



Research Annual Report 2011



Other research

Hospital pharmacy and clinical pharmacology

The allergic immune response and related allergic reactions

Determinants of falls and disability in older persons

Neurotraumatology and intracranial hydrodynamics

The urinary tract: Physics, function, epidemiology and reconstructive surgery

Pathophysiology and treatment of asthma, atopy and cystic fibrosis

Applied social and forensic psychiatry

Biology and technology of oral and maxillofacial surgery

Diagnosis and treatment of disorders of the posterior segment and strabismus

Otorhinolaryngology and head & neck surgery

Addiction research

Philosophy of Medicine

Palliative and supportive care

Surgery, non-programme associated

Pediatrics, non-programme associated

EMC OR-01-34-01 - Hospital pharmacy and clinical pharmacology

Programme (brief description)

Pharmacokinetics describe how the organism handles a drug, pharmacodynamics deal with the effects of the drug on the organism. Pharmacogenetics studies the genetic factors that contribute to variability in both pharmacokinetics and pharmacodynamics.

In our research programme we are active in all three topics. At present the focus of our activities is in the following areas of interest:

1. Transplantation. For several immunosuppressive drugs therapeutic drug monitoring is routinely applied. Pharmacogenetic studies aim to identify genes causing interindividual variability in drug exposure, population pharmacokinetic studies focus on non-genetic factors. Studies are ongoing in kidney, liver and heart transplantation. In kidney transplantation a multinational randomized trial investigating the value of therapeutic drug monitoring for mycophenolate mofetil is coordinated by our center. In 2005 a new PK/PD study was initiated, looking at IMPDH monitoring.

2. Oncology. Population pharmacokinetic studies are performed to identify patient factors that explain interpatient variability in exposure to anticancer drugs (e.g. docetaxel).

3. Psychiatry. Several anti-depressive and anti-psychotic drugs are critical dose drugs, for which dose adjustment based on drug levels is applied. In several clinical trials PK/PD analysis is part of the outcome parameters. In 2005 a prospective trial was started, to evaluate the impact of genotyping CYP2D6 before initiating imipramine therapy.

4. Pediatrics. Limited sampling strategies are developed to allow pharmacokinetic studies in neonates. Methods for determination of renal function in pediatric patients are evaluated.

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
internal medicine	0,1				0,1								
pharmacy	2,7	1			3,7								
Total	2,8	1			3,8	0	0	0	0	42	11	11	64

Article/Letter to the editor

Akker, C.H.P. van den, Schierbeek, H., Minderman, G, Vermes, A., Schoonderwaldt, E.M., Duvekot, J.J., Steegers, E.A.P. & Goudoever, J.B. van (2011). Amino Acid Metabolism in the Human Fetus at Term: Leucine, Valine, and Methionine Kinetics. *Pediatric Research*, 70(6), 566-571.

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Part of book - abstract

Bemt, P.M.L.A. van den (2011). Veiliger gebruik van orale oncolytica in de dagelijkse praktijk. In E.J. van Hoogdalem & K.H. Hoogendoorn (Eds.), *Kanker, over leven; nieuwe inzichten in de oncologie*. Houten: Anselmus Colloquim.

EMC OR-01-39-07 - The allergic immune response and related allergic reactions

Programme (brief description)

The program is focused on the allergic immune response, the pathophysiological backgrounds of allergic disorders and the different forms of intervention. The induction of the allergic reaction by allergens is the basis of research to characterization of allergens, the relation between inhalant and food allergics and the modulation of the allergic reactions. Apart from classical pharmacotherapy and environmental control immunomodulation of the allergic response is the most promising form of intervention.

The main components of research comprise:

Occupational allergy. Erasmus MC has a unique location in a heavily industrialize environment, providing an excellent opportunity for research in occupational allergy. This is reflected by research to allergy in horticulture and health professionals. Patient care in occupational allergy is organized in a multidisciplinary initiative instituted by the section of Allergology in collaboration with the departments of Dermatology, Pulmonology, ENT and Public health. There is a close connection between this initiative and our research. Currently, two large longitudinal studies among bell pepper and health care workers have been finalised. The focus of research is on the incidence of occupational disease and the effects of occupational disease on sick leave, work performance, costs and health related quality of life. The results of these studies will be published as PhD thesis in the beginning of 2012.

Food allergy. Research focuses on the optimization of diagnostic procedures.

