3D PRINTING BODY PARTS: AN OVERVIEW OF HOW ADDITIVE MANUFACTURING IS SHAPING TISSUE ENGINEERING

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ABSTRACT

Additive manufacturing, commonly referred to as 3D printing, is a manufacturing technique that emerged in the 1980's mainly focused on engineering prototyping. Recent advances in the precision and cost of the techniques, as well as the widespread use of 3D modeling have increased 3D printing's scope of use from high-end engineering prototypes to a large variety of uses in manufacturing. 3D printing has been shown to improve the processing time, reduce waste, and increase the level of customization of certain products by eliminating the need for the specialty tooling and dies that are traditionally used in manufacturing. Additionally, the ability to physically print complex shapes based on a computer model has given rise to new products that would otherwise be simply impossible to create. The field of tissue engineering has taken advantage of this technology by printing organic cells and inorganic biomaterials with levels of control and precision that surpass previous production techniques. Furthermore, 3D printing allows for applications that can be customized on a case-by-case basis to fit the needs of individual patients. This review will overview the 3D printingbased techniques being used in tissue engineering, and discuss the current applications and challenges of this rapidly advancing field.

INTRODUCTION

Tissue engineering was conceptualized in 1987 at a National Science Foundation committee meeting¹. The concept is defined as "an interdisciplinary field that applies the principle of engineering and the life sciences toward the development of biological substitutes that restore, maintain, or improve tissue function."²

Main components of tissue engineering

- **Cells** The patient's own cells can be harvested and grown outside the body
- Scaffold Made out of bio-compatible materials, the scaffold supports the tissue and allows for cell permeation.³

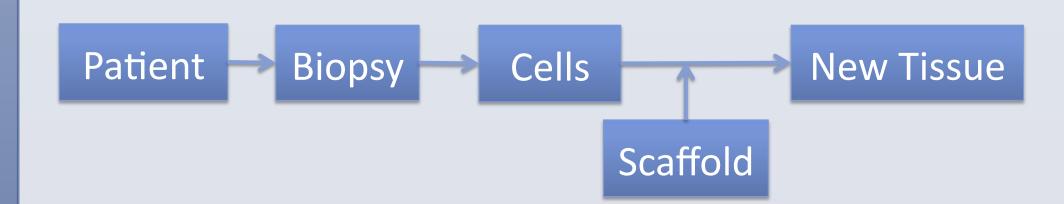


Figure 1. Flowchart representing the basic components used in tissue engineering.

Using the patient's own cells

- Eliminates the risk of disease transmission
- Eliminates the risk of immunogenic incompatibility
- Increases the availability of donor tissue³

CURRENT APPLICATIONS

ORGANS

- The following structures have been successfully created using 3D printed bio-scaffolds⁴
 - Skin

• 3D Printing and Tissue Engineering⁵

- Esophagus
- Blood Vessels
- Bladder

- Trachea
- Automates the most tedious parts of the process, decreasing lead time
- Increases precision and uniformity of the micro-scale structures of the bio-scaffolding
- Allows for custom tailored treatments built directly from 3D MRI models of the patient's existing tissue structures
- When cells are removed from the host and assembled together on support structures, a few things happen naturally:
 - The tissues will fuse together⁶
 - The tissues actually adopt the form and the function of the original tissue structure⁷
- The biological mechanisms of cell permeation happen naturally, but it is up to the scientists to develop the bio-scaffold for the cells to grow on
 - The scaffold creates the form of the new tissue or organ
 - The scaffold ensures that the tissue or organ is structurally sound

BONES

- Washington State University researchers are using 3D printed bio-scaffolds to grow bones9
 - Created 3D scaffolds made of bio-compatible material
 - Submerged the scaffold into immature human bone cells
 - A week later, the bone cells had attached to the scaffold and fused together
 - Can use this technique with a computed tomography scan to replicate a patient's own bone
- Oxford Performance Materials¹⁰
 - Received the first FDA approval to use 3D printed polymer implants to replace damaged bone structures in the human skull
 - Made out of PEKK, the implant acts as a very sturdy bio-scaffold that will allow the surrounding bone structure to grow into and fuse with the implant
- Other materials for bone implants are being researched including polyethylene, glass, and starch-based plastics like PLA
- Scaffolds for bones can also be designed for drug delivery by implanting the necessary drugs inside the scaffold itself, the drug will be released as the regenerating tissue fuses with that section of the scaffold¹¹

(d) 15kV X2,300 10μm UoC

Figure 2⁵. PDMS bio-scaffold created using stereolithography techniques.

