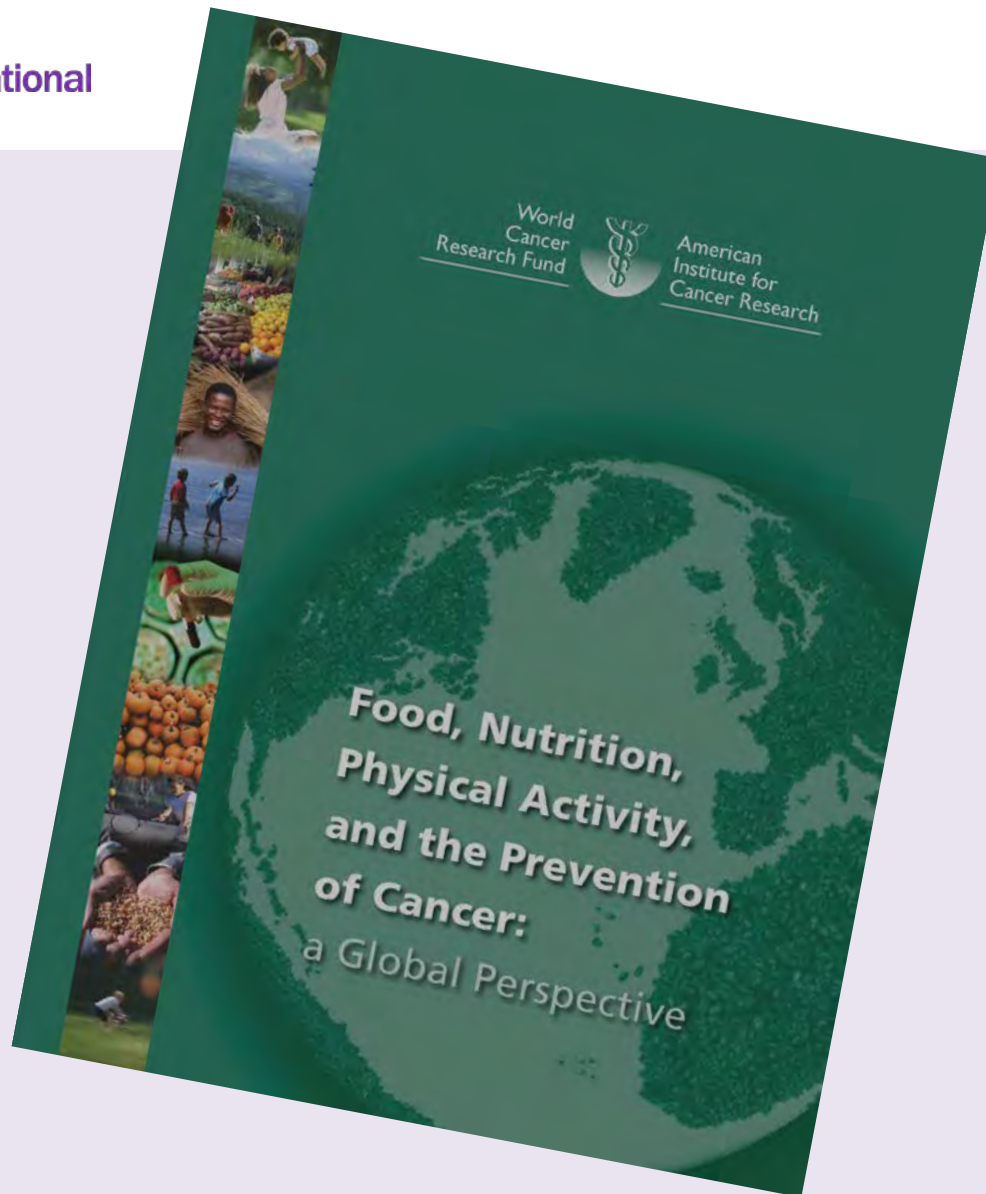


Two enabling characteristics for acquiring hallmarks

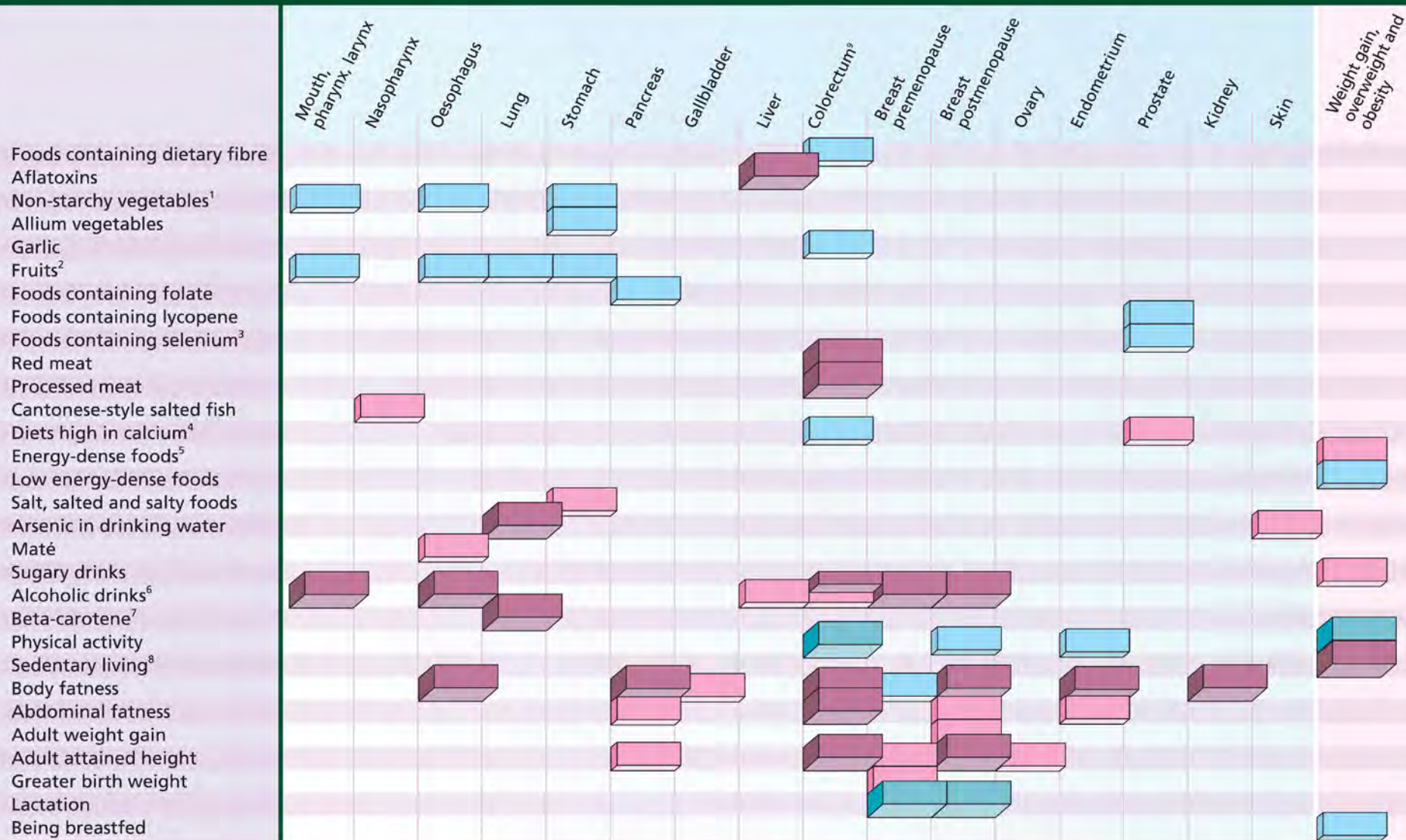


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Summary of 'convincing' and 'probable' judgements



