

### Cancer Sites Linked to Overweight/Obesity

- Esophagus
- Gallbladder
- Pancreas
- Liver
- Colon
- Rectum
- Post-menopausal Breast
- Endometrial
- Aggressive Prostate
- Non-Hodgkin's Lymphoma
- Leukemia
- Multiple Myeloma
- Renal\*
- Melanoma
- Thyroid

The Clinical Guide to Oncology Nutrition/Oncology Nutrition DPG, 2<sup>nd</sup> ed. 2008. American Dietetic Association.  
\*Renal thyroid and melanoma cited by Wolin, Carson, and Colditz.

### Obesity and Cancer

Wolin, K.Y., Carson, K, Colditz, G.A. Obesity and Cancer. *The Oncologist* 2010; 15:556-565.

- 20% cancer cases caused by obesity
- Complicates chemo dosing & tx toxicity
- Poor treatment outcomes
- Higher mortality rates
- Less disease free time for survivors
- Current Research
  - What is the underlying biology of the relationship between obesity and the various cancer sites?

### Hormones, Growth Factors, and Cytokines, Oh My!

- Blood glucose & Insulin (c-peptide)
  - ↑ Colon, Prostate, Endometrial
  - ? Kidney and Pancreatic
- Estrogen
  - ↑ Endometrial and Breast
  - ? Colon
- Leptin
  - ↑ Colorectal and Prostate
- TNF, Interleukin, C-RP
  - ↑ Pro-inflammatory factors produced by adipocytes
- Insulin-like growth factor



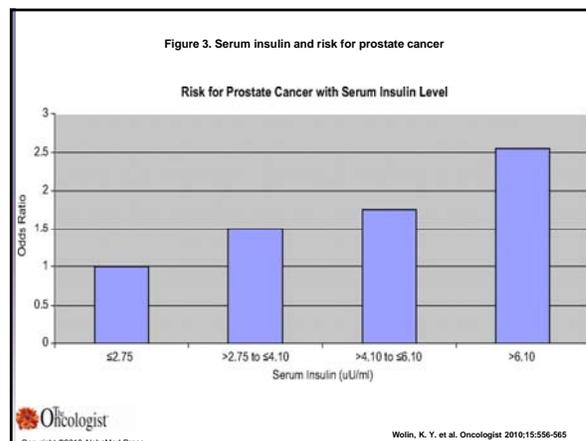
World Cancer Fund & American Institute for Cancer Research 2<sup>nd</sup> Report on Food, Nutrition, Physical Activity, and the Prevention of Cancer: A Global Perspective. P. 39. Washington DC AICR 2007.

**Table 1. RR for cancer per 5 kg/m<sup>2</sup> higher BMI and most likely causal mechanism: Males**

Cancer type	RR	Causal mechanism
Esophageal adenocarcinoma	1.52 <sup>a</sup>	Reflux esophagitis and chronic irritation
Thyroid	1.33 <sup>a</sup>	Unknown
Colon	1.24 <sup>a</sup>	Insulin
Renal	1.24 <sup>a</sup>	In part though hypertension
Liver	1.24	Fatty liver cirrhosis
Malignant melanoma	1.17 <sup>b</sup>	?
Multiple myeloma	1.11 <sup>a</sup>	Inflammatory pathways—IL-6
Rectum	1.09 <sup>a</sup>	?
Gallbladder	1.09	Chronic secretion-gallstones and irritation
Leukemia	1.08 <sup>b</sup>	?
Pancreas	1.07	Possible insulin pathway
Non-Hodgkin's lymphoma	1.06 <sup>a</sup>	Inflammatory pathways—IL-6
Prostate <sup>c</sup>	1.03	?
Lung	0.76 <sup>a</sup>	Smoking leads to leanness and causes lung cancer
Esophageal squamous	0.71 <sup>a</sup>	Smoking leads to leanness and causes squamous esophageal cancer

Shown is the RR for a five-point greater BMI—for example, the RR linked to a BMI of 28 compared with a BMI of 23, or a BMI of 32 compared with a BMI of 27.  
<sup>a</sup>p < .0001.  
<sup>b</sup>p < .01.  
<sup>c</sup>p < .05.  
<sup>d</sup>Biased to null because this includes predominantly low-grade lesions.  
Abbreviations: BMI, body mass index; IL, interleukin; RR, relative risk.  
Based on Figure 3 of Renehan AG, Tyson M, Egger M et al. Body-mass index and incidence of cancer: A systematic review and meta-analysis of prospective observational studies. *Lancet* 2008;371:569–578.

