

RESEARCH FELLOW 公募
国立循環器病研究センター研究所 各部紹介

部・室名	循環動態制御部
責任者(部長名・室長名)	西中 知博・朔 啓太
研究内容 循環動態制御部では、循環の仕組みをシンプルに数理モデル化することで理解し、その知見を活かした最適な循環制御方法を模索しています。また、医療機器開発を最重要課題とし、多くの企業と共同研究・開発を実施しています。 1. 循環の仕組みを知る 循環の仕組みを知る手段として、シンプルな数理モデルを用いた循環の理解や病態の解明に取り組んでいます。そのために、圧容積関係(Pressure-volume loop: PV loop)や循環平衡理論、自律神経解析などを駆使し、正常と病態の循環制御的違いを同定しています。 2. シミュレーションで医療のミライを拓く 循環器系の数理モデル化により、コンピュータ上に任意の循環をシミュレーションすることが可能になります。将来的には、仮想治療や治療効果予測による診療支援、自動治療への発展、医療機器の In-silico 検証など幅広い用途で医療に貢献するツールに成長させたいと考えています。 3. 循環を操り、治療に生かす 培った循環に関わる器官の機能や循環調節に関する研究結果を基に循環を最適にコントロールする手法を開発することで病気の克服を目指しています。具体的には、薬物や機械で循環の制御機構を再建する研究・開発を行っています。	
直近 3 年間の業績 1. Hiraki N, Morita H, Nakamura Y, Ohba K, Sato K, Otake M, Yoshida Y, Nishikawa T, Fukumitsu M, Uemura K, Kawada T, Maruhashi T, Saku K*. Understanding the hemodynamic effects of milrinone in normal and failing hearts: an experimental study using the circulatory equilibrium framework in canines. <i>Am J Physiol Heart Circ Physiol.</i> 2026, 330: H1241-H1256. 2. Teramoto K*, Saku K*, Ikeda Y, Nakata J, Unoki T, Yamamoto T, Sakamoto T, Ako J. Temporal Hemodynamic Patterns in Cardiogenic Shock Treated With Isolated Percutaneous Ventricular Assist Device and Combined Therapy With Venoarterial Extracorporeal Membrane Oxygenation. <i>J Card Fail.</i> 2026, S1071-9164(26)00073-4. 3. Matsushita H, Oba K, Kurono K, Nakabayashi M, Yoshida Y, Sato K, Morita H, Otake M, Hiraki N, Fukumitsu M, Nishikawa T, Uemura K, Kawada T, Ichinose M, Ono Y, Saku K*. Diffuse Correlation Spectroscopy for Non-invasive Monitoring of Peripheral Perfusion During Cardiogenic Shock and VA-ECMO Support: An Experimental Study. <i>Shock.</i> 2026, in press	

