

GERMAN CANCER RESEARCH CENTER (DKFZ)

DKFZ Clinician Scientist Program

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supported by the





DKFZ Clinician Scientist Program Fellows Brochure

FROM BEDSIDE TO BENCH AND BACK.

TRANSLATING RESEARCH CAREERS





Scientist Fellowships we provide the much needed time for Clinician Scientists in training to focus on a research project for an extended period of time. This will foster the uptake of new research results into the clinical practice.

The DKFZ is one of the leading cancer research centers worldwide and provides a nurturing environment in which translational cancer research can thrive. We have a strong and long-standing reputation of training excellent scientists at all career stages, and our research programs naturally integrate both basic and clinical research. In order to directly support researching clinicians in their professional development, we have established a dedicated program to foster the careers of Clinician Scientists at DKFZ and beyond. While they are part of the DKFZ Clinician Scientist Program, our Fellows receive state-of-the art research training to work on interdisciplinary projects at the interface of basic and clinical oncology research.

The protected time at our Center allows them to exclusively focus on a research project that is relevant for their medical specialization. The Fellows gain hands-on laboratory experience, enhance their portfolio of research skills and establish their own network within the scientific community. The Fellows are therefore well equipped to maintain the ongoing transition

between the two worlds and ensure a fruitful exchange between the disciplines.

From my own experience I know that the treatment of patients and parallel lab work are not always easy to combine. However, I am convinced that in addition to the direct impact on scientific output, there is also a great personal benefit associated with the profession as a Clinician Scientist. I am therefore very happy that, due to the great donation of the Dieter Morszeck Foundation, we are in a pole position to support the career development of the future generation of Clinician Scientists.

We are truly impressed by the high level of qualification, enthusiasm and commitment to research that our first cohort of Fellows have shown since being competitively selected. Therefore, I want to cordially invite you to find out more about the Fellows 2019 and their individual research projects on the following pages of this brochure.

Prof. Dr. med. Michael Baumann Chairman and Scientific Director German Cancer Research Center

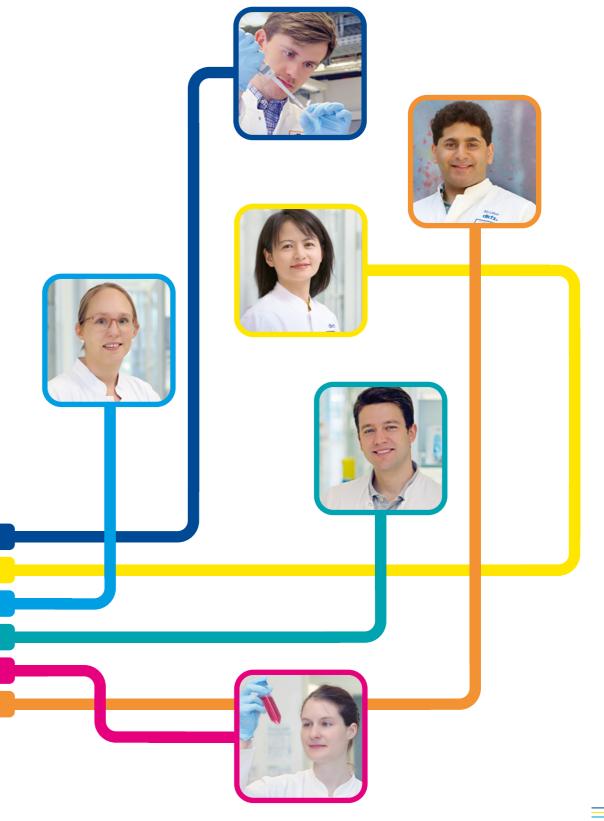




CONTENTS

Clinician Scientist training at the German Cancer Research Center **Translational Cancer Research** 7 The DKFZ Clinician Scientist Fellows 2019 8 Maximilian Blank 8 Xiangli Jiang 10 Laura Klotz 12 Raphael Lutz 14 Verena Panitz 16 Mohammad Rahbari 18

20



Imprint

Clinician Scientist training at the **German Cancer Research Center**

More than 450,000 people are diagnosed with cancer each year in Germany. Cancer is a disease that poses enormous challenges to research, because every cancer is different and its course can vary immensely even from one patient to the next. To perform research into cancer is the task of the German Cancer Research Center (Deutsches Krebsforschungszentrum, DKFZ) according





From bedside to bench and back – the DKFZ Clinician Scientist Program addresses physicians who show enthusiasm for science and who have a wish to link basic and clinical research.

