# Cancer Biology Lecture No. 3

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<u>Lecture's Structure:</u>

Definition of Epidemiology Cancer is a global problem Epidemiology identifies causes of cancer Epidemiological Data for some types of cancer Role Of Various Factors In The Development Of Cancer

**Risk Factors** 

**Epidemiology** is the study of the patterns, causes, and effects of health and disease conditions in defined populations.

It is the cornerstone of public health, and informs policy decisions by identifying risk factors for disease and targets for preventive medicine.

#### **Cancer Is a Global Problem**

The incidence and mortality rates for various cancers are similar, though not identical, among developed countries.

In the developing world, as countries become more westernized and their populations achieve longer life expectancy, cancer rates are increasing.

#### **Cancer Is a Global Problem**





#### Epidemiology identifies the etiology of cancer

Regional differences in the distribution of various cancers in different regions of the world reflect differing etiologic factors.

#### **Examples:**

Schistosomiasis infections and bladder cancer in parts of Africa.

Hepatitis B infections in liver cancer in China

Epidemiological data for some types of cancer

Lung Cancer and Cigarett Smoking



# **Epidemiology of Cancer** Epidemiological data for some types of cancer Breast Cancer

Over 210,000 new cases of invasive breast cancer and over 40,000 deaths.

Although breast cancer is more common in developed Western societies, incidence rates are increasing in the developing world. **Epidemiology of Cancer** Epidemiological data for some types of cancer Breast Cancer risk factors

Duration of exposure to female hormones

Dietary and low physical activity factors (obesity; high-fat diet).

Ionizing radiation during breast development.

Chronic use of hormone replacement therapy.

Genetic inheritance (family history) of breast cancer such as brca1, brca2, or p53 germline mutations.

Epidemiological data for some types of cancer

#### **Colorectal Cancer**

Although cancers of the colon and rectum are relatively rare in developing countries, they are the second-most frequent malignancies in the developed world.

This discrepancy appears to be largely due to the conditions of an affluent lifestyle, because the major risk factors are a diet abundant in fat, refined carbohydrates, and animal protein and low in fiber, combined with physical inactivity.

### **Epidemiology of Cancer** Role Of Various Factors In The Development Of Cancer

Incidence rates of migrants changes to match the new location (food intake habits and life styles).

Environmental and lifestyle factors play the predominant role in cancer causation.

Thus a high percentage of cancers are preventable, or at least ''delayable.''

### **Epidemiology of Cancer** Role Of Various Factors In The Development Of Cancer

*Table 3–5.* Estimated Percentage of Total Cancer Deaths Attributable to Established Causes of Cancer

Risk Factor	Percentage
Tobacco	30
Adult diet and obesity	30
Sedentary lifestyle	5
Occupational factors	5
Family history of cancer	5
Viruses and other biologic agents	5
Perinatal factors and growth	5
Reproductive factors	3
Alcohol	3
Socioeconomic status	3
Environmental pollution	2
Ionizing and ultraviolet radiation	2
Prescription drugs and medical procedures	1
Salt, other food additives, and contaminants	1

#### Risk factors

Cigarette Smoking (Lung Cancer) Alcohol (oral and esophageal cancers, liver cancers) Note: Alcohol appears to be synergistic with tobacco in causing cancers of the mouth, pharynx, larynx, and esophagus, but not that of the lung.

#### Mechanisms

(1) a carcinogenic effect of other chemicals such as Nnitrosamines in alcoholic

Beverages.

- (2) a solvent action that facilitates absorption of carcinogens found in tobacco smoke.
- (3) a carcinogenic effect due to acetaldehyde,
- a major metabolite of ethanol.

#### Risk factors

**Diet** Second leading risk factor after tobacco.

Correlation of obesity with the increased incidence of various cancers such as esophagus, colorectum, breast,

endometrium, and kidney.

Differences in cancer rates among various countries

Cancer	Dietary and Diet-Related Risk Factors	Dietary Protective Factors	Other Major Risk Factors
Oral cavity, pharynx and esophagus	Alcohol Very hot drinks Obesity (adenocarcinoma of the oesophagus) Chinese-style salted fish (nasopharyngeal cancer)	Probably fruit and vegetables	Smoking
Stomach	Probably high intake of salt-preserved foods and salt	Probably fruit and vegetables	Infection by Helicobacter pylori
Colorectum	Obesity Possible red and processed meat	Probably fruit and vegetables and other plant foods rich in fiber	Sedentary lifestyle
Liver	High alcohol intake Foods contaminated with aflatoxins	None established	Hepatitis viruses
Pancreas	None established	None established	Smoking
Larynx	Alcohol	None established	Smoking
Lung	None established	Possibly fruit & vegetables	Smoking
Breast	Obesity after menopause Alcohol	None established	Reproductive and hormonal factors
Endometrium	Obesity	None established	Low parity
Cervix	None established	None established	Human papillomavirus
Prostate	None established	None established	None established
Kidney	Obesity	None established	None established

Table 3–7. Dietary Risk Factors, Dietary Protective Factors, and Other Major Risk Factors for Common Cancers

From Key et al.,<sup>68</sup> with permission.

