

REFERENCES

- Aaltonen LA, Peltomaki P, Leach FS, Sistonen P, Pylkkanen L, Mecklin J-P, et al. Clues to the pathogenesis of familial colorectal cancer. *Science* (Washington DC) 1993; 260: 812-16.
- Abderrahim F and Esteban C. Biology and treatment of nasopharyngeal cancer. In: Abeloff MD, Klastersky J, editors. *Current opinion in oncology*. Philadelphia: Current Science, 1995; 255-63.
- Ah-See KW, Cooke TG, Pickford IR, Soutar D, and Balmain A. An allelotype of squamous carcinoma of the head and neck using microsatellite markers. *Cancer Res* 1994; 54: 1617-21.
- Armstrong RW, Armstrong MJ, Yu MC, and Henderson BE. Salted fish and inhalants as risk factors for nasopharyngeal carcinoma in Malaysian Chinese. *Cancer Res* 1983; 43: 2967-70.
- Barrett JC. Mechanisms of action of known human carcinogens. In: Vainio H, Magee PN, McGregor DB, and McMichael AJ, editors. *Mechanisms of carcinogenesis in risk identification*. New York : Oxford University Press, 1992 : 115-34.
- Benedict WF, Srivatsan ES, Mark C, Banerjee A, Sparkes RS, and Murphree AL. Complete or partial homozygosity of chromosome 13 in primary retinoblastoma. *Cancer Res* 1987; 47: 4189-91.
- Bishop JM. The molecular genetics of cancer. *Science* 1987; 235: 305-11.
- Bos JL, Van Kreijl CF. Genes and gene products that regulate proliferation and differentiation : critical targets in carcinogenesis. In : Vainio H, Magee PN, McGregor DB, and McMichael AJ, editors. *Mechanisms*

- of carcinogenesis in risk identification. New York: Oxford University Press, 1992 :57-85.
- Brauch H, Tory K, Kotler F, Gazdar AF, Pettengill OS, Johnson B, et al. Molecular mapping of deletion sites in the short arm of chromosome 3 in human lung cancer. *Genes Chrom Cancer* 1990; 1: 240-6.
- Bronner CE, and 17 others. Mutation in the DNA mismatch repair gene homologue *hMLH1* is associated with hereditary nonpolyposis colon cancer. *Nature(Lond.)* 1994; 368: 258-61.
- Buetow KH, Murray JC, Iorael JL, et al. Loss of heterozygosity suggests tumour suppressor gene responsible for hepatocellular carcinoma. *Proc Natl Acad USA* 1992; 52: 5368-72.
- Burt RD, Vaughan TL, Nisperos B, Swanson M, Berwick M. A prospective association between the HLA-A2 antigen and nasopharyngeal carcinoma in US whites. *Int J Cancer* 1994; 56: 465-7.
- Cairns P, Tokino K, Eby Y, and Sidransky D. Homozygous deletions of 9p21 in primary human bladder tumours detected by comparative multiplex polymerase chain reaction. *Cancer Res* 1994; 54: 1422-4.
- , Shaw ME, and Knowles MA. Preliminary mapping of the deleted region of chromosome 9 in bladder cancer. *Cancer Res* 1993; 53: 1230-32.
- Callahan R, and Campbell G. Mutations in human breast cancer : an overview. *J Natl Cancer Inst* 1989; 81: 1780-6.
- Carter SL, Negrini M, Baffa R, Gillum DR, Rosenberg AL, Schwartz GF, et al. Loss of heterozygosity at 11q23 in breast cancer. *Cancer Res* 1994; 54: 6270-4.

- Cavenee WK, Dryja TP, Phillips RA, Benedict WF, Godbout R, Gallie BL, et al. Expression of recessive alleles by chromosomal mechanism in retinoblastoma. 1983; 305: 779-84.
- Cawkwell L, Bell SM, Lewis FA, Dixon MF, Taylor GR, and Quirke P. Rapid detection of allele loss in colorectal tumours using microsatellites and fluorescent DNA technology. Br J Cancer 1993; 67: 1262-7.
-, Lewis FA & Quirke P. Frequency of allele loss of DCC, p53, RB1, WTI, NF1, NM23 and APC / MCC in colorectal cancer assayed by fluorescent multiplex polymerase chain reaction. Br J Cancer 1994; 70: 813-8.
- Chan May KM, Huang Dolly WSP, Ho YH, Lee JCK. Detection of Epstein-Barr virus-associated antigen in fine needle aspiration smears from cervical lymph nodes in the diagnosis of nasopharyngeal carcinoma. Acta Cytol 1989; 33: 351-4.
- Chang WY-H, Cairns P, Schoenberg MP, Polascik TJ, and Sidransky D. Novel suppressor loci on chromosome 14q in primary bladder cancer. Cancer Res 1995; 55: 3246-49.
- Cheng JQ, Jhanwar SC, Lu YY, and Testa JR. Homozygous deletions with in 9p21-22 identify a small critical region of chromosomal loss in human malignant mesotheliomas. Cancer Res 1993; 53: 4761-63.
- Chin TL, Chan WY, Chen W, and Shew JH. Nasopharyngeal carcinoma and retinoblastoma gene expression. Labolatory Invest 1992; 67: 56-70.
- Choi PHK, Suen MWM, Huang HP, Lo K-W and Lee JCK. Nasopharyngeal carcinoma: genetic changes, Epstein-Barr virus infection, or both. Cancer 1993; 72: 2873-8.

