





# KERIC

KERIC is a core facility for preclinical imaging and surgery at Karolinska.

Equipped with 9.4T MRI, PET-CT and PET-MR for small animal applications.

Angiography-CT and fully equipped Surgery room for several large animal application.

We have Application Scientists for running the equipment and assist with data handling.

Animal Technicians with expertise in the above modalities that can assist in the procedures.

We can **Host animals** direct or indirect with collaboration within KU (AKM).

https://ki.se/keric





### MRI

Name: Magnetic Resonance Imaging

MR, magnetic gradients and RF Source: 22.6MHz @ 1T, 400MHz @ 9.4T

Soft tissue: brain, organs, fat, Targets: blood. Other nuclei (19F)

Bone, metal, regions close to material with different magnetic properties.

> Can show diffusion and flow, anatomical, paramagnetic, functional MRI (fMRI)

### CT

X-ray, gamma

Computed Tomography CT (x-ray)

Hard tissue: bone, vague organ images

Water, not best for soft tissue

Relatively fast, can be used with contrast to view veins and organs, good contrast for hard matters, good with metals.

### PET

Radioactivity

Position Emission Tomography PET (radioactive substances)

Only radioactive source. Requires a reference images.

Only detects radioactive sources

Can view uptake and distribution. Can follow over time. Wide range of ligands.

Cannot detect:

Other benefits:

# **MRI 9.4T**

Preclinical 9.4T MRI equipped with three gradients with different bore sizes.

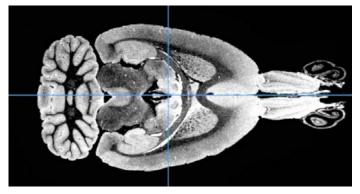
20 cm bore size for large larger animals such as rabbits and small piglets.

12 cm bore size for medium size animals such as rats.

6 cm bore size for smaller animals such as mice.



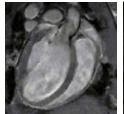




Rat brain (cordinal view)

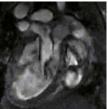
We can do anatomical imaging of brain, spine, heart, legs or whole body in various animal models. Arterial Spin Labelling, (ASL) in rabbit brain applications, fMRI, pharmacological MRI and cardiac imaging are established applications.

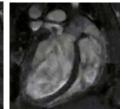
With monitoring system to follow biological data allow us triggering scanning to allow images with minimal motion artefacts for breathing or heart beat.











Rat heart. (reduced framerate)



We are installing a new 9.4T MRI system (Q4 2022)

The system will have a cryo-coil for mouse brain applications and keep us updated with that latest experiments and features with preclinical MRI

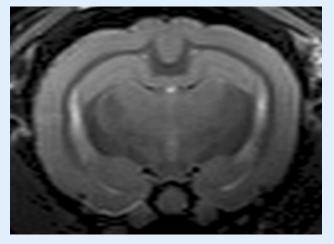
# **MRI 9.4T**



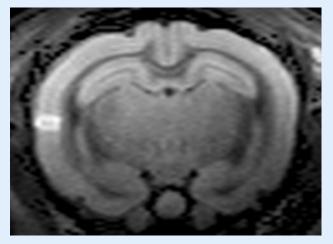
1000



### Rabbit brain. Induces stroke model

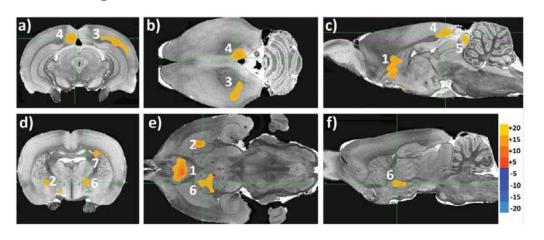


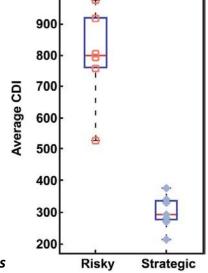
S0, T2 image (Standad MR image)



Difusion Weighted MR image

### Gambling behaviour in rat model studied with fMRI





Tjernström N, Tie-Qiang L, Holst S, Roman E. Functional connectivity in rewardrelated networks is associated with individual differences in gambling strategies in male Lister hooded rats. Addiction Biology. 2022 Jan 17;27(2):e13131 link

### Parkinson study in rat model with treatment of L-DOPA investigated with fMRI and dwMRI

Monnot, C., Zhang, X., & Svenningsson, P. et. Al. (2017). Asymmetric dopaminergic degeneration and levodopa alter functional corticostriatal connectivity bilaterally in experimental parkinsonism. Exp Neurol, 292, 11-20. <u>link</u>