Research objectives are closely related with patient care initiatives. In particular, the new initiative "Kinderhaven" - collaboration between Allergology, Paediatric Pulmonology and Paediatric Dermatology provides an environment for research to this topic. Several studies from the section of Allergology focus on the discrepancy between sensitization and clinical allergy. A large study to the positioning of MRA's (mediator release assays) and other tools in the diagnosis of cashew nut allergy will be financially supported by STW/NWO. The study will start in 2012.

Immunomodulation. The section Allergology is taking part in large scale RCT's to the effectiveness of sublingual immunotherapy. There is collaboration with the dept. of General Practice. A study to the efficacy of sublingual immunotherapy in children allergic to house dust mites has recently been finalised. Another immunotherapy study in children with a grass pollen allergy will result in a PhD thesis in the beginning of 2012. A large cost- effectiveness study with regard to allergen immunotherapy among adults has been started in 2009. The section of Allergology is taking the lead in this multi-centre study financially supported by ZonMW. The study is still ongoing.

The section co-operates with the dept. of Parasitology, LUMC to the role of regulatory B-cells in subcutaneous immunotherapy in asthmatics. At the Erasmus MC the section is collaborating with the dept. of Immunology in a study to the role of IgE bearing B-cells in treatment of patients with either anti-IgE or immunotherapy. With prof. Barbara Bohle (department of Pathophysiology, Medical University of Vienna) the section is studying the clinical and immunological effects of inhalant immunotherapy on cross-reactive food allergy. Currently, the section is involved in new studies to the use of anti-IgE and epicutaneous immunotherapy in patients with a peanut allergy.

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
family practice		0,7			0,7								
internal medicine	0,5				0,5								
Total	0,5	0,7			1,2	0	0	0	0	9	3	6	18

Article/Letter to the editor

Al Hadithy, A.F.Y., Maaren, M van & Vermes, A. (2011). Anaphylactic reactions following Kenacort-A (R) injection: carboxymethylcellulose is involved once again. *Contact Dermatitis*, 64(3), 179-180.

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Part of book - abstract

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Book Editor

Gerth van Wijk, R. (Ed.). (2011). *Het kinderallergie formularium 2e editie*. Houten: Bohn Stafleu van Loghum.

EMC OR-01-39-08 - Determinants of falls and disability in older persons

Programme (brief description)

- A - Determinants of falls
- B - Drugs: effects and side effects, drug-related hospital admission, drug-related falls.
- C - Vascular ageing.
- D - Dementia - Co-morbidity in Alzheimer's disease
- E - Delirium - Determinants and prognosis
- F - Transmural projects with Stichting 'De StroomOpmaatGroep' in Rotterdam, Stichting Elisabeth in Breda and with the Department of Industrial Design of Delft Technical University

Key Figures

Department	Sp1	Sp2	Sp3	Sp4	SpTot	T1	T2	T3	Ttot	IS	FS	OP	Ptot
internal medicine	1,3				1,3								
Total	1,3				1,3	2	0	0	2	14	3	1	20

Thesis

Hartholt, K.A. (2011, september 21). *Falls and drugs in the older population: medical and societal consequences*. EUR. Prom./coprom.: Dr. T.J.M. van der Cammen & Prof. Dr. P. Patka.

Sterke, C.S. (2011, november 25). *A study of balance, gait and psychotropic drug use in relation to fall risk in nursing home residents with dementia*. EUR. Prom./coprom.: Prof. Dr. Ir. A. Burdorf & Dr. T.J.M. van der Cammen.

Article/Letter to the editor

Adams, D., Jonge, R. de, Cammen, T.J.M. van der, Zietse, R. & Hoorn, E.J. (2011). Acute kidney injury in patients presenting with hyponatremia. *Journal of Nephrology*, 24(6), 749-755.

Blommers, E., Klimek, M., Hartholt, K.A., Cammen, T.J.M. van der, Klein, J. & Noordzij, P.G. (2011). Perioperative care of the older patient. *Zeitschrift für Gerontologie und Geriatrie*, 44(3), 187-191.

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Book Editor

Mattace Raso, F.U.S. (Ed.). (2011). *Guidelines Cardiovascular Riskmanagement*. Den Haag: CBO.

