

## **Corporate Overview**

July 2024



### **Forward-Looking Statements**

This presentation contains forward-looking statements that involves substantial risks and uncertainties. Any statements about the company's future expectations, plans and prospects, including statements about its strategy, future operations, development of its product candidates, and other statements containing the words "believes," "anticipates," "plans," "expects," "estimates," "intends," "predicts," "projects," "targets," "could," "may," and similar expressions, constitute forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995, although not all forward-looking statements regarding: expectations regarding the timing for the commencement and completion of product development or clinical trials; the rate and degree of market acceptance and clinical utility of the company's products; the company's commercialization, marketing and manufacturing capabilities and strategy; the company's intellectual property position and strategy; the company's estimates regarding expenses, future revenue, capital requirements and needs for additional financing; developments relating to the company's competitors and industry; and the impact of government laws and regulations.

Actual results may differ materially from those indicated by such forward-looking statements as a result of various important factors, including: the ability to develop commercially viable product formulations; the sufficiency of the company's cash resources; the ability to obtain necessary regulatory and ethics approvals to commence additional clinical trials; whether data from early clinical trials will be indicative of the data that will be obtained from future clinical trials; whether the results of clinical trials will warrant submission for regulatory approval of any investigational product; whether any such submission will receive approval from the United States Food and Drug Administration or equivalent foreign regulatory agencies and, if we are able to obtain such approval for an investigational product, whether it will be successfully distributed and marketed. These risks and uncertainties, as well as other risks and uncertainties that could cause the company's actual results to differ significantly from the forward-looking statements contained herein, are discussed in our annual report on Form 10-K for the year ended December 31, 2023, and other filings with the SEC which can be found at www.sec.gov. Any forward-looking statements contained in this presentation speak only as of the date hereof and not of any future date, and the company expressly disclaims any intent to update any forward-looking statements, whether as a result of new information, future events or otherwise.

Photo: Gertrude "Trudy" Elion, inventor of azathioprine and recipient of Nobel Prize in Medicine in 1988.

## **Eledon Company Highlights**



#### Optimized & Differentiated Lead Asset

- Tegoprubart has demonstrated the ability to:
  - Engage B and T Cell targets
  - Decrease pro-inflammatory biomarkers
  - Prevent both human-to-human and pigto-human organ transplant rejection
  - Achieve high kidney function (eGFR) levels post-transplant
- 100+ subjects of safety data
- Engineered to increase potency, half-life and manufacturability vs. other anti-CD40 approaches



#### Strong Financial Profile

- \$42.9M in cash, cash equivalents and short-term investments as of March 31, 2024, and subsequently completed \$52M in Private Placement financing
- Potential for up to \$58M in additional financing from the 2023 Private Placement
- Expected sufficient to fund completion of the Phase 2 BESTOW trial



2H2024 Expected Milestones

- Full enrollment in Phase 2 BESTOW trial of tegoprubart in kidney transplantation expected at year end
- Continued enrollment in both the kidney transplant Phase 1B as well as in the investigator sponsored Phase 2 trial in islet cell transplantation for Type 1 diabetes



## Tegoprubart is a Pipeline in a Product Opportunity

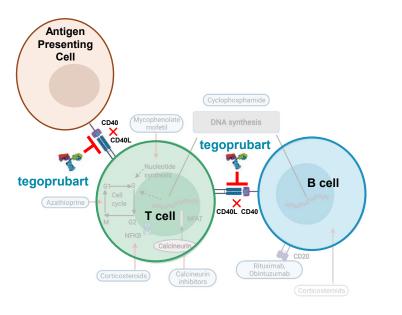
	DEVELOPMENT STAGE				
INDICATIONS	PRE-CLINICAL	Early Human Trials/ PHASE 1	PHASE 2	PHASE 3	NOTES
TRANSPLANTATION					
Kidney					<ul> <li>Phase 2 BESTOW, ex-US Phase 1b &amp; Long-Term Extensions trials ongoing</li> </ul>
Islet Cell					<ul><li>U. Chicago investigator sponsored trial</li><li>Funded by JDRF &amp; The Cure Alliance</li></ul>
Liver					
XENOTRANSPLANTATION					
Kidney					<ul> <li>Performed under U.S. FDA Expanded Access Protocol (EAP)</li> </ul>
Heart					<ul> <li>Performed under U.S. FDA Expanded Access Protocol (EAP)</li> </ul>
NEUROINFLAMMATION					
Amyotrophic Lateral Sclerosis (ALS)					<ul> <li>Seeking non-equity dilutive financing to advance program to Phase 3</li> </ul>





