# ■ SUPPLEMENT ■

# Complete Model Equations and Parameters

for

Sex-Specific Computational Models for Blood Pressure Regulation in the Rat

#### Cardiovascular Function

$$P_{mf} = \left(\frac{1}{SF_V} \times 7.436 \times V_b - 30.18\right) \times \epsilon_{aum} \tag{S1}$$

$$P_{ra} = \max \left\{ 0.2787 e^{SF_R \times 0.2281\Phi_{co}} - 0.9119, 0 \right\}$$
 (S2)

$$P_{ma} = \Phi_{co} \times R_{tp} \tag{S3}$$

$$\Phi_{vr} = \frac{P_{mf} - P_{ra}}{R_{vr}} \tag{S4}$$

$$\Phi_{co} = \Phi_{vr} \tag{S5}$$

$$\frac{dvas}{dt} = vas_f - vas_d \tag{S6}$$

$$vas_f = \frac{11.312 \times e^{-\Phi_{co} \times (SF_R \times 0.4714)}}{100000}$$
(S7)

$$vas_d = vas \times K_{vd} \tag{S8}$$

$$R_a = R_{ba} \times \epsilon_{aum} \tag{S9}$$

$$R_{ba} = K_{bar}/vas (S10)$$

$$R_{vr} = (8R_{bv} + R_a)/31 (S11)$$

$$R_{tp} = R_a + R_{bv} \tag{S12}$$

$$\varepsilon_{aum} = \frac{4}{5}(a_{chemo} + a_{baro}) \tag{S13}$$

$$a_{chemo} = \frac{1}{4} a_{auto} \tag{S14}$$

$$\frac{da_{baro}}{dt} = \frac{3}{4} \left\{ \frac{da_{auto}}{dt} - 0.0000667(a_{baro} - 1) \right\}$$
 (S15)

$$a_{auto} = 3.0042e^{-0.0107P_{ma}} (S16)$$

# Renal Hemodynamics

$$\Phi_{rb} = P_{ma}/R_r \tag{S17}$$

$$\Phi_{gfilt} = P_f \times C_{gcf} \tag{S18}$$

$$P_f = P_{gh} - (P_B + P_{go}) (S19)$$

$$P_{gh} = P_{ma} - \Phi_{rb} \times R_{AA} \tag{S20}$$

$$R_r = R_{AA} + R_{EA} \tag{S21}$$

$$R_{AA} = R_{aa-ss} \times \beta_{rsna} \times \Sigma_{tqf} \times \Sigma_{myo} \times \Psi_{AT1R-AA} \times \Psi_{AT2R-AA}$$
 (S22)

$$R_{EA} = R_{ea-ss} \times \Psi_{AT1R-EA} \times \Psi_{AT2R-EA} \tag{S23}$$

$$\beta_{rsna} = \frac{2}{1 + e^{-3.16(rsna - 1)}} \tag{S24}$$

$$\Sigma_{tgf} = \begin{cases} 0.3408 + \frac{3.449}{3.88 + e^{(\Phi_{md-sod} - SF_S \times 3.891)/(SF_S \times -0.9617)}} & \text{in males} \\ 0.3408 + \frac{3.449}{3.88 + e^{(\Phi_{md-sod} - SF_S \times 6.591)/(SF_S \times -0.9617)}} & \text{in females} \end{cases}$$
(S25)

$$\Sigma_{myo} = 0.75 + \frac{1.2}{1 + 3.8e^{-0.6(P_{gh} - 63.8)}}$$
 (S26)

$$\Psi_{AT1R-AA} = 0.8 + 0.2092 \times \frac{[AT1R]}{[AT1R]_{eq}} - 0.0092 \div \frac{[AT1R]}{[AT1R]_{eq}}$$
 (S27)

$$\Psi_{AT1R-EA} = 0.925 + 0.0835 \frac{[AT1R]}{[AT1R]_{eq}} - 0.0085 \div \frac{[AT1R]}{[AT1R]_{eq}}$$
 (S28)

$$\Psi_{AT2R-AA} = \begin{cases} 0.9 + 0.1e^{-\frac{[AT2R]}{[AT2R]_{eq}} - 1} & \text{in females} \\ 1 & \text{in males} \end{cases}$$

$$(S29)$$

$$\Psi_{AT2R-EA} = \begin{cases} 0.9 + 0.1e^{-\frac{[AT2R]}{[AT2R]_{eq}} - 1} & \text{in females} \\ 1 & \text{in males} \end{cases}$$
 (S30)

