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Circulating Biomarkers in Patients with Head and Neck Cancer and the Influence of Cigarette Smoking

2019-11-19

head and neck cancer hnc is a collective name for heterogeneous tumors located in the head and neck regions for which smoking alcohol and human papillomavirus hpv are documented risk factors the survival of hnc patients has only improved marginally during the last decade the most important prognostic factors are tumor size local spread and distant metastases tumor node metastasis tnm staging prognostic biomarkers are needed as a complement to tnm staging the aim for this thesis was to investigate rapid and low cost blood based biomarkers which could indicate the risk of hnc recurrence of the disease or the survival of hnc patients furthermore the aim was to examine how cigarette smoking influences the levels of biomarkers in paper i a possible role of plasma cytokines or proteins associated with immune response or inflammation as biomarkers for the survival of hnc patients was investigated higher levels of c reactive protein crp and tumor necrosis factor alpha tnf \alpha were detected in plasma of the patients compared with the levels in the controls the elevated levels of these two biomarkers detected in patients were associated with decreased survival in paper ii the influence of 45 single nucleotide polymorphisms snps located in 41 genes associated with cell cycle progression cell death dna repair or immune response on cancer risk tumor recurrence and survival in hnc patients were investigated snps in immune response genes were associated with risk for hnc an elevated risk for recurrence and a decreased survival in hnc patients in paper iii the influence of cigarette smoking on levels of inflammatory cells proteins or cytokines chemokines micrornas mirnas and snps was analysed in healthy smokers and non smokers higher levels of total white blood cells wbcs neutrophils monocytes lymphocytes neutrophil to lymphocyte ratio nlr crp monocyte chemoattractant protein 1 mcp 1 and interferon gamma ifn y were detected in smokers compared to non smokers and indicate an inflammatory response also a lower level of oncomirna mir 21was detected in smokers this alteration in combination with the elevated levels of ifn y in smokers could be a protective response to cigarette smoke the higher levels of ifn y in smokers compared to non smokers were however only detected in individuals with snp rs2069705 genotype ag gg this indicates a genetic association of the levels of ifn y in paper iv the separate effects of cigarette smoking and hnc on inflammatory or immune biomarkers and the impact of high risk human papillomavirus age and gender were investigated comparisons of circulating levels of wbcs and its subpopulations plasma proteins or cytokines chemokines between smoking and non smoking patients smoking and non smoking controls and between the patient and control groups were analysed smoking had highest impact on elevated levels of wbcs ifn y and mcp 1 and hnc had highest impact on elevated levels of neutrophils monocytes nlr crp macrophage inflammatory protein 1 beta and tnf α in conclusion host immune response associated parameters could be suitable as biomarkers for the risk of hnc risk of recurrence or in predicting survival of hnc patients this thesis show that hnc are associated with systemic inflammatory response and upregulated crp and tnf α is related to shorter survival in hnc patients additionally snps in immune response genes such as rs1800629 in the tnf α gene indicates a risk for hnc or an elevated risk for recurrence and a decreased survival in hnc patients these rapid and low cost blood based biomarkers could be used in combination or as a supplement to established biomarkers in the clinic for a more personalized treatment modality huvud och halscancer hh cancer innefattar tumörer belägna i huvud och halsområdet tobaksrökning ökar risken dramatiskt för olika sjukdomar knappt hälften av rökande patienter dör i de av rökning orsakade sjukdomarna cancer är orsaken till en tredjedel av de rökrelaterade dödsfallen av vilka hh cancer är en en andel av patienterna med hh cancer dör på grund av att tumören varit för stor redan vid diagnos eller att tumören har spridit sig till övriga delar av kroppen men för många patienter är det mycket oklart vad som bestämmer behandlingsresultatet tumörerna är till synes lika och behandlingen standardiserad målet med denna doktorsavhandling var att undersöka billiga och lättillgängliga biologiska markörer som kan indikera risk för att drabbas av hh cancer