KEY



Convincing decreased risk



Probable decreased risk



Probable increased risk



Convincing increased risk

¹ Includes evidence on foods containing carotenoids for mouth, pharynx, larynx; foods containing beta-carotene for oesophagus; foods containing vitamin C for oesophagus

² Includes evidence on foods containing carotenoids for mouth, pharynx, larynx and lung; foods containing beta-carotene for oesophagus; foods containing vitamin C for oesophagus

³ Includes evidence from supplements for prostate

⁴ Evidence is from milk and studies using supplements for colorectum

⁵ Includes 'fast foods'

⁶ Convincing harm for men and probable harm for women for colorectum

⁷ The evidence is derived from studies using supplements for lung

⁸ Includes evidence on television viewing

⁹ Judgement for physical activity applies to colon and not rectum



The Panel emphasises the importance of not smoking and of avoiding exposure to tobacco smoke

RECOMMENDATIONS

BODY FATNESS

Be as lean as possible within the normal range of body weight

PHYSICAL ACTIVITY

Be physically active as part of everyday life

FOODS AND DRINKS THAT PROMOTE WEIGHT GAIN

Limit consumption of energy-dense foods
Avoid sugary drinks

PLANT FOODS

Eat mostly foods of plant origin

ANIMAL FOODS

Limit intake of red meat and avoid processed meat

ALCOHOLIC DRINKS

Limit alcoholic drinks

PRESERVATION, PROCESSING, PREPARATION

Limit consumption of salt
Avoid mouldy cereals (grains) or pulses (legumes)

DIETARY SUPPLEMENTS

Aim to meet nutritional needs through diet alone

BREASTFEEDING

Mothers to breastfeed; children to be breastfed

CANCER SURVIVORS

Follow the recommendations for cancer prevention



SUMMARY OF STRONG EVIDENCE ON DIET, NUTRITION, PHYSICAL ACTIVITY AND PREVENTION OF CANCER

	MOUTH, PHARYNX, LARYNX (2007)	NASOPHARYNX (2007)	ESOPHAGUS (2007)	LUNG (2007)	STOMACH (2007)	PANCREAS (2012)	GALLBLADDER (2015)	LIVER (2015)	COLORECTUM (2011)	BREAST PREMENOPAUSE (2010)	BREAST POSTMENOPAUSE (2010)	OVARY (2014)	ENDOMETRIUM (2013)	PROSTATE (2014)	KIDNEY (2015)	SKIN (2007)
Foods containing dietary fibre									Convincing decreased risk							
Aflatoxins								Convincing increased risk								
Non-starchy vegetables¹	Probable decreased risk		Probable decreased risk		Probable decreased risk											
Allium vegetables					Probable decreased risk											
Garlic									Probable decreased risk							
Fruits²	Probable decreased risk		Probable decreased risk	Probable decreased risk	Probable decreased risk											
Red meat									Convincing increased risk							
Processed meat									Convincing increased risk							
Cantonese-style salted fish		Probable increased risk														
Diets high in calcium³									Probable decreased risk							
Salt, salted and salty foods						Probable increased risk										
Glycaemic load													Probable increased risk			
Arsenic in drinking water				Convincing increased risk												Probable increased risk
Maté			Probable increased risk													
Alcoholic drinks⁴	Convincing increased risk		Convincing increased risk					Convincing increased risk	Probable increased risk	Convincing increased risk	Convincing increased risk				Probable increased risk	Probable decreased risk
Coffee						Substantial effect on risk unlikely		Probable decreased risk					Probable decreased risk			
Beta-carotene⁵				Convincing increased risk										Substantial effect on risk unlikely		Substantial effect on risk unlikely
Physical activity⁶									Convincing decreased risk		Probable decreased risk		Probable decreased risk			
Body fatness⁷			Convincing increased risk			Convincing increased risk	Probable increased risk	Convincing increased risk	Convincing increased risk	Probable decreased risk	Convincing increased risk	Probable increased risk	Convincing increased risk	Convincing increased risk	Convincing increased risk	Convincing increased risk
Adult attained height⁸						Probable increased risk			Convincing increased risk	Probable increased risk	Convincing increased risk	Convincing increased risk		Probable increased risk	Convincing increased risk	
Greater birth weight										Probable increased risk						
Lactation										Convincing decreased risk	Convincing decreased risk					



NUTRITION AND CANCERS

- **ADIPOSIITY**
 - BREAST (PM), COLORECTUM, ENDOMETRIUM, OESOPHAGUS, PANCREAS, GALLBLADDER, KIDNEY, OVARY, PROSTATE (ADVANCED), LIVER
- **PHYSICAL (IN)ACTIVITY**
 - COLON, BREAST, ENDOMETRIUM
- **MEAT – RED AND PROCESSED**
 - COLON, RECTUM, STOMACH (non-cardia)
- **ALCOHOL**
 - MPL, BREAST, COLORECTUM, LIVER, OESOPHAGUS
- **PLANT FOODS (F&V, PULSES, WHOLEGRAINS)**
 - MPL, OESOPHAGUS, STOMACH, COLORECTUM (DF), LUNG
- **BREASTFEEDING**
 - BREAST (MOTHER), OBESITY (CHILD)

Publications timetable

Year	Publication
2010	Breast
2011	Colorectum
2012	Pancreas
2013-14	Endometrium, ovary, breast cancer survivors, prostate
2014-15	Bladder, kidney, liver, gallbladder
2016	Stomach, oesophagus
2017	Mouth, pharynx, larynx, nasopharynx, lung, breast, colorectum

2017 **Review Recommendations for Cancer Prevention**

Cancer	Exposure	2007	New
Colorectum	Dietary fibre	Prob ↓	Convincing ↓
Endometrium	Coffee	-	Prob ↓
	Glycaemic load	-	Prob ↑
Liver	Body fatness	LS ↑	Convincing ↑
	Coffee	-	Prob ↓
Kidney	Height	LNC	Prob ↑
	Alcohol	Effect unlikely	Prob ↑
Bladder	Arsenic	LS ↑	Prob ↑
Ovary	Body fatness	LNC	Prob ↑
Pancreas	Folate	Prob ↓	LNC
Prostate	Body fatness	LNC	Prob ↑ (adv)
	Height	LNC	Prob ↑
Oesophagus	Fruit/veg/ β carotene/vit C	Prob ↓	LS↓/LNC
Stomach	Body fatness	LNC	Prob ↑ (cardia)
	Alcohol	LNC	Prob ↑
	Processed meat	LNC	Prob ↑ (non-cardia)
	Fruit/veg	Prob ↓	LNC/LS ↓

**Estimates of
cancer
preventability by
appropriate diet,
nutrition,
physical activity
and body fatness**

	USA	UK	BRAZIL	CHINA
Mouth, pharynx, larynx	63	67	63	44
Oesophagus	63	71	50	33
Lung	36	33	36	38
Stomach	47	45	41	33
Pancreas	19	15	11	8
Gallbladder	21	16	10	6
Liver	30	24	13	7
Colorectum	50	47	41	22
Breast	33	38	22	11
Ovary	5	4	3	1
Endometrium	59	44	37	21
Prostate (advanced)	11	9	5	4
Kidney	24	19	13	8
Total for these cancers	31	32	25	24
Total for all cancers	21	24	18	20

