

2019

RESEARCH REPORT

ASKLEPIOS KLINIKEN HAMBURG GMBH



ASKLEPIOS

Gesund werden. Gesund leben.



INTRODUCTION

DEAR COLLEAGUES, DEAR READERS,

as in the previous years, we at the Asklepios Kliniken Hamburg look back on a successful year in the area of clinical, translational and healthcare research. This is reflected by the numerous top-class publications with the participation of Hamburg scientists, which we would like to present to you. Not included in this report, but equally vital for the success of the scientific work of our departments are the numerous active contributions at national and international congresses, which are important for the high international reputation of our clinics and departments.

In this year's research report, we would like to invite you to look back on 20 years of ASKLEPIOS proresearch and, in addition, present a project that shows the outstanding scientific work being done in a way that is representative of many others.

Dr. Pawel Kuklik and Dr. Christian Eickholt from Prof. Dr. Stephan Willems' team will give you an outlook on upcoming randomized studies on atrial fibrillation, explain the pathophysiological background to this subject area and provide an insight into the potential of the new high-resolution 3-D mapping methods.

As usual, we have included a summary of the research projects overview for the entire AKHH.

Thanks to the internal research funding provided by Asklepios Kliniken GmbH, we were able to realize numerous translational and medical care projects. One focus of our research continues to be the promotion of young scientists. The achievements of our students, but also the support of their supervisors is again reflected in an exciting, thematically highly diverse range of diploma theses. In this year's Asklepios Medical School article, Mrs. Grimm reports on the possibility of submitting a scientific paper as a substitute for the diploma thesis.

In brief, this year's research report once again mirrors the great commitment and motivation for scientific work, in addition to patient care at the highest level, at the Asklepios Kliniken Hamburg. We would like to express our great praise and thanks to all employees.

The team of ASKLEPIOS proresearch is looking forward to supporting you in your projects in the coming years.

Hamburg, April 2020
Dr. Nele Geßler
Management ASKLEPIOS proresearch



INTERNAL RESEARCH FUNDING

INTERNAL RESEARCH FUNDING OF THE ASKLEPIOS KLINIKEN HAMBURG GMBH

The management of the AKHH provided funds from the operating profit of the AKHH for internal research

funding for innovative projects in patient-oriented research. Of the 12 applications received, the ones listed below were newly approved.

Clinic	Department	Employee	Title
AKA	Oncology	PD Dr. G. Schilling Prof. Dr. D. Arnold	Development and Evaluation of Patient-oriented Treatment Algorithms at the Asklepios Tumor Center Hamburg
AKB	Visceral Surgery	Dr. G. Makridis	Perioperative Assessment of Residual Liver Function Capacity with the 13C Methacetin Breath Test (ReLiF Study)
AKB	Visceral Surgery	Dr. C. Wagner	miRNA as Diagnostic Marker in Pancreatic Adenocarcinoma
AKB	Cardiology	Dr. K. Schenke	CORRECT Radial RCT – CORonaRy angiography and intErventions via distal vs proximal aCcess – a randomized Trial of different RADIAL puncture sites
AKNH	Ophthalmology	PD Dr. M. Schargus	Development of a Hamburg IVOM Register
AKNO	Psychiatry	Dr. M. Hagemann-Goebel	Evaluation of Metacognitive Training for Depression (D-MTK): Randomized Controlled Trial (RCT) in Outpatient Psychiatric/Psychotherapeutic Care (application for renewal)

Overview of approved internal research funding projects 2019



ASKLEPIOS PRORESEARCH – MEDICAL RESEARCH WITH TRADITION



For more than 20 years, „ASKLEPIOS proresearch – Clinical Research and Development“ has brought together medicine and science and stands for research and innovation in the Asklepios Clinics in Hamburg. In April 1999 ASKLEPIOS proresearch started with only two employees in Hamburg-Barmbek - at that time still under the aegis of the Hamburg State Hospitals (LBK, Landesbetrieb Krankenhäuser Hamburg). Today, 16 study coordinators in the seven houses of the Hamburg Asklepios clinics oversee the practical implementation of clinical research and development projects. In Building J, the core facility on the Asklepios Klinik St. Georg campus, another 16 employees coordinate projects with industry and support self-initiated study projects.

ASKLEPIOS proresearch acts as Site Management Organization (SMO) of AKHH. This includes a full service for researching physicians and for external industry and research partners – both in contract research and in self-initiated studies in all seven hospitals in Hamburg.

ASKLEPIOS proresearch provides support in the acquisition of industrial clinical studies, offers consultancy on the planning and implementation of innovation and research projects and assists with cost calculation and the application for subsidies. ASKLEPIOS proresearch also handles the application for authorizations and ethics committee approvals. Researching physicians at the Asklepios Kliniken Hamburg benefit from tailored study logistics.

The proresearch team's core tasks encompass advising, supporting and thus relieving the burden on the study centers, as well as providing regulatory and legal protection for all those involved.

Central elements of the service profile of ASKLEPIOS proresearch include Project service for scientists including project management in the conduct of clinical trials according to the German Medicines Act (AMG, Arzneimittelgesetz), the German Medical Products Directive (MPG, Medizinproduktegesetz) and professional law,

contract management, patent and innovation management, project controlling, central quality management as well as support and relief of the trial centers by trained personnel, the trial coordinators. ASKLEPIOS proresearch naturally applies Good Clinical Practice (GCP) and offers semi-annual GCP training courses for scientific personnel. When required, the biometrics staff take over the statistical evaluation of the data collected.

MEDICAL PROGRESS AND INNOVATION AT ASKLEPIOS

Every year, the AKHH management supports high-quality research projects by allocating financial resources for internal research funding. ASKLEPIOS proresearch accompanies the process of application selection by an interdisciplinary research commission and the subsequent implementation of all funded projects. In addition to start-up financing, the emphasis is on the expansion of a sustainable research infrastructure and the strengthening of the Group's research priorities. To this end, a call for projects is issued to all Group employees in the Fall of each year. A cooperation links ASKLEPIOS proresearch with the Asklepios Medical School (AMS) in the field of publicly funded projects. Due to its non-profit status, AMS can apply for funds from foundations or other public institutions that require the applicant to be non-profit. In addition, there are numerous links through students of the Asklepios Medical School, who are involved in research and other scientific activities of various departments as part of their diploma theses.

RESEARCH IN EVERYDAY CLINICAL PRACTICE

The challenges in clinical research are manifold, as the complexity of research projects increases – both due to regulatory requirements and a growing number of partners involved. The pharmaceutical and medical device industry is interested in carrying out clinical studies





The Asklepios proresearch team

efficiently and successfully; and researching physicians need support in carrying out their own research projects. Investigator initiated trials (IITs) are essential for optimizing treatment and therapy concepts. However, IITs represent a special challenge, since the same administrative and legal requirements apply to these studies as to comparable studies by industrial sponsors. Even studies that do not fall under the German Medicines Act must comply with GCP criteria. For all these studies it is imperative to find, with limited financial and personnel resources, the optimum solution to keep the burden on researching physicians as low as possible while maintaining the best possible data quality and highest patient safety. ASKLEPIOS proresearch is a reliable partner in all projects, takes over the administration of the projects and provides personnel support as well so that research can be carried out alongside everyday clinical practice.

BIOMETRICS AND DATA ANALYSIS

ASKLEPIOS proresearch accompanies scientific projects through the entire project cycle and offers statistical support for clinical studies, registries or the collection of clinical routine data. Prior to the beginning of a study,

our biometrician supports interested physicians in the preparation of a suitable project design and case number estimation and, on request, creates randomization lists. Upon study completion, the biometrician advises on results analysis and implements measures to prevent statistical bias.

Data analysis is followed by the results output with suitable key figures as well as the graphical representation of interrelationships between therapy, risk factors and treatment success. Results are prepared for publication in scientific journals and statistical questions from the article review are answered.

CLINICAL RESEARCH AT THE HIGHEST QUALITY LEVEL

Supported by a centrally controlled, supra-regional quality management, ASKLEPIOS proresearch guarantees the implementation of studies at the highest quality level in accordance with the German Medicines and Medical Devices Act and Good Clinical Practice. Based on the requirements of the DIN ISO 9001:2015 certification as well as the legal regulations, ASKLEPIOS proresearch has established a quality management system which is

continuously maintained and expanded. The QMS specifications of ASKLEPIOS proresearch are binding for all activities performed by departments, institutes and clinics of the AKHH within the scope of research projects. The ASKLEPIOS proresearch QMS thus lays the foundation for uniformly high standards in the realization of clinical research and development projects and provides legal protection for researching physicians and departments.

ACHIEVING AMBITIOUS GOALS TOGETHER

Over the past 20 years and in cooperation with ASKLEPIOS proresearch, physicians have made clinical research at the Asklepios Kliniken Hamburg an integral part of everyday hospital life. Our goal has always been to support ambitious and research-oriented physicians in their work in order to jointly advance medical progress and to excel in future-oriented, unparalleled patient care. Together with you, we would like to continue implementing innovative research projects in the coming years.

Questions? Please feel free to reach out to ASKLEPIOS proresearch by phone at +49 40-181885-3160 or by e-mail at info.proresearch@asklepios.com.

LISTS OF PUBLICATIONS

BY SPECIALIZATION

ANESTHETICS AND INTENSIVE CARE

Best practice & research clinical anaesthesiology: Advances in haemodynamic monitoring for the perioperative patient: Perioperative cardiac output monitoring.

Bein B, Renner J.

Best Pract Res Clin Anaesthesiol. 2019 Jun;33(2):139-153. doi: 10.1016/j.bpa.2019.05.008. Epub 2019 Jul 22. Review.

Perioperative myocardial ischemia: Current aspects and concepts.

Bein B, Schiewe R, Renner J.

Anaesthesist. 2019 Aug;68(8):497-508. doi: 10.1007/s00101-019-0605-z. Review. German.

