

Redefining Kidney Disease Treatment with A1M Therapies

Non-confidential summary

June 2025



PIONEERING TRANSFORMATIVE MEDICINES FOR KIDNEY DISEASE






















RMC-035 for kidney protection in open-heart surgery

- > Clinical proof-of-concept established in Phase 2a AKITA study with 177 patients
 - > 59% reduction vs placebo (MAKE, regulatory endpoint)
- > Phase 2b *POINTER* study ongoing – > enrolment completed, results expected in Q4 2025
- > **Granted FDA Fast Track Designation**; eligible for Breakthrough Therapy Designation
- > **First-to-market potential**; >USD 1 billion market – no approved therapies

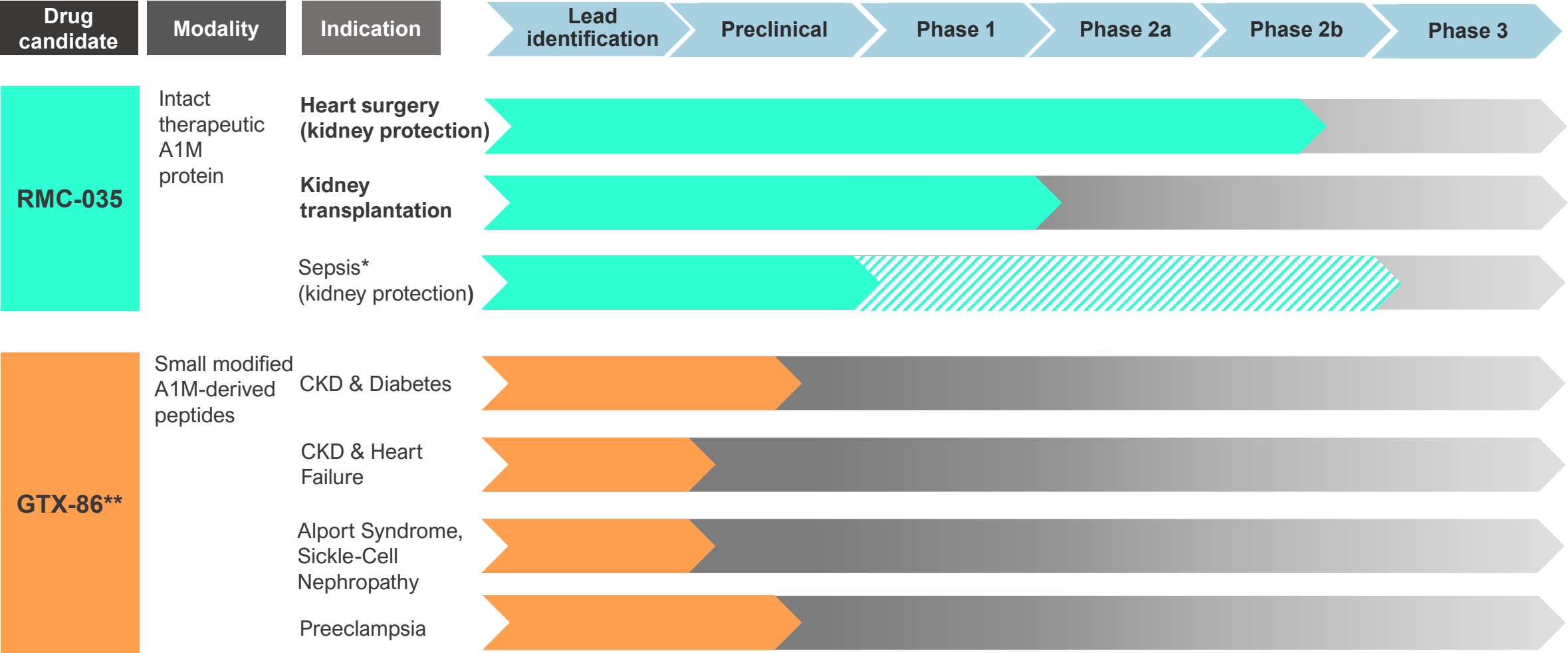
Additional opportunities with RMC-035 & GTX peptides

- > **Phase 3 ready sepsis program** and **Phase 2a/b ready kidney transplantation program** for >USD 5.6 billion market
- > **Unique positioning of preclinical GTX peptides** in chronic kidney disease for >USD 8 billion market
- > Listed in Stockholm with top shareholders including Industrifonden and Swedbank Robur [Nasdaq FN Growth Market: GUARD]

EXPERIENCED MANAGEMENT TEAM WITH STRONG & PROVEN TRACK RECORD IN DRUG DEVELOPMENT

NAME / POSITION	EXPERIENCE		NAME / POSITION	EXPERIENCE	
 TOBIAS L. AGERVALD MD, PhD, CEO	+10 years in industry		 KARIN BOTHA MSc, CFO	+20 years in industry	   GlaxoSmithKline
 MICHAEL REUSCH MD, CMO	+30 years in industry	 	 PETER GILMOUR MSc, PhD, CSO/Head of Preclinical	+20 years in industry	 
 TORBJÖRN LARSSON BSc, Head of CMC	+30 years in industry	  	 SARA THURESSON MSc, Head of Clinical Operations	+15 years in industry	   

DIFFERENTIATED PIPELINE BASED ON A1M MECHANISM



* Opportunity to initiate pivotal Phase 3 study in sepsis following results in ongoing Phase 2b study (POINTER) in open-heart surgery.

** Multiple GTX peptides fulfill criteria for candidate drug nomination. GTX-86 at nomination stage.

A1M, alpha-1-microglobulin.

CHRONIC KIDNEY DISEASE & END-STAGE RENAL DISEASE – A GLOBAL HEALTH CONCERN

Acute Kidney Injury (AKI):

- Multiple causes, often resulting from in-hospital complications like severe infections & sepsis and major surgeries (e.g., open-heart surgery, kidney transplantation)
- 50% or more of high-risk open-heart surgery patients develop AKI; addressable patient population ~100,000-120,000 cases per year in the US alone (~30,000 patients with pre-operative CKD)

Progression to Chronic Kidney Disease (CKD):

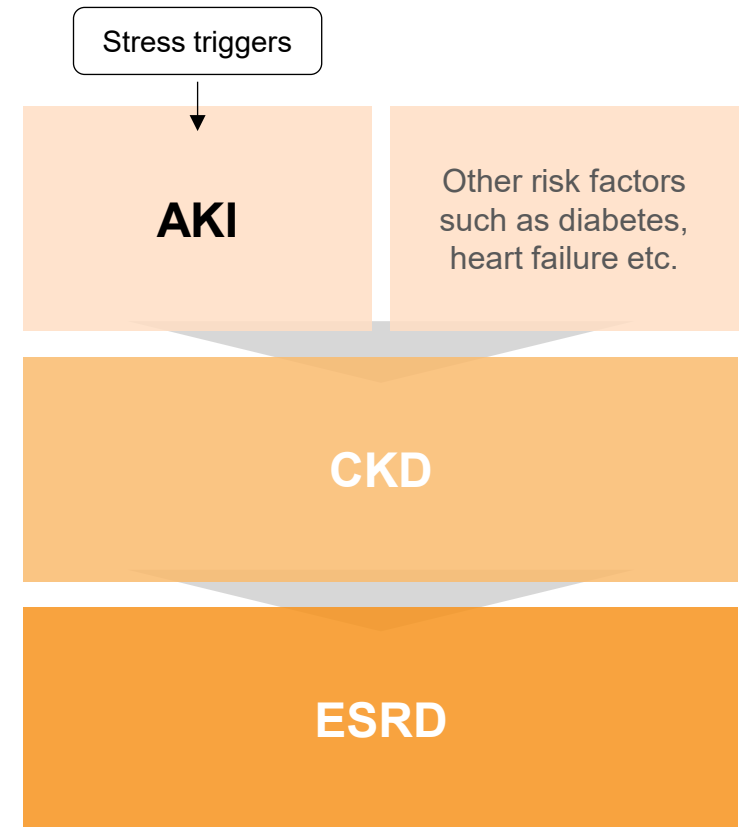
- AKI raises the risk of CKD; 15-20% progress to advanced CKD within 24 months
- CKD leads to severe complications, e.g., cardiovascular disease and kidney failure
- Years of life lost (YLL) from CKD are expected to surpass diabetes by 2040

