

High pressure tissue decellularization offers promise for tissue engineering applications

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Q Your work involves decellularized tissues. What does this mean, and why is it important for the field of tissue engineering?

A: From the late 1990s, it has been reported that human decellularized tissue-extracellular matrix from which cellular components have been removed-works well as a substitute for living tissue. Decellularized tissue exhibits good biocompatibility, and the lack of donor cellular material minimizes the risk of graft rejection. Importantly, the removal of donor cellular material may allow for the use of animal tissue as a source for xenotransplantation into humans, so this technology has the potential to provide a high-performance scaffold for tissue engineering.

Q Could you explain the technique you have developed, and any challenges you had to overcome?

A: The main existing approach for decellularizing tissues involves using detergents. Such techniques may leave residual cytotoxic chemicals or change the mechanical properties of the tissue. We developed an alternative method of decellularizing tissues using high-

Dr. Kishida graduated from the Faculty of Engineering at Kyoto University, where he received his MD and PhD. He became Assistant Professor in 1992, and Associate Professor in 1994 at Kagoshima University. He joined TMDU as Professor of Material-based Medical Engineering at the Institute of Biomaterials and Bioengineering in 2004. In 2014, TMDU established the Department of Acellular Biomaterials and Regenerative Medicine and Dr. Kishida became the field manager there.

hydrostatic pressure (HHP, >600 MPa). The cell debris can then be removed by washing, without chemicals or detergents.

Q What are the benefits of this method of decellularization?

A: *In vivo* grafts of HHP-decellularized tissues show minimal inflammation and exhibit good long-term stability. Grafted decellularized tissues can also strongly recruit host cells, facilitating rapid integration into host tissue. An additional benefit is that HHP has a sterilizing effect: Because the treatment disrupts lipid bilayers, it can destroy pathogens such as bacteria, fungi, and some viruses.

Q What kinds of applications for these decellularized tissues have you explored?

A: We transplanted decellularized porcine

corneas into rabbits, and achieved transparent corneas that lasted for at least twelve months after transplantation.

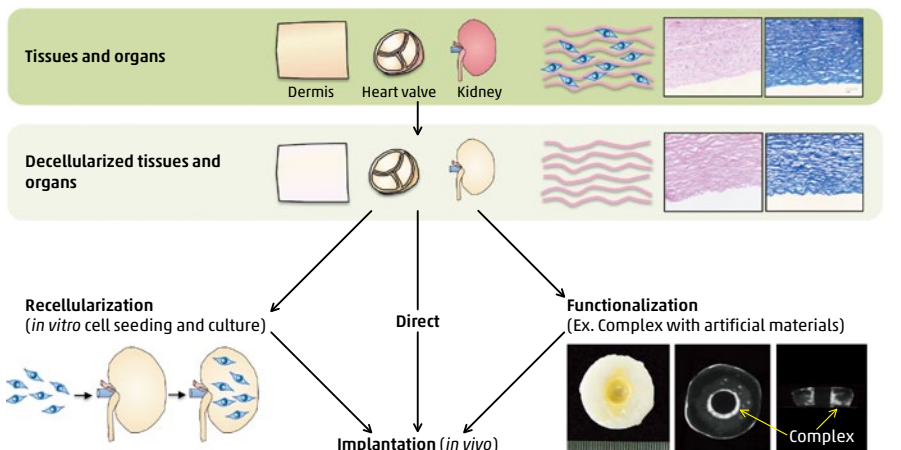
Another challenge we tackled was developing graft materials that can function as an interface between hard and soft tissues. We have used decellularized tissues in conjunction with a synthetic polymer to develop a hybrid biomaterial. This material can function as an interface between different tissue types, or between a tissue and an implanted synthetic device.

We also explored using powders made from decellularized tissues for treating myocardial infarction in a rat model. Decellularized liver powder promoted cell integration and neovascularization both *in vitro* and *in vivo*, and suppressed myocardial necrosis.

Q How do you expect your research to develop in the future?

A: We are working on regenerating bone marrow, brain, and cartilage using the relevant decellularized tissues. Decellularized tissue is emerging as superior to many artificial materials currently used in tissue engineering. We will continue working with our research partners to explore the potential of decellularized tissue.

Overview of applications of decellularized tissues and organs



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