Adipositas in der Anästhesie – Big is beautiful?

Bein B, Scholz J.

Anesthesiol Intensivmed Notfallmed Schmerzther. 2019 Apr;54(4):240-241. doi: 10.1055/a-0820-6582. Epub 2019 Apr 12. German.

Factor XIII activity in patients requiring surgical re-exploration for bleeding after elective cardiac surgery – a prospective case control study

Elisabeth H. Adam, Jens Meier, Bernd Klee, Kai Zacharowski, Christian Weber, Andreas Pape

Journal of Critical Care 2019, accepted

Anaesthesia in Adult Patients with Obesity.

Fudickar A, Bein B.

Anesthesiol Intensivmed Notfallmed Schmerzther. 2019 Apr;54(4):242-254. doi: 10.1055/a-0636-2782. Epub 2019 Apr 12. German.

Prehospital non-invasive ventilation in acute respiratory failure is justified even if the distance to hospital is short.

Hensel M, Strunden MS, Tank S, Gagelmann N, Wirtz S, Kerner T.

Am J Emerg Med. 2019 Apr;37(4):651-656. doi: 10.1016/j.ajem.2018.07.001.

Granisetron reduces the need for uterotonics but not sympathomimetics during cesarean delivery

Raimann FJ, Baldauf HP, Louwen F, Jennewein L, Fischer D, Zacharowski K, Weber CF

Int J Gynaecol Obstet. 2019 Jun;145(3):361-366

Influence of the WOMAN Trial on national wide standard operating procedures for treatment of postpartum hemorrhage

Raimann FJ, Jennewein L, Sonntagbauer M, Raddatz LM, Möllmann CJ, Louwen F, Zacharowski K, Weber CF

J Gynecol Obstet Hum Reprod. 2019 Apr;48(4):269-273

Comparing four video laryngoscopes and one optical laryngoscope with a standard macintosh blade in a simulated trapped car accident victim

Raimann FJ, Tepperis D, Meininger D, Zacharowski K, Schalk R, Byhahn C, Weber CF, Mutlak H

Emergency Medicine International; Volume 2019, Article ID 9690839

Fast interpretation of thrombelastometry and aggregometry in patients suffering from chronic liver disease

Stegewerth K, Weber CF, Moehlmann M, Adam EH, Zacharowski Z, Zeuzem S, Weiler N

Clin Lab. 2019 Aug 1;65(8) 1513-1521

Mobile phone-based alerting of CPR-trained volunteers simultaneously with the ambulance can reduce the resuscitation-free interval and improve outcome after out-of-hospital cardiac arrest: A German, population-based cohort study.

Stroop R, Kerner T, Strickmann B, Hensel M.

Resuscitation. 2019 Dec 27. pii: S0300-9572(19)30735-X. doi: 10.1016/j.resuscitation.2019.12.012. [Epub ahead of print]

Genome-wide association study of myocardial infarction, atrial fibrillation, acute stroke, acute kidney injury and delirium after cardiac surgery - a sub-analysis of the RIPHeart-Study.

Westphal S, Stoppe C, Gruenewald M, Bein B, Renner J, Cremer J, Coburn M, Schaelte G, Boening A, Niemann B, Kletzin F, Roesner J, Strouhal U, Reyher C, Laufenberg-Feldmann R, Ferner M, Brandes IF, Bauer M, Kortgen A, Stehr SN, Wittmann M, Baumgarten G, Struck R, Meyer-Treschan T, Kienbaum P, Heringlake M, Schoen J, Sander M, Treskatsch S, Smul T, Wolwender E, Schilling T, Degenhardt F, Franke A, Mucha S, Tittmann L, Kohlhaas M, Fuernau G, Brosteanu O, Hasenclever D, Zacharowski K, Meybohm P; RIPHeart-Study Collaborators. BMC Cardiovasc Disord. 2019 Jan 24;19(1):26. doi: 10.1186/s12872-019-1002-x.

Genome-wide association study of myocardial infarction, atrial fibrillation, acute stroke, acute kidney injury and delirium after cardiac surgery – a subanalysis of the RIPHeart-Study

Westphal S, Stoppe C,... Zacharowski K, Meybohm P and RIPHeart-Study Collaborators (amongst others: Weber C)

BMC Cardiovasc Disord. 2019 Jan 24;19(1):26. doi: 10.1186/s12872-019-1002-x

Epinephrine in Out-of-Hospital Cardiac Arrest.

Wnent J, Bein B, Fischer M; German Resuscitation Registry Steering.

N Engl J Med. 2019 Jan 24;380(4):396-7. doi: 10.1056/NEJMc1816187.

CARDIOLOGY AND CARDIAC SURGERY

Treatment with mononuclear cell populations improves post-infarction cardiac function but does not reduce arrhythmia susceptibility

Andrié RP, Beiert T, Knappe V, Linhart M, Stöckigt F, Klein AM, Ghanem A, Lübkemeier I, Röhl W, Nickenig G, Fleischmann BK, Schrickel JW.

PLoS One. 2019 Feb 14;14(2):e0208301. doi: 10.1371/journal.pone.0208301. eCollection 2019

Cardiac contractility modulation improves long-term survival and hospitalizations in heart failure with reduced ejection fraction

Anker SD, Borggrefe M, Neuser H, Ohlow MA, Röger S, Goette A, Remppis BA, Kuck KH, Najarian KB, Gutterman DD, Rousso B, Burkhoff D, Hasenfuss G.

Eur J Heart Fail. 2019 Sep;21(9):1103-1113. doi: 10.1002/ehf.1374. Epub 2019 Jan 16

Low systemic arterial compliance is associated with increased cardiovascular morbidity and mortality in aortic valve stenosis.

Bahlmann E, Cramariuc D, Saeed S, Chambers JB, Nienaber CA, Kuck KH, Lønnebakken MT, Gerds E

Heart. 2019;105:1507-1514.

Evaluating Real-World Clinical Outcomes in Atrial Fibrillation Patients Receiving the WATCHMAN Left Atrial Appendage Closure Technology

Boersma LV, Ince H, Kische S, Pokushalov E, Schmitz T, Schmidt B, Gori T, Meincke F, Protopopov AV, Betts T, Mazzone P, Foley D, Grygier M, Sievert H, De Potter T, Vireca E, Stein K, Bergmann MW; following investigators and institutions participated in the EWOLUTION study. Circ Arrhythm Electrophysiol. 2019 Apr;12(4):e006841. doi: 10.1161/CIRCEP.118.006841.

Five-year clinical outcome of multicenter randomized trial comparing amphilimus - with paclitaxel-eluting stents in de novo native coronary artery lesions

Carrié D, Berland J, Verheye S, Hauptmann KE, Vrolix M, Musto C, Berti S, Dibié A, Maupas E, Antonucci D, Schofer J.

Int J Cardiol. 2019 Nov 15. pii: S0167-5273(18)35877-7. doi: 10.1016/j.ijcard.2019.10.058.

Inadvertent transseptal puncture into the aortic root: the narrow edge between luck and catastrophe in interventional cardiology

Chen H, Fink T, Zhan X, Chen M, Eckardt L, Long D, Ma J, Rosso R, Mathew S, Xue Y, Ju W, Wasmer K, Ma C, Yang J, Maurer T, Yang B, Heeger CH, Ho SY, Kuck KH, Wu S, Ouyang F.

Europace. 2019 Jul 1;21(7):1106-1115. doi: 10.1093/europace/euz042.

Safety and efficacy of endovascular ultrasound renal denervation in resistant hypertension: 12-month results from the ACHIEVE study

Daemen J, Mahfoud F, Kuck KH, Andersson B, Böhm M, Graf T, Sievert H, Kahlert P, Iyer M, Zeller T.

J Hypertens. 2019 Sep;37(9):1906-1912. doi: 10.1097/HJH.0000000000002120

A meta-analysis of the effect of stent design on clinical and radiologic outcomes of carotid artery stenting.

de Vries EE, Meershoek AJA, Vonken EJ, den Ruijter HM, van den Berg JC, de Borst GJ; ENDORSE Study Group.

J Vasc Surg. 2019 Jun;69(6):1952-1961.e1. doi: 10.1016/j.jvs.2018.11.017.

Renewal Theory as a Universal Quantitative Framework to Characterize Phase Singularity Regeneration in Mammalian Cardiac Fibrillation

Dharmaprani D, Schopp M, Kuklik P, Chapman D, Lahiri A, Dykes L, Xiong F, Aguilar M, Strauss B, Mitchell L, Pope K, Meyer C, Willems S, Akar FG, Nattel S, McGavigan AD, Ganesan AN.

Circ Arrhythm Electrophysiol. 2019 Dec;12(12):e007569. doi: 10.1161/CIRCEP.119.007569. Epub 2019 Dec 9.

Outcomes of transcatheter aortic valve replacement without predilation of the aortic valve: Insights from 1544 patients included in the SOURCE 3 registry.

Dumonteil N, Terkelsen C, Frerker C, Collart F, Wöhrle J, Butter C, Hovorka T, Pinaud F, Baumgartner H, Tarantini G, Wendler O, Lefèvre T; SOURCE 3 Investigators.

Int J Cardiol. 2019 Dec 1;296:32-37. doi: 10.1016/j.ijcard.2019.06.013. Epub 2019 Jun 12.

Does valve in valve TAVR carry a higher risk for thromboembolic events compared to native valve TAVR?

Eitan A, Brinkmann C, Haselbach T, Witt J, Schofer J.

Catheter Cardiovasc Interv. 2019 Jul 9. doi: 10.1002/ccd.28391.

Atrial fibrillation ablation strategies and outcome in patients with heart failure: insights from the German ablation registry

Eitel C, Ince H, Brachmann J, Kuck KH, Willems S, Gerds-Li JH, Tebbenjohanns J, Richardt G, Hochadel M, Senges J, Tilz RR.

Clin Res Cardiol. 2019 Jul;108(7):815-823. doi: 10.1007/s00392-019-01411-3. Epub 2019 Feb 20.

