GeneTether Increasing Efficiency in Gene Editing

A final prospectus containing important information relating to the securities described in this document has been filed with the securities regulatory authorities in each of the provinces of British Columbia, Alberta and Ontario. A copy of the final prospectus, and any amendment, is required to be delivered with this document. This document does not provide full disclosure of all material facts relating to the securities offered. Investors should read the final prospectus and any amendment for disclosure of those facts, especially risk factors relating to the securities offered, before making an investment decision.

Corporate Presentation | March 21, 2022

Disclaimer

Reference to Prospectus

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These forward looking statements are based on a number of assumptions which may prove to be incorrect including, but not limited to: general economic, market and business conditions, the outcome of research studies, the ability to obtain certain approvals, the accuracy of cost estimates, ability to obtain sufficient capital on satisfactory terms, availability of equipment and supplies, changes in customer demand, currency exchange rates and the impact of changes in applicable laws and regulations. The forward looking statements contained in this presentation are made as of the date hereof or the dates specifically referenced in this presentation, where applicable. Except as required by law, GeneTether undertakes no obligation to update publicly or to revise any forward looking statements that are contained or incorporated in this presentation. All forward looking statements contained in this presentation are expressly qualified by this cautionary statement.

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GeneTether We are on a mission to develop new, curative therapies for patients with devastating genetic diseases using our GeneTether platform technology

Investment Highlights

Experienced Team

Disruptive Platform Technology

Rare Genetic Diseases

IP Portfolio

Focused on harnessing next generation technology to significantly increase efficiency of gene editing and potentially cure serious and life threatening genetic diseases Extensive public life science company experience Global capital markets experience and extensive investor network

> Highly efficient insertion of DNA into the genome for gene correction and complementation strategies Proof of concept studies showed ~7x higher gene editing efficiency using GeneTether compared to unmodified Cas9 > Expected to result in superior efficacy, safety, and flexibility

Pursuing curative therapies for rare genetic diseases Genetic kidney diseases that progress to chronic and end-stage kidney disease Life-threatening genetic skin diseases

> Wholly-owned intellectual property; no 3rd party financial obligations > 1 issued patent (Australia) Notice of Allowance from USPTO (February 2022) > 7 others pending

Experienced Life Sciences Team



Roland Boivin, MBA Chief Executive Officer & Director



R. Geoffrey Sargent, PhD Co-Founder & Chief Scientific Officer

Jean Jen, CPA, CA, MPAcc Chief Financial Officer





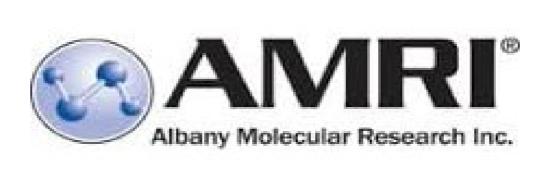
Peter Sampson, PhD Vice President, R&D

Kuldeep Neote, PhD Chair – Scientific Advisory Board Innovation/Strategy Consultant





























Experienced Board of Directors



William J. Garner, MD Co-Founder & Executive Director



Director

Director

Chairperson

Andre Fraga, Int. MBA

P. Gage Jull, PEng, MBA, CFA

Daren Graham, JD











Bordeaux Capital









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edp renewables



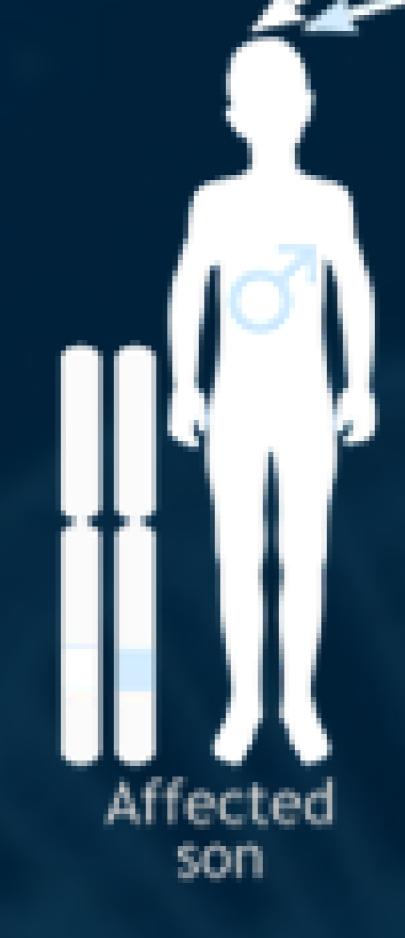


Approximately 10,000 diseases are known to be caused by aberrant DNA sequences that are inherited by one or both biological parents.

Traditional small molecule and biologic therapies have had limited success in treating many of these diseases because they fail to address the underlying genetic causes.

Recent advances in gene editing technologies provide the potential for curative therapies for many genetic diseases.

