

Supplementary Methods

Calculation of F-measure

The F-measure can be calculated for a given trade-off between precision and recall as

$$F_{x/y} = \frac{\left(1 + \left(\frac{x}{y}\right)^2\right) \times (P \times R)}{\left(\frac{x}{y}\right)^2 \times P + R} \quad \text{where } x/y \text{ times as much weight is given to recall as precision. We set}$$

$x=1$ and $y=5$. This weighs precision five times as much as recall when selecting the appropriate cut-off for experimental and clinical validation.

Correlation based prediction of cell-lineage-specific genes

We used a correlation-based strategy for predicting cell-lineage specific expression as a baseline method. We used the Sleipnir library for functional genomics to calculate a z-transformed correlation of the gene expression for all genes in the micro-dissected human kidney data. We then ranked genes by their average z-transformed correlation to all genes in the podocyte gold standard. While the SVM was able to successfully separate different glomerular compartments (Fig. 2), the baseline correlation approach did not achieve useful separation (SI Fig. 1).

Gene expression data extraction from the GUDMAP

Briefly, podocytes, mesangial cells and endothelial cells were isolated from kidneys of adult Mafb-EGFP transgenic mice (Tg(Mafb-EGFP)79Gsat), Meis-EGFP transgenic mice (Tg(Meis1-EGFP)FO156Gsat), and Tie2-GFP transgenic mice (Tg(TIE2GFP)287Sato) respectively, using a combination of collagenase A digestion, sieving and trypsinization followed by Fluorescence Activated Cell Sorting (FACS). Affymetrix Mouse Gene 1.0 ST Arrays were used by GUDMAP to obtain gene expression data following the Affymetrix standard protocol.

The robustness *in silico* nano-dissection to poor or limited standards

To assess the robustness of our method when our lineage-specific positive markers are limited or of varying quality, we examine the performance of nano-dissection under two sets of conditions. In the first case, we subsample from our gold standard to create a smaller gold standard containing only 50-90% of the genes in our full standard (SI Fig 4). In the second case, we add 1 to 4 identically sized standards made up of randomly selected genes to our gold standard. This generates new standard sets 2-5 times the size of our gold standard. We then compare nano-dissection's performance to that of linear SVM using these sets of gold standards (SI Fig 5). In both cases, we observe that nano-dissection is robust to such perturbations and effectively separates podocyte and non-podocyte genes, even under such sub-optimal conditions.

Comparison of *in silico* nano-dissection to genome-wide GFR correlation

To evaluate whether nano-dissection provides information not available from a genome-wide GFR correlation analysis when the goal is to identify genes associated with diseases that show a cell lineage specific etiology, we performed a systematic analysis focused on hereditary glomerular disease. We curated genes associated with hereditary FSGS in prior literature and excluded any that were included within the positive gold standard for nano-dissection. These thirteen FSGS-associated genes (*APOL1*, *ARHGAP24*, *COL4A3*, *COL4A4*, *INF2*, *ITGB4*, *LAMB2*, *MYH9*, *MYO1E*, *PLA2R1*, *SCARB2*, *SMARCAL1*, and *TSPAN13*) do not overlap with our podocyte positive gold standards.

First, we evaluated, for all genes, the absolute value of their expressions' correlation with the square root of GFR measurements, termed the "GFR correlation score". The absolute value of this correlation is used because genes associated with a disease may either have a positive or negative relationship with renal function. Genes associated with hereditary FSGS (red, SI Fig. 8A) were observed to have a greater GFR correlation score than genes without demonstrated FSGS association (blue, SI Fig. 8A), but the relationship was not statistically significant (t-test $p = 0.17$). Thus a genome-wide assessment of GFR

correlation alone was not capable of identifying genes associated with this hereditary renal disease. In contrast to the results for the GFR correlation score, genes associated with FSGS were given significantly higher scores by podocyte nano-dissection (red, SI Fig. 8B) (t-test $p = 0.005$), as compared to nano-dissection scores of genes without known FSGS association (blue, SI Fig. 8B).

Figure legends for supplementary figures:

Supplementary Fig. 1: The distribution of tissue specific genes by percentile using classical correlation approach. This approach was not able to separate known podocyte genes from genes specific to the other glomerular cell-lineages and tubular cells.

Supplementary Fig. 2: Nano-dissection is applicable to non-renal cell lineages. Here we separate astrocyte (green) genes from microglia (blue) and oligodendrocyte (red) genes. Cell-type-specific standards for each type were obtained from HPRD, and nano-dissection was applied to the hand-curated compendium of brain samples available in the nano-dissection webserver (analysis URL: <http://nano.princeton.edu/analysis/view/8c02f58c-10c0-4693-832b-f40a0e8c9cca/>). The top 150 predictions (dotted line), as used in our podocyte study, provide a highly accurate separation of astrocytes from the other lineages.

Supplementary Fig. 3: Nano-dissection is applicable to non-renal cell lineages, even on whole-organism data compendia. Here we separate fibroblast (red) genes from keratinocyte (green) and melanocyte (blue) genes. Cell-type-specific standards for each type were obtained from HPRD, and nano-dissection was applied to the entire human gene-expression compendium available in the nano-dissection webserver (analysis URL: <http://nano.princeton.edu/analysis/view/f8e6845c-ed51-433e-b558-5782ac885154/>). The top 150 predictions (dotted line), as used in our podocyte study, provide a highly accurate separation of fibroblasts from the other lineages.

Supplemental Fig. 4: Nano-dissection's performance is robust to limited gold standards and those of varying quality. The observed performance is shown at 100% of standard (blue dot). The performance of 50 trials sub-sampled to 50-90% of the existing standard size is shown to the left of 100% and red bars indicate 95% confidence intervals.

Supplementary Fig. 5: Iterative nano-dissection approach is robust to low quality gold standards, whereas SVM is not. We evaluate both our method and the linear SVM in the context of adding a contaminating random gold standard. The plot shows the AUC of each approach when a random standard of 1, 2, 3, or 4 times the size of our gold standard is added as an additional standard. The iterative nano-dissection approach identifies that the standard is useless and avoids it, while the SVM's performance deteriorates substantially. The shaded regions show 95% confidence intervals (nonparametric bootstrap with 10,000 resamples) around the mean.

Supplementary Fig. 6: Nano-dissection is applicable to non-podocyte renal lineages. Here we separate mesangial (light blue) genes from endothelial (dark blue), podocyte (green) and tubular (red) genes. Nano-dissection was applied to the renal microdissection compendium available in the nano-dissection webserver (analysis URL: <http://nano.princeton.edu/analysis/view/Mesangial-nano-dissection-hUjTVxrPBnP38T2c/>).

Supplementary Fig. 7: Pseudocode for our algorithm showing the nano-dissection decision rule. The array $d[]$ represents the results of the chosen classification algorithm, here the distance to the hyperplane from the SVM. The lineage-specific validation procedure is used when the validation standard is specific to the lineage of interest, which also occurs when no external validation standard is available. The compartment validation procedure is used when the validation standard includes lineages expected to occur in both the positive and negative standards. Also see the Methods section for an in depth description.

Supplementary Fig. 8: Genes associated with FSGS do not have a significantly different correlation to GFR (A, red bar) than genes with unknown status (A, blue bar), but FSGS associated genes do have significantly higher nano-dissection scores (B, red bar) than genes not known to be associated with FSGS (B, blue bar). Error bars indicate 95% confidence intervals as assessed by 1000 bootstrap samples.

Supplementary Table 1. Positive standards for podocytes selected by literature search.

Supplementary Table 2. Negative standards for podocytes selected by literature search.

Supplementary Table 3. Top 136 predicted genes by *in silico* nano-dissection approach.

Supplementary Table 4. GUDMAP gene expression datasets have been used for this study.

Supplementary Table 5. 102 genes exhibit significantly higher expression in podocytes in comparison with the other two major cell types in mouse glomerulus.

Supplementary Table 6. Demographics of 139 CKD patients.

