

**Table S1.** Exploratory analyses of differential abundance according to dual analytic approaches\*.

Cellular proliferation (glomerular compartment)			Interstitial fibrosis (tubulointerstitial compartment)		
Binary	Semi-quantitative		Binary	Semi-quantitative	
SPP1	TMPRSS11D	SNAI2	LOC102724777	FGF1	LOC441204
RAP1B	STX2	LOC101929058	NPHS2	ITGA1	FRG2
SNORD42B	LOC105376917	LRRC37A	PODXL	LOC101927189	CTTNBP2
	TAB3	FN1	LOC101927972	APELA	
	PPFIA2	LRR1	LOC105374312	CLIC5	
	FBXO5	RIT2	TRV-AAC2-1	NPL	
	NRN1	ZNF876P	YPEL2	LOC730202	
	MIR635	TRAV8-6	SNORA1	TLR4	
	LOC101928131	PABPC5	ADGRF5	EXOC8	
	PRAMEF2	C2orf68	SCGB2B3P	PPIAL4A	
	MARCH8	LINC01534	IL1RL1	S1PR4	
	LINC00871	GJA5	LOC101927811		
	EPDR1	CSNK1A1P1	LOC441204		
	NTS	PPP4R4	MIR30C1		
	CTD-2350J17.1	OR2B6	RPL23AP7		
	NAP1L2	OR4B1	MIR4636		
	NPAS3	SERPINA9	IGHV7-81		
	EIF1AX-AS1	DPH3	NCF1C		
	ANP32AP1	TSPYL4	LOC101928386		
	TSC22D2	ANKRD30BP3	FRG2		
	SPP1		DDX58		
Crescent formation (glomerular compartment)			Sclerosis score (glomerular compartment)		
Binary	Semi-quantitative		Binary	Semi-quantitative	
ASF1A	TAB3	FBXO48	SCARNA9		NDUFA3
HSPA5	AKAP14	CXorf66	TP53TG1		LINC02060
TRGJP2	ARMC10	LOC283922	OR4M2		OR4M2
C19orf73	MIR4484	LGALS3			USP6NL
RASA4B	NAP1L2	LINC00871			SMCP

SNORD54	LRRC37	—	AP1S2
	A		
SNORD4A	DPH3		NEAT1
TRC-GCA4-1	SPDYE5		
LINC02297	C2orf68		

\* The lists represent all transcripts (n=102) considered in an exploratory analysis according to sub-threshold criteria of unadjusted  $p < 0.01$  and |median RNA abundance difference|  $> 0.5$  or |Spearman's  $\rho$ |  $> 0.5$  in each of the analyses performed on samples from the Toronto Cohort. The transcripts in red are interferon regulated genes (46 out of 102).

**Table S2.** Exploratory pathway analysis of describing histological lesions of activity. Endocapillary hypercellularity in (a), crescent formation in (b).

**(a) Endocapillary hypercellularity**

	Pathway name	P-value	Adjusted p-value	Z-score	Combined score
1	GRB2:SOS provides linkage to MAPK signaling for Integrins_Homo sapiens_R-HSA-354194	0.0004658	0.03447	-2.3	17.62
2	p130Cas linkage to MAPK signaling for integrins_Homo sapiens_R-HSA-372708	0.0004658	0.03447	-2.15	16.46
3	Integrin alphaIIb beta3 signaling_Homo sapiens_R-HSA-354192	0.001532	0.07556	-2.13	13.83
4	MAP2K and MAPK activation_Homo sapiens_R-HSA-5674135	0.003022	0.08945	-2.3	13.33
5	Platelet Aggregation (Plug Formation)_Homo sapiens_R-HSA-76009	0.002867	0.08945	-1.95	11.43
6	Signalling to p38 via RIT and RIN_Homo sapiens_R-HSA-187706	0.01463	0.2297	-2.43	10.25
7	Signalling to ERKs_Homo sapiens_R-HSA-187687	0.01701	0.2297	-2.44	9.96
8	Integrin cell surface interactions_Homo sapiens_R-HSA-216083	0.009137	0.2254	-1.95	9.18
9	Signaling by PDGF_Homo sapiens_R-HSA-186797	0.0432	0.2492	-2.42	7.6
10	NGF signalling via TRKA from the plasma membrane_Homo sapiens_R-HSA-187037	0.04619	0.2492	-2.37	7.3
11	Fc epsilon receptor (FCERI) signaling_Homo sapiens_R-HSA-2454202	0.05279	0.2492	-2.33	6.85
12	Signaling by Interleukins_Homo sapiens_R-HSA-449147	0.05182	0.2492	-2.26	6.68
13	Synthesis of diphthamide-EEF2_Homo sapiens_R-HSA-5358493	0.01707	0.2297	-1.54	6.29
14	Fibronectin matrix formation_Homo sapiens_R-HSA-1566977	0.01283	0.2297	-1.43	6.24
15	Gastrin-CREB signaling pathway via PKC and MAPK_Homo sapiens_R-HSA-881907	0.06548	0.2492	-2.28	6.21