Educational needs among physicians treating patients with atrial fibrillation: lessons for Poland from the European Society of Cardiology international educational needs assessment study

Farkowski MM, Karliński MA, Sterliński M, Tomasik T, Antz M, Vahanian A, Kuck KH, Hindricks G, Dagres N, Heidbuchel H.

Pol Arch Intern Med. 2019 Sep 30;129(9):586-591. doi: 10.20452/pamw.14924. Epub 2019 Aug 5.

Procedural success, safety and patients satisfaction after second ablation of atrial fibrillation in the elderly: results from the German Ablation Registry

Fink T, Metzner A, Willems S, Eckardt L, Ince H, Brachmann J, Spitzer SG, Deneke T, Schmitt C, Hochadel M, Senges J, Rillig A.

Clin Res Cardiol. 2019 Dec;108(12):1354-1363. doi: 10.1007/s00392-019-01471-5. Epub 2019 Apr 5

Management of arrhythmia recurrence in patients with pulmonary vein stenosis following atrial fibrillation ablation

Fink T, Tilz RR, Heeger CH, Schlüter M, Feickert S, Rottner L, Reissmann B, Lemes C, Alessandrini H, Maurer T, Mathew S, Rillig A, Metzner A, Ouyang F, Kuck KH.

Europace. 2019 Oct 1;21(10):1494-1501. doi: 10.1093/europace/euz187

Higher left ventricular mass-wall stress-heart rate product and outcome in aortic valve stenosis.

Gerds E, Saeed S, Midtbø H, Rossebø A, Chambers JB, Einarsen E, Bahlmann E, Devereux R

Heart. 2019;105:1629-1633.

Cytosorb Adsorption During Emergency Cardiac Operations in Patients at High Risk of Bleeding.

Hassan K, Kannmacher J, Wohlmuth P, Budde U, Schmoeckel M, Geidel S.

Ann Thorac Surg. 2019 Jul;108(1):45-51. doi: 10.1016/j.athoracsur.2018.12.032. Epub 2019 Jan 23

Efficacy and Safety of Cryoballoon Ablation in Patients With Heart Failure and Reduced Left Ventricular Ejection Fractio - A Multicenter Study

Heeger CH, Abdin A, Mathew S, Reissmann B, Yalin K, Liosis S, Fink T, Proietti R, Eitel C, Vogler J, Lemeš C, Maurer T, Rillig A, Meyer-Saraei R, Graf T, Wohlmuth P, Goldmann B, Ouyang F, Kuck KH, Metzner A, Tilz RR.

Circ J. 2019 Jul 25;83(8):1653-1659. doi: 10.1253/circj.CJ-19-0151. Epub 2019 Jun 28

Left Atrial Appendage Isolation in Patients Not Responding to Pulmonary Vein Isolation

Heeger CH, Rillig A, Geisler D, Wohlmuth P, Fink T, Mathew S, Tilz RR, Reissmann B, Lemes C, Maurer T, Santoro F, Inaba O, Sohns C, Huang Y, Alessandrini H, Dotz I, Schlüter M, Metzner A, Kuck KH, Ouyang F.

Circulation. 2019 Jan 29;139(5):712-715. doi: 10.1161/CIRCULATIONAHA.118.037451.

Response by Heeger et al to Letter Regarding Article, „Left Atrial Appendage Isolation in Patients Not Responding to Pulmonary Vein Isolation: Benefit and Risks“.

Heeger CH, Rillig A, Geisler D, Wohlmuth P, Fink T, Mathew S, Tilz RR, Reissmann B, Lemes C, Maurer T, Santoro F, Inaba O, Sohns C, Huang Y, Alessandrini H, Dotz I, Schlüter M, Metzner A, Kuck KH, Ouyang F. *Circulation*. 2019 Jul 23;140(4):e169-e170. doi: 10.1161/CIRCULATIONAHA.119.040943. Epub 2019 Jul 22.

Outcomes of cryoballoon or radiofrequency ablation in symptomatic paroxysmal or persistent atrial fibrillation

Hoffmann E, Straube F, Wegscheider K, Kuniss M, Andresen D, Wu LQ, Tebbenjohanns J, Noelker G, Tilz RR, Chun JKR, Franke A, Stellbrink C, Garcia-Alberola A, Dorwarth U, Metzner A, Onarrak T, Brachmann J, Kuck KH, Senges J; FREEZE Cohort Study Investigators. *Europace*. 2019 Sep 1;21(9):1313-1324. doi: 10.1093/europace/euz155.

Valve-in-Valve Implantation Using the ACURATE Neo in Degenerated Aortic Bioprostheses: An International Multicenter Analysis

Holzamer A, Kim WK, Rück A, Sathananthan J, Keller L, Cosma J, Bauer T, Nef H, Amat-Santos IJ, Brinkert M, Husser O, Pellegrini C, Schofer J, Nerla R, Montorfano M, Giannini F, Stella P, Kuwata S, Hilker M, Castricola F, Ussia GP, Webb JG, Nietlisbach F, Toggweiler S. *JACC Cardiovasc Interv*. 2019 Nov 25;12(22):2309-2316. doi: 10.1016/j.jcin.2019.07.042.

Mechanisms of sex differences in atrial fibrillation: role of hormones and differences in electrophysiology, structure, function, and remodelling

Katja E. Odening, Sebastian Deiß, Dagmara Dilling-Boer, Maxim Didenko, Urs Eriksson, Sotirios Nedios, Fu Siang Ng, Ivo Roca Luque, Pepa Sanchez Borque, Kevin Vernooij, Adrianus P. Wijnmaalen, and Hikmet Yorgun *Europace* (2019) 21, 366–376

Impact of an interatrial shunt device on survival and heart failure hospitalization in patients with preserved ejection fraction.

Kaye DM, Petrie MC, McKenzie S, Hasenfubeta G, Malek F, Post M, Dougherty RN, Trochu JN, Gustafsson F, Lang I, Kolodziej A, Westenfeld R, Penicka M, Rosenberg M, Hausleiter J, Raake P, Jondeau G, Bergmann MW, Spelman T, Aytug H, Ponikowski P, Hayward C, investigators RL-Hs. *ESC Heart Fail* 2019;6:62-69

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Predictive impact of previous coronary artery bypass grafting on mortality after MitraClip implantation for ischemic functional mitral regurgitation

Kitamura M, Kaneko H, Schlüter M, Schewel D, Schmidt T, Alessandrini H, Kreidel F, Okamoto M, Neuss M, Butter C, Kuck KH, Frerker C. *Int J Cardiol*. 2019 Jun 15;285:21-26. doi: 10.1016/j.ijcard.2019.02.045. Epub 2019 Mar 1

Design and rationale for the „Me & My Heart“ (eMocial) study: A randomized evaluation of a new smartphone-based support tool to increase therapy adherence of patients with acute coronary syndrome

Krackhardt F, Maier LS, Appel KF, Köhler T, Ghanem A, Tschoepe C, Dahl JV, Degenhardt R, Niklasson A, Ahlqvist M, Waliszewski MW, Jörnten-Karlsson M. *Clin Cardiol*. 2019 Nov;42(11):1054-1062. doi: 10.1002/clc.23254. Epub 2019 Sep 6.

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Kuck KH, Albenque JP, Chun KJ, Fürnkranz A, Busch M, Elvan A, Schlüter M, Braegelman KM, Kueffer FJ, Hemingway L, Arentz T, Tondo C, Brugada J; FIRE AND ICE Investigators. *Circ Arrhythm Electrophysiol*. 2019 May 22;12(6):e007247. doi: 10.1161/CIRCEP.119.007247

Catheter Ablation Versus Best Medical Therapy in Patients With Persistent Atrial Fibrillation and Congestive Heart Failure: The Randomized AMICA Trial

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Double Filtration During Carotid Artery Stenting Using a Novel Post-Dilation Balloon With Integrated Embolic Protection

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Internal Versus External Electrical Cardioversion of Atrial Arrhythmia in Patients With Implantable Cardioverter-Defibrillator: A Randomized Clinical Trial

Lüker J, Kuhr K, Sultan A, Nölker G, Omran H, Willems S, Andrié R, Schrickel JW, Winter S, Vollmann D, Tilz RR, Jobs A, Heeger CH, Metzner A, Meyer S, Mischke K, Napp A, Fahrige A, Steinhauser S, Brachmann J, Baldus S, Mahajan R, Sanders P, Steven D. *Circulation*. 2019 Sep 24;140(13):1061-1069. doi: 10.1161/CIRCULATIONAHA.119.041320. Epub 2019 Aug 30

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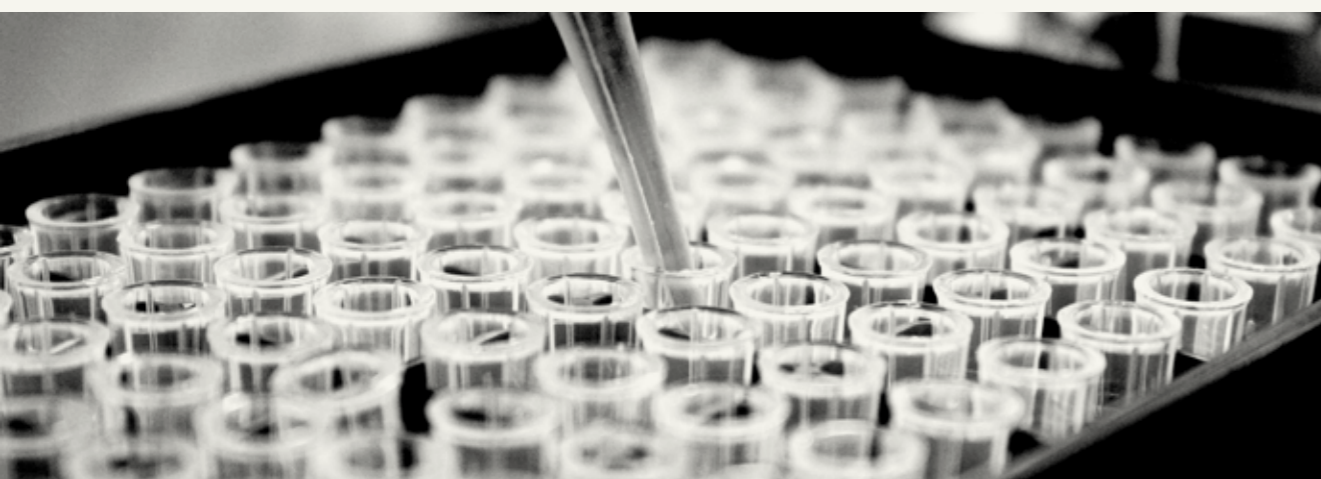
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ASKLEPIOS MEDICAL SCHOOL PUBLICATIONS AS A SUBSTITUTE FOR DIPLOMA THESES

DR. MONIKA GRIMM
DR. CHRISTOPH JERMANN



Students of the Asklepios Campus Hamburg (ACH) of Semmelweis University also engage in scientific research at Asklepios. They are obliged to write a scientific paper, the diploma thesis, as part of their studies. The majority of the students do this at the Asklepios clinics in Hamburg, thereby supporting ongoing research projects. In order to increase the visibility of the students' research activities, they are welcome to submit an accepted publication as first author in a peer-reviewed journal instead of a thesis.