Genetic Diseases



Affected

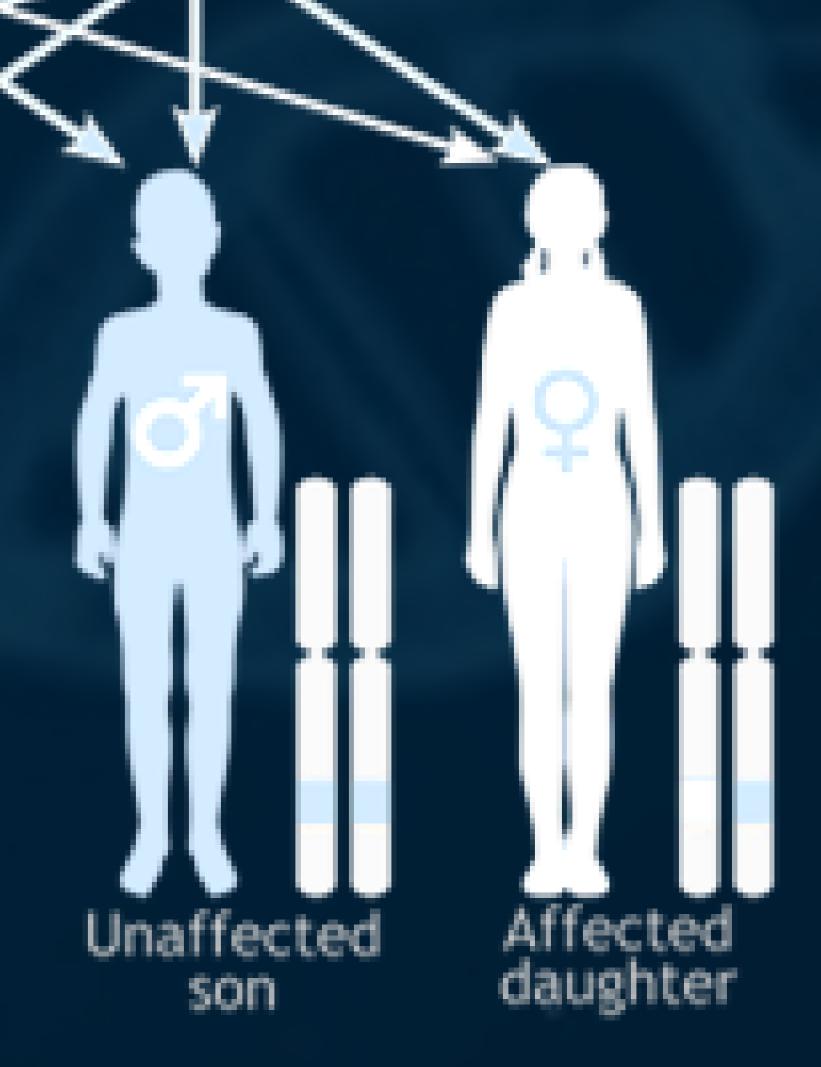
father



daughter

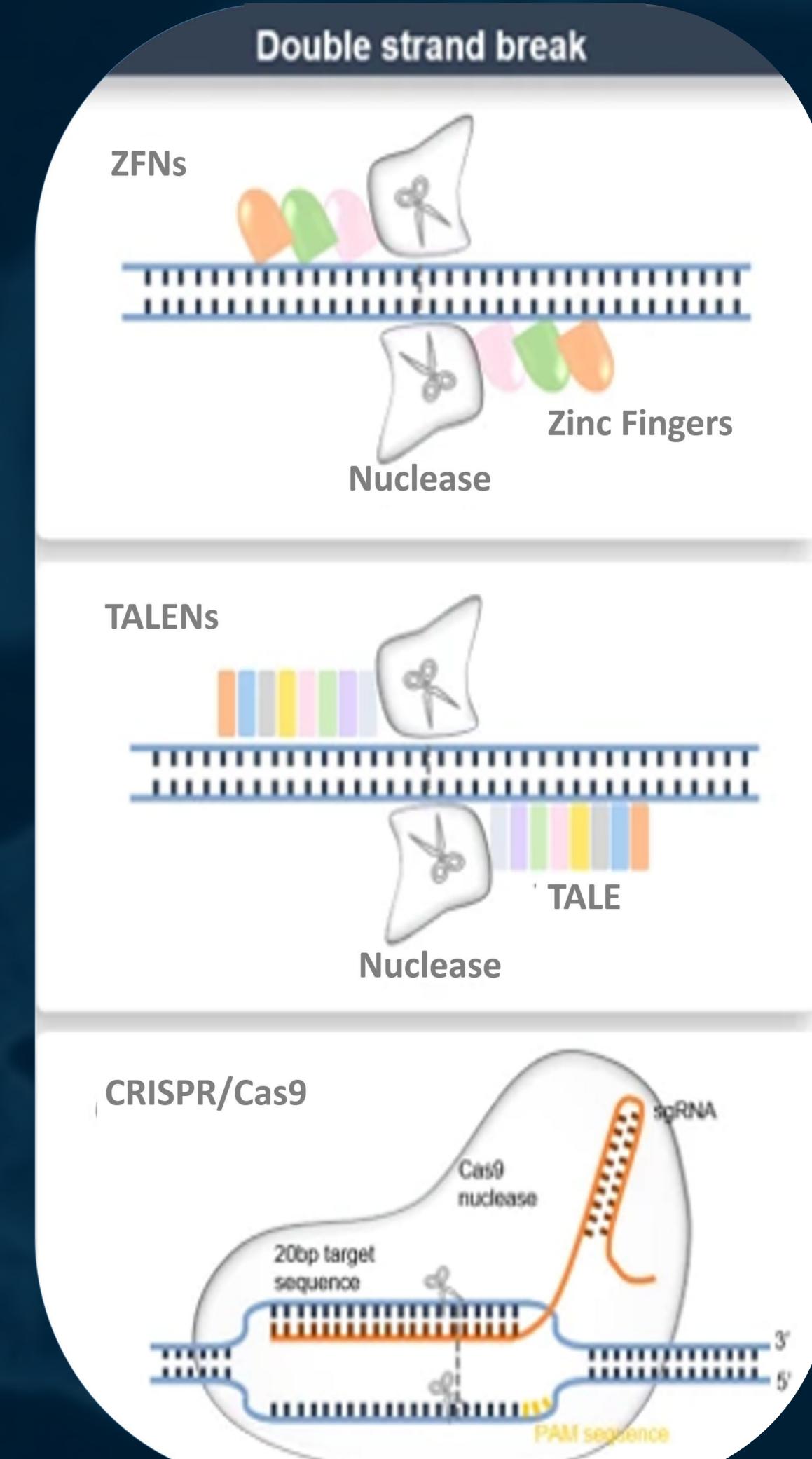
GeneTether Therapeutics **Corporate Presentation**

Unaffected mother



Unaffected Affected

Gene Editing – How it Works Creating double-strand breaks



A gene editing nuclease, CRISPR/Cas9 for example, is guided to a precise, predefined location in a cell's DNA where it creates a double-strand break (DSB).

Creating double-strand breaks is like a biological "find and delete" function.

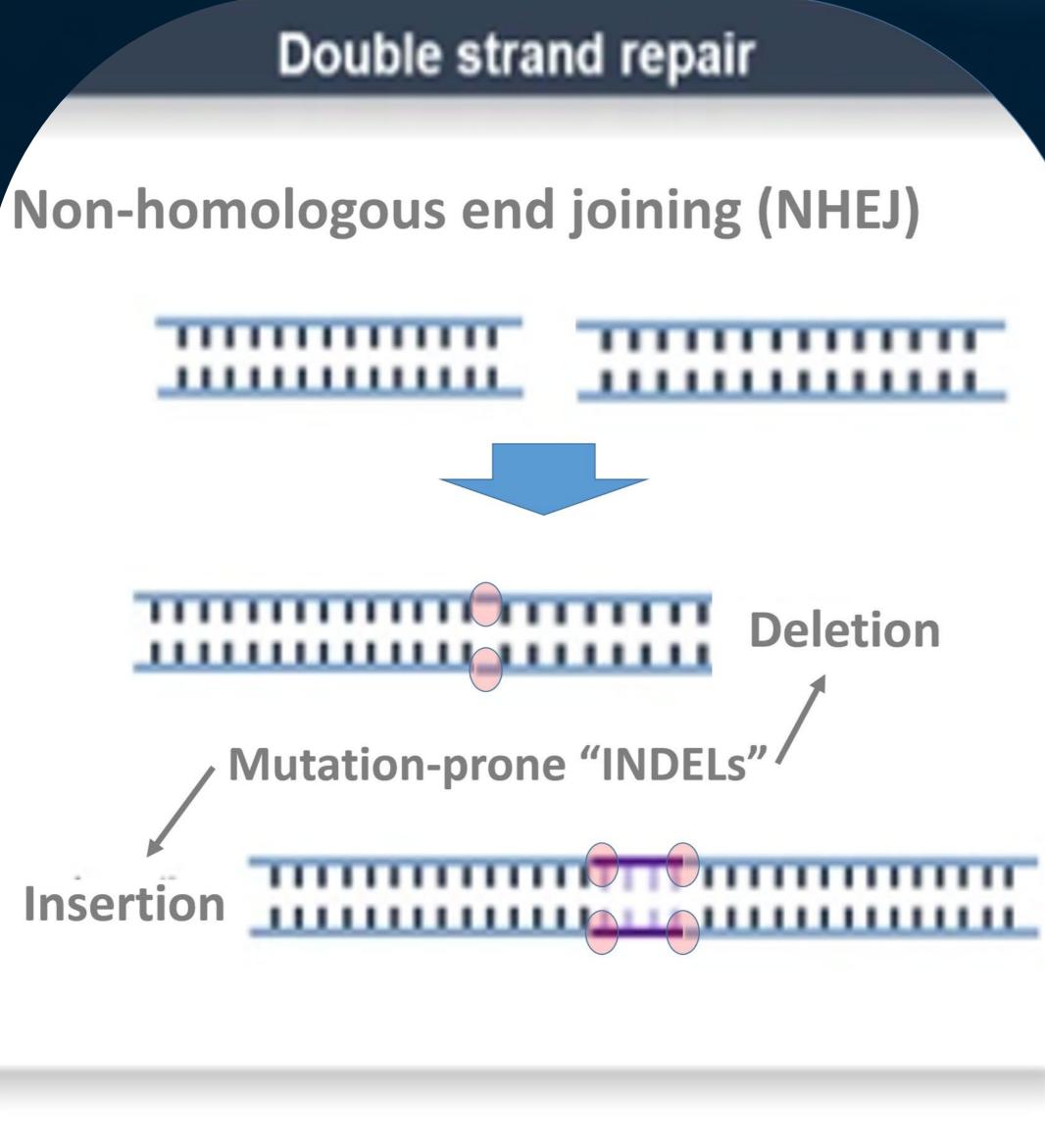
Major Gene Editing Technologies For creating double-strand breaks

CRISPR – A component of certain bacterial immune systems that is capable of guiding the system to matching sequences of DNA.

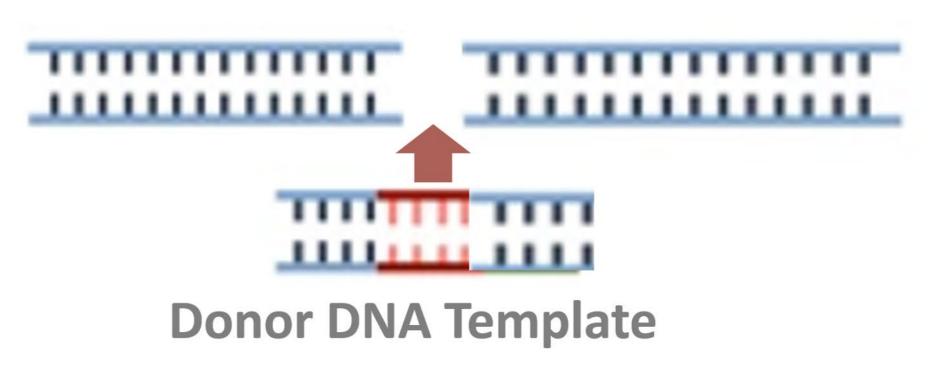
ZFN – Sequence-specific restriction enzymes generated by fusing a zinc finger DNA-binding domain to a DNA-cleavage domain.

TALEN – Sequence-specific restriction enzymes generated by fusing a transcription activator-like effector domain to a DNA-cleavage domain.

Gene Editing – How it Works Repairing double-strand breaks



Homology-directed repair (HDR)





Double-strand breaks are repaired by one of two competing cellular repair mechanisms: non-homologous end joining (NHEJ) or, in the presence of a DNA repair template, homology-directed repair (HDR).

Repair via HDR is like a biological "find and replace" function.

The Gene Editing Ecosystem Altering a DNA Sequence in an Endogenous Gene

.....