AKI in patients with pre-existing CKD:

- CKD is a strong risk factor for AKI
- AKI in CKD accelerates progression to ESRD – high unmet need

CKD to End-Stage Renal Disease (ESRD):

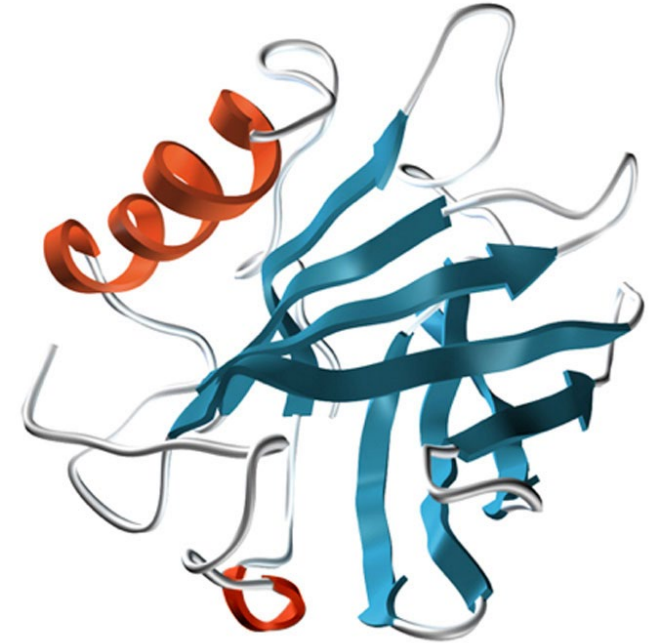
- 10-15% of CKD patients advance to ESRD; requires dialysis or kidney transplant
- High mortality rate (15-20%), worse than many cancers
- Represents 7% of Medicare costs but affects 1% of the population



THERAPEUTIC A1M DELIVERY – A NOVEL PARADIGM WITH CLINICALLY VALIDATED MECHANISM

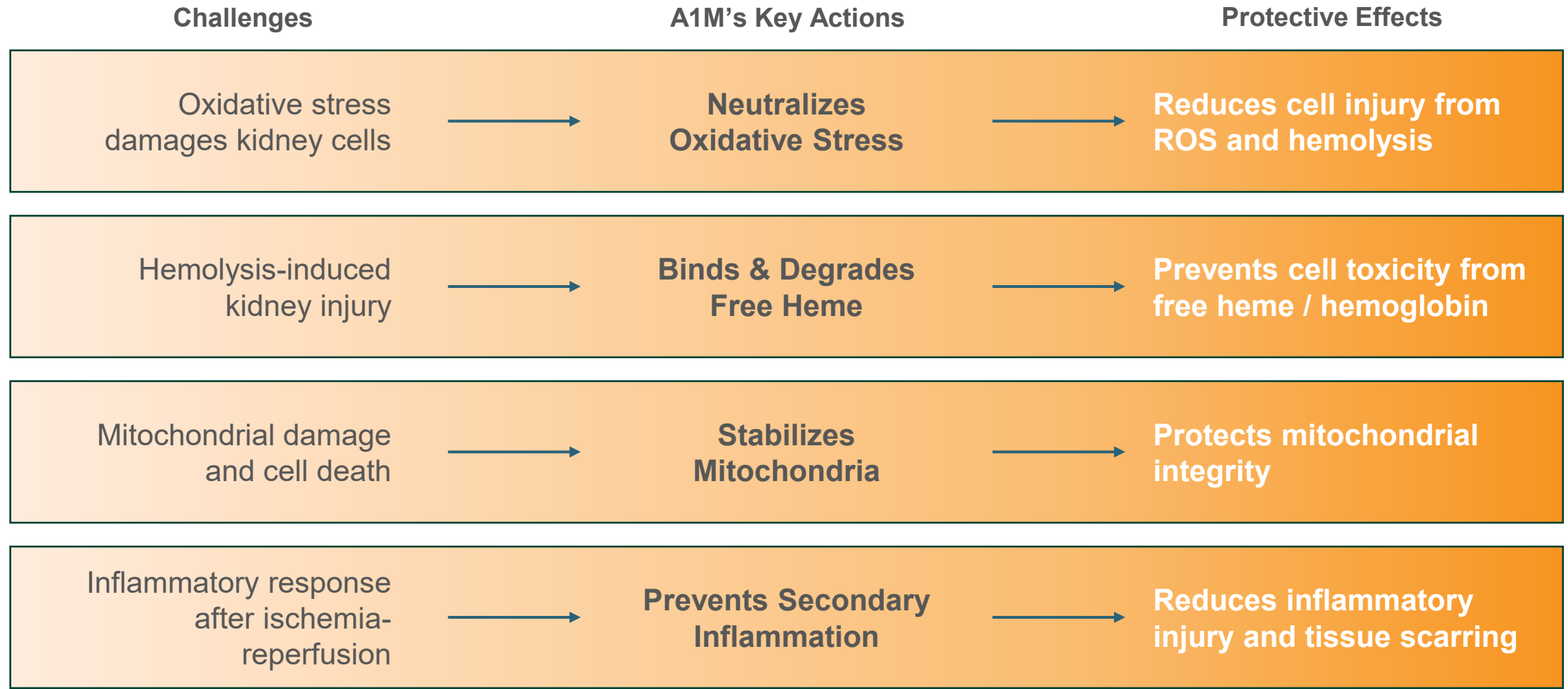
Harnessing the natural properties of A1M

- Endogenous ~22 kDa circulating glycoprotein
 - Liver main source of expression
- Ubiquitous distribution & cellular uptake
- Renal clearance
- *Evolutionary conserved house-keeping mechanism:*
 - **Potent dual-action inhibitor of oxidative stress**
(reductase, radical scavenging)
 - **Heme binding**
 - **Mitochondrial protection**