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Our Cancer Prevention Recommendations

Be a healthy weight

Keep your weight as low as you can within the healthy range

Move more

Be physically active for at least 30 minutes every day, and sit less

Avoid high-calorie foods and sugary drinks

Limit high-calorie foods (particularly processed foods high in fat or added sugar, or low in fibre) and avoid sugary drinks

Enjoy more grains, veg, fruit and beans

Eat a wide variety of whole grains, vegetables, fruit and pulses such as beans

Limit red meat and avoid processed meat

Eat no more than 500g (cooked weight) a week of red meat, such as beef, pork and lamb. Eat little, if any, processed meat such as ham and bacon

For cancer prevention, don't drink alcohol

For cancer prevention, it's best not to drink alcohol. If you do, limit alcoholic drinks to two for men and one for women a day

Eat less salt and avoid mouldy grains & cereals

Limit your salt intake to less than 6g (2.4g sodium) a day by adding less salt and eating less food processed with salt

Avoid mouldy grains and cereals as they may be contaminated by aflatoxins

For cancer prevention, don't rely on supplements

Eat a healthy diet rather than relying on supplements to protect against cancer

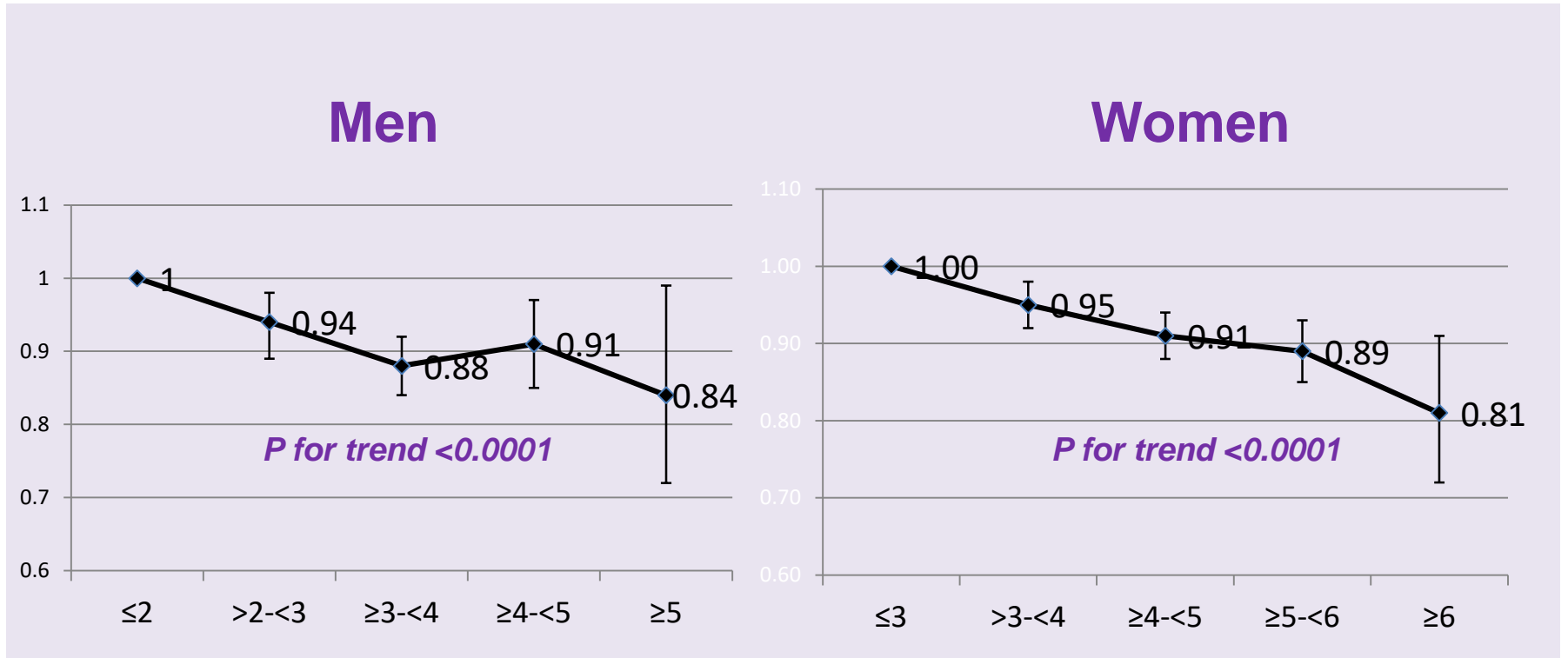
If you can, breastfeed your baby

If you can, breastfeed your baby for six months before adding other liquids and foods

Cancer survivors should follow our Recommendations (where possible)

After cancer treatment, the best advice is to follow the Cancer Prevention Recommendations. Check with your health professional

Association between the WCRF/AICR score and total cancer risk

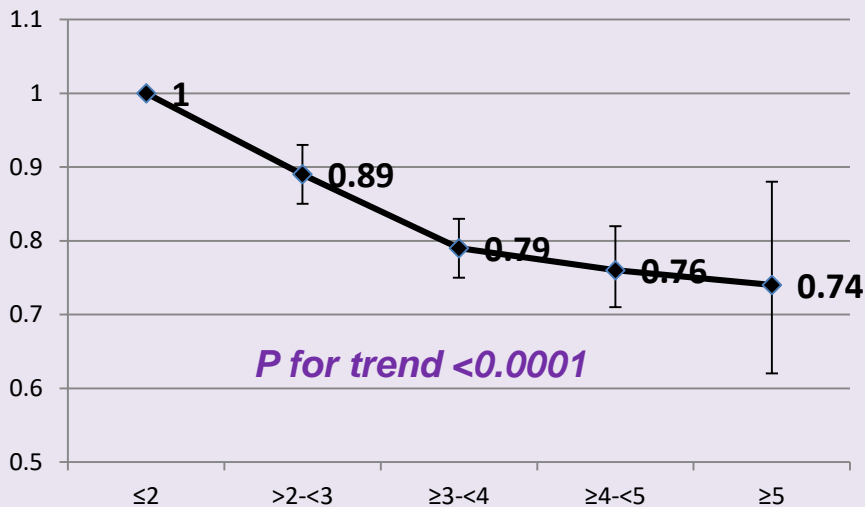


Cox regression model stratified by centre and age, and adjusted by energy intake, level of school, smoking status, presence of chronic diseases at baseline, ever use of contraceptive pills, ever use of HRT, age at first menarche, age at first pregnancy, and menopausal status

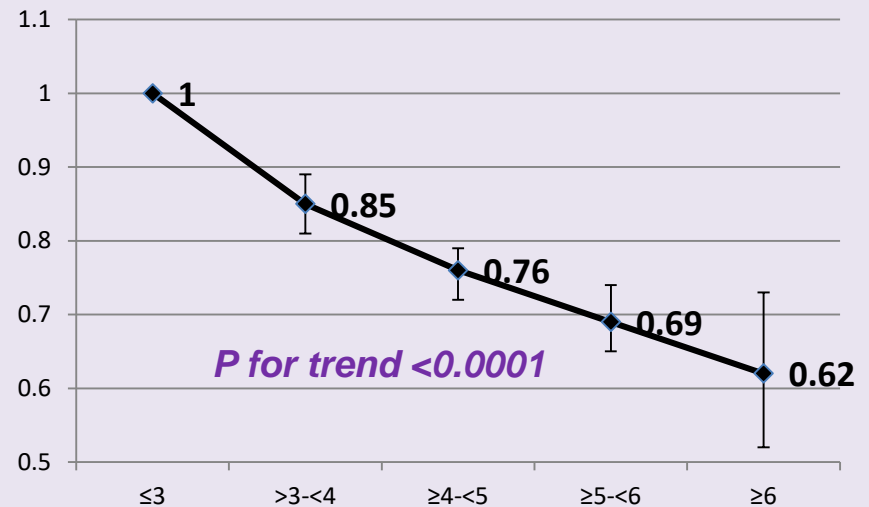
Romaguera D et al, AJCN 2012

Association between the WCRF/AICR score and total mortality

Men



Women



Cox regression model stratified by centre and age, and adjusted by level of school, smoking status, smoke intensity, and menopausal status