GeneTether's current focus

Correct a mutated gene to repair a

dysfunctional protein

Complement by inserting a missing gene to produce a needed protein

Delete

a mutated gene to halt the production of a diseasecausing protein

.........

GeneTether Therapeutics Corporate Presentation

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The Problem Current technologies for correcting or complementing aberrant genes are inherently inefficient

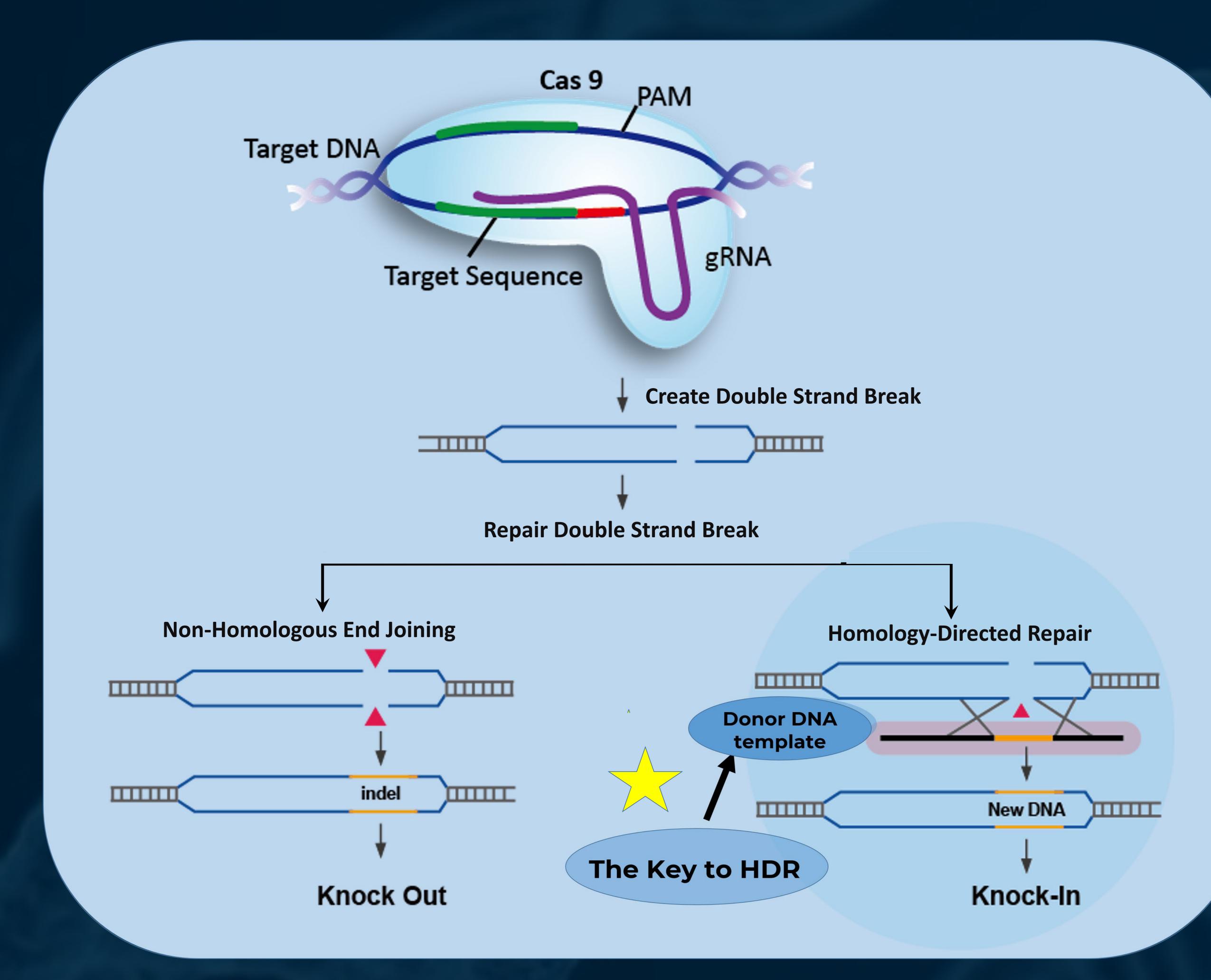
0.5 -20% Efficiency Rates

Efficiency: The ratio of gene edits actually made versus the maximum number that *could* have been made through the insertion of donor DNA templates.

Correcting or complementing with a donor DNA template requires that a strand break be repaired via HDR. HDR requires a donor DNA template in the immediate vicinity of a break.

Efficiency rates vary from gene to gene and from cell type to cell type, but all are currently below rates that make large scale, cost-effective commercialization feasible.

Gene Editing Efficiency Homology-Directed Repair vs Non-Homologous End Joining



Correcting and complementing genes requires delivery of a donor double strand break.

If a donor DNA template is not located near a double strand break, repair will not incorporate the donor DNA template via HDR.

The result is error-prone repair via NHEJ, leading to low gene editing efficiency, DNA mutation and rearrangements, and cell death.

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DNA template to the site of a DNA

The GeneTether Solution Proximity Matters

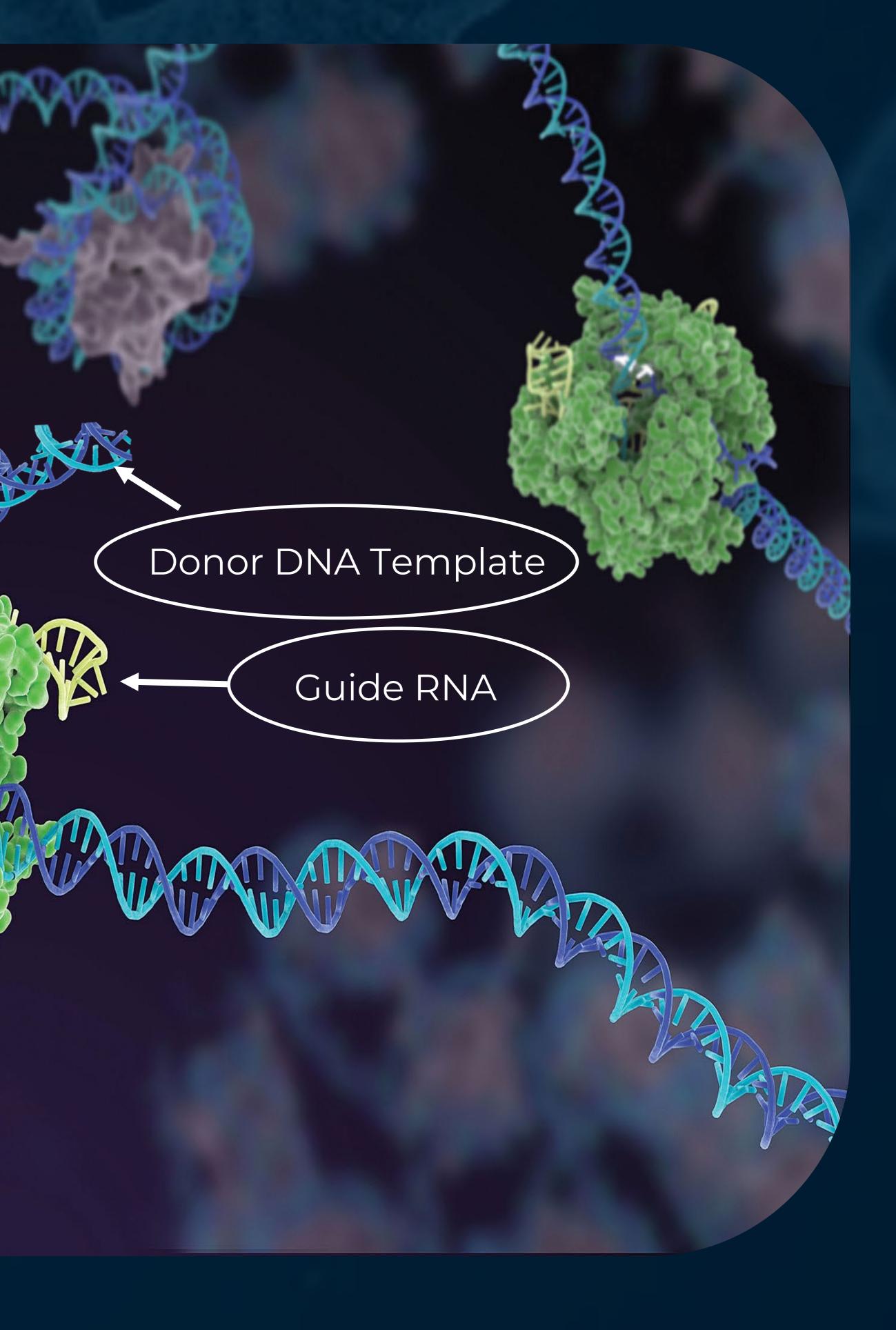
GeneTether has developed a proprietary method to attach, or "tether," donor DNA templates to gene editing nucleases.