EMC OR-01-45-01 - Neurotraumatology and intracranial hydrodynamics

Programme (brief description)

The program is devoted to experimental and clinical research in the field of traumatic brain injury (TBI) and hydrocephalus. The program on TBI focuses on three themes: 1) Trial methodology, guidelines and evidence based medicine, 2) Intensive care monitoring/therapy and neuroprotection, 3) Ethics and outcome. The program on intracranial hydrodynamics mainly concentrates on normal pressure hydrocephalus.

Head injury is the most common cause of death and disability in young adults in the Western world. As such it constitutes a major health and socio-economic problem. The initial trauma initiates complex pathophysiologic mechanisms, leading to secondary brain damage. Basic research has elucidated many of these mechanisms and has resulted in the development of promising neuroprotective agents. Various of these agents are being tested in our in vitro model of experimental TBI. We have participated in the clinical evaluation of neuroprotective agents. This translational research has proved complex, and none of the randomised controlled trials on neuroprotective agents with demonstrated efficacy in the experimental situation, has shown convincing benefit in the overall population of TBI. Problems in clinical trial design and analysis, specific to the field of TBI may have contributed to this failure. Methodologic challenges are related to the heterogeneity of the population, outcome assessment, detection of treatment effects, the use of surrogate markers and interim analyses. These observations have prompted a significant research effort, organised as an international collaboration based on extensive analyses of data available from studies and trials in TBI encompassing approximately 10.000 patients.

International recognition of the importance of this study was illustrated by the receipt of a NIH RO1 grant for the period 2003-2006.

In the clinical situation cerebral ischemia is considered the central factor in the development of secondary damage. Experimental and clinical programs focus on techniques for monitoring cerebral oxygenation and metabolism. Previous work has demonstrated the value of clinical monitoring of local brain oxygen tension in patients with TBI, and the current research focuses on possibilities to target therapy towards cerebral oxygenation

The third theme of the TBI program focuses on outcome assessment, including evaluation of cognitive functioning, and the development and evaluation of a disease specific scale for quality of life assessment in TBI. Ethical considerations are relevant to neuro-intensive care in general and particularly to the field of neuro-traumatology. The main focus of ethical research and evaluation concentrates on informed consent procedures in acutely incapacitated patients following TBI.

Normal pressure hydrocephalus is a treatable disease common to elderly patients with symptomatology often akin to Alzheimer's disease or vascular dementia. In this theme specific attention is paid to clinical diagnostic criteria and to shunt hydrodynamics. The department of neurosurgery currently co-ordinates a national multi-center study evaluating the use of a programmable shunt device for treatment of normal pressure hydrocephalus.

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
neurosurgery	4,81				4,81								
Total	4,81				4,81	0	0	0	0	6	0	1	7

Article/Letter to the editor

Andriessen, TMJC, Horn, J, Franschman, G, Naalt, J van der, Haitsma, I., Jacobs, B.C., Steyerberg, E.W. & Vos, P.E. (2011). Epidemiology, Severity Classification, and Outcome of Moderate and Severe Traumatic Brain Injury: A Prospective Multicenter Study. *Journal of Neurotrauma*, 28(10), 2019-2031.

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Zheng, P., Romme, E., Spek, P.J. van der, Dirven, C.M.F., Willemsen, R. & Kros, J.M. (2011). Defect of development of ocular vasculature in Glut1/SLC2A1 knockdown in vivo. *Cell Cycle*, 10(11), 1871-1872.

EMC OR-01-49-02 - The urinary tract: Physics, function, epidemiology and reconstructive surgery

Programme (brief description)

The program concentrates on physical and (neuro)physiological modelling of the urinary tract. Models are developed on the basis of measurements in single smooth muscle cells, muscle fibres and complete organ systems in vitro. Passive and active mechanical properties, electrophysiological properties and excitation-contraction coupling are described in this way. The innervation and function of the organ systems are studied in situ in guinea pigs and rats using morphological and neurophysiological methods. The models are applied for the development of diagnostic and therapeutic methods in a clinical setting. Epidemiological studies such as 'A longitudinal non-invasive study of changes in urinary bladder contractility secondary to benign prostatic hyperplasia' provide normative data that are important for modelling and clinical analysis as well. In this way function and dysfunction are described quantitatively and new treatment modalities are evaluated objectively. Automated analysis of clinical data and non-invasive diagnostic methods are important issues in the diagnostic field. In the therapeutic field, emphasis is placed on the development and evaluation of neural stimulation techniques to modulate organ (dys)function based on neurophysiologic principles and on stemcell therapy of incontinence.