Vergnaud AC et al, AJCN 2013

WCRF and ACS recommendations and cancer – systematic review

- Ten large prospective studies; 12 publications
- Strong and consistent evidence
- Greater adherence to score associated with lower overall cancer incidence and mortality
- Both men and women
- Also breast, colorectal endometrium

Kohler LN et al, CEBP 2016, 25, 1-11

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Inferring causality

- **Strength**
- **Consistency**
- **Specificity**
- **Timing**
- **Dose
Response**
- **Plausibility**
- **Coherence**
- **Experiment**
- **Analogy**

Bradford Hill

Reasons for uncertainty

- Measurement error
 - Diet, activity, anthropometry (cf adiposity)
 - Random error, systematic bias
- Study design
 - RCT vs cohort vs case control
 - Mechanistic
 - Population
 - Study size
- Confounding
 - Smoking
 - Nutrient vs food
 - Multiple collinearity eg PA
- Exposure homogeneity
- Small effect size

Certainty is unattainable – degrees of uncertainty

Is the evidence strong enough to take action?

Evidence accrues and conclusions may change

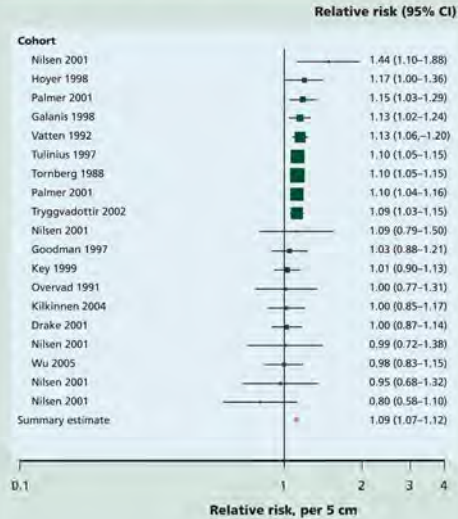
Convincing is not the same as proof

Significant shifts in emphasis

- Adiposity and activity vs foods and drinks
- Foods vs nutrients
- Whole diets vs individual foods
- Plant foods vs fruit and veg
- Lifecourse (height)

IMPACT OF OFFSPRING SIZE AND GROWTH ON CANCER RISK - 2007

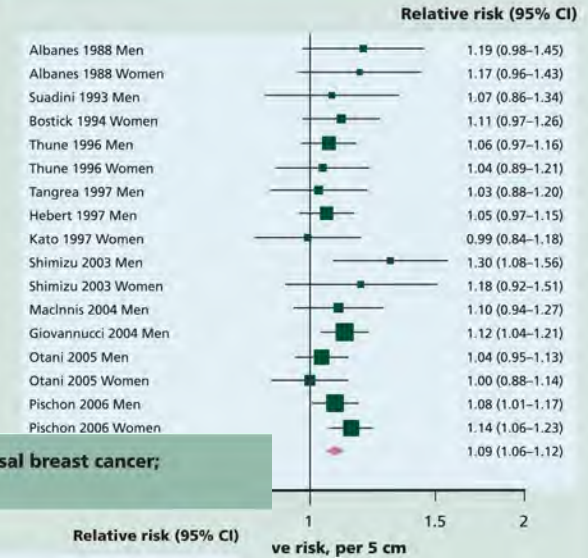
Height and breast cancer (age unspecified); cohort studies



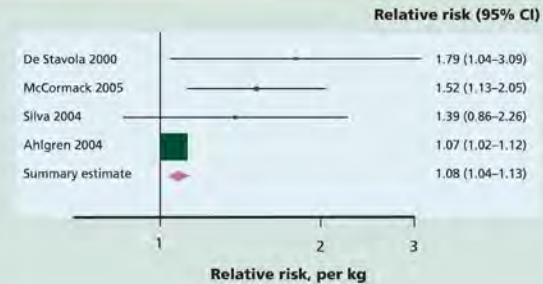
Height and postmenopausal breast cancer; cohort studies



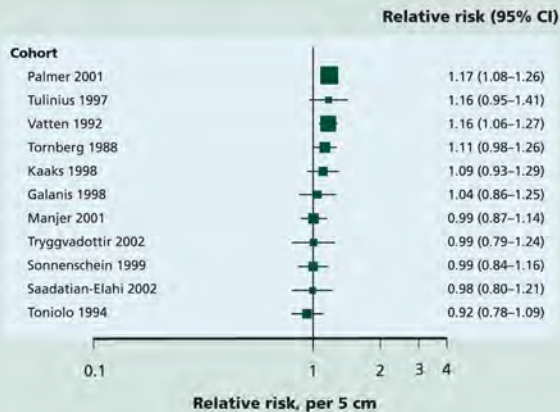
Height and colorectal cancer; cohort studies



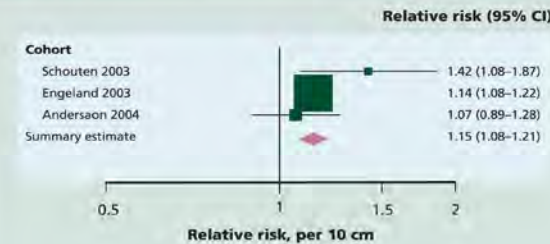
Birth weight and premenopausal breast cancer; cohort studies



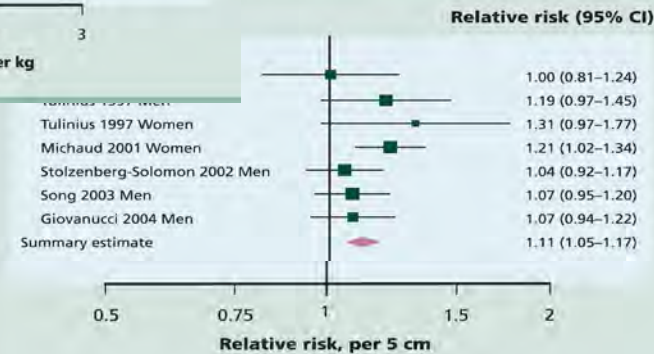
Height and premenopausal breast cancer; cohort studies