Copyright ©2010 AlphaMed Press Wolin, K. Y. et al. *Oncologist* 2010;15:556-565



**Table 2. RR for cancer per 5 kg/m<sup>2</sup> higher BMI and most likely causal mechanism: Females**

Cancer type	RR	Causal mechanism
Endometrium	1.59*	Endogenous estrogen
Gallbladder	1.59*	Chronic secretion-gallstones and irritation
Esophageal adenocarcinoma	1.51*	Reflux esophagitis and chronic irritation
Renal	1.34*	In part through hypertension
Leukemia	1.17*	Unknown
Thyroid	1.14*	Unknown
Breast (postmenopausal)	1.12*	Endogenous estrogen
Pancreas	1.12*	Possible insulin pathway
Multiple myeloma	1.11*	Inflammatory pathways—IL-6
Colon	1.09*	Insulin
Non-Hodgkin's lymphoma	1.07	Inflammatory pathways—IL-6
Liver	1.07	Fatty liver cirrhosis
Breast (premenopausal)	0.92*	Irregular menstrual cycles, hormones
Lung	0.8*	Smoking leads to leanness and causes lung cancer
Esophageal squamous	0.57*	Smoking leads to leanness and causes squamous esophageal cancer

RR for a five-point greater BMI—for example, the RR linked to a BMI of 28 compared with a BMI of 23, or a BMI of 32 compared with a BMI of 27.  
 \*p < .0001.  
 †p < .01.  
 ‡p < .05.  
 Abbreviations: BMI, body mass index; IL, interleukin; RR, relative risk.  
 Based on Figure 4 of Reichen AG, Tyson M, Egger M et al. Body-mass index and incidence of cancer: A systematic review and meta-analysis of prospective observational studies. Lancet 2008;371:569–578.

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### Breast Cancer and Weight

- Risk of breast cancer ↓ 50% in women who lost >10kg and maintained (Eliassen et. al. 2006.)
- Nutrition goals during tx focus on avoiding weight gain
- Nutrition goals after tx focus on weight loss and building lean mass
- Physical activity and wt. loss reduce risk of recurrence



### Behavior Change Goals

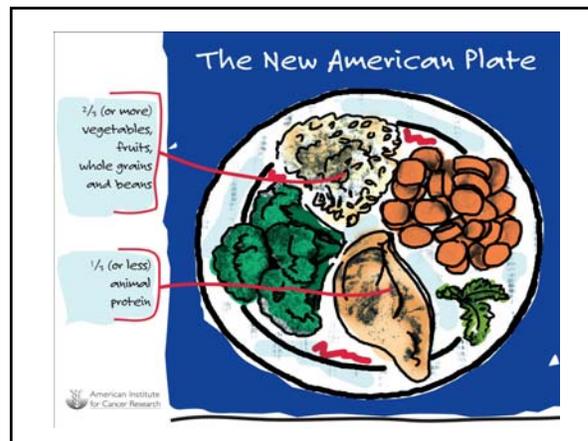
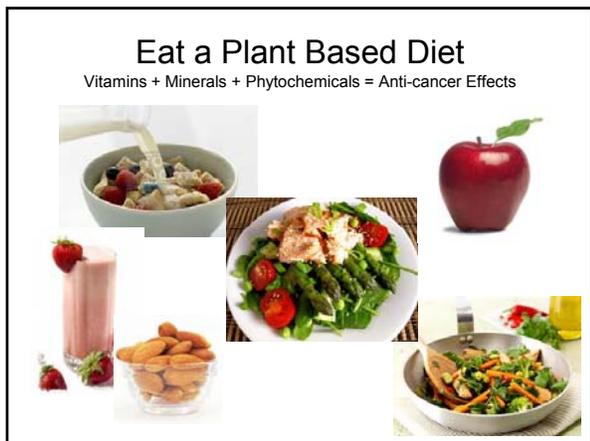
- Focus on behavior changes that result in creating an energy deficit
- Help individuals find motivation
- Maintain progress and motivation
- Wolin recommends applying the “Five A’s”
  - Assess, advise, agree, assist, arrange

### Meet Sarah

- 26 year-old teacher with back pain
- Started with lunch...
- Next dinner and sweets...
- Exercise
- Reinforcements
  - Consistent weight loss and shopping
- Continuous maintenance required

### AICR Recommendations for Cancer Prevention

1. Be as lean as possible within the normal range of body weight.
2. Be physically active as part of every day life.
3. Eat a plant-based diet.
  - Get at least 5 servings per day of non-starchy vegetables and fruits.
  - Eat unprocessed grains and beans with each meal.