<b>(b) Crescent formation</b>					
	Pathway name	P-value	Adjusted p-value	Z-score	Combined score
1	Synthesis of diphthamide-EEF2_Homo sapiens_R-HSA-5358493	0.009165	0.1221	-2.41	11.33
2	ATF6-alpha activates chaperone genes_Homo sapiens_R-HSA-381183	0.01144	0.1221	-2.09	9.36
3	IRAK2 mediated activation of TAK1 complex upon TLR7/8 or 9 stimulation_Homo sapiens_R-HSA-975163	0.01144	0.1221	-1.76	7.89
4	TRAF6 mediated induction of TAK1 complex_Homo sapiens_R-HSA-937072	0.01825	0.1221	-1.94	7.76
5	JNK (c-Jun kinases) phosphorylation and activation mediated by activated human TAK1_Homo sapiens_R-HSA-450321	0.01825	0.1221	-1.93	7.71
6	IRAK2 mediated activation of TAK1 complex_Homo sapiens_R-HSA-937042	0.01144	0.1221	-1.72	7.7
7	ATF6-alpha activates chaperones_Homo sapiens_R-HSA-381033	0.01372	0.1221	-1.8	7.7
8	Formation of Senescence-Associated Heterochromatin Foci (SAHF)_Homo sapiens_R-HSA-2559584	0.01938	0.1221	-1.93	7.59
9	TAK1 activates NFkB by phosphorylation and activation of IKKs complex_Homo sapiens_R-HSA-445989	0.02949	0.1428	-2.06	7.24
10	Advanced glycosylation endproduct receptor signaling_Homo sapiens_R-HSA-879415	0.01485	0.1221	-1.7	7.16
11	activated TAK1 mediates p38 MAPK activation_Homo sapiens_R-HSA-450302	0.01938	0.1221	-1.77	7
12	NOD1/2 Signaling Pathway_Homo sapiens_R-HSA-168638	0.03507	0.1473	-2.07	6.94
13	PERK regulates gene expression_Homo sapiens_R-HSA-381042	0.03173	0.1428	-1.96	6.78
14	Antigen Presentation: Folding, assembly and peptide loading of class I MHC_Homo sapiens_R-HSA-983170	0.02837	0.1428	-1.77	6.32
15	TNFR1-induced NFkappaB signaling pathway_Homo sapiens_R-HSA-5357956	0.02949	0.1428	-1.76	6.21

Pathway analysis of transcripts that achieved statistical significance based on a threshold of  $p < 0.01$  and  $|\text{median RNA abundance difference}| > 0.5$   $|\text{Spearman's } \rho| > 0.5$ . Analysis was done using Enrichr and Reactome 2016 database. The top 15 pathways involved in endocapillary hypercellularity (a), and crescent formation (b) are displayed. Z-score indicates the Enrichr Z score. The combined score is calculated from the p-value obtained from Fisher's exact test and the Enrichr z-score (combined score =  $-\log(p) * z$ ). Pathways are ordered based on the combined score in descending order.

**Table S3.** Exploratory pathway analysis of transcripts describing histological lesions of chronic damage. Interstitial fibrosis in (a), global and segmental sclerosis (b).