Height and ovarian cancer; cohort studies



Height and colorectal cancer; cohort studies



Height and cancer

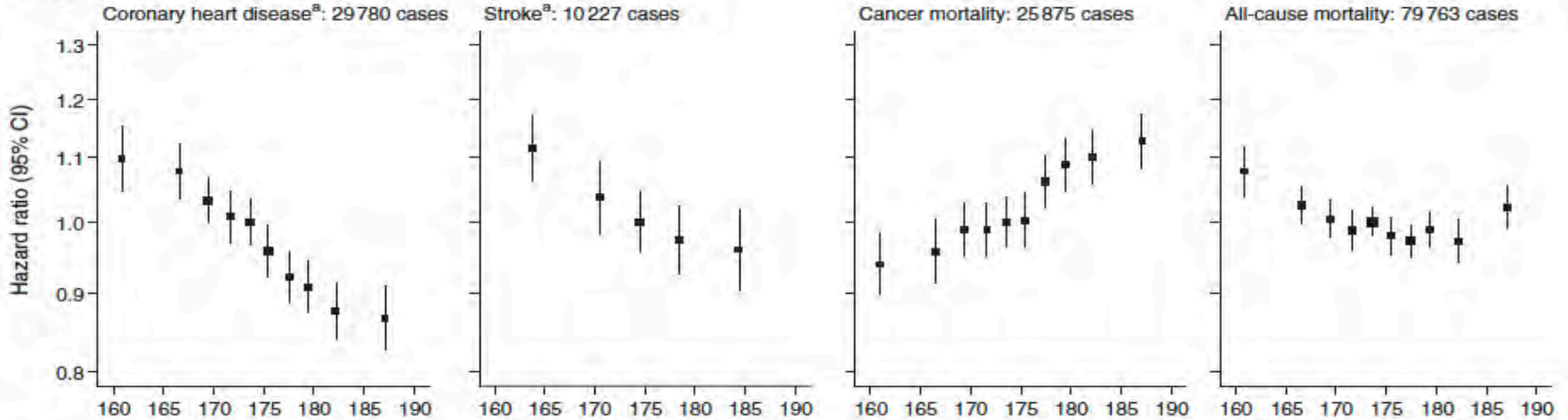
CUP 2015

Every 5 cm increment in height increases risk of cancers of:

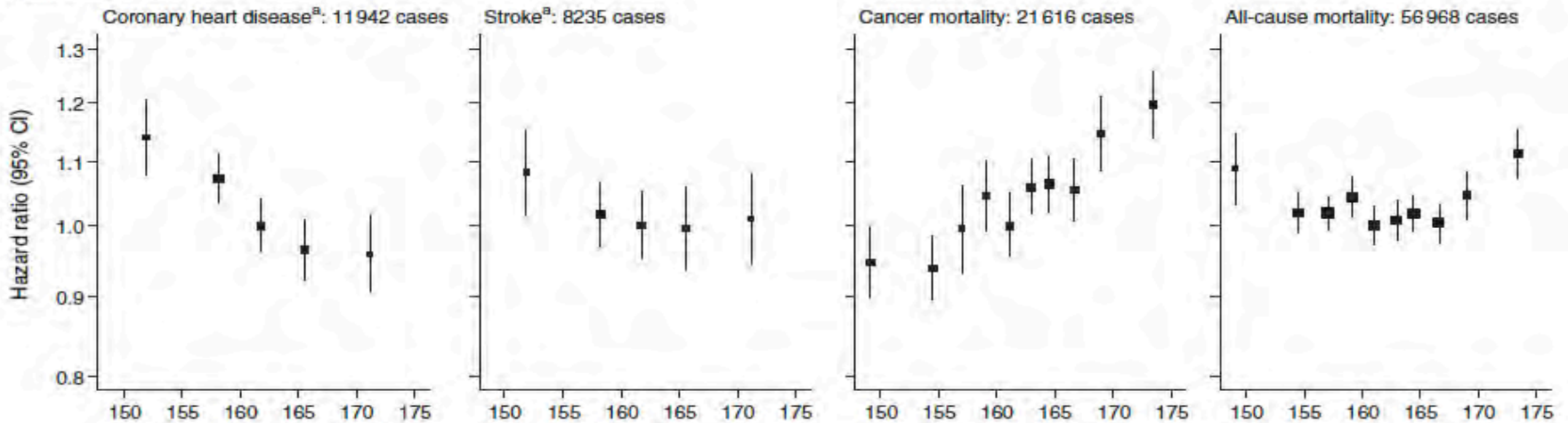
- Kidney -10%
- Breast (pre-menopausal) - 9%
- Breast (post-menopausal) -11%
- Ovary - 8%
- Pancreas - 7%
- Colorectum - 5%
- Prostate – 4%

Height and risk of CVD and cancer

MALES:



FEMALES:



CVD

Mean height (cm)

Cancer

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Bottom Line

The key messages are robust and generally agreed

- Be active, and don't be sedentary – and keep it up as long as possible
- Eat enough but not too much – don't get too thin or fat
- Eat food not pills
- Mostly from plants, emphasise wholegrains and pulses
- Avoid highly processed energy dense foods and sugar sweetened beverages (and alcohol, processed meat and salty foods)
- Grow appropriately from conception to adulthood
- Get your mother to be well nourished before getting pregnant. And to breastfeed you.

Bottom Line

If you already have a diagnosis of cancer, the evidence on nutrition and outcome is not strong, but the best advice is to follow (as far as possible) the recommendations for cancer prevention

Cancer & Nutrition NIHR infrastructure collaboration

Improving cancer prevention and care.
For patients. For clinicians. For researchers

Aim:

To help facilitate translational research in cancer and nutrition which will generate the evidence to improve cancer prevention and care

Objectives:

To bring coherence to existing activities by

- creating a framework for future research
- establishing better networks between cancer and nutrition stakeholders



NHS
National Institute for
Health Research

Cancer and Nutrition NIHR infrastructure collaboration

Improving cancer prevention and care.
For patients. For Clinicians. For researchers.



Full Report of Phase One July 2015

JOHN TUKEY

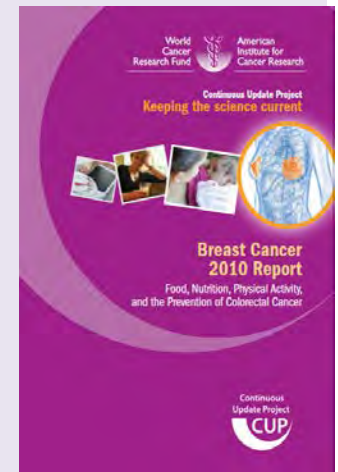
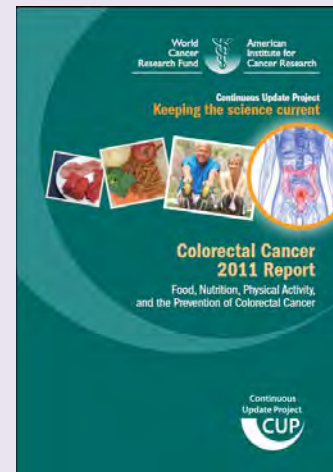
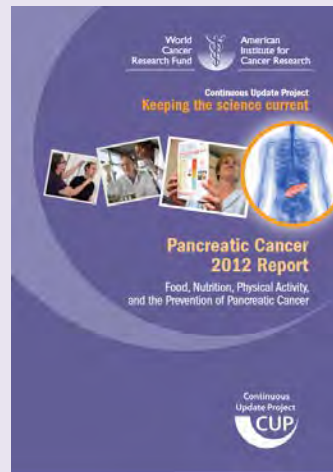
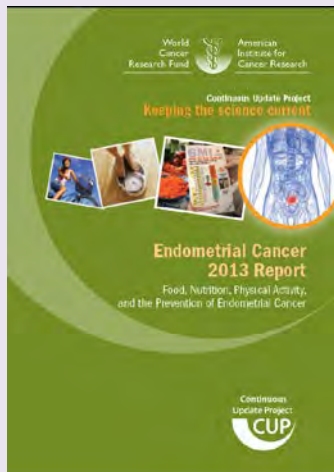


Far better an approximate answer to the *right* question, which is often vague, than an *exact* answer to the wrong question, which can always be made precise.