The result is that the donor DNA template is nearby at the time a strand break is induced.

Correspondingly, there is an enhanced likelihood that **repair of the break will take place via HDR**, thereby allowing a far **greater number of gene edits** per payload delivery and **reducing the risk of mutagenesis or off-target gene edits**.

LacR/Cas9 Fusion

Target Genomic DNA



Proof of Concept Study Design Introducing the delF508 mutation with GeneTether platform + CRISPR/Cas9 versus untethered CRISPR/Cas9

Delivered to target cell

Edited CFTR Genomic Locus with delF508 deletion

"Cas9 – GeneTether – Donor DNA" Construct

EXON 11

EXON 11

Lactose repressor-Cas9 fusion protein

Unedited CFTR Genomic Wild Type Locus

500 bp donor fragment with delF508 deletion and 5' or 3' lacO sequence

Cas9 Directed Double Strand Break

Homology Directed Repair using the tethered 500 bp donor fragment with delF508 deletion

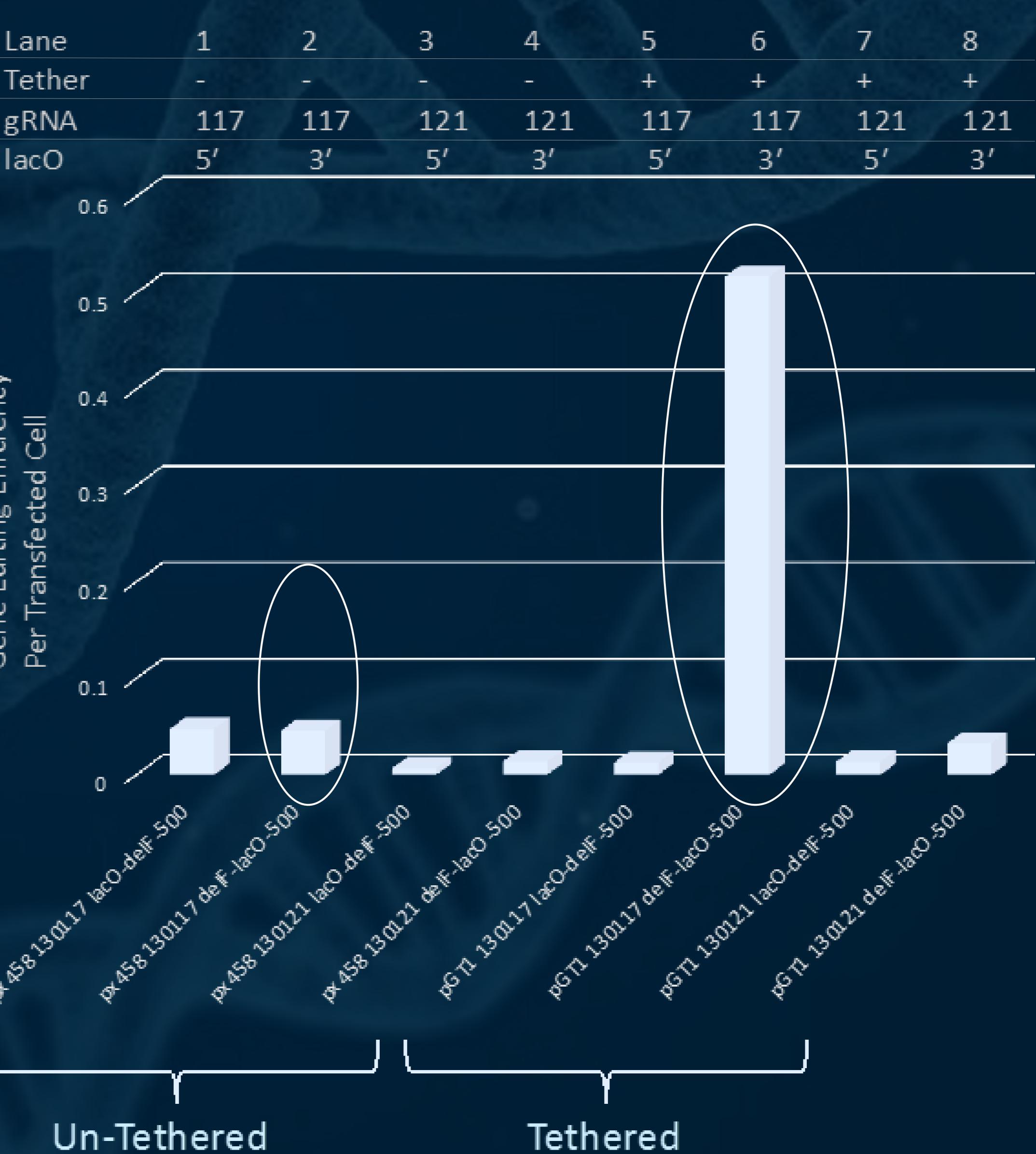






As shown, the lactose repressor-Cas9/GeneTether plasmid (pGT1) with the 130117 guide (lane 6) demonstrated a robust editing efficiency, resulting in ~7x more edits than the px458 vector with unmodified Cas9 and the same donor DNA fragment (lane 2).

Proof of Concept Study Results



5	6	7	8
+	+	+	+
117	117	121	121
5′	3′	5'	3′

Our Research Pipeline Therapeutic Programs

Nephrology

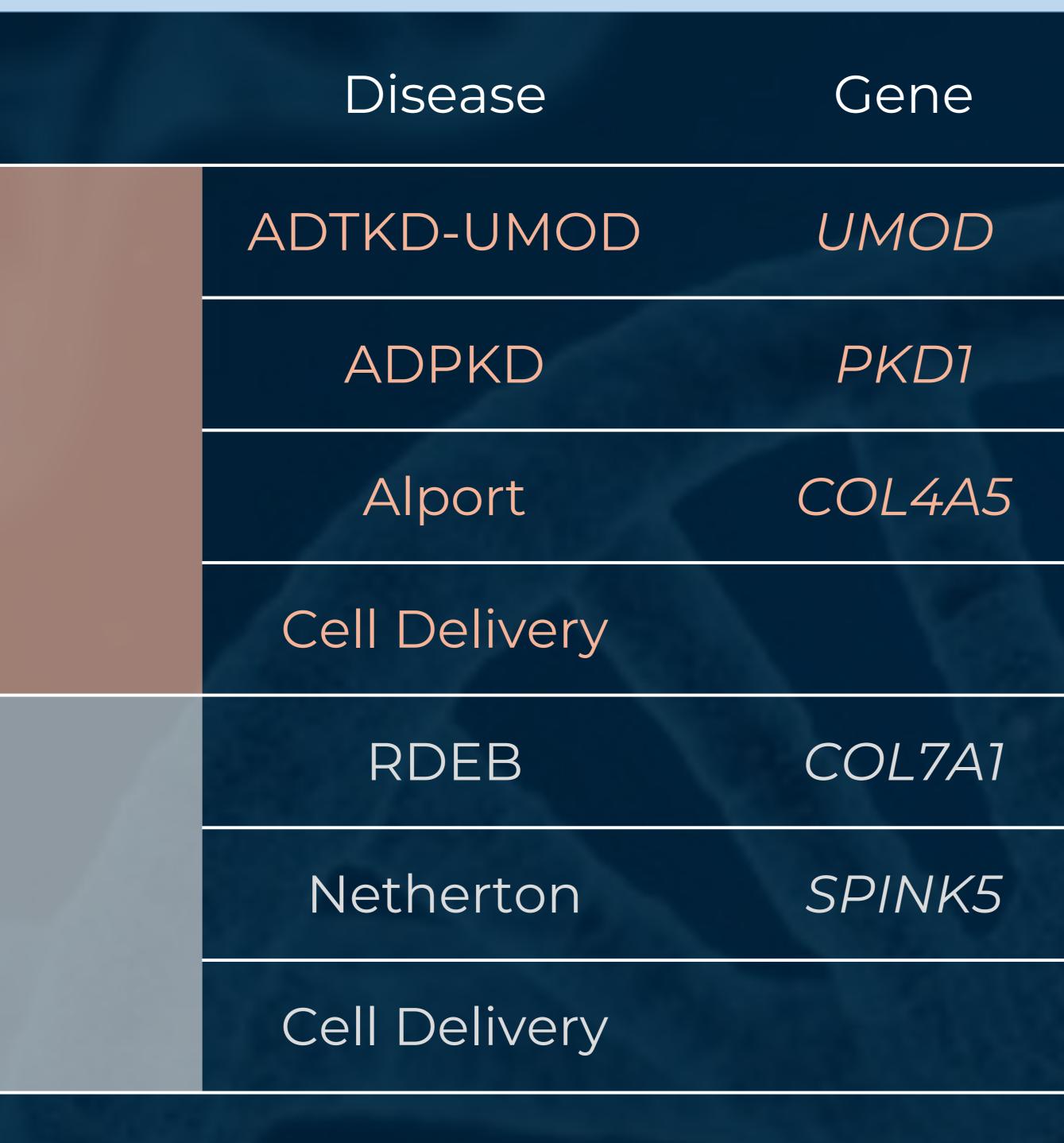
Dermatology

Platform & Intellectual Property Expansion

Large animal cell lines

Zebrafish

In vitro editing in human cell lines



Initiated

 \checkmark

Target Completion

Study Site

Q2 2022

Q2 2022

Ongoing

Target Discovery

Lead Target Selection



¹Target discovery includes identifying and/or developing cell line and animal models, conducting proof-of-concept studies, and identifying and/or developing tissue selective delivery vehicles.