German Cancer Research Center's official program to foster innovative translational research by strengthening the research profile of clinician scientists. The program deepens the link between basic and clinical research, ensuring efficient transfer from bedside to bench and back.

We aim to recruit highly talented physicians in training who wish to gain basic and translational cancer research experience in a worldclass scientific setting. Through the research groups in DKFZ's Research Programs we provide access to cutting-edge research facilities and state-of-the-art labs.

The DKFZ Clinician Scientist Program is the Program fellows are supported by dedicated fellowships encompassing protected research time, tailored training and intersectoral mentoring. Extensive network-building and continued institutional support lead the way to long-term career perspectives.



Information on the DKFZ Clinician Scientist Program, upcoming calls for application and eligibility criteria can be found at:

www.dkfz.de/clinicianscientist

Contact: Dr. Philipp Gebhardt **DKFZ Clinician Scientist** Program







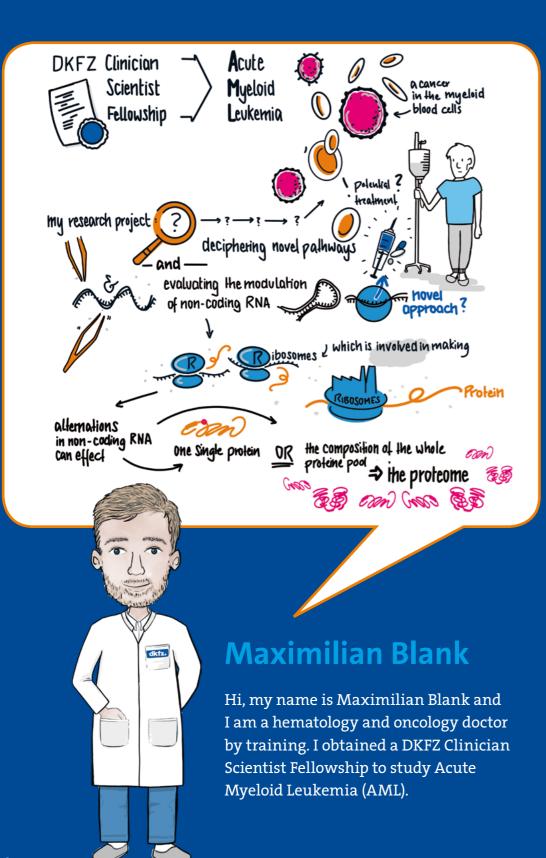
Translational Cancer Research at the DKFZ

The DKFZ is a prime location to pursue basic and translational oncology research in one of its six research programs.

In order to ensure that novel results from the "lab bench" and newly developed methodologies can be put into practice more quickly for the benefit of future patient care, we draw on a wide range of knowhow and highly specialized, networked infrastructures. This is complemented by the close collaboration of DKFZ researchers and clinicians, such as the DKFZ Clinician Scientist Fellows, who are performing cutting-edge research on a plethora of key oncology topics.

The pursuit of translation is, so to speak, embedded in our institutional DNA and we are constantly striving to further develop career options for the bright minds driving the innovative research projects DKFZ is known for.







incompletely understood. Therefore, deciphering important pathways in the progress of AML might reveal novel and improved treatment options, especially with regard to the ability to induce remission and to prevent relapse of this severe disease.

Clinical work is closely intertwined with both clinical and basic research. The DKFZ Clinician Scientist Fellowship offers me the unique opportunity to perform cutting-edge research while still pursuing my residency in the Department of Hematology and Oncology at the University Hospital Heidelberg. And my work will hopefully result in the identification of new therapeutic insights that will help us to translate our findings from bench to bedside.



"The DKFZ Clinician Scientist Fellowship offers two years of protected research time and I am flexible to intersperse clinical work if needed. It provides an excellent opportunity to pursue my research project with the aim to contribute to the better understanding of pathogenetic mechanisms in AML."

that are not directly translated into proteins. The ones I am focusing on are involved in the process of making ribosomes, the cellular factories where proteins are synthesized. Alterations of ncRNAs could thus affect synthesis of individual proteins and even the overall composition of the pool of proteins that are produced by a cell or an organism. The latter is called the proteome. To date, the mechanisms underlying

Acute Myeloid Leukemia (AML), is

a cancer of the myeloid blood cells.