The journal's impact factor is not decisive for recognition as a diploma thesis. A publication with shared first authorship is also recognized as a diploma thesis, yet only by one of the first authors.

The first student to take advantage of the opportunity to submit a publication as a substitute for the diploma thesis was Jakob Feldt who, in 2019, wrote his diploma thesis in the Cardiology Department of the Asklepios Klinik St. Georg under the supervision of Dr. Thomas Fink and PD Dr. Andreas Metzner. His paper was published in the journal *Current Opinion in Cardiology*, titled „Catheter ablation in elderly patients: a review of current studies and pending questions“ (*Curr Opin Cardiol.* 2019, 34(1): 29–34). In his review article, Mr. Feldt provides an overview of current studies dealing with the risk of catheter ablation in elderly patients. He concludes that, in most cases, the patient's general condition plays a more decisive role for a well-tolerated and effective catheter ablation than the patient's age.

In 2020, Jendrik Becker-Assmann was the next to seize the opportunity to submit a publication as a replacement for his thesis. He wrote his thesis under the supervision of Mohammad-Hossein Fard-Aghaie, MD, in the Department of Prof. Karl J. Oldhafer, MD. The work was published

in the journal *Der Chirurg* with the title „Diagnostic and prognostic significance of the α -feto protein in hepatocellular carcinoma“ (*Chirurg.* 2020. doi: 10.1007/s00104-020-01118-6). In his study, Mr. Becker-Assmann concludes that the α -feto-protein (AFP) level can be attributed a therapeutic utility and that the measured AFP concentration has therapeutic consequences regarding therapy strategy.

Publishing at an early stage as first author helps students increase the visibility of their research work while for the Departments, the supervision efforts invested pay off in the form of impact points earned. In this respect, the possibility of submitting a publication as a substitute for a diploma thesis is a win-win situation for all parties involved. We hope that many departments will make use of this possibility in the future. We already know of nine planned theses of this kind that shall be completed within the next three years. And more will certainly follow.



Figure 1: Graduates of the Asklepios Campus Hamburg of Semmelweis University, 2019

DIPLOMA THESES

YEAR 8

Name	Vorname	Thema	Betreuer
Arnold	Danielle Louisa	Drei Jahre Peritonektomie und HIPEC-Therapie an der Asklepios Klinik Barmbek Einzeitige und zweizeitige HIPEC-Applikation im deskriptiven Vergleich	Dr. med. Michael Lipp
Basedau	Hauke	Zentrale Verarbeitung nozizeptiver Reize – Bildgebende Methodiken zur Untersuchung der Neuromatrix	Dr. rer. nat. Jan Mehnert
Beckelmann	Stefanie Kerstin	Die kleinen Speicheldrüsenkarzinome – Epidemiologie, Therapie, Prognose	Prof. Dr. med. Jens E. Meyer
Benger	Konstantin	Winkelstabile Plattenosteosynthese am distalen Radius: Retrospektiver Vergleich von zwei unterschiedlichen Plattensystemen: Aptus Radius vs. VariAx DR	PD Dr. med. Marc Schult
Bjoernsgard	Karen Elisabeth	Geschlechtskranke nach dem zweiten Weltkrieg – Formen und Institutionen der Versorgung in Hamburg	Univ.-Prof. Dr. Florian Steger
Bohnen	Jan-Eric	Wirksamkeit und Sicherheit der Kyroballonablation bei Patienten über 75 Jahren	Dr. med. Christian-Hendrik Heeger

Brückner	Fabian	Der „erweiterte Suizid“ und seine Charakteristika im Vergleich – eine retrospektive Auswertung der Obduktionsfälle aus dem IfR Hamburg über den Zeitraum von 01.01.2005 - 31.03.2018	Prof. Dr. med. Jan Peter Spermhake
Burmeister	Martin	Kreativität und Morbus Parkinson – Eine Untersuchung im Asklepios Klinikum Harburg	Prof. Dr. med. Rudolf Töpfer
Cichutek	Sophia Marlene	Konstruktion und Funktionalitätstest lentiviraler und gammaretroviraler Vektoren zur Expression Nanobody-basierender chimärer Antigenrezeptoren (CAR) gegen CD38	PD Dr. rer. nat. Dipl.-Biochem. Kristoffer Riecken
Danzer	Maximilian	Evaluation der Klassifikation der patellofemorale Instabilität nach Frosch am Beispiel der Typen 3e und 5	Prof. Dr. med. Christian Kühne
Daß	Christian Camillo	Korrelation des late enhancements im linksventrikulären Myokard mit der Myokardbiopsie	PD Dr. med. Dietmar Kivelitz
Dekorsy	Antonia	Perioperative Schmerztherapie bei Tonsillektomie: Eine Literaturrecherche	Dr. med. Catharina Meyer zu Natrup
Diener	Leopold	Einfluss von technischen und operativen Parametern auf das postoperative Hörvermögen bei Elektroakustischer Stimulation	Prof. Dr. med. Jens E. Meyer
Duske	Vincent Sean	Der Barmbeker Frühwarnscore (BFS) – ein Pilotprojekt	Dr. med. Sebastian Wirtz
Feldt	Jakob	Catheter ablation in elderly patients: A review of current studies and pending questions	Dr. med. Thomas Fink
Firla	Jan	Adjuvante Chemotherapie beim duktalem Pankreaskarzinom	Prof. Dr. med. Karl J. Oldhafer
Geisler	Dominic Nicholas	Katheterablation basierte Isolation des linken Vorhofes: Klinischer Nutzen, Komplikationen und Prävention von Thrombembolien	Dr. med. Christian-Hendrik Heeger
Gerlitzky	Franziska Hedwig	Antibiotische und antimykotische Prävention postoperativer Komplikationen in der kolorektalen Chirurgie – Eine Untersuchung der Anastomoseninsuffizienzen und Wundinfektionen im Asklepios Klinikum Barmbek	Dr. med. Atingwa Tasi
Hagen	Marius	Der Einfluss von software-basierter Metallartefaktreduktion (iMAR) auf den klinischen Konturierungsprozess am Beispiel zweier Patientenfälle	Prof. Dr. Florian Würschmidt
Haneke	Mirko	Analyse eines Rotormappingsystems bei persistierendem Vorhofflimmern	Dr. med. Shibu Mathew
Heckschen	Konstantin	Auswahl der postoperativen Bildgebung zur Lageüberprüfung einer Stapesprothese	Prof. Dr. med. Jens E. Meyer
Hommelsheim	Stephan Alexander	Prognostische Bedeutung einer präprozeduralen Anämie bei hochgradiger Aortenstenose nach kathetergeführtem Aortenklappenersatz (TAVI)	Dr. med. Dmitry Schewel
Huber	Tessa Maria Christina	Aktuelle Behandlungskonzepte intrahepatischer cholangiozellulärer Karzinome in einem Leberzentrum	Dr. med. Gregor Stavrou
Hüttemann	Eva Lisa	Charakterisierung des Kallmann-Syndroms	Prof. Dr. med. Onno E. Janßen
Janzen	Jara	Patientenzufriedenheit auf der geschlossenen Akutstation der Psychiatrie: Sind Patienten zufrieden und wo besteht Verbesserungspotenzial, vor allem unter Berücksichtigung von Zwangsmaßnahmen?	Prof. Dr. med. Matthias Nagel
John	Alexander	Die CT-gesteuerte Radiofrequenzablation (RFA) zur minimal-invasiven Behandlung von Tumoren	Prof. Dr. med. W. Gross-Fengels
Korthaus	Alexander Sebastian	Unterschied zwischen der biplanaren aufsteigenden und der absteigenden medial öffnenden Tibiakopf Osteotomie	PD Dr. med. Matthias Krause