- Chow WH, McLaughlin JK, Hrubec Z, Nam JM, Blot WJ. Tabacco use and nasopharyngeal carcinoma in a cohort of US veterans. *Int J Cancer* 1993; 55: 59-40.
- Cliby W, Ritland S, Hartmann L, Dodson M, Halling KC, Keeney G, et al. Human epithelial ovarian cancer allelotype. *Cancer Res* 1993; 53: 2393-8.
- Cleton-Jansen A-M, Collins N, Lakhani SR, Weissenbach J, Devilee P, Cornelisse CJ, et al. Loss of heterozygosity in sporadic breast tumours at the BRAC2 locus on chromosome 13q12-13. *Br J Cancer Res* 1995; 72: 1241-4.
- Coleman A, Fountain JW, Nobori T, Olopade OI, Robertson G, Housman DE, et al. Distinct deletions of chromosome 9p associated with melanoma versus glioma, lung cancer, and leukemia. *Cancer Res* 1994; 54: 344-8.
- Decker H-JH, Li FP, Bixenman HA, and Sandberg AA. Chromosome 3 and 12p rearranged in a well-differentiated peritoneal mesothelioma. *Cancer Genet Cytogenet* 1990; 46: 135-7.
- Diaz MO, Rubin CM, Harden A, Ziemin S, Le Beau RA, and Rowley JD. Deletions of interferon genes in acute lymphoblastic leukemia. *N Eng J Med* 1990; 322: 77-82.
- El-Naggar AK, Lee M-S, Wang G, Luna MA, Goepfert H, Batsakis JG. Polymerase chain reaction based restriction fragment length polymorphism analysis of the short arm of chromosome 3 in primary head and neck squamous carcinoma. *Cancer* 1993; 72: 881-6.

- Fearon ER, Vogelstein B. A genetic model for colorectal tumourigenesis. *Cell* 1990; 61: 759-67.
- Ferti-Passantopoulou A, Panani AD, Raptis S. Preferential involvement of 11q23-24 and 11p15 in breast cancer. *Cancer Genet Cytogenet*. 1991; 51: 183-8.
- Field JK, Kiaris H, Howard P, Vaughan ED, Spandidos DA, and Jones AS. Microsatellite instability in squamous cell carcinoma of head and neck. *Br J Cancer* 1995; 71: 1065-9a.
- Foulkes WD, Campbell IG, Stamp GWH, Allan GJ, and Trowsdale J. Loss of heterozygosity and amplification on chromosome 11q in human ovarian carcinoma. *Br J Cancer Res* 1993; 67: 551-9.
- , Ragoussis J, Stamp GWH, Allan GL, and Trowsdale J. Frequent loss of heterozygosity on chromosome 6 in human ovarian carcinoma. *Br j Cancer* 1993; 67: 551-9.
- Freeman JL, McIvor NP, Feinmesser R, Cheung RK, Dosch HM. Epstein-Barr virus and nasopharyngeal carcinoma : bringing molecular genetics strategies to head and neck oncology. *J Otolaryn* 1994; 23: 130-4.
- Fujimori M, Tokino T, Hino O, Kitagawa T, Imamura T, Okamoto E, et al. Allelotype study of primary hepatocellular carcinoma. *Cancer res* 1992; 51: 89-93.
- , Well SA, and Nakamura Y. Fine scale mapping of the gene responsible for multiple endocrine neoplasia type 1 (MEN1) *Am J Hum Gent* 1992 50: 399-403.
- Fujino T, Risinger JI, Collins NK, Liu F-S, Nishii H, Takahashi H, et al. Allelotype of endometrial carcinoma. *Cancer Res* 1994; 54: 4294-8.

- Fults D, Pedone CA, Thomas GA, and White R. Allelotype of human malignant astrocytoma. *Cancer Res* 1990; 0: 5784-9.
- Gatti RA, Bercel I, Boder E, et al. Localization of an ataxia telangiectasia gene to chromosome 11q12-23. *Nature* 1988; 366: 577-80.
- Goddard AD and Solomon E. Genetic aspects of cancer. In : Harris H and Hirschhorn K, editors. *Advances in human genetics*. New York : Plenum Press, 1993: 321-76.
- Gudmundsson J, Johannesson G, Bergthorsson JT, et al. Different tumour types from BRCA2 carriers show wild-type chromosome deletions on 13q12-q13. *Cancer Res* 1995; 55: 4830-2.
- Han HJ, Yanagisawa A, Kato Y, Park JG, and Nakamura Y. Genetic instability in pancreatic cancer and poorly differentiated type of gastric cancer. *Cancer Res* 1993; 53: 5087-9.
- Harris CC. Tumour suppressor genes, multistage carcinogenesis and molecular epidemiology. In: Vainio H, Magee PN, McGregor DB, and McMichael AJ, editors. *Mechanisms of carcinogenesis in risk identification*. New York : Oxford University Press, 1992 : 115-7.
- Henning H. Malignant conversion: the first stage in progression from benign to malignant tumours. In: Slaga TJ, Klein-Szanto AJP, Boutwell RK, Stevenson DE, Spitzer HL, and D'Motto B, editors. *Skin carcinogenesis*. New York: Alan R Liss, 1989: 95-105.
- , Shores R, Wenk ML, Spangler EF, Tarone R, and Yuspa SH. Malignant conversion of mouse skin tumours is increased by tumour initiation and unaffected by tumour promoters. *Nature* 1983; 67-9.