Supplementary figure 1

density

0

20

40

60

80

100

percentile

0.020

0.015

0.010

0.005

0.000

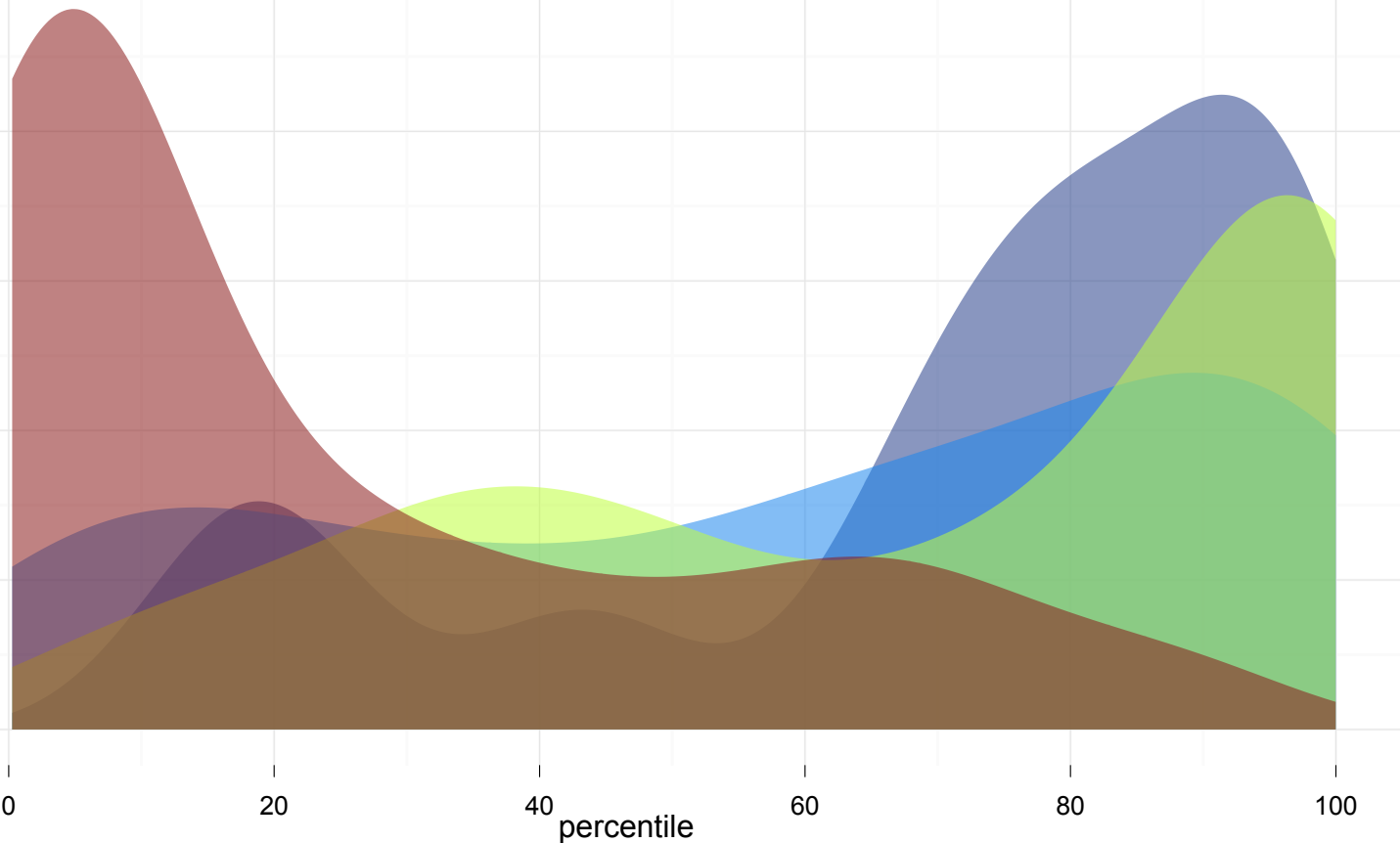
cell type

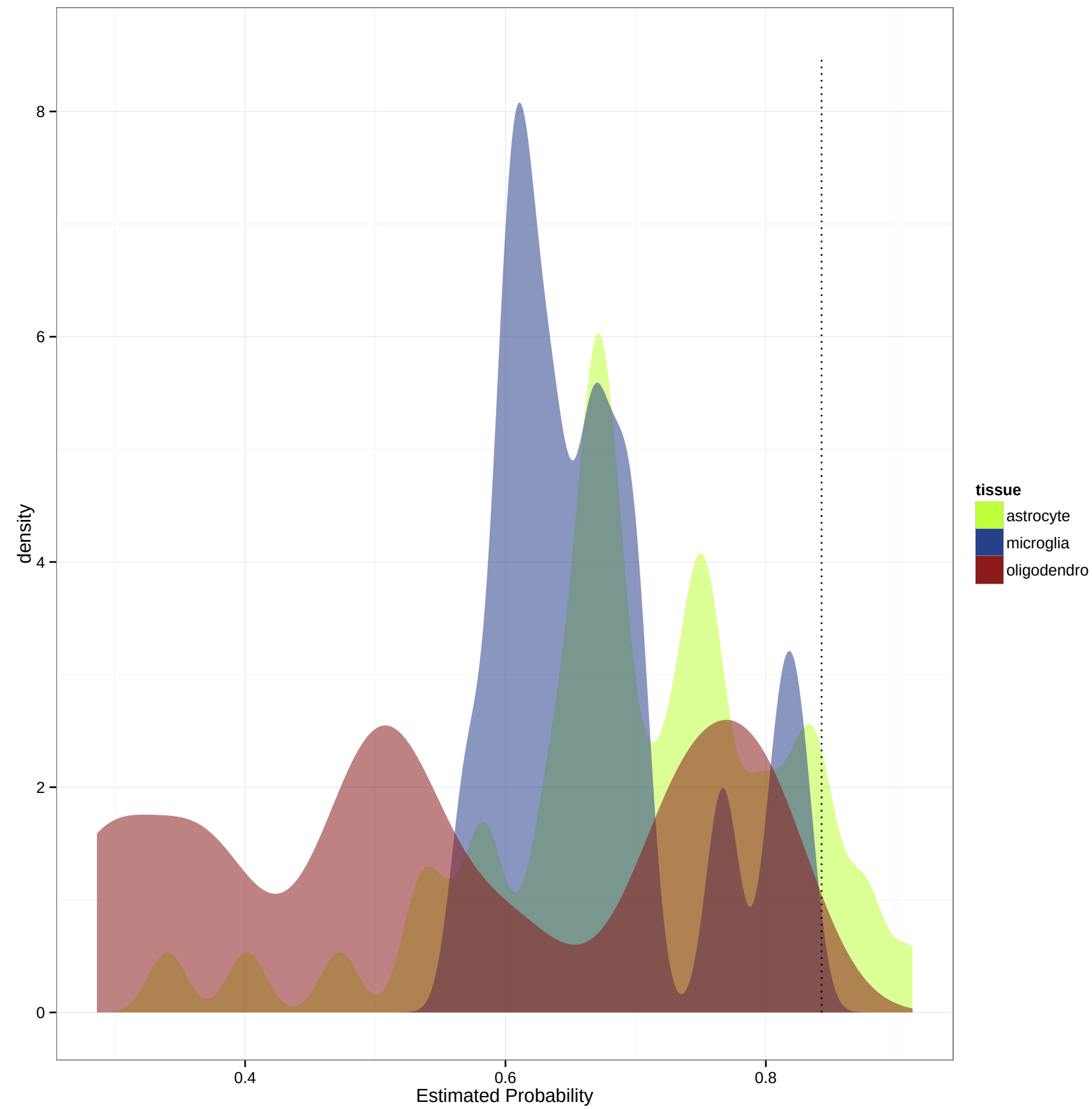
podocyte

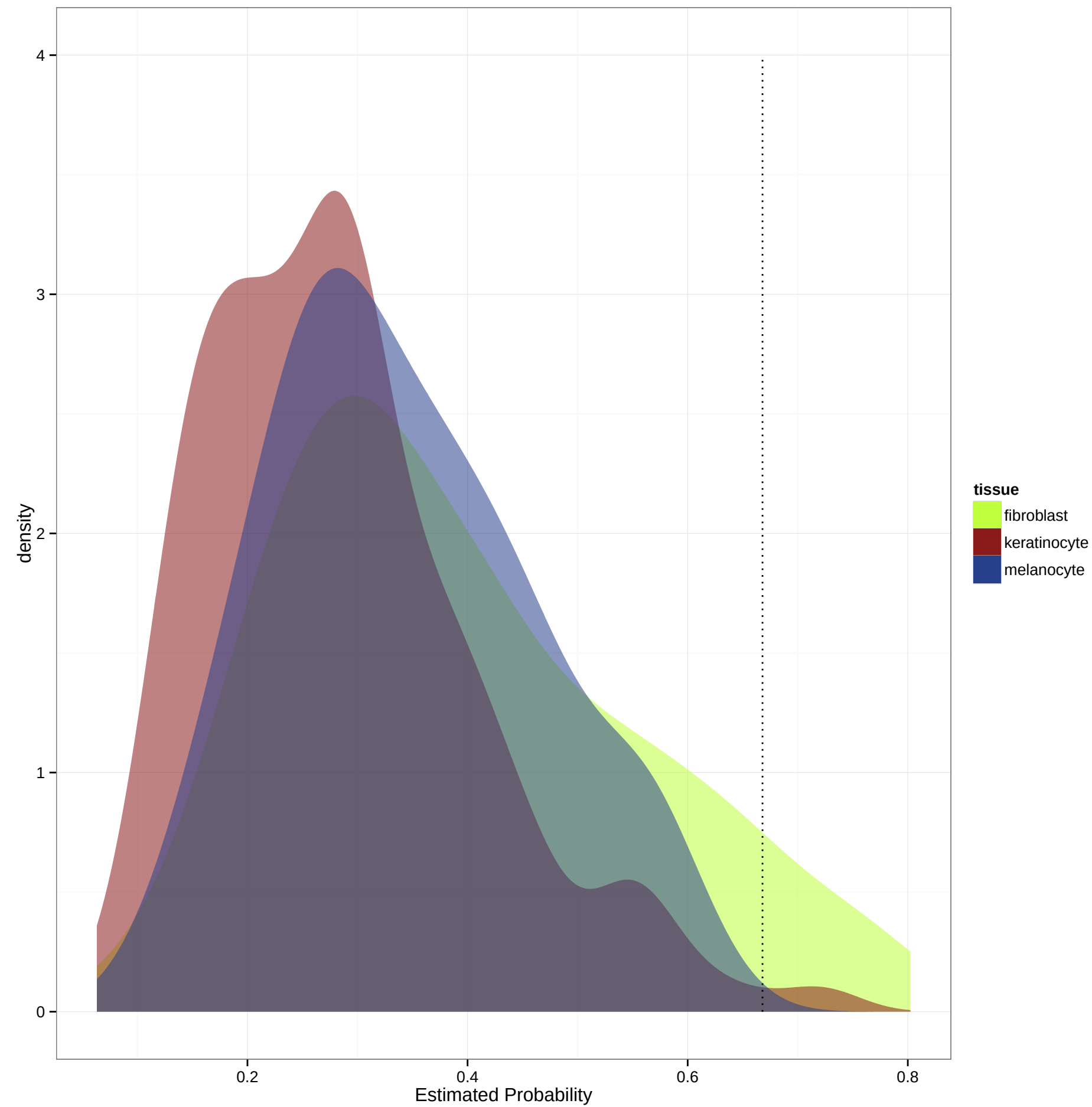
endothelial

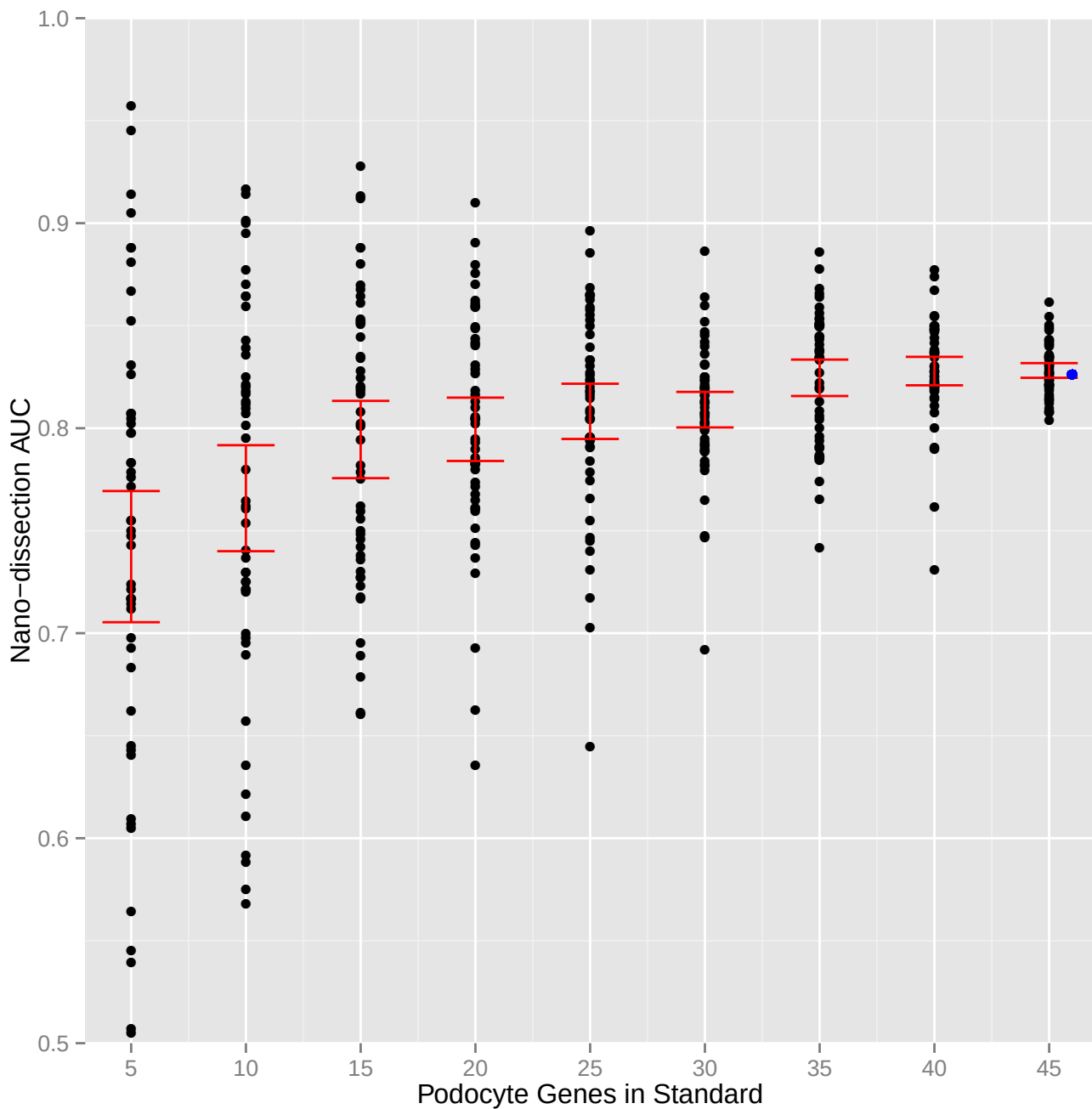
messangial

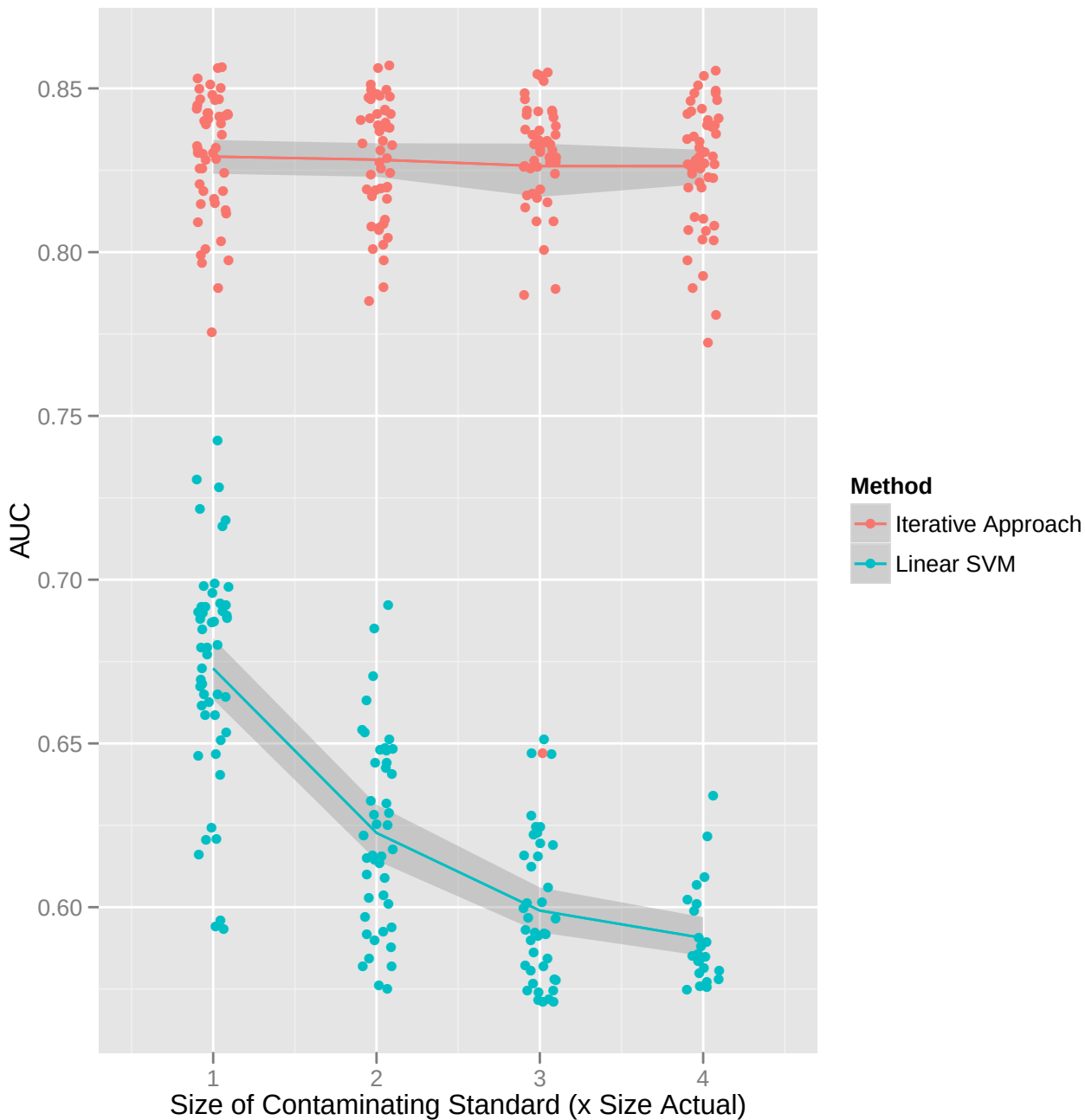
tubular

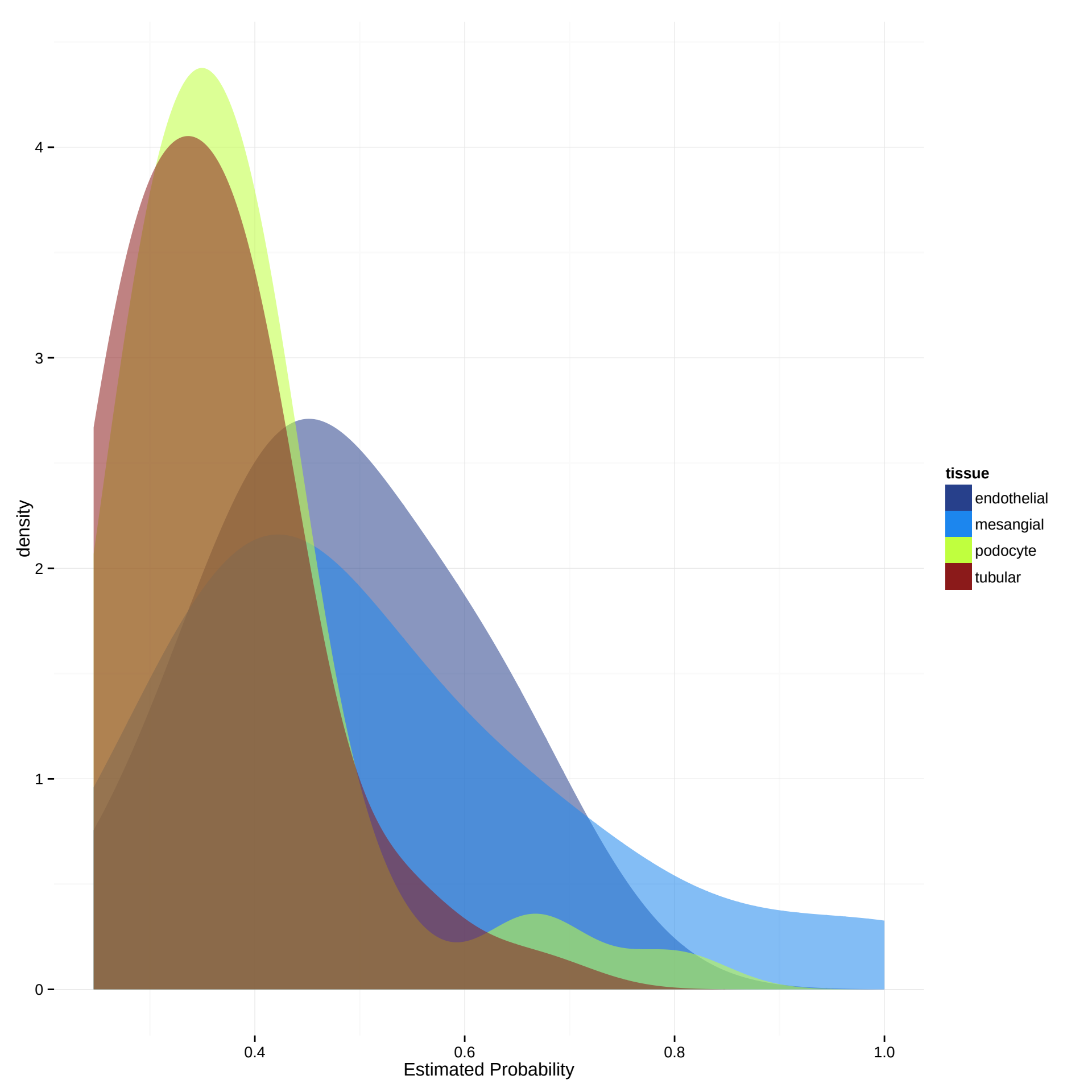






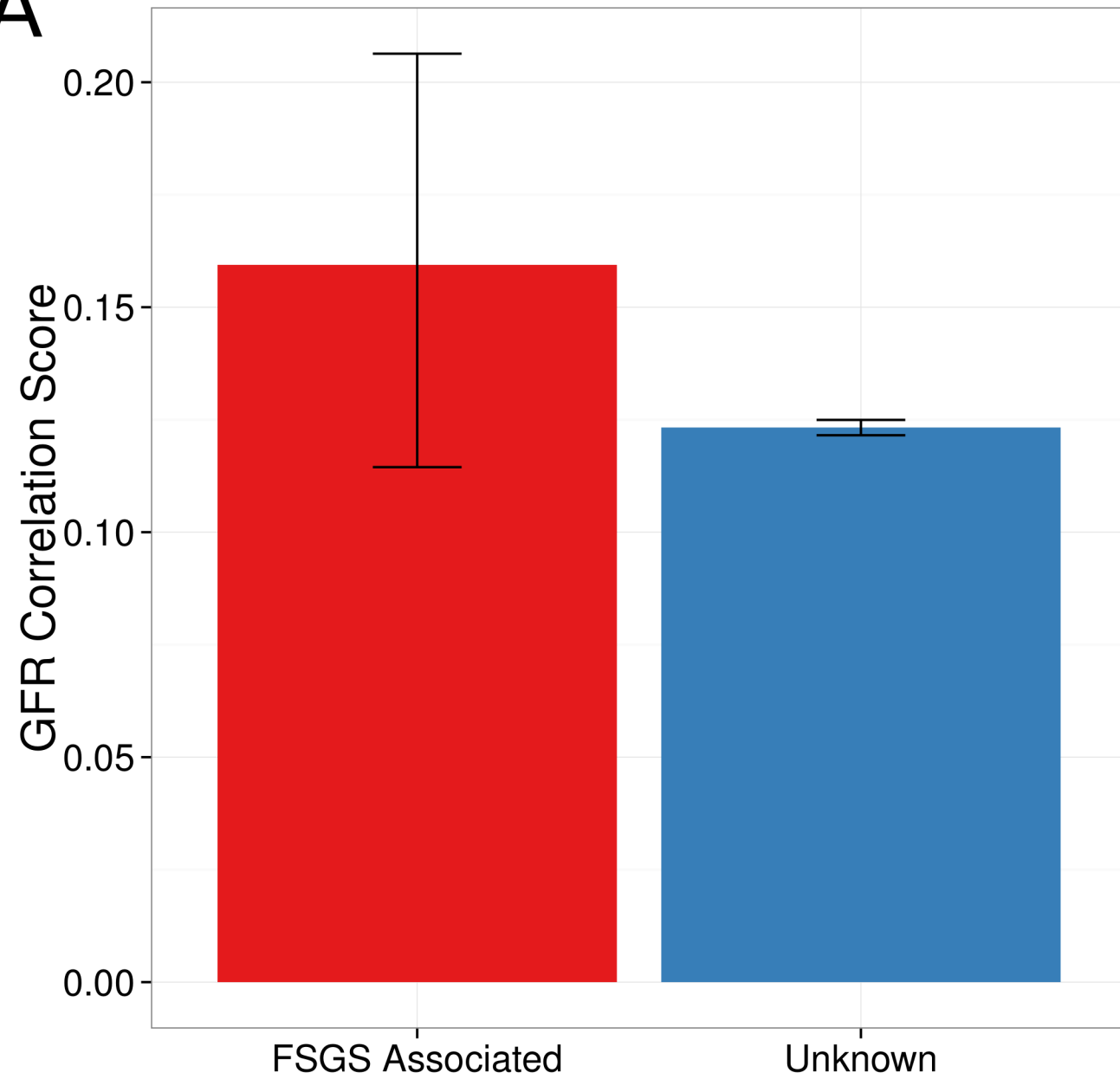
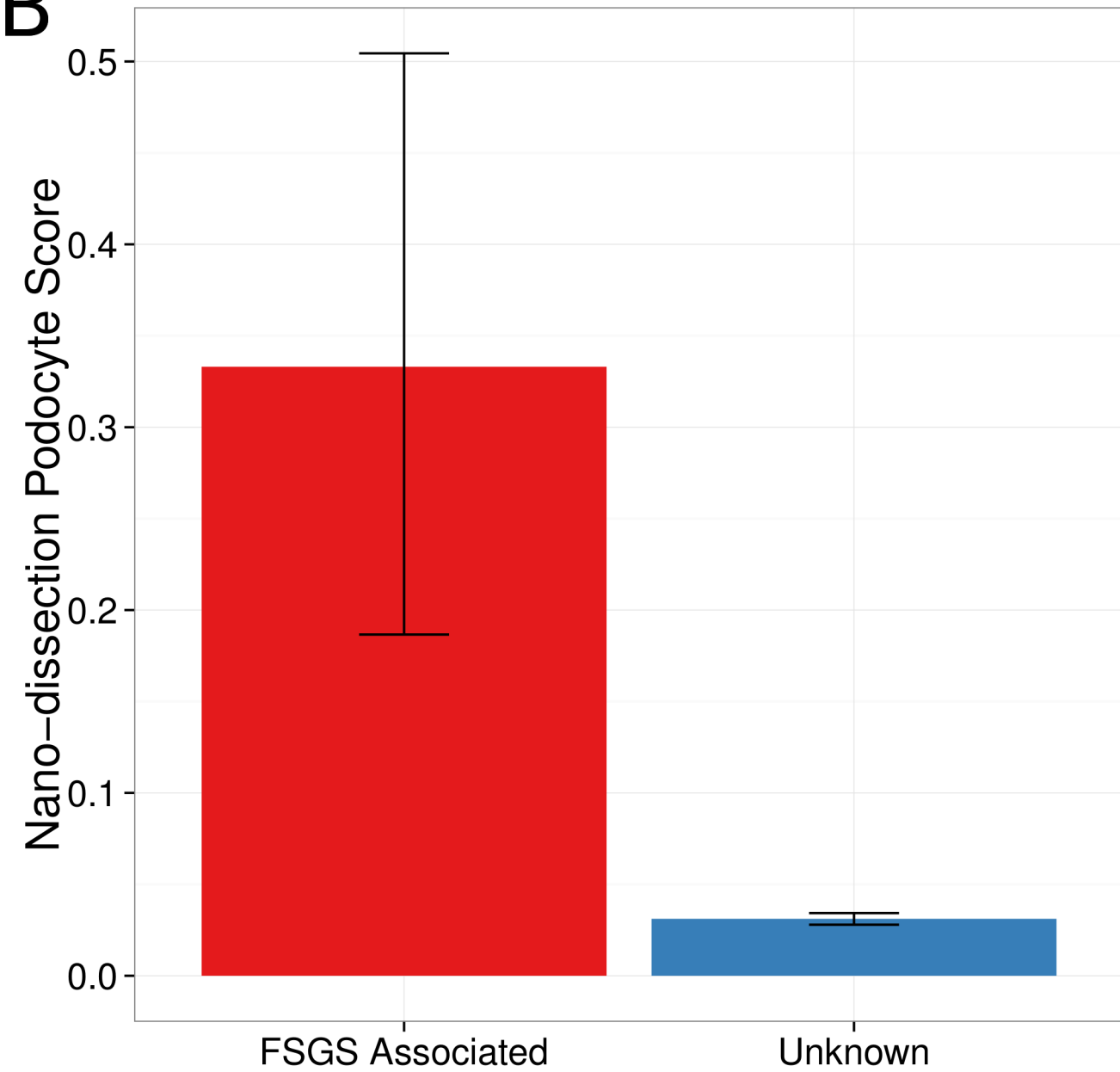






Given: m user supplied standards in ordering of increasing specificity

```
10.     set all p[k] := 1.0