3-D structure of A1M protein

A1M PROTECTS KIDNEY FUNCTION BY TARGETING CRITICAL DISEASE PATHWAYS





RMC-035 – recombinant human intact A1M

Kidney protection in open-heart surgery

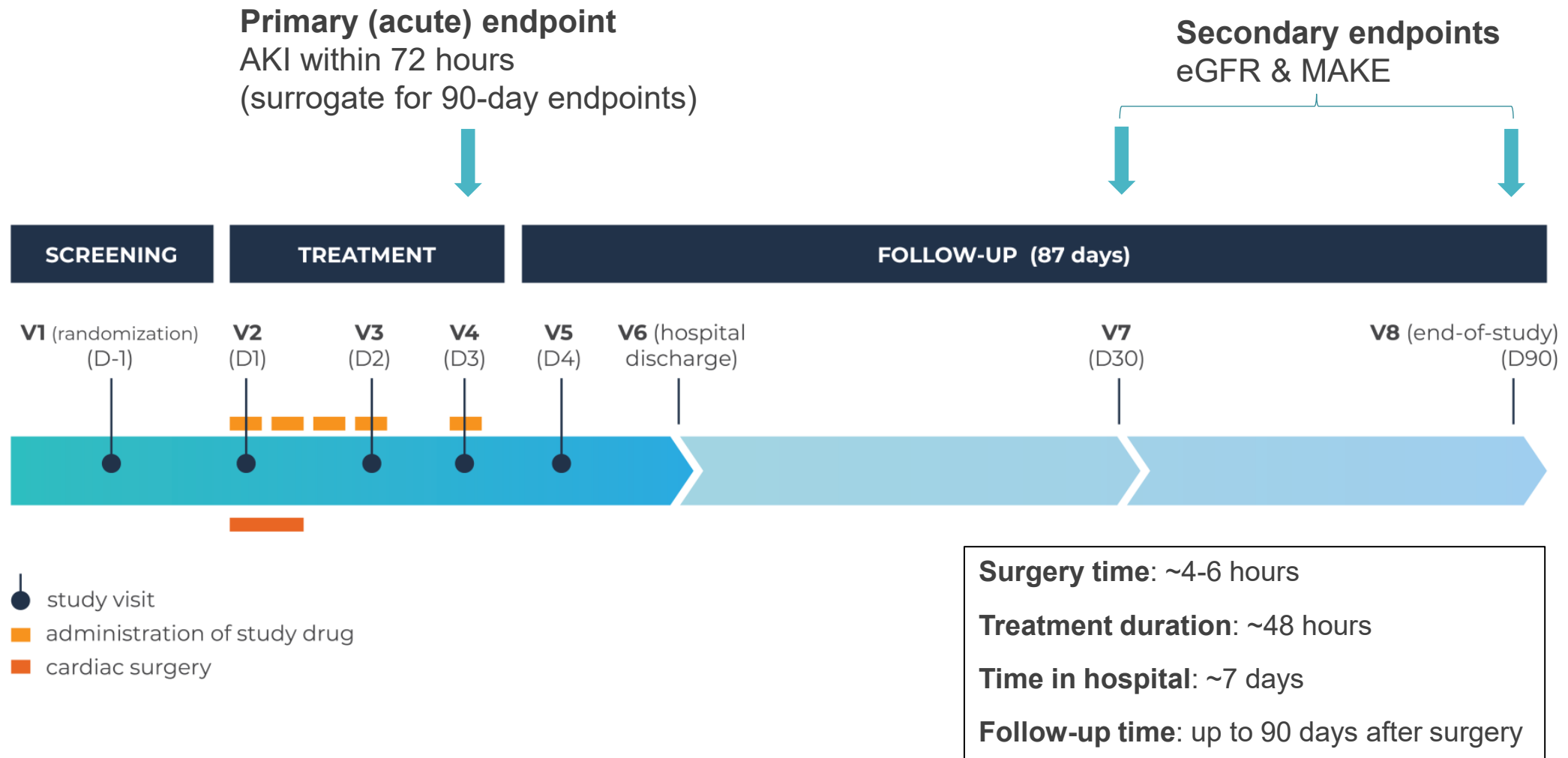
COMPLETED PHASE 2a AKITA STUDY

– OUTLINE & OBJECTIVES



- Recruitment in the **U.S., Canada & Europe**
- Double-blind, placebo-controlled study in patients undergoing **open-heart surgery at increased risk for kidney injury**
- **177 patients** randomized and dosed
- Main objective: **proving efficacy & safety with the maximum possible dose**
 - Primary endpoint: acute SCr change meeting AKI criteria
(surrogate for long-term renal outcomes, allowing for lower sample size. Not accepted regulatory endpoint)
 - Secondary endpoints: eGFR change and Major Adverse Kidney Events (MAKE) at Day 90
(clinically meaningful outcomes, **MAKE is the accepted regulatory endpoint**)
- Start dose 1.3 mg/kg
 - 0.65 mg/kg to subjects with low pre-operative renal function due to risk for overexposure associated with transient SCr increase (tubular overload)

COMPLETED PHASE 2a AKITA STUDY – FLOWCHART





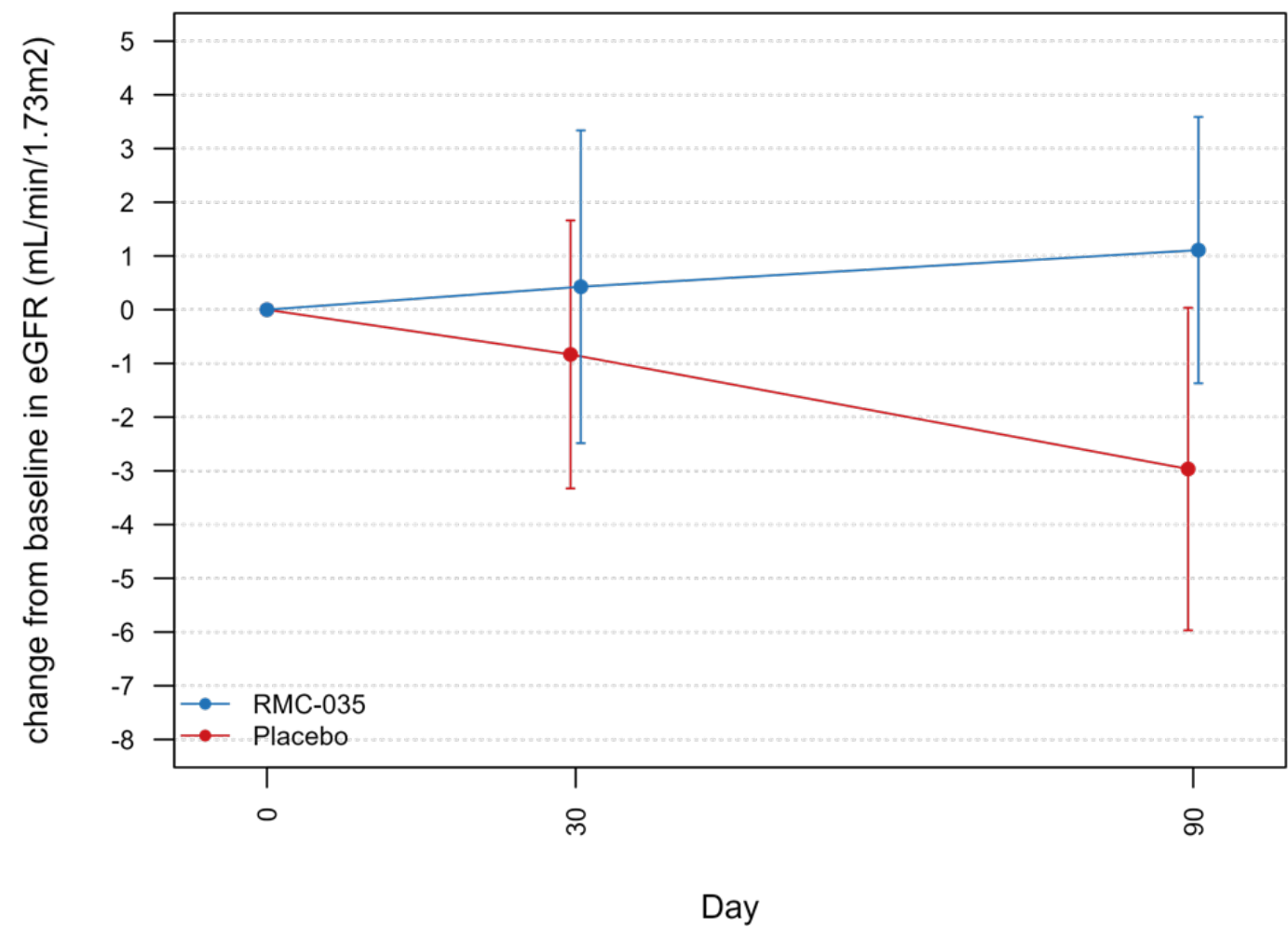
PHASE 2a RESULTS SUPPORT ROBUST EFFICACY ON HARD KIDNEY ENDPOINTS

Efficacy stronger than required for regulatory approval based on renal function (eGFR) & Major Adverse Kidney Events (MAKE) on Day 90

- **Statistically significant & clinically relevant improvement of eGFR vs placebo**
 - 4.3 mL/min (full population)
 - 7.9 mL/min (pre-defined subgroup of patients with chronic kidney disease [CKD])
- **Reduced proportion of patients with MAKE90 (i.e., severe loss of kidney function)**
 - 59% risk reduction vs placebo for composite endpoint MAKE90 (death, dialysis or $\geq 25\%$ eGFR loss)
 - FDA recommends **MAKE90 as primary endpoint in Phase 3 – 20% risk reduction sufficient for approval**