- Hearne CM, Ghosh S, Todd JA. Microsatellites for linkage analysis of genetic traits. *Trends Genet* 1992; 288-94.
- Herbst H, Niedobitek G, Kneba M, et al. High incidence of Epstein-Barr virus genomes in Hodgkin's disease. *Am J Pathol* 1990; 137: 13-8.
- Hibi K, Tkahashi T, Yamakawa K, Ueda R, Sekido Y, Ariyoshi Y, et al. Three distinct regions involved in 3p deletion in human lung cancer. *Oncogene* 1992; 7: 445-9.
- Hildesheim A, West S, De Veyra E, De Guzman MF, Jurado A, Jones G, et al. Herbal medicine use, Epstein-Barr virus, and risk of nasopharyngeal carcinoma. *Cancer Res* 1992; 52: 3048-51.
- Hollywood DP, Barton CM. Oncogenes and tumour suppressor genes. In: Lemonie N, Neoptolemos J, Cooke T, editors. *Cancer : A molecular approach*. Oxford: Blackwell Scientific, 1994: 13-35.
- Hording U, Nielsen HW, Albeck H. Nasopharyngeal carcinoma: histopathological types and association with Epstein-Barr virus. *Oral Oncol Eur J Cancer* 1993; 29B: 137-9.
- Horii A, Han H-J, Shimada M, Yanagisawa A, Kato Y, Ohta H, et al. Frequent replication errors at microsatellite loci in tumours of patients with multiple primary cancers. *Cancer Res* 1994; 54: 3373-5
- Huang DP, Lo K-W, Choi PHK, Ng AYT, Tsao S-Y, Yiu GKC, et al. Loss of heterozygosity on the short arm of chromosome 3 in nasopharyngeal carcinoma. *Cancer Genet Cytogenet* 1991;54; 91-9.
- , Andrew van Hasselt C, et al. A region of homozygous deletion on chromosome 9p21-22 in primary nasopharyngeal

- carcinoma. *Cancer Res* 1994; 54: 4003-6.
- Huang THM, Quesenberry JT, Martin MB, Loy TS, and Diaz-Arias AA. Loss of heterozygosity detected in formalin-fixed, paraffin-embedded tissue of colorectal carcinoma using a microsatellite located within the deleted in colorectal carcinoma gene. *Diagnos Molec Patho* 1993; 2: 90-3.
- Hubert A, Jeannel D, Tuppin P, and De The G. Anthropology and epidemiology: a pluridisciplinary approach of environmental factors of nasopharyngeal carcinoma. In: Turoz T, Pagano JS, Ablashi G, De The G, Lenoir G & Pearson GR, editors. *The associated Epstein-Barr virus and diseases*. John Libbey Eurotext Ltd, 1993: 777-90.
- Ionov Y, Peinado MA, Malkhosyan S, Shibata D, and Perucho M. Ubiquitous somatic mutations in simple repeated sequences reveal a new mechanism for colonic carcinogenesis. *Nature(Lond.)* 1993; 363: 558-61.
- James CD, Collins VP, Allalunis-Turner MJ, and Days RS. Localization of chromosome 9p homologous deletions in glioma cell lines with markers constituting a continuous linkage map. *Cancer res* 1993; 53: 3674-6.
- Kamb A, Gruis N, Feldhaus J, Liu Q, Harshnman K, Tavtigian S, et al. A cell cycle regulator potentially involved in genesis of many tumour types. *Science(Washington DC)* 1994; 264: 436-9.
- Knowles MA, Elder PA, Williamson M, Cairns JP, Shaw ME, and Law MG. Allelotyping of human bladder cancer. *Cancer Res* 1994; 54: 531-8.
- Knudson AG. All in the (cancer) family. *Nature Genetics* 1993; 5: 103-4.

- Knudson AG. Mutation and cancer : statistical study of retionblastoma. Proc Natl Acad Sci USA 1971; 68: 820-3.
- Kohno T, Takayama H, Hamaguchi M, Takano H, Yamaguchi N, Tsuda H, et al. Deletion mapping of chromosome 3p in human uterine cervical cancer. Oncogene 1993; 5: 1825-32.
- Kok K, Osinga J, Carrit B, Davis MB, Van Der Hout AH, Vander Veen AY, et al. Deletion of a DNA sequence at chromosomal region 3p21 in all major types of lung cancer. Nature 1987; 330: 578-81.
- Kovacs G, and Frisch S. Clonal chromosome abnormalities in tumour cells from patients with sporadic renal cell carcinomas. Cancer Res 1989; 49: 651-9.
- Lammie GA and Peter G. Chromosome 11q13 abnormalities in human cancer. Cancer Cells 1991; 3: 414-20.
-, Fante V, Smith R, et al. D11S287, a putative oncogene on chromosome 11q13, is amplified and expressed in squamous cell and mammary carcinomas and linked to BCL-1. Oncogene 1991; 6: 439-44.
- Largey JS, Meltzer SJ, Yin J, Norris K, sauk JJ, and Archibald DW. Loss of heterozygosity of p53 in oral cancers demonstrated by the polymerase chain reaction. Cancer(Phila) 1993; 71: 1933-7.
- Larsson C, Skogseid B, Oberg K, Nakamura Y, and Nordonskjold N. Multiple endocrine neoplasia type1 gene maps to chromosome 11 and is lost in insulinoma. Nature 1988; 332: 85-7.
- Latif F, Fivash M, Glenn G, Tory K, Orcutt M, Hampsch K, et al. Chromosome 3p deletions in head and neck carcinomas: statistical ascertainment