#### Renal Function

$$\Phi_{filsod} = \Phi_{afilt} \times C_{sod} \tag{S31}$$

$$\Phi_{pt-sodreab} = \Phi_{filsod} \times \eta_{pt-sodreab} \tag{S32}$$

$$\Phi_{md-sod} = \Phi_{filsod} - \Phi_{nt-sodreab} \tag{S33}$$

$$\Phi_{dt-sodreab} = \Phi_{md-sod} \times \eta_{dt-sodreab} \tag{S34}$$

$$\Phi_{dt-sod} = \Phi_{md-sod} - \Phi_{dt-sodreab} \tag{S35}$$

$$\Phi_{cd-sodreab} = \Phi_{dt-sod} \times \eta_{cd-sodreab} \tag{S36}$$

$$\Phi_{u-sod} = \Phi_{dt-sod} - \Phi_{cd-sodreab} \tag{S37}$$

$$\eta_{pt-sodreab} = \eta_{pt-sodreab}^{eq} \times \gamma_{filsod} \times \gamma_{AT1R} \times \gamma_{rsna}$$
 (S38)

$$\eta_{dt-sodreab} = \eta_{dt-sodreab}^{eq} \times \psi_{al} \tag{S39}$$

$$\eta_{cd-sodreab} = \eta_{cd-sodreab}^{eq} \times \lambda_{dt} \times \lambda_{anp} \times \lambda_{al}$$
 (S40)

$$\eta_{cd-sodreab} = \eta_{cd-sodreab}^{eq} \times \lambda_{dt} \times \lambda_{anp} \times \lambda_{al}$$

$$\gamma_{filsod} = \begin{cases}
0.8 + \frac{0.3}{1 + e^{1 + [(\Phi_{filsod} - SF_S \times 113.7)/(SF_S \times 138)]}} & \text{in males} \\
0.8 + \frac{0.3}{1 + e^{1 + [(\Phi_{filsod} - SF_S \times 108.3)/(SF_S \times 138)]}} & \text{in females}
\end{cases}$$
(S41)

$$\gamma_{AT1R} = 0.92 + \frac{0.136}{1 + e^{-1.7983(\frac{[AT1R]}{[AT1R]eq} - 0.8017)}}$$

$$\gamma_{rsna} = 0.72 + \frac{0.56}{1 + e^{(1-rsna)/2.18}}$$
(S43)

$$\gamma_{rsna} = 0.72 + \frac{0.56}{1 + e^{(1 - rsna)/2.18}} \tag{S43}$$

$$\psi_{al} = \frac{11.55}{1 + 0.1e^{-0.007937C_{al}}} - 10.5 \tag{S44}$$

$$\lambda_{dt} = \begin{cases} 0.8 + \frac{0.275}{1 + e^{\frac{1}{SF_S} \times 2.31(\Phi_{dt-sod} - SF_S \times 2.224)}} & \text{in males} \\ 0.8 + \frac{0.2417}{1 + e^{\frac{1}{SF_S} \times 2.86(\Phi_{dt-sod} - SF_S \times 3.699)}} & \text{in females} \end{cases}$$
(S45)

$$\lambda_{anp} = -0.1 \times \hat{C}_{anp} + 1.1 \tag{S46}$$

$$\lambda_{al} = \frac{1}{C_{al}^{eq0.06}} C_{al}^{0.06} \tag{S47}$$

$$\Phi_{pt-wreab} = \Phi_{gfilt} \times \eta_{pt-wreab} \tag{S48}$$

$$\Phi_{md-u} = \Phi_{qfilt} - \Phi_{pt-wreab} \tag{S49}$$

$$\Phi_{dt-wreab} = \Phi_{md-u} \times \eta_{dt-wreab} \tag{S50}$$

$$\Phi_{dt-u} = \Phi_{md-u} - \Phi_{dt-wreab} \tag{S51}$$

$$\Phi_{cd-wreab} = \Phi_{dt-u} \times \eta_{cd-wreab} \tag{S52}$$

$$\Phi_u = \Phi_{dt-u} - \Phi_{cd-wreab} \tag{S53}$$

$$\eta_{pt-wreab} = \eta_{pt-wreab}^{eq} \times \mu_{pt-sodreab} \tag{S54}$$

$$\eta_{dt-wreab} = \eta_{dt-wreab}^{eq} \times \mu_{dt-sodreab} \tag{S55}$$

$$\eta_{cd-wreab} = \eta_{cd-wreab}^{eq} \times \mu_{cd-sodreab} \times \mu_{adh}$$
 (S56)