Results support progression to Phase 2b

RMC-035 PREVENTS LOSS OF RENAL FUNCTION AFTER OPEN-HEART SURGERY



MMRM model
4.3 mL/min
p=0.06*

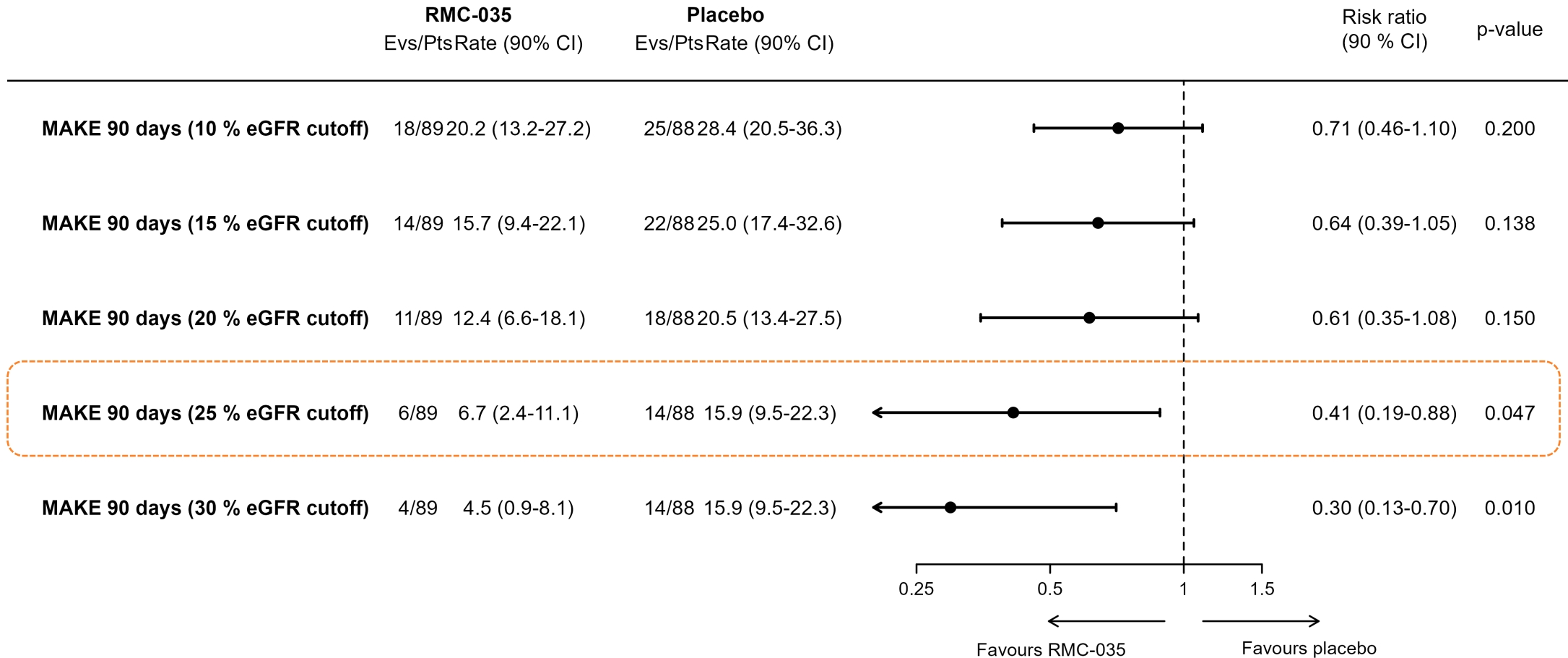
Pre-defined alpha level was 0.1.
P-values < 0.1 are statistically significant.

Source: Study ROS-05 Clinical Study Report. MAKE, major adverse kidney events; eGFR, estimated glomerular filtration rate

RMC-035 CONSISTENTLY REDUCES MAKE90 USING VARIOUS THRESHOLDS OF eGFR LOSS



Secondary endpoint **MAKE90** (death, dialysis, $\geq 25\%$ eGFR loss) met





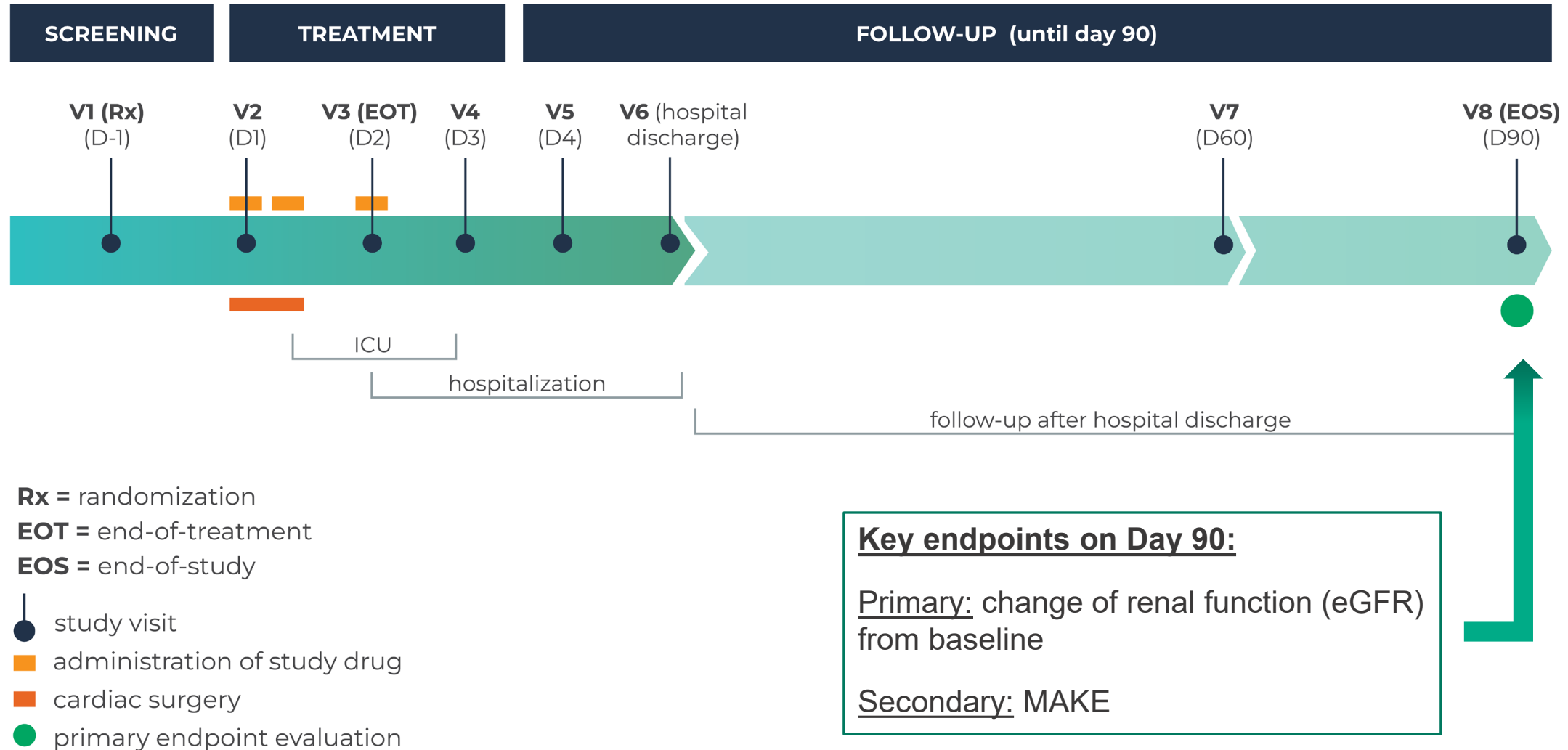
PHASE 2b *POINTER* STUDY – STATUS UPDATE

- **Final optimization step before Phase 3**
- **170 patients** randomized and dosed (Europe & North America)
- Two RMC-035 dose arms (30 & 60 mg) and Placebo (2:2:3 randomization)
 - Simplified dosing: 3 infusions within 24 hours, switch from weight-based to flat dose
- **Primary endpoint:** change in eGFR from before surgery to Day 90

Important study milestones

- ✓ **Positive outcome of DSMC safety reviews (no safety signals identified)**
- ✓ **Enrolment completed in June 2025**, ahead of schedule (9 months enrollment time)
- *Top-line results anticipated in Q4 2025*

ONGOING PHASE 2b *POINTER* STUDY – FLOWCHART



CLEAR PATH TOWARDS MARKET APPROVAL FOR RMC-035

- **Fast Track Designation** granted by the US FDA
 - *Reducing risk for death, dialysis or irreversible loss of kidney function in patients undergoing open-chest cardiac surgery at high risk for acute kidney injury*
- Indication eligible for **Breakthrough Therapy Designation**
- **Single pivotal Phase 3 study** sufficient to support market approval
 - *Primary endpoint MAKE at Day 90 after surgery (~600 patients)*
 - *Potential for accelerated approval based on interim analysis of eGFR (~300 patients)*
 - *First-to-market potential*
- **Phase 3 Expansion Opportunities**
 - ***Sepsis** Phase 3 ready and **Kidney Transplant** Phase 2a/b ready, following successful Phase 2b POINTER results*