11.     for k := 1 to m do begin
12.         create k_standard containing all genes from standards k
13.         d[] = results of classification with k_standard
14.         if (lineage-specific validation)
15.             for each negative standard j do begin
16.                  $sr\_j := \sum_{i=1}^{n_p} R(d_i)$ 
17.                  $C\_j := n_p n_j + \frac{n_p(n_p+1)}{2} - SR_j$ 
18.                  $U\_j := \max(C_j, n_p n_j - C_j)$ 
19.                  $Z\_j := \frac{U_j - \frac{n_j n_p}{2}}{\sqrt{\frac{n_j n_p (n_j + n_p + 1)}{12}}}$ 
20.                 p[k] := p[k] * (1 - cdf(Z_j)) // normal CDF
21.             end;
22.             p[k] :=  $\sqrt[j]{p[k]}$ 
23.         else if (compartment validation)
24.              $sr\_j := \sum_{i=1}^{n_v} R(|d_i|)$ 
25.              $C\_j := n_p n_j + \frac{n_p(n_p+1)}{2} - SR_j$ 
26.              $U\_j := \max(C_j, n_p n_j - C_j)$ 
27.              $Z\_j := \frac{U_j - \frac{n_j n_p}{2}}{\sqrt{\frac{n_j n_p (n_j + n_p + 1)}{12}}}$ 
28.             p[k] := 1 - cdf(Z_j) // normal CDF
29.         endif
30.     select the kth standard that results in the lowest p[k]
```

A**B**

Supplementary Table 1. Positive standards for podocytes selected by literature search.