Krause	Janina Maren	Häodynamische Unterschiede und Einflussfaktoren biologischer Aortenklappenprothesen	Dr. med. Nicolai Bayer
Krümmel-bein	Michel Pierre	Vergleichende Mutationsanalyse mittels konventioneller molekulargenetischer Testung und Amplikon-basierter Next-Generation-Sequenzierung bei Adenokarzinomen der Lunge in einem Hamburger Patientenkollektiv	Dr. med. Markus Tiemann
Lehner	Louisa Antonella	Patientenscreening auf Familiäre Hypercholesterinämie: Mutationsfrequenz und Mutationsspektrum	Hon. Prof. Dr. med. Martin Merkel
Lehnert	Franziska	Antidepressant Effect of Botulinum Toxin A Injections in Outpatients with Affective Disorders – A Practical Experience	PD Dr. med. Marc Axel Wollmer
Lohmann	Lisa Andrea	Der Einfluss von Botulinumtoxin auf die emotionale Bewertung von Bildern und Wörtern	PD Dr. med. Marc Axel Wollmer
Malan	Marcel	Klinische Ergebnisse nach kombinierter distaler Femurosteotomie zur Behandlung der Patellainstabilität bei Genu Valgum	Dr. med. Jannik Frings
Mohr	Ann-Christin	Die Bedeutung eines umfassenden somato-psychischen Behandlungskonzeptes anhand des Krankheitsbildes Brustkrebs im Hinblick auf das Outcome	Dr. med. Goetz Broszeit
Obert	Jan Daniel Alexander	Eine retrospektive Analyse von 93 Patienten mit einem perihilärem Gallengangskarzinom	Hon. Prof. Dr. med. Siegbert Faiss
Paaß	Hendrik	Drei Jahre Peritonealdialyse-Katheter-Implantation an der Asklepios Klinik Barmbek – Ein erstes Follow-Up	Dr. med. Michael Lipp
Panzer	Juliane	Testosteronsubstitution bei Transmännern: Auswirkungen auf Hormonstatus, Metabolismus und die Häufigkeit der Autoimmunthyreoiditis Hashimoto	Prof. Dr. med. Onno E. Janßen
Preyer	Caroline Elisabeth Marie	Schnittführung der Cochlea Implantation – unter besonderer Berücksichtigung des minimalinvasiven Zugangs	Prof. Dr. med. Jens E. Meyer
Rieß	Juliane Christine	Der Einfluss von Übergewicht auf den Krankheitsverlauf der juvenilen idiopathischen Arthritis	Dr. med. Ivan Foeldvari
Sagasser	Joana	Konzeption und Entwicklung eines Lehrplan begleitenden Histopathologieskriptes für das 5. und 6. Semester der Asklepios Medical School – Pathologische Präparate der Kardiologie, Orthopädie, Urologie, Pulmologie und Gynäkologie	Prof. Dr. med. Katharina Tiemann
Salloum	Omar	Effektivität und Sicherheit der Cryoballon basierten PVI bei Patienten mit einem implantierten kardialen System.	Dr. med. Christian-Hendrik Heeger
Scheer	Tristan Ruben	Validierung und Weiterentwicklung eines Screening-Instrumentes zur Verkürzung von Überweisungszeiten bei Kindern mit Verdacht einer juvenilen idiopathischen Arthritis	Dr. med. Ivan Foeldvari
Schöppach	Leonie	Vergleich verschiedener Untersuchungsverfahren zur Differenzialdiagnose HPVpositiver und HPV-negativer Oropharynxkarzinome	Prof. Dr. med. Jens E. Meyer
Torster	Leopold	Konzeption und Entwicklung eines Lehrplan begleitenden Histopathologieskriptes für das 5. und 6. Semester des Asklepios Campus Hamburg	Prof. Dr. med. Katharina Tiemann
Traupe	Florian Maximilian	Auswirkung von Kryotherapie auf Laktatentwicklungen – ein zeit- und temperaturabhängiger Vergleich des Laktatabbaus am Beispiel von Wasserballspielern	Michael Ehnert
Valentin	Lisa	Neue Therapieoptionen bei kompliziertem nephrotischen Syndrom im Kindesalter	Prof. Dr. med. Markus J. Kemper

Wesseler	Antje	Diagnostik des Morbus Menière	PD Dr. med. Dietmar Kivelitz
Westphal	Christina	Die unterschiedlichen Arten der Cochleotomie	Prof. Dr. med. Jens E. Meyer
Wittmann	Kira Felicitas	Plattenostearythese bei Weber-B-Fraktur: Retrospektiver Vergleich von zwei unterschiedlichen Plattensystemen: Konturenplatte nach Brug vs. VariAx	PD Dr. med. Marc Schult
Wolbert	Lara	Hart- und Weichgewebsveränderungen nach chirurgischer Dysgnathiebehandlung in der Mund-, Kiefer- und Gesichtschirurgie anhand von kephalometrischer Fernröntgenseitenbildanalyse und Profildfotoauswertung	Dr. med. Dr. med. dent. N. B. Liebehenschel
Zäck	Matthias Julian	Prognostische Bedeutung des R2-CHA2DS2-VASc-Scores bei Patienten mit hochgradiger Aortenstenose nach TAVI	Dr. med. Jury Schewel

YEAR 9

Name	Vorname	Thema	Betreuer
Amian	Jacqueline	Einsatzmöglichkeiten des mikrovaskulären Unterarmklappens in der Kopf-Hals-Onkologie	Prof. Dr. Thomas Verse
Arbabzadah	Sahar	Synchrone kolorektale Lebermetastasen – Behandlungsstrategien und Überleben	Dr. Dr. habil. Gregor A. Stavrou
Bartels	Inga	Prädiktoren für postoperative Komplikationen nach zytoreduktiver Chirurgie (CRS) und hyperthermer intraperitonealer Chemotherapie (HIPEC) eine retrospektive Analyse	Dr. med. Michael Lipp
Baumgart	Elisabeth Maria	Der Einfluss des posterioren tibialen Slopes auf die Prävalenz von peripheren Bandinstabilitäten bei vorderen Kreuzbandrupturen	Dr. med. Lena Alm
Becker	Johannes David	Retrospektive Analyse von CT-gestützten transthorakalen Lungenpunktionen in den Jahren 2016/2017“	Prof. Dr. med. Roman Fischbach
Becker-Assmann	Jendrik	Diagnostische und prognostische Bedeutung des Alpha-Feto-Proteins (AFP) beim Hepatozellulären Karzinom.	Dr. med. Mohammad-Hossein Fard-Aghaie
Bertels	Jana	Durchführbarkeit und akuter Erfolg der Katheterablation atrialer Linien bei Patienten nach Mitralklappenoperationen	Prof. Dr. med. Stephan Willems/Christine Lemes
Bozcicek	Esra	Klinische und hormonelle Charakteristika von Patienten mit PubertätsgynäkomastieVergleich der Diagnostik mit den Empfehlungen der Leitlinie	Prof. Dr. Carl-Joachim Partsch
Braun	Florian	Spektrum des von Willebrand Syndroms mit besonderer Berücksichtigung des erworbenen von Willebrand Syndroms. Eine retrospektive Auswertung von 2010 – 2017.	Prof. Dr. med. Ulrich Budde
Drews	Jan	Stellenwert der Diagnostik der Endothelfunktion in der kardiologisch-angiologischen Praxis	Dr. Boris Leithäuser
Drexler	Richard	Die Etablierung einer immunhistochemischen Färbung des Hitzeschockproteins-27 im dukalen Adenokarzinom des Pankreas und eine Analyse des Expressionsmusters am Patientenkollektiv	Prof. Dr. K.-J. Oldhafer

Ellendt	Christina	Echokardiografische Analysen zur Rolle des linksventrikulären Ausflustrakts in Bezug auf die Schrittmacherpflichtigkeit nach TAVI	Prof. Dr. med. Derk Frank
Frank	Victoria Julia	Der Einfluss des Gelenkflächenwinkels bei Umstellungsosteotomien des Kniegelenkes	Dr. med. Lena Alm
Fröschle	Felix Florian	Interleukin 6 (IL-6) in der Diagnostik einer Anastomoseninsuffizienz am Ösophagus – eine retrospektive Analyse von 23 Patienten.	Dr. Tina Maghsoudi
Gémes	Kristóf György	Die PD-L1 Expression und lymphozytäre Infiltration bei Triple-negativen-Mammakarzinomen	Prof. Dr. med. Katharina Tiemann
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Hühnlein	Lisa	Sicherheit, audiologische Ergebnisse und gesundheitsbezogene Lebensqualität nach Cochlea Implantation mit HiRes 90K und HiFocus™ Mid-Scala Elektrode	Dr. med. Attila Övári
Katz	Jérôme	Die Ergebnisse der Behandlung der akuten Cholezystitis bei Patienten im Alter über 85 Jahren – eine retrospektive Studie	Dr. Thomas Mansfeld
Keller	Peter Sebastian	Aufbau eines Sarkomzentrums an der Asklepios Klinik Sankt Georg in Hamburg	Prof. Dr. med. habil. Carolin Tonus
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Kölzer	Charlotte	Verbesserung der Compliance von Patienten mit Hilfe der Smartphone App „Stone MD: Nierensteine“	Dr. med. Benedikt Becker
Kölzer	Hannah	Qualitative und quantitative Analyse von computertomographisch gesteuerten Knochenbiopsien bei Patientinnen mit Mammakarzinom	Prof. Dr. med. Roman Fischbach
Kuck	Marie Philine	Die hochauflösende Bildgebung der linksatrialen Anatomie des Herzens mittels eines neuen dielektrischen Breitband-Mapping-Systems	Dr. med. Tilmann Maurer
Labonte	Lukas Antonius Josef	Dehnen im Sport: Sinnhaftigkeit und Auswirkungen auf das Leistungsvermögen	Michael Ehnert
Leitner	Katharina Elisabeth	Surgical Risk Calculator – Evaluierung von Effizienz und Nutzen bezogen auf das Patientenkollektiv einer deutschen chirurgischen Abteilung	Prof. Dr. Dr. habil Thomas Carus
Lentz	Yannick	Postoperative Komplikation nach offener vs. laparoskopischer Resektion für Kolonkarzinome	Dr. med. Atingwa Tasi
Lorenz	Nikolas	Prävalenz der koronaren Herzkrankheit bei chronisch obstruktiver Lungenerkrankung – pathophysiologische Mechanismen und empirische Belege der Komorbidität	PD Dr. med. Hans-Peter Hauber
Ludwig	Julia	Die Sonographie in der Diagnostik eines Abszesses in der Kopf- und Halsregion – eine systematische Übersichtsarbeit	Prof. Dr. Jens Eduard Meyer
Martin	Isabell	Durchführbarkeit, Sicherheit und Akuteffektivität des Kryobalons der vierten Generation im Rahmen der Katheterablation von Vorhofflimmern	PD Dr. med. Andreas Rillig / Dr. med. Laura Rottner

