

Electronic Supplementary Information: Application of *in vitro* bioassays for water quality monitoring in three drinking water treatment plants using different treatment processes including biological treatment, nanofiltration and ozonation coupled with disinfection

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Table S1: Selected water quality parameters of the source water feeding the three WTPs.

Parameter	WTP	May 2018	July 2018	October 2018	December 2018
Turbidity (NTU)	Méry-sur-Oise*	5.1	5.3	3.2	14.3
	Choisy-le-Roi	5.0	5.0	2.0	4.0
	Neuilly-sur-Marne	10.0	12.6	8.0	6.0
Total organic carbon (mg/L)	Méry-sur-Oise	2.7	2.0	2.9	3.7
	Choisy-le-Roi	2.8	2.7	2.3	2.9
	Neuilly-sur-Marne	2.5	2.4	2.5	3.2
Temperature (°C)	Méry-sur-Oise	17.8	23.7	15.3	9.2
	Choisy-le-Roi	18.2	24.4	16.6	9.9
	Neuilly-sur-Marne	18.1	24.4	16.2	8.9
Flow rate (m <sup>3</sup> /s)	Méry-sur-Oise	78	54	36	47
	Choisy-le-Roi	214	106	90	93
	Neuilly-sur-Marne	110	42	46	50

\*units of formazin nephelometric unit (FNU).

Table S2: Selected water quality parameters of the produced water from the three WTPs.

<b>Parameter</b>	<b>WTP</b>	<b>May 2018</b>	<b>July 2018</b>	<b>October 2018</b>	<b>December 2018</b>
Turbidity (NTU)	Méry-sur-Oise	0.02	0.02	0.02	0.04
	Choisy-le-Roi*	0.03	0.03	0.03	0.03
	Neuilly-sur-Marne	0.03	0.03	0.04	0.04
Total organic carbon (mg/L)	Méry-sur-Oise	0.36	0.48	0.3	0.4
	Choisy-le-Roi	1.3	1.0	1.0	1.2
	Neuilly-sur-Marne	1.0	0.9	1.1	1.3
Temperature (°C)	Méry-sur-Oise	14.1	24.5	16.7	9.3
	Choisy-le-Roi	14.5	24.9	16.3	9.1
	Neuilly-sur-Marne	14.3	24.5	16.3	9.5
Residual free chlorine (mg/L)	Méry-sur-Oise	0.25	0.35	0.35	0.25
	Choisy-le-Roi	0.30	0.40	0.40	0.30
	Neuilly-sur-Marne	0.40	0.47	0.51	0.50
Conductivity (µS/cm)	Méry-sur-Oise	457	520	485	460
	Choisy-le-Roi	570	580	495	430
	Neuilly-sur-Marne	573	632	500	633
pH	Méry-sur-Oise	7.71	7.55	7.84	7.94
	Choisy-le-Roi	7.58	7.40	7.61	7.74
	Neuilly-sur-Marne	7.38	7.28	7.54	7.70

\*units of formazin nephelometric unit (FNU).

Table S3: IC<sub>10</sub> and EC<sub>10</sub> values for ERα GeneBLAzer (agonist mode) in units of relative enrichment factor (REF).

WTP	Sample Name	May 2018		July 2018	
		Cytotoxicity	Activation of ER	Cytotoxicity	Activation of ER
		IC <sub>10</sub> (REF)	EC <sub>10</sub> (REF)	IC <sub>10</sub> (REF)	EC <sub>10</sub> (REF)
Méry-sur-Oise	Source	N/A	N/A	74.0	26.0 ± 8.8
	After nanofiltration	>100	>100	>100	>100
	After biological treatment	68.5	Cytotoxic	77.6	Cytotoxic
Choisy-le-Roi	Produced water	>100	>100	>100	>100
	Source	>100	2.56 ± 0.2	51.4 ± 5.3	11.2 ± 4.9
	After UV treatment	>100	>100	>100	>100
Neuilly-sur-Marne	Produced water	>100	>100	>100	>100
	Source	55.5	1.09 ± 0.2	65.6	3.12 ± 0.6
	After UV treatment	>100	>100	>100	>100
Control	Produced water	>100	>100	>100	>100
	Bottled water with sodium thiosulphate	>100	2.34 ± 0.6	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	>100	>100
October 2018					
WTP	Sample Name	Cytotoxicity	Activation of ER	Cytotoxicity	Activation of ER
		IC <sub>10</sub> (REF))	EC <sub>10</sub> (REF)	IC <sub>10</sub> (REF)	EC <sub>10</sub> (REF))
		IC <sub>10</sub> (REF))	EC <sub>10</sub> (REF)	IC <sub>10</sub> (REF)	EC <sub>10</sub> (REF))
Méry-sur-Oise	Source	38.4	23.2 ± 1.5	37.5	14.4 ± 0.7
	After nanofiltration	>150	>150	>150	>150
	After biological treatment	>150	>150	N/A	N/A
Choisy-le-Roi	Produced water	>150	>150	>150	>150
	Source	40.2	15.9 ± 0.6	41.6	4.45 ± 0.21
	After UV treatment	>150	>150	>150	132 ± 13
Neuilly-sur-Marne	Produced water	>150	>150	>150	>150
	Source	30.9	12.8 ± 0.8	33.3	4.78 ± 0.21
	After UV treatment	>150	>150	>150	>150
Control	Produced water	>150	>150	>150	110 ± 9.0
	Bottled water with sodium thiosulphate	>150	>150	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	>150	>150

Figure S1: Example full concentration-effect curves for induction (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in ER $\alpha$  GeneBLAzer (agonist mode) (left plot), along with linear concentration-effect curves for induction (centre plot) and cytotoxicity (right plot).