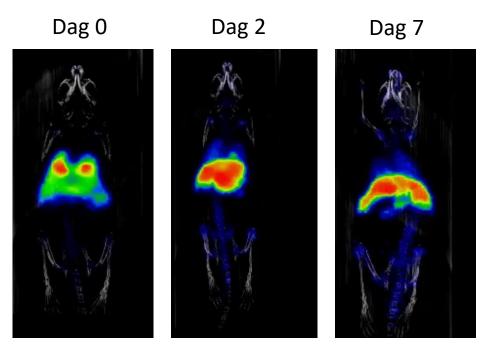
### Comparing rat model to human model with fMRI

Sierakowiak, A., Monnot, & Brené, S. et. Al. (2015). Default mode network, motor network, dorsal and ventral basal ganglia networks in the rat brain: comparison to human networks using resting state-fMRI. PLoS One, 10(3), e0120345. link

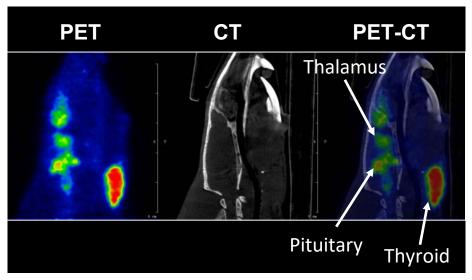
# PET-CT / PET-MR 2x PET-CT and 1 PET-MR (1T)



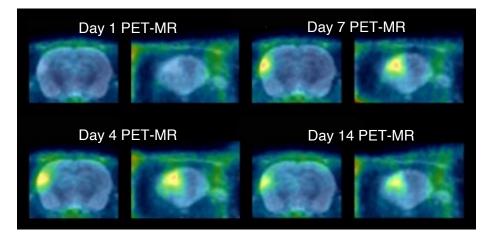
Collaboration with Radio pharmacy Core Facility (RCF) at KI We can do PET with tracers containing: Short half time such as <sup>18</sup>F, <sup>11</sup>C or <sup>123</sup>I Long half time such as <sup>89</sup>Zr, <sup>68</sup>Ga or <sup>125</sup>I More complicated ligands that can be activated in-vivo



[89Zr]Zr-oxine MOLM-14 Mus #38: 600 kBq, 0.8 x 10^6 cells



PET-CT tracer [11C] in rat

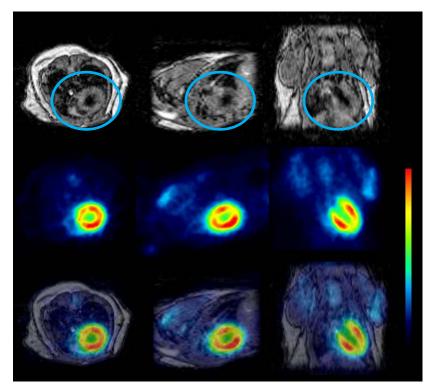


PET-MR of uptake from day 1 to day 14 of using [11C]PBR28 tracer in M2CAO animal.

# PET-CT / PET-MR 2x PET-CT and 1 PET-MR (1T)

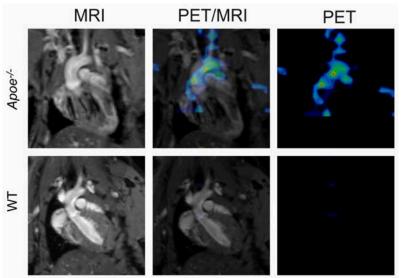


## PET-MR mouse heart (FTG)



Strong PET signal from 18F (FTG), but an avranged image of the heart

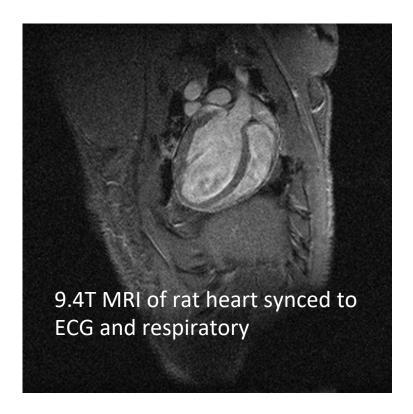
## PET + 9.4T MRI mouse heart (89Zr)

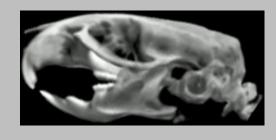


Rather weak signal from PET, but clear MR signal from synchronized ECG + resp



2x PET-CT and 1 PET-MR (1T)





CT rat skull (reconstructed 3D)