FIRST-TO-MARKET POTENTIAL WITH NO APPROVED THERAPIES

– COMPETITOR LANDSCAPE

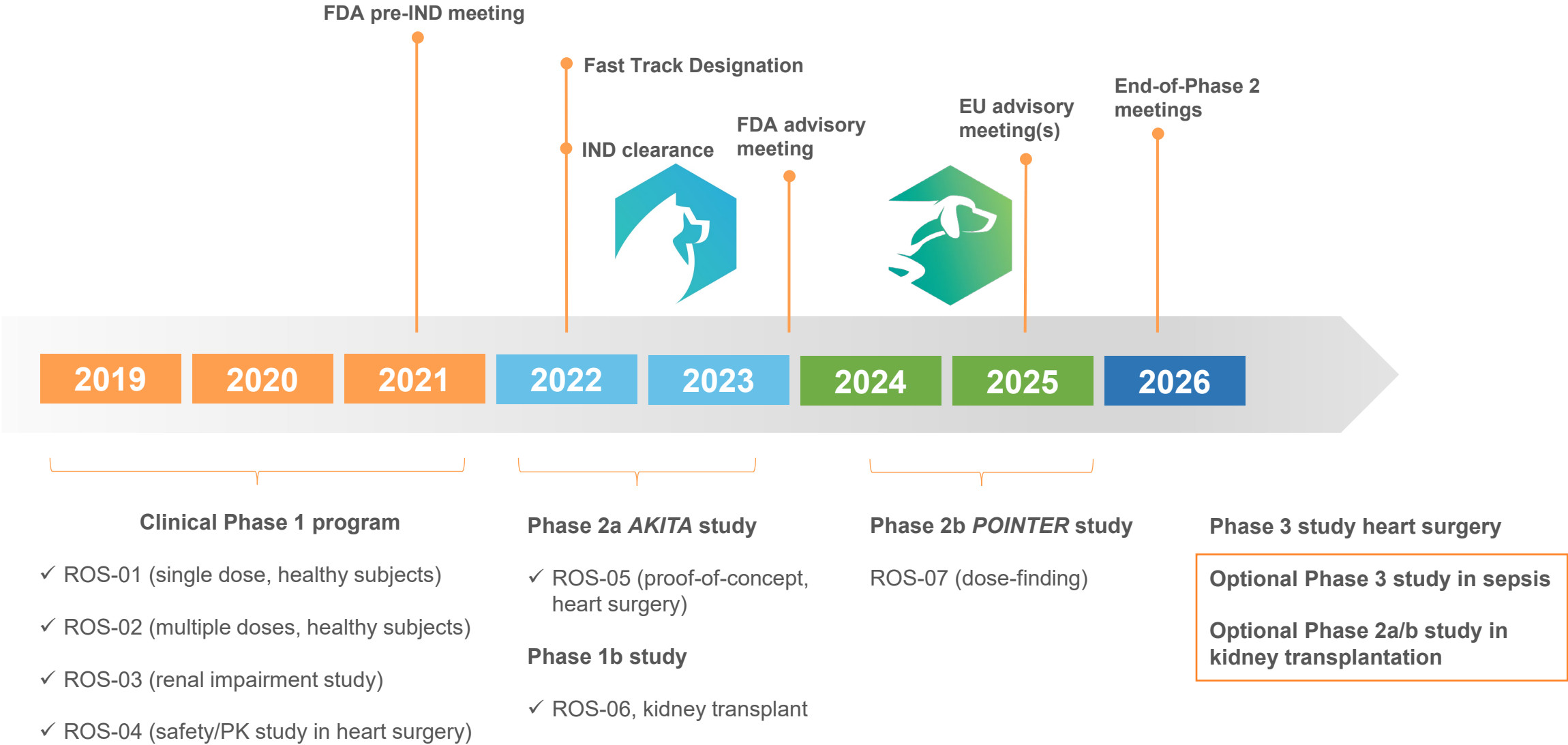
COMPANY (DRUG)	PHASE	MODE OF ACTION	POC DATA HEART SURGERY	COMMENT
Guard Therapeutics (RMC-035)	2b	A1M analog	Yes	Ongoing – expected results year-end 2025
Novartis (TIN-816)	2a	Human CD39 enzyme	-	Ongoing – expected results Q3 2025 N=120, acute primary endpoint <i>Study did not reach primary endpoint and stopped for futility</i>
AstraZeneca / Alexion (Ultomiris)	3	Complement 5 inhibitor	-	Ongoing – expected results Q1 2027 N=736, MAKE is primary endpoint No efficacy data available in open-heart surgery
Genentech (GDC-8264)	2	RIP-1 inhibitor	-	Ongoing – expected results Q4 2027 N=404, MAKE is primary endpoint No efficacy data available
AM Pharma (Ilofotase alfa)	2a	ALP analog	-	Ongoing – results expected Q4 2025 N=250, acute primary endpoint No efficacy data available in open-heart surgery
Renibus Therapeutics (RBT-1)	3	Iron sucrose + stannus protoporphyrin	-	Targets acute endpoints like length of hospital stay & hospital re-admission rate. Did not show efficacy on renal endpoints in Phase 2a study.


RECENT PHARMA DEALS IN NEPHROLOGY

– TOTAL DEAL VALUE OVER \$12BN 2023-25 YTD

TARGET	ACQUIRER	YEAR	DEAL VALUE	STAGE	LEAD ASSET	INDICATION
Regulus Tx	Novartis	2025	\$800m + \$900m milestones	Phase 1b	Farabursen	Autosomal Dominant Polycystic Kidney Disease
Alpine Immune Sciences	Vertex Pharma	2024	\$4.9bn	Phase 2	Povetacicept	IgAN
Human Immunology Biosciences	Biogen	2024	\$1.15bn + milestones	Phase 2	Feltzartamab	IgAN, Primary membranous nephropathy & antibody-mediated rejection
Jnana Tx	Otsuka	2024	\$800m	Preclinical	Panel of solute carrier inhibitors	Ion transporter kidney disease
Calliditas	Asahi Kasei	2024	\$1.1bn	Marketed	Tarpeyo (Budesonide)	IgAN
Chinook Tx	Novartis	2023	\$3.5bn	Phase 3	Atrasentan & Zigakibart	IgAN
CinCor Pharma	AstraZeneca	2022	\$1.8bn	Phase 2	Baxdrostat	Treatment-resistant hypertension, primary aldosteronism and CKD
Vifor Pharma	CSL	2021	\$12.3bn	-	Product portfolio in nephrology	-
Sanifit Tx	Vifor Pharma	2021	\$205m + milestones	Phase 3	SNF472	Treatment for calciphylaxis ESRD patients
Corvidia Tx	Novo Nordisk	2020	\$2.1bn	Phase 2	Zilitivekimab	Therapies within CKD segments

KEY MILESTONES & DELIVERY ACCORDING TO PLAN





GTX platform – small A1M-derived peptides

Treatment of chronic kidney disease

GTX PEPTIDES – 2nd GENERATION A1M DRUGS

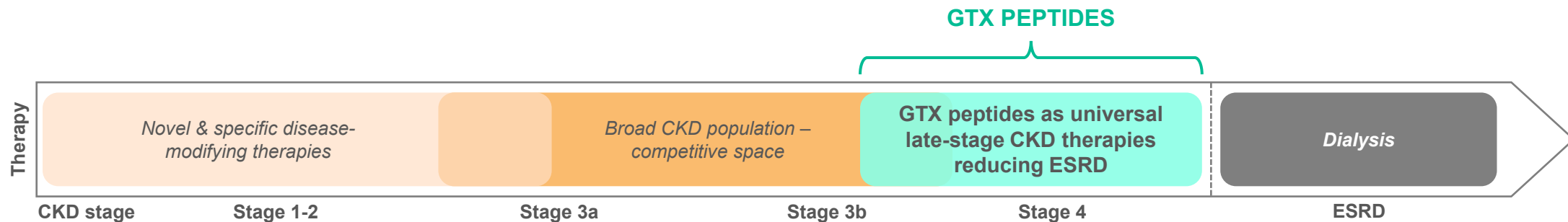
Delivered SC

Intended as chronic therapy with intermittent dosing (e.g., for CKD)