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Pleye	Lars Nikolas	Wie unterscheiden sich Eingriffszeiten und Strahlenexposition bei chirurgischer und perkutaner EVAR?	Prof. Dr. Klaus Mathias
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Schinzel	Annika Lisa Marie	Effektivität und Risiken minimalinvasiver Stents in der Glaukomchirurgie	PD Dr. Marc Schargus
Sturm	Berit	Die COPD und ihre Komorbiditäten am Beispiel der Herzinsuffizienz	PD Dr. med. Hans-Peter Hauber
Töckelt	Feri	Die Wertigkeit der einmaligen intravenösen Gabe von Tranexamsäure beim unilateralen endoprothetischen Hüftgelenkersatz	Dr. Oliver Niggemeyer
Ullmann	Octavia	Vergleich der laparoskopischen Resektion mit der offenen Resektion des Rektumkarzinoms im Hinblick auf postoperative Komplikationen	Dr. med. Atingwa Tasi
Ustorf	Klara Marie	Ist die laparoskopische Appendektomie ein ausreichendes Verfahren bei perforierter Appendizitis – eine retrospektive Auswertung des Patientenguts eines städtischen Erstversorgerkrankenhauses in Hamburg.	Dr. Thomas Mansfeld
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RESEARCH PROJECTS OVERVIEW AKHH AGGREGATION LEVEL AKHH

In 2019, there are 392 active research projects and clinical studies, which represents a decrease compared to the previous year (440 active projects). The largest decline is seen at the Asklepios Westklinikum, while the number of projects in St. Georg and Altona has risen slightly. The number of events held within the AKHH last year dropped slightly from 136 to 115.

Most of the projects are still being implemented in the field of hemato-oncological/oncological and cardiovascular diseases. As in 2018, research in this area is still leading - with almost 140 active projects, the figures of the previous year have been exceeded (2018: 110 active projects). With a total of 96 projects, cardiology remains number two and, in contrast to the previous year, is also growing. For the first time, neurology and urology rank third and fourth with slightly increased project numbers. Gastroenterology, last year in 3rd place with over 70 projects, meanwhile shows a massive drop in study activity.

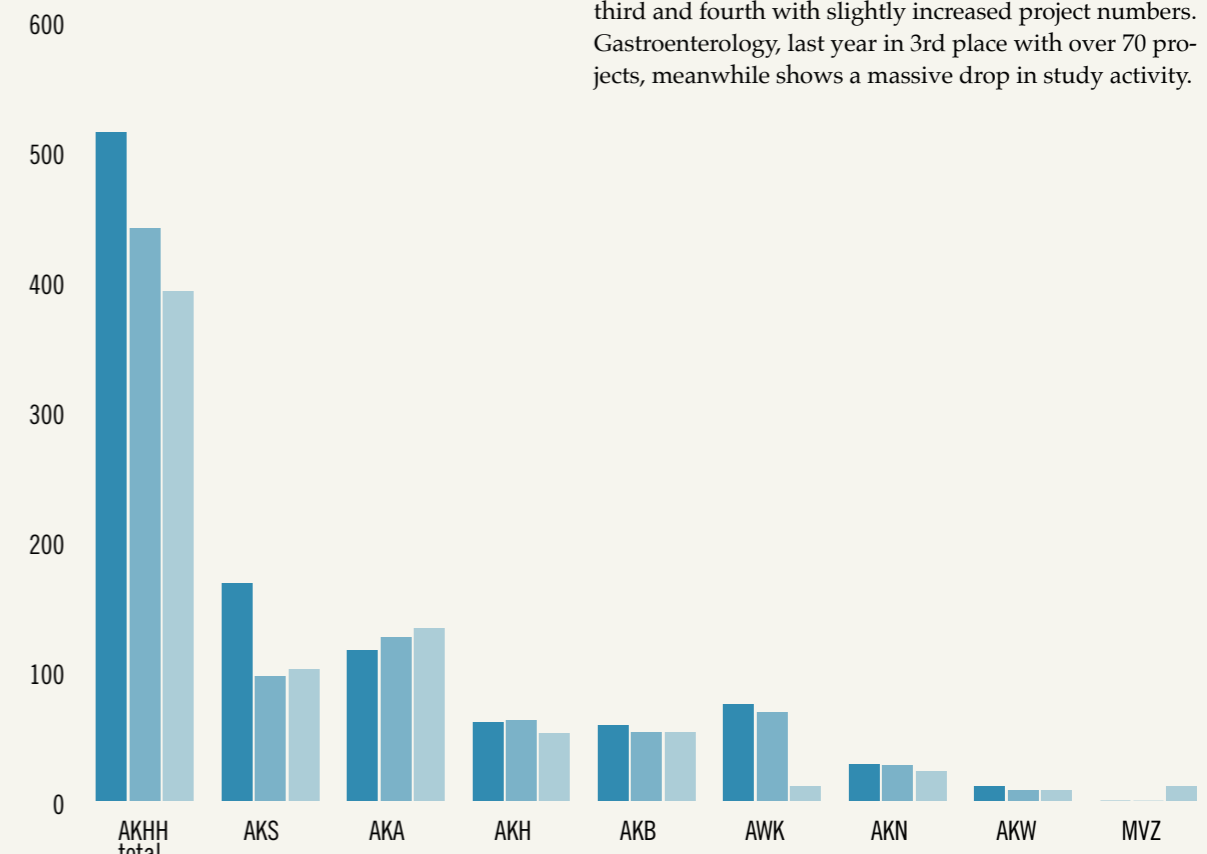


Figure 1: Active research projects AKHH 2017 to 2019



Main sites with a focus on

- Hemato-oncology research: AK Altona, AK St. Georg, AK Harburg (Lunge), AK Barmbek
- cardiovascular research: AK St. Georg, AK Harburg (Angiologie), AK Barmbek
- neurological research: AK Nord, AK Wandsbek
- urology research: AK Altona, AK Harburg, AK Barmbek

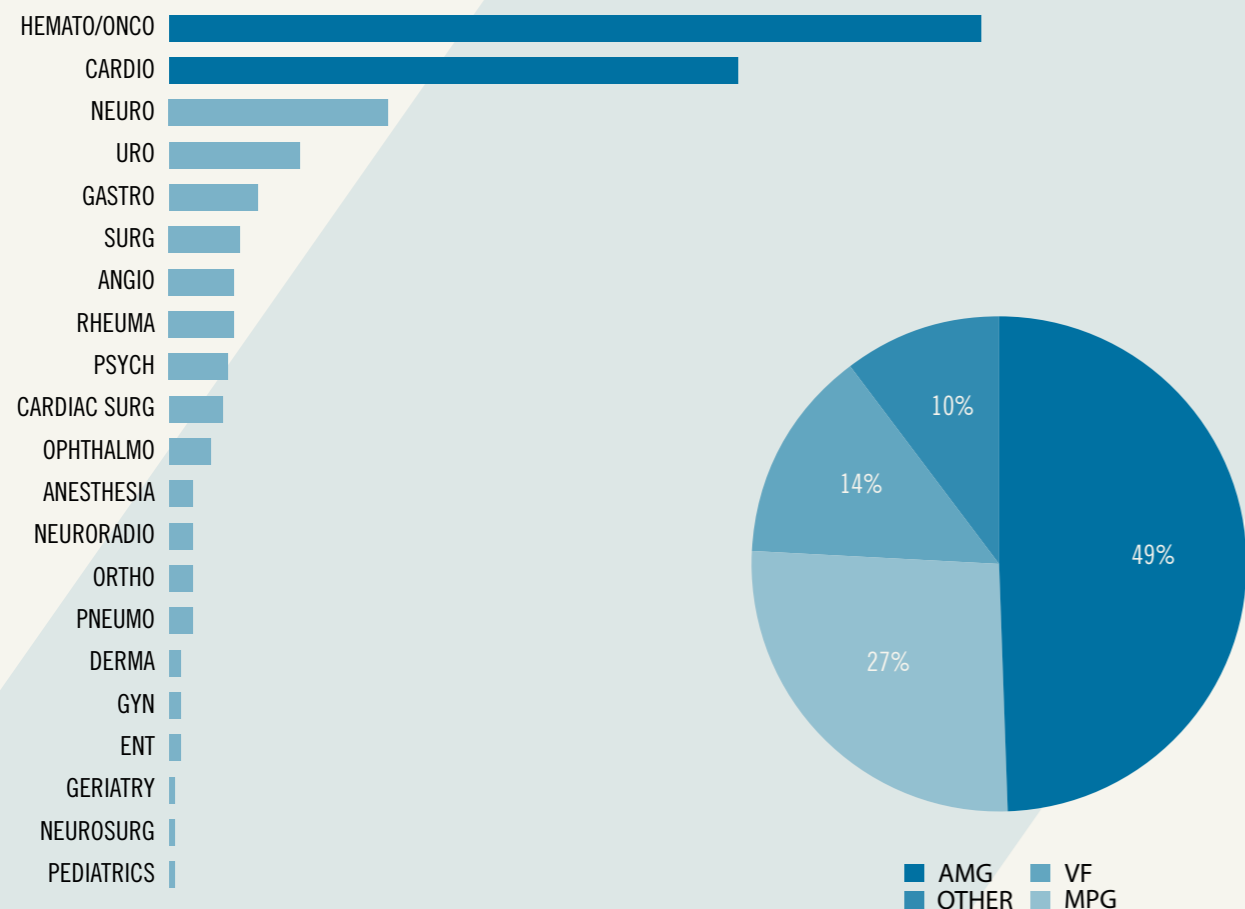


Figure 2: Research projects by specialization - AKHH research projects

The project portfolio largely corresponds to that of previous years. Almost 50 % of all projects are clinical trials under the German Medicines Act (AMG) - 53 % in the previous year - and 27 % under the German Medical Devices Act (MPG). A slight increase of 5 % was observed