Foods	Phytochemicals	Vitamins & Fiber	Antioxidant Action	Enzyme Formation	Cancer Inhibitor
Cruciferous veg, broccoli, cauliflower, cabbage	Indoles, sulforaphane, isothiocyanate	Folate, Vitamins A & C, Insoluble fiber			X
Allium veg, garlic, onion	Organosulfur compounds	Soluble and insoluble fiber	X		X
Berries, grapes, apples, nuts	Ellagic acid	Vitamins E & C, beta-carotene, Soluble fiber	X	X	X
Soybeans, beans, peas, lentils	Isoflavones (genistein), saponins, phytosterols	Soluble fiber			X
Citrus fruit	Terpenes, coumarins, flavanoids	Vitamin C, folate, Soluble fiber	X	X	X

Foods	Phytochemicals	Vitamins & Fiber	Antioxidant Action	Enzyme Formation	Cancer Inhibitor
Carrots, yams, canteloupe, butternut squash	Beta-carotene	Soluble and Insoluble fiber	X	X	X
Flaxseed, whole wheat, barley, brown rice	Lignans	Soluble and insoluble fiber	X		X
Tomatoes, tomato products, red grapefruit	Lycopene	Vitamin C, Soluble fiber	X		
Green tea, grapes, cocoa beans	Polyphenols	Soluble fiber			X
Hot peppers	Capsaicin	Vitamin C	X		X

### Tomato and Broccoli Study

- A lycopene supplement may not hurt you, but the whole tomato can help you more. A whole tomato may help you, but a tomato eaten with broccoli will help you more. Tomato with broccoli may help you, but a medley of different vegetables eaten together will bolster the body's different defenses against chronic disease."

--Jeff Prince, Vice-President for Education at AICR

- ### AICR Recommendations for Cancer Prevention
- Limit consumption of energy dense foods, avoid sugary drinks, and fast foods.
  - Limit intake of red meat (<18oz per week) and avoid processed meats.
  - Limit alcohol intake.
    - Men: 2 drinks or less per day
    - Women: 1 drink or less per day
  - Limit consumption of salt and avoid moldy grains and legumes.
  - Aim to meet nutrient needs through diet alone.

## Recommendations for Special Populations

9. Mothers to breastfeed and children to be breastfed.
10. Cancer survivors should aim to follow the recommendations for nutrition, physical activity, and weight.

## Physical Activity & Colon Cancer

Physical Activity, Sedentary Behavior, and the Risk of Colon and Rectal Cancer in the NIH-AARP Diet and Health Study.

- 488,720 Subjects age 50-71, 7 year follow up
- Observed 18% lower risk of colon cancer exercising 5x per week Vs. rarely/never
  - Men “significant decrease” (  $p = 0.001$ , 95% CI)
  - Women “suggested decrease” ( $p = 0.376$ , 95% CI)
- Inverse relations with low and moderate-vigorous intensity
- Positive association with sedentary behavior and colon cancer

## Study Conclusion

*“Engaging in physical activity of any intensity is associated with reductions in colon and rectal cancer risk. Conversely, time spent sedentary is associated with increased colon cancer risk.”*

Physical Activity, Sedentary Behavior, and the Risk of Colon and Rectal Cancer in the NIH-AARP Diet and Health Study.

Exercise as often as possible  
as intensely as possible.

## Hot Topics: Vitamin D

- Research and Review of RDAs
  - Current RDA: adults 50-70 200 IU; >70 400IU
- Colon cancer risk reduced with adequacy
- Emerging evidence suggests protection against cancer, metabolic syndrome, diabetes, HTN, MS, and others
- Questions remain...
  - How much? For whom?
  - Consequences of high Vitamin D status?

## Vitamin D: Where do we go from here?



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