- of allelic loss. *Cancer Res* 1992; 52: 1451-6.
- Leach SF, Nicolaides NC, Papadopoulos N, Liu B, Jen J, Parsons R, et al. Mutations of a mutS homolog in hereditary nonpolyposis colorectal cancer. *Cell* 1993; 75: 1215-25.
- Lee HP, Gourley L, Duffy SW, Esteve J, Lee J, Day NE. Preserved foods and nasopharyngeal carcinoma : a case-control study among Singapore Chinese. *Int J Cancer* 1994; 59: 585-90.
- Levine AJ. The tumour suppressor genes. *Annu Rev Biochem* 1993; 62: 623-51.
- Levine PH, Pocinki AG, Madigan P, Nad Bale S. Familial nasopharyngeal carcinoma in patients who are not Chinese. *Cancer* 1992; 70: 1024-9.
- Litt M and Luty JA. A hypervariable microsatellite revealed by *in vitro* amplification of a dinucleotide repeat within the cardiac muscle actin gene. *Am J Hum Genet* 1989; 44: 397-401.
- Lo K-W, Huang DP, and Lau K-M. p16 gene alterations in nasopharyngeal carcinoma. *Cancer Res* 1995; 55: 2039-43.
- Loeb LA. Microsatellite instability : marker of a mutator phenotype in cancer. *Cancer Res* 1994; 54: 5059-63.
- Lothe RA, Peltomaki P, Meling GI, Aaltonen LA, Nystrom-Lahit M, Pyikkanen L, et al. Genomic instability in colorectal cancer : relationship to clinicopathological variables and family history. *Cancer Res* 1993; 53: 5849-52.
- Louis DN, Deimling AV, and Seinyinger BR. A (CA)n dinucleotide repeat assay for evaluating loss of allelic heterozygosity in archival human brain tumour specimens. *Am J Pathol* 1992; 141: 777-82.

- Lu SJ, Day NE, Degos L, et al. Linkage of nasopharyngeal carcinoma susceptibility locus to the HLA region. *Nature* 1990; 346: 470-1.
- Maestro R, Gasparotto D, Vukosavljevic T, Barzan L, Sulfara SI, Boiocchi M. Three discrete regions of deletion at 3p in head and neck cancers. *Cancer Res* 1993; 53: 5775-9.
- Marshall CJ. Tumour suppressor genes. *Cell* 1991; 64: 313-26.
- Meltzer SJ, Yin J, Manin B, Rhyu M-G, et al. Microsatellite instability occurs frequently and in both diploid and aneuploid cell populations of Barrett's-associated esophageal adenocarcinomas. *Cancer Res* 1994; 54: 3379-82.
- Merlo A, Gabrielson E, Aski F, Baylin S, and Sidransky D. Frequent loss of chromosome 9 in human primary non-small cell lung cancer. *Cancer Res* 1994; 54: 640-2.
- , Mabry M, Vollmer R, Baylin SB and Sidrasky D. Homozygous deletion on chromosome 9p and loss of heterozygosity on 9q, 6p, and 6q in primary human small cell lung cancer. *Cancer Res* 1994; 54: 2322-6.
- Misra BC and Srivatsan ES. Localization of HeLa cell tumour suppressor gene to the long arm of chromosome 11. *Am J Human Genet* 1989; 45: 567-77.
- Mitra AB, Murty VVVS, Li RG, Pratap M, Luthra UK, and Chaganti RSK. Allelotyping analysis of cervical carcinoma. *Cancer Res* 1994; 54: 4481-7.
- Modrich P. DNA mismatch correction. *Annu Rev Genet* 1991; 25: 229-53.

- Mori N, Yokota J, Oshimura M, Cavenee WK, Mizoguchi H, Noguchi M, et al. Concordant deletions of chromosome 3p and loss of heterozygosity for chromosome 13 and 17 in small cell lung carcinoma. *Cancer Res* 1989; 49: 5130-5.
- Mori T, Aoki T, Matsubara T, Iida F, Xi Qun D, et al. Frequent loss of heterozygosity in the region including BRCA1 on chromosome 17q in squamous cell carcinomas of the esophagus. *Cancer Res* 1994; 54: 1638-40.
- Morita R, Ishikawa J, Tsutsumi M, Hikiji K, Tsukada Y, Kamidono S, et al. Allelotype of renal cell carcinoma. *Cancer res* 1991; 51: 820-3.
- Mulligan LM, Mole SE. Strategies for isolating genes in hereditary and sporadic tumours. In: Levine AJ, Schmidek HH, editors. *Molecular genetics of nervous system tumours*. New York: Wiley-Liss, 1993: 195-208.
- Muir C, Waterhouse J, Mack T, Powell J, and Whelan S. Cancer incidence in five continents, Vol 5. IARC, Lyon : IARC Scientific Publication 1987; 88.
- Nakashima H, Inoue H, Mori M, Ueo H, Ikeda M, and Akiyoshi T. Microsatellite instability in Japanese gastric cancer. *Cancer* 1995; 75: 1503-7.
- Nakashima H, Mori M, Mimori K, Inoue H, Shibuta K, et al. Microsatellite instability in Japanese esophageal carcinoma. *Int J Cancer Res* 1995; 64: 286-9.
- Naoki M, Lun Y, Mitsuo O, et al. Concordant deletions of chromosome 3p and loss of heterozygosity for chromosome 13 and 17 in small cell