- Panel of novel **A1M-derived peptides with preserved functionality** vs native A1M
 - ~15-35 aa, chemical synthesis
- Enables **clinically validated A1M mechanism** in **non-acute settings**
- **Robust preclinical efficacy** across diverse acute and chronic kidney models
- **Strong IP**, composition of matter until 2044
- Broad clinical development opportunity with **unique positioning in CKD**
 - **High degree of optionality** – strategy yet to be refined
- Candidate drug nomination pending; transfer into slow-release formulation
- ~2 years to IND

GTX PEPTIDES – MASSIVE OPPORTUNITY IN LATE-STAGE CKD

- High potency and efficacy, comparable to RMC-035
- Broad impact across CKD etiologies, including orphan diseases
 - *Demonstrated robust efficacy in a wide range of preclinical disease models*
- Unique opportunity in late-stage CKD patients:
 - *Highest risk for progression to ESRD*
 - *Often excluded from clinical trials*
 - *Current CKD therapies ineffective or contraindicated*



PIONEERING TRANSFORMATIVE MEDICINES FOR KIDNEY DISEASE

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Appendix

ADDITIONAL OPPORTUNITY FOR RMC-035

– SEPSIS

- Sepsis is **leading cause of acute kidney injury (AKI)**. Multifactorial etiology, kidney cell stress due to reduced perfusion, oxidative stress, endotoxins and inflammation
- **RMC-035 efficacious in preclinical sepsis models**
- In the US, **~1.7 million** patients develop sepsis each year; **~800,000 patients** with sepsis develop AKI; and **~250,000 patients** develop CKD
- Dosing regimen: once daily up to 5 days; First dose given at sepsis diagnosis (ICU admission)
- Clear regulatory path to market approval – **Major Adverse Kidney Events (MAKE) at 90 days**
- **Single confirmatory Phase 3 study** sufficient for approval
 - *Sample size ~400-600 patients depending on eligibility criteria*
 - *Recruitment time ~2 years*
- **Pivotal Phase 3 study in sepsis enabled by Phase 2b POINTER study (heart surgery)**
 - *Interim analysis with sample size re-estimation to be built in in the absence of preceding efficacy study*
 - *Should be preceded by a Phase 1b study of approximately 15-20 patients to evaluate exposure & safety*

ADDITIONAL OPPORTUNITY FOR RMC-035

– KIDNEY TRANSPLANTATION

- **Phase 1b study of RMC-035 completed**
- Acute kidney graft dysfunction & impaired long-term dysfunction in deceased donor transplantation due to **graft ischemia & ischemia-reperfusion injury**, and **inflammatory / fibrotic response**
- Approximately **20,000 deceased donor kidney transplantation** performed annually in US – eligible for **orphan drug designation**
- Treatment goal: **protect long-term graft function in recipient**, avoiding need for re-transplantation
- Dosing regimen: once daily up to 5 days; First dose given intra-operatively to graft recipient
- **Clear regulatory path** to market approval – **eGFR at 1 year** after transplantation
- **Single confirmatory Phase 3 study** sufficient for approval
 - *Sample size 300-600 patients depending on eligibility criteria*
 - *Recruitment time 2-3 years*
- New formulation considered, may enable higher price point than in open-heart surgery

GLOBAL MARKET OPPORTUNITY FOR RMC-035 SUPPORTS

BLOCKBUSTER POTENTIAL

- **Open-heart surgery: >USD 1 billion (global)**
 - *100,000 patients annually in the US (~30,000 with CKD). Total US population ~250,000 patients.*
 - *Price estimate: USD 5,000 – 10,000 per patient*
 - *Total US market potential: USD 0.5 – 1 billion*
- **Sepsis: >USD 5 billion (global)**
 - *~1.7 million adults in the US develop sepsis each year*
 - *~500,000 patients in scope for treatment*
 - *Total US market potential: USD 2.5 – 5 billion*
- **Kidney transplantation: >USD 0.6 billion (global)**
 - *~20,000 patients in the US undergoing deceased donor kidney transplantation each year*
 - *Total US market potential: USD 0.3 – 0.6 billion (estimated price USD 15,000 – 30,000 per patient)*

STRONG VALUE PROPOSITION – EFFICACY & COST-BENEFIT PROFILE

Robust kidney protection in AKITA study

- *~5 x greater eGFR effect than one year of standard-of-care CKD treatment*
- *3 x greater risk reduction of MAKE than required for FDA approval*

Strong evidence for eGFR in Cost Effectiveness Models

- *Value dossier supported by available health economic data in patients with CKD*

Short-term therapeutic benefit (AKI, dialysis, length of hospital stay & re-admission)

- *Acute benefits offer direct & indirect hospital savings*

Attractive cost-benefit profile

- *Anticipated formulary inclusion with marginal impact on hospital cost*
- *Fulfils NTAP criteria*

Value dossier based on HEOR & available Cost Effectiveness Models in AKI & CKD

Approach P&T Committees & Payers via FDA-approved AMCP dossier

Launch

Critical Pre- and Post-Launch activities

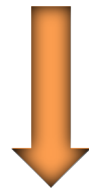
Cost Effectiveness Analysis: quality-adjusted life years (QALYs)