The future of data analysis. Annals of Mathematical Statistics 1962

An approximate answer to the right question is worth a great deal more than a precise answer to the wrong question.

– *Super Freakonomics*



http://www.wcrf.org/cancer_research/cup/key_findings/index.php

Watch this space....

2017 update is coming...



http://www.wcrf.org/cancer_research/cup/key_findings/index.php



World
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Breast Cancer (C50): 2014

Number of New Cases, Crude and European Age-Standardised (AS) Incidence Rates per 100,000 Population, UK

		England	Wales	Scotland	Northern Ireland	UK
Female	Cases	46,085	2,872	4,583	1,293	54,833
	Crude Rate	167.3	182.8	166.6	137.9	167.2
	AS Rate	173.4	176.1	164.1	151.4	172.1
	AS Rate - 95% LCL	171.8	169.7	159.3	143.2	
	AS Rate - 95% UCL	175.0	182.5	168.8	159.7	173.6
						170.7

AS rate 173.4 182.8 **166.6** 151.4 172.1

95% LCL and 95% UCL are the 95% lower and upper confidence limits around the AS Rate

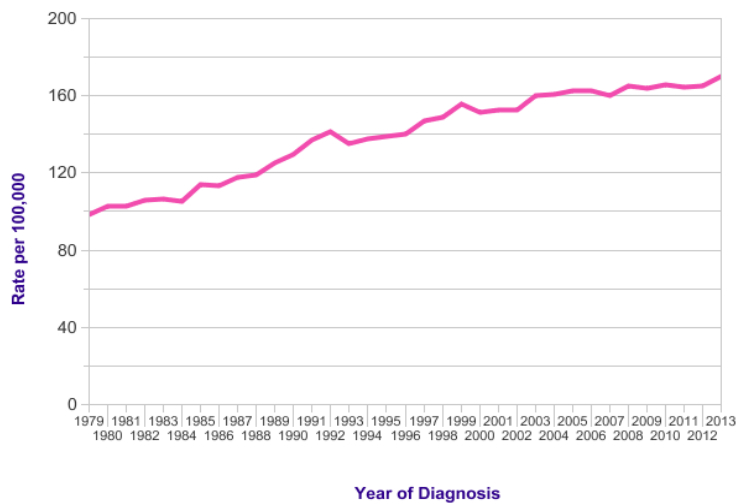
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Breast Cancer (C50): 1979-2013
European Age-Standardised Incidence Rates
per 100,000 Population, Females, Great Britain

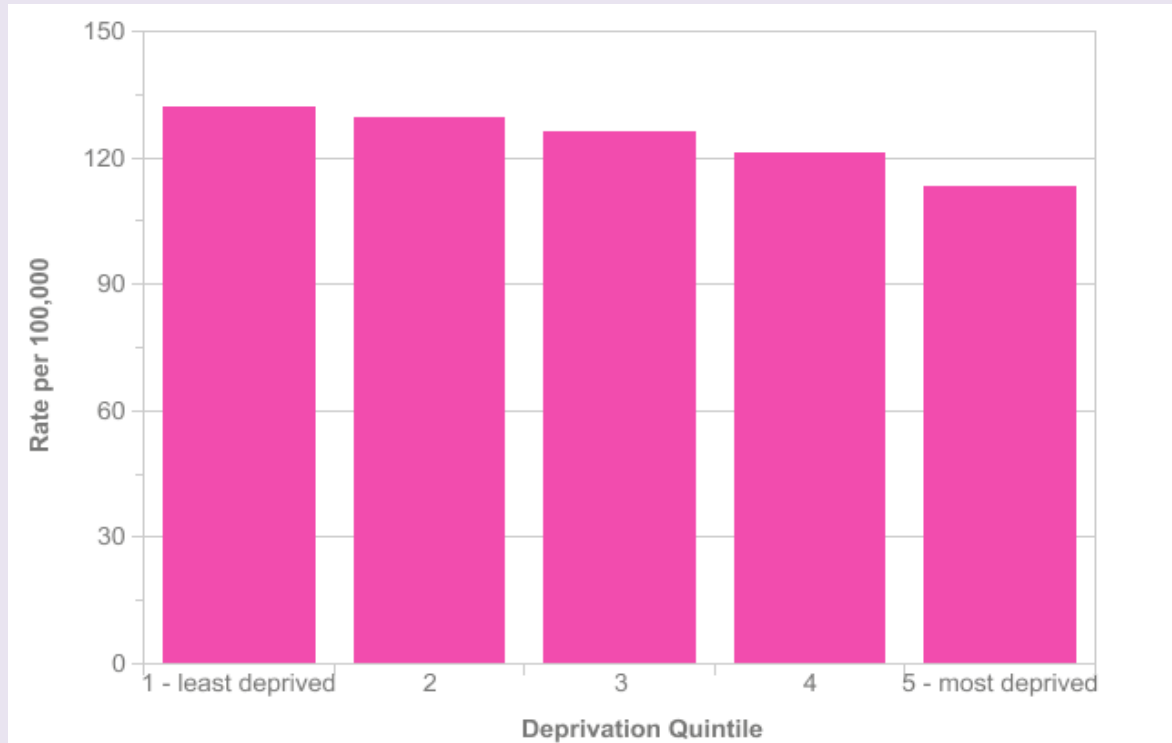


Breast Cancer (C50): 1971-2014
European Age-Standardised Mortality Rates
per 100,000 Population, Females, UK



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Breast Cancer (C50): 2006-2010
European Age-Standardised Incidence Rates by Deprivation Quintile, England



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Breast Cancer

FOOD, NUTRITION, PHYSICAL ACTIVITY AND BREAST CANCER (PREMENOPAUSE) 2010

	DECREASES RISK	INCREASES RISK
Convincing	Lactation	Alcoholic drinks
Probable	Body fatness	Adult attained height ¹ Greater birth weight
Substantial effect on risk unlikely	None identified	

¹ Adult attained height is unlikely directly to modify the risk of cancer. It is a marker for genetic, environmental, hormonal, and also nutritional factors affecting growth during the period from preconception to completion of linear growth (see chapter 6.2.13 – Second Expert Report).

