# **CORPORATE PRESENTATION**JUNE 2014

WWW.DIAMEDICA.COM





## FORWARD LOOKING STATEMENT

This presentation may contain forward-looking statements, which reflect the Company's current expectation regarding future events. These forward-looking statements involve risks and uncertainties that may cause actual results, events or developments to be materially different from any future results, events or developments expressed or implied by such forward-looking statements. Such factors include, but are not limited to, changing market conditions, the successful and timely completion of clinical studies, the establishment of corporate alliances, the impact of competitive products and pricing, new product development, uncertainties related to the regulatory approval process or the ability to obtain drug product in sufficient quantity or at standards acceptable to health regulatory authorities to complete clinical trial or to meet commercial demand, and other risks detailed from time to time in the Company's ongoing quarterly and annual reporting. Certain of the assumptions made in preparing forward-looking statements include but are not limited to the following: that DM199, DM204 and other programs will generate positive efficacy and safety data in preclinical and future clinical trials; that DiaMedica will complete preclinical and clinical trials within the timelines communicated. Except as required by applicable securities laws, we undertake no obligation to publicly update or revise any forward-looking statements, whether as a result of new information, future events or otherwise.



## INVESTMENT HIGHLIGHTS

#### DEVELOPING BREAKTHROUGH TREATMENTS FOR METABOLIC SYNDROME

#### DM199: Recombinant human tissue kallikrein-1 (KLK-1)

- Novel insulin sensitizer, vasodilator & improves kidney function
- Type 2 diabetes in Phase 2
- Diabetic kidney disease preparing for Phase 2

#### DMDx: Diabetic Kidney disease diagnostic assay

Currently in assay development

#### DM204: mAb designed to activate the BK2 receptor

- Novel insulin sensitizer and vasodilator
- Preclinical development: Type 2 diabetes & blood pressure control

#### **Extensive intellectual property rights worldwide**

20-F has been filed to register the common shares with the SEC



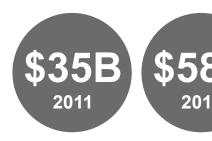
# TYPE 2 DIABETES (T2D): BACKGROUND

## One of the Largest Opportunities in Health Care:

Type 2 Diabetes: insulin resistance resulting in inefficient glucose control



90% are T2D patients1



Market for diabetes drug treatments<sup>2</sup>



T2D Patients not achieving glycemic control target<sup>3</sup>



Total adult (ages 20 - 79) health care expenditure<sup>1</sup>

#### Peferences

- 1. IDF Diabetes Atlas, 6th edn. 2013
- 2. Standard & Poor's 2012
- 3. Adelphi Disease Specific Program DSP III + IV



# DM199: TYPE 2 DIABETES PRODUCT POSITIONING



DM199 restores the body's natural ability to respond to insulin



<sup>\*</sup> Majority in drug classes not recommend for use with diabetic kidney disease

<sup>1.</sup> Datamonitor 2018 Forecast

<sup>2.</sup> Barclays 2020 Forecast

# GLP-1 A RAPIDLY GROWING TREATMENT FOR T2D

2013

2020

\$2.5 billion

\$7.3 billion

# DM199 is the next breakthrough beyond GLP-1's\*

\* Market leaders – Victoza® & Byetta®



# DM199: TARGET PRODUCT PROFILE VS GLP-1

<b>Target Product Profile</b>	DM199 (rKLK1)	GLP-1 Analog
Primary Mechanism	Insulin Sensitizer	Insulin Secretagogue
Glucose Control	✓	✓
Blood Pressure Control	✓	×
Kidney Protection	✓	×
Endothelial Function	✓	×
Weight loss	TBD	✓ (minor)
Dosing	1x weekly s.c.	2x daily – 1x weekly s.c.
# Competing Products	0 (first-in-class)	≥ 26