$$\mu_{pt-sodreab} = 0.12 \tanh \left( 10 \left( \frac{\eta_{pt-sodreab}}{\eta_{pt-sodreab}^{eq}} - 1 \right) \right) + 1 \tag{S57}$$

$$\mu_{dt-sodreab} = 0.12 \tanh \left( 10 \left( \frac{\eta_{dt-sodreab}}{\eta_{dt-sodreab}^{eq}} - 1 \right) \right) + 1$$

$$\mu_{cd-sodreab} = 0.12 \tanh \left( 10 \left( \frac{\eta_{cd-sodreab}}{\eta_{cd-sodreab}^{eq}} - 1 \right) \right) + 1$$

$$(S58)$$

$$\mu_{cd-sodreab} = 0.12 \tanh \left( 10 \left( \frac{\eta_{cd-sodreab}}{\eta_{cd-sodreab}^{eq}} - 1 \right) \right) + 1 \tag{S59}$$

$$\mu_{adh} = 1.0328 - 0.1938e^{-0.4441C_{adh}} \tag{S60}$$

# Renin-Angiotensin-Aldosterone System

$$R_{sec} = N_{rs} \times \nu_{md-sod} \times \nu_{rsna} \times \nu_{AT1R} \tag{S61}$$

$$\frac{d[PRC]}{dt} = R_{sec} - \frac{\ln(2)}{h_{renin}}[PRC] \tag{S62}$$

$$PRA = [PRC] \times X_{PRC-PRA} \tag{S63}$$

$$\frac{d[AGT]}{dt} = k_{AGT} - PRA - \frac{\ln(2)}{h_{AGT}}[AGT] \tag{S64}$$

$$\frac{d[AngI]}{dt} = PRA - (c_{ACE} + c_{Chym} + c_{NEP})[AngI] - \frac{\ln(2)}{h_{AngI}}[AngI]$$

$$(S65)$$

$$\frac{d[AngII]}{dt} = (c_{ACE} + c_{Chym})[AngI] - (c_{ACE2} + c_{AII=AIV} + c_{AT1R} + c_{AT2R})[AngII] - \frac{\ln(2)}{h_{AngII}}[AngII]$$
(S66)

$$\frac{d[Ang(1-7)]}{dt} = c_{NEP}[AngI] + c_{ACE2}[AngII] - \frac{\ln(2)}{h_{Ang(1-7)}}[Ang(1-7)]$$
(S67)

$$\frac{d[AngIV]}{dt} = c_{AII=AIV}[AngII] - \frac{\ln(2)}{h_{AngIV}}[AngIV]$$
 (S68)

$$\frac{d[AT1R]}{dt} = c_{AT1R}[AngII] - \frac{\ln(2)}{h_{AT1R}}[AT1R]$$
 (S69)

$$\frac{d[AT2R]}{dt} = c_{AT2R}[AngII] - \frac{\ln(2)}{h_{AT2R}}[AT2R]$$
 (S70)

$$\nu_{md-sod} = \begin{cases} 0.2262 + \frac{28.04}{11.56 + e^{(\Phi_{md-sod} - SF_S \times 1.659)/(SF_S \times 0.6056)}} & \text{in males} \\ 0.2262 + \frac{28.04}{11.56 + e^{(\Phi_{md-sod} - SF_S \times 4.359)/(SF_S \times 0.6056)}} & \text{in females} \end{cases}$$
(S71)

$$\nu_{rsna} = 1.822 - \frac{2.056}{1.358 + e^{(rsna - 0.8662)}} \tag{S72}$$

$$\nu_{AT1R} = \left(\frac{[AT1R]}{[AT1R]_{eq}}\right)^{-0.95} \tag{S73}$$

$$N_{als} = N_{als}^{eq} \times \xi_{k/sod} \times \xi_{map} \times \xi_{AT1R} \tag{S74}$$