- lung carcinoma. *Cancer Res* 1989; 49: 5130-5.
- Nawroz H, Van Der Riet P, Hruban RH, Koch W, and Sidrasky D. Allelotyping of head and neck squamous cell carcinoma. *Cancer Res* 1994; 54: 1152-5.
- Naylor SL, Johnson BE, Minna JD, and Sakaguchi AY. Loss of heterozygosity of chromosome 3p markers in small cell lung cancer. *Nature(Lond.)* 1987; 329: 451-4.
- Nicolaides NC, Papadopoulos N, Liu B. Mutation of two PMS homologous in hereditary nonpolyposis colon cancer. *Nature(Lond.)* 1994; 371: 75-80.
- Niedobitek G, Agathanggelou A, Barber P, Smallman LA, Jones EL, and Young LS. p53 overexpression and Epstein-Barr virus infection in undifferentiated and squamous cell nasopharyngeal carcinomas. *J Patho* 1993; 170: 457-61.
- Nobori T, Miura K, Wu DJ, Lois A, Takabayashi K, and Carson DA. Deletions of the cyclin-dependent kinase-4 inhibitor gene in multiple human cancer. *Nature(Lond.)* 1994; 368: 753-6.
- Olopade OI, Buchhagen DL, Malik K, Sherman J, et al. Homozygous loss of interferon genes defines the critical region on 9p that is deleted in lung cancers. *Cancer Res* 1993; 53: 2410-5.
- Osborne RJ, and Leech V. Polymerase chain reaction allelotyping of human ovarian cancer. *Br J Cancer* 1994; 69: 429-38.
- Papadopoulos N, Nicolaides NC, Wei Y-F, Ruben SM, Carter KC, Rosen CA, et al. Mutation of a *mutL* homologous in hereditary colon cancer. *Science(Wash.)* 1994; 263: 1625-9.

- Patel P, O' Rahilly S, Buckle V, Nakamura Y, Turner RC, and Wainscoat JS.
Chromosome 11 allele loss in sporadic insulinoma. Clin Pathol 1990;
43; 377-8.
- Peiffer SL, Herzog TJ, Tribune DJ, Mutch DG, Gersell DJ, and Goodfellow PJ.
Allelic loss of sequences from the long arm of chromosome 10 and
replication errors in endometrial cancers. Cancer Res 1995; 55:
1922-6.
- Peltomaki P, Aaltonen LA, Sistonen P, Pylkkanen L, Mecklin J-P, Jarvinen
H, et al. Microsatellite instability is associated with tumours that
characterize the hereditary nonpolyposis colorectal carcinoma
syndrome. Cancer Res 1993; 53: 5853-5.
- Popescu NC, Chahinian AP, and Dipaola JA. Non random chromosome
alterations in human malignant mesothelioma. Cancer Res 1988; 48:
142-7.
- Rajadurai P, Prasad U, Chandrika G, Sadler R, Flynn K, and Raab-Traub N.
Undifferentiated, Nonkeratinizing, and Squamous cell carcinoma of
the nasopharynx ; variants of Epstein-Barr virus infected neoplasia.
Am J Patho 1995; 146: 1355-67.
- Ragoussic J, Senger G, Trowsdale J, and Campbell I. Genomic organization
of the human folate receptor genes on chromosome 11q13.
Genomics 1992; 14: 423-30.
- Risinger JL, Berchuck A, Kohler MF, Watson P, Lynch HT, and Boyd J.
Genetic instability of microsatellites in endometrial carcinoma. Cancer
Res 1993; 53: 5100-3.

- Rosenberg CL, Kim HG, Shows TB, Kronenberg HM, and Arnold A.
Rearrangement and overexpression of D11S287E, a candidate
oncogene on chromosome 11q13 in benign parathyroid tumours.
Oncogene 1991; 6: 449-53.
- Rowly JD. Molecular cytogenetics: Rosetta stone for understanding cancer
twenty-nineth G.H.A. Clowes Memorial Award lecture. *Cancer Res*
1990; 50: 3816-25.
- Ruley HE. Oncogenes. In: Levine AJ, Schmidke HH, editors. *Molecular
genetics of nervous system tumours*. New York : Wiley-Liss , 1993:
89-100.
- Ryberg D, Lindstedt BA, Zienolddiny S and Haugen A. A hereditary genetic
marker closely associated with microsatellite instability in lung cancer.
Cancer Res 1995; 55: 3996-9.
- Sager R. Tumour suppressor genes : the puzzle and the promise. *Science*
(Washington DC) 1989; 246: 1406-12.
- Sato T, Saito H, Morita R, Koi S, Lee JH & Nakamura Y. Allelotype of human
ovarian cancer. *Cancer Res* 1991; 51: 5118-22.
....., Tanigami A, Yamakawa K, Akiyama F, Kasumi F, Sakamoto G, et al.
Allelotype of breast cancer : cumulative allele losses promote tumor
progression in primary breast cancer. *Cancer Res* 1990; 50: 7184-9.
- Seizinger BR, Rouleau GA, Ozelius LJ, et al. Von Hippel-Lindau disease
maps to the regions of chromosome 3 associated with renal cell
carcinoma. *Nature* 1988; 332: 268-9.
- Serrano M, Hannon GJ, and Beach D. A new regulatory motif in cell-cycle
control causing specific inhibition of cyclin D/CDK4. *Nature(Lond.)*