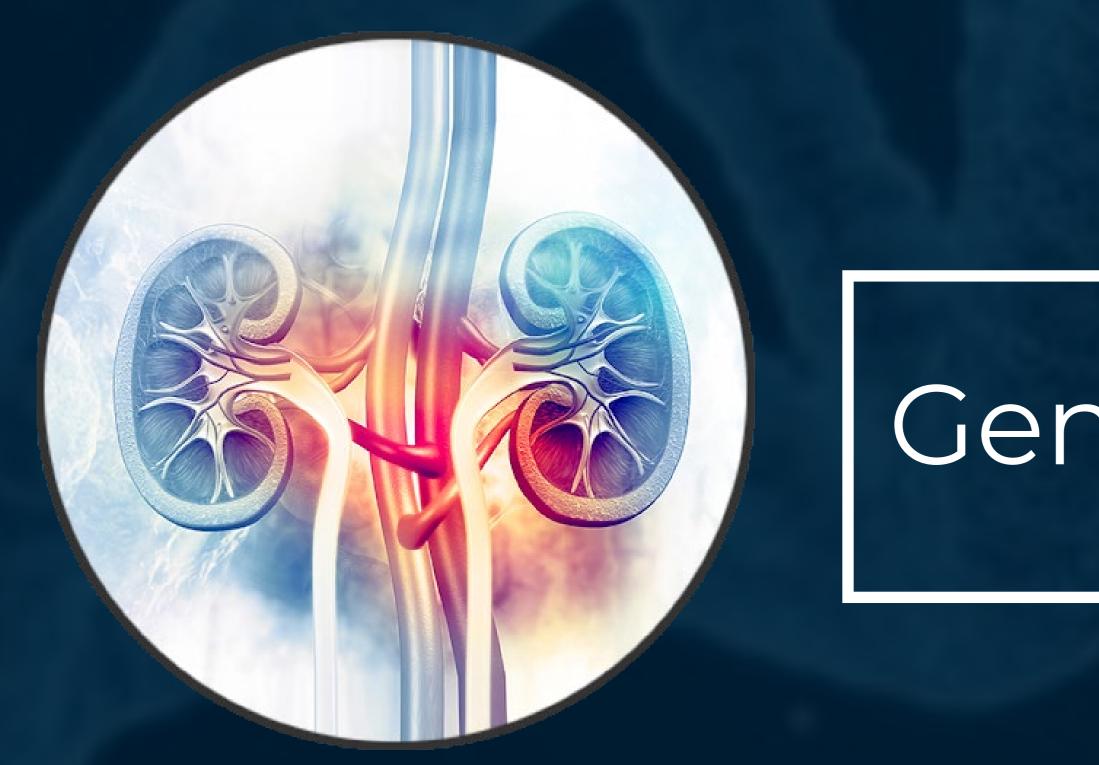
²Lead optimization includes refinement of GeneTether construct and delivery formulation, and demonstrating efficacy and tolerability in animal models.



We are developing gene correction and complementation therapies for the treatment of patients with rare, monogenic kidney diseases that lead to chronic kidney disease (CKD).

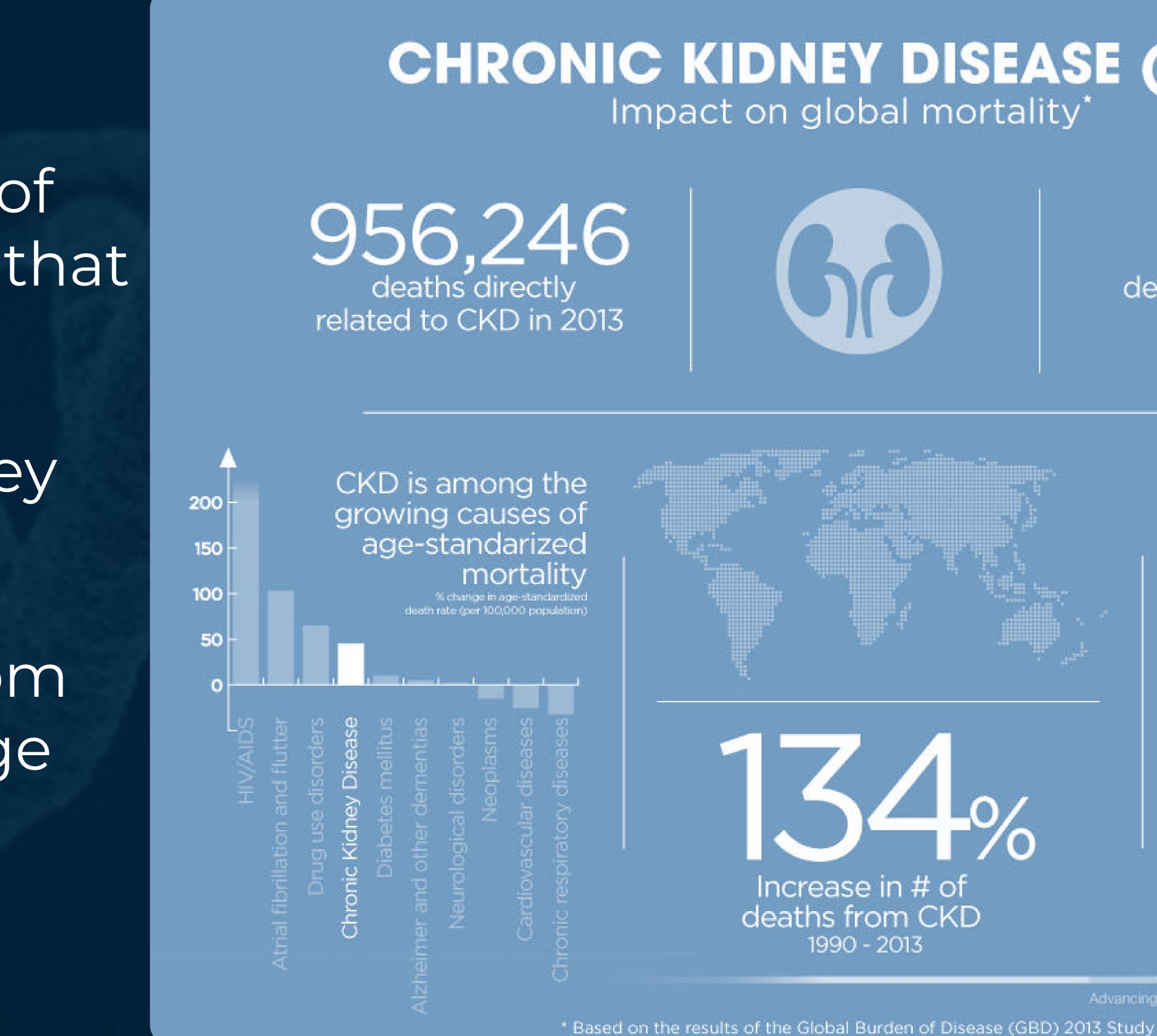
625 monogenic disorders associated with kidney and urological traits have been identified.

CKD treatment strategies only address symptom management and prolonging time to end-stage kidney disease (ESKD) and kidney transplant.



Genetic Kidney Diseases & CKD

Gene editing offers the possibility of a permanent curative therapy.