$$\frac{dN_{al}}{dt} = \frac{1}{T_{al}}(N_{als} - N_{al}) \tag{S75}$$

$$C_{al} = \begin{cases} 379N_{al} & \text{in females} \\ 395N_{al} & \text{in males} \end{cases}$$
 (S76)

$$\xi_{k/sod} = \frac{5}{1 + e^{0.265(C_k/C_{sod} - 23.7)}} \tag{S77}$$

$$\xi_{map} = \begin{cases} 70.1054e^{-(0.0425) \times P_{ma}} & \text{if } P_{ma} \le 100\\ 1 & \text{if } P_{ma} > 100 \end{cases}$$

$$\xi_{AT1R} = 0.1 + \frac{2.9}{1 + e^{-2\left(\frac{[AT1R]}{[AT1R]eq} - 1.399\right)}}$$

$$(S78)$$

$$\xi_{AT1R} = 0.1 + \frac{2.9}{1 + e^{-2\left(\frac{[AT1R]}{[AT1R]eq} - 1.399\right)}}$$
(S79)

# Miscellaneous

 $rsna_0 = N_{rsna} \times \alpha_{map} \times \alpha_{rap}$ 

$$rsna = \begin{cases} rsna_0^{\frac{1}{rsna_0}} & \text{in females} \\ rsna_0 & \text{in males} \end{cases}$$
 (S80)

$$\alpha_{map} = 0.5 + \frac{1}{1 + e^{(P_{ma} - 103)/15}} \tag{S81}$$

$$\alpha_{rap} = 1 - 0.008P_{ra} \tag{S82}$$

$$\Phi_{win} = \frac{SF_U \times 0.002313}{1 + e^{-0.8(C_{adh} - 4.3404)}}$$
 (S83)

$$\frac{dV_{ecf}}{dt} = \Phi_{win} - \Phi_u \tag{S84}$$

$$V_b = SF_V \times 4.5479 + \frac{SF_V \times 2.4312}{1 + e^{-(V_{ecf} - SF_V \times 18.1128) \times (\frac{1}{SF_V} \times 0.4744)}}$$
 (S85)

$$N_{adhs} = \begin{cases} \left[ \max\{ (C_{sod} - 140), 0 \} + \max\{ (\epsilon_{aum} - 1), 0 \} - \delta_{ra} \right] / 3 & \text{in males} \\ \left[ \max\{ (C_{sod} - 143), 0 \} + \max\{ (\epsilon_{aum} - 1), 0 \} - \delta_{ra} \right] / 3 & \text{in females} \end{cases}$$
(S86)

$$\frac{dN_{adh}}{dt} = \frac{1}{T_{adh}}(N_{adhs} - N_{adh}) \tag{S87}$$

$$C_{adh} = 4N_{adh} (S88)$$

$$\frac{d\delta_{ra}}{dt} = 0.2 \frac{dP_{ra}}{dt} - 0.0007\delta_{ra} \tag{S89}$$

$$\frac{dM_{sod}}{dt} = \Phi_{sodin} - \Phi_{u-sod} \tag{S90}$$

$$C_{sod} = \frac{M_{sod}}{V_{ecf}} \tag{S91}$$

$$\hat{C}_{anp} = 7.4052 - \frac{6.554}{1 + e^{(P_{ra} - 3.762)/(1)}}$$
(S92)

### Scaling Factors

Rat Value = Human Value  $\times SF$ 

Table S1: Baseline values of all model variables and parameters for male (M) and female (F). ADH, antidiuretic hormone; ALD, aldosterone; ANP, atrial natriuretic peptide; CD, collecting duct; DT, distal tubule; MAP, mean arterial pressure; PT, proximal tubule; RSNA, renal sympathetic nerve activity.

