

Department of Cardiac Surgery

Chair of Cardiac Surgery

Address

Krankenhausstraße 12
91054 Erlangen
Phone: +49 9131 8533319
Fax: +49 9131 8532768
www.herzchirurgie.uk-erlangen.de

Director

Prof. Dr. med. Michael Weyand

Contact

Prof. Dr. med. Michael Weyand
Phone: +49 9131 8533319
Fax: +49 9131 8532768
herz-sekretariat@uk-erlangen.de

Research Focus

- Chronic rejection of allografts
- Therapy of end-stage heart failure: Heart transplantation or support with a left or right ventricular assist device
- Electromechanical coupling in heart failure
- Development of a non-blood contacting heart actor
- High speed camera investigations on heart valves in a pulse duplicator

Structure of the Department

Professorships: 1

Personnel: 100

- Doctors (of Medicine): 15
- Scientists: 4 (thereof funded externally: 1)

Clinical focus areas

- Adult cardiac surgery
- Heart transplantation in adults and children
- Minimally invasive valve surgery
- Mechanical circulatory support
- Wound management
- Heart insufficiency therapy
- Rhythm surgery
- Surgery in grown-up with congenital heart disease
- Interventional heart valve surgery
- Interventional aortic surgery

Research

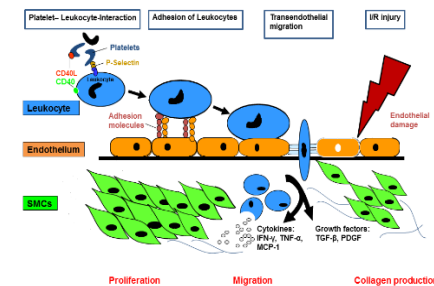
Main research topics are on the one hand basic research in transplantation and on the other hand clinical research in mechanical circulatory support and the development of new heart assist devices in close cooperation with the Faculty of Engineering.

Chronic rejection of allografts

PI: PD Dr. C. Heim

Transplant vasculopathy is the main reason for late graft failure after heart transplantation. In order to develop effective therapeutic strategies and translate them into clinical success, a detailed understanding of the mechanisms responsible for the development of transplant vasculopathy is essential. We have recently established and characterized the abdominal aortic allograft model as a suitable tool to study the development of transplant vasculopathy. Ongoing projects involve

the role and importance of platelets and their inhibition in the development of transplant vasculopathy. Immunomodulatory effects of Clopidogrel could be shown in small animal models. The results of these preclinical studies could be translated into a multi-center study (CEDRIC). Additionally, microvascular integrity of pulmonary grafts was shown to be essential for the long-term success of animal transplant models. In cooperation with the Department of Medicine, another major aim of this working group is the use of antiproliferative substances to explore potential strategies to avoid the development of transplant vasculopathy in experimental transplant models. Additionally an ISHLT grant investigates the serostatus of cytomegalovirus as risk factor for transplant vasculopathy in a international heart transplant register.



Mechanisms in CAV

After I/R injury, endothelial damage is likely to occur. Adhesion molecules are upregulated and after platelet-leukocyte interaction, leukocytes transmigrate through the endothelial layer. There they produce several cytokines and growth factors. As a result, SMC produce collagen, proliferate, and migrate into the neointima. This finally leads to a progressive narrowing of the transplanted vessels and to subsequent graft failure. CAV, cardiac allograft vasculopathy; IFN- γ , interferon- γ ; I/R, ischemia/reperfusion; MCP-1, monocyte chemoattractant protein-1; PDGF, platelet-derived growth factor; SMC, smooth muscle cell; TGF- β , transforming growth factor- β ; TNF- α , tumor necrosis factor α . (Reproduced from Heim et al., Thorac Cardiovasc Surg 2018, with permission from Thieme)

Therapy of end-stage heart failure: Heart transplantation or support with a left or right ventricular assist device

PI: Dr. R. Tandler

Orthotopic cardiac transplantation is the therapy of choice for cardiac insufficient patients. Due to an increasing shortage of donor organs, these cardiac insufficient patients need to be bridged with an implantable ventricular assist device until a suitable donor organ is available. In some cases heart disease has already progressed to such an extent that the patients need to be stabilized with a left ventricular assist device or – in case of additional right heart failure – with a biventricular assist device.

