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The development of bioresorbable composite polymeric scaffolds with high mechanical strength

Upma Sharma¹ PhD, Danny Concagh¹ MS, Lee Core¹ MSE, Yina Kuang¹ PhD, Changcheng You¹ PhD, Quynh Pham¹ PhD, Greg Zugates¹ PhD, Rany Busold¹ BS, Stephanie Webber¹ BS, Jonathan Merlo¹ BS, Robert Langer PhD³, George Whitesides PhD², Maria Palasis¹ PhD

¹480 Biomedical, Inc., Watertown, MA 02472, USA
²Massachusetts Institute of Technology, Cambridge, MA
³Harvard University, Cambridge, MA

Supporting Information: Supplementary Figures and Tables



Supplemental Figure 1. Example radial force curve of the elastomer coated braid. The data is of the first two cycles. The shaded region indicates target vessel diameter range for implantation.



Supplemental Figure 2. Schematic illustrating the mechanisms of polymer degradation. Hydrolysis of (a) ester, (b) urethane, and (c) urea segments of polyester urethane/urea elastomers.



Supplemental Figure 3. Retention of mechanical properties of 75:25/PLCL and 85:15/PLCL implant designs. (A) Radial resistive force as a percentage of baseline over time. (B) Chronic outward force as a percentage of baseline over time. Each data point represents the mean and standard deviation of an n=5 samples.



Supplemental Figure 4. Cyclic bend fatigue testing on 85:15/PLCL implants. Each data point represents the mean and standard deviation of an n=3 samples. The arrow indicates the point at which mechanical integrity is loss.



Supplemental Figure 5. Effect of coating weight on implant mechanical properties. Data shown represents the mean and standard deviation on an n=3 samples.



Supplemental Figure 6. *In vitro* degradation testing of implants. Data shows comparison of resorption under physiological conditions (pH = 7.4) and accelerated conditions (pH=12). Data shown represents the mean and standard deviation on an n=3 samples.

Supplemental Table 1. Impact of PGCL molecular weight and cross-link density on material properties of elastomer films

| PGCL Molecular Weight | PGCL:HDI Ratio (wt:wt) | E (MPa) | $\sigma_{ m max}$ (MPa) | & (%) |
|--------------------------|---------------------------|---------|-------------------------|-------|
| 20,000 | 12:1 | 5.7 | 5.4 | 350 |
| 20,000 | 12:4 | 43.2 | 7.5 | 320 |
| 100,000 | 20:1 | 4.2 | 7.3 | 820 |
| 100,000 | 20:2 | 3.9 | 9.0 | 860 |
| 100,000 | 20:3 | 6.0 | 6.7 | 700 |
| 100,000 | 20:4 | 10.1 | 9.6 | 810 |

E = Young's Modulus, σ_{max} = Maximum tensile strength, ε_{b} = Break strain All measurements carried out at room temperature in air

SUPPLEMENTARY INFORMATION

Supplemental Table 2. Details of implantation procedure results and comparison of histological scores between 10:90/PGCL, 75:25/PGCL, and 75:25/PLCL implants in swine.

| Parameter | Timepoint | Group | | | |
|----------------------------|-----------|---------------|-------------|---------------|--|
| | | 10:90/PGCL | 75:25/PGCL | 75:25/PLCL | |
| Balloon-to-artery ratio | 1 mo | 1.20 ± 0.04 | 1.21 ± 0.02 | 1.12 ± 0.03 | |
| | 3 mo | 1.25 ± 0.04 | 1.23 ± 0.02 | 1.12 ± 0.04 | |
| | 6 mo | N/A | 1.19 ± 0.02 | 1.11 ± .03 | |
| Stent-to-artery ratio | 1 mo | 0.98 ± 0.06 | 1.01 ± 0.09 | 1.02 ± 0.09 | |
| | 3 mo | 1.04 ± 0.11 | 1.01 ± 0.07 | 1.01 ± 0.05 | |
| | 6 mo | N/A | 1.04 ± 0.10 | 1.00 ± 0.04 | |
| Injury score | 1 mo | 0.4 ± 0.4 | 0.3 ± 0.4 | 0.3 ± 0.1 | |
| | 3 mo | 1.5 ± 0.5 | 2.3 ± 0.4 | 0.6 ± 0.7 | |
| | 6 mo | N/A | 0.7 ± 0.1 | 1.0 ± 1.1 | |
| Inflammation score | 1 mo | 3.7 ± 0.7 | 3.1 ± 1.0 | 0.7 ± 0.3 | |
| | 3 mo | 3.1 ± 0.2 | 3.7 ± 0.5 | 2.0 ± 1.0 | |
| | 6 mo | N/A | 3.0 ± 0.0 | 2.8 ± 0.3 | |
| Fibrin score | 1 mo | 1.3 ± 0.9 | 1.1 ± 0.6 | 0.0 ± 0.0 | |
| | 3 mo | 0.0 ± 0.0 | 0.0 ± 0.0 | 0.0 ± 0.0 | |
| | 6 mo | N/A | N/A | 0.0 ± 0.0 | |