FOOD, NUTRITION, PHYSICAL ACTIVITY AND BREAST CANCER (POSTMENOPAUSE) 2010

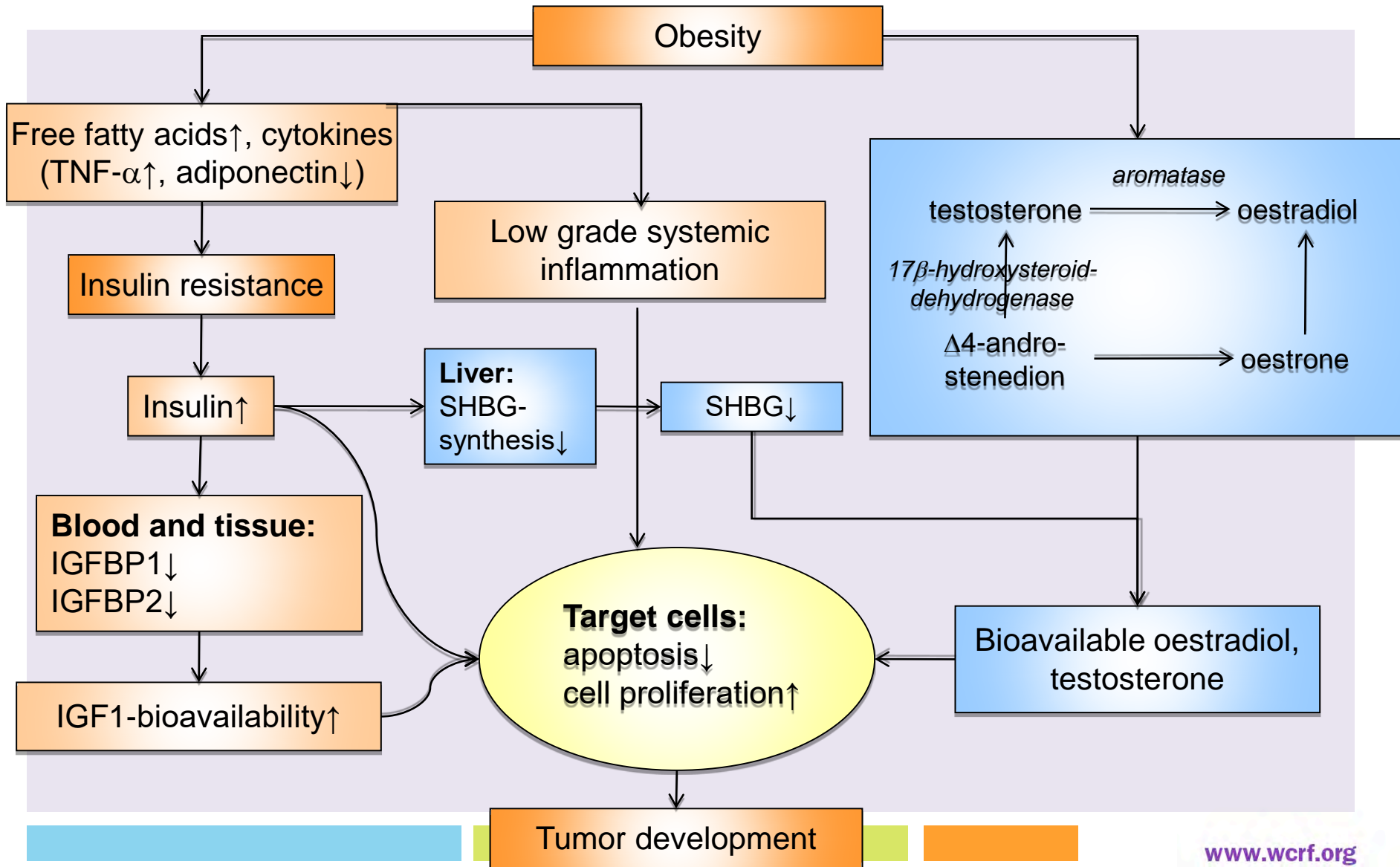
	DECREASES RISK	INCREASES RISK
Convincing	Lactation	Alcoholic drinks Body fatness Adult attained height ¹
Probable	Physical activity ²	Abdominal fatness Adult weight gain
Substantial effect on risk unlikely	None identified	

¹ Adult attained height is unlikely directly to modify the risk of cancer. It is a marker for genetic, environmental, hormonal, and also nutritional factors affecting growth during the period from preconception to completion of linear growth (see chapter 6.2.13 – Second Expert Report).

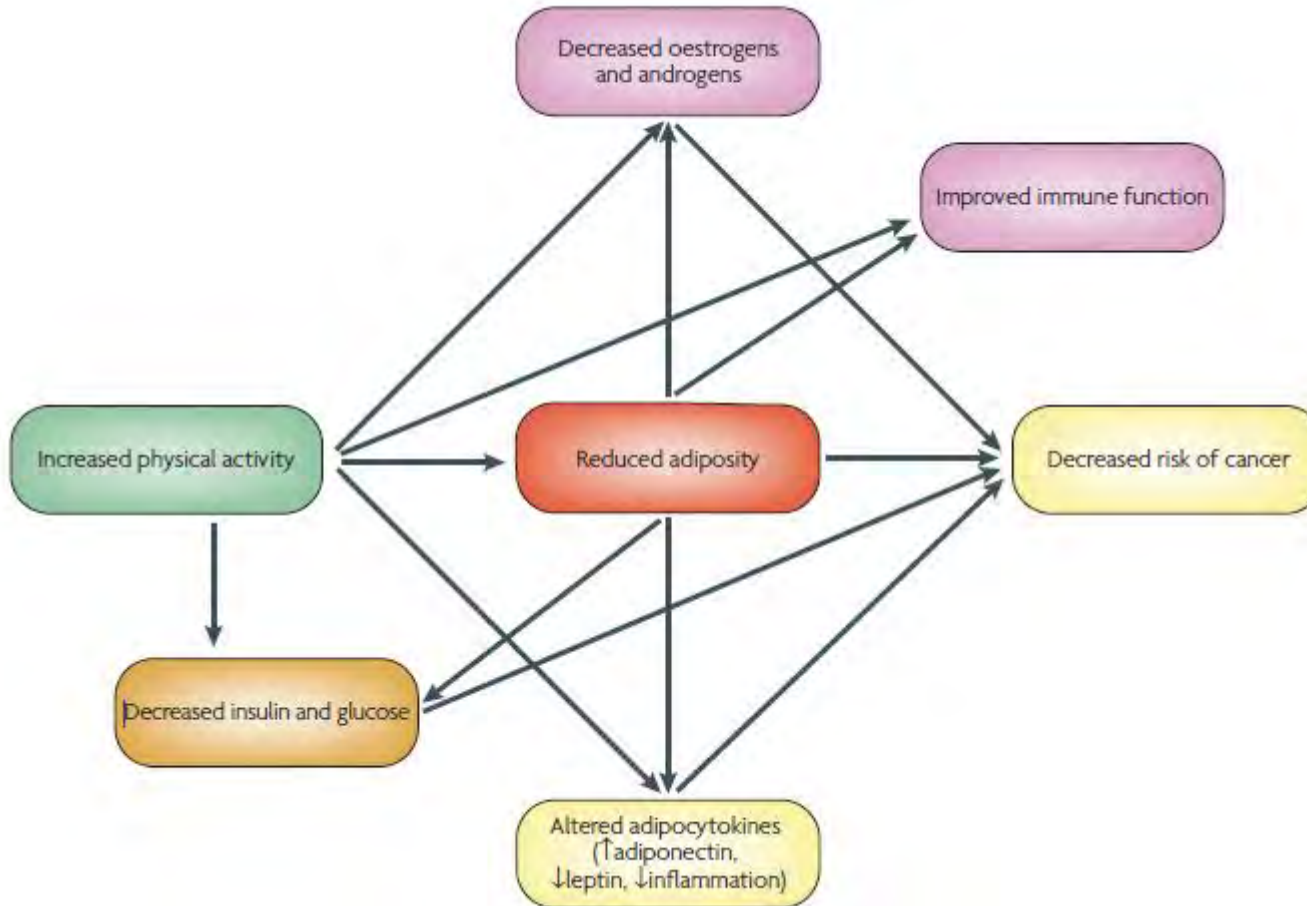
² Physical activity of all types: occupational, household, transport and recreational.