Budget Impact Model & Budget Analysis Tool

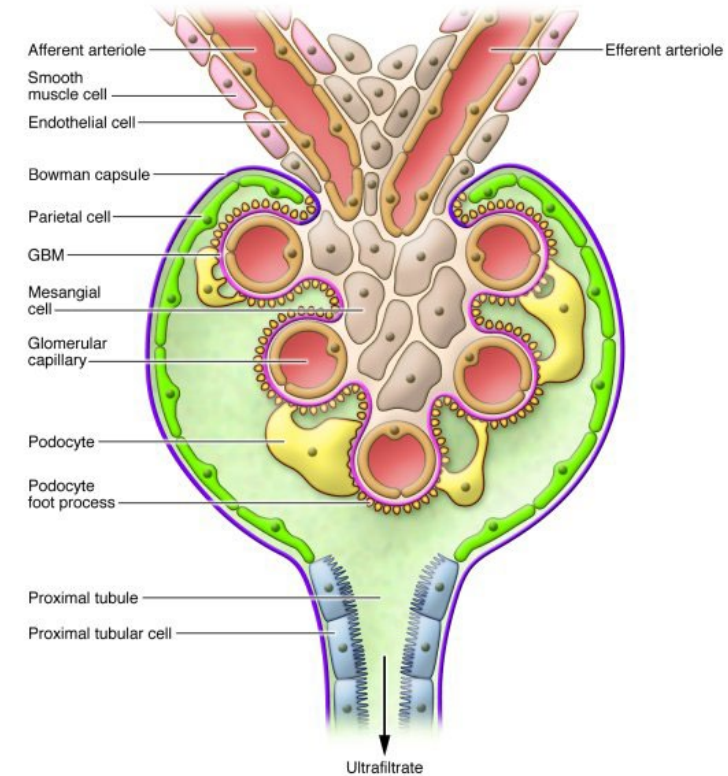
Preclinical GTX data

THERAPEUTIC A1M RAPIDLY DISTRIBUTES TO KIDNEYS

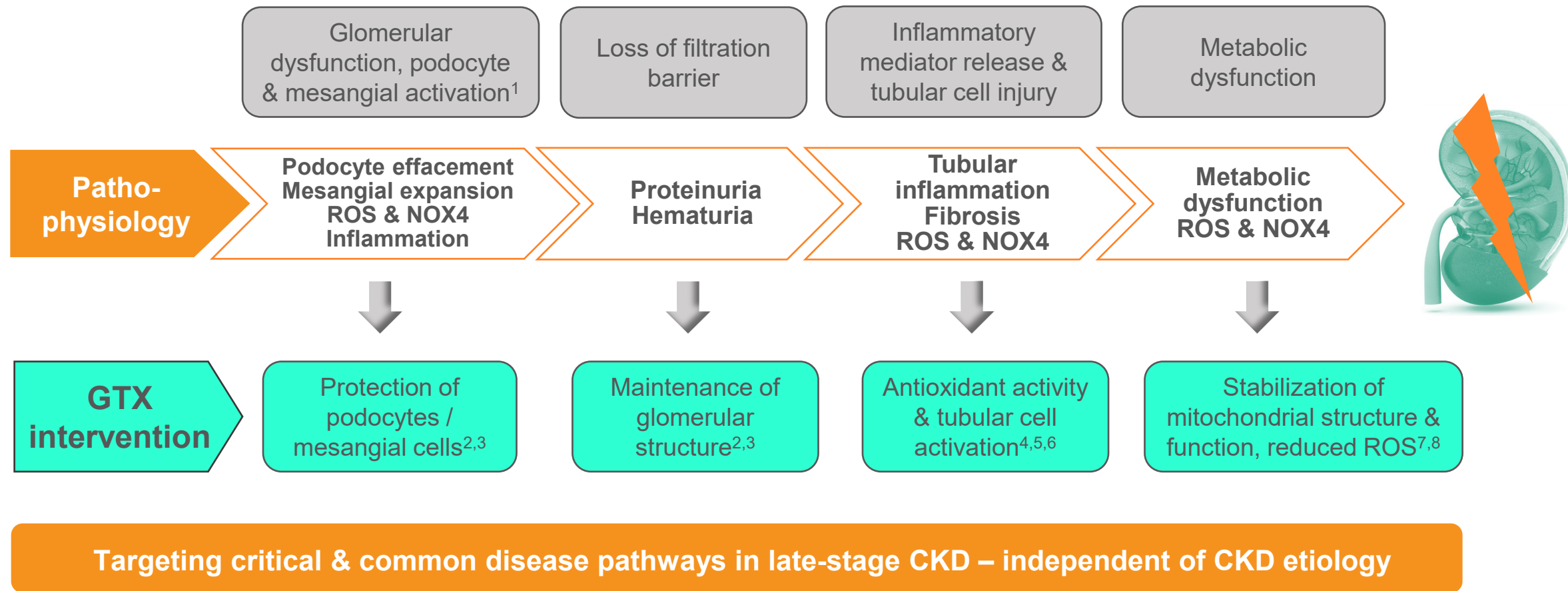
- **Renal clearance** of LMW proteins and peptides (glomerular filtration)
- **Targeted uptake in proximal tubules**
 - Tubular injury is a hallmark of AKI & progressive CKD, driving tubulointerstitial & glomerular fibrosis
- **Distribution profile enables global kidney protection**
 - Glomerular barrier (endothelium, glycocalyx, podocytes, basal membrane)
 - Proximal tubules – primary site of intracellular uptake in kidneys



Combined glomerular & tubular protection against albuminuria & hematuria
(not addressed by available therapies)



THERAPEUTIC POTENTIAL ACROSS THE SPECTRUM OF KIDNEY DISEASE



ROS, reactive oxygen species; NOX4, nicotinamide adenine dinucleotide phosphate oxidase 4

¹ Wickman et al 2016. PLoS One. 11(5): e0155255.

² Nääv et al 2015. PLoS One. 0125499

³ Wester-Rosenlöf et al 2014. 9(1): e86353.

⁴ Åkerström et al 2019. Antioxidants & Redox Signal. 30(4): 489

⁵ Burmakin et al. 2024 Am J Physiol Renal Physiol. doi: 10.1152/ajprenal.00067.2024

⁶ Study reports with in vitro and in vivo GTX activity on file

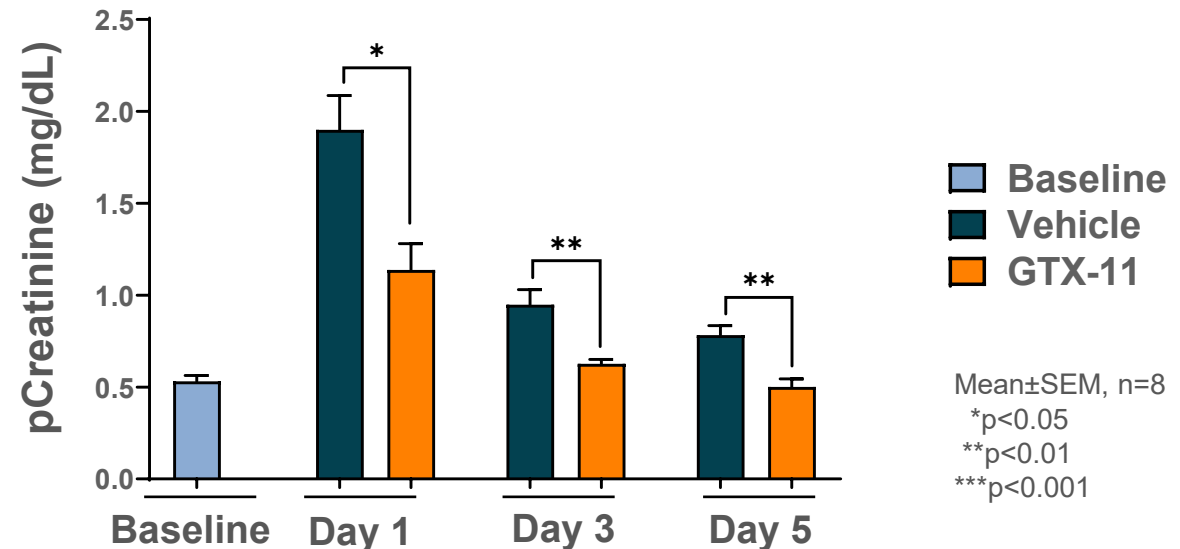
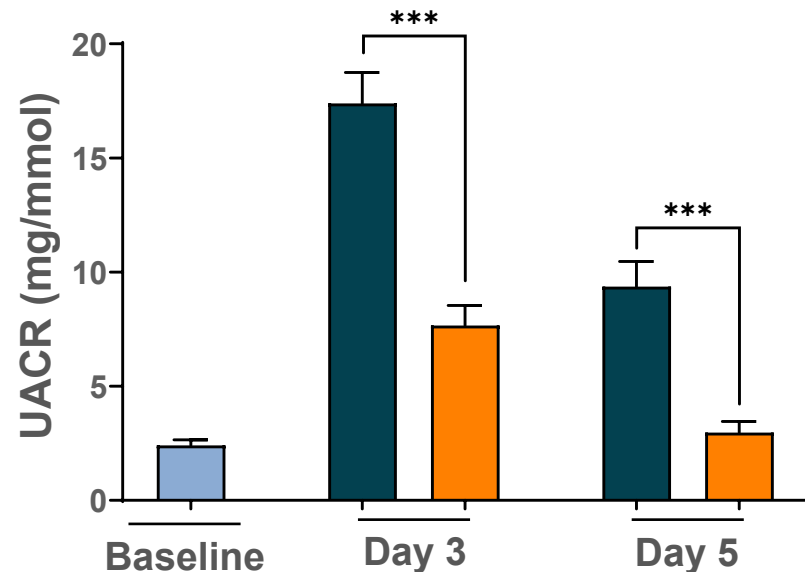
⁷ Kristiansson et al. 2020. Int J Mol Sci. 21(6): 5825

⁸ Olsson et al 2013. Antiox & Redox Signal. 18(6): 2017

GTX-11 IMPROVES KIDNEY FUNCTION & REDUCES ALBUMINURIA

Rat kidney IRI model

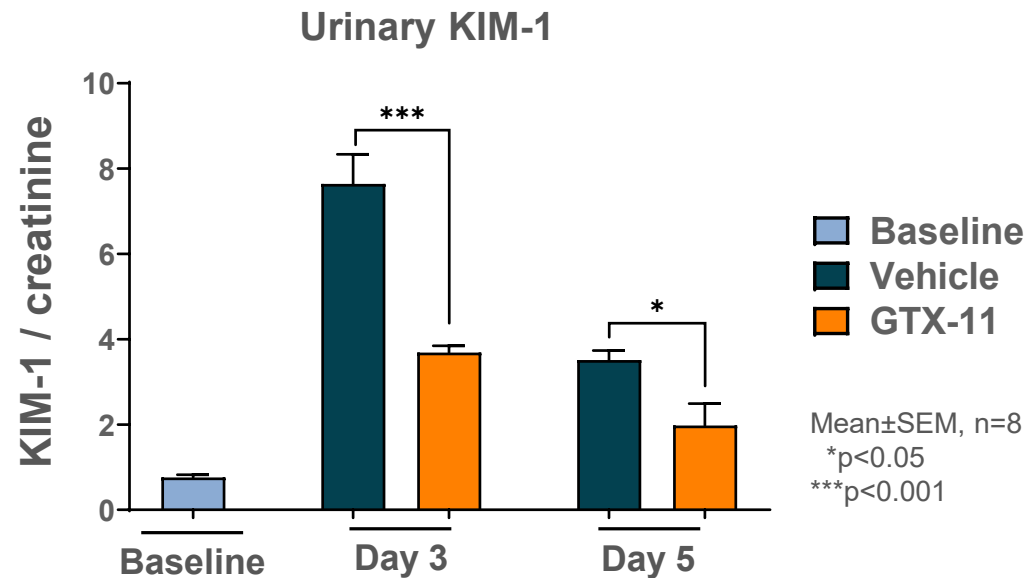
- GTX-11 dosed by IV injection at 5 mg/kg
 - 30 min before renal pedicle clamp ischemia & 4, 8, 24, 48 hours post-reperfusion