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The Lancet

Kidney CHRONIC KIDNEY DISEASE (CKD) Campaign Impact on global mortality* $1_{in}57$ deaths worldwide due to CKD of all cardiovascular deaths are attributable to CKD .207,453 cardiovascular deaths were attributed to one of the principal CKD markers, Increase in # of low Glomeular Filtration Rate deaths from CKD 1990 - 2013

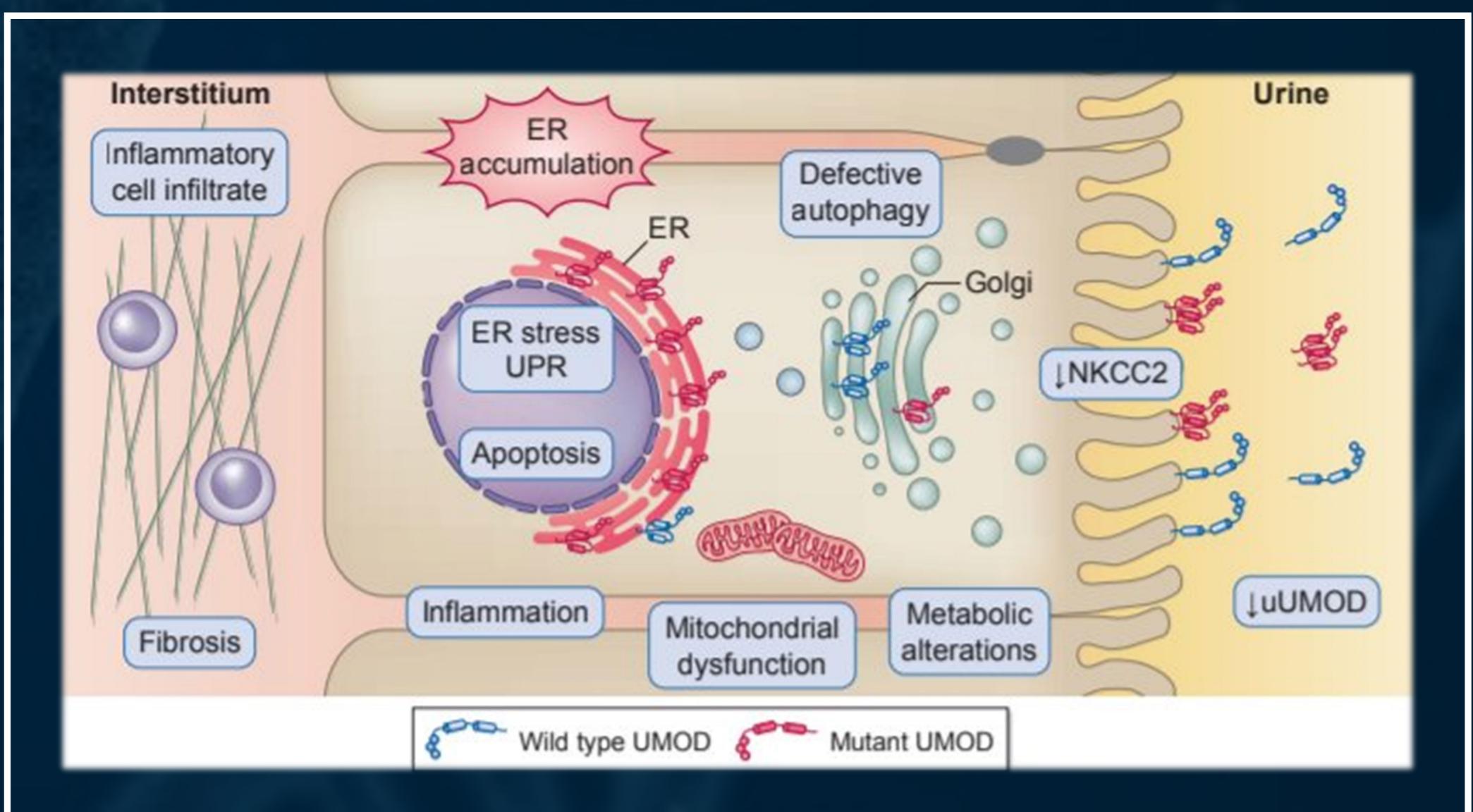
Advancing Nephrology Around the

Genetic Kidney Diseases Autosomal Dominant Tubulo-Interstitial Kidney Disease Autosomal dominant tubulo-interstitial kidney disease (ADTKD) is a group of rare genetic diseases that affect the tubules of the kidney. The most common form is ADTKD-UMOD.

CKD Patients

~3% = ADTKD-UMOD ~9,000 global Rare Disease

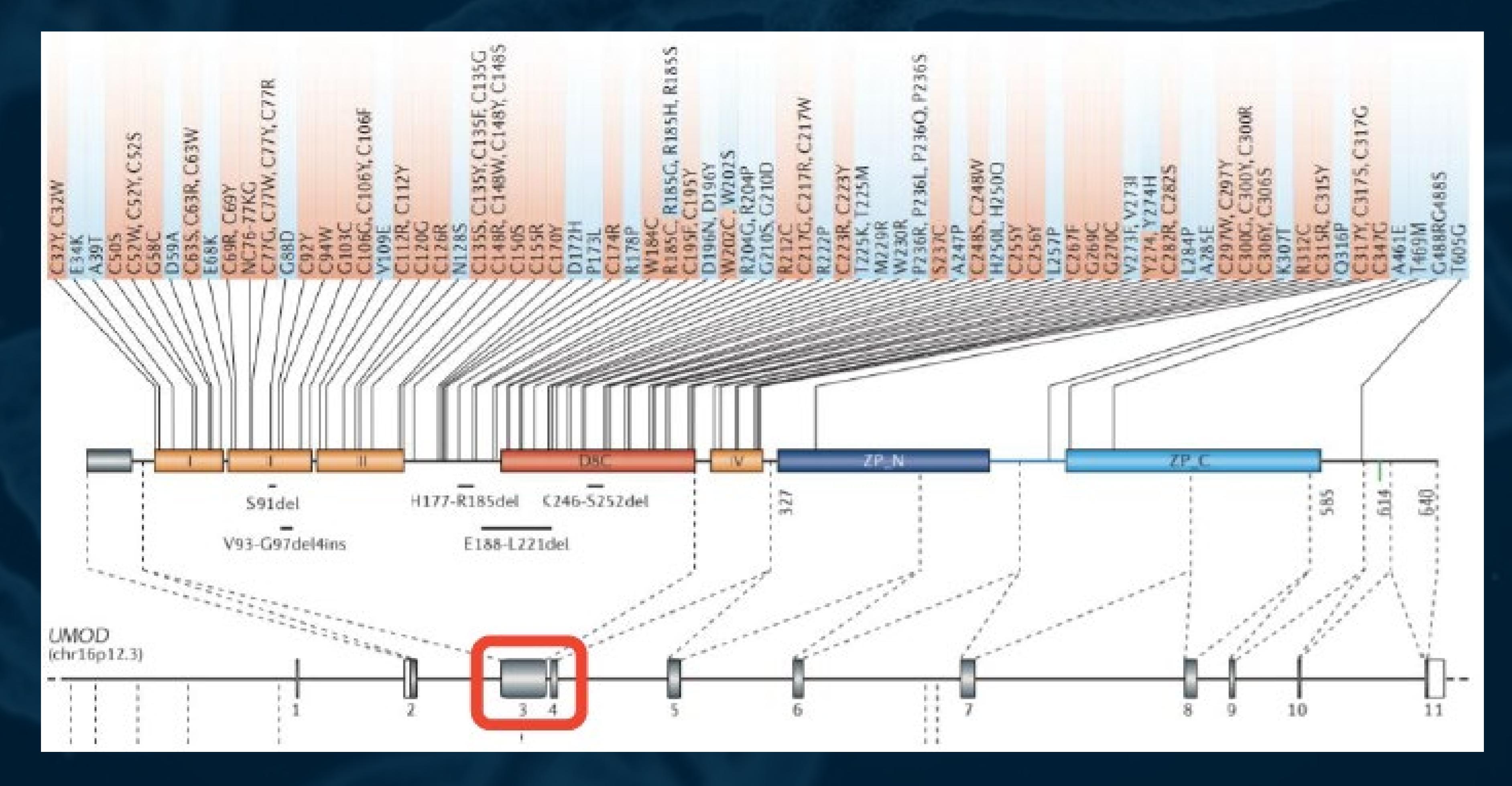
No approved therapy Symptom management



The UMOD gene encodes for production of uromodulin, a protein which protects against kidney stones and urinary tract infection. Misfolded uromodulin accumulates in the endoplasmic reticulum of tubulin cells, which leads to CKD, followed by ESKD.

GeneTether for ADTKD-UMOD

Over 95% of known UMOD mutations underlying ADTKD are reported in a small segment of the UMOD gene known as exon 3 and exon 4 (highlighted in red below). Our GeneTether technology may allow the development of a single treatment for ADTKD-UMOD resulting from mutations in this region by correcting a locus that fully encompasses exons 3 and 4.



Genetic Kidney Diseases Autosomal Dominant Polycystic Kidney Disease

Autosomal dominant polycystic kidney disease (ADPKD) is a rare genetic disorder characterized by the growth of numerous cysts in the kidneys.

> Affects ~140,000* people in the U.S. and ~12.5 million people worldwide

No cure approved **Treatment limited** to symptom management

*Qualifies as a rare disease

Responsible for up to 10% of all cases of ESKD



- 15% of cases: mutations in PKD2

 - Renal
 - Cyst formation



+

Increased kidney volume



Kidney stones



Urinary tract infection



+

Abdominal pain

End-stage kidney disease GeneTether Therapeutics **Corporate Presentation**

What are the causes of ADPKD?