- 1993; 336: 704-7.
- Simons MJ, Day NE, Wee GB, Shanmugaratnam K, Ho HC, Wong SH, et al. Nasopharyngeal carcinoma V: Immunogenetic studies of Southeast Asian ethnic groups with high and low risk for the tumor. *Cancer Res* 1974; 34: 1192-5.
- Somers KD, Merrick MA, Lopez ME, Incognito LS, Schechter GL, and Casey G. Frequent p53 mutations in head and neck cancer. *Cancer Res* 1992; 52: 5997-6000.
- Srivatsan FS, Misra BC, Venugopalan M, and Wilczynski SP. Loss of heterozygosity for alleles on chromosome 11 in cervical carcinoma. *Am J Hum Genet* 1991a; 49: 868-77.
- Strand M, Prolla TA, Liskay RM, and Petes TD. Destabilization of simple repetitive DNA in yeast by mutations affecting DNA mismatch repair. *Nature(Lond.)* 1993; 365: 274-6.
- Suzuki T, Yokota J, Mugishima H, Okabe I, et al. Frequent loss of heterozygosity on chromosome 14q in neuroblastoma. *Cancer Res* 1989; 49: 1095-8.
- Swift M, Morrell D, Massey RB, and Case CL. Incidence of cancer in 161 familial affected by ataxia telangiectasia. *New Eng J Med* 1991; 325: 1831-6.
- Thibodeau SN, Bren G, and Schaid D. Microsatellite instability in cancer of the proximal colon. *Science(Washi.)* 1993; 260: 816-9.
- Tronick SR, Aaronson SA. Growth factors and signal transduction. In: Mendelsohn J, Howley PM, Israel MA, Liotta LA, editors. *The molecular basis of cancer*. Philadelphia: Saunders, 1995: 117-40.

- Tsuchiya E, Nakamura Y, Weng S-Y, Nakagawa K, Tsuchiya S, Sugano H, et al. Allelotype of non-small cell lung carcinoma comparison between loss of heterozygosity in squamous cell carcinoma and adenocarcinoma. *Cancer Res* 1992; 52: 2478-81.
- Van der Hout AH, Kok K, Van den Berg A, Oosterhuis W, Carritt B, Buys CH. Direct molecular analysis of a deletion of 3p in tumours from patients with sporadic renal cell carcinoma. *Cancer Genet Cytogenet* 1988; 32: 281-5.
- Van der Riet P, Karp D, Farmer E, Wei Q, Grossman L, Tokino K, et al. Progression of basal cell carcinoma through loss of chromosome 9q and inactivation of a single p53 allele. *Cancer res* 1994; 54: 25-7.
- , Nawroz H, Hruban RH, Corio R, et al. Frequent loss of chromosome 9p21-22 early in head and neck cancer progression. *Cancer Res* 1994; 54: 1156-8.
- Varmus HE. The molecular genetics of cellular oncogenes. *Annu Rev Genet* 1984; 18: 553-612.
- Vogelstein B, Fearon ER, Kern SE, Hamilton SR, Preisinger AC, Nakamura Y, et al. Allelotype of colorectal carcinomas. *Science* 1989; 244: 207-11.
- Watanabe M, Imai H, Shiraishi T, Shimazaki J, Kotake T and Yatani R. Microsatellite instability in human prostate cancer. *Br J Cancer* 1995; 72: 562-4.
- Weber JL. Informativeness of human (dC-dA)_n, (dG-dT)_n polymorphisms. *Genomics* 1990; 7: 524-30.
- , and May PE. Abundant class of human DNA polymorphisms

- which can be types using the polymerase chain reaction. Am J Hum Genet 1989; 44: 388-96.
- Weinberg RA. Tumor suppressor genes. Science 1991; 254: 1138-46.
- Weinstein BI. Mitogenesis is only one factor in carcinogenesis. Science 1991; 251: 387-8.
- Weiss LM, Movahed LA, Butler AE, et al. Analysis of lymphoepithelioma and lymphoepithelioma-like carcinomas for Epstein-Barr virus by *in situ* hybridization. Am j Surg Pathol 1989; 13: 625-31.
- Weston A, Willey JC, Modali R, Sugimura H, McDowell EM, Resau J, et al. Differential DNA sequence deletions from chromosome 3, 11, 13, and 17 in squamous-cell carcinoma, and adenocarcinoma of the human lung. Proc Natl Acad Sci USA 1989; 86: 5099-103.
- Yamaguchi T, Toguchida J, Yamamura T, et al. Allelotype analysis in osteosarcomas : frequent allele loss on 3q, 13q, 17p, and 18q. Cancer Res 1992; 52: 2419-23.
- Yee CJ, Roodi N, Verrier CS, and Parl FF. Microsatellite instability and loss of heterozygosity in breast cancer. Cancer Res 1994; 54: 1641-4.
- Yokota J, Tsukada Y, Nakajima T, et al. Loss of heterozygosity on the short arm of chromosome 3 in carcinoma of the uterine cervix. Cancer Res 1989; 49: 3598-601.
- , Wada M, Shimosato Y, Terada M, and Sugimura T. Loss of heterozygosity on chromosome 3, 13, and 17 in small-cell lung carcinoma and on chromosome 3 in adenocarcinoma of the lung. Proc Natl Acad Sci USA 1987; 84: 9252-6.

