


CANCER REHABILITATION

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Co-morbidities

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Adiposity


- Reeves et al. (2007):
 - Strong evidence that overweight individuals have greater chance of developing cancer and reduced survival;
 - Direct correlation between BMI and endometrial, post-menopausal breast cancer, kidney, myeloma, pancreatic, ovarian and colorectal cancer.
- Many studies show increased survival with increased weight-loss following therapy.

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Body Mass

- Dignam et al. (2006):
 - 4,310 breast & bowel patients between 1989-94;
 - Those underweight (BMI < 19) and obese (BMI > 35);
 - Worse survival rates than healthy weight individuals;
 - More relapse and non-cancer related death.
- Gross et al. (2006):
 - 1069 prostate patients 1994-2002;
 - obese men had a higher risk of early disease recurrence.

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Men

- Obese men were shown to be 33% more likely to die of cancer compared to those of a normal weight and obese women had a staggering 55% increased risk of dying from cancer.

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Mechanisms of being overweight

- ↑ Oestradiol promotes tumour growth in ovarian, endometrial and breast cancers;
- ↑ Leptin promotes proliferation, reduces apoptosis and stickiness of cancer cells.;
- Correlation between Leptin and IGF-1 and the protein Cyclo-oxidase 2 (COX-2);
- ↓ Progesterone has defensive effect on cancer cells;
- Dietary fat and fibre: low fat and high fibre = ↑ oestrogen excretion.

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Insulin and IGF-1

- Higher in overweight;
- Higher in sedentary;
- Promotes proliferation, impairs apoptosis, promotes metastasis.

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Weight gain with cancer, why?

- Chemotherapy encourages snacking;
- Steroids given with chemo...
- Fatigue promotes sedentary behaviour;
- Hormone therapies (tamoxifen, aromatase inhibitors) encourage fat storage and are taken for prolonged periods;
- Cancer-related attitude.

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Diabetes mellitus

- Marble (1934):
 - Of 10000 diabetics 256 developed cancer;
 - Average age of diabetes at diagnosis was 55.8, average age on death 62.9 (7 years);
 - Average survival after cancer diagnosis 1.8 years.
- *More of the surviving subjects expected to contract malignant disease.*

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Hyperinsulinemia and Diabetes mellitus

- Vigneri et al. (2009):
 - Hyperinsulinemia affects IGF-1 mechanisms promoting proliferation and metastasis;
 - Risk of several cancers increased (pancreas, liver, breast, colorectal, urinary tract, and female reproductive organs) in diabetic patients.
- Factors include:
 - diabetes duration, varying levels of metabolic control, different drugs used for therapy, the possible presence of chronic complications, obesity, hyperglycaemia, and increased oxidative stress

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Diabetes medication

- Bowker et al. (2011):
 - Cancer mortality over 5.4 years: 4.9%, Sulphonylureas; 3.5% Metformin; 5.8% insulin.
- *“Patients with type 2 diabetes exposed to sulphonylureas and exogenous insulin had a significantly increased risk of cancer-related mortality compared with patients exposed to metformin”.*

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Cardiovascular disease

- *“Elderly breast cancer patients with a history of cardiac disease and/or diabetes treated with trastuzumab (Herceptin) have an increased incidence of cardiotoxicity. Continuous cardiac monitoring is especially advised in this population” (Serrano et al, 2011).*
- 33% ↑ CV disease
- 33.3% ↑ diabetes

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CV disease

- Yeh et al. (2004):
 - Cardiotoxicity during or following anthracycline therapy (chemo) had many drug-related risk factors, but also presence of CV disease and ischaemia.

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CV disease continued...

- Berthe et al. (2006):
 - “We showed that classical risk factors for cardiovascular diseases, except hypertension, increased the incidence of CVDs”....
 - “Patients with newly diagnosed H-Lymphoma and survivors of HL, especially when treated at young ages, should strongly be advised to refrain from smoking, to maintain a healthy body weight, and to exercise regularly”.

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Summary of mechanisms

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graph LR; COX-2 --> Binds[Binds to cancer cells]; COX-2 --> Prolif[Increases prolif + metastasis]; Binds --- NSAID[Asprin + NSAID]; Prolif --- NSAID; Insulin[Insulin and glucose] --> IGF1[Increases IGF-1 activity]; IGF1 --- Diet[Diet and exercise]; FAT --> Oestradiol[Increases Oestradiol]; FAT --> Leptin[Increases leptin]; FAT --> Progesterone[Reduces Progesterone]; Oestradiol --- Diet; Leptin --- Diet; Progesterone --- Diet;
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Summary of mechanisms

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graph LR; Treatments[Herceptin/Anthracyclines] --> Cardio[Cardio-toxicity/CHF]; Treatments --> CVD; Cardio --- Screen[Screen history, Monitor, Diet, light/mod exercise]; CVD --- Screen;
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