





Press release dated May 31, 2022

# A world premiere: for the first time, a human liver was treated in a machine and then successfully transplanted

The multidisciplinary Zurich research team Liver4Life has succeeded in doing something during a treatment attempt that had never been achieved in the history of medicine until now: it treated an originally damaged human liver in a machine for three days outside a body and then implanted the recovered organ into a cancer patient. One year later, the patient is doing well.

The Liver4Life research team owes this success to its perfusion machine, which was developed in house, making it possible to implant a human organ into a patient after a storage period of three days outside a body. The machine mimics the human body as accurately as possible, in order to provide ideal conditions for the human livers. A pump serves as a replacement of the heart, an oxygenator replaces the lungs and a dialysis unit takes over the functions of the kidneys. In addition, numerous hormone and nutrient infusions perform the functions of the intestine and pancreas. Like the diaphragm in the human body, the machine also moves the liver to the rhythm of human breathing. In January 2020, the multidisciplinary Zurich research team – involving the collaboration of University Hospital Zurich (USZ), ETH Zurich and the University of Zurich (UZH) – demonstrated for the first time that perfusion technology makes it possible to store a liver outside the body for several days (see press release of January 13, 2020).

## From poor to good in three days

The team prepared the liver in the machine with various drugs. In this way, it was possible to transform the liver into a good liver graft, even though it was originally not approved for transplantation. Perfusion over several days, i.e. the mechanical perfusion of the organ, enables, for example, antibiotic or hormonal therapies or the optimization of liver metabolism. In addition, lengthy laboratory or tissue tests can be carried out without time pressure. Under normal circumstances, this is not possible because organs can only be stored for 12 hours if they are stored conventionally on ice and in commercially available perfusion machines.

## Treatment attempt successful

As part of an approved individual treatment attempt, the doctors gave a cancer patient on the Swisstransplant waiting list the choice of using a treated human liver. Following his consent, the organ was transplanted in May 2021. The patient was able to leave hospital a few days after the transplantation and is now doing well: "I am very grateful for the life-saving organ. Due to my rapidly progressing tumor, I had little chance of getting a liver from the waiting list within a reasonable period of time."

## Saving more lives

The article on the first transplantation of a liver preserved for several days on a perfusion machine was published in one of the most renowned scientific journals, Nature Biotechnology, on May 31, 2022. "Our therapy shows that by treating livers in the perfusion machine, it is possible to alleviate the lack of functioning human organs and save lives," explains Prof. Pierre-Alain Clavien, Director of the Department of Visceral Surgery and Transplantation at the University Hospital Zurich (USZ). Prof. Mark Tibbitt, Professor of Macromolecular Engineering at ETH Zurich, adds: "The interdisciplinary approach to solving









complex biomedical challenges embodied in this project is the future of medicine. This will allow us to use new findings even more quickly for treating patients."

The next step in the Liver4Life project is to review the procedure on other patients and to demonstrate its efficacy and safety in the form of a multicenter study. Its success would mean that in the future, a liver transplantation, which usually constitutes an emergency procedure, would be transformed into a plannable elective procedure. At the same time, a next generation of machines is being developed. In addition, those involved in basic research continue to look for ways of treating other liver diseases outside the body with drugs, molecules or hormones.

## Liver4Life: a Wyss Zurich project

The <u>Liver4Life</u> project was launched in 2015 under the umbrella of the Wyss Zurich Translational Center (Wyss Zurich). It brings together the highly specialized technical knowhow and biomedical knowledge of around ten medical professionals, biologists and engineers. The project is being financed with donations from the initiator of Wyss Zurich, Dr. h.c. mult. Hansjörg Wyss.

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Clavien PA, Dutkowski P et al. Transplantation of a human liver following 3 days of ex situ normothermic preservation. Nature Biotechnology 2022. Doi: 10.1038/s41587-022-01354-7 https://www.nature.com/articles/s41587-022-01354-7

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#### Images

*Photo 1*: The Wyss Zurich Team connects the donor liver to the perfusion machine in the clean room.



*Photo 2*: Prof. Pierre-Alain Clavien and Prof. Philipp Dutkowski during the transplantation of the liver treated in the machine.



*Photo 3*: The surgeon, Prof. Pierre-Alain Clavien, and the patient when the latter was discharged from hospital after the successful transplantation.



*Photo 4*: Anniversary celebration of the Wyss Zurich Team with the patient. From left to right: Matteo Müller, Prof. Mark Tibbitt, the patient, Prof. Pierre-Alain Clavien, Lucia Bautista Borrego, Max Hefti and Richard Sousa Da Silva.











*Diagram 1*: The perfusion machine replaces the functions of various organs in order to keep the donor liver alive outside of the body.

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