Symbol	Description	Units	Baseline M Value	Baseline F Value
	Cardiovascular Function		wi value	r value
D .	Mean filling pressure	mmHg	7 20	7.26
$P_{mf}$ $P_{ra}$	Right atrial pressure	mmHg		0
$P_{ma}$	Mean arterial pressure	mmHg		103
$\Phi_{vr}$	Venous return	ml	15.0	9.57
$\Phi_{co}$	Cardiac output	$rac{min}{ml}$	15.0	9.57
vas	Vascularity	min	1	1
$vas_f$	Vascularity formation rate	_	$1 \times 10^{-5}$	$1 \times 10^{-5}$
$vas_d$	Vascularity destruction rate	_	$1 \times 10^{-5}$	$1 \times 10^{-5}$
$K_{vd}$	Vascularity destruction coefficient	_	$1 \times 10^{-5}$	$1 \times 10^{-5}$
$K_{bar}$	Coefficient relating basic arterial resistance to vascularity	$\frac{mmHg}{ml/min}$	5.69	8.91
$R_a$	Arterial resistance	mmHg	5.69	8.91
$R_{ba}$	Basic arterial resistance	$\frac{ml/min}{mmHg}$	5.69	8.91
$R_{vr}$	Resistance to venous return	$\frac{ml/min}{mmHg}$	0.485	0.759
	Basic venous resistance	$ml/min \ mmHg$	1.17	1.83
$R_{bv}$		$\frac{\overline{ml/min}}{mmHg}$		
$R_{tp}$	Total peripheral resistance	$\frac{ml/min}{ml}$	6.86	10.7
$\varepsilon_{aum}$	Autonomic multiplier effect	_	1	1
$a_{chemo}$	Chemoreceptor activity	_	0.25	0.25
$a_{baro}$	Baroreceptor activity	_	1	1 1
$a_{auto}$	Autonomous system activity		1	1
-	Renal Hemodynamics	ml	2.00	2.22
$\Phi_{rb}$	Renal blood flow	$\frac{min}{min}$	3.60	2.30
$\Phi_{gfilt}$	Glomerular filtration rate	$\frac{\overline{min}}{ml/min}$	1.22	0.833
$C_{gcf}$	Glomerular capillary filtration coefficient	mmHg	0.068	0.047
$P_f$	Net filtration pressure	mmHg		17.7
$P_{gh}$	Glomerular hydrostatic pressure	mmHg	63.9	63.7
$P_B$	Bowman hydrostatic pressure	$mmHg \\ mmHg$	18 28	18 28
$P_{go}$	Glomerular osmotic pressure Renal vascular resistance	mmHg	28.6	44.8
$R_r$		$\frac{ml/min}{mmHg}$		
$R_{AA}$	Afferent arteriolar resistance	$\frac{ml/min}{mmHg}$	10.9	17.0
$R_{EA}$	Efferent arteriolar resistance	$\frac{mmng}{ml/min}$	17.7	27.8
$\beta_{rsna}$	Effect of RSNA on afferent arteriolar resistance	_	1	1
$\Sigma_{tgf}$	Tubuloglomerular feedback signal	_	1	1
$\Sigma_{myo}$	Myogenic response	_	1	1
$\Psi_{AT1R-AA}$	Effect of AT1R-bound Ang II on afferent resistance	_	1	1
$\Psi_{AT1R-EA}$	Effect of AT1R-bound Ang II on efferent resistance Effect of AT2R-bound Ang II on afferent resistance	_	1	1 1
	Effect of AT2R-bound Ang II on afferent resistance  Effect of AT2R-bound Ang II on efferent resistance	_	1	1
+ A12K-EA	Renal Function			1
Φ	Filtered Na <sup>+</sup> load	$\mu eq$	174	122
$\Phi_{filsod}$ $\Phi_{pt-sodreab}$	Proximal tubule Na <sup>+</sup> reabsorption rate	$_{\mu eq}^{min}$	140	61.1
$\Phi_{pt-sodreab}$ $\Phi_{md-sod}$	Macula densa Na <sup>+</sup> flow	$_{\mu eq}^{min}$	34.9	61.1
$\Phi_{md-sod}$ $\Phi_{dt-sodreab}$	Distal tubule Na <sup>+</sup> reabsorption rate	$_{\mu eq}^{min}$	17.4	30.5
$\Phi_{dt-sod}$	Distal tubule Na <sup>+</sup> flow	$\frac{min}{\mu eq}$	17.4	30.5
$\Phi_{cd-sodreab}$	Collecting duct Na <sup>+</sup> reabsorption rate	$\frac{min}{\mu eq}$	16.2	29.3
$\Phi_{u-sod}$	Urine Na <sup>+</sup> flow	$\frac{min}{\mu eq}$	1.22	1.22
$\eta_{pt-sodreab}$	Fractional proximal tubule Na <sup>+</sup> reabsorption	$\stackrel{min}{-}$	0.80	0.50
$\eta_{dt-sodreab}$	Fractional distal tubule Na <sup>+</sup> reabsorption	_	0.50	0.50
$\eta_{cd-sodreab}$	Fractional collecting duct Na <sup>+</sup> reabsorption	_	0.93	0.96
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$\gamma_{filsod}$	Effect of the filtered Na <sup>+</sup> load on fractional PT Na <sup>+</sup> reabsorption	_	1	1
$\gamma_{AT1R}$	Effect of AT2R-bound Ang II on fractional PT Na <sup>+</sup> reabsorption	_	1	1
$\gamma_{rsna}$	Effect of RSNA on fractional PT Na <sup>+</sup> reabsorption	_	1	1
$\psi_{al}$	Effect of ALD on fractional DT Na <sup>+</sup> reabsorption	_	1	1
