

Supplemental Table 1. Summary of study characteristics.

Study	Population	Technique	Significant Findings
Mischak et al., 2004 ¹	<p><u>Case:</u> 112 T2D adults with varying UAE</p> <p>46 normal (< 20 mg/L)</p> <p>45 low grade (> 20 mg/L)</p> <p>21 high grade (> 100 mg/L)</p> <p><u>Control:</u> 39 healthy non-smoker adults</p> <p><u>All:</u> had baseline SCr concentrations of < 1.3 mg/dl; no heart failure, cancer, and chronic inflammatory disease</p>	CE-MS/MS; >800 Da cut-off with ≥ 2 charge states; MosaiquesVisu and Mascot software	<p><i>Associated with renal as indicated by increased UAE</i></p> <ul style="list-style-type: none"> - ↑ ALB - ↓ INSL3, UMOD
Meier et al., 2005 ²	<p><u>Case:</u> 44 T1D adolescents</p> <p><u>Control:</u> 9 healthy age-matched volunteers</p>	CE-MS; >800 Da cut-off with ≥2 charge states; MosaiquesVisu software	Identified a urinary protein pattern for early nephropathy that was different from that of controls
Jain et al., 2005 ³	<p><u>Case:</u> 100 T2D adults with microalbuminuria</p> <p><u>Control:</u> 20 healthy adults</p> <p>18 with diabetes and normoalbuminuria</p> <p>1 without diabetes and with microalbuminuria</p> <p>1 without microalbuminuria and diabetes</p>	2DE and MALDI-MS/MS; 5-mL cut-off; ‘Ettan MALDI Software’ with ‘proteo Metrics LLC’ search engine	<p><i>T2D with microalbuminuria v. controls</i></p> <ul style="list-style-type: none"> - ↑ AMBP, AZGP1, ORM
Sharma et al., 2005 ⁴	<p><u>Case:</u> 3 adults with macroalbuminuria and impaired renal function</p> <p>2 T1D and 1 T2D</p> <p><u>Control:</u> 5 healthy adults</p>	Fluorescence-based DIGE and SELDI-MS/MS; Profound search engine and Mascot software	<p><i>Diabetic nephropathy v. controls</i></p> <ul style="list-style-type: none"> - ↑ SERPINA1 <p>Validated SERPINA1 via ELISA</p>
Dihazi et al., 2007 ⁵	<p><u>Case:</u> 117 adults with varying disease</p> <p>38 with T2D nephropathy</p> <p>45 with uncomplicated T2D</p> <p>34 with non-diabetic nephropathy</p> <p><u>Control:</u> 45 healthy adults</p>	GE and SELDI-MS/MS; >1kDa cut-off; MSDB and SwissProt dabatases; ProteinChip array	<p><i>Diabetic nephropathy v. controls</i></p> <ul style="list-style-type: none"> - ↑ B2M - ↓ UBA52

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Otu et al., 2007 ⁶	<u>Case</u> : 31 T2D adults with nephropathy <u>Control</u> : 31 T2D adults with normoalbuminuria; matched for diabetes duration, age, sex, and BMI <u>All</u> : had baseline SCr concentrations of ≤ 1.2 mg/dl; Pima Indians	SELDI-MS; 2-40 kDa range with ≥ 2 signal-to-noise ratio CIPHERgen Biomarker Wizard software	Identified a urinary protein pattern for T2D nephropathy cases that was different from that of controls Survivor bias due to case-control design; unknown stability of urine samples stored for 10 years
Rao et al., 2007 ⁷	<u>Case</u> : 33 T2D adults with varying UAE 10 with normoalbuminuria 13 with microalbuminuria 10 with macroalbuminuria <u>Control</u> : 9 healthy adults	DIGE-MS/MS; immunodepletion of abundant serum proteins; Protein-Lynx Global Server software using SwissProt database	<i>T1D with macroalbuminuria v. normoalbuminuria</i> - \uparrow A1BG, AHSG, AZGP1, GC, HPX, LRG1, SERPINA1, S100A9 - \downarrow AMBP, APOA1, RBP, TTR
Bellei et al., 2008 ⁸	<u>Case</u> : 24 T2D adults with varying UAE 10 with normoalbuminuria 13 with micro- or macroalbuminuria <u>Control</u> : 12 healthy adults	2DE and LC-MS/MS; Mascot software using SwissProt Database	<i>T2D nephropathy v. T2D and controls</i> - \uparrow APOH, B2M, CAH, IGKC, RBP, TTR - \downarrow PPAP, RNA3, KLK3
Rossing et al., 2008 ⁹	<u>Case</u> : 89 T1D adults with varying UAE 30 with normoalbuminuria 29 with microalbuminuria 30 with macroalbuminuria <u>Control</u> : 30 healthy adults	CE-MS and LC-MALDI-MS/MS; <30 kDa cut-off; MosaiquesVisu software	<i>T1D with normoalbuminuria v. controls</i> - \downarrow COL1A1, UMOD <i>T1D with macroalbuminuria v. microalbuminuria</i> - \uparrow ALB, AHSG, B2M, SERPINA1, TTR - \downarrow COL (1A1, 1A2, 3A1), FGB, PGRMC1, PSORS1C2, UMOD
Lapolla et al., 2009 ¹⁰	<u>Case</u> : 30 adults with varying diseases 10 T2D without nephropathy 10 T2D with nephropathy 10 non-diabetic nephropathy <u>Control</u> : 10 healthy adults	LC-MS/MS; <30 kDa cut-off; ProteinPilot v2.1 using Uniprot Database	<i>T2D nephropathy v. T2D and controls</i> - \downarrow COL1A1, UMOD