Electromechanical coupling in heart failure

PI: PD Dr. C Heim

Remodelling of cardiomyocytes in heart failure patients is well described in the literature, but not completely understood. The calcium delivery in cardiomyocytes may be altered in heart failure patients. In previous studies the remodeling of the T-system of the cardiomyocytes was discussed as responsible for cardiac recovery in ventricular assist device patients. Therefore the aim of the ongoing projects in cooperation with the Institute of Physiology and Pathophysiology is to further analyze underlying mechanisms of the T-system remodeling using human heart tissue from VAD or transplant patients.

Development of a non-blood contacting heart actor

PI: Prof. Dr. M. Weyand

The support of the insufficient heart muscle function by artificial support systems is worldwide an intensive field of research and an aim sought for for about 60 years. Rising life expectancy and the growing number of heart-insufficient patients on the one hand as well as restricted availability of donor organs and damping of the increase of the health costs will further raise the need in innovative support systems in the future. On account of the risks of the existing, invasive, clinical methods, a carefully implantable technology is necessary. It must be functioning reliably as well as permanently and intervene not invasive in the heartcirculatory system. Within a clinical-medical setting, the investigation of a research project pursues from the interpretation over the production up to the clinical validity of the system function more new, actoric, and patient-individual heart muscle support systems for the purposes of an external compression of the heart. Therefore the main focuses are the investigation of a biomechanically efficient, mechanical system as well as the development of dior piezoelectric based actor material patterns.

High speed camera investigations on heart valves in a pulse duplicator

PI: Dr. M. Kondruweit

High-speed camera investigations on heart valves in an animal model are an already established model. In this project these proceedings are applied into a pulse duplicator to be able to compare several heart valve types in a standardized procedure. Special situations, as for example the Ventricle Assist Devices support and the effect on the hemodynamic on the heart valves, are examined. The results should show possible reasons for heart valve attrition by measuring power vectors. If possible, these reasons shall be corrected by changing the valve design.

Teaching

The Department of Cardiac Surgery takes part

in compulsory and elective subjects for the curricular teaching of the human medicine and dentistry.

Bachelor's and Master's theses especially from the Faculty of Engineering are supervised as well as MD and PhD theses.

Selected publications

Heim C, Kuckhahn A, Ramsperger-Gleixner M, Nicolls MR, Weyand M, Ensminger SM. Microvasculature in murine tracheal allografts after combined therapy with clopidogrel and everolimus. *Interact Cardiovasc Thorac Surg.* 2020

Fritz NM, Stamminger T, Ramsperger-Gleixner M, Kuckhahn A, Müller R, Weyand M, Heim C. Cytomegalovirus chemokine receptor M33 knockout reduces chronic allograft rejection in a murine aortic transplant model. *Transplant Immunology.* 2020

Rivinius R, Kaya Z, Boeken U, Provaznik Z, Heim C, Knosalla C, Schoenrath F, Rieth A, Berchtold-Herz M, Barten MJ, Rauschnig D, Grinninger C, Warnecke G, Schulze PC, Katus HA, Kreusser MM, Raake PW. COVID-19 among heart transplant recipients in Germany: A multicenter survey. *Clinical Research in Cardiology.* 2020; 109(12):1531-1539

Abu-Khousa M, Fiegler DJ, Sommer ST, Minabari G, Milting H, Heim C, Weyand M, Tomasi R, Dendorfer A, Volk T, Seidel T. The degree of t-system remodeling predicts negative force-frequency relationship and prolonged relaxation time in failing human myocardium. *Frontiers Physiology.* 2020; 11:182

Seidel T, Fiegler DJ, Baur TJ, Ritzer A, Nay S, Heim C, Weyand M, Oakley RH, Cidlofski JA, Volk T. Glucocorticoids preserve the t-tubular system in ventricular cardiomyocytes by upregulation of autophagic flux. *Basic Res Cardiol.* 2019; 114(6):47

Heim C, Khan MA, von Silva-Tarouca B, Kuckhahn A, Stamminger T, Nicolls MR, Weyand M, Ensminger SM. Preservation of microvascular integrity in murine orthotopic tracheal allografts by clopidogrel. *Transplantation.* 2019; 103:899-908

International cooperations

Dr. M. Nicolls, Professor in Pulmonary and Critical Care Medicine, Stanford University, CA: USA

Dr. J. Stehlik, Director of ISHLT Registry, University of Utah, UT: USA

Dr. K. Kush, Professor in Cardiology and Cardiac Transplantation, Stanford University, CA: USA