$\lambda_{dt}$	Effect of DT Na <sup>+</sup> outflow on fractional CD Na <sup>+</sup> reabsorption	_	1	1
$\lambda_{anp}$	Effect of ANP on fractional CD Na <sup>+</sup> reabsorption	_	1	1
$\lambda_{al}$	Effect of ALD on fractional CD Na <sup>+</sup> reabsorption	— m1	1	1
$\Phi_{pt-wreab}$	Proximal tubule water reabsorption rate	$rac{ml}{min} \ ml$	1.05	0.417
$\Phi_{md-u}$	Macula densa ultrafiltrate flow	$\frac{ml}{min} \\ ml$	0.171	0.417
$\Phi_{dt-wreab}$	Distal tubule water reabsorption rate	$rac{ml}{min} \ \frac{ml}{ml}$	0.102	0.250
$\Phi_{dt-u}$	Distal tubule ultrafiltrate flow	$rac{ml}{min} \ ml$	0.0682	0.168
$\Phi_{cd-wreab}$	Collecting duct water reabsorption rate	$rac{ml}{min} \ ml$	0.0532	0.152
$\Phi_u$	Urine flow	$\frac{ml}{min}$	0.0150	0.0150
$\eta_{pt-wreab}$	Fractional proximal tubule water reabsorption	_	0.86	0.50
$\eta_{dt-wreab}$	Fractional distal tubule water reabsorption	_	0.60	0.60
$\eta_{cd-wreab}$	Fractional collecting duct water reabsorption	_	0.78	0.91
$\mu_{pt-sodreab}$	Effect of osmotic gradient in PT	_	1	1
$\mu_{dt-sodreab}$	Effect of osmotic gradient in DT	_	1	1
$\mu_{cd-sodreab}$	Effect of osmotic gradient in CD	_	1	1
$\mu_{adh}$	Effect of ADH on CD fractional water reabsorption	_	1	1
	Renin-Angiotensin-Aldosterone System			
$R_{sec}$	Normalized renin secretion rate	_	1	1
[PRC]	Plasma renin concentration	$\underline{fmol}$	17.3	17.3
PRA	Plasma renin activity	$_{fmol}^{ml\cdot min}$	136	114
[AGT]	Angiotensinogen concentration	$_{fmol}^{ml\cdot min}$	$5.76 \times 10^5$	$5.76 \times 10^5$
		$\frac{\overline{ml}}{fmol}$	90	75
[AngI]	Angiotensin I concentration	$\frac{\overline{ml}}{fmol}$		
[AngII]	Angiotensin II concentration	$\frac{\overline{ml}}{fmol}$	6.0	6.0
[Ang(1-7)]	Angiotensin (1-7) concentration	$\frac{fml}{fmol}$	50	25
[AngIV]	Angiotensin IV concentration	$\frac{fmol}{fmol}$	1.29	1.29
[AT1R]	AT1R-bound Angiotensin II concentration	ml	20	20
[AT2R]	AT2R-bound Angiotensin II concentration	$\frac{fmol}{ml}$	6.8	6.8
$X_{PRC-PRA}$	Ratio of PRA to [PRC]	-,	7.83	6.60
$k_{AGT}$	AGT production rate	$\frac{fmol}{\substack{ml \cdot min \\ 1}}$	801	780
$c_{ACE}$	Reaction rate of Ang converting enzyme	$\frac{1}{min}$	0.0968	0.116
$c_{Chym}$	Reaction rate of chymase	$\frac{1}{min}$	0.0108	0.0128
$c_{NEP}$	Reaction rate of neutral endopeptidase	$\frac{1}{min}$	0.0127	0.00767
$c_{ACE2}$	Reaction rate of Ang converting enzyme 2	$\frac{1}{min}$	$2.67 \times 10^{-3}$	$4.33 \times 10^{-4}$
$c_{AII=AIV}$	Reaction rate of conversion of Ang II to Ang IV	$\frac{1}{min}$	0.298	0.298
$c_{AT1R}$	Reaction rate of binding of Ang II to AT1R	$\frac{1}{m_i i n}$	0.197	0.197
$c_{AT2R}$	Reaction rate of binding of Ang II to AT2R	$\frac{1}{min}$	0.0657	0.0657
$h_{renin}$	Half life of renin	min	12	12
$h_{AGT}$	Half life of AGT	min	600	600
$h_{AngI}$	Half life of Ang I	min	0.5	0.5
$h_{AngII}$	Half life of Ang II	min	0.66	0.66
$h_{Ang(1-7)}$	Half life of Ang (1-7)	min	30	30
$h_{AngIV}$	Half life of Ang IV	min	0.5	0.5
$h_{AT1R}$	Half life of AT1R-bound Ang II	min	12	12
$h_{AT2R}$	Half life of AT2R-bound Ang II	min	12	12
$\nu_{md-sod}$	Effect of macula densa Na <sup>+</sup> flow on renin secretion rate	_	1	1
$ u_{rsna}$	Effect of RSNA on renin secretion rate	_	1	1
$\nu_{AT1R}$	Effect of AT1R-bound Ang II on renin secretion rate	_	1	1
$N_{als}$	Normalized ALD secretion rate	_	1	1
$N_{al}$	Normalized ALD concentration	_	1	1
$C_{al}$	Aldosterone concentration	$\frac{ng}{l}$	395	379
$T_{al}$	Time constant for ALD hormone secretion	$\min^{l}$	30	30
$\xi_{k/sod}$	Effect of K to Na <sup>+</sup> concentration ratio on ALD secretion rate	_	1	1
$\xi_{map}^{\kappa/soa}$	Effect of MAP on ALD secretion rate	_	1	1
Smap			-	-