Supplemental Table 3. Details of implantation procedure results and comparison of histological scores between 75:25/PLCL and 85:15/PLCL implants in sheep.

| Parameter | Timepoint | Group | | |
|-----------------------|-----------|---------------|-----------------|--|
| | | 75:25/PLCL | 85:15/PLCL | |
| Dellassia | 1 mo | 1.15 ± 0.07 | 1.13 ± 0.04 | |
| | 3 mo | 1.16 ± 0.05 | 1.13 ± 0.03 | |
| ratio | 6 mo | 1.16 ± 0.05 | 1.17 ± 0.07 | |
| Tallo | 12 mo | 1.17 ± 0.05 | 1.14 ± 0.08 | |
| | 18 mo | N/A | 1.17± 0.07 | |
| | 1 mo | 1.05 ± 0.06 | 1.01 ± 0.05 | |
| | 3 mo | 1.11 ± 0.20 | 1.02 ± 0.09 | |
| Stent-to-artery ratio | 6 mo | 1.02 ± 0.04 | 1.02 ± 0.03 | |
| | 12 mo | 1.05 ± 0.06 | 1.04 ± 0.08 | |
| | 18 mo | N/A | 1.09 ± 0.12 | |
| | 1 mo | 0.1 ± 0.1 | 0.1 ± 0.1 | |
| | 3 mo | 0.1 ± 0.1 | 0.1 ± 0.1 | |
| Injury score | 6 mo | 0.1 ± 0.1 | 0.1 ± 0.1 | |
| | 12 mo | 0.3 ± 0.3 | 0.6 ± 0.3 | |
| | 18 mo | N/A | 0.5 ± 0.2 | |
| Inflammation score | 1 mo | 1.1 ± 0.2 | 0.6 ± 0.6 | |
| | 3 mo | 2.4 ± 0.4 | 1.2 ± 0.3 | |
| | 6 mo | 1.5 ± 0.6 | 1.5 ± 0.2 | |
| | 12 mo | 1.8 ± 0.7 | 2.4 ± 0.3 | |
| | 18 mo | N/A | 2.1 ± 0.7 | |
| Fibrin score | 1 mo | 0.0 ± 0.0 | 0.0 ± 0.0 | |
| | 3 mo | 0.0 ± 0.0 | 0.0 ± 0.0 | |
| | 6 mo | 0.0 ± 0.0 | 0.0 ± 0.0 | |
| | 12 mo | 0.0 ± 0.0 | 0.0 ± 0.0 | |
| | 18 mo | N/A | 0.0 ± 0.0 | |

Supplemental Table 4. Summary of polymer characterization information.

| Implant designation | Base braid | | Elastomer coating | | |
|------------------------|-------------|-----------------------|------------------------|--|---------------------------------|
| | Composition | Inherent Viscosity | Prepolymer composition | Prepolymer molecular weight (Mn) | Prepolymer/HDI ratio (wt/wt) |
| 10:90/PGCL | 10:90 PLGA | 1.53 | 50:50 PGCL | 49600 | 10:1 |
| 75:25/PGCL | 75:25 PLGA | 1.91 | 50:50 PGCL | 49600 | 10:1 |
| 75:25/PLCL | 75:25 PLGA | 1.91 | 40:60 PLCL | 54100 | 10:1 |
| 85:15/PLCL | 85:15 PLGA | 1.80 | 40:60 PLCL | 54100 | 10:1 |

Note: No residual solvents or residual HDI were detected using gas chromatography. The residual ethylene oxide levels after sterilization was less than 100 ppm.

Supplemental Table 5. Semi-quantitative scoring scheme for histological analysis.

| Attribute | Score | Description of assigned weight | | |
|--------------|-------|---|--|--|
| Injury score | 0 | IEL intact, endothelium typically denuded, media may be compressed but not lacerated | | |
| | 1 | IEL lacerated, media typically compressed but not lacerated | | |
| | 2 | IEL lacerated, media visibly lacerated, EEL intact but may be compromised | | |
| | 3 | EEL lacerated, typically large lacerations of media extending through Eel, coil wires sometimes residing in adventitia | | |
| Inflammation | 0 | <25% struts with fewer than 10 inflammatory cells | | |
| | 1 | Up to 25% struts with greater than 10 inflammatory cells | | |
| | 2 | 25-50% struts with greater than 10 inflammatory cells | | |
| score | 3 | >50% struts with greater than 10 inflammatory cells | | |
| | 4 | 2 or more loops (or 4 or more struts) with associated granulomatou inflammatory reaction | | |
| Fibrin Score | 0 | No fibrin is appreciated (or only small strands) | | |
| | 1 | At least 25% of struts involving confluent fibrin that surrounds up to 25% of the strut circumference | | |
| | 2 | At least 50% of struts involving confluent fibrin that surrounds >25% of strut circumference | | |
| | 3 | ALL struts with confluent fibrin surrounding >50% of strut circumference OR (2) Confluent fibrin involving >25% of strut circumference with involvement >50% of struts AND extension between struts or bridging. | | |

IEL – Internal elastic lamina; EEL- External elastic lamina