eller om dessa markörer kan förutspå behandlingsresultat och överlevnad hos de drabbade patienterna dessutom undersöktes hur cigarettrökning påverkade nivåerna av markörerna i studie i undersöktes om molekyler i blodet biomarkörer förknippade med immunförsvaret kunde förutsäga överlevnaden hos hh cancerpatienter i jämförelse med friska individer sågs högre nivåer av molekylerna tnf α och crp hos patienterna och dessa förhöjningar var relaterade till förkortad överlevnad hos patienterna i studie ii var målet att undersöka om variationer i gener förknippade med immunförsvaret celldelning celldöd eller enzymer som reparerar skadat dna kunde påverka risk och prognos för hh cancer resultatet visade framför allt att små ärftliga variationer i gener som reglerar immunförsvaret kunde påverkade risk för hh cancer risk för återfall i sjukdomen samt överlevnaden hos patienterna i studie iii jämfördes inflammatoriska och immunförknippade biomarkörer som kunde påverkas av cigarettrökning mellan friska rökare och friska icke rökare rökarna hade en högre inflammatorisk aktivitet med högre nivåer av totalt antal vita blodkroppar och tre av dess olika undergrupper neutrofiler monocyter och lymfocyter samt av biomarköerna crp mcp 1 och ifn y de funna lägre nivåerna av den cancerförknippade biomarkören mir 21 och högre nivåer av den förmodat skyddande biomarkören ifn y hos rökarna kan vara ett uttryck för kroppens försvar mot den cancerframkallade cigarettröken Ärftliga faktorer tycks kunna påverka de högre nivåerna av ifn y hos rökarna eftersom ökningen endast fanns i en grupp individer med viss typ av genetisk uppsättning eftersom både rökning och hh cancer ger upphov till inflammation undersöktes i studie iv hur dessa var för sig påverkade nivåerna av inflammatoriska biomarkörer detta för en bättre förståelse hur immunförsvaret reagerar på rökning och hh cancer jämförelser av inflammatoriska markörer från rökande och icke rökande patienter och rökande och ickerökande friska individer genomfördes rökning hade störst påverkan på de högre nivåerna av totalt antal vita blodkroppar och signalmolekylerna mcp 1 och ifn y hh cancer hade störst påverkan på högre nivåerna av neutrofiler monocyter kvoten mellan neutrofiler och lymfocyter crp mip 1b och tnf α uppkomsten av hh cancer behandlingsresultat och överlevnad bland patienterna kan antas inte bara bero på tumörens egenskaper utan även på värdfaktorer hos patienten dessa kan vara ärftliga eller bero på reglering av gener eller tumörens omgivning av t ex immunceller och inflammatoriska molekyler och hur dessa samverkar med miljöfaktorer som tobaksrökning i denna avhandling presenteras biomarkörer som kan bidra med information om risk och prognos för hh cancer samt hur tobaksrökning påverkar dessa markörer

Indicators and Test-papers

1904

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Proceedings of the American Academy of Arts and Sciences

1893

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what is the difference between an academic and professional qualification who should get a professional qualification did you know that some professions can not be legally practised with a degree alone why get a uk qualification is it expensive to gain a british qualification what is a chartered institute or society and is it better than a non chartered body what is the difference between a professional body and a trade union these are all questions answered in this book which is designed to help individuals choose a career path and the right professional organisation in today s world it isn t enough to have a qualification you need to be able to meet with peers and use the valuable networks that are already in place to foster your professional qualification provides a comprehensive survey of the qualifications available in the uk along with guidance on where they lead entry requirements where to apply and where to study derived from the vast and authoritative british qualifications database this important publication provides the first easily accessible guide to qualifications and how to get them in the uk built around a comprehensive directory of professional qualifying bodies each professional area is described in depth and its qualifications identified and explained the book is supported by a simple website which ensures purchasers of the book are kept up to speed with new developments