#### **Risk factors**

Sexual Development, Reproductive Patterns, and Sexual Behavior The duration of hormonal exposure appears to play a role in the susceptibility to breast cancer in women.

These factors include early age of menarche, delayed age of first pregnancy, and delayed menopause, suggesting longer duration of exposure to hormonal stimulation as an etiologic agent in breast cancer.

#### **Risk factors**

Industrial Chemicals and Occupational Cancers

Industrial levels of 4-aminobiphenyl have a higher incidence of bladder cancer

Occupational exposure to asbestos fibers results in a higher incidence of lung cancer, mesotheliomas, gastrointestinal tract cancers, and laryngeal cancers.

Leukemia in workers exposed to benzene

#### **Risk factors**

Herbicides

Air and Water Pollutants

Radiation

sunlight as a cause of skin Cancer

Evidences:

1. Skin cancer occurs primarily on exposed areas.

- 2. Skin cancer is relatively rare in dark skinned races in whom skin pigment filters out UV radiation.
- 3. Skin cancer frequency and the intensity of solar radiation are related.

4. Skin cancer can be induced in laboratory animals by repeated exposure to UV radiation.

5. The inability to repair DNA damaged by UV radiation is associated with skin cancer

# Epidemiology of Cancer Risk factors

**Ionizing Radiation** increased incidence of leukemia among radiologists was recognized.

Chernobyl disaster

Epidemiology of Cancer Risk factors

Infection Cancer is not an infectious disease Infection with certain viruses probably acts in concert with other carcinogenic agents or processes.

**Epidemiology of Cancer Risk factors GENETIC FACTORS IN CANCER** Somatic mutations Germline mutations Inherited Cancers (1% to 2%, of total cancers) The probability that an individual carrying the retinoblastoma gene will develop a tumor is about 95% and an average of three to four tumors occur in such a gene carrier.

Syndrome	Gene	Location	Cancer Site and Cancer Type
Familial retinoblastoma	RB1	13q14	Retinoblastoma, osteosarcoma
Multiple endocrine neoplasia II	RET	10q11	Medullary thyroid carcinoma, pheochromocytoma
Multiple endocrine neoplasia I	MEN1	11q13	Adrenal, pancreatic islet cells
Neurofibromatosis type I	NF1	17q11	Neurofibromas, optic gliomas, pheochromocytoma
Neurofibromatosis type II	NF2	22q2	Bilateral acoustic neuromas, meningiomas, cerebral astrocytomas
Bloom syndrome	BLM	15q26	Leukemia, lymphoma
Familial adenomatous polyposis	APC	$5q\overline{2}1$	Colorectal, thyroid
Von Hippel-Lindau	VHL	3p25	Renal cell carcinoma, pheochromocytoma
Familial Wilm's tumor	WT1	11q	Wilms tumor (kidney)
Xeroderma pigmentosum	XP(A–D)	9q,3p,19q,15p	Basal cell carcinoma, squamous cell carcinoma, melanoma (skin)
Fanconi anemia	FAC	16q, 9q, 3p	Acute leukemia
Li-Fraumeni syndrome	p53	17p13	Breast and andrenocortical carcinomas, bone and soft-tissue sarcomas, brain tumors, leukemia
Cowden syndrome	PTEN	10q22	Breast, thyroid
Gorlin syndrome	PTCH	9q31	Basal cell carcinoma
X-linked proliferative disorder	XLP	Xq25	Lymphoma
Peutz-Jeghers syndrome	LKB1	19p	Breast, colon
Ataxi telangiectasia	ATM	11q22	Leukemia, lymphoma

#### Table 3–9. Inherited Cancer Syndromes Caused by a Single Genetic Defect\*

### *Table 3–10.* High-risk Susceptibility Genes and Their Chromosomal Location\*

Gene	Location	Associated Tumors
BRCA1	17q	Breast, ovary, colon, prostate
BRCA2	13q	Breast, ovary, pancreas, prostate
p16 INK4A	9p	Melanoma, pancreas
CDK4	6q	Melanoma, other tumors (rarely)
hMLH1	Зр	Colorectal, endometrial, ovarian cancer
hMSH2	2р	Colorectal, endometrial, ovarian cancer
hMSH6	2р	Colorectal, endometrial, ovarian cancer
PMS1	2q	Colorectal cancer, other tumors (rarely)
PMS2	7p	Colorectal cancer, other tumors (rarely)
HPC2	17p	Prostate (rarely)

# QUESTIONS