- Yu MC, Garabrant DH, Huang TB, and Henderson BI. Occupational and other non-dietary risk factors for nasopharyngeal carcinoma in Guangzhou, China. *Int j Cancer* 1990; 45: 1033-9.
- Yuspa SH, Hennings H, Roop D, Strickland J, and Greenhalgh DA. Genes and mechanism involved in malignant conversion. In: Harris CC and Liotta LA, editors. *Genetic mechanisms in carcinogenesis and tumour progression*. New York: Wiley-Liss, 1990: 115-26.
- Zhang S-Y, Klein - Szanto AJP, Sauter ER, et al. Higher frequency of alterations in the p16/CDKN2 gene in squamous cell carcinomas cell lines than in primary tumours of the head and neck. *Cancer Res* 1994; 54: 5050-3.
- Zheng YM, Tuppin P, Hubert A, Jeannel D, Pan YJ, Zeng Y, et al. Environmental and dietary risk factors for nasopharyngeal carcinoma : a case-control study in Zangwu county, Guangxi, China. *Br J Cancer* 1994; 69: 508-14.
- Zur Hausen H, Schulte-Holthausen H, Klein G, et al. EBV DNA in biopsies of Burkitt tumours and anaplastic carcinomas of the nasopharynx. *Nature* 1970; 228: 1056-8.

APPENDIX

APPENDIX A

American Joint Committee for Cancer

Staging System for Nasopharyngeal Carcinoma

Primary Tumor (T)

Tx Primary cannot be assessed

Tis Carcinoma in situ

T0 No evidence of primary tumor

T1 Tumor limited to one subsite of the nasopharynx

T2 Tumor invades more than one subsite of the nasopharynx

T3 Tumor invades nasal cavity and oropharynx or both

T4 Tumor invades skull or cranial nerves or both

Regional Lymph Nodes (N)

Nx Regional lymph nodes cannot be assessed

N0 No regional lymph node metastasis

N1 Metastasis in a single ipsilateral lymph node, 3 cm or less in greatest dimension

N2 Metastasis in a single ipsilateral lymph node, more than 3 cm but not more than 6 cm in greatest dimension; or in bilateral or contralateral lymph nodes, none more than 6 cm in greatest dimension.

N2a Metastasis in a single ipsilateral lymph node more than 3 cm but not more than 6 cm in greatest dimension

N2b Metastasis in multiple ipsilateral lymph nodes, none more than 6 cm in greatest dimension

N2c Metastasis in bilateral or contralateral lymph nodes, none more than 6 cm in greatest dimension

Distant Metastasis (M)

Mx Presense of distant metastasis cannot be assessed

M0 No distant metastasis

M1 Distant metastasis

Stage Grouping

0	Tis	N0	M0
I	T1	N0	M0
II	T2	N0	M0
III	T3	N0	M0
	T1	N1	M0
	T2	N1	M0
	T3	N1	M0
IV	T4	N0	M0
	T4	N1	M0
	Any T	N2	M0
	Any T	N3	M0
	Any T	Any N	M1

APPENDIX B

BUFFERS AND REAGENTS

1. 1M Tris (pH 7.0)

Tris base	121.1	g
dH ₂ O	700	ml.

* adjust the pH to 7.0 by adding conc. HCl

Adjust volume to 1.0 litre with dH₂O, and sterilize by autoclaving.

2. 0.5 M EDTA (pH 8.0)

Disodium ethylenediamine tetraacetate. 2H ₂ O	186.6	g
dH ₂ O	700	ml.

* adjust the pH to 8.0 with conc. NaOH

Adjust volume to 1.0 litre with dH₂O and sterilize by autoclaving.

3. 10M Ammonium acetate (CH₃COONH₄)

Ammonium acetate	770	g
dH ₂ O	700	ml

Adjust the volume to 1.0 litre with dH₂O and sterilize by filtration

4. 10X TBE buffer

Tris base	104	g
Boric acid	55	g
0.5 M EDTA	40	ml.

Adjust the volume to 1.0 litre with dH₂O and filter through Whatman paper no.1

5. 6M NaCl

NaCl	350.64	g
dH ₂ O	600	ml.

Adjust the volume to 1.0 litre with dH₂O and sterilize by autoclaving.

6. 25%Ammonium Persulfate : (NH₄)₂S₂O₈

Ammonium persulfate	2.5	g
add dH ₂ O to	4	ml.
* store at 4°C		

7. Proteinase K (stock solution)

Dissolve Proteinase K	20	mg
dH ₂ O	1	ml.
* store at -20 °C		

8. 10% Sodium dodecyl sulfate (SDS)

SDS (eletrophoresis grade)	100	g
dH ₂ O	870	ml.