**DM199** is a full spectrum metabolic syndrome treatment

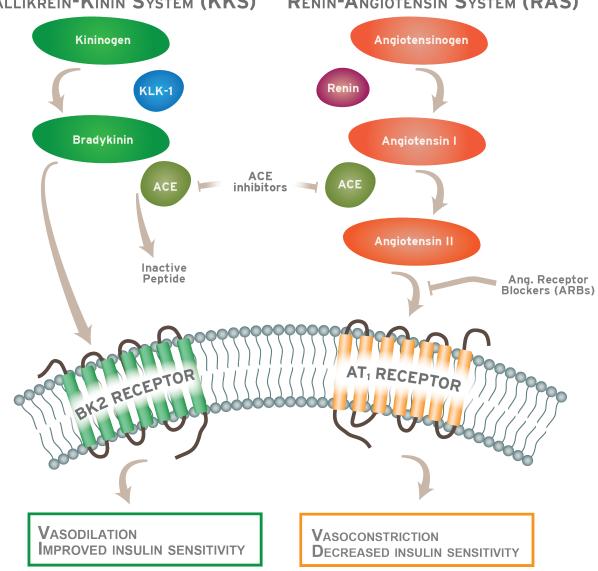


## DM199: MECHANISM OF ACTION

#### IMBALANCE OF RAS SYSTEM CONTRIBUTES TO METABOLIC SYNDROME

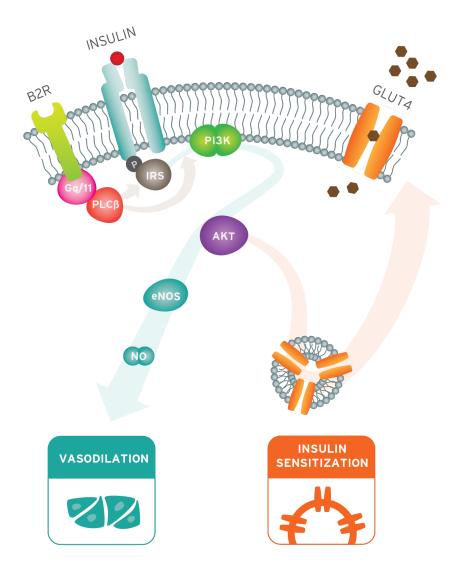
KALLIKREIN-KININ SYSTEM (KKS) RENIN-ANGIOTENSIN SYSTEM (RAS)