Besides deciphering novel pathways in the development of leukemia, my

research project aims to evaluate the

modulation of specific non-coding

RNAs – which could potentially be

used as a novel therapeutic approach

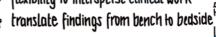
for the treatment of AML. Non-

coding RNAs (ncRNAs) are RNAs

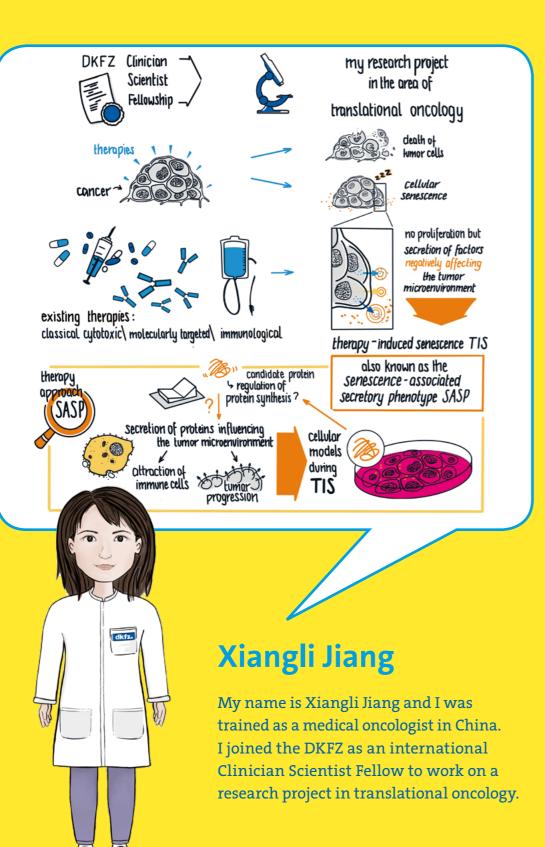
formation of blood cancer remain



2 years of protected research time flexibility to intersperse clinical work







Cancer therapies are commonly applied to induce the death of tumor cells. But, in addition, they can also trigger so-called cellular senescence. This is a cellular state in which cancer cells are not proliferating but they can still secrete factors that might negatively affect the direct environment of the tumor – the tumor microenvironment. Known as therapy-induced senescence (TIS), this state can be induced by treating cancers with therapies such as classical cytotoxic therapies (e.g. chemotherapy), molecularly targeted therapies and also immunotherapies. The resulting senescent cells are characterized by the secretion of high amounts of protein. This phenomenon is known as the senescence-associated secretory phenotype (SASP). The secreted proteins influence the tumor microenvironment by attracting immune cells and are even promoting tumor progression. Therefore, targeting the SASP could be a promising approach for cancer therapy. In this research project I will study, together with colleagues, how senescent cells synthesize large amounts of protein despite being in

found that a protein involved in the regulation of protein synthesis – its name is eIF5A - might be a critical player in this process. Using cellular models and cutting-edge approaches, we will characterize the role of this molecule during TIS. Through this research project, we hope to potentially uncover a novel molecular pathway that can be exploited for a more effective cancer treatment. The DKFZ Clinician Scientist Fellowship directly supports my ambitions to gain international research experience and to build up my personal research network. In the era of precision medicine and rapidly evolving medical science, physicians not only need to have profound experience in clinical practice, but also need a deep understanding of basic research. The fellowship is therefore helping me to further deepen the link between clinical and basic cancer research in a world-class scientific setting. I am convinced that this experience will have a significant impact on my long-term career perspective as a