British Qualifications

2004

the kemp and young series provides a general introduction to a number of subject areas in a style that will be ideally suited for those wishing to learn more the concise presentation of the subject matter is made possible by the reduction of the work to its simplest terms this is achieved through the omission of unnecessary mathematics or mathematical concepts and the generous use of diagrams and illustrations rapid reference to the substance of each topic can be made by use of the carefully constructed index the third edition of ship stability notes and examples has been updated by dr c b barrass who has wide experience in both industry and the academic field the book has been thoroughly revised and expanded to be more in line with current examinations and now covers topics such as ship squat angle of heel whilst turning and moments of inertia via simpson s rules also included is a diagram showing deadweight moment ship stability notes and examples is an invaluable tool to aid in the passing of maritime examinations updated volume of the popular kemp and young series for the new millennium 66 fully worked examples with a further 50 giving final answers

Journal of the Royal Institute of Chemistry

1956

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1973

nanostructures for oral medicine presents an up to date examination of the applications and effects of nanostructured materials in oral medicine with each chapter addressing recent developments specific applications and uses of nanostructures in the oral administration of therapeutic agents in dentistry the book also includes coverage of the biocompatibility of nanobiomaterials and their remarkable potential in improving human health and in reducing environmental pollution emerging advances such as dr franklin tay s concept of a new

nanotechnology process of growing extremely small mineral rich crystals and guiding them into the demineralized gaps between collagen fibers to prevent the aging and degradation of resin dentin bonding is also discussed this work will be of great value to those who work in oral medicine providing them with a resource to gain a greater understanding of how nanotechnology can help them create more efficient cost effective products in addition it will be of great interest to those who work in materials science who wish to gain a greater appreciation of how nanostructured materials are applied in this field outlines the major uses of nanostructured materials for oral medicine including the properties of each material discussed and how it should best be applied explores how nanostructured materials enable the creation of more effective drug delivery systems in oral medicine discusses how novel uses of nanostructured materials may be applied in oral medicine to create more effective devices

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1956

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2002

tides a primer for deck officers and officer of the watch exams prepares the reader for the officer of the watch and master mate certificates required by all officers on commercial seagoing vessels from the formation of tides and tidal stream data right through to practice questions with answers and even mock exam papers this book will provide you with all the reference material you need in order to pass your exams

Ship Stability: Notes and Examples

2000-12-12

includes annual report

Successful ICT Projects in Excel

2002

this book is designed to help students on an a level advanced vce or similar course to design and implement a site using microsoft frontpage 2002 frontpage is a suitable software package for the project component in as level and advanced vce courses parts 1 to 2 of the book take the reader through the steps in creating and publishing a site both with and without the use of a wizard part 3 gives advice on all stages of project work from the definition of a suitable problem through to documentation the book was written primarily for as and a level information and communication technology students and contains in appendix b the aga mark scheme for as module 3 coursework it will also be suitable for students on many

other courses at different levels since the mark scheme with minor variations is one which applies to projects in many ict courses for a second year project in an a level course data collected from a site can be exported and used in an access database this aspect could be developed further to fulfil the requirements of for example module 6 of the aqa ict specification a sample project is included to show students how a complete project report may be laid out

British Welding Journal

1964

terrestrial navigation a primer for deck officers and officer of the watch exams prepares the reader for the officer of the watch and master mate certificates required by all officers on commercial seagoing vessels revision and self test guide to all navigation related elements contained within the officer of the watch exams are included top tips are highlighted throughout the book the case studies and checklists have been designed to add context and aid recall from basic trigonometry and plane sailings plotting right through to practice questions with answers and mock exam papers this book will provide you with all the reference material you need to pass your exams