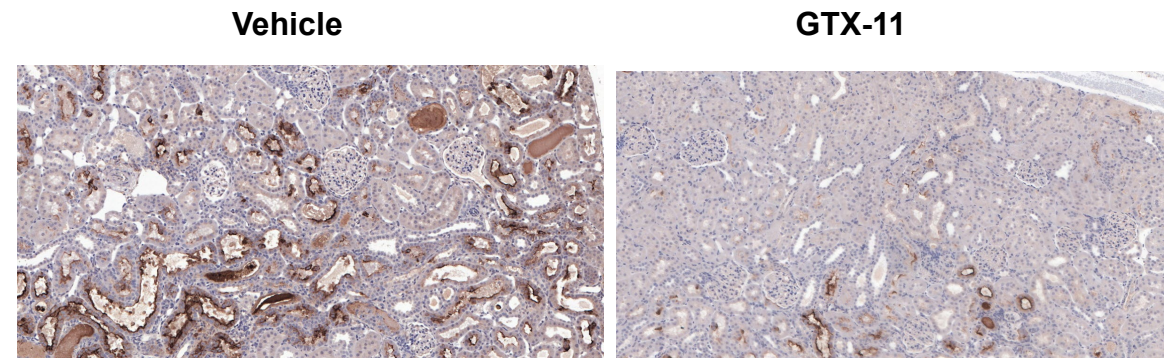
GTX-11 REDUCES KIDNEY INJURY (TISSUE & URINE BIOMARKERS)

Rat kidney IRI model

- KIM-1 (tubular cell injury marker) expression assessed at sacrifice
 - KIM-1 protein assessed in urine by ELISA & tissue expression by IHC
- GTX-11 dosed by IV injection at 5 mg/kg
 - 30 min before ischemia & 4, 8, 24, 48 hours post-reperfusion



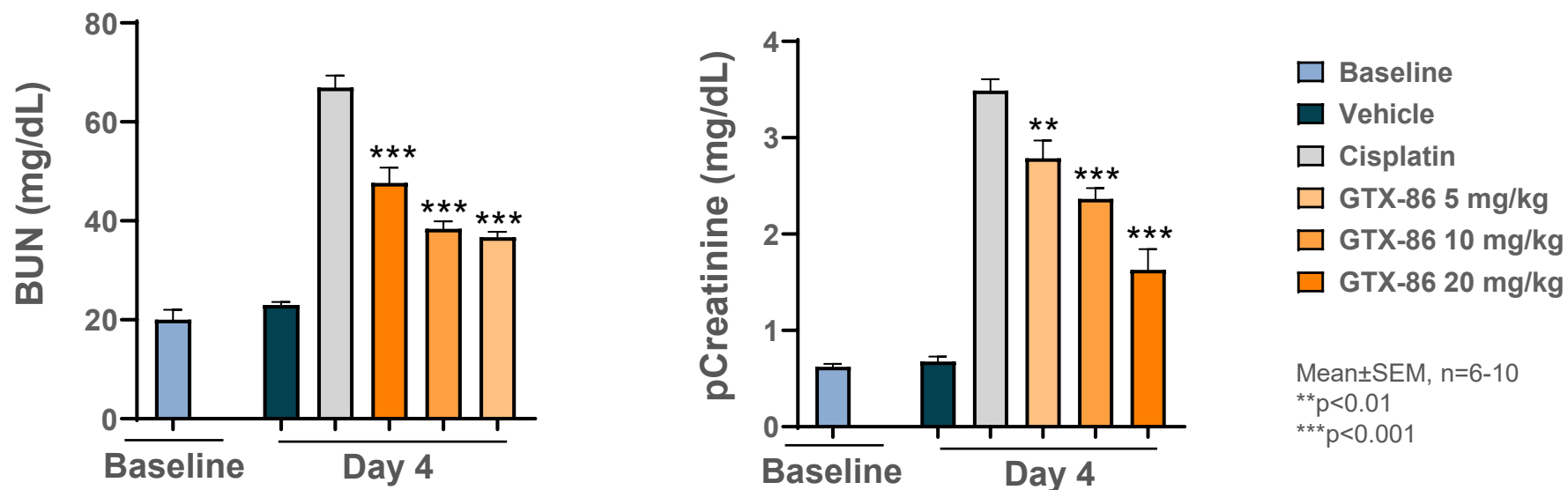
Kidney tissue KIM-1 expression



GTX-86 IMPROVES KIDNEY FUNCTION – DOSE RESPONSE

Mouse cisplatin nephropathy model

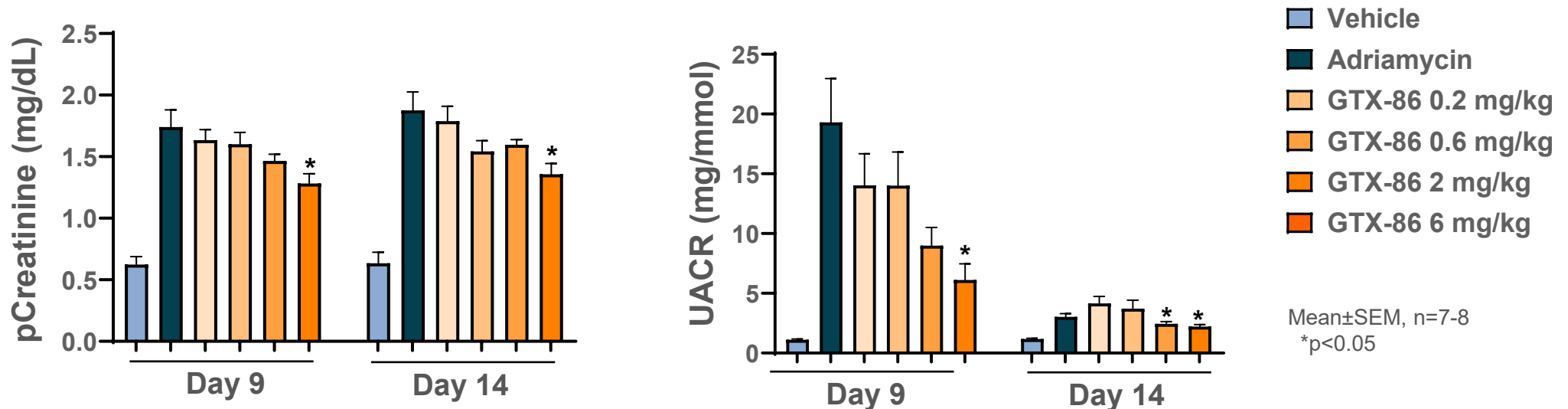
- GTX-86 dosed by SC injection at 5, 10 or 20 mg/kg
 - Once daily injection until sacrifice (first dose in conjunction with cisplatin induction)



GTX-86 IMPROVES KIDNEY FUNCTION & REDUCES ALBUMINURIA – DOSE RESPONSE

Mouse FSGS model

- GTX-86 administered SC at 0.2, 0.6, 2 or 6 mg/kg once daily (initiated before induction with adriamycin)



GTX-86 REDUCES ALBUMINURIA & IMPROVES KIDNEY FUNCTION – DOSE RESPONSE

Mouse diabetic kidney disease model

- Diabetes and renal dysfunction apparent by Day 21 after streptozocin (STZ) administration
- At that point (Day 0), GTX-86 injected SC once daily at either 0.2, 0.6 or 2 mg/kg for 28 days