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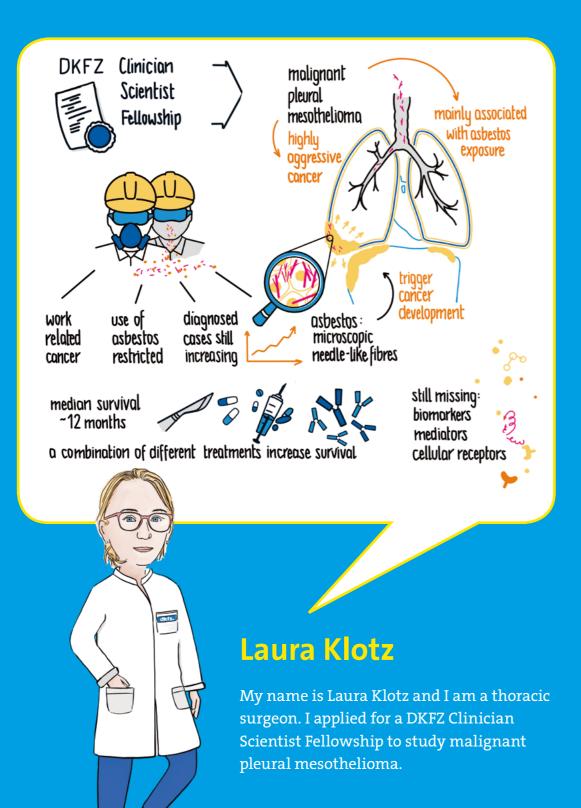


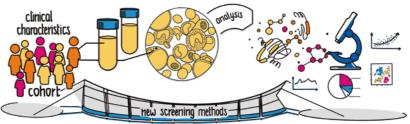
an arrested state. Previous research

A gaining international research experience

clinician scientist.

- a deepening the link between clinical and basic research
- ⇒ building a personal research network





the survival rate for patients with early stage disease.

More detailed analyses of clinical parameters and the correlation with molecular biomarkers might help to improve the knowledge of the complex pathobiology of the disease and to refine current treatment possibilities. To achieve this, it is essential to identify mediators and cellular receptors that play a crucial role for tumor development and tumor growth.

Hence, the DKFZ Clinician Scientist for our Fellowship allows me to link the clinical characteristics of our large patient cohort to biomarkers which are identified by analyzing human tissue and blood samples. Within this translational research approach, my aim is to identify biomarkers that might be relevant for screening high-risk patients after work-related asbestos exposure and to predict their response to treatment.



"By building a bridge from thoracic surgery to basic research, I aim to learn cutting-edge methods and techniques in order to optimize future treatment possibilities mesothelioma patients."



Pleural mesothelioma is a locally

invasive and highly aggressive cancer

of the thorax. This tumor entity is

mainly associated with asbestos

exposure and actually represents

a common work-related cancer.

Despite the restricted use of asbestos

in the industry in recent years, the

number of newly diagnosed cases is

still increasing. Asbestos consists of

fibrous minerals that are composed

of microscopic, needle-like fibers.

It can trigger the development of

mesothelioma up to 50 years after

exposure. Unfortunately, the median

survival for unresectable disease is

about 12 months. To date, no specific

biomarker has been identified for

screening or early diagnosis – which

would in turn enable treatment at an early stage of the disease

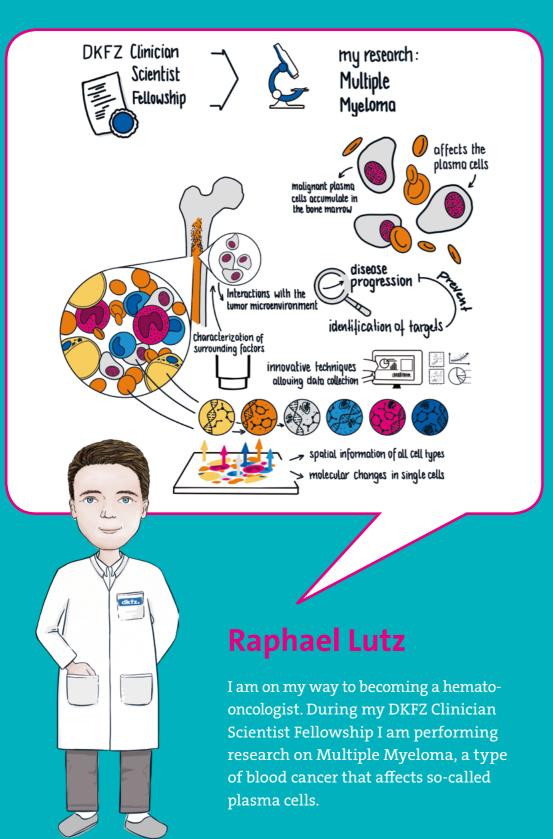
and therefore increase survival. A

combination of surgery, chemotherapy

and potentially immunotherapy

plays an important role in increasing

- DKFZ Clinician & build a bridge between thoracic surgery and basic oncology research Scientist acquire new technical skill and expand research methods portfolio
 - Fellowship & gain better understanding of cancer biology to improve treatment of mesothelioma patients





During my DKFZ Clinician Scientist

Fellowship I am performing research

on Multiple Myeloma (MM), which

is a type of blood cancer that affects

so-called plasma cells. Malignant

plasma cells typically accumulate in

the bone marrow. In recent years it

became clear that the interactions

of these specific cells with the

tumor microenvironment play an

important role in the progression of

Thus, there is an urgent need to

deeply characterize the relationship

of plasma cells with surrounding

factors in order to gain a global

understanding of MM development.