DM199 COUNTER BALANCES RAS SYSTEM THROUGH KKS SYSTEM





# DM199: MOLECULAR MECHANISM OF ACTION



#### Nitric Oxide vasodilation

Increase eNos activity

#### Insulin sensitization

 Insulin signaling pathway via GLUT4 translocation

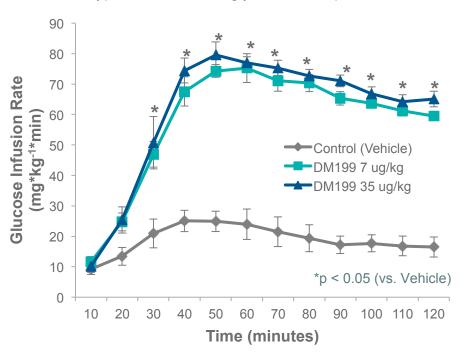




# COMPELLING PRECLINICAL DIABETES DATA

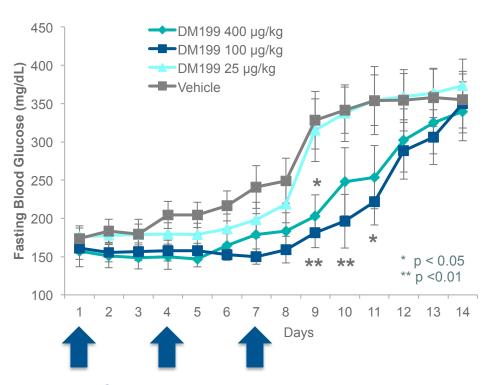
#### SUSTAINED GLUCOSE CONTROL IN TYPE 2 DIABETES MODELS

Study 1: SD Rat Study – single dose Hyperinsulinemic Euglycemic Clamp



Improvement in glucose control

Study 2: ZDF fa/fa Rat Study - 2 weeks DM199 dosed days 1, 4 & 7



**Sustained glucose control** 



# DM199: PHASE 1 CLINICAL TRIALS DATA





**Double-blinded** 

3
PHASE I
TRIALS

Placebo-controlled

**SAFETY: Well-Tolerated** 

**EXTENDED PK PROFILE:** Possible 1x week dosing (Tmax ~24 hr & T<sub>1/2</sub> ~60 hr)



DOSE LIMITING TOLERANCE: Orthostatic Hypotension at 50 μg/kg

# DM199: PHASE 1B TYPE 2 DIABETES EARLY SIGNS OF ACTIVITY

**Double-blinded** 



Placebo-controlled

**SAFETY: Well-Tolerated** 

**REDUCED INSULIN RESISTANCE:** 

Reduced Insulin AUC & HOMA2-IR levels

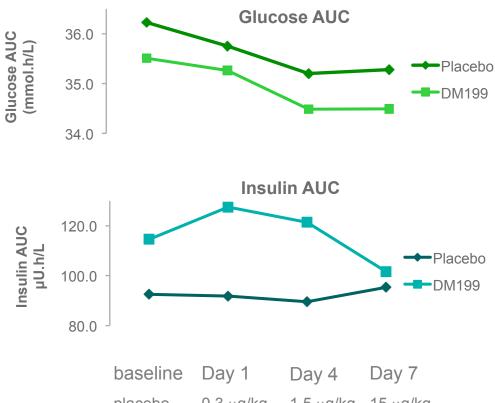


# DM199: PHASE 1B IMPROVEMENT IN INSULIN LEVELS

#### PATIENT BASELINE DIABETIC DEMOGRAPHICS

	Placebo (N=3)	DM199 (N=7)
HbA1c (%)	6.9 (6.6-7.2)	7.0 (6.5-8.8)
HOMA2-IR	1.2 (1.0-1.5)	1.9 (1.1-3.2)
Fasting Glucose (mmol/L)	9.1 (8.7-9.4)	9.1 (7.4-13.0)
Fasting Insulin (µU/mL)	9.8 (8.6-12.1)	14.0 (8.9-22.7)

# **POST PRANDIAL GLUCOSE AND INSULIN AUC**







# **CURRENT PHASE 2: TYPE 2 DIABETES**







# Meal Tolerance Test (MTT), HbA1c, & 7-Point Glucose Profile

**ENDPOINTS** 







# DIABETIC KIDNEY DISEASE MARKETPLACE



Diabetes patients with DKDs<sup>1</sup>



Current treatments limited to anti-hypertensive meds



GLP-1 agonists + other diabetic meds contraindicated for kidney disease



Up to \$100K

Cost of kidney dialysis per patient substantial impact on quality of life

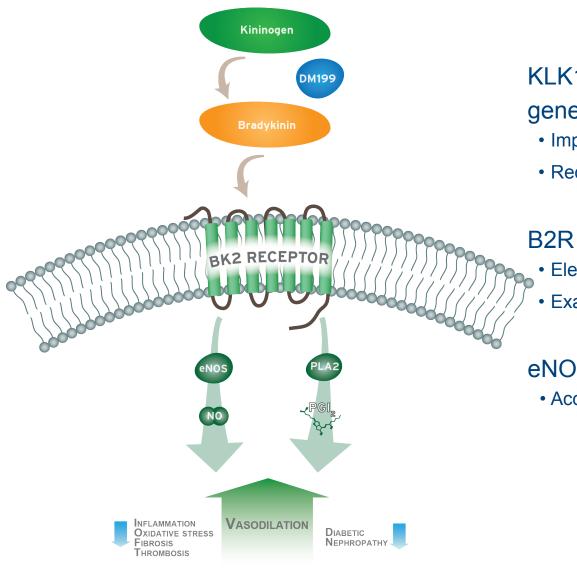
# **DM199** is Positioned to Treat Diabetic Kidney Disease



#### References

- 1. Reutens, AT. Med Clin N Am 2013: 97(1-18)
- National Chronic Kidney Disease Fact Sheet 2014. CDC DDT

# DM199 MECHANISM - DIABETIC KIDNEY DISEASE



## KLK1 (DM199) gene therapy in diabetic rats

- Improved glomerular, tubular functions
- Reduced collagen synthesis, deposition

#### B2R knock out diabetic mice

- Elevated albuminuria
- · Exacerbated fibrosis, GBM thickening

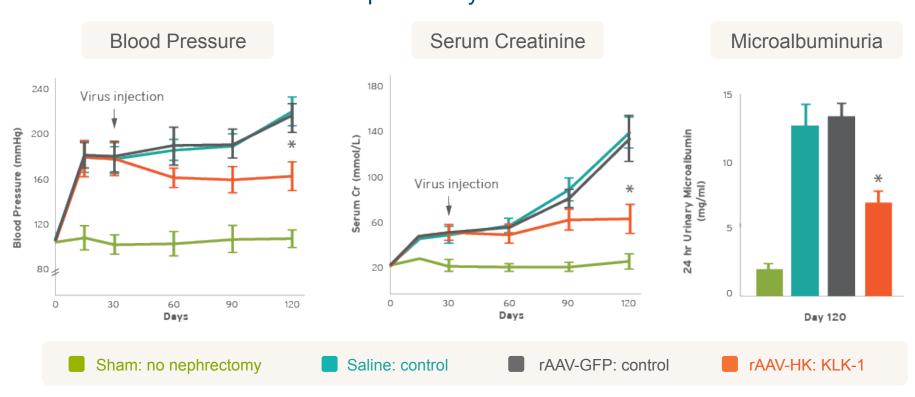
#### eNOS knock out diabetic rats & mice