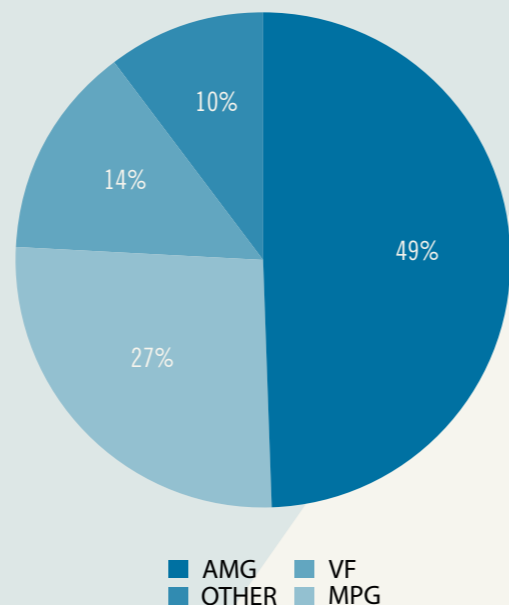


Figure 3: Project portfolio 2019 AKHH total (n=514)

in MPG studies. With a 76 % share, external contract research projects in cooperation with the pharmaceutical industry accounted for the largest part. The shares of health services research, sample collections and register studies are comparable to those of the preceding year.

The project portfolio of the clinics in the AKHH reflects the range of fundamental and patient-oriented research required by the Science Council. In this way, we continue

to meet our demand for university research-oriented science.

Number of projects at the AKHH 2019	392
Number of events at the AKHH 2019	117

Table 1: Number of projects and events in 2019 at the AKHH

EXTERNAL FUNDS

Considering all internal and external revenues, the total research revenue will amount to € 3,402,637 in 2019 and thus fall below the previous year's total (2018: € 4,028,873). With an amount of € 1,834,355, revenues from external contract research also lag behind the previous year's figures (€ 2,145,131). This can be mainly explained by the decline in industrial contract research in the field of gastroenterology, which has enjoyed rather high study activity for years. An increase in external revenues was recorded in the area of event administration (2018: 219,364 €).

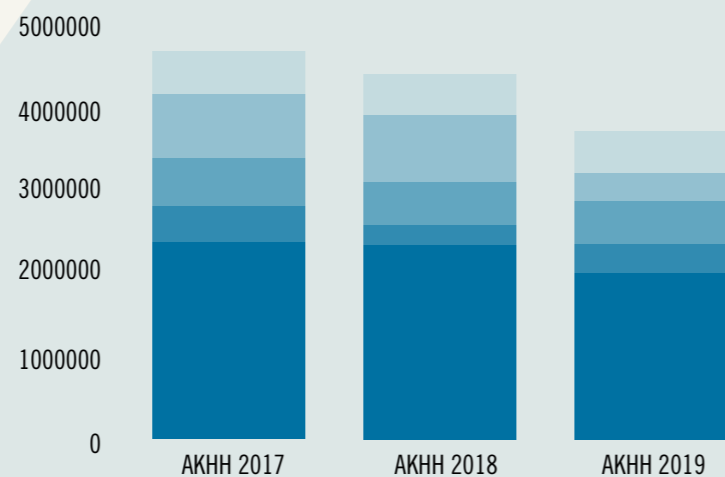
PROMEDIG gGmbH has recorded a slight increase in donations compared to 2018 (€ 472,801). Internal revenues (research funding and AMS apprenticeship bonus) were significantly lower than in the previous year due to a reduction in research funding. The third-party funding ratio (DMQ, Drittmittelquotient), which is calculated as the ratio of external revenues from contract research and events as well as donations received by PROMEDIG gGmbH to total research revenues, is 7 percentage points higher than in the previous year.

Total external funds AKHH	2019
Ext: contract research (incl. other operations)	1.726.626,29 €
Ext: public funds	107.728,80 €
Int: research funding	310.510,00 € ¹
Int: AMS apprenticeship bonus (incl. other operations)	463.277,82 € ²
Ext: events/event surplus	317.490,87 €
Ext: PROMEDIG gGmbH (stock/acquisition)	477.002,07 €
Total DMQ (= external revenues)	77,3 %
Total research revenue AKHH	3.402.636,62 €

Table 2: Total research revenue AKHH 2019

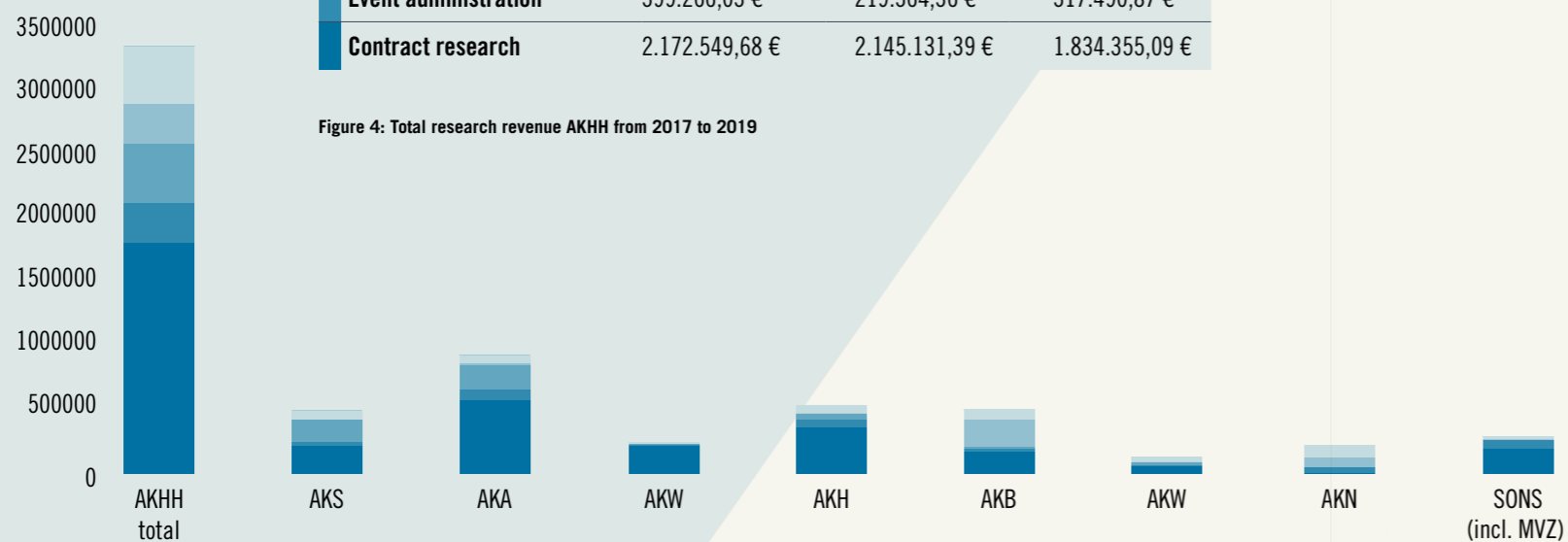
¹ incl. research funding by ASKLEPIOS proresearch

² only clinics of the AKHH



	AKHH 2017	AKHH 2018	AKHH 2019
DMQ*	72%	70%	77%
Pool money (n/a)	n/a	n/a	n/a
AMS apprenticeship premiums	470.293,75 €	450.509,19 €	463.277,82 €
Research funding	710.000,00 €	741.068,00 €	310.510,00 €
PROMEDIG	531.756,47 €	472.800,55 €	477.002,07 €
Event administration	399.266,03 €	219.364,36 €	317.490,87 €
Contract research	2.172.549,68 €	2.145.131,39 €	1.834.355,09 €

Figure 4: Total research revenue AKHH from 2017 to 2019



	AKHH total	AKS	AKA	AKW	AKH	AKB	AKW	AKN	SONS (incl. MVZ)
DMQ*	77,3%	85,1%	91,7%	93,6%	88,3%	41,2%	69,3%	23,4%	90,9%
Pool money (n/a)	0,00 €	0,00 €	0,00 €	0,00 €	0,00 €	0,00 €	0,00 €	0,00 €	0,00 €
AMS apprenticeship premium	463.277,82 €	74.845,30 €	62.960,54 €	15.873,74 €	63.486,81 €	84.450,33 €	40.449,90 €	94.327,34 €	26.883,86 €
Research funding	310.510,00 €	0,00 €	15.000,00 €	0,00 €	0,00 €	218.000,00 €	0,00 €	77.510,00 €	0,00 €
PROMEDIG	477.002,07 €	174.783,91	197.548,33	9.000,00 €	45.936,63	14.815,13	29.595,10 €	5.322,97	0,00 €
Event administration	317.490,87 €	29.929,51 €	80.450,29	3.106,37 €	68.558,59 €	19.895,27 €	1.973,26 €	42.209,83 €	71.367,75 €
Contract research	1.834.355,09 €	222.403,89 €	586.366,48 €	221.507,12 €	363.666,88 €	177.318,01 €	59.942,88 €	4.873,53 €	198.276,30 €

Figure 5: Total research revenue AKHH 2019, distribution by hospitals of the AKHH

*DMQ = Share of external revenues in total revenues = total revenues minus research funding, AMS and pool funds

STAFF AND SCIENTIFIC PERSONNEL

Staff financed by research funds are divided into different categories. Our study coordinators, with 15.4 full-time employees (FT), continue to make up the majority. These staff are employed exclusively on research projects, and neither involved in routine care nor included in the indi-

vidual clinics' or departments' personnel budget. Staff who otherwise support the research projects in the departments can be financed by proresearch in two ways: by cost absorption (4.5 FT) or by direct employment through ASKLEPIOS proresearch (scientific staff: 5.2 FT).