## Mechanism Overview of CD40L Inflammatory Signaling

#### CD40/CD40L Pathway and Tegoprubart Site of Action



- Interaction of CD40 with CD40L on immune cells mediates activation of the co-stimulatory immune pathway, controlling "cross talk" between the adaptive and innate immune systems
- Maximal activation of inflammatory system is a 3-step process requiring co-stimulatory signaling
  - Step 1: Major histocompatibility complexes (MHC) and CD3/TCR engagement
  - Step 2: CD40 and CD40L binding resulting in cell division and clonal expansion
  - **Step 3:** Pro-inflammatory response by polarized T cells expressing inflammatory chemokines and cytokines
- Blocking CD40L shifts polarization away from proinflammatory signaling to T cell anergy, apoptosis, and polarization to a Treg environment
  - Blocking CD40L thus does not generally result in lymphopenia often seen with immunosuppressive agents



## Tegoprubart: Potential Best-in-Class Anti-CD40/CD40L Antibody

Targeting CD40 Ligand vs. CD40 Receptor		lgG1 vs. fusion protein	
CD40L and CD40	CD40L only	or pegylated FAB	
Targeting both anti-CD40L and anti-CD40 inhibits	<ul> <li>Blocking anti-CD40L also inhibits CD11 costimulatory receptors on antigen presenting cells, thus blocking the pro-inflammatory polarization of CD8<sup>+</sup> Cytotoxic T cells</li> </ul>	✓ Up to over 2x times longer half-life	
B cell polarization and class switching, as well as inhibits the pro-inflammatory polarization of CD4 <sup>+</sup>	<ul> <li>Blocking CD40L also polarizes CD4<sup>+</sup> lymphocytes to FoxP3<sup>+</sup> Regulatory T cells (Tregs), thus creating a potentially more tolerogenic environment</li> </ul>	✓ Manufacturing advantages	
Helper T cells	<ul> <li>CD40L is more selectively expressed, providing the potential for additional safety and PK/PD/dosing advantages</li> </ul>	✓ Less anti-drug antibodies	





## Our Mission: One Transplant for Life

Eledon is committed to ensuring the greatest gift will keep on giving

#### We are tackling an urgent & significant need...

25,000 kidney transplants per year in the U.S. with 90,000+ Americans on the kidney transplant waiting list

Average age at transplant is 50 years old but an average kidney graft only functions 10-15 years

1 in 6 of the 90,000+ Americans on the kidney transplant waiting list have already had a transplant

Alternative to kidney transplant is dialysis which has an under 50% five-year survival rate

5,000 Americans per year die waiting for a kidney transplant

#### ... with tegoprubart, our lead asset in Phase 2

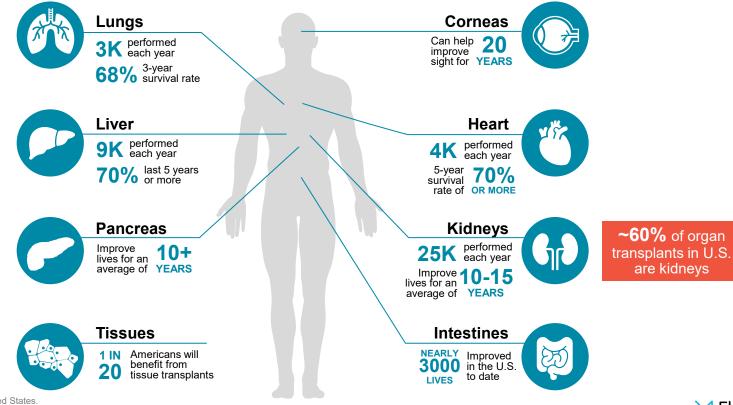
Demonstrated ability in a Phase 1b clinical trial to protect transplanted kidneys and achieve high kidney function (eGFR)