Accelerated nephropathy



# PRECLINICAL NEPHRECTOMY MODEL

# SUSTAINED KIDNEY PROTECTION IN PRECLINCAL MODEL 5/6<sup>th</sup> Nephrectomy animal model



KLK1 gene therapy stabilized kidney function as measured by significant improvements in:

• Blood pressure, serum creatinine & microalbuminuria vs. untreated control



## PLANNED PHASE 2: DIABETIC KIDNEY DISEASE

# Rationale for DM199 in Diabetic Kidney Disease (DKD)

#### Vasodilator

ACE inhibitors used to treat DKD (complementary mechanism to DM199)

#### **Renal Function**

KLK-1 gene therapy improves blood pressure, creatinine & microalbumin

# Phase 2 Clinical Trial Being Planned

- Up to 150 patients
- 3 6 months treatment
- Clinical endpoints: kidney function biomarkers



# DMDx: DIABETIC KIDNEY DISEASE DIAGNOSTIC

- Current disease diagnosis relies primarily on eGFR for staging disease
  - No diagnostic tools to predict rate of disease progression to end stage renal disease
- KLK-1 levels in urine are potentially correlated with severity of kidney disease
- DMDx is an assay to detect KLK-1 levels in urine and predict disease progression:
  - Enables treatment regime based on rate of progression
  - Identify patients likely to respond to treatment
  - Identify patients best suited for DM199 therapy
    - Benefits reimbursement





# DM204: MECHANISM OF ACTION

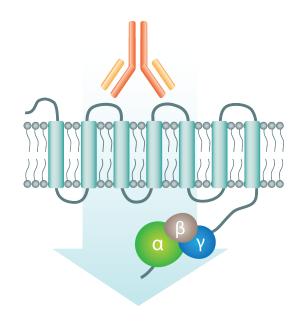
#### mAb designed to activate the BK2 receptor

#### **Advantages:**

- Superior glucose control
- Blood pressure control
- Total cholesterol improvement
- Anticipated infrequent delivery

#### **Status**

- Preclinical development
- Further elucidating the mechanism of action

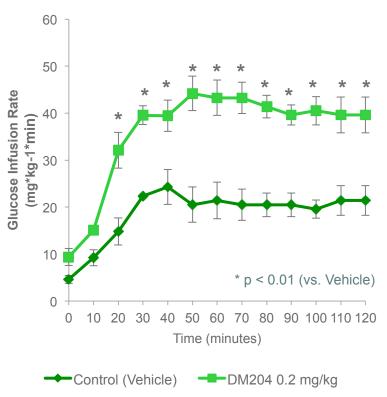




# DM204: PRECLINICAL EFFICACY

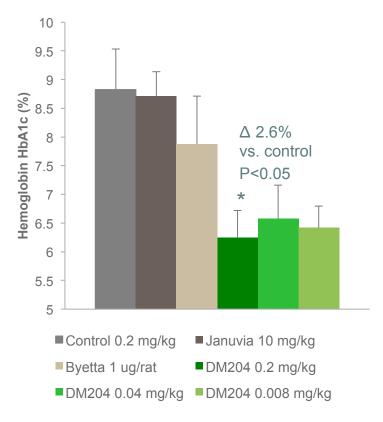
#### ENHANCED GLUCOSE CONTROL

Study 1: SD Rat Study – single dose Hyperinsulinemic Euglycemic Clamp



Improvement in glucose control

Study 2: ZDF fa/fa Rat Study - 3 weeks DM204 dosed 2x week



Improvement in HbA1c



# PRODUCT PIPELINE

#### **Therapeutics INDICATION DISCOVERY** PRE-CLINICAL PHASE I PHASE 2 PHASE 3 DM199 Type 2 Diabetes Diabetic Kidney DM199 Disease Alport Syndrome DM199 DM199 Type 1 Diabetes Other Metabolic DM199 **Indications** Type 2 Diabetes DM204 **Diagnostics ANALYTICAL CLINICAL DISCOVERY DEVELOPMENT** INDICATION **VALIDATION VALIDATION** Diabetic Kidney **DMD**x Disease