<b>(a) Interstitial fibrosis</b>					
	Pathway name	P-value	Adjusted p-value	Z-score	Combined score
1	FCERI mediated MAPK activation_Homo sapiens_R-HSA-2871796	0.02125	0.1656	-2.48	9.55
2	Axon guidance_Homo sapiens_R-HSA-422475	0.02075	0.1656	-2.3	8.93
3	Innate Immune System_Homo sapiens_R-HSA-168249	0.02382	0.1656	-2.3	8.61
4	VEGFA-VEGFR2 Pathway_Homo sapiens_R-HSA-4420097	0.02764	0.1656	-2.34	8.38
5	Signaling by VEGF_Homo sapiens_R-HSA-194138	0.02945	0.1656	-2.26	7.98
6	Downstream signal transduction_Homo sapiens_R-HSA-186763	0.0325	0.1656	-2.32	7.94
7	CHL1 interactions_Homo sapiens_R-HSA-447041	0.0183	0.1656	-1.92	7.69
8	Signaling by PDGF_Homo sapiens_R-HSA-186797	0.03831	0.1656	-2.23	7.27
9	MAPK1 (ERK2) activation_Homo sapiens_R-HSA-112411	0.0183	0.1656	-1.71	6.84
10	Cytokine Signaling in Immune system_Homo sapiens_R-HSA-1280215	0.03748	0.1656	-2.07	6.79
11	MAPK3 (ERK1) activation_Homo sapiens_R-HSA-110056	0.02032	0.1656	-1.72	6.72
12	FGFR1b ligand binding and activation_Homo sapiens_R-HSA-190370	0.01224	0.1656	-1.52	6.71
13	Interleukin-6 signaling_Homo sapiens_R-HSA-1059683	0.02233	0.1656	-1.66	6.33
14	Fc epsilon receptor (FCERI) signaling_Homo sapiens_R-HSA-2454202	0.04692	0.1656	-2.06	6.3
15	Signaling by Interleukins_Homo sapiens_R-HSA-449147	0.04604	0.1656	-2.03	6.26
<b>(b) Global and segmental sclerosis</b>					
	Pathway name	P-value	Adjusted p-value	Z-score	Combined score
1	Nef mediated downregulation of MHC class I complex cell surface expression_Homo sapiens_R-HSA-164940	0.004492	0.07213	-2.12	11.44
2	Nef-mediates down modulation of cell surface receptors by recruiting them to clathrin adapters_Homo sapiens_R-HSA-164938	0.009412	0.07213	-2.18	10.15
3	The role of Nef in HIV-1 replication and disease pathogenesis_Homo sapiens_R-HSA-164952	0.01253	0.07213	-2.31	10.13
4	Membrane Trafficking_Homo sapiens_R-HSA-199991	0.01436	0.07213	-2.12	9.01
5	Lysosome Vesicle Biogenesis_Homo sapiens_R-HSA-432720	0.01564	0.07213	-2.06	8.54
6	Vesicle-mediated transport_Homo sapiens_R-HSA-5653656	0.01939	0.07213	-2.12	8.38
7	Complex I biogenesis_Homo sapiens_R-HSA-6799198	0.02184	0.07213	-2	7.64
8	Golgi Associated Vesicle Biogenesis_Homo sapiens_R-HSA-432722	0.02404	0.07213	-1.9	7.07
9	Retrograde transport at the Trans-Golgi-Network_Homo sapiens_R-HSA-6811440	0.02184	0.07213	-1.78	6.82
10	Clathrin derived vesicle budding_Homo sapiens_R-HSA-421837	0.03107	0.07626	-1.78	6.19
11	Host Interactions of HIV factors_Homo sapiens_R-HSA-162909	0.05659	0.1019	-2.15	6.19
12	trans-Golgi Network Vesicle Budding_Homo sapiens_R-HSA-199992	0.03107	0.07626	-1.77	6.16
13	Respiratory electron transport_Homo sapiens_R-HSA-611105	0.03892	0.08757	-1.88	6.1
14	Respiratory electron transport, ATP synthesis by chemiosmotic coupling, and heat production by uncoupling proteins_Homo sapiens_R-HSA-163200	0.048	0.09258	-1.89	5.73
15	MHC class II antigen presentation_Homo sapiens_R-HSA-2132295	0.04542	0.09258	-1.84	5.7
Pathway analysis of transcripts that achieved statistical significance based on a threshold of $p < 0.01$ and $ \text{median RNA abundance difference}  > 0.5$ $ \text{Spearman's } \rho  > 0.5$ . Analysis was done using Enrichr and Reactome 2016 database. The top 15 pathways involved in interstitial fibrosis (a), and global and segmental sclerosis (b) are displayed. Z-score indicates the Enrichr Z score. The combined score is calculated from the p-value obtained from Fisher's exact test and the Enrichr z-score (combined score = $\log(p) * z$ ). Pathways are ordered based on the combined score in descending order.					

**Table S4.** Clinical profiles of patients from the Longitudinal Cohort. Values displayed as mean  $\pm$ SD or median (IQR) as appropriate.

	CR (n=27)	NR (n=9)	P value
Age (years)	30.9 $\pm$ 9	31.4 $\pm$ 10	NS
Female (%)	24 (89)	4 (55)	0.02
Class IV (%)	17 (63)	5 (56)	NS
MMF Induction (%)	14 (52)	3 (33)	NS
Serum creatinine ( $\mu$ mol/L)			
at Bx <sub>1</sub>	61.9 $\pm$ 17.7	114.9 $\pm$ 88.4	0.004
at Bx <sub>2</sub>	61.9 $\pm$ 8.8	88.4 $\pm$ 39.8	0.003
Proteinuria (g/d)			
at Bx <sub>1</sub>	3.0 $\pm$ 1.6	3.1 $\pm$ 3.0	NS
at Bx <sub>2</sub>	0.2 $\pm$ 0.1	2.2 $\pm$ 1.7	<0.0001
Median time to Follow up (months)	8 (5-15)	7 (6-13)	NS
NIH indices:			
Activity Index (max 24)			
Bx <sub>1</sub>	8 (2-15)	7 (4-12)	NS
Bx <sub>2</sub>	3.5 (0-6)	3 (0-9)	NS
Chronicity Index (max 12)			
Bx <sub>1</sub>	3 (0-6)	3 (0-6)	NS
Bx <sub>2</sub>	4 (2-7)	4 (1-8)	NS