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
radiology	0,3				0,3								
urology	1,7		0,7		2,4								
Total	2		0,7		2,7	0	0	0	0	1	0	0	1

Article/Letter to the editor

Groen, J., Blok, B.F.M. & Bosch, J.L.H.R. (2011). Sacral Neuromodulation as Treatment for Refractory Idiopathic Urge Urinary Incontinence: 5-Year Results of a Longitudinal Study in 60 Women. *Journal of Urology*, 186(3), 954-959.

EMC OR-01-54-02 - Pathophysiology and treatment of asthma, atopy and cystic fibrosis

Programme (brief description)

This programme aims (1) to elucidate the etiology and pathogenesis of childhood asthma and asthma-like lung disease and allergy and determining the effects of chronic inflammation on the development and function of the respiratory system and (2) to determine the effects of therapeutic and preventive interventions in chronic inflammatory lung disease and atopy in children.

Chronic lung disease, especially asthma, is one of the leading causes of chronic morbidity in children. Childhood asthma and other obstructive lung diseases predispose for obstructive lung disease in adults, an important health care issue with great socioeconomic impact. The search for causal and contributing factors, development of new diagnostic techniques and the determination of optimal treatment modalities for childhood asthma and allergy has the potential to improve morbidity and prognosis, and thus to reduce the burden of these diseases. In this respect, the development and validation of new techniques to measure lung function and airway inflammation in infants and young children has priority.

Infant lung function testing and new image analysis techniques using CT scan are developed as sensitive, objective endpoints for assessment of airway inflammation and damage. The development of structural lung damage is studied in a cohort of CF patients using CT and extensive lung function testing.

Inhalation of aerosolized medication is an important aspect of the treatment of lung disease, and poses specific problems in the pediatric age group. In this project, we examine the factors that determine lung deposition and effects of inhaled drugs in young children to optimize the effects of treatment.

The therapeutic and preventive intervention studies carried out within this project are important for specific patient groups (asthma, cystic fibrosis), and have impact for the treatment of childhood obstructive lung disease in primary and secondary health care practice.

The development of allergic diseases with age is studied in two large birth cohorts (Prevention and Incidence of Asthma and Mite Allergy, "PIAMA", and "Generation R"). These focus on interaction of genetic and environmental factors in relation to immunopathology and clinical phenotypes.

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
gastroenterology & hepatology	0,4				0,4								
neurosciences	0,11				0,11								
psychiatry	5,98	3,31		8,5	17,79								
Total	6,49	3,31		8,5	18,3	1	0	0	1	35	0	4	40

Thesis

Hol, J. *Mucosal immune regulation in early infancy: monitoring and intervention*. EUR (167 pag.) (Enschede: Ipskamp Drukkers BV). Prom./coprom.: Prof. Dr. J.C. de Jongste, E.E.S. Nieuwenhuis, Dr. J.N. Samsom & E.H.G. van Leer.

Article/Letter to the editor

Bakker, E.M., Volpi, S., Salonini, E., Wiel - Kooij, E.C. van der, Sintnicolaas, C.J.J.C.M., Hop, W.C.J., Assael, B.M., Merkus, P.J.F.M. & Tiddens, H.A.W.M. (2011). Improved treatment response to dornase alfa in cystic fibrosis patients using controlled inhalation. *European Respiratory Journal*, 38, 1335.

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Part of book - abstract

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Editor board / refereeship

Vrijlandt, E.J.L.E. & Jongste, J.C. de (Eds.). (2009). *Tijdschrift voor Kindergeneeskunde*, 77(6).

EMC OR-01-58-01 - Applied social and forensic psychiatry

Programme (brief description)

Project A: Neurobiology of mood disorders

Main participants: Prof. Dr. S.A. Kushner, Prof. Dr. W.J.G. Hoogendijk, Prof. Dr. W.M.A. Verhoeven, Prof. Dr. C.L. Mulder

