

Dr. Stanley Ho Medical Development Foundation Symposium



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- Top 10 Cancers in Hong Kong
- Trend & Risk Factors of Cancers
- Updates of Diagnostics & Therapeutics
- Personalized Medicine to Precision Medicine
- Prevention & Early Detection of Cancers
- Conclusion

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Hong Kong Cancer Registry

- The Hong Kong Cancer Registry (HKCaR) is a population-based cancer registry
- collecting the basic demographic data, information of the cancer site, and histology of all cancer patients from both the public and private medical institutions in Hong Kong.

About Us	• FA	• Statistics	 Publ 	ication	 Link 	s • Contact Us	1	
Statistics	Тор	ſen Cancers i	n 2013	(Download ful	ll details <u>her</u>	2 3 1)		
•Top Ten Cancers		Incidence	e			Mortality	1	
•Cancer Facts	Ma Rank	le Female Site	Both No.	Rel. Freq.	Ma Rank	le Female Site	Both No.	Rel, Freq.
•Cancer Statistics Query System(CanSQS)	1 2 3 4	Colorectum Lung Breast Liver	4,769 4,631 3,544 1,852	16.5% 16.0% 12.2% 6.4%	1 2 3 4	Lung Colorectum Liver Stomach	3,867 1,981 1,542 625	28.5% 14.6% 11.2% 4.6%
All Ages	6	Stomach	1,655	5.7% 3.8%	5 6 7	Breast Pancreas	584	4.4%
And a second sec	7	skin	997	3.4%	8	Non-Hodgkin	372	2.6%
	8	Corpus uteri	942	3.3%	9	Oesophagus	329	2.4%
(covers 30 years of data)	9	Non-Hodgkin lymphoma	877	3.0%	10	Nasopharynx	312	2.3%
Children and Adolescents	10	Nasopharynx All sites	841 28,936	2.9% 100%		All sites	13,589	100%

Cancer Facts

A brief overview of the incidence and mortality of cancer in Hong Kong according to types of cancer.



• Local cancer burden continues to rise, reaching a new record of 28,936 in 2013



Hong Kong Cancer Registry 2013⁵



 These 5 leading cancers comprised of nearly 60% of all new cancers diagnosed in Hong Kong.

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Major Findings from 2012...

 Among the top 5 cancer sites, prostate cancer had advanced 3 places compared to 2002. Number of new cases of lung, colorectal, breast and prostate cancers saw a substantial rise largely due to ageing and growing population, while that of liver cancer showed a relatively small increase.





Major Findings from 2013...

- Colorectal cancer has reclaimed the top spot in the cancer incidence ranking, after overtaking lung for the first time in history by in 2011, but the difference was only around 0.5%.
- Compared to the previous year, most of the increase was attributed to the growing number of cancers in women, particularly in cancers of the lung, corpus uteri and breast.

Hong Kong Cancer Registry 2013⁹



Major Findings since 2012...

In 2012, cancers of corpus uteri and cervix showed a marked increase in incidences, by 18.2% to 810 cases and 16.9% to 457 cases respectively.

A modest increase of 2.1% in ovarian cancers was also observed

Ranked No. 8 in 2013 942 new cases











2011年女性乳腺癌按年齡組別及類型之分佈 **Distribution of Female Breast Cancer** by Age Group & Type in 2011

Distribution of Fe 2011 年初年期目前	male B 及感激:	reast C 子和 [0]	Cancer (?)(Ca ir	by Ag	e Grou L入前的	ip and (Invasia	Type (e)] 泥 e	Jewasin Birit (Ja G	ve and 南守	Ca in-	situ) in	20 11				Inv	asive 818/	BC a 3419 :	ged >=65" = 23.9%
Age (yrs) Irille(st)	0-	5-	10-	15-	20-	25-	30-	35-	40-	45-	50-	55-	60-	65-	70-	75-	80-	85+	All ages 形形化曲
Invasive 人役性	0	0	0	0	3	22	68	173	360	620	524	434	397	225	170	154	123	146	3,419
Cain-situ III (Cain-	σ	0	0	1	0	t	9	26	66	103	64	69	42	30	22	16	7		484
Total URID	0	0	0	1		25	77	199	436	723	608	503	435	265	192	170	130	154	3 903