4. Matsushita H, Nishikawa T, Uemura K*, Sasaki K, Nakamura Y, Sato K, Morita H, Ohba K, Kawada T, Kataoka Y, Alexander J Jr, Saku K. Automated Hemodynamic Management System for the Prevention of Hypotension and Cardiac Output Reduction During General Anesthesia: Preclinical Experimental Validation. *Journal of Anesthesia*. 2026, in press
5. Ohuchi H, Saku K, Miyazaki A, Mori A, Kato Y, Sakaguchi H, Shiraishi I, Kurosaki K. Benefits of Chronic Atrial Pacing in Fontan Physiology with Sinus Node Dysfunction and Preserved Ejection Fraction: Novel Therapeutic Approach for Failing Fontan. *Pediatr Cardiol*. 2026, 47: 666-674.
6. Nishikawa T*, Matsushita H, Otake M, Yokota S, Kakuuchi M, Sato K, Morita H, Hiraki N, Ohba K, Yoshida Y, Fukumitsu M, Uemura K, Kawada T, Kusano K, Saku K*. Impaired atrioventricular time interaction contributes to tachycardia-induced hemodynamic deterioration: In vivo and in silico hybrid approach. *Int J Cardiol*. 2026, 12: 134192.
7. Yokota S, Maruhashi T, Kakuuchi M, Nishikawa T, Matsushita H, Morita H, Sato K, Yoshida Y, Kawada T, Uemura K, Nishinaka T, Saku K*. Impella preserves haemodynamics with adequate stressed blood volume and normal pulmonary vascular resistance in a goat model of ventricular fibrillation. *Eur Heart J Open*. 2025, 6: oeaf173.
8. Kataoka Y*, Uemura K, Sampei M, Fukuda Y, Peterson J, Funada R, Saku K, Joe Alexander, Sunagawa K. A Discrete Hemodynamic Control Framework: Proof-of-Concept Study for Autonomous Drug Therapy in Acute Heart Failure. *IEEE Trans Biomed Eng*. 2025, 12. in press
9. Nishikawa M, Kawada T, Saku K, Kinoshita H, Kawahito S*. Effects of sugammadex on heart rate variability in anesthetized male rats. *Physiol Rep*. 2025, 13: e70507.
10. Nishikawa M, Kawada T, Hiraki N, Fukumitsu M, Sato K, Kinoshita H, Kawahito S, Saku K*. Acute effects of vericiguat on urine excretion during baroreflex-mediated sympathetic arterial pressure regulation in male rats. *Autonomic Neuroscience*. 2025, 262: 103363..
11. Ikeda Y*, Saku K, Nakata J, Unoki T, Nakahara S, Iwaya T, Iikura S, Takigami Y, Yamamoto T, Sakamoto T, Ako J. UNLOADERS-PVAD Investigators. UNLOADERS-PVAD Weaning Score: predicting post-weaning adverse events in cardiogenic shock patients supported by microaxial flow pump. *Journal of Intensive Care*. 2025, 13(1): 62.
12. Kamada K, Mannoji H, Tohyama T, Nishikawa T, Tsutsui H, Sunagawa K, Saku K. Simulated microgravity exposure markedly attenuates the baroreflex total loop gain and induces orthostatic hypotension in male rats. *Physiol Rep*. 2025, 13 : e70637.
13. Yokoi A, Kawada T, Fukumitsu M, Hiraki N, Sato K, Kakuuchi M, Joe Alexander Jr, Nishikawa T, Tanaka R, Saku K. Vericiguat attenuates the dynamic gain of open-loop baroreflex function in a low-frequency range. *Autonomic Neuroscience: Basic and Clinical*. 2025, 262: 103354.
14. Sato K, Peterson J, Uemura K, Ohba K, Nishikawa T, Sasaki K, Yokota S, Matsushita H, Morita H, Yoshida Y, Hiraki N, Otake M, Nakamura Y, Fukumitsu M, Kawada T, Kataoka Y, Alexander J, Sunagawa K, Saku K. One Surface Fits All: Validating the Venous Return Model Across Species and Scales in Circulatory Equilibrium. *Am J Physiol Heart Circ Physiol*. 2025, 329(4): H853-H866.
15. Fukumitsu M*, Hotta N, Kawada T, Nishikawa T, Saku K. A rapid increase in left

atrial pressure reduces pulmonary arterial compliance in rats: Insights from pulmonary arterial impedance analysis. *Journal of Applied Physiology*. 2025, 1985.