Used to prevent rejection in pig-to-human cardiac & kidney xenotransplantation, opening a potential new source of organs

100+ subjects of safety data in multiple indications



# Every Transplant Begins with Altruism & Each Donor Can Provide Multiple Organ Types

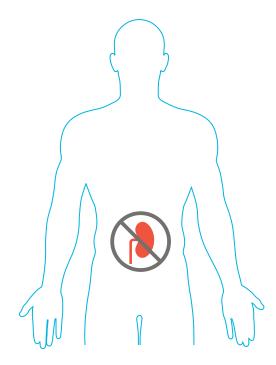


Note: Numbers are for the United States. Sources: golm.org; USDHHS.



8

## Immunosuppression is Necessary to Protect Transplanted Organs



Without immunosuppression, the **host** sees **donor** kidney as "foreign" and attacks (i.e., rejects) it

Immunosuppression must be taken for life or the organ will be rejected even years after the transplant



Kidney Transplantation is a Large but Highly Concentrated Market

Large patient population				
	25,000+	21,000+	Kidney transplants annually	
<b><sup></sup><u>mnnnnnnnnnnnnnnnn</u>nnn</b>	255,000+	188,000+	People living with a functioning kidney transplant	

90,000+ Americans on transplant waiting list

5,000 Americans per year die waiting for a kidney transplant
~15% of U.S. adults on waitlist are waiting for repeat transplants

Limited expert centers and doctors



~250 transplant centers in the U.S.



~1,200 surgeons perform abdominal organ transplants in the U.S.



10

Kidney Transplantation Immunosuppression Market Represents a Multi-Billion Dollar Commercial Opportunity

### End stage renal disease & transplant

**\$50+ Billion** annual U.S. Medicare expenditure including Kidney Transplantation costs of **\$440,000+ / transplant** 

Medicare covers cost of immunosuppressive transplant drugs, regardless of patient age, if patient does not have other insurance Global organ transplant immunosuppressant market size estimated \$5.3+ billion



Astellas reported tacrolimus global revenues of over \$1.4B in FY2023 (Prograf, first FDA approval 1994)

### Graft failure of transplanted kidneys is the norm and expensive



Average age of transplant is **50 years old** <u>but</u>



Average organ only functions 10-15 years so patients return to dialysis or need a repeat transplant Patients returning to dialysis: costs \$100,000+, ▼quality of life, <50% 5-year survival rate

**Re-transplants deplete** an already inadequate **donor organ pool** 

#### \$150,000+

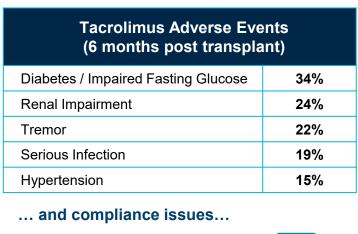
average incremental U.S., medical costs / patient year after graft failure



Over 30-50% of Kidney Transplants Fail Within 10 Years on Current Transplant Standard of Care Immunosuppression

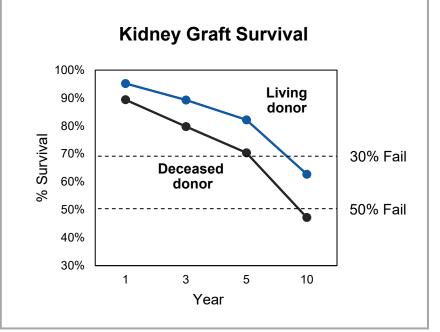
# Transplant immunosuppression has high toxicity...