Programme description: Despite remarkable treatment advances over the past century, mood disorders remain among the most severe and intractable mental illnesses. Undoubtedly, the incomplete clinical repertoire for the diagnosis, treatment and prevention of mood disorders results from our historically limited understanding of the relevant neurobiology. However, recent advances in brain imaging, epidemiology, and genetics have laid a strong foundation for a detailed neurobiology. Accordingly, our research programme comprises both highly focused clinical studies and translational neurobiology. The four core elements of our programme include the translational neurobiological research laboratory of Steven Kushner, the O3 mental health network of Professor Niels Mulder, the Women's Mental Health program and a prospective biobank. Importantly, a key strategy in our efforts is the synergy between scientific research (knowledge generation), education (knowledge transfer) and patient care activities (knowledge application). Further, our research focus is also aligned with the core strengths of our clinical department, and regards neurobiological studies of mood disorders with an emphasis on serious, treatment-resistant and peripartum uni- and bipolar depression. This sharp focus is being coordinated with the highly successful epidemiological cohorts of the Erasmus MC (Generation R, ERGO) and work closely with strong (pre) clinical departments such as Neuroscience, Immunology, Obstetrics, Anaesthesia and Genetics. Importantly, our programme is also synergistic within 3 major domains of the Erasmus MC academic component: weesziekten, obesity and "klaar voor een kind". Together, these elements are mutually reinforcing and contribute to the central focus of our research programme.

Using an endophenotype-based approach in collaboration with the department of Neuroscience, we use state-of-the-art methods for temporally and spatially-restricted manipulations of gene expression and electrophysiological activity within defined neural circuits to uncover the causal links between neurobiology and behavior. In particular, we have focused our efforts on the translational neurobiology of social behavior and emotional learning. Thus far, we have demonstrated a novel mechanism by which neurons are selected for participation in a memory network based upon a competitive selection process driven by neuronal excitability. Further, the GEZIN studie (GEnetische Zoektocht In Neuropsychiatrische stoornissen) utilizes patient-derived neurons to investigate the molecular and cellular mechanisms of severe mental illness, in close collaboration with the department of Clinical Genetics and the Erasmus MC Stem Cell Institute. To avoid heterogeneity caused by the wide range of clinical phenomenology, we have focused our efforts on families with a high incidence of severe mental illness, identified through a large collaboration with psychiatric institutions throughout The Netherlands. Next-generation deep sequencing methods are being applied to identify the specific genetic variants underlying the psychiatric disturbance in each individual family. In parallel, patient-specific induced pluripotent stem cell (iPS) cultures provide the highly unique opportunity to study neural cell cultures derived from each individual participant. For the first time in the field of psychiatry, we can now study the neural development and physiology of cells that are entirely derived from patients, to develop neurobiologically-inspired treatments for severe mental illness.

Project B: Applied social psychiatry

Main participants: Prof. Dr. C.L. Mulder, Prof. Dr. H.J.C. van Marle, Prof. Dr. W.J.G. Hoogendijk

Programme description: This research programme comprises applied social and forensic psychiatry. Since 2007, the Research Center O3 (a collaboration between mental health organisations in the Rijnmond region, the municipal health center and the department of Psychiatry) carries out studies in the field of public mental health. The main aim of the research center is to study difficult to engage populations with severe mental illness in the public domain as well as in the mental health services domain. This includes studies related to public mental health issues such as social breakdown and self neglect, and help seeking behaviour (using the Rijnmond Case Register data), effects of different organisational models for assertive outreach, interventions to increase compliance with treatment, the epidemiology and prevention of involuntary admissions and coercive measures in psychiatry and

victimisation of psychiatric patients. Populations studied include the general population, and outpatient in inpatient patients with severe mental illness. A strong focus is on developing and testing new interventions, especially for difficult to engage patients with severe mental illness.

Within the forensic psychiatric research, determinants of violent behaviour, persistent delinquency in adolescents, treatment effects, and the epidemiology of criminality in combination with mental health care are studied. Diagnosis, treatment and risk profiling are the main areas of research in forensic psychiatry. The group has two main themes in this field, the development of criminal careers for juvenile and adult offenders and psychiatric determinants of violent behaviour, with special interest for the translation of empirical research findings into clinical practice. By means of a multidisciplinary approach, psychotic offenders, antisocial personality disordered patients and psychopaths are discriminated regarding the start of their offending behaviour related to the onset of their mental disorder and regarding psychiatric precursors of violence. Results should lead to a better indication and programming of forensic psychiatric treatment. Special attention is given to the complexity of sexual offending, as a phenomenon limited to adolescence, persistent adult offending with or without other non-sexual offenses, and the specific influence of cognitive distortions on sexual offending. The focus of our research group is on how this subtype differs from other subtypes within sex offenders with respect to psychiatric disorders. Further studies are directed at the role of attachment problems and personality disorders in domestic violence offenders.