	Total pro rata FT 2019
Study coordinators	15,4
Cost absorption	4,5
Scientific staff	5,2
Total	25,1

Table 3: Aggregated personnel costs FoBe 2019, AKHH

IN SEARCH OF A SUITABLE ABLATION THERAPY FOR PATIENTS WITH RECURRENT ATRIAL FIBRILLATION AFTER INITIAL PULMONARY VEIN ABLATION

DR. PAWEL KUKLIK
DR. CHRISTIAN EICKHOLT
PROF. DR. STEPHAN WILLEMS

Catheter ablation of atrial fibrillation (AF) has become first-line therapy in patients unresponsive to electrical or pharmacological cardioversion. In case of paroxysmal AF, pulmonary vein isolation (PVI) aiming at complete electrical isolation is recommended due to high acute and long-term success rates. However, once AF becomes persistent, ablation efficiency dramatically decreases, most likely due to advanced electrical and structural remodeling of the atria (1) resulting from unaddressed risk factors. This leads to the recurrence of AF in patients who previously underwent a PVI procedure. Such patients are scheduled for a consecutive procedure during which the quality of the initial PVI is assessed and gaps in ablation lesions are corrected. However, a considerable proportion of patients remains in AF despite complete and verified PVI. There, two approaches can be considered: either to perform electrical cardioversion and terminate the procedure, or, to continue ablation within the atrial body searching for drivers maintaining arrhythmia (2,3).

Some studies, most notably STAR AF II (4), suggested that additional ablation beyond PVI may not benefit patients as expected. However, several studies explored promising novel techniques, e.g. high frequency source ablation (5), electrogram complexity-guided ablation, low-voltage amplitude targeting (6), activation-dispersion-guided ablation (7), stepwise ablation approach (8,9) and recently proposed techniques of identification of rotating waves and point sources (10), to name just few.

Such a great variety of approaches may stem from the heterogeneity of the atrial pathophysiological phenotype presented during the ablation procedure (so called 'AF substrate'). Some patients in an early stage of electrical and structural remodeling of the atria may have localized sites maintaining arrhythmia with a clearly defined pattern of electrical activity – then, localized ablation can

remove the AF driver and benefit the patient in the long term. However, patients with an advanced stage of remodeling may lack such clearly defined spots and may require more extensive ablation or, in case of very advanced remodeling, the abandonment of further ablation attempts. Since additional catheter ablation is related with a decrease of atrial function and an increased risk of procedural complications, it is important to identify an optimum approach to maximize patient benefit and minimize the extent of unnecessary catheter ablation.

The most important imaging technique used to assess the degree of structural and electrical remodeling of the atria is 3D electro-anatomical mapping. During such mapping, hundreds of electrograms are recorded across the entire atrial endocardial surface. Then, these electrograms are processed to extract various parameters shedding light on the state of the underlying myocardial tissue. One of the most obvious parameters is the amplitude of the electrical signal collected at a given site. Diseased, fibrotic tissue generates low-amplitude electrical activity, whereas healthy, bulky areas generate a high amplitude. In our approach, we look at several electrogram-derived parameters such as the degree of electrogram morphology "fractionation" (see Figure 1), presence of rotating waves or percent of time the electrical activity is present in small area of the atrium. Those parameters are then projected onto the 3D geometry of the atria and used to identify targets for ablation (see Figure 2).



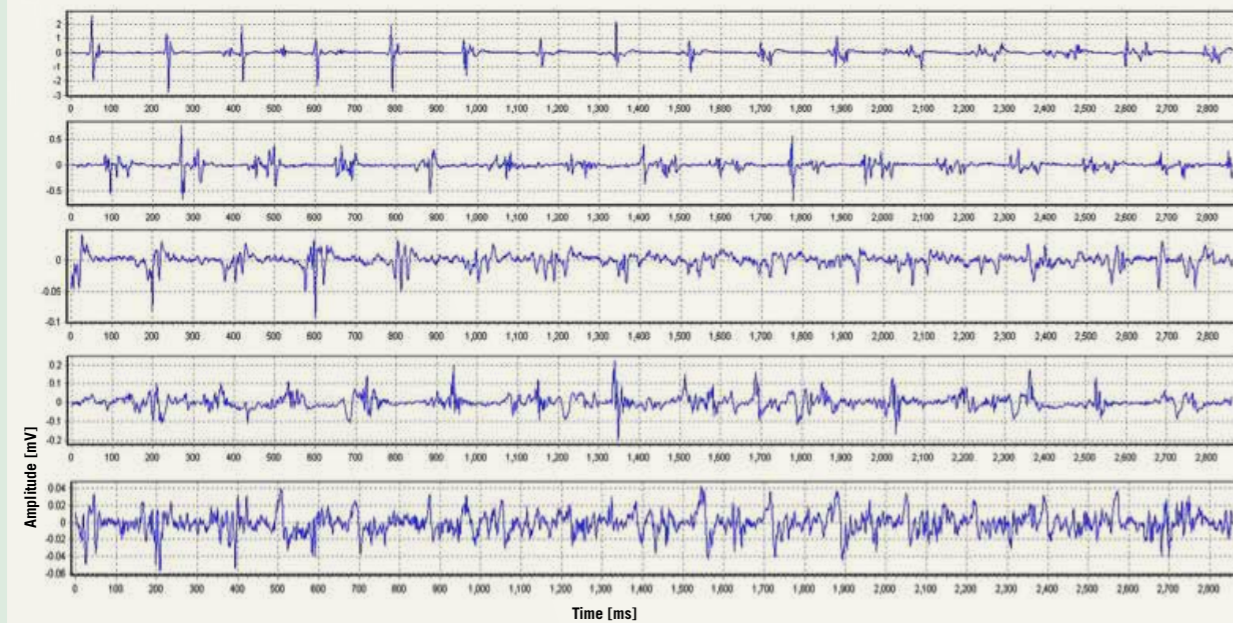


Figure 1 Sample intracardiac electrograms collected during Atrial Fibrillation with a progressive degree of morphology fractionation.

Our goal is to create a patient-tailored ablation strategy that would consider various biomarkers (ECG parameters, intracardiac electrograms, AF history, etc.) and suggest an optimal catheter ablation strategy (see Figure 3). Our work focuses on:

- the development of signal processing algorithms extracting various complexity-related parameters from ECG and intracardiac electrograms,
- the identification of the clinically most relevant parameters in the context of catheter ablation,
- the estimation of the thresholds in decision trees leading to optimal ablation strategy and

To evaluate the long-term efficacy of developed ablation strategies high-quality clinical trials are necessary, which are designed and conducted at our site. One important

project was planned together with the Asklepios Campus Hamburg. The start of this multicenter prospective randomized study will be in 2020. We are looking forward to the results.

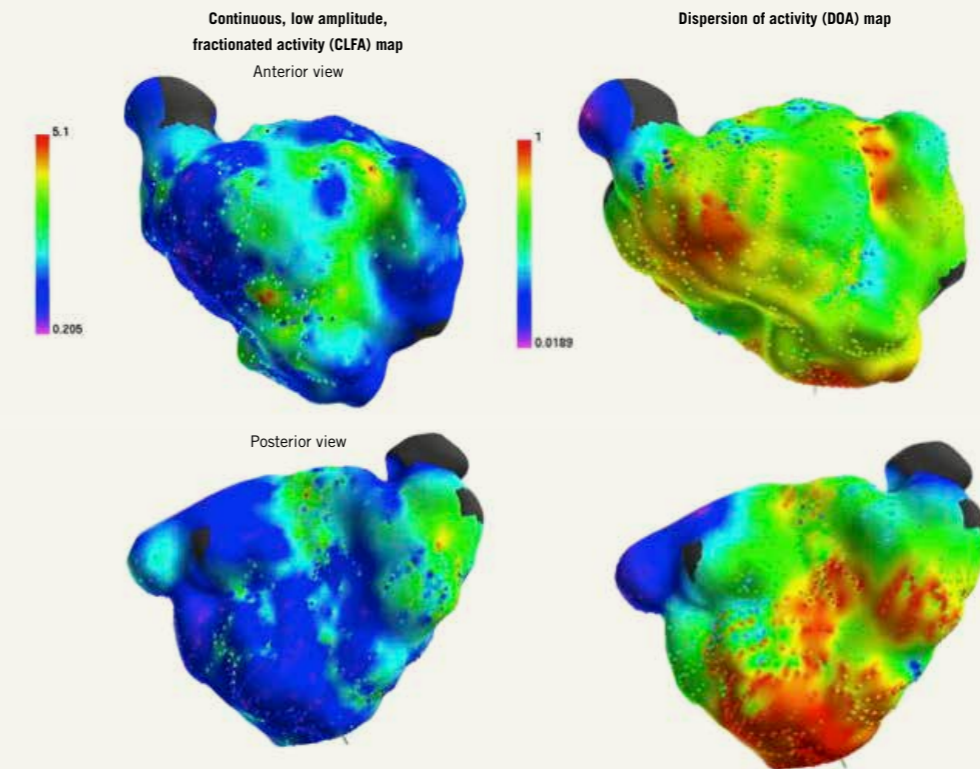


Figure 2 Left Atrium. Sample 3D maps of parameters related with arrhythmia complexity. Areas of the greatest complexity (red) are targeted during catheter ablation.

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The references have been abridged for space reasons. For a complete list of the literature used please contact the author.

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LIST OF ABBREVIATIONS

ACH	Asklepios Campus Hamburg of Semmelweis University Budapest
MD	Medical Direktor
AKHH	Asklepios Kliniken Hamburg GmbH
AMG	Arzneimittelgesetz – German Medicines Act
AMS	Asklepios Medical School gGmbH
DMQ	Drittmittelquotient – Third party funding ratio
n/a	not applicable
CD	commercial director
MPG	Medizinproduktegesetz – German Medical Devices Act
FT	fulltime employee