Age-specific Incidence rates of Female Breest Cancer (per 100,000 women) by Type (Invasive and Ca in-situ) in 2011 2011 年初毎期分類 (原位(Ca in-situ)法人信性(Invasive)]現 通行各年齢近別目前年(初十第名支性人口注意)

Age (ym) 印刷(訊)	0-	5.	10-	15-	20-	25	30-	35-	40-	45-	50-	55-	60-	65-	70-	75-	-90-	85+	Lifetime risk* 一生武权知道*	Crude rate 租發病率	ASR (World)* 印創標準(日 (世界)**
Invasive 人授性	0.0	0.0	0.0	0.0	1.3	7.2	21.0	52.7	109.1	174.0	161.3	167.4	194.1	199.5	149.5	142.3	146.8	172.4	1 in 17	90.7	61.0
Ca in-situ 哥位朝	0.0	0.0	0.0	0.5	0.0	0.3	2.8	7.9	20.0	28.9	25.9	26.6	20.5	26.6	19.3	14.8	8.4	9.4	1 in 112	12.8	8.8
Total 總數	0.0	0.0	0.0	0.5	1.3	7.6	23.8	60.7	1,29.1	202.9	187.1	194.1	214.7	226.1	168.9	157.1	155.1	181.8	1 in 15	103.6	69.9

Cumulative Metme mix betters the age of 75 - mit statistics F4 alls Rates are standardized to the age distillation of the WHO 2007 World Standard Population. Companisons with these rates from other sources are valid only if they use the same standard population for calculations in the the University of American Research and Rese Research and Research a

Source: Hong Kong Cancer Registry, Hospital Authority 資料來源:醫院管理局許達感性資料統計中心

Nov 2013

年齡超過75歲之乳腺癌患者: Invasive BC aged >= 75: 423/3419 = 12.37%

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Combating cancer with CANCER!

- C-<u>C</u>onfirm the diagnosis (Histology)
- A-<u>A</u>ssess the disease status (Staging)
- N-Identify unmet <u>Needs</u> (physical, Psycho-social)
- C-<u>C</u>ombined Expertise (MDT approach)
- E-Evaluate the disease (serial scans...)
- R-<u>R</u>elive (survivors) or <u>R</u>ecurrence



Genetic mutations predisposing to cancer

- Oncogenes (activation)
- Tumor suppressor genes (deletion)
- DNA repair genes (deletion)





- Cancer is clonal in nature
- All cancer arises from genetic mutation
- Multi-step process leading to cancer formation
 multifactorial
- Genetic hereditary vs sponataneous
- Environmental 2/3







Risk Factors

- Unhealthy lifestyle
- Environmental factors...















Breakthroughs in Cancer

- Screening & early detection
- Diagnosis
- Treatment
 - Surgery
 - Radiotherapy
 - Medical Treatment chemotherapy, hormonal therapy & targeted therapy
 - Personalized Treatment



Understanding Treatment Guidelines

- Treatment Guidelines
 - Put extensive evidence-based data into best possible clinical practice
 - To enhance quality of care, to prevent undertreatment, over-treatment and wrong treatment.
 - Compliance with guidelines improve patient outcome

- e.g. St. Gallen recommendation in breast cancer



- With the exception of a few solid tumors and hematological malignancies, chemotherapy alone does not cure cancer
- Chemotherapy is mainly employed in combination with primary treatment (surgery or radiotherapy), i.e. as an adjunctive Rx, or as palliative Rx for recurrent and metastatic disease



Neoadjuvant Therapy

- Make inoperable primary tumour become operable
- Down-size, down-stage tumour
- Control occult metastasis
- Assess tumour response

e.g. aim for breast-conserving therapy in breast cancer





Palliative Therapy

- Relieve symptoms
- Control disease

Neuropathy

Improves quality of life of patients





Phlebitis



The Era of Molecular Targeted Therapy





Molecular Targeted Therapies





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- Drugs that block the growth & spread of cancer cells
- Interfere specific molecules involved in carcinogenesis
 - Process by which normal cells become cancer cells, leading to tumor growth



Molecular Targeted Therapy

- Scientists call these molecules, "molecular targets"
- Molecular Targeted Drugs / Therapy
 - Focusing on the molecules and cellular changes
- Not chemotherapy
- More effective
- Less harmful to normal cells









BREAST CANCER





COLORECTAL CANCER





LUNG CANCER





LIVER CANCER (HEPATOCELLULAR CARCINOMA)





Nasopharyngeal Carcinoma (NPC)





Adoptive T-cell Immunotherapy for NPC







Epidermal Growth Factor Receptor Pathway Map	
Pegradation Recycling	
Small GTPases	
MAPK Cascade	
Transcription Control of Control	

K. Oda et al., Mol. Syst. Biol. 1:8-24 (2005)

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Breast cancer was initially thought as a strictly *local* disease...