Key Figures

Department	Sp1	Sp2	Sp3	Sp4	SpTot	T1	T2	T3	Ttot	IS	FS	OP	Ptot
gastroenterology & hepatology	0,4				0,4								
neurosciences	0,11				0,11								
psychiatry	5,98	3,31		8,5	17,79								
Total	6,49	3,31		8,5	18,3	1	0	0	1	51	10	19	81

Thesis

Mies, G.W. (2011, december 06). *Performance monitoring in depression. Behavioural and neurophysiological correlates of feedback processing*. EUR (172 pag.). Prom./coprom.: Prof. Dr. M.W. Hengeveld, M.W. van der Molen & Dr. F.M. van der Veen.

Article/Letter to the editor

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EMC OR-01-59-01 - Biology and technology of oral and maxillofacial surgery

Programme (brief description)

In Oral and Maxillofacial Surgery, the reconstruction of defects of either teratological, traumatological or pathological disposition, is always directly related to optimal function and esthetics of the face. The regeneration of hard and soft tissues can be initiated with the biological process of distraction osteogenesis. This process is utilized in a clinical setting and further elucidated in an animal study. The purpose of the projects is the development and evaluation of this process.

1. Various rhythms and forces in distraction osteogenesis: an animal experimental study. In a rabbit model different rhythms and forces are clinically, radio graphically (including micro-CT) and histologically tested to investigate ideal parameters for distraction osteogenesis.
2. Surgically-assisted rapid maxillary expansion: surgical and orthodontic aspects. In a prospective randomized clinical trial two surgical methods of widening the narrow maxilla are tested. The bone-borne distraction is compared with the tooth-borne distraction.
3. Orbital wall fractures: a prospective clinical trial. In a prospective randomized clinical trial two surgical endoscopic-assisted methods of orbital wall fractures are explored: autogenous bone from the iliac crest is compared with an alloplastic material.
4. Midface-hypoplasia: various peri-operative aspects. In patients with congenital craniofacial anomalies the severe hypoplastic midface can be corrected by means of distraction osteogenesis. In this study short-term and long-term results are evaluated including the usage of modern 3D-CT scanning.

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
oral and maxillofacial surgery	1,6			1,75	3,35								
Total	1,6			1,75	3,35	0	0	0	0	3	0	2	5

Inaugural speech

Wolvius, E.B. (2011). *MKA en de juiste maat*. Oratie leerstoel mondziekten, kaak- en aangezichtschirurgie en bijzondere tandheelkunde. Rotterdam: ErasmusMC.

Article/Letter to the editor

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Veldhuis, A.H. te, Lobbezoo, F., Veldhuis, E.C. te, Naeije, M. & Selms, M. van (2011). Synoviale chondromatose van het temporomandibulaire gewricht, een systematisch literatuuronderzoek naar de kenmerken. *Nederlands Tijdschrift Tandheelkunde*.

EMC OR-01-60-01 - Diagnosis and treatment of disorders of the posterior segment and strabismus

Programme (brief description)

Studies of cornea, uveal melanoma, macular diseases and strabismus

Ways are sought to prolong the survival of corneal grafts by local modulation of the immune response. Therefore, cellular and humoral immunity is examined in experimental corneal allograft rejection.

In a prospective and retrospective study of uveal melanoma patients the relation between chromosomal abnormalities and survival are analysed with molecular biology techniques. The detection of these chromosomal abnormalities in fine needle biopsies is validated with fluorescent in-situ hybridisation in collaboration with the Universities of Sheffield and Liverpool. Target genes like p16, p73 and pten are investigated. In a prospective study the control rate and complication rate is studied in uveal melanoma patients treated by stereotactic radiotherapy.

The pathogenesis of age-related macular degeneration (AMD) was studied in een group of very old. Macaque monkeys. Molecular mechanisms that causes senescence in the retina, apoptosis and the expression of somatostatine receptor 2A are studied. Somatostatin receptor binding may inhibit angiogenesis in AMD. A pilot study was performed with octreotide treatment for the control of subfoveal choroidal neovascularisation in AMD in men. A two-center prospective randomized masked trial is initiated to investigate its clinical effect. The study of the effect of adjuvant hormonal treatment in surgery for macular holes is ongoing.