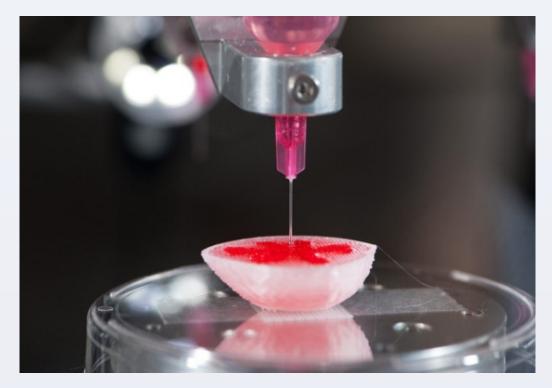


Figure 38. Prototype kidney being created on a 3D printer.



Figure 4⁴. Functional bladder created using printed bio-scaffolds.



Figure 5¹⁰. Example of an FDA approved 3D printed PEKK cranial implant.

CONCLUSION

Tissue engineering and 3D printing are both emerging fields that are gaining a lot of attention in the scientific world. By incorporating 3D printing techniques, tissues engineering is reaching a point that therapies can be created quickly, precisely, and with a level of detailed customization that could never before have been imagined. The possibility of having replacement organs or bone created using tissue from the same patient, formed to the exact structure of the organ or bone being replaced sounds like the work of science fiction, but it is actually close to reality. There is still work to be done, but it is clear that these two relatively new fields have amazing potential together.

REFERENCES

- 1. Nerem, R.M. (2006) Tissue engineering: the hope, the hype, and the future. Tissue Eng., 12(5), 1143-1150.
- Langer, R.; Vacanti, J.P. (1993) Tissue engineering. Science, 260(5110), 920-926.
 Lewandowska-Szumiel M, Wójtowicz J. (2011) Bone tissue engineering a
- field for new medicinal products?. *Current Pharmaceutical Biotechnology*;12(11):1850-1859.

 4. "Replacement Organs and Tissue: Overview", Wake Forest School of
- Medicine, (2013). http://www.wakehealth.edu/Research/WFIRM/ Projects/Replacement-Organs-and-Tissue.htm

 5. S. Rekstyte, M. Malinauskas, and S. Juodkazis. (2013), "Three-
- dimensional laser micro-sculpturing of silicone: towards bio-compatible scaffolds," Opt. Express 21, 17028-17041

 6. Lauffenburger, D.A. and Griffith, L.G. (2001) Who's got pull around here?
- Cell organization in development and tissue engineering. Proc. Natl. Acad. Sci. U. S. A. 98, 4282-4284
 Thompson, R., Reckova, M., deAlmeida, A., Bigelow, M., Stanley, C., Spruill, J., & Sedmera, D. (2003). The oldest, toughest cells in the heart.
- Novartis Foundation Symposium, 250157.8. 3D Printing in Medicine: How Technology Will Save Your Life. (2013), http://www.cgtrader.com/blog/wp-content/uploads/2013/08/
- rapid_prototyping_biology.jpg

 9. "3D Printer Used to Make Bone-Like Material" (2011). http://
 news.wsu.edu/2011/11/29/3d-printer-used-to-make-bone-like-material-
- video/#.UqPNfI0THLI

 10. "OsteoFab Patient Specific Cranial Device Receives 510(k) Approval OsteoFab Implants Ready for US Market and Beyond" (2013). Oxford
 Performance Materials Press Release, http://www.oxfordpm.com/news/
 article/2013-02-18_osteofab_patient_ specific_cranial_device_receives_

510k_approval_-_osteofab_implants_ready_for_us_market_and_beyond

- 11. Bose S, Vahabzadeh S, Bandyopadhyay A. (2013) Bone tissue engineering using 3D printing. *Materials Today*;16(12):496-504.
- 12. "Replacement Organs and Tissue: Anal Sphincters", Wake Forest School of Medicine, (2013). http://www.wakehealth.edu/Research/WFIRM/Projects/Anal-Sphincters.htm
- 13. "Replacement Organs and Tissue: Engineering a Kidney", Wake Forest School of Medicine, (2013). http://www.wakehealth.edu/Research/WFIRM/Research/Engineering-A-Kidney.htm
- 14. "Organovo Achieves One Month Performance, Drug Responsiveness for 3D Bioprinted Liver Tissues" (2013). http://ir.organovo.com/news/press-releases/press-releases-details/2013/Organovo-Achieves-One-Month-Performance-Drug-Responsiveness-for-3D-Bioprinted-Liver-Tissues/default.aspx

FUTURE TRENDS AND CHALLENGES

- The next step for 3D printing body parts is the creation of more complicated solid organs, specifically with large internal vascular networks. The following organs are currently in development:
 - Sphincters 3D printed sphincters have been implanted into mice, and have developed their own native blood supply and function 12
 - **Kidneys** Artificial kidneys made with kidney cells have been shown to be functional in mice¹³
 - Livers In late 2013, it was demonstrated that 3D printed liver tissue maintained key liver function for 40 days 14

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