Manifestations of ADPKD include



Extrarenal

Cysts in the liver, pancreas, spleen, and central nervous system



Cerebral aneurysms



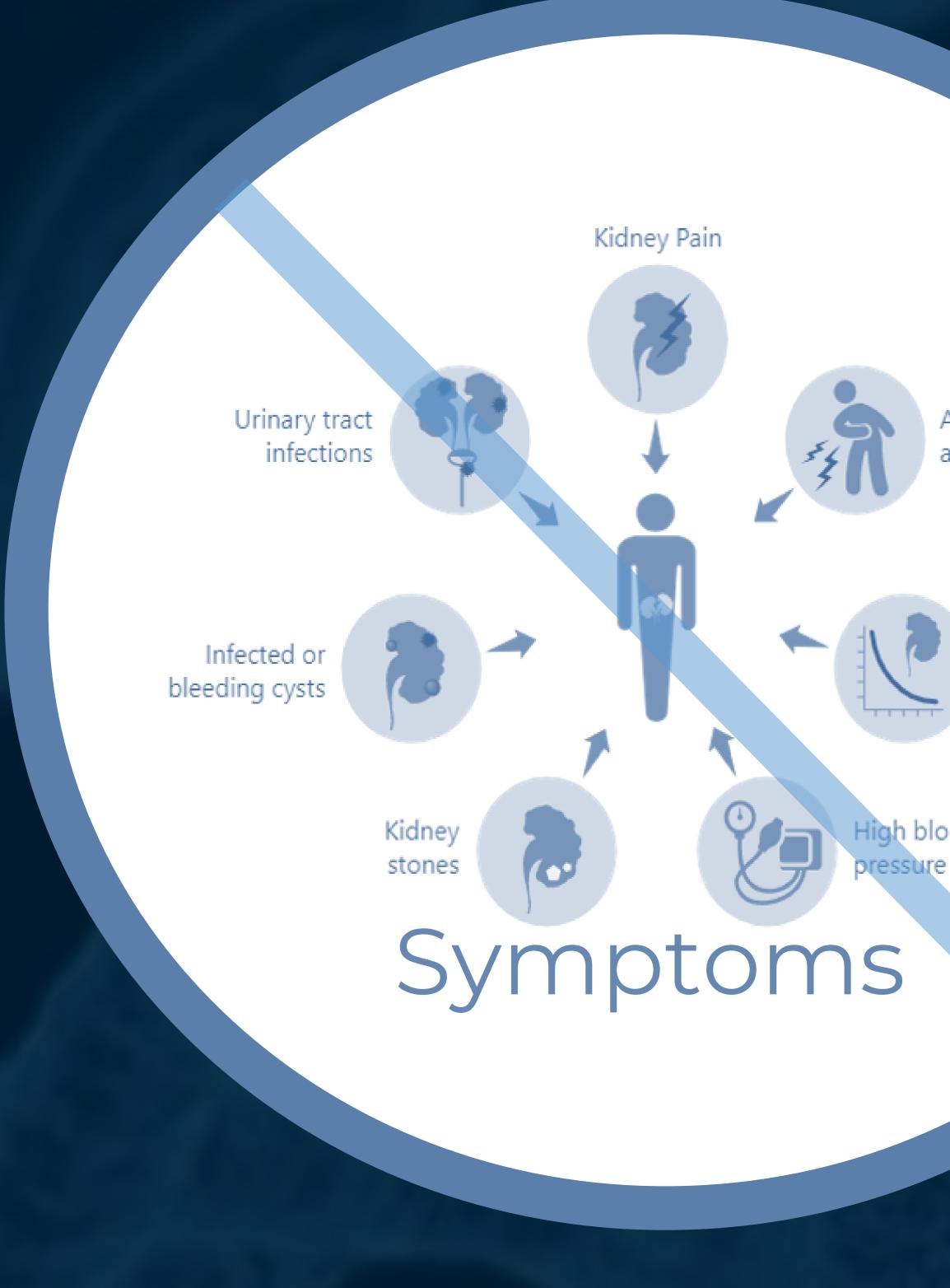
F Polycystic liver disease



Diverticular disease and mitral valve prolapse

GeneTether for ADPKD-PKD1

Because mutations of the PKD1 gene account for ~85% of ADPKD cases, we intend to investigate the use of our GeneTether technology to correct or complement PKD1 gene function. We believe this may enable the restoration of functional polycystin 1 protein with an objective of developing a potentially permanent cure.



Abdominal discomfort and bloating

> Reduced kidney unction



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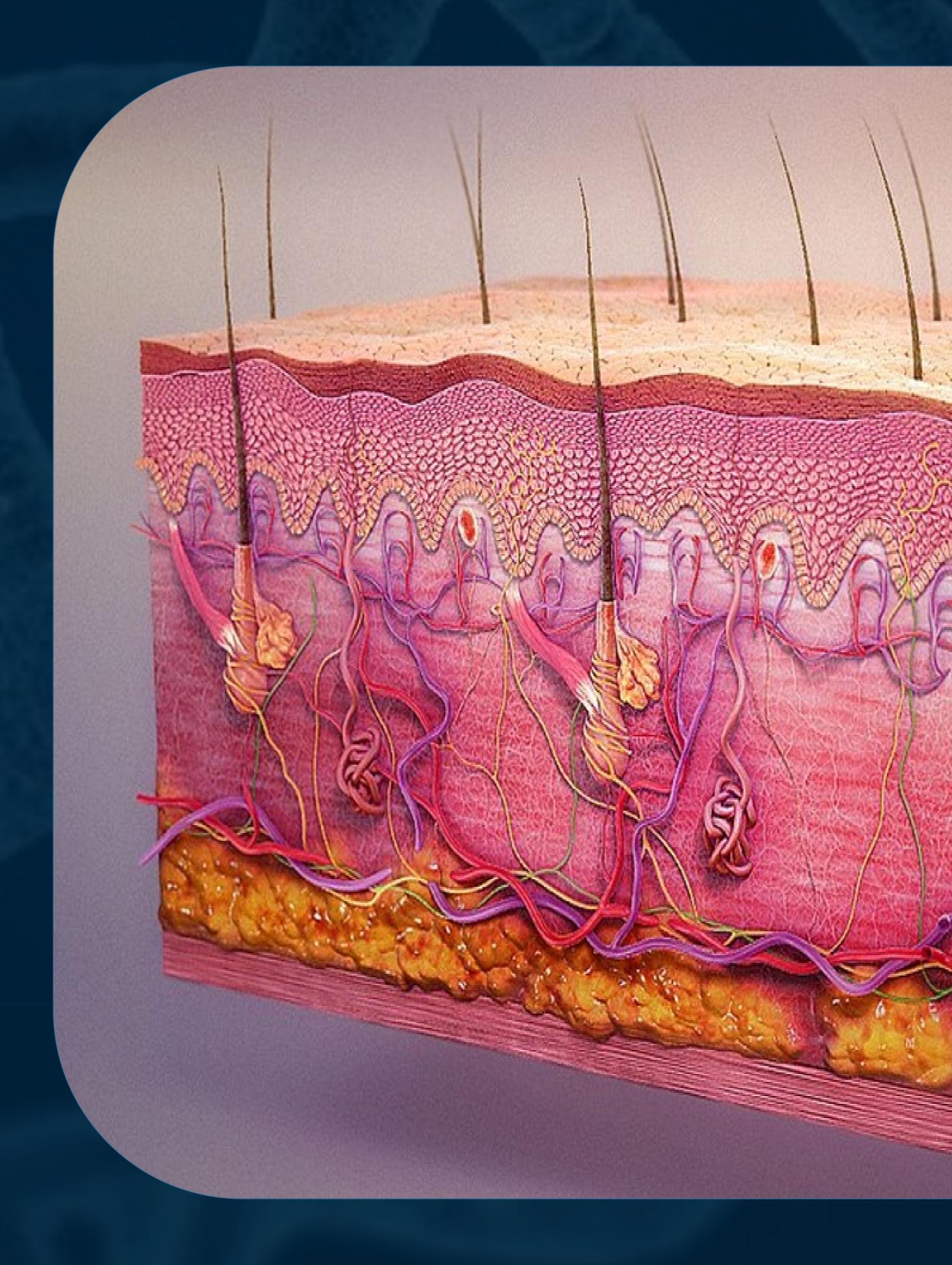
Transplant

Genetic skin diseases represent a broad class of rare diseases with confluent and overlapping phenotypes. We have identified genetic skin diseases as one of our initial discovery targets due to:

- > the significant unmet need for curative treatments, and
- our therapies.

Genetic Skin Diseases

> the well-characterized underlying genetic mutations of certain diseases, > the potential for topical delivery of



GeneTether Therapeutics **Corporate Presentation**

(Subcutaneous Tissue)

Genetic Skin Diseases Recessive Dystrophic Epidermolysis Bullosa

Recessive Dystrophic Epidermolysis Bullosa (RDEB) is a rare, often fatal, genetic skin condition caused by a mutation in the COL7A1 gene.

Lack of collagen protein makes the skin incredibly fragile, leading to blistering or skin loss at the slightest friction.

There is currently no cure for RDEB. The standard-ofcare includes wound care, pain management, prevention of skin trauma, and early detection and treatment of squamous cell carcinoma.