# **Angiography CT**



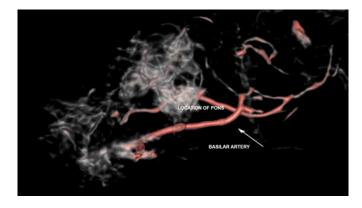
Angio-CT room



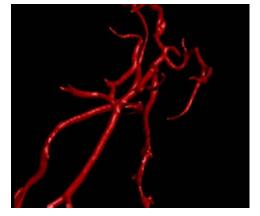


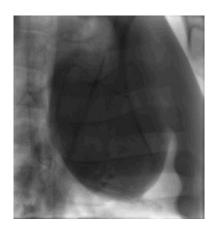
The equipment is continuously upgraded and used in activities like:

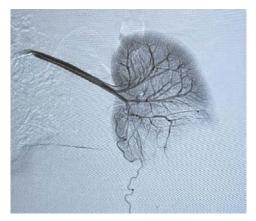
- Research in endovascular devices.
- Experimental stroke research, combined.
- Courses e.g. endovascular stroke treatment.
- Development of direct puncture devices.
- Super selective stem cell installation.
- Radiation reduction in angiography.



Angio-CT using contrast to visualize vessels in pig brain







Angio-CT using contrast to visualize vessels in brain, heart and kidney



# **Experimental Surgery**



Four fully equipped operating units for surgical training, courses, and research projects.



- Operating table (x4)
- Anesthesia equipment (Dräger Primus Infinity)
- Monitoring equipment (Datex / Philips MX800)
- Surgical diathermy (Valleylab, ForceTriad)
- Surgical light (Maquet Volista)
- Patient warming units (Mistral Air)
- Surgical suction
- Standard surgical instruments

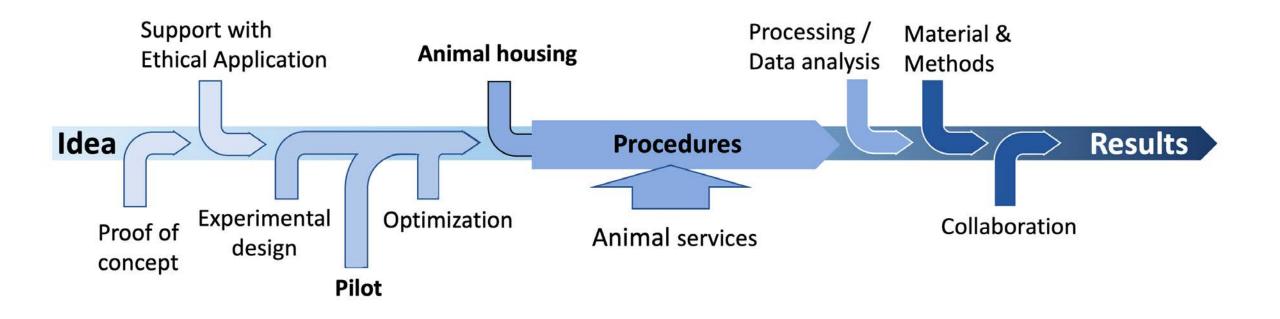
Our operating units are regularly used for surgical training in areas such as transplantation, neuro/stroke, specialist trauma expertise, otorhinolaryngology, gynecology, pediatric surgery and more. We also have ongoing research projects for new and refined medical equipment and methods.



## From Idea to Results







We can support preclinical projects from the initial **Idea** with a **Proof of concept** and setup of a **Pilot**.

Assist with the **Ethical Application** and take **care of the Animals** before, during and after the **Procedures**.

In imaging projects we also can assist with data analysis and data processing.

It is our mission that you reach your goal with the work performed at KERIC.

## https://ki.se/keric







Research areas Researchers Research funding Infrastructure

🏫 / Research / Infrastructure / Core facilities for research / Karolinska Experimental Research and Imaging Centre (KERIC)

## Karolinska Experimental Research and Imaging Centre (KERIC)

KERIC is a core facility offering different imaging analyses in experimental systems in combination with experimental surgery in large animals like swine as well as in rodents. We offer a fully equipped facility for experimental surgery with four full size operation units equipped

Booking is done via iLab or in communication with staff.

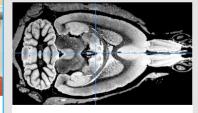
about KERIC

Contact

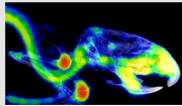
Collaborations







Preclinical MRI 9.4T



PET-CT/MR



Angio CT



**Experimental Surgery** 



#### Selected Research at KERIC

Some of our most successful customers that uses KERIC for their research.



KAROLINSKA







We can help you get your ideas to scientific results

#### Example of Workflow

How we can help in different stages of a research project. Not just images.



#### News for KERIC

https://ki.se/keric webpage update 2022-04-12





### Selected projects and publications (scroll right)

Publications list from projects that have used KERIC facilities or collaborated direct with KERIC. List is in chronological order. Starting with the latest at top.