Symbol	Gene Name
<i>ACTN4</i>	actinin, alpha 4
<i>NPHS1</i>	nephrosis 1, congenital, Finnish type (nephrin)
<i>NPHS2</i>	nephrosis 2, idiopathic, steroid-resistant (podocin)
<i>SYNPO</i>	synaptopodin
<i>PTPRO</i>	protein tyrosine phosphatase, receptor type, O
<i>NES</i>	nestin
<i>DDN</i>	dendrin
<i>WT1</i>	Wilms tumor 1
<i>TCF21</i>	transcription factor 21
<i>MAGI1</i>	membrane associated guanylate kinase, WW and PDZ domain containing 1
<i>MAGI2</i>	membrane associated guanylate kinase, WW and PDZ domain containing 2
<i>RAB3A</i>	RAB3A, member RAS oncogene family
<i>KIRREL</i>	kin of IRRE like (Drosophila)
<i>CD2AP</i>	CD2-associated protein
<i>CASK</i>	calcium/calmodulin-dependent serine protein kinase (MAGUK family)
<i>IQGAP1</i>	IQ motif containing GTPase activating protein 1
<i>TJP1</i>	tight junction protein 1 (zona occludens 1)
<i>TRPC6</i>	transient receptor potential cation channel, subfamily C, member 6
<i>DES</i>	desmin
<i>PDPN</i>	podoplanin
<i>PODXL</i>	podocalyxin-like
<i>LMX1B</i>	LIM homeobox transcription factor 1, beta
<i>FAT1</i>	FAT tumor suppressor homolog 1 (Drosophila)
<i>CDH3</i>	cadherin 3, type 1, P-cadherin (placental)
<i>PLCE1</i>	phospholipase C, epsilon 1
<i>CLIC5</i>	chloride intracellular channel 5
<i>CDH13</i>	cadherin 13, H-cadherin (heart)
<i>MME</i>	membrane metallo-endopeptidase
<i>UCHL1</i>	ubiquitin carboxyl-terminal esterase L1 (ubiquitin thiolesterase)
<i>CD80</i>	CD80 molecule
<i>SV2B</i>	synaptic vesicle glycoprotein 2B
<i>PALLD</i>	palladin, cytoskeletal associated protein
<i>FYN</i>	FYN oncogene related to SRC, FGR, YES
<i>PLAUR</i>	plasminogen activator, urokinase receptor
<i>DAG1</i>	dystroglycan 1 (dystrophin-associated glycoprotein 1)
<i>AGRN</i>	agrin
<i>EFNB1</i>	ephrin-B1
<i>EZR</i>	ezrin
<i>FOXC2</i>	forkhead box C2 (MFH-1, mesenchyme forkhead 1)
<i>MAFB</i>	v-maf musculoaponeurotic fibrosarcoma oncogene homolog B (avian)
<i>UTRN</i>	utrophin
<i>MYOC</i>	myocilin, trabecular meshwork inducible glucocorticoid response
<i>BASP1</i>	brain abundant, membrane attached signal protein 1
<i>SULF1</i>	sulfatase 1
<i>SCEL</i>	sciellin
<i>CX3CL1</i>	chemokine (C-X3-C motif) ligand 1

Supplementary Table 2. Negative standards for podocytes selected by literature search.

Symbol	Name
<i>ACTA2</i>	actin, alpha 2, smooth muscle, aorta
<i>ADAM10</i>	ADAM metallopeptidase domain 10
<i>AGTR1</i>	angiotensin II receptor, type 1
<i>ALDOB</i>	aldolase B, fructose-bisphosphate
<i>ALOX12</i>	arachidonate 12-lipoxygenase
<i>ANGPT2</i>	angiopoietin 2
<i>APOA4</i>	apolipoprotein A-IV
<i>AQP1</i>	aquaporin 1 (Colton blood group)
<i>AQP11</i>	aquaporin 11
<i>AQP2</i>	aquaporin 2 (collecting duct)
<i>AQP3</i>	aquaporin 3 (Gill blood group)
<i>AQP4</i>	aquaporin 4
<i>AQP6</i>	aquaporin 6, kidney specific
<i>AQP7</i>	aquaporin 7
<i>AQP8</i>	aquaporin 8
<i>BDKRB1</i>	bradykinin receptor B1
<i>BMP7</i>	bone morphogenetic protein 7
<i>CA2</i>	carbonic anhydrase II
<i>CALB1</i>	calbindin 1, 28kDa
<i>CD34</i>	CD34 molecule
<i>CDH16</i>	cadherin 16, KSP-cadherin
<i>CLCNKB</i>	chloride channel Kb
<i>CLDN16</i>	claudin 16
<i>CLDN4</i>	claudin 4
<i>CLDN5</i>	claudin 5
<i>COL6A1</i>	collagen, type VI, alpha 1
<i>CXCL16</i>	chemokine (C-X-C motif) ligand 16
<i>CYP4B1</i>	cytochrome P450, family 4, subfamily B, polypeptide 1
<i>DPYS</i>	dihydropyrimidinase
<i>EDN1</i>	endothelin 1
<i>EHD3</i>	EH-domain containing 3
<i>EMCN</i>	endomucin
<i>ENG</i>	endoglin
<i>ETS1</i>	v-ets erythroblastosis virus E26 oncogene homolog 1 (avian)
<i>EXOC3L2</i>	exocyst complex component 3-like 2
<i>F11R</i>	F11 receptor
<i>F2R</i>	coagulation factor II (thrombin) receptor
<i>FBN1</i>	fibrillin 1
<i>FLT1</i>	fms-related tyrosine kinase 1 (vascular endothelial growth factor/vascular permeability factor receptor)
<i>FOXC1</i>	forkhead box C1
<i>FXYD2</i>	In multiple Geneids
<i>GAS6</i>	growth arrest-specific 6
<i>GGT1</i>	gamma-glutamyltransferase 1
<i>GLI2</i>	GLI family zinc finger 2
<i>HAVCR1</i>	hepatitis A virus cellular receptor 1
<i>HGD</i>	homogentisate 1,2-dioxygenase
<i>HOXB7</i>	homeobox B7
<i>HPGD</i>	hydroxyprostaglandin dehydrogenase 15-(NAD)

Symbol	Name
<i>HUNK</i>	hormonally up-regulated Neu-associated kinase
<i>ICAM2</i>	intercellular adhesion molecule 2
<i>ID1</i>	inhibitor of DNA binding 1, dominant negative helix-loop-helix protein
<i>IGFBP5</i>	insulin-like growth factor binding protein 5
<i>ITGA5</i>	integrin, alpha 5 (fibronectin receptor, alpha polypeptide)
<i>ITGA8</i>	integrin, alpha 8
<i>KDR</i>	kinase insert domain receptor
<i>KHK</i>	ketoheokinase (fructokinase)
<i>KLF6</i>	Kruppel-like factor 6
<i>LDB2</i>	LIM domain binding 2
<i>LMO7</i>	LIM domain 7
<i>LRP2</i>	low density lipoprotein receptor-related protein 2
<i>LTA</i>	lymphotoxin alpha (TNF superfamily, member 1)
<i>MEIS2</i>	In multiple Geneids
<i>MEST</i>	mesoderm specific transcript homolog (mouse)
<i>MPV17L</i>	MPV17 mitochondrial membrane protein-like
<i>PAX2</i>	paired box 2
<i>PAX8</i>	paired box 8
<i>PCDH12</i>	protocadherin 12
<i>PCK2</i>	phosphoenolpyruvate carboxykinase 2 (mitochondrial)
<i>PDGFB</i>	platelet-derived growth factor beta polypeptide
<i>PDGFRB</i>	platelet-derived growth factor receptor, beta polypeptide
<i>PECAM-1</i>	platelet/endothelial cell adhesion molecule
<i>PKD1</i>	polycystic kidney disease 1 (autosomal dominant
<i>PKD2</i>	polycystic kidney disease 2 (autosomal dominant)
<i>PPAP2A</i>	phosphatidic acid phosphatase type 2A
<i>PTGS1</i>	prostaglandin-endoperoxide synthase 1 (prostaglandin G/H synthase and cyclooxygenase)
<i>SCNN1G</i>	sodium channel, nonvoltage-gated 1, gamma
<i>SERPINB7</i>	serpin peptidase inhibitor, clade B (ovalbumin), member 7
<i>SERPINE2</i>	serpin peptidase inhibitor, clade E (nexin, plasminogen activator inhibitor type 1), member 2
<i>SFRP2</i>	secreted frizzled-related protein 2
<i>SLC12A1</i>	solute carrier family 12 (sodium/potassium/chloride transporters), member 1
<i>SLC12A3</i>	solute carrier family 12 (sodium/chloride transporters), member 3
<i>SLC2A2</i>	solute carrier family 2 (facilitated glucose transporter), member 2
<i>SLC2A9</i>	solute carrier family 2 (facilitated glucose transporter), member 9
<i>SLC34A1</i>	solute carrier family 34 (sodium phosphate), member 1
<i>SLC4A4</i>	solute carrier family 4, sodium bicarbonate cotransporter, member 4
<i>SLC5A2</i>	solute carrier family 5 (sodium/glucose cotransporter), member 2
<i>SLC9A3</i>	solute carrier family 9 (sodium/hydrogen exchanger), member 3
<i>SPP1</i>	secreted phosphoprotein 1
<i>STARD5</i>	In multiple Geneids
<i>TEK</i>	TEK tyrosine kinase, endothelial
<i>THBD</i>	thrombomodulin
<i>THBS1</i>	thrombospondin 1
<i>Thy1</i>	Thy-1 cell surface antigen
<i>TNS1</i>	tensin 1
<i>TNS3</i>	tensin 3
<i>UMOD</i>	uromodulin
<i>VWF</i>	von Willebrand factor

Supplementary Table 3. Top 136 predicted genes by *in silico* nano-dissection approach.