No cure



WHAT IS EB?

Epidermolysis bullosa (EB) is a painful genetic skin condition that causes the skin to tear and blister at the slightest touch.





Affects an estimated 500,000 people worldwide

GeneTether Therapeutics Corporate Presentation



Skin as fragile as a butterfly wing



Layers of skin tear and bliste



internal organs

GeneTether for RDEB

Skin grafts engineered to include normal copies of the COL7A1 gene complimentary DNA has shown promise for improved wound healing.

As with other non-integrating gene therapies, the effects were not long lasting, as cell division reduces expression of the COL7A1 gene over time.

We believe that RDEB is a candidate for *in vivo* gene correction, as a single donor DNA template inserted via HDR may permanently restore COL7A7 functionality across multiple mutations.







Cell Delivery

Cellular delivery of gene editing payloads is an important and difficult component of a viable therapeutic.

We are currently evaluating multiple viral and non-viral delivery technologies that have enhanced kidney and skin tropism, high levels of functional transduction, and improved manufacturability.

Technologies we are evaluating include next generations of lipid nanoparticles, helper-dependent adenoviruses, and adeno-associated viruses.

Intellectual Property Patents and Pending Applications¹

Wholly-owned patent portfolio; no 3rd party financial obligations

We will seek to continue to innovate and strategically protect our innovations in the following three main areas:

- editing systems;
- Uses in monogenic kidney disorders,

Cell delivery into tissues and cells of interest.

¹There is no guarantee that new patents will issue or effectively protect the commercial prospects of GeneTether's assets if they do. GeneTether has not received any written legal opinion in relation to patentability of the subject matter disclosed and claimed in its patent applications. ²In February 2022, USPTO issued a Notice of Allowance for a patent entitled

"Modified Nucleic Acid Editing Systems for Tethering Donor DNA" related to GeneTether's platform technology. Upon issuance, it is expected that the standard 20 year patent term will extend to March 2039

Composition of matter claims combining components of our GeneTether platform with other components of various gene

monogenic skin disorders, and other nonkidney and non-skin disease targets; and

Granted Australia

Notice of Allowance from USPTO (February 2022)²

USA









Japan



Canada







GeneTether Therapeutics **Corporate Presentation**

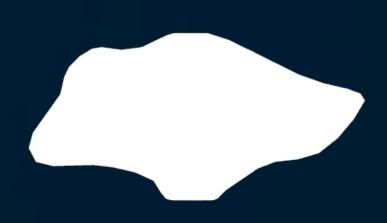
Israel







Singapore



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Development Timeline/Catalysts

Objective

Continued validation of GeneTether platform technology and expansion of portfolio

Identification of lead development program(s)

¹Dates generally represent anticipated completion of activities and are subject to factors that may be beyond our control, including the availability of thirdparty collaborators and contractors. The activities included are summaries only and are subject to change at management's discretion. Many of the activities listed above will be ongoing for the duration of our development programs.

Activities

	Identify and engage qualified con
	Non-cGMP manufacturing of key GeneTether-based gene editing s
ofIP	Editing in: - large animal cell lines - zebrafish - human cell lines
	Identify and engage key opinion potential disease targets
	Initiate and/or complete <i>in vitro</i> c genetic disease targets as descrik under:
	- "Our GeneTether Platform for F Diseases"
	- "Our GeneTether Platform for F
	- Complete assessment of variou model

ntract research organizations

/ components of our system

leaders in the areas of our

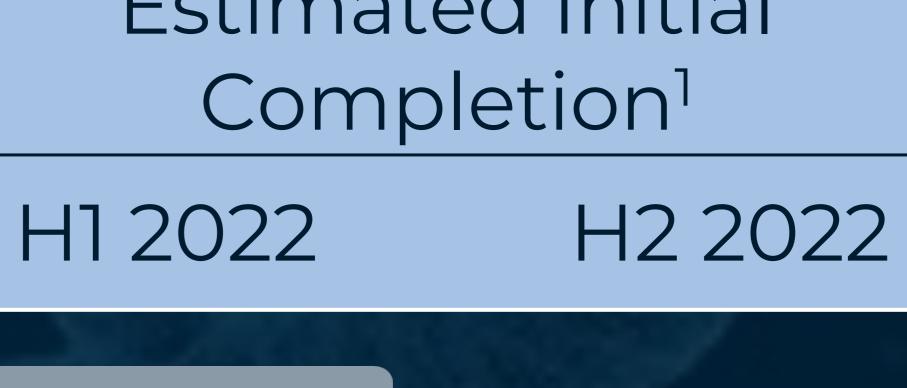
cell line editing in potential rare, bed in the final prospectus

Rare, Monogenic Kidney

Rare, Monogenic Skin Diseases"

us delivery platforms in kidney





Peer Group Comparables

In accordance with Section 13.7(4) of National Instrument 41-01 – General Prospectus Requirements, all the information relating to GeneTether's comparables and any disclosure relating to the comparables, which is contained in the presentation to be provided to potential investors, has been removed from this template version for purposes of its filing on the System for Electronic Document Analysis and Retrieval (SEDAR).

GeneTether Therapeutics Corporate Presentation Capital Structure | Sources & Uses of Funds

Pre-Offering Shares Outstanding¹ Common Stock **Options to Purchase Com** Pre-Offering FD Shares Owned by Insiders

Shares Offered^{2, 3} Post-Offering FD Shares Post-Offering Market Cap

¹On a post-Reorganization basis following GeneTether's reorganization event ²Assuming no exercise of Agent's overallotment option. ³7.5M Units are being offered, assuming minimum Offering of C\$4,500,000. Each Unit is comprised of one common share purchase warrant. Each warrant is exercisable to purchase one common share at a price of C\$0.72. Warrant shares are not included in Post-Offering "Fully Diluted (FD) Shares" above. ⁴Unaudited

⁵Unallocated working capital is to provide additional contingency for additional research & development, overhead and general and administrative expense overrun

	41.7M	
mmon Stock	9.8M	
	51.5M	
	~79%	
	7.5M	
	59.0M	
p at C\$0.60	C\$35.4M	

- Source of Funds
- Working Capital as at February 28, 2022
- Estimated Net Proceeds from the Offering²
- Total Available Funds⁴
- Use of Available Funds
- GeneTether technology R&D
- General and Administrative Expenses
- Unallocated Working Capital⁵
- Total Available Funds⁴

C\$3,545K C\$4,910K C\$3,640K C\$5,005K Minimum Maximum Offering Offering C\$1,842K C\$2,842K C\$1,350K C\$1,350K C\$448K C\$813K			
C\$95K C\$95K C\$3,545K C\$4,910K C\$3,640K C\$5,005K Minimum Maximum Offering Offering C\$1,842K C\$2,842K C\$1,350K C\$1,350K C\$448K C\$813K			
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C\$3,640K C\$5,005K Minimum Maximum Offering Offering C\$1,842K C\$2,842K C\$1,350K C\$1,350K C\$448K C\$813K	,	C\$95K	C\$95K
Minimum Offering C\$1,842KMaximum Offering C\$2,842KC\$1,350KC\$1,350KC\$448KC\$813K		C\$3,545K	C\$4,910K
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C\$448K C\$813K		C\$1,842K	C\$2,842K
		C\$1,350K	C\$1,350K
C\$3,640K C\$5,005K		C\$448K	C\$813K
		C\$3,640K	C\$5,005K

Terms of the Offering

Company	GeneT
Offering	Prospe for ma
Issue Price	C\$0.6C
Units	Eachl
Warrant	Each w month
Agent's Option	The Ag
Use of Proceeds	To con for ger
Eligibility	The Ur
Offering Jurisdictions	British
Closing Date	Onora
Lead Agent	Resear

- ether Therapeutics Inc.
- per Unit
- Unit is comprised of one Common Share and one common share purchase warrant
- varrant is exercisable to purchase one Common Share at a price of C\$0.72 for a period of 36 ns following the closing of the Offering
- nduct certain R&D activities related to the GeneTether platform technology and neral and administrative purposes
- nits will be eligible for registered plans²
- Columbia, Alberta, and Ontario
- about March 29, 2022
- rch Capital Corporation
- ¹Gross proceeds before deducting Agent's commission and expenses of the Offering ²See the final prospectus for full disclosure regarding holding Offering securities within a registered plan

ectus offering of 7.5M Units to raise a minimum of C\$4.5M¹ and up to a maximum of 10M Units ximum of up to C\$6.0M¹ on a commercially reasonable efforts basis

gent shall have the option to increase the size of the offering by up to 11.5M Units (C\$6.9M¹)

Investment Highlights

Experienced Team

Disruptive Platform Technology

Rare Genetic Diseases

IP Portfolio

Focused on harnessing next generation technology to significantly increase efficiency of gene editing and potentially cure serious and life threatening genetic diseases Extensive public life science company experience Global capital markets experience and extensive investor network

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