16. Hiraki N, Kawada T, Fukumitsu M, Nishikawa T, Matsushita H, Yoshida Y, Sato K, Morita H, Otake M, Ohba K, Uemura K, Joe Alexander Jr., Saku K*. Impact of dopamine on baroreflex-mediated sympathetic arterial pressure regulation in rats: an open-loop analysis. *Am J Physiol Regul Integr Comp Physiol*. 2025, 329(2): R329-R339.
17. Takata M, Nishikawa M, Hayashi H, Yamamura A, Saku K, Kawada T, Sato M, Kawahito S, Kinoshita H*. Lack of Effect of Omecamtiv Mecarbil on Smooth Muscle Myosin Phosphatase Target Subunit-1 (MYPT1) Phosphorylation and Hemodynamics in Rats. *Cureus*. 2025, 17(7): e87854.
18. Matsushita H, Kurono K, Nakabayashi M, Sato K, Morita H, Yoshida Y, Fukumitsu M, Uemura K, Kawada T, Ichinose M, Ono Y, Saku K. Non-invasive monitoring of microcirculation dynamics in hypovolemic shock: a novel application of diffuse correlation spectroscopy. *Intensive Care Medicine Experimental*. 2025, 13: 53
19. Ikeda Y, Saku K, Nakata J, Unok T, Yamamoto T, Sakamoto T, Ako J. Phase-Specific Hemodynamic Criteria and Outcomes in Patients With Cardiogenic Shock Receiving Percutaneous Ventricular Assist Devices. *J Am Heart Assoc*. 2025, 14: e042249.
20. Tadokoro N*, Saku K, Tonai K, Tadokoro Y, Kutsuzawa R, Fukushima S. Rapid-onset postoperative acute kidney injury is associated with mortality in patients with postcardiotomy cardiogenic shock. *Frontiers in Cardiovascular Medicine*. 2025, 13.
21. Sato K, Yoshida Y, Yokota S, Matsushita H, Morita H, Fukumitsu M, Nishikawa T, Uemura K, Kawada T, Saku K. Pressure-strain product reflects left ventricular stroke work under a wide range of left ventricular assist device support levels. *Frontiers in Cardiovascular Medicine*. 2025, 12.
22. Kataoka Y, Fukuda Y, Shelly I, Peterson J, Uemura K, Saku K, Sampei M, Alexander J, Suna. Discrete Two-Degree-of-Freedom Control for Hemodynamic Optimization: Circulatory Simulation Study with Baroreflex Variability. *Annu Int Conf IEEE Eng Med Biol Soc*. 2025, 1-6.
23. Uemura K, Nishikawa T, Matsushita H, Sasaki K, Sato K, Yokota S, Morita H, Yoshida Y, Fukumitsu M, Kawada T, Kataoka Y, Saku K. Minimally invasive monitor of cardiac output based on the machine-learning analysis of the pulse contour of the peripheral arterial pressure. *Annu Int Conf IEEE Eng Med Biol Soc*. 2025, 1-4.
24. Nishikawa T, Uemura K, Matsushita H, Morita H, Sato K, Yoshida Y, Fukumitsu M, Kawada T, Saku K. Development of a Framework for the Hemodynamic Impact of Positive End-Expiratory Pressure in Normal and Heart Failure Conditions. *Am J Physiol Heart Circ Physiol*. 2025, 328: H361-H376.
25. Unoki T, Nakayama T, Saku K, Matsushita H, Inamori T, Matsura J, Toyofuku T, Sato T, Konami Y, Suzuyama H, Inoue M, Horio E, Kodama K, Taguchi T, Nishikawa T, Sawamura T, Nakao K, Sakamoto T, Okumura K, Koyama J. Impact of ECPELLA Support on 1-Year Outcomes and Myocardial Damage in Patients with Acute Coronary Syndrome and Refractory Cardiogenic Shock: A Single-Center Retrospective Observational Study. *J Cardiol*. 2025, 85: 352-359.
26. Unoki T, Uemura K, Yokota S, Matsushita H, Kakuuchi M, Morita H, Sato K, Yoshida Y, Sasaki K, Kataoka Y, Nishikawa T, Fukumitsu M, Kawada T, Sunagawa K, Alexander J, Saku K. Closed-loop automated control system of