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Jiang et al., 2009 ¹¹	<u>Case</u> : 162 T2D adults with varying UAE 54 with normoalbuminuria 54 with microalbuminuria 54 with macroalbuminuria or nephropathy <u>Control</u> : 82 healthy adults; matched for age and sex <u>All</u> : Of Chinese Han descent	Fluorescence-based DIGE and MALDI-MS/MS; Mascot software using Swiss-Prot/TrEMBL protein database	<i>T1D with micro- or macroalbuminuria v. controls</i> - ↑ ORM Validated ORM via immunoturbidimetry: increased urinary ORM levels was associated with worsening diabetic kidney disease
Jiang et al., 2009 ¹²	<u>Case</u> : 12 adults with nephropathy 6 T1D and 6 T2D <u>Control</u> : 6 healthy adults; matched for age and sex <u>All</u> : Of Chinese Han descent	2DE and MALDI-MS/MS; Mascot software using Swiss-Prot/TrEMBL protein database;	<i>Diabetic nephropathy v. controls</i> - ↑ ALB, AZGP1, ECAD, IGKC, KNG, PTGDS, ORM, RBP - ↓ AMBP, HP, TTR, UMOD Validated ECAD via ELISA: increased urinary ECAD levels was associated with worsening diabetic kidney disease
Merchant et al., 2009 ¹³	<u>Case</u> : 21 T1D progressors with declining renal function <u>Control</u> : 40 stable T1D non-progressors <u>All</u> : from the Joslin Study of the Natural History of Microalbuminuria in Type 1 Diabetes	LC-MALDI-MS and MALDI-MS/MS; <10 kDa cut-off; Mascot Software using NCBI nr 20060712 database	<i>Progressors v. non-progressors</i> - ↑ FAT2, IPPK, TJP3 - ↓ COL4A1, COL5A1, TNX
Snell-Bergeon et al., 2009 ^{14a}	<u>Case</u> : 19 adults with coronary artery disease 12 with and 4 without T1D <u>Control</u> : 19 adults without coronary artery disease; matched for age, diabetes status and duration, and sex <u>All</u> : from the Coronary Artery Calcification in Type I Diabetes (CACTI) study	CE-MS; <20 kDa cut-off MosaiquesVisu software	- Validated urinary protein patterns identified by and described in Rossing et al 2008 for T1D and T1D nephropathy
Thraikill et al., 2009 ¹⁵	<u>Case</u> : 24 T1D adults with varying UAE 12 normoalbuminuria 12 microalbuminuria <u>Control</u> : 12 healthy adults	Fluorescence-based DIGE and LC-MS/MS; >3 kDa; Mascot and Scaffold software	<i>T1D with microalbuminuria v. other groups</i> - ↑ CLU, CUBN, GC, LRP2, RBP4, EGF, TF, ALB