Gene Symbol	Entrez ID	Gene Name
<i>CHI3L1</i>	1116	chitinase 3-like 1 (cartilage glycoprotein-39)
**PCOLCE2	26577	procollagen C-endopeptidase enhancer 2
**PLA2R1	22925	phospholipase A2 receptor 1, 180kDa
<i>NDNF</i>	79625	neuron-derived neurotrophic factor
**FGF1	2246	fibroblast growth factor 1 (acidic)
*CORO2B	10391	coronin, actin binding protein, 2B
<i>TPPP3</i>	51673	tubulin polymerization-promoting protein family member 3
<i>SPOCK1</i>	6695	sparc/osteonectin, cwcv and kazal-like domains proteoglycan (testican) 1
<i>CA10</i>	56934	carbonic anhydrase X
<i>DPP6</i>	1804	dipeptidyl-peptidase 6
<i>TNNI1</i>	7135	troponin I type 1 (skeletal, slow)
<i>LRRC2</i>	79442	leucine rich repeat containing 2
<i>TNNT2</i>	7139	troponin T type 2 (cardiac)
<i>ARHGAP19</i>	84986	Rho GTPase activating protein 19
<i>KLK7</i>	5650	kallikrein-related peptidase 7
<i>KLK6</i>	5653	kallikrein-related peptidase 6
<u><i>TSPAN2</i></u>	<u>10100</u>	<u>tetraspanin 2</u>
<i>TMEM45A</i>	55076	transmembrane protein 45A
<i>BMP2</i>	650	bone morphogenetic protein 2
<i>ZNF804A</i>	91752	zinc finger protein 804A
*F2R	2149	coagulation factor II (thrombin) receptor
<i>ANXA1</i>	301	annexin A1
<i>DYNC1I1</i>	1780	dynein, cytoplasmic 1, intermediate chain 1
**NFASC	23114	neurofascin
<i>TYRO3</i>	7301	TYRO3 protein tyrosine kinase
<i>USP46</i>	64854	ubiquitin specific peptidase 46
*F3	2152	coagulation factor III (thromboplastin, tissue factor)
<i>PTGDS</i>	5730	prostaglandin D2 synthase 21kDa (brain)
<i>AMIGO2</i>	347902	adhesion molecule with Ig-like domain 2
<i>MCM6</i>	4175	minichromosome maintenance complex component 6
<i>PHYHIP</i>	9796	phytanoyl-CoA 2-hydroxylase interacting protein
<i>MYL9</i>	10398	myosin, light chain 9, regulatory
<i>TNNC1</i>	7134	troponin C type 1 (slow)
<i>MTHFD2</i>	10797	methylenetetrahydrofolate dehydrogenase (NADP+ dependent) 2, methenyltetrahydrofolate cyclohydrolase
<i>ST3GAL6</i>	10402	ST3 beta-galactoside alpha-2,3-sialyltransferase 6
<i>GPRC5A</i>	9052	G protein-coupled receptor, family C, group 5, member A
*GJA1	2697	gap junction protein, alpha 1, 43kDa
<i>CDC42EP3</i>	10602	CDC42 effector protein (Rho GTPase binding) 3
*DPYSL3	1809	dihydropyrimidinase-like 3
*ZNF185	7739	zinc finger protein 185 (LIM domain)
<i>POLA2</i>	23649	polymerase (DNA directed), alpha 2 (70kD subunit)
<i>MYO1D</i>	4642	myosin ID
<i>RGS10</i>	6001	regulator of G-protein signaling 10
<i>SNCA</i>	6622	synuclein, alpha (non A4 component of amyloid precursor)
<i>SPOCK2</i>	9806	sparc/osteonectin, cwcv and kazal-like domains proteoglycan (testican) 2
<u><i>B3GALT2</i></u>	<u>8707</u>	<u>UDP-Gal:betaGlcNAc beta 1,3-galactosyltransferase, polypeptide 2</u>
<i>NFE2L3</i>	9603	nuclear factor (erythroid-derived 2)-like 3
<i>ZFPM2</i>	23414	zinc finger protein, multitype 2
*SRGAP2	23380	SLIT-ROBO Rho GTPase activating protein 2
<i>SEMA3G</i>	56920	sema domain, immunoglobulin domain (Ig), short basic domain, secreted, (semaphorin) 3G
<i>PPFIA4</i>	8497	protein tyrosine phosphatase, receptor type, f polypeptide (PTPRF), interacting protein (liprin), alpha 4
<i>RASL11B</i>	65997	RAS-like, family 11, member B
<i>ITGA3</i>	3675	integrin, alpha 3 (antigen CD49C, alpha 3 subunit of VLA-3 receptor)
<i>HBEGF</i>	1839	heparin-binding EGF-like growth factor

<i>LOXL1</i>	4016	lysyl oxidase-like 1
<i>ROBO1</i>	6091	roundabout, axon guidance receptor, homolog 1 (Drosophila)
<i>EGR3</i>	1960	early growth response 3
<i>HTRA1</i>	5654	HtrA serine peptidase 1
<i>PLTP</i>	5360	phospholipid transfer protein
<i>FRY</i>	10129	furry homolog (Drosophila)
<i>SST</i>	6750	somatostatin
<i>PPP2R2B</i>	5521	protein phosphatase 2, regulatory subunit B, beta
<i>MYOZ2</i>	51778	myozenin 2
<i><u>PDE4B</u></i>	<u>5142</u>	<u>phosphodiesterase 4B, cAMP-specific</u>
<i>PNMA2</i>	10687	paraneoplastic antigen MA2
<i>ITIH5</i>	80760	inter-alpha (globulin) inhibitor H5
<i>*SLK</i>	9748	STE20-like kinase
<i>ME1</i>	4199	malic enzyme 1, NADP(+)-dependent, cytosolic
<i><u>**ARHGAP28</u></i>	<u>79822</u>	<u>Rho GTPase activating protein 28</u>
<i>PAMR1</i>	25891	peptidase domain containing associated with muscle regeneration 1
<i>**PRKAR2B</i>	5577	protein kinase, cAMP-dependent, regulatory, type II, beta
<i>CHST2</i>	9435	carbohydrate (N-acetylglucosamine-6-O) sulfotransferase 2
<i>GNMB</i>	10457	glycoprotein (transmembrane) nmb
<i>*COLGALT2</i>	23127	collagen beta(1-O)galactosyltransferase 2
<i>MYO1E</i>	4643	myosin IE
<i>PTGER4</i>	5734	prostaglandin E receptor 4 (subtype EP4)
<i>WFS1</i>	7466	Wolfram syndrome 1 (wolframin)
<i>APOD</i>	347	apolipoprotein D
<i>FZD2</i>	2535	frizzled family receptor 2
<i>CCDC91</i>	55297	coiled-coil domain containing 91
<i><u>HS3ST3A1</u></i>	<u>9955</u>	<u>heparan sulfate (glucosamine) 3-O-sulfotransferase 3A1</u>
<i>KIAA0040</i>	9674	KIAA0040
<i>*CBLB</i>	868	Cas-Br-M (murine) ecotropic retroviral transforming sequence b
<i>*ZDHHC6</i>	64429	zinc finger, DHHC-type containing 6
<i>AIF1</i>	199	allograft inflammatory factor 1
<i>MXRA8</i>	54587	matrix-remodelling associated 8
<i>MFSD6</i>	54842	major facilitator superfamily domain containing 6
<i>H2AFJ</i>	55766	H2A histone family, member J
<i><u>CDKN1C</u></i>	<u>1028</u>	<u>cyclin-dependent kinase inhibitor 1C (p57, Kip2)</u>
<i>C1orf21</i>	81563	chromosome 1 open reading frame 21
<i>GMDS</i>	2762	GDP-mannose 4,6-dehydratase
<i>ARHGEF3</i>	50650	Rho guanine nucleotide exchange factor (GEF) 3
<i>KDELRL3</i>	11015	KDEL (Lys-Asp-Glu-Leu) endoplasmic reticulum protein retention receptor 3
<i>NELL1</i>	4745	NEL-like 1 (chicken)
<i>CACNB2</i>	783	calcium channel, voltage-dependent, beta 2 subunit
<i>RECK</i>	8434	reversion-inducing-cysteine-rich protein with kazal motifs
<i>ELOVL4</i>	6785	ELOVL fatty acid elongase 4
<i>TIMP1</i>	7076	TIMP metalloproteinase inhibitor 1
<i>KBTBD11</i>	9920	kelch repeat and BTB (POZ) domain containing 11
<i>MAFF</i>	23764	v-maf musculoaponeurotic fibrosarcoma oncogene homolog F (avian)
<i>CAND2</i>	23066	cullin-associated and neddylation-dissociated 2 (putative)
<i>S100A8</i>	6279	S100 calcium binding protein A8
<i>COL4A5</i>	1287	collagen, type IV, alpha 5
<i>CTGF</i>	1490	connective tissue growth factor
<i>IGFBP2</i>	3485	insulin-like growth factor binding protein 2, 36kDa
<i>F5</i>	2153	coagulation factor V (proaccelerin, labile factor)
<i>ATP9A</i>	10079	ATPase, class II, type 9A
<i>TUSC3</i>	7991	tumor suppressor candidate 3
<i>UGGT2</i>	55757	UDP-glucose glycoprotein glucosyltransferase 2
<i>TMOD1</i>	7111	tropomodulin 1
<i>EIF1AY</i>	9086	eukaryotic translation initiation factor 1A, Y-linked