British Education Index

1975

it is not unusual for patients with head and neck cancer hnc to suffer from both tumor and treatment related pain that is difficult to alleviate despite individualized pharmacological treatment the presence of concomitant various dimensions of pain in patients during the often difficult period following radiotherapy rt has not been elucidated several aspects concerning the importance of relatives for hnc patients have been addressed however little attention has been given to how relatives perceive patients with hnc pain and it is important to further elucidate this sparsely studied topic knowledge about health related quality of life hr gol in this patient group during early rt is limited and needs to be assessed in relation to diagnosis and treatment self care sc refers to what patients do on their own to achieve maintain and promote optimal health and may help reduce pain for several pain conditions the impact of patient education and sc on pain and other common hnc symptoms need further clarification the aims of this thesis were to describe experiences and perceptions of pain in patients with hnc shortly after rtto describe how relatives perceived the patient's situation especially concerning pain and how they experienced their own situation to identify factors that impact hr gol during early rtto develop effective pain management strategies maintain activities of daily living and promote hr gol in patients with hnc undergoing rt using patient education and sc instruction in paper i patients with hnc described existential pain expressed as fear of death meaninglessness and guilt already during and shortly after rt physical pain psychological distress and social withdrawal played a significant role patients with hnc who were treated with rt should also proactively be offered treatment for the various dimensions of pain in paper ii relatives described their mental stress in response to a challenging situation that required their active support to help ease the patient s difficult condition the interviews with relatives also revealed a lack of personal knowledge and frustration over the inability to participate in patient care as well as inadequate support from the healthcare system early interventions from the healthcare system on behalf of the relatives may be necessary to meet these needs in paper iii regression models revealed that pain intensity and symptoms of depression adversely affected hr qol in patients with hnc during early rt customized prehabilitation programs aimed at preventing pain and symptoms of depression could help preserve good hr gol paper iv assessed individual patient education and sc initiatives that resulted in a tendency for lower pain intensity during a portion of rt one way to potentially enhance the benefits of education and sc could be to improve for example patient motivation and self efficacy as well as to optimize supportive efforts from caregivers in conclusion by identifying factors that can impact hr gol and evaluating the effect of patient education and sc this thesis contributes to knowledge on perceived pain and the patient s situation during and shortly after rt in patients with hnc and their relatives this thesis points to the need for evaluation and further development of patient education and effective sc

strategies for pain in patients with hnc undergoing rt as well as for development and evaluation of support strategies for patients with pain and their families during and after rt

Paper

1978

a comprehensive annually updated guide to higher education offering practical advice on courses and places to study the book deals with the mechanics of applying to college and also information on matters from finance and accommodation to a glossary of unfamiliar terms

The Test of War

1988

timed to coincide with the icc cricket world cup 2003 in south africa this book begins with an account of the 2003 final in johannesburg edward griffiths then goes back to the beginning the genesis of the one day game with the launch of the gillette cup in 1963 and traces the development of the game over four decades there are some accounts of the first and subsequent cricket world cup tournaments which highlight the changes in the game over the years heroic performances triumphs and defeats

Journal of the Institute of Fuel

1960

the field of professional academic and vocational qualifications is ever changing the new edition of this highly successful and practical guide provides thorough information on all developments fully indexed it includes details on all university awards and over 200 career fields their professional and accrediting bodies levels of membership and qualifications it acts as an one stop guide for careers advisors students and parents and will also enable human resource managers to verify the qualifications of potential employees

Nanostructures for Oral Medicine

2017-04-11

plant engineers are responsible for a wide range of industrial activities and may work in any industry this means that breadth of knowledge required by such professionals is so wide that previous books addressing plant engineering have either been limited to only certain subjects or cursory in their treatment of topics the plant engineering handbook offers comprehensive coverage of an enormous range of subjects which are of vital interest to the plant engineer and anyone connected with industrial operations or maintenance this handbook is packed with indispensable information from defining just what a plant engineer actually does through selection of a suitable site for a factory and provision of basic facilities including boilers electrical systems water hvac systems pumping systems and floors and finishes to issues such as lubrication corrosion energy conservation maintenance and materials handling as well as environmental considerations insurance matters and financial concerns one of the major features of this volume is its comprehensive treatment of the maintenance

management function in addition to chapters which outline the operation of the various plant equipment there is specialist advice on how to get the most out of that equipment and its operators this will enable the reader to reap the rewards of more efficient operations more effective employee contributions and in turn more profitable performance from the plant and the business to which it contributes the editor keith mobley and the team of expert contributors have practiced at the highest levels in leading corporations across the usa europe and the rest of the world produced in association with plant engineering magazine this book will be a source of information for plant engineers in any industry worldwide a flagship reference work for the plant engineering series provides comprehensive coverage on an enormous range of subjects vital to plant and industrial engineer includes an international perspective including dual units and regulations

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