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Alkhalaf et al., 2010 ^{16b}	<u>Case</u> : 64 T2D adults with nephropathy and retinopathy <u>Control</u> : 82 T2D adults with normoalbuminuria <u>All</u> : from the Prevention of Diabetic Complications (PREDICTIONS) study; aged 35-75 years and had diabetes for ≥5 years	CE-MS and CE-MS/MS; <20 kDa cut-off; Mascot software using the SwissProt database	<i>Diabetic nephropathy v. controls</i> - ↑ AHSB, ALB, B2M, SERPINA1, TTR - ↓ CD99, COL (1A1, 1A2, 3A1), UMOD
Maahs et al., 2010 ^{17a}	<u>Case</u> : 587 diabetic adults (299 T1D) 369 with and 218 without impaired renal function (ACR >30 mg/g or GFR >60 ml/min) <u>Control</u> : 315 healthy adults <u>All</u> : from 10 different hospital centers in the US, Europe and Australia	CE-MS/MS; <20 kDa cut-off; Mascot and MDSB Protein database	<i>T1D and T2D v. controls</i> - ↑ FG (A, B), SERPINA1, - ↓ COL (1A1, 1A2, 2A1, 3A1, 8A2), PGRMC1 <i>T2D v. T1D and controls</i> - ↓ COL (1A1, 1A2) <i>T1D v. controls</i> - ↓ UMOD
Riaz et al., 2010 ¹⁸	<u>Case</u> : 100 T2D patients <u>Control</u> : 43 healthy adults; matched for age and sex <u>All</u> : from a double-blind placebo-controlled randomized clinical trial in Lahore, Pakistan	SDS-PAGE and LC-MALDI-MS/MS; Mascot software using the SwissProt database; ELISA for candidate validation	<i>T2D v. controls</i> - ↑ ALB, AZGP1, ECAD, RBP4 - ↓ AMBP, HP, TTR Verified all 7 candidates via ELISA
Wu et al., 2011 ¹⁹	<u>Case</u> : 75 T2D adults with varying UAE 30 normoalbuminuria 25 microalbuminuria 20 macroalbuminuria <u>Control</u> : 20 healthy sex-matched adults	ProteinChip H50 array; SELDI-MS; < 80 kDa; support vector machine learning	Identified 4 ion fragments with differential excretion between T2D adults with microalbuminuria and normoalbuminuria
Jin et al., 2012 ²⁰	<u>Case</u> : 43 T2D adults with diabetic retinopathy and persistent microalbuminuria <u>Control</u> : 43 T2D healthy adults; matched for age, sex, BMI, and diabetes duration	LC-MS/MS; >3 kDa cut-off; ProteinPilot v.2.0.1 and the Paragon algorithm; iTRAQ for quantification; MRM for candidate validation	<i>T2D with microalbuminuria v. T2D controls</i> - ↑ CP, GC, HP, PSCA, ORM1, SERPINA1, TF - ↓ FABP, HSPG2, MASP2 Verified HP, ORM, PSCA, SERPINA1, and TF via MRM

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Schlatzer et al., 2012 ²¹	<u>Case</u> : 13 T1D progressors who developed micro- or macroalbuminuria <u>Control</u> : 11 T1D non-progressors; matched for diabetes duration and age <u>All</u> : from CACTI study	LC-MS/MS; unknown cut-off; Proteomarker and Mascot softwares	<i>Progressors v. non-progressors</i> - ↑ ORM - ↓ CLU, GRN, UMOD Verified all 4 candidates via ELISA
Soggiu et al., 2012 ²²	<u>Case</u> : 20 T1D adults with varying UAE 16 normoalbuminuria 4 microalbuminuria <u>Control</u> : 10 healthy adults	MALDI-MS and LC-MS/MS; >10 kDa cut-off; Mascot software	<i>T1D (particularly with microalbuminuria) v. controls</i> - ↑ A1BG, AMBP, AZGP1, RBP4 - ↓ APO (A1, E), CD59, HMWK, UMOD
Zürbig et al., 2012 ^{23c}	<u>Case</u> : 15 diabetic progressors who developed macroalbuminuria 6 T1D and 9 T2D <u>Control</u> : 20 diabetic non-progressors who remained normoalbuminuric 10 T1D and 10 T2D	CE-MS; <20 kDa cut-off; MosaiquesVisu software	<i>Progressors v. non-progressors</i> - ↑ ALB, SERPINA1 - ↓ COL1A1, CD99, CLU, PIGR, UMOD
Bhensdadia et al., 2013 ²⁴	<u>Case</u> : 4 T2D progressors (UAE increased by >60% by the end of follow-up) with worsening SCr <u>Control</u> : 4 T2D non-progressors with stable SCr; matched for baseline SCr and UAE <u>All</u> : from Veterans Affairs Diabetes Trial (VADT)	LC-MS/MS; Mascot software and Scaffold; selected reaction monitoring (SRM) and ELISA for validation	<i>Progressors v. non-progressors</i> - ↑ AGRN, AGT, HP, MASP2 - ↓ LAMP Verified the 5 markers above, UMOD, and NGAL in 30 adults via SRM; validated HP via ELISA in 204 adults
Chu et al., 2013 ²⁵	<u>Case</u> : 28 uncomplicated T2D adults <u>Control</u> : 29 healthy adults	LC-MALDI-MS/MS; 10 kDa cut-off; Bioworks Browser	<i>T2D v. controls</i> - ↓ CLU, EPRS, HINT1
Roscioni et al., 2013 ^{26c}	<u>Case</u> : 44 T2D progressors with worsening UAE <u>Control</u> : 44 T2D non-progressors <u>All</u> : from the Prevention of Renal and Vascular End-stage Disease (PREVEND) study	CE-MS/MS; <20 kDa cut-off; MosaiquesVisu software	<i>Progressors v. non-progressors</i> - ↑ AHSG - ↓ COL1A1 <i>Associated with UAE</i> - ↑ AHSG, ALB, SERPINA1 - ↓ COL1A1, UMOD