*adjust the pH to 7.2 by adding conc. HCl (a few drops)

Adjust the volume to 1.0 litre with dH₂O

9. Digestion buffer

100 mM NaCl
 10 mM Tris-Cl (pH 8.0)
 25 mM EDTA (pH 8.0)
 0.5% SDS
 0.1 mg/ml PK*

* PK is labile and must be added fresh with each use.

10. T₁₀E₁₀

1 M Tris	1	ml
0.5 M EDTA	2	ml
add dH ₂ O to	97	ml

11. T₂₀E₅

1 M Tris	2	ml
0.5 M EDTA	1	ml
add dH ₂ O to	97	ml

12. T₁₀E₁

1 M Tris	1	ml
0.5 M EDTA	0.2	ml
add dH ₂ O to	100	ml

13. Loading dye

0.025% Xylene cyanol FF	2.5	mg
0.025% Bromophenol blue in dH ₂ O	2.5	mg
98% Deionized formamide	9.8	ml
10 mM EDTA	0.2	ml

14. 6% Denaturing gel

Urea (utrapure)	25.2	g
38% acrylamide/ 2% bisacrylamide	9	ml
10 X TBE	6	ml
adjust the volume to 60 ml with dH ₂ O		
* TEMED	60	μl
* 10% (NH ₄) ₂ S ₂ O ₈	0.6	ml

* Must be added fresh just before use.

APPENDIX C

List of STRP markers for alleotyping study

Chromosome arm	Loci	Location	Size(bp)
1P	D1S243	1p36.1-36.2	142-170
1Q	D1S103	1q31-q32	82-102
2P	D2S131	2p	229-247
	D2S405	2p	275
2Q	D2S102	2q33-q37	138-162
3P	D3S1038	3p25VHL	115
	D3S192	3p25VHL	96-118
	D3S1600	3p14	182-198
	D3S966	3p21.3	147
	D3S1480	3p	87-125
	D3S1255	3p25VHL	150
	D3S1217	3p13-14	170
	D3S1481	3p14	120
	D3S1214	3p21.3	170
3Q	GLUT2	3q26.1-q26.3	184-222
	D3S1744	3q23-q24	170
4P	D4S174	4p11-p15	175-195
	D4S1599	4q	142-156
4Q	D4S1554	4q11-q35	184-208
	D4S1625	4q	194
5P	D5S392	5p	83-117
	D5S819	5p	281
5Q	D5S82	5q14-21	169-179
	D5S346	5q21-q22	96-122
6P	D6S309	6p	254-272
	D6S477	6p	238-280
6Q	IGF2R	6q27	158-166
	D6S503	6q	270
	D6S292	6q	141-161
7P	D7S517	7p	239-257
	D7S460	7p	234
7Q	D7S486	7q31	133-146
	D7S821	7q	281
8P	NEFL	8p	137-147
	D8S87	8p21.3-p22	170
	D8S110	8p	281
8Q	D8S88	8q22	76-100

List of STRP markers for alleotyping study (continue)

Chromosome arm	Loci	Location	Size(bp)
8Q	MCC	8q24	87-125
9P	D9S169	9p21	259-275
	D9S165	9p	250
	IFNA	9p22	130
9Q	D9S51	9q	135-159
	D9S290	9q34	148-160
	ABLI	9q34	130
10P	D10S89	10p11.2-pter	140
	D10S249	10p	118-134
10Q	D10S169	10q11.2-qter	99-117
	D10S677	10q	271
11P	WTI	11p13	150
	D11S554	11p	234
11Q	D11S534	11q13	228-244
	D11S956	11q13	247-308
	D11S976	11q23	130
	D11S897	11q23	98-120
	INT2	11q13.3	161-177
12P	D12S341	12p	114-130
	D12S62	12p	194
12Q	MFD133	12q	161-175
	D12S86	12q	124-160
13Q	D13S284	13q13RBI	197-227
	D13S119	13q	197-213
14P	TCRD	14q11.2	118-128
	D14S118	14q	230
15Q	GABRB3	15q11-13	181-201
	D15S131	15q	238-274
	D15S123	15q	191-207
16P	D16S287	16p13.11	201-225
	D16S748	16p13.2-13.13	230
16Q	D16S511	16q22-24	182-222
	D16S539	16q24.2-q24.3	194
17P	D17S520	17p12	130-144
	D17S945	17p13	186-208
	D17S1176	17p	95-109
17Q	KRT9	17q21NMEI	182-198
	NFI	17q11.2	234
18P	D18S59	18pter-p11.22	148-164
18Q	D18S35	18q21.2-q21.3	104-124

List of STRP markers for allelotyping study(continue)

Chromosome arm	Loci	Location	Size(bp)
18Q	DCC	18q21.1	106-160
	D18S535	18q	150
19P	D19S221	19p	191-211
	D19S412	19q	89-113
19Q	D19S246	19q	270
	D20S470	20p	280
20P	D20S27	20p12	128-138
	D20S17	20q12-q13.1	130-140
21Q	D21S258	21q	184-206
	D21S11	21q21	271
22Q	IL2RB	22q	149-163
	D22S446	22q11NF2	260

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