We are currently using innovative

techniques that enable us to

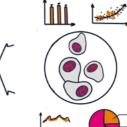
describe molecular changes in MM on the level of single cells and at

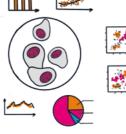
the same time keep the spatial

information of all cell types within the

bone marrow. This approach has the

the disease.





potential to unravel the key drivers of MM pathogenesis on a cellular and molecular level. A global and profound understanding of these processes might subsequently lead to the identification of drug targets in order to prevent disease progression.

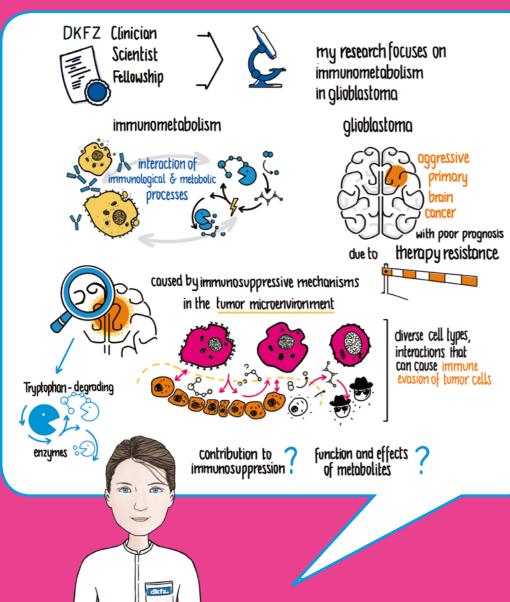
The fellowship provided by the DKFZ Clinician Scientist Program gives me the opportunity to perform state of the art translational research. My goal is to generate scientific results that will have a major impact on future patient outcomes. In the future, I want to combine my expertise in clinical and translational research and develop novel therapeutic strategies to prevent disease progression of Multiple Myeloma. I am confident that this research can contribute to our ultimate goal of finding a cure for this specific cancer entity.



"The fellowship provided by the **DKFZ** Clinician Scientist Program gives me the opportunity to perform state of the art translational research."



- A opportunity for state-of-the-art translational research
- impact on future patient outcomes
- A combination of expertise in clinical and basic research



During my Clinician Scientist Fellowship at the DKFZ my research focuses on the complex topic of immunometabolism in glioblastoma.



research that is dedicated to the interaction of immunological and metabolic processes. Glioblastoma is the most common and also most aggressive primary brain cancer in adults with a median overall survival of only 15 months after diagnosis despite current standard of care.

One reason for this very poor prognosis is the glioblastoma's resistance to therapy, to which immunosuppressive mechanisms in the tumor microenvironment contribute relevantly. The tumor microenvironment consists diverse cell types, including different immune cell subtypes. Metabolic processes play a relevant role in the interaction between immune cells and tumor cells as they can, for example, foster the immune evasion of the tumor cells, i.e. the tumor cells escape the control by the immune system.