<i>PVRL2</i>	5819	poliovirus receptor-related 2 (herpesvirus entry mediator B)
<i>SLAMF8</i>	56833	SLAM family member 8
<i>GNPTAB</i>	79158	N-acetylglucosamine-1-phosphate transferase, alpha and beta subunits
<i>HSPB8</i>	26353	heat shock 22kDa protein 8
<i>PMEPA1</i>	56937	prostate transmembrane protein, androgen induced 1
<i>TCEAL2</i>	140597	transcription elongation factor A (SII)-like 2
<i>LARGE</i>	9215	like-glycosyltransferase
<i>C1S</i>	716	complement component 1, s subcomponent
<u><i>SCHIP1</i></u>	<u>29970</u>	<u>schwannomin interacting protein 1</u>
**PDLIM2	64236	PDZ and LIM domain 2 (mystique)
<u><i>MYOF</i></u>	<u>26509</u>	<u>myoferlin</u>
<i>PLP2</i>	5355	proteolipid protein 2 (colonic epithelium-enriched)
<i>IGFBP1</i>	3484	insulin-like growth factor binding protein 1
<i>PIEZO1</i>	9780	piezo-type mechanosensitive ion channel component 1
*NXN	64359	nucleoredoxin
<i>TMCO3</i>	55002	transmembrane and coiled-coil domains 3
<i>EGR2</i>	1959	early growth response 2
<u><i>COL4A3</i></u>	<u>1285</u>	<u>collagen, type IV, alpha 3 (Goodpasture antigen)</u>
<i>HPS5</i>	11234	Hermansky-Pudlak syndrome 5
<i>RNF220</i>	55182	ring finger protein 220
<i>NDN</i>	4692	neccdin homolog (mouse)
<i>CMAHP</i>	8418	cytidine monophospho-N-acetylneuraminic acid hydroxylase, pseudogene
<i>AXL</i>	558	AXL receptor tyrosine kinase
<i>HOXA1</i>	3198	homeobox A1
*MPP5	64398	membrane protein, palmitoylated 5 (MAGUK p55 subfamily member 5)

* denotes genes specifically expressed in podocytes in glomeruli; **denotes genes exclusively expressed in podocytes in whole kidney. This evaluation is based on the IHC images provided by HPA. Genes underlined are also predicted by data analysis using mouse data downloaded from GUDMAP.

Supplementary Table 4. Genes with sufficiently specific staining in HPA to identify localization to renal cell lineages. Bold genes indicate genes that stained only to podocytes within the glomerulus, bold and underlined genes showed podocyte specific staining within the entire kidney.

Gene Symbol	Nano-dissection Score	HPA Identified Lineage	Gene Name
<u>PCOLCE2</u>	<u>1.08</u>	<u>Podocyte in Kidney</u>	<u>procollagen C-endopeptidase enhancer 2</u>
<u>PLA2R1</u>	<u>1.05</u>	<u>Podocyte in Kidney</u>	<u>phospholipase A2 receptor 1, 180kDa</u>
<u>FGF1</u>	<u>1.00</u>	<u>Podocyte in Kidney</u>	<u>fibroblast growth factor 1 (acidic)</u>
CORO2B	0.95	Podocyte in Glomerulus	coronin, actin binding protein, 2B
ZNF804A	0.825	Other Renal	zinc finger protein 804A
F2R	0.815	Podocyte in Glomerulus	coagulation factor II (thrombin) receptor
ANXA1	0.81	Other Renal	annexin A1
<u>NFASC</u>	<u>0.80</u>	<u>Podocyte in Kidney</u>	<u>neurofascin</u>
USP46	0.79	Other Renal	ubiquitin specific peptidase 46
F3	0.79	Podocyte in Glomerulus	coagulation factor III (thromboplastin, tissue factor)
PTGDS	0.78	Other Renal	prostaglandin D2 synthase 21kDa (brain)
GJA1	0.74	Podocyte in Glomerulus	gap junction protein, alpha 1, 43kDa
DPYSL3	0.73	Podocyte in Glomerulus	dihydropyrimidinase-like 3
ZNF185	0.73	Podocyte in Glomerulus	zinc finger protein 185 (LIM domain)
SRGAP2	0.70	Podocyte in Glomerulus	SLIT-ROBO Rho GTPase activating protein 2
SEMA3G	0.70	Other Renal	sema domain, immunoglobulin domain (Ig), short basic domain, secreted, (semaphorin) 3G
SLK	0.65	Podocyte in Glomerulus	STE20-like kinase
<u>ARHGAP28</u>	<u>0.65</u>	<u>Podocyte in Kidney</u>	<u>Rho GTPase activating protein 28</u>
<u>PRKAR2B</u>	<u>0.64</u>	<u>Podocyte in Kidney</u>	<u>protein kinase, cAMP-dependent, regulatory, type II, beta</u>
GPNUMB	0.63	Other Renal	glycoprotein (transmembrane) numb
COLGALT2	0.63	Podocyte in Glomerulus	collagen beta (1-O) galactosyltransferase 2
PTGER4	0.63	Other Renal	prostaglandin E receptor 4 (subtype EP4)
CBLB	0.61	Podocyte in Glomerulus	Cas-Br-M (murine) ecotropic retroviral transforming sequence b
ZDHHC6	0.61	Podocyte in Glomerulus	zinc finger, DHHC-type containing 6
MFSD6	0.61	Other Renal	major facilitator superfamily domain containing 6
S100A8	0.58	Other Renal	S100 calcium binding protein A8
PVRL2	0.57	Other Renal	poliovirus receptor-related 2 (herpesvirus entry mediator B)
LARGE	0.56	Other Renal	like-glycosyltransferase
<u>PDLIM2</u>	<u>0.55</u>	<u>Podocyte in Kidney</u>	<u>PDZ and LIM domain 2 (mystique)</u>
NXN	0.55	Podocyte in Glomerulus	nucleoredoxin
MPP5	0.53	Podocyte in Glomerulus	membrane protein, palmitoylated 5 (MAGUK p55 subfamily member 5)

Supplementary Table 5. GUDMAP gene expression datasets used in this study.

GUDMAP ID	GEO Sample ID	Sample ID	Sample Description
12336	GSM429038	Adult_MafB_GFP1	Visceral epithelium (syn: podocytes layer)
12337	GSM429039	Adult_MafB_GFP2	Visceral epithelium (syn: podocytes layer)
12338	GSM429040	Adult_MafB_GFP3	Visceral epithelium (syn: podocytes layer)
13521	GSM519446	Adult_Meis_GFP1	Glomerular mesangial
13522	GSM519447	Adult_Meis_GFP2	Glomerular mesangial
13523	GSM519448	Adult_Meis_GFP3	Glomerular mesangial
13694	GSM559844	Tie2_Glom1_GFP1	Glomerular Capillary System
13695	GSM559845	Tie2_Glom1_GFP2	Glomerular Capillary System
13696	GSM559846	Tie2_Glom1_GFP3	Glomerular Capillary System

Supplementary Table 6. 102 highly expressed genes in podocytes in comparison with the other two major cell types in mouse glomerulus. Analysis is based on array data from GUDMAP (see website).

Mus musculus gene_ID	Mus musculus gene_symbol	Homolo-Gene-ID	Homo sapiens gene_ID	Homo sapiens gene_symbol	podo vs endo (LogFC)	podo vs endo (FC)	podo vs mes (logFC)	podo vs mes (FC)
13482	<i>Dpp4</i>	3279	1803	<i>DPP4</i>	6.26	76.38	6.59	96.59
18479	<i>*Pak1</i>	1936	5058	<i>*PAK1</i>	5.75	53.87	5.98	63.01
13809	<i>*Enpep</i>	68216	2028	<i>*ENPEP</i>	6.26	76.49	5.37	41.38
15229	<i>Foxd1</i>	3290	2297	<i>FOXD1</i>	5.66	50.60	5.65	50.14
14181	<i>Fgfbp1</i>	3766	9982	<i>FGFBP1</i>	5.57	47.42	5.68	51.44
103142	<i>Rdh9</i>	74505	8608	<i>RDH16</i>	5.68	51.30	5.38	41.58
<u>15478</u>	<u><i>Hs3st3a1</i></u>	<u>88735</u>	<u>9955</u>	<u><i>HS3ST3A1</i></u>	<u>5.46</u>	<u>44.04</u>	<u>5.48</u>	<u>44.78</u>
69908	<i>Rab3b</i>	2149	5865	<i>RAB3B</i>	5.17	35.91	5.16	35.68
269181	<i>Mgat4a</i>	8153	11320	<i>MGAT4A</i>	5.06	33.47	4.96	31.23
66260	<i>Tmem54</i>	11937	113452	<i>TMEM54</i>	4.87	29.24	5.02	32.55
320910	<i>Itgb8</i>	18567	3696	<i>ITGB8</i>	4.81	27.98	4.86	29.00
71934	<i>Car13</i>	75207	377677	<i>CA13</i>	4.84	28.66	4.82	28.31
241447	<i>Cers6</i>	72228	253782	<i>CERS6</i>	4.81	27.98	4.81	28.09
17930	<i>Myom2</i>	2953	9172	<i>MYOM2</i>	4.75	26.96	4.86	29.06
20349	<i>Sema3e</i>	8247	9723	<i>SEMA3E</i>	4.72	26.38	4.74	26.68
78910	<i>Asb15</i>	43797	142685	<i>ASB15</i>	4.62	24.51	4.72	26.42
<u>12577</u>	<u><i>Cdkn1c</i></u>	<u>58</u>	<u>1028</u>	<u><i>CDKN1C</i></u>	<u>4.69</u>	<u>25.82</u>	<u>4.64</u>	<u>25.00</u>
544963	<i>**Iqgap2</i>	101543	10788	<i>**IQGAP2</i>	4.70	26.08	4.31	19.86
210530	<i>Leprel1</i>	10062	55214	<i>LEPREL1</i>	6.43	85.95	2.59	6.01
54366	<i>Cttnal1</i>	2815	8727	<i>CTNNAL1</i>	5.05	33.24	3.94	15.30
226352	<i>Epb4.1l5</i>	32492	57669	<i>EPB41L5</i>	5.02	32.42	3.66	12.66
212531	<i>Sh3bgrl2</i>	12876	83699	<i>SH3BGRL2</i>	4.34	20.31	4.26	19.19
231532	<i>Arhgap24</i>	32754	83478	<i>ARHGAP24</i>	4.86	29.05	3.73	13.26
54670	<i>Atp8b1</i>	21151	5205	<i>ATP8B1</i>	3.66	12.65	4.92	30.31
<u>268970</u>	<u><i>**Arhgap28</i></u>	<u>18264</u>	<u>79822</u>	<u><i>**ARHGAP28</i></u>	<u>4.78</u>	<u>27.38</u>	<u>3.79</u>	<u>13.83</u>
109222	<i>Rarres1</i>	2166	5918	<i>RARRES1</i>	4.24	18.85	4.32	20.03
11535	<i>Adm</i>	873	133	<i>ADM</i>	4.65	25.18	3.84	14.33
69354	<i>Slc38a4</i>	75077	55089	<i>SLC38A4</i>	4.04	16.43	4.27	19.31
<u>226101</u>	<u><i>Myof</i></u>	<u>40882</u>	<u>26509</u>	<u><i>MYOF</i></u>	<u>4.36</u>	<u>20.58</u>	<u>3.58</u>	<u>11.93</u>
72960	<i>Top1mt</i>	43082	116447	<i>TOP1MT</i>	3.72	13.15	4.19	18.23
117600	<i>Srgap1</i>	56898	57522	<i>SRGAP1</i>	3.49	11.27	4.41	21.22