- extracorporeal membrane oxygenation and left ventricular assist device support in cardiogenic shock. *ASAIO J.* 2025, 71(6): 461-471.
27. Maruhashi T, Saku K, Maruki H, Oi M, Asari Y. Potential new treatment for inferior vena cava injury using extracorporeal membrane oxygenation applying flow diversion effect. *Trauma Surg Acute Care Open.* 2024, 9: e001618.
 28. Fukumitsu M, Kawada T, Nishikawa T, Yokota S, Matsushita H, Morita H, Sato K, Yoshida Y, Uemura K, Saku K. Effects of nitric oxide inhalation on pulmonary arterial impedance: differences between normal and pulmonary hypertension male rats. *American Journal of physiology. Heart and Circulatory Physiology.* 2024, 327: H287–H297.
 29. Kawada T*, Yamamoto H, Fukumitsu M, Nishikawa T, Matsushita H, Yoshida Y, Sato K, Morita H, Alexander J Jr, Saku K. Acute effects of empagliflozin on open-loop baroreflex function and urine output in streptozotocin-induced type 1 diabetic rats. *J Physiol Sci.* 2024, 74: 48.
 30. Kawada T*, Fukumitsu M, Matsushita H, Yoshida Y, Sato K, Morita H, Nishikawa T, Suehara S, Sawada S, Saku K. Effects of bilateral renal denervation on open-loop baroreflex function and urine excretion in spontaneously hypertensive rats. *Hypertension research.* 2024, 11: 3255-3266.
 31. Matsushita H, Saku K*, Nishikawa T, Unoki T, Yokota S, Sato K, Morita H, Yoshida Y, Fukumitsu M, Uemura K, Kawada T, Kikuchi A, Yamaura K. Impact of right ventricular and pulmonary vascular characteristics on Impella hemodynamic support in biventricular heart failure: A simulation study. *J of Cardiol.* 2024, 85: 100-107.
 32. Fukuda Y*, Kawada T, Kataoka Y, Peterson J, Saku K, Alexander J, Sunagawa K. Influence of angiotensin II and telmisartan on in vivo high-resolution renal arterial impedance in rats. *Am J Physiol Regul Integr Comp Physiol.* 2024, 327: R349-R361.
 33. Nishikawa T, Kamada K, Morita H, Matsushita H, Yokota S, Sato K, Unoki T, Tsutsui H, Sunagawa K, Saku K*. Automated control of Impella maintains optimal left ventricular unloading during periods of unstable hemodynamics and prevents myocardial damage in acute myocardial infarction. *Int J Cardiol.* 2024, 410: 132244.
 34. Yokota S, Uemura K*, Unoki T, Matsushita H, Kakuuchi M, Yoshida Y, Sasaki K, Kawada T, Nishikawa T, Kataoka Y, Peterson J, Sunagawa K, Alexander J, Saku K. Novel closed-loop control system of dual rotary blood pumps in total artificial heart based on the circulatory equilibrium framework: a proof-of-concept in vivo study. *IEEE Trans Biomed Eng.* 2024, 71: 3358-3369.
 35. Sasaki K*, Kawada T, Matsushita H, Yokota S, Kakuuchi M, Yokoi A, Yoshida Y, Morita H, Sato K, Nishikawa T, Annette P N Kutter, Kataoka Y, Joe Alexander, Saku K, Ishikawa T, Uemura K*. Computer-controlled closed-loop norepinephrine infusion system for automated control of mean arterial pressure in dogs under isoflurane-induced hypotension: A feasibility study. *Frontiers in Veterinary Science, section Anesthesiology and Animal Pain Management.* 2024, 11: 1374356.
 36. Li M*, Zheng C, Kawada T, Uemura K, Yokota S, Matsushita H, Saku K. Donepezil Attenuates Progression of Cardiovascular Remodeling and Improves Prognosis in Spontaneously Hypertensive Rats with Chronic Myocardial Infarction. *Hypertension Research.* 2024, 47: 1298-1308.
 37. Matsushita H, Saku K*, Nishikawa T, Yokota S, Sato K, Morita H, Yoshida Y, Fukumitsu M, Uemura K, Kawada T, Yamaura K. The impact of ECPELLA on

haemodynamics and global oxygen delivery: a comprehensive simulation of biventricular failure. Intensive Care Medicine Experimental. 2024, 12:13.

38. Unoki T*, Konami Y, Nakayama T, Suzuyama H, Horio E, Taguchi E, Saku K, Sawamura T, Nakao K, Sakamoto T. Efficacy and safety of post-closure technique using Perclose ProGlide/ProStyle device for large-bore mechanical circulatory support access sites. Cardiovasc Revasc Med. 2024, 62:60-65.
39. Kawada T*, Matsushita H, Yokota S, Yoshida Y, Fukumitsu M, Alexander J, Saku K. Short-term dynamic characteristics of diuresis during exogenous pressure perturbations with and without arterial baroreflex control. Am J Physiol Regul Integr Comp Physiol. 2024, 326: R230-R241.

その他 情報

総合内科専門医、循環器専門医、麻酔科専門医、心臓外科専門医、集中治療専門医が所属しており、臨床に直結する研究や開発を実施しています。

- 循環動態の臨床教育を目的とした循環動態アカデミーという活動をしています：
<https://circ-dynamics.jp/>
- 迷走神経刺激カテーテルの開発と実用化を目指しています：
<http://www.circucon.jp/ivns.html>
- NTT Research 社とともに心血管バイオデジタルツインの構築を目指した共同研究をしています：<https://www.ncvc.go.jp/res/divisions/cv-biodt/>
- 国循環ベンチャーを創業し、AI による医療連携の円滑化に関する試みをしています：<https://cubec.jp/>
- 心原性ショックレジストリー UNLOADERS 研究を推進しています：
<https://sk-kumamoto.jp/unloaders/>