Effect of AT1R-bound Ang II on ALD secretion rate	_	1	1				
Miscellaneous							
Renal sympathetic nerve activity	_	1	1				
Effect of MAP on RSNA	_	1	1				
Effect of right atrial pressure on RSNA	_	1	1				
Water intake	$\frac{ml}{min}$	0.0150	0.0150				
Na <sup>+</sup> intake	$\frac{\mu eq}{min}$	1.22	1.22				
Extracellular fluid volume	ml	46	36				
Blood volume	ml	15	12				
Time constant for ADH secretion	min	6	6				
Normalized ADH secretion rate	_	1	1				
Normalized ADH concentration	_	1	1				
Antidiuretic hormone concentration	$\frac{\mu units}{min}$	4	4				
Effect of right atrial pressure on ADH secretion rate	_	1	1				
Total amount of Na <sup>+</sup>	$\mu eq$	6540	5340				
Plasma Na <sup>+</sup> concentration	$\frac{\mu eq}{ml}$	143	147				
Normalized atrial natriuretic peptide concentration	_	1	1				
Plasma K concentration	$\frac{\mu eq}{ml}$	5	5				
Scaling Factors							
Human to rat sodium flow scaling factor	$\frac{\mu eq}{mea}$	9.69	9.69				
Human to rat urine flow scaling factor	$\frac{ml}{l}$	15	15				
Human to rat volume scaling factor	$\frac{ml}{l}$	3	2.4				
Human to rat resistance scaling factor	$\frac{l}{ml}$	0.343	0.537				
	Renal sympathetic nerve activity Effect of MAP on RSNA Effect of right atrial pressure on RSNA Water intake Na <sup>+</sup> intake Extracellular fluid volume Blood volume Time constant for ADH secretion Normalized ADH secretion rate Normalized ADH concentration Antidiuretic hormone concentration Effect of right atrial pressure on ADH secretion rate Total amount of Na <sup>+</sup> Plasma Na <sup>+</sup> concentration Normalized atrial natriuretic peptide concentration Plasma K concentration Scaling Factors Human to rat sodium flow scaling factor Human to rat volume scaling factor	Miscellaneous         Renal sympathetic nerve activity       —         Effect of MAP on RSNA       —         Effect of right atrial pressure on RSNA       —         Water intake $\frac{ml}{min}$ Na <sup>+</sup> intake $\frac{ml}{min}$ Extracellular fluid volume $ml$ Blood volume $ml$ Time constant for ADH secretion $min$ Normalized ADH secretion rate       —         Normalized ADH concentration       —         Antidiuretic hormone concentration $\frac{\mu units}{min}$ Effect of right atrial pressure on ADH secretion rate       —         Total amount of Na <sup>+</sup> $\frac{\mu eq}{ml}$ Plasma Na <sup>+</sup> concentration $\frac{\mu eq}{ml}$ Normalized atrial natriuretic peptide concentration       —         Plasma K concentration $\frac{\mu eq}{ml}$ Scaling Factors         Human to rat sodium flow scaling factor $\frac{meq}{ml}$ Human to rat volume scaling factor $\frac{ml}{l}$ Human to rat volume scaling factor $\frac{ml}{l}$	MiscellaneousRenal sympathetic nerve activity—1Effect of MAP on RSNA—1Effect of right atrial pressure on RSNA—1Water intake $\frac{ml}{min}$ 0.0150 $\frac{ml}{min}$ 1.22Na <sup>+</sup> intake $\frac{\mu eq}{min}$ 1.221.22Extracellular fluid volume $ml$ 46 $ml$ 15Blood volume $ml$ 1515Time constant for ADH secretion $min$ 6 $min$ 6Normalized ADH secretion rate—1Normalized ADH concentration $\frac{\mu units}{min}$ 4 $\frac{\mu units}{min}$ 4Effect of right atrial pressure on ADH secretion rate—1Total amount of Na <sup>+</sup> $\frac{\mu eq}{ml}$ 6540 $\frac{\mu eq}{ml}$ 143Normalized atrial natriuretic peptide concentration $\frac{\mu eq}{ml}$ 143 $\frac{1}{ml}$ 143Normalized atrial natriuretic peptide concentration—1Plasma K concentration $\frac{\mu eq}{ml}$ 5 $\frac{1}{ml}$ 5Human to rat sodium flow scaling factor $\frac{\mu eq}{ml}$ 9.69Human to rat volume scaling factor $\frac{ml}{l}$ 15Human to rat volume scaling factor $\frac{ml}{l}$ 3Human to rat volume scaling factor $\frac{ml}{l}$ 3				