Obesity and Cancer – Potential Mechanisms



Mechanisms linking physical activity and cancer



McTiernan 2008



Obesity, physical inactivity and cancer Mechanisms - Summary

- Insulin resistance
 - abnormal insulin/IGF axis
 - excess growth factors
- Inflammation
 - promotes oncogenic genetic signature
 - increases proliferation, angiogenesis
 - reduces apoptosis
- Excess oestrogen
 - promotes proliferation and cancer in sensitive tissues
- Reduced immune function

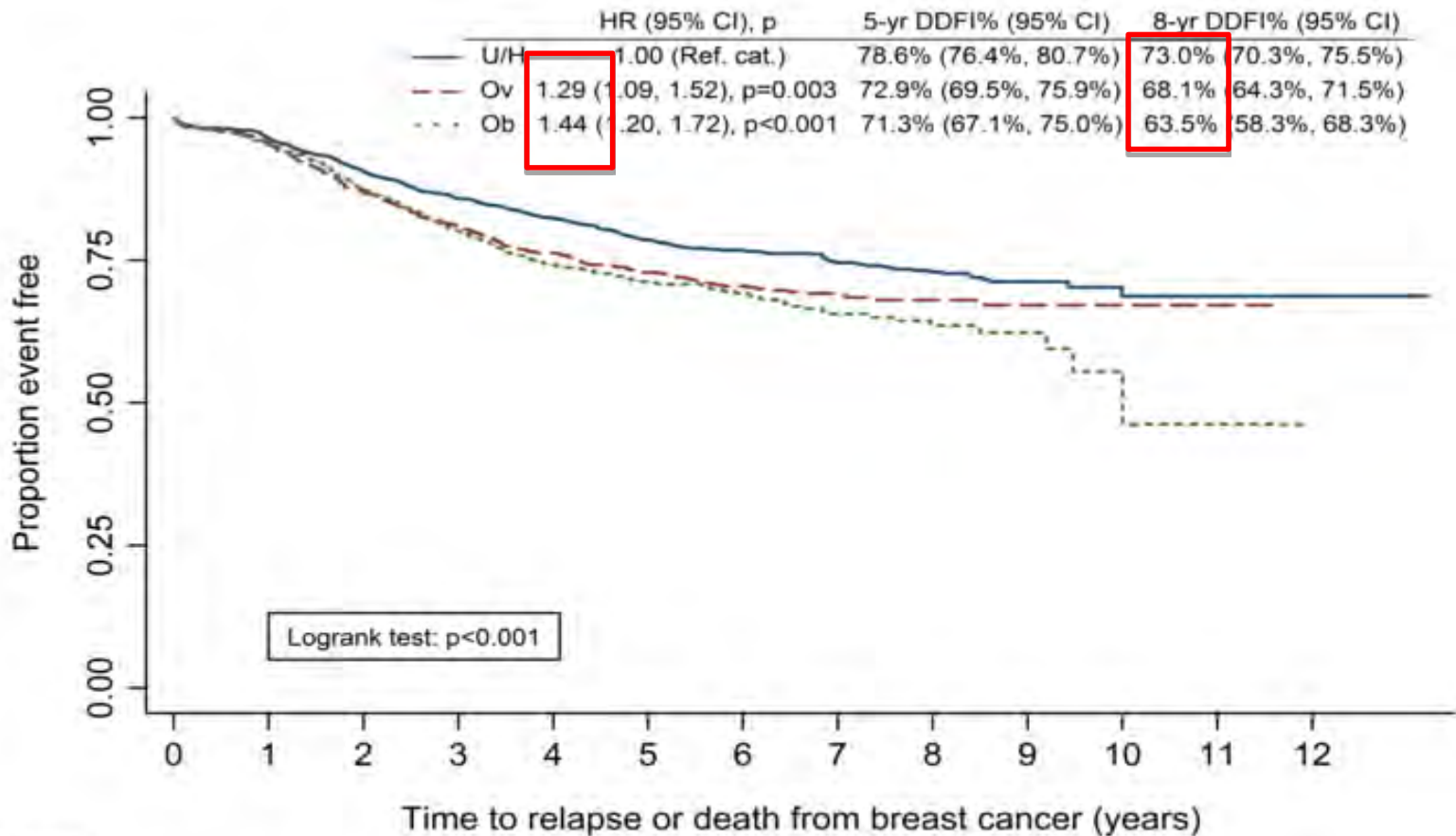


Pathological features - POSH

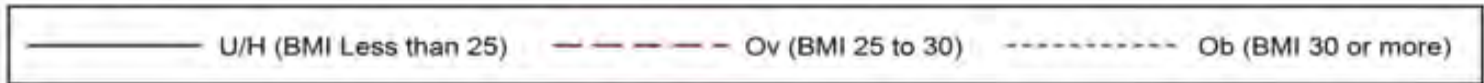
	Underweight or Healthy weight n=1526 (54.0%)	Overweight n=784 (27.6%)	Obese n=533 (18.8%)	
Mean tumour size/ mm	20 (0-170)	24 (0-199)	26 (0.5-130)	U/H vs. Ov: p<0.0001 U/H vs. Ob: p<0.0001
Multifocal	12 (30.6%)	220 (30.4%)	130 (27.2%)	NS
Grade 3	879 (59.0%)	485 (63.6%)	331 (63.9%)	U/H vs. Ov: p=0.034 U/H vs. Ob: p =0.048
Node positive	736 (49.0%)	419 (54.2%)	284 (54.6%)	U/H vs. Ov: p=0.019 U/H vs. Ob: p=0.027
ER negative	483 (31.7%)	273 (34.9%)	213 (40.1%)	U/H vs Ob: p<0.001
HER 2 positive	381 (28.2%)	180 (26.4%)	129 (27.3%)	NS
ER/ PR/ HER 2 negative	305 (20.8%)	176 (23.4%)	136 (26.8%)	U/H vs. Ob: p=0.005



Distant disease free survival – POSH



	0	1	2	3	4	5	6	7	8	9	10	11	12
U/H (BMI Less than 25)	1518	1459	1355	1256	1144	938	679	462	284	137	44	14	3
Ov (BMI 25 to 30)	778	738	663	611	544	462	321	206	116	42	14	2	0
Ob (BMI 30 or more)	525	498	453	409	358	291	199	131	74	30	6	1	0



DIET, NUTRITION, PHYSICAL ACTIVITY AND BREAST CANCER SURVIVAL (BY OUTCOME)

Outcome	ALL CAUSE MORTALITY		BREAST CANCER MORTALITY		SECOND PRIMARY BREAST CANCER	
	DECREASED RISK	INCREASED RISK	DECREASED RISK	INCREASED RISK	DECREASED RISK	INCREASED RISK
	Exposure Timeframe	Exposure Timeframe	Exposure Timeframe	Exposure Timeframe	Exposure Timeframe	Exposure Timeframe
STRONG EVIDENCE	Convincing					
	Probable					
LIMITED EVIDENCE	Limited-suggestive	Physical activity Before diagnosis >12 months after diagnosis	Body fatness Before diagnosis <12 months after diagnosis >12 months after diagnosis	Physical activity Before diagnosis	Body fatness ¹ Before diagnosis <12 months after diagnosis	Body fatness Before diagnosis <12 months after diagnosis
		Foods containing fibre Before diagnosis >12 months after diagnosis	Total fat Before diagnosis			
		Foods containing soy >12 months after diagnosis	Saturated fatty acids Before diagnosis			
STRONG EVIDENCE	Substantial effect on risk unlikely					

STRONG: Evidence strong enough to support a judgement of a convincing or probable causal relationship and generally justify making recommendations
LIMITED: Evidence that is too limited to justify making specific recommendations

¹ Post menopause only

Summary of Breast Cancer Survivors Report

- Although there were significant associations between some exposures and outcomes, incomplete adjustment for potential confounders restricted the ability to ascribe causality
- CUP Panel concluded that evidence is limited