# BROAD INTELLECTUAL PROPERTY PROTECTION

Composition of Matter	<ul> <li>DM199: Glycosylation &amp; sialic acid content – novel structure (2033)</li> <li>DM204: Complementarity determining regions for binding to BK2R (2031)</li> </ul>
Methods of Use	<ul> <li>DM199 subcutaneous delivery with improved PK profile (2033)</li> <li>DM199: For treatment of Type 2 Diabetes (2030)</li> <li>DM199: For treatment of Type 1 Diabetes (2032)</li> <li>DM199 for treatment of chronic kidney disease including DKD (2033)</li> <li>DM199 for treatment of NAFLD and NASH (2034)</li> <li>DM204 for Type 2 Diabetes and various other diseases (2031)</li> </ul>
Combination Products	<ul> <li>DM199+GLP-1 analog (2027)</li> <li>DM199+Insulin (2033)</li> <li>DM204+diabetes medications (2031)</li> </ul>
Freedom to Operate DM199 & DM204	No third party patents detected that could block commercialization



## **EXPERIENCED TEAM**

#### **MANAGEMENT**

#### Rick Pauls, MBA

PRESIDENT & CEO

Past co-founder & Managing Director of early stage life sciences venture capital fund

#### Mark Robbins, PhD, JD

VP, CLINICAL & REGULATORY

30+ years biopharma & drug development, 11 successful NDA/BLA approvals

#### Dennis Kim, MD, MBA

CONSULTING CHIEF MEDICAL OFFICER
Previously with Amylin, Orexigen & Enteromedics

#### John Savage, CPA

CHIEF FINANCIAL OFFICER

17+ years with pubic co's. Previously with UnitedHealth Group, Best Buy & Golf Galaxy

#### Mark Williams, PhD

VP, RESEARCH

Co-inventor of DM199 & DM204, 4 phase II trials

#### **SCIENTIFIC ADVISORS**

#### John Amatruda, MD

Former Sr. VP & Franchise Head for Diabetes and Obesity at Merck

Lead development & regulatory approvals for Januvia<sup>™</sup> and Janumet<sup>™</sup> - 1st compounds in the DPP-IV inhibitor class for Type 2 diabetes

#### Paul Burn, PhD

Chair Sanford Project, former SVP at JDRF, Hoffman-La Roche Global Head Metabolic, Bayer Director Metabolic and Lilly Director

#### Alan Cherrington, PhD

Past President of America Diabetes Assoc. Chairman of Molecular Physiology and Biophysics at Vanderbilt U Medical Center Fredrick Banting Award Winner from the ADA

#### **Daniel Porte Jr**, MD

Past President of American Diabetes Assoc. Prof of Medicine, U of California San Diego Council member of National Institute of Diabetes

#### Ralph DeFronzo, MD

Professor of Medicine and Chief of the Diabetes Division at the U of Texas Health Science Center Deputy Director of the Texas Diabetes Institute Banting Award , Lily Award, & Albert Renold Award Winner from the ADA Claude Bernard Award Winner EASD Novartis Award Winner

#### **BOARD OF DIRECTORS**

#### Richard Pilnik, MBA

President, Innovex (a Quintiles Company)
Former VP & Chief Marketing Officer at Eli Lilly
Former Board of Elan Pharma (acquired)

#### Dr. Michael Giuffre, MD, MBA

Clinical Professor of Cardiac Sciences at University of Calgary Board of UNICEF Canada & AB Medical Assoc

#### Rick Pauls, MBA (chairman)

President & CEO

#### **Dawson Reimer, MEAS**

CentreStone Ventures L.P. advisor

#### **Thomas Wellner**

CEO Revera Inc., former CEO CML Healthcare Ltd, (acquired) Senior positions including Brand, Marketing and General Management at Eli Lilly



# **CORPORATE INFORMATION**

Share Price	June 2, 2014	\$0.69
<b>Total Shares Outstanding</b>	June 2, 2014	~61M
Market Capitalization (CDN)	June 2, 2014	~\$42M
Warrants (w/ early expiry clauses)	June 2, 2014	~6.5M
Cash	March 31, 2014 May 2014 capital raise	\$1.6M \$2.2M



# **UPCOMING MILESTONES**

#### DEVELOPING BREAKTHROUGH TREATMENTS FOR METABOLIC SYNDROME

Status
✓
✓
✓
2014 Fall
2014 H2
2015
2014
2014
2015
2014 H1



# CONTACT

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