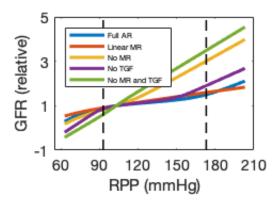


Figure S1: Renal autoregulation. Following the same experimental protocol as in Ref. (14), renal perfusion pressure is perturbed from its baseline value of 103 mmHg within a wide range. Male GFR is plotted as relative change from baseline value for full autoregulation, no myogenic response, no TGF, and no myogenic response and TGF. Dashed vertical lines are at the end points of the autoregulatory range of 90-170 mmHg. Female GFR curves are the same.

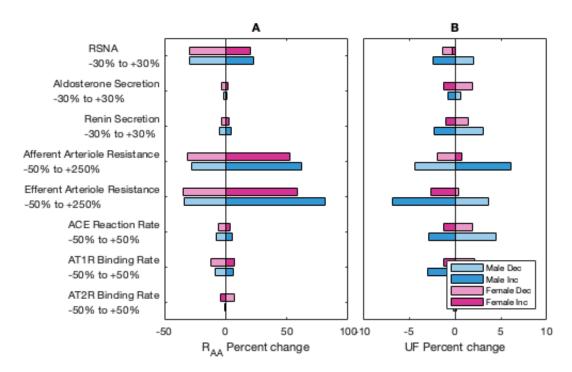


Figure S2: Sensitivity analysis. Percent change in afferent arteriolar resistance and urine flow due to various perturbations.