68813	Dock5	57016	80005	DOCK5	4.00	16.04	3.82	14.14
214345	Lrrc1	44445	55227	LRRC1	3.82	14.11	3.74	13.35
14400	Gabrb1	20221	2560	GABRB1	3.65	12.52	3.91	15.02
70415	2610018G03Rik	84402	51765	MST4	3.66	12.63	3.83	14.25
69454	Clic3	3432	9022	CLIC3	3.67	12.76	3.80	13.94
208890	Slc26a7	13770	115111	SLC26A7	3.64	12.49	3.75	13.43
195208	Dcdc2a	9483	51473	DCDC2	3.59	12.08	3.79	13.81
26878	B3galt2	74512	8707	B3GALT2	3.76	13.53	3.61	12.21
140792	Colec12	34248	81035	COLEC12	3.58	11.99	3.72	13.13
170743	Tlr7	75060	51284	TLR7	3.68	12.85	3.55	11.70
13134	*Dach1	7288	1602	*DACH1	3.97	15.66	3.22	9.29
56318	Acpp	55552	55	ACPP	3.56	11.82	3.61	12.19
14397	Gabra4	631	2557	GABRA4	3.46	11.03	3.68	12.85
109323	C1qtnf7	12900	114905	C1QTNF7	3.56	11.81	3.52	11.48
54381	Cpq	9401	10404	CPQ	4.77	27.22	2.26	4.78
333883	Cd59b	56386	966	CD59	3.42	10.68	3.59	12.04
13197	Gadd45a	1449	1647	GADD45A	3.59	12.07	3.39	10.49
68178	*Cgnl1	41901	84952	*CGNL1	3.93	15.27	3.05	8.28
170772	Glcci1	15773	113263	GLCCI1	3.57	11.86	3.40	10.59
72607	Usp13	68372	8975	USP13	3.39	10.45	3.58	11.93
330267	*Thsd7a	46582	221981	*THSD7A	2.91	7.54	4.05	16.52
217265	Abca5	10263	23461	ABCA5	3.76	13.56	3.14	8.79
107515	Lgr4	10226	55366	LGR4	3.74	13.34	3.10	8.57
16173	Il18	1200	3606	IL18	3.15	8.89	3.44	10.83
81907	Tmem108	18688	66000	TMEM108	2.93	7.63	3.63	12.41
107895	Mgat5	1808	4249	MGAT5	3.30	9.83	3.24	9.46
231503	Tmem150c	18818	441027	TMEM150C	3.21	9.25	3.31	9.90
73094	Sgip1	13001	84251	SGIP1	3.46	11.01	3.05	8.27
30953	Schip1	32200	29970	SCHIP1	2.68	6.40	3.80	13.97
76459	Car12	20327	771	CA12	3.21	9.24	3.26	9.56
56312	Nupr1	8229	26471	NUPR1	3.83	14.20	2.63	6.21
242122	Vtcn1	11627	79679	VTCN1	2.97	7.86	3.41	10.61
68024	Hist1h2bc	113763	8970	HIST1H2BJ	2.91	7.54	3.38	10.42
70747	Tspan2	21169	10100	TSPAN2	2.60	6.05	3.62	12.32
108897	Aif1l	12863	83543	AIF1L	2.87	7.31	3.34	10.16
14420	Galc	124	2581	GALC	3.82	14.16	2.39	5.24

107351	Kank1	17706	23189	KANK1	3.62	12.29	2.59	6.03
20971	Sdc4	31121	6385	SDC4	3.40	10.58	2.77	6.84
114249	Npnt	14138	255743	NPNT	2.56	5.88	3.55	11.68
218518	Marveld2	27037	153562	MARVELD2	2.91	7.51	3.14	8.83
97114	Hist2h3c2	110949	8350	HIST1H3A	2.71	6.54	3.32	9.96
74051	Steap2	17682	261729	STEAP2	3.02	8.11	2.96	7.76
13645	Egf	1483	1950	EGF	3.20	9.20	2.73	6.63
22339	Vegfa	2534	7422	VEGFA	2.24	4.71	3.69	12.86
102545	Cmtm7	15882	112616	CMTM7	2.72	6.61	3.19	9.10
19271	Ptprj	2130	5795	PTPRJ	2.68	6.39	3.15	8.85
338365	Slc41a2	9870	84102	SLC41A2	3.02	8.11	2.76	6.75
12955	Cryab	68209	1410	CRYAB	3.43	10.76	2.35	5.08
140580	Elmo1	56685	9844	ELMO1	2.57	5.96	3.18	9.09
239083	Ccnb1ip1	18819	57820	CCNB1IP1	2.83	7.12	2.92	7.55
66848	Fuca2	119	2519	FUCA2	3.18	9.06	2.52	5.75
211949	Spsb4	25315	92369	SPSB4	2.73	6.62	2.94	7.66
54003	Nell2	4488	4753	NELL2	2.72	6.59	2.94	7.69
67451	Pkp2	3364	5318	PKP2	2.88	7.39	2.68	6.41
353169	Slc2a12	59263	154091	SLC2A12	2.71	6.53	2.85	7.22
110310	Krt7	4058	3855	KRT7	2.80	6.96	2.72	6.61
210029	Metrn1	16993	284207	METRNL	2.93	7.62	2.43	5.40
12828	Col4a3	68033	1285	COL4A3	2.75	6.71	2.58	5.98
16419	Itgb5	20511	3693	ITGB5	2.96	7.79	2.31	4.97
18578	Pde4b	1953	5142	PDE4B	2.50	5.66	2.76	6.78
100039795	Ildr2	52388	387597	ILDR2	2.54	5.83	2.70	6.50
72634	Tdrkh	4999	11022	TDRKH	2.41	5.33	2.66	6.34
244141	Nars2	57002	79731	NARS2	2.57	5.92	2.51	5.68
320685	Dctd	55618	1635	DCTD	2.45	5.47	2.62	6.13
71648	Optn	11085	10133	OPTN	2.50	5.65	2.41	5.32
12829	Col4a4	20071	1286	COL4A4	2.46	5.49	2.38	5.21
22094	Tshb	463	7252	TSHB	2.28	4.84	2.56	5.90
207965	Mettl21d	56995	79609	METTL21D	2.45	5.48	2.38	5.20
16398	Itga2	1662	3673	ITGA2	2.34	5.05	2.46	5.51
76969	Chst1	2711	8534	CHST1	2.28	4.86	2.45	5.47
11443	Chrn1	594	1140	CHRN1	2.22	4.66	2.28	4.85

Abbreviations are: Podo, podocytes (visceral epithelial cells); Endo, glomerular endothelial cells; Mes, glomerular mesangial cells. * denotes genes specifically expressed in podocytes in glomeruli; **denotes genes exclusively expressed in podocytes in whole kidney. This evaluation is based on the IHC images provided by HPA. Genes underlined are also predicted by *in silico* nano-dissection.

Supplementary Table 7. Demographics of 139 patients. Age and eGFR are presented as means $\pm$ standard deviation. eGFR was calculated using the MDRD equation.

Disease type	CKD patients (n=139)	eGFR (mL/min per 1.73 m ²)	Age (years)	Gender (male/female)
LN	30	63.7 $\pm$ 29.4	34.7 $\pm$ 13.4	7m/23f
IgAN	26	73.9 $\pm$ 37.0	36.4 $\pm$ 14.1	19m/7f
MGN	21	84.8 $\pm$ 40.1	53.7 $\pm$ 18.0	12m/9f
FSGS	17	70.9 $\pm$ 37.0	44.6 $\pm$ 15.8	9m/8f
HTN	15	40.9 $\pm$ 23.8	57.1 $\pm$ 11.8	12m/3f
DN	12	52.9 $\pm$ 30.3	54.8 $\pm$ 10.9	8m/4f
MCD	12	97.3 $\pm$ 45.7	37.4 $\pm$ 17.7	7m/5f
FSGS-MCD	6	85.7 $\pm$ 22.0	47.9 $\pm$ 15.7	2m/4f