The study of strabismus consists of three multicenter and cohort studies: Together with the University of Heidelberg a study is organised to address the question whether children with congenital strabismus should be operated early or late, involving 54 European clinics (1992-2002). A second randomized study is organised that compares the two surgical techniques that are most commonly used in congenital strabismus, in 9 German and 4 Dutch clinics (1997-2002). A new microelectronic device has been developed that measures compliance in patch wearing in children with amblyopia; a cohort study has been initiated to measure compliance in the treatment of amblyopia.

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
ophthalmology	1,8	0,4	1,4		3,6								
<i>Total</i>	<i>1,8</i>	<i>0,4</i>	<i>1,4</i>		<i>3,6</i>	<i>3</i>	<i>0</i>	<i>0</i>	<i>3</i>	<i>38</i>	<i>1</i>	<i>0</i>	<i>42</i>

Thesis

Ramdas, W.D. (2011, november 09). *Epidemiologic and genetic insights into open-angle glaucoma*. EUR (287 pag.). Prom./coprom.: Prof. Dr. J.R. Vingerling, Prof. Dr. Ir. C.M. van Duijn & N.M. Jansonius.

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Article/Letter to the editor

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EMC OR-01-62-02 - Otorhinolaryngology and head & neck surgery

Programme (brief description)

Can we improve early detection of disease?
Do prognostic models improve the quality of health care?

Head and neck cancer

Can we identify the role of organ-sparing strategies in head and neck cancer?
Do modern optical imaging and molecular biology add to the current diagnostic armamentarium?
What are key factors in optimal palliation of head neck cancer patients?
How can we improve Quality of Life in head and neck cancer?

Audiology/otology

Can we adequately describe epidemiology of hearing loss and can we identify etiological factors?
Can we improve diagnostics in hearing loss and central auditory processing disorders?
How can we effectively evaluate and optimise the result of Cochlear Implantation?

Speech

Is the Lidcombe programme for early stuttering intervention more (cost-)effective than the Demands and Capacities Model based treatment (Dutch standard) ?
Can risk factors for a Speech Sound Disorder (SSD) in preterm children at age 4 be identified? Is a short, parent based speech intervention programme for preterm children with a SSD at 2 years of age effective ?

Tissue engineering

Find methods to repair large tissue defects after oncologic surgery

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
orthopaedics	0,22				0,22								
otorhinolaryngology	8,16	0,37		0,37	8,9								
Total	8,38	0,37		0,37	9,12	3	0	0	3	23	4	1	31

Thesis

Coenraad, S. (2011, september 14). *Auditory maturation and congenital hearing loss in NICU infants*. EUR (Ridderkerk: Ridderprint BV). Prom./coprom.: Prof. Dr. R.J. Baatenburg de Jong, Prof. Dr. J.B. van Goudoever, Dr.ir. A. Goedegebure & Dr. L.J. Hoeve.

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Verseijden, F. (2011, januari 07). *Engineering vascularized adipose tissue*. EUR (144 pag.). Prom./coprom.: Prof. Dr. S.E.R. Hovius, Prof. Dr. G.J.V.M. van Osch & Dr. S.O.P. Hofer.

Article/Letter to the editor

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Book - monography - book editors

Franken, M.C.J.P. & Hakkesteegt, M.M. (2011). *Eldar: Spreken en Zingen*. Houten: Van Gorcum.

Part of book - abstract

Buul, G.M. van & Osch, G.J.V.M. van (2011). Musculoskeletal stem cells. In G. Steinhoff (Ed.), *Regenerative Medicine: from protocol to patient*. London: Springer Publ..

EMC OR-01-65-03 - Addiction research

Programme (brief description)

What is the prevalence and incidence of drug use, alcohol consumption and other lifestyles and addiction? Which factors are the determinants of lifestyles and addiction? What are the consequences of lifestyles and addiction in terms of health, quality of life, environment (a.o. nuisance). What is the quality, efficiency of addiction care, prevention and policy? What is the interaction of lifestyles and addiction with society

Key Figures

Department	Sp1	Sp2	Sp3	Sp4	SpTot	T1	T2	T3	Ttot	IS	FS	OP	Ptot
addiction research	0,79			5	5,79								
Total	0,79			5	5,79	1	0	0	1	4	1	16	22

Thesis

Rooij, A.J. van (2011, mei 11). *Online video game addiction. Exploring a new phenomenon*. EUR. Prom./coprom.: Prof. Dr. H. van de Mheen.

Onderzoekscasus

Risselada, A & Schoenmakers, T.M.