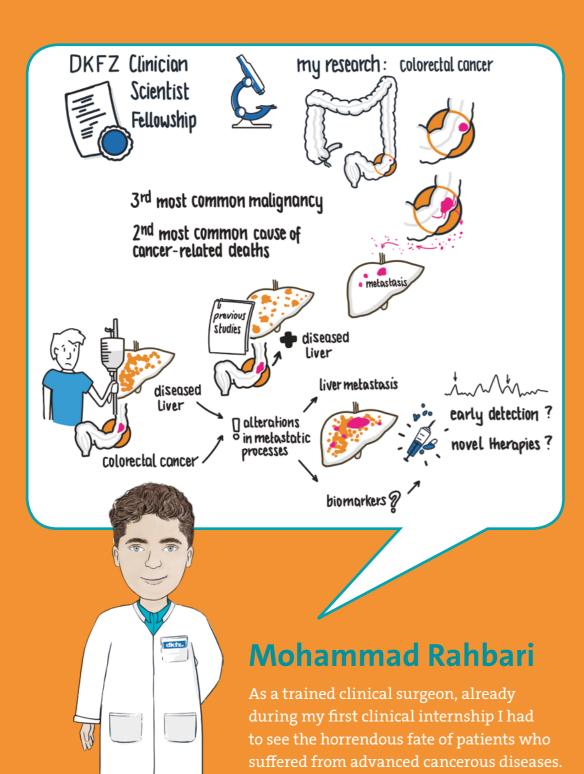
Immunometabolism is a field of Therefore, my special interest lies in the regulation of tryptophandegrading enzymes in glioblastoma cells and in different immune cell subsets, as these enzymes contribute to immunosuppression in the tumor microenvironment. I also want to study the metabolites that are produced by the metabolic enzymes and their effects and functions.

On the clinical side, I am training as a resident in Neurology at Heidelberg University Hospital. The DKFZ Clinician Scientist Fellowship provides me with the opportunity to conduct research over a timespan of two years. During this protected research time I am able to specifically focus on my research topic immunometabolism in glioblastoma. In the future, I hope that a better understanding of the resistance mechanisms in glioblastoma will help to improve treatment of glioblastoma patients.

"The concept of a career as a Clinician Scientist fascinates me because it allows the deep integration of clinical and scientific knowledge and experience and hence facilitates the translation from bench to bedside. "



- A Integration of clinical and scientific knowledge and experience
- ☆ Focus on research project over a two-year period
- ☆ Translation of findings for the well-being of cancer patients



This motivated me to focus my career on

the improvement of clinical cancer care.



State day.

Pursuing my career, I soon realized that it will be more than helpful to integrate the perspective of a basic scientist into my clinician's viewpoint. Thus, I decided to dive deeper into the biology of cancer initiation, progression and finally metastasis, the process which ultimately causes the death of cancer patients.

In my research project I am working on colorectal cancer. This cancer develops from the colon or rectum and it represents the third most common malignancy and the second most common cause of cancerrelated deaths worldwide. Patients suffering from colorectal cancer die of cancer manifestations at distant organs, so-called metastasis. The most frequent organ in which metastases occur is the liver. Previous studies have shown that

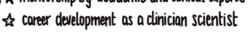
patients with colorectal cancer often concurrently suffer from fatty liver disease. Fatty liver disease is characterized by the accumulation of excess fat in the liver, which can lead to chronic inflammation and tissue damage.

Most importantly, patients who suffer at the same time from colorectal cancer and fatty liver disease seem to have alterations in the processes leading to metastasis in the liver. We aim to better understand the influence of fatty liver disease on liver metastasis, and may therefore not only be able to develop new clinical biomarkers for early detection, but also novel therapies to treat this devastating disease. In this regard, the DKFZ Clinician Scientist Fellowship represents a unique opportunity for me to combine both worlds.

"Dedicated mentorship by selected academic and clinical experts will help me to further develop my career as a clinician scientist."



☆ combination of both worlds ☆ mentorship by academic and clinical experts





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TRANSLATING RESEARCH CAREERS





www.dkfz.de/clinicianscientist

DKFZ
Clinician Scientist
Fellowships

Are you considering a career as a clinician scientist?

The DKFZ in Heidelberg belongs to the leading cancer research centers worldwide. Team up with a strong partner to support your translational research ambitions!

Expand your research experience and complement your clinical training with a DKFZ Clinician Scientist Fellowship featuring protected time for research, comprehensive training and mentoring. You are are a physician conducting specialized medical training and wish to enhance your career? Apply now!

More info at www.dkfz.de/clinicianscientist

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Published by

German Cancer Research Center (DKFZ) in the Helmholtz-Association

Edited by

German Cancer Research Center DKFZ Clinician Scientist Program Im Neuenheimer Feld 280 69120 Heidelberg, Germany clinicianscientist@dkfz.de ww.dkfz.de/clinicianscientist

Editorial responsibility Dr. Philipp Gebhardt

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Infographics:

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Design concept and layoutDagmar Anders

Printed by

CITY-DRUCK Offsetdruck GmbH

© Deutsches Krebsforschungszentrum (DKFZ), Heidelberg, 2020

Image credits

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