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Siwy et al., 2014 ^{27c}	<u>Case:</u> 87 T2D adults with nephropathy <u>Control:</u> 78 T2D adults with normoalbuminuria <u>All:</u> from the Proteomic Prediction and Renin Angiotensin Aldosterone System Inhibition Prevention of Early Diabetic Nephropathy In Type 2 diabetic adults with normoalbuminuria (PRIORITY) trial	CE-MS; <20 kDa cut-off; MosaiquesVisu software	<i>T2D nephropathy v. control</i> - ↑ A1BG, AHSG, ALB, APOA1, B2M, SERPIN (A1, C1), TTR - ↓ CD99, COL (1A1, 1A2), FXYD2, FGA, PGRMC1, PIGR
Lewandowicz et al., 2015 ²⁸	<u>Case:</u> 72 T2D adults 33 without retinopathy or nephropathy 15 with diabetic retinopathy 24 with diabetic nephropathy (and retinopathy) <u>Control:</u> 27 healthy adults matched for age and sex	LC-MS/MS; unknown cut-off; iTRAQ quantitation; Mascot Software	<i>T2D nephropathy v. control</i> - ↑ ALB, SERPINA1 - ↓ COL1A1, HSPG2 <i>T2D v. control</i> - ↑ ALB, PTGDS, SPP1 - ↓ ACTB, APOA1, COL1A1, HSPG2
Suh et al., 2015 ²⁹	<u>Case:</u> 40 T1D children and adolescents <u>Control:</u> 41 healthy age-matched siblings	SDS-PAGE and FASP; LC-MS/MS; > 30 kDa; Mascot Software and Protein Prophet	<i>T1D v. control</i> - ↑ ENPEP, NAGA, MAN2B1, CTSC, FUCA1, ASAH1, GNS, FUCA2, DPP7, CPQ, HEXB, CTSB, LRG1, CST2, RBP, APOM, GAS6, GP5, TIMP1, SLC3A2, SELL, CDH5, MCAM, MSLN, PI16, VCAM1, COLEC12, ALCAM, ACE2, ERP44, HK3, GSN - ↓ LAMP, KNG1, ACY3, AMN, ERP MGAM, IGFLR1, MADCAM1, CPM, RNF149, TOLLIP, HRSP12, CTTN Verified 6 proteins (FUCA2, NAGA, COLEC-12, CD166, TIMP1, and APOM) via Western Blot. (These proteins were subsequently used into our bioinformatic analyses.)

Study	Population	Technique	Significant Findings
Zhang et al., 2015 ³⁰	<u>Case</u> : 49 T2D adults divided into groups based HbA1c levels [BM] and fasting plasma glucose levels [GM] <u>Control</u> : 29 healthy adults matched for age and sex	LC-MALDI-MS/MS; <10 kDa cut-off; ClinProt software using the IPI human database	<i>T2D v. control</i> - ↑ FGA, F2 <i>T2D in high BM group v. low BM group</i> - ↑ FGA, F2
Fu et al., 2016 ³¹	<u>Case</u> : 28 T2D adults with microalbuminuria <u>Control</u> : 30 T2D adults with normoalbuminuria	LC-MALDI-MS/MS; <10 kDa cut-off; ClinProt software using the IPI human database	<i>T2D with microalbuminuria v. T2D controls</i> - ↑ F2, FGA, VTN - ↓ F2, FGA ITIH4, VTN Peptides of F2, FGA, and VTN were increased and decreased Verified via MALDI-TOF-MS/MS

^aStudies that also validated the diabetes7 model¹⁴

^bStudies that also validated the Rossing peptidome⁹

^cStudies that also validated the CKD273 classifier³²

2DE, two-dimensional gel electrophoresis; CE, capillary electrophoresis; DIGE, difference gel electrophoresis; ELISA, enzyme-linked immunosorbent assay; GE, gel electrophoresis; LC, liquid chromatography; MALDI, Matrix-assisted laser desorption/ionization; MRM, multiple reaction monitoring; SRM, selected reaction monitoring; T1D, type 1 diabetes; T2D, type 2 diabetes; UAE, urinary albumin excretion.