Article/Letter to the editor

Baars, J.E., Boon, B., Garretsen, H.F.L. & Mheen, D. van de (2011). The reach of a hepatitis B vaccination programme among men who have sex with men. *European Journal of Public Health*, 21(3), 333-337.

Bommel , J. (2011). Legalisering van online gokken in Nederland: kansen of bedreigingen. *TSG*, 89(8), 362-363.

Labree, L.J.W., Mheen, H. van de, Rutten, F.F.H. & Foets, M.M.E. (2011). Differences in overweight and obesity among children from migrant and native origin: a systematic review of the European literature. *Obesity Reviews*, 12(5), 535-547.

Naaborgh, LA & Schoenmakers, T.M. (2011). Signalering, screening en kortdurende interventie bij jongeren met risicovol alcoholgebruik,. *Verslaving*, 7, 3.

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EMC OR-01-74-01 - Philosophy of Medicine

Programme (brief description)

Are the obese the lepers of modern time? When is suffering unbearable? Is it okay to enhance one's memory by taking a pill? Can doctors still do the best for individual patients in health care systems organised by market principles? Is there an obligation to donate a kidney to one's close ones? Just some examples of questions we are working on now.

In our research the following themes play a central role:

- Ethics and individual responsibility for health
- Ethics, beauty and appearance
- Ethics and fiction
- Ethics and paediatrics
- Ethics and technology assessment and professional morality
- Ethics and transplantation

These are the themes that we have been working on for at least 5 years. More recent subjects we are working on are: medical decisions regarding the end of life, neuroethics and ethical aspects of pandemics.

We work in interdisciplinary contexts we; are not afraid of practical solutions, but also fascinated by underlying fundamental ethical issues. We strive for international cooperation, particularly via our EU projects, but we also participate in the Dutch debate.

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
medical history	6,46	3,1	1	1,22	11,78								
Total	6,46	3,1	1	1,22	11,78	1	0	0	1	12	12	45	70

Article/Letter to the editor

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Rapport

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EMC OR-01-86-13 - Palliative and supportive care

Programme (brief description)

Palliative care aims to improve the quality of life of patients and their relatives in case of life-threatening disease. Because of the complexity of problems, multidisciplinary partnership is considered of utmost importance. When the underlying disease is still responsive to specific therapy, supportive care will be given to enhance the patient's quality of life by (symptomatic) treatment of concurrent disease- or therapy-induced complications. The research programme includes studies on the prevalence of symptoms and psychosocial problems in cancer patients at different stages during the course of disease, their needs and perceived quality of life and the development of clinical interventions to achieve symptom control in these patients. Specific attention is given to the development of effective interventions for chemotherapy-induced complications, medical and nurse-initiated interventions in pain and fatigue management, the use of a care pathway for the dying and end-of-life decisions.

Key Figures

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
medical oncology	0,7			1	1,7								
medical psychology & psychotherapy	0,2				0,2								
neurology	0,03				0,03								
public health	0,04	2,14			2,18								
Total	0,97	2,14		1	4,11	0	0	0	0	7	3	5	15

Article/Letter to the editor

Baan, M.A.G., Mesa, AA, Steen, AL Adriaens-van de & Rijt, C.C.D. van der (2011). Survey on the Prevalence of Pain in Dutch District Nursing Care. *Journal of Palliative Medicine*, 14(6), 696-699.

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EMC OR-02-47-12 - Surgery, non-programme associated**Key Figures**

<i>Department</i>	<i>Sp1</i>	<i>Sp2</i>	<i>Sp3</i>	<i>Sp4</i>	<i>SpTot</i>	<i>T1</i>	<i>T2</i>	<i>T3</i>	<i>Ttot</i>	<i>IS</i>	<i>FS</i>	<i>OP</i>	<i>Ptot</i>
surgery	2,9	1		1	4,9								
Total	2,9	1		1	4,9	0	0	0	0	10	3	0	13

Article/Letter to the editor

Berkowska, M.A., Driessen, G.J.A., Bikos, V, Grosserichter, C., Stamatopoulos, K, Cerutti, A, He, B, Biermann, K., Lange, J.F., Burg, M. van der, Dongen, J.J.M. van & Zelm, M.C. van (2011). Human memory B cells originate from three distinct germinal center-dependent and -independent maturation pathways. *Blood*, 118(8), 2150-2158.

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Thesis

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Article/Letter to the editor

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Part of book - abstract

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