## ...so Kidney Transplants only typically function 10-15 years



#### Up to 15 pills/day for life

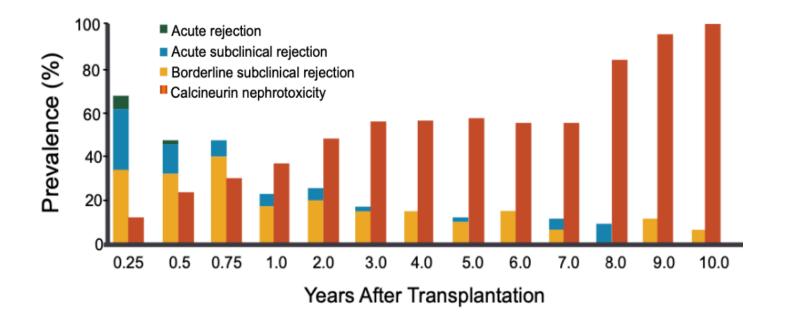
Nearly all patients miss doses and ~25% of patients are considered non-adherent





12

# Calcineurin Inhibitor (CNI) Side Effects Play a Leading Role in Kidney Graft Pathology Over Time







# Removing CNIs May Ameliorate the Kidney Transplantation Graft Failure Cycle

### Kidney Transplantation and Graft Failure Cycle

#### Inflammation & CNI Associated Kidney Damage

- Nephrotoxicity
- Hypertension
- Diabetes

#### Transplant

\$440,000+ avg. cost per U.S. patient

#### **Graft Failure**

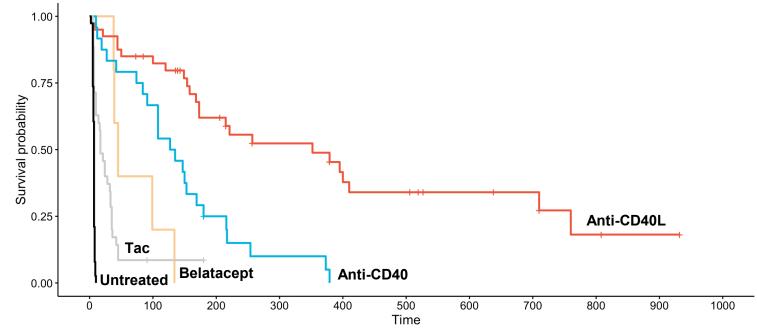
\$150,000+ avg. incremental medical costs per patient post graft failure

#### Dialysis & Kidney Wait List

- ~15% of adults on waitlist are for repeat transplants
- ~15% to 20% mortality rate in 1<sup>st</sup> year of dialysis



## Inhibition of CD40L Improved Survival vs. Other Approaches in Non-Human Primate (NHP) Kidney Transplantation Monotherapy Studies



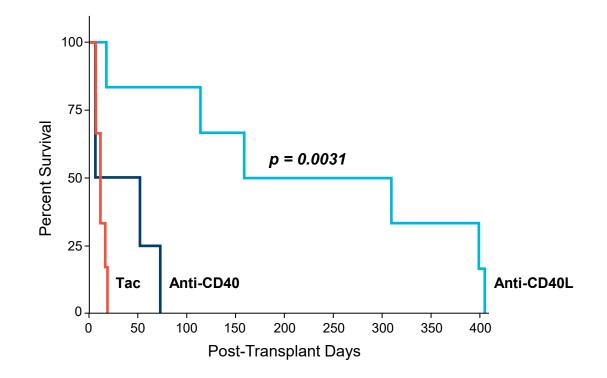
#### **NHP Survival Post Kidney Transplant**

Sources: Perrin, 2022; Song, 2014; Song, 2016; Duan, 2017.

Note: In aggregated data from published studies, NHPs receiving anti-CD40L (e.g., 5c8, AI794, IDEC-131) immunomodulation monotherapy post kidney transplantation had longer average survival than those receiving anti-CD40 monotherapy (e.g., 4D11, cH5D12, Chi220, ASKP1240), tacrolimus monotherapy or untreated controls. Meta-analysis not based on head-to-head studies. Differences between any individual programs may vary. Tac = tacrolimus.



NHP Renal Xenotransplantation Experience has Demonstrated Advantage of Blocking CD40L vs. Tacrolimus or CD40





# Phase 1b and Phase 2 Kidney Transplantation Studies are Enrolling in Parallel

Dhaaa 4h	52-week, open label, single arm study	Primary endpoints:
Phase 1b	ATG induction therapy plus	<ul> <li>Safety &amp; tolerability</li> </ul>
Up to 24 participants undergoing kidney transplantation	CNI-free maintenance therapy with tegoprubart	<ul> <li>Secondary endpoints:</li> <li>Graft function (eGFR)</li> <li>Participant and graft survival</li> </ul>
Canada, UK and Australia	(as a replacement for tacrolimus) as part of a maintenance immunosuppressive regimen including mycophenolate and a corticosteroid taper	<ul> <li>Biopsy proven acute rejection (BPAR)</li> <li>Immune cell infiltrate of graft biopsy</li> <li>Biomarker measures of kidney injury and rejection ris</li> </ul>

#### Phase 2 "BESTOW"

~120 participants (60/arm) undergoing kidney transplantation

U.S. and other countries

#### 52-week, head-to-head, superiority study

ATG induction therapy plus

#### CNI-free maintenance therapy with tegoprubart or tacrolimus

as part of a maintenance immunosuppressive regimen including mycophenolate and a corticosteroid taper

#### **Primary endpoints:**

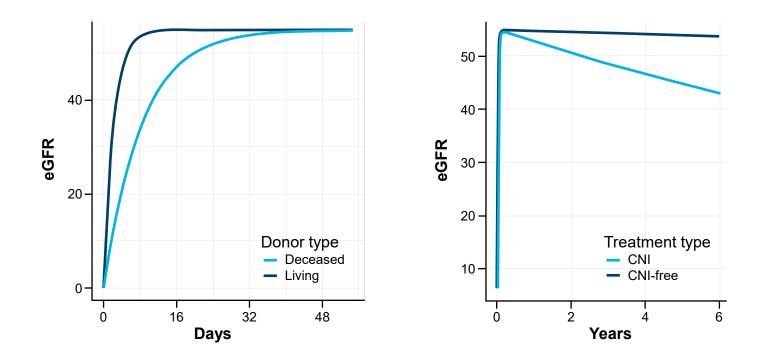
- Graft function (eGFR)
- Safety & tolerability

#### Secondary endpoints:

- · Participant and graft survival
- Biopsy proven acute rejection (BPAR)
- Immune cell infiltrate of graft biopsy
- Rate of new onset diabetes mellitus (NODAT)
- Biomarker measures of kidney injury and rejection risk



## eGFRs Over Time Post Transplant: Mean ~53 mL/min/1.73m<sup>2</sup> After 12 Months Using CNIs

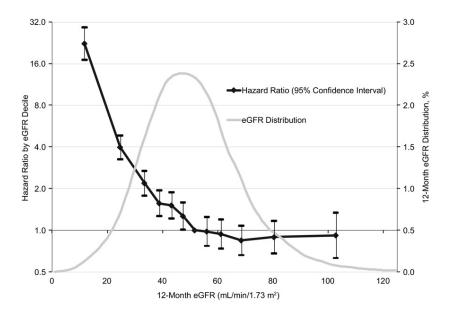


Note: **n = 4,868 patients**. eGFR estimated using MDRD 4-variable GFR Equation. Source: Kosinski et al. *Clin Transl Sci*. 2023 Nov; 00:1–11.



Kidney Allograft Function is an Early Predictor of Future Graft Failure

eGFR at 12 months is associated with 10-year death-censored graft failure (n = 13,661 patients)



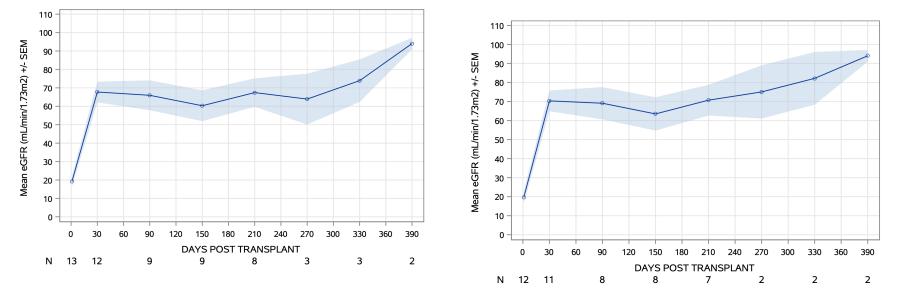
- Median eGFR of 50 mL/min/1.73m2 at 12 months (95th percentile of 83 mL at 12 months)
- Graft function measured using eGFR at 12 months post transplant is associated independently with subsequent graft failure
- Of multiple covariates,12-month eGFR is the strongest predictor of graft failure



## Phase 1b Kidney Transplantation Aggregate Mean eGFR

#### **All Subjects**

#### Excluding Subject with Surgical Complication (ATN) on Day 0



Note: ATN = Acute Tubular Necrosis. Estimated glomerular filtration rate (eGFR) as of March 2024, calculated using the chronic kidney disease epidemiology collaboration (CKD-EPI) creatinine equation. N is the number of participants at that time contributing data to mean eGFR calculation. Graphs use end of treatment last value.

Source: Company data.



## Phase 1b Kidney Transplantation: Demographics and Disposition

Age/ Gender	Ethnicity	Donor	Underlying Disease	Donor Age	HLA Mismatch
60/F	White	Living	Polycystic Kidney Disease	40	5
77/F	White	Deceased	Diabetes	42	5
62/M	White	Living	Cystic Disease	55	4
68/M	White	Living	Diabetes	57	4
23/F	Asian	Living	Glomerulonephritis	49	2
44/M	White	Deceased	Polycystic Kidney Disease	42	4
65/M	White	Living	Diabetes	64	2
57/F	White	Living	Diabetes	45	6
35/M	Other	Living	Glomerulonephritis	59	5
56/F	White	Living	Cystic Disease	54	4
59/M	White	Living	Diabetes	63	6
38/M	Asian	Deceased	FSGS	32	5
68/M	Other	Deceased	Diabetes	38	5

• Median recipient age: 59 y.o. (vs. CPI median of 51 y.o.)

• Median donor age: 49 y.o. (vs. CPI median of 44. y.o.)

• Mean HLA mismatch: 4.4 / 6



Note: Enrollment cut off for Demographics and Disposition is April 2024. Source: Kosinski (CPI) et al. *Clin Transl Sci.* 2023 Nov; 00:1–11.

## Phase 1b Kidney Transplantation: Treatment Emergent Adverse Events\*

System Organ	Preferred Term	N (%)
	Diarrhea	5 (38%)
Gastrointestinal	Constipation	5 (38%)
	Nausea	4 (31%)
	Vomiting	3 (23%)
	Polyomavirus (BK) viremia	5 (38%)
Infections	Cytomegalovirus viremia	2 (15%)
Infections	Upper respiratory tract infection	2 (15%)
	Urinary tract infection	2 (15%)
Presedural Complication	Transplant surgery complications	3 (23%)
Procedural Complication	Procedural pain	3 (23%)
Plead and Lymphotic System	Leukopenia	4 (31%)
Blood and Lymphatic System	Neutropenia	4 (31%)
Cardiac	Tachycardia	4 (31%)
0	Peripheral edema	2 (15%)
General	Pyrexia	2 (15%)
Matakaliana	Hypophosphatemia	3 (23%)
Metabolism	Hypoglycemia	2 (15%)
Musculoskeletal and Connective Tissue	Pain	4 (31%)
Skin and Subcutaneous tissue	Alopecia	2 (15%)
Vaccular	Hypertension	2 (15%)
Vascular	Hypotension	3 (23%)

\*Occurring in 2 or more study subjects as of April 2024. Of all the reported TEAEs, 7 events experienced by 3 subjects are reported as serious. These SAEs include neutropenia, acute kidney injury, T-cell rejection, Polyomavirus viremia, anterior abdominal wall collection, and hyperkalemia

## No cases of hyperglycemia, new onset diabetes or tremor

- 1 participant discontinued study on day 217 due to alopecia and fatigue, 1 participant discontinued study on day 54 due to Polyomavirus viremia, and 1 participant discontinued study on day 176 due to rejection (Banff score 2B)
- 1 participant experienced a surgical procedure related kidney damage (Acute Tubular Necrosis) on day 0, prior to tegoprubart administration, which impacted their kidney function. The subject remains in the study
- BK infections were controlled by temporarily decreasing immunosuppression, and CMV infections were controlled by antivirals



## Islet Cell Transplant Opportunity



~1.6M Americans live with Type 1 diabetes



~70,000 (5%) estimated to have "BT1D", the Brittle form of Type 1 Diabetes



BT1D patients have difficult-to-manage glucose levels with severe blood glucose fluctuations despite treatment and higher risk of diabetes related death



Currently, **islet cell transplantation is underutilized** in part due to immunosuppressive regimens with CNIs that may be toxic to transplanted insulin producing islet cells **Tegoprubart may unlock the islet cell transplant market** by potentially:

- Improving islet cell graft survival regardless of cell source (e.g., manufactured) and/or the use of a pouch
- 2. Reducing side effects associated with standard of care regimens including islet cell death



Phase 1/2 Study Assessing the Use of Tegoprubart to Prevent Islet Cell Transplant Rejection in Participants with Type 1 Diabetes

#### DESIGN

#### **PLANNED DATA GENERATION**

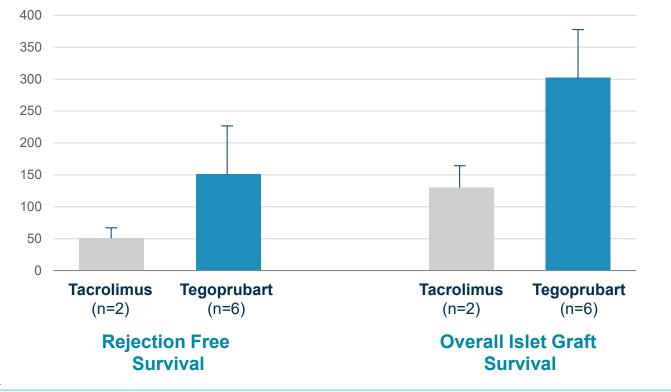
- 52-week, open label, single dose level study
- Initial group of up to 3 participants with Type 1 Diabetes (T1D) at the University of Chicago
- Islet cell transplant combined with induction therapy plus tegoprubart and mycophenolate mofetil (MMF) every third week by IV infusion
- Financing principally from the Juvenile Diabetes Research Foundation (JDRF) and The Cure Alliance

- Safety & tolerability
- Graft function & insulin independence
  - e.g., C-peptide, HbA1C
- Number of hypoglycemic events
- Need for repeat islet cell transplant(s)



Tegoprubart Prolonged Graft Survival vs. Tacrolimus Containing Regimen (CIS) in a Non-Human Primate Islet Cell Transplant Model...

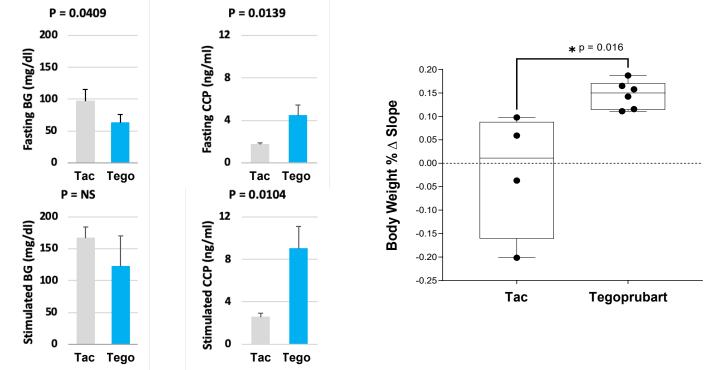
#### Mean Survival (Days)





... And Tegoprubart was Associated With Better Metabolic Control as Well as Healthier Animals Overall as Demonstrated by Body Weight

**Tegoprubart vs. Tacrolimus Regimens in NHP Islet Cell Transplantation Model** 





## **Eledon Pharmaceuticals**

19900 MacArthur Blvd., Suite 550 Irvine, California 92612, USA info@eledon.com +1 949-238-8090