Based on William Stewart Halsted (1852-1922)'s theory: Breast cancer is a strictly local disease, only curable by radical surgery...

Early breast cancer used to be managed exclusively by surgeons...





Breast cancer was then thought as a *systemic* disease...

The work of **Bernard Fisher**, breast surgeon & chairman of the National Surgical Adjuvant Breast & Bowel Project (NSABP) in the 1960s has led to the development of breast cancer being a systemic disease...

Leading to a paradigm shift in breast cancer management with the introduction of adjuvant therapy...



Personalized treatment of breast cancer started in the 1960s...

Professor Elwood V. Jensen, first identified the Estrogen receptor (ER) in 1958...

Introduction of **TAMOXIFEN** as a first "targeted agent" in the 1970s...





Success Story of identifying Anti-HER-2 in the 1990s...HER2型乳癌個人化治療





MBC : metastatic breast cancer; MoAb : monoclonal antibody

Immuno-oncology Anti-PD1 and Anti-PDL1 Monoclonal Antibodies







 Breast cancer is a heterogeneous disease comprised of different molecular subtypes based on gene/protein expression





Changing Portraits of Breast Cancer 乳癌的個人化治療





claudin low Lum A Lum B Basal Her2



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Sparano JA, Fazzari M, Kenny PA. Clinical application of gene expression profiling in breasts cancer. Sur Oncol Clin N Am. 2010;19:581-606







Individualized Test Categories

Drug Selection	HER2 (Anti-HER-2 targeted agents) ER+ (Tamoxifen/Aromatase Inhibitors)
Drug Dosage	• Not yet available
Drug Efficacy	• Not yet available
Disease Status	• Not yet available
Recurrence Risk	Oncotype DX / Mammaprint Multivariate analysis uPA/PAI-1 (node negative)
Predisposition	• BRCA-1/2

















Source: ONS



高齡化人士最主要之恐懼為健康衰退 Greatest fear in old age is poor health

- 大概三分之一之高齡人士因身體及腦部不健康
 而引致「殘弱」
- In one-third of older people, unhealthy body and brain results in frailty
- 癌症 (Cancer)
- 中風 (Stroke)
- 認知障礙症 (Dementia)
- 心臟病 (Heart disease)
- 呼吸系統疾病 (Lung disease)



Geriatric Oncology - Population Perspective

- Urban vs Rural Areas
 - Lung, CRC and Breast vs Lung, Stomach, Esophagus
- Different ethnic groups
 - Stomach cancer in N. Chinese vs NPC in S. Chinese
- Liver cancer (HBV-related HCC) and NPC
- National Central Cancer Registry
 - Made up from different local registries







大多數高齡人士都身體健壯 Many older people are physically fit





大多數高齡人士都在 家庭中擔當重要角色 <u>Many have</u> a vital family role





大多數高齡人士 都持續進修學習 Many continue to learn





大多數高齡人士都 享受良好的生活質素 Many have a good quality of life





幫助高齡人士更加強壯 Helping older people become stronger

- · 適量運動及均衡營養有助高齡人士身心 靈更加壯健以接受醫學治療
- Exercise and nutrition can make older people stronger for medical treatments





一般醫學治療目的 Aim of Medical Treatment

- 促進身體及心靈健康 Better physical and psychological well-being
- 提升生活質素 Better good quality of life
- 不論年齡 No matter what age





Why screening

Two major / unique objectives:

- To <u>detect</u> (asymptomatic/early) cancers before they metastasize, so that surgical cure is possible
 - for most common types of cancers
- 2. To <u>prevent</u> cancer by detection and removal of premalignant adenomas
 - for CRC and cervical cancer



Why screening

- Cancer screening aims to detect cancers when an individual does not present with any symptoms.
- Ideally, these cancer screening tests are used to detect cancers at an early stage or even prevent cancers from developing.
- Yet, not all tests are proven to be effective in decreasing the mortality rate of cancers, some results may even be misleading, and taking some of the tests may possess risks.



Why screening

 Therefore there are a lot to consider before deciding whether to receive a screening test or not.

Research has been done to find out the overall effectiveness and worthiness of various kinds of cancer screening tests...





Cancer Screening

Cancer	Tests	Target Population
Colorectal	-Annual Faecal Occult Blood Test (FOBT) -Colonoscopy every 10 years	Individuals aged 50-75 with average risk
Cervical	Cervical Pap Smear	Women aged 25-64 who have ever had sexual contacts
Epithelial Ovarian	-Serum CA125 measurement -transvaginal ultrasound	Women with positive family history of ovarian cancer
Liver	-HBV vaccination -Periodic screening by AFP and ultrasound	-General public -Chronic hepatitis carrier
Nasopharyngeal	-EBV serology blood test -nasopharyngoscopy	Aged 30-70 family members of nasopharyngeal cancer patients

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Cancer Screening

Tests with insufficient evidence

Cancer	Test	Target Population
Breast	-Mammography -Routine breast selfexamination	General female population
Prostate	Routine digital rectal examination	Asymptomatic men
Cervical	HPV DNA test	Women aged 25-64 who have ever been sexually active
Lung	Routine screening with low- dose spiral CT	General population



Cancer Screening

Tests NOT recommended

Cancer	Test	Target Population
Prostate	Routine use of PSA, especially for men over 70 years old	Asymptomatic men
Liver	Routine screening with AFP or ultrasound	General population
Lung	Routine screening with chest x-ray or sputum cytology	General population
Stomach	Routine endoscopic examination	General population

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- Cancer is a disease of the DNA
 - Structural alterations in genes (mutations)
 - Changes in gene transcription
 - Abnormal post-transcriptional protein synthesis
 - Proliferation of cells with abnormal growth regulation which invade normal tissues
- At least 90% of cancer results from DNA alterations
 - Upon exposure to environmental carcinogens



Cancer Prevention in the 21st Century

- At least 90% of cancer results from DNA alterations
 - Upon exposure to environmental carcinogens
- Less than 10% of cancer caused by inheritance of mutated "cancer predisposition genes".



- Extraordinary advances in knowledge of
 - Epidemiology
 - Molecular genetics
 - Molecular biology
 - Cellular biology of cancer
- Much contribution to prevention and cure of cancer



Cancer Prevention in the 21st Century

- · Primary and secondary prevention
- Examples:
 - Immunization against hepatoma
 - HPV vaccination for cervical cancer
 - Chemoprevention to reduce high-risk women for breast cancer
 - Genetic testing for mutations
 - Breast cancer predisposition genes BRCA-1, BRCA-2
 - Bilat subcutaneous mastectomy for positive mutant carriers



Cancer Prevention in the 21st Century

- Our lifestyle is the MAJOR CAUSE of the noncommunicable diseases
 - Cardiovascular disease
 - Cancer
 - Chronic respiratory disease
- About 50% cancer is potentially preventable
 - By population-based primary & secondary prevention



Cancer Prevention in the 21st Century

- Identify high-risk group
 - Hereditary or familial background
 - Personal history
- Control risk factors
 - Host
 - Environment



Cancer Prevention in the 21st Century

- Population screening for the common disease with effective treatment in early stage
- Early detection / screening
- Life-style modification
 - Weight control
 - Healthy and balanced diet
 - Healthy psychosocial status





Cancer Prevention in the 21st Century

Primary & secondary prevention the most cost-effective strategies for reducing morbidity and mortality from cancer.







Conclusion

- Cancer is an important public health problem with increasing incidence and impact in the 21st century
- It is a genetic disease with multi-step process and interaction with the environment.
- Common cancers in Hong Kong include breast, colon, lung, liver and NPC, some of which meet the criteria for population screening and early detection of the high-risk groups.



Conclusion

- There has been major breakthroughs and advances in the early detection, diagnosis and treatment of common cancers in the past decade.
- Multidisciplinary approach with personalized treatment is the trend of strategy.
- Primary prevention and public health education is of paramount importance to increase the awareness of this major illness in the community and the healthcare workers.