Please include KERIC in the acknowledgement, in publications based on results obtained from the any of our modalities or collaburated direct with

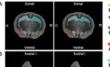
"The """ was performed at the Department of Comparative Medicine/Karolinska Experimental Research and Imaging Centre at Karolinska University Hospital, Solna, Sweden."

/Thank you

COMMUNICATIONS

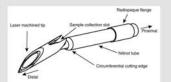
#### COVID-19 pathophysiology may be driven by an imbalance in the reninangiotensin-aldosterone system 🗹

SARS-CoV-2 uses ACE2, an inhibitor of the Renin-Angiotensin-Aldosterone System (RAAS), for cellular entry. Studies indicate that RAAS imbalance worsens the prognosis in COVID-19. We present a consecutive retrospective COVID-19 cohort with findings of frequent pul-monary thromboembolism (17%), high pulmonary artery pressure (60%) and lung MRI perfusion disturbances. We demonstrate, in swine, that infusing angiotensin II or blocking ACE2 induces increased pulmonary artery pressure, reduces blood oxygenation, increases coagulation, disturbs lung perfusion, induces diffuse alveolar damage, and acute tubular necrosis compared to control animals.



#### Changes in brain architecture are consistent with altered fear processing in domestic rabbits &

The most characteristic feature of domestic animals is their change in behavior associated with selection for tameness. Here we show, using high-resolution brain magnetic resonance imaging in wild and domestic rabbits, that domestication reduced amygdala volume and enlarged medial prefrontal cortex volume, supporting that areas driving fear have lost volume while areas modulating negative affect have gained volume during domestication. ...



#### Myocardial micro-biopsy procedure for molecular characterization with increased precision and reduced trauma 27

Endomyocardial biopsy is a valuable tool in cardiac diagnostics but is limited by low diagnostic yield and significant complication risks. Meanwhile recent developments in transcriptomic and proteomic technologies promise a wealth of biological data from minimal tissue samples. To take advantage of the minimal tissue amount needed for molecular analyses, we have developed a sub-millimeter endovascular biopsy device, considerably smaller than current clinical equipment, and devised a low-input RNAsequencing protocol for analyzing small tissue

### **KERIC Calendar**



### **Publications KERIC**

Publications list from projects that have used KERIC facilities or collaborated direct with KERIC in reverse chronological order. Starting with the latest at top. (60 Publications)

Tjernström N, Tie-Qiang L, Holst S, Roman E. Functional connectivity in rewardrelated networks is associated with individual differences in gambling strategies in male Lister hooded rats. Addiction Biology. 2022 Jan 17;27(2):e13131https://doi.org/10.1111/adb.13131 🗹

Rysz, S. Al-Saadi J., Sjöström A., Farm M., Jalde F. C., Plattén M., Eriksson H., Klein M., Vargas-Paris R., Nyrén S., Abdula G., Ouellette R., Granberg T., Jonsson Fagerlund M., Lundberg J. COVID-19 pathophysiology may be driven by an imbalance in the renin-angiotensin-aldosterone system. Nat Commun 12, 2417 (2021). https://doi.org/10.1038/s41467-021-22713-z ば

Savva, C., Helguero, L. A., González-Granillo, M., Couto, D., Melo, T., Li, X., Angelin, B., Domingues, M. R., Kutter, C., & Korach-André, M. (2021). Obese mother offspring have hepatic lipidic modulation that contributes to sex-dependent metabolic adaptation later in life. Commun Biol, 4(1), 14. https://doi.org/10.1038/s42003-020-01513-z &

Rysz, S., Jalde F. C., Oldner A., Eriksson L. I., Lundberg J. and Jonsson Fagerlund M. Treatment with angiotensin II in COVID-19 patients may not be beneficial. Crit Care 24, 546 (2020). https://doi.org/10.1186/s13054-020-03233-6 ♂

Rysz, S. et al. The effect of levosimendan on survival and cardiac performance in an ischemic cardiac arrest model - A blinded randomized placebo-controlled study in swine. Resuscitation 150, 113–120 (2020). https://doi.org/10.1053/j.jvca.2013.03.027 ♂

Ahmed, M., Tegnebratt, T., Tran, T. A., Lu, L., Damberg, P., Gisterå, A., Tarnawski, L., Bone, D., Hedin, U., Eriksson, P., Holmin, S., Gustafsson, B., & Caidahl, K. (2020). Molecular Imaging of Inflammation in a Mouse Model of Atherosclerosis Using a Zirconium-89-Labeled Probe. Int J Nanomedicine, 15, 6137-6152. https://doi.org/10.2147/IJN.S256395 ♂

González-Granillo, M., Savva, C., Li, X., Ghosh Laskar, M., Angelin, B., Gustafsson, J., & Korach-André, M. (2020). Selective estrogen receptor (ER)β activation provokes a redistribution of fat mass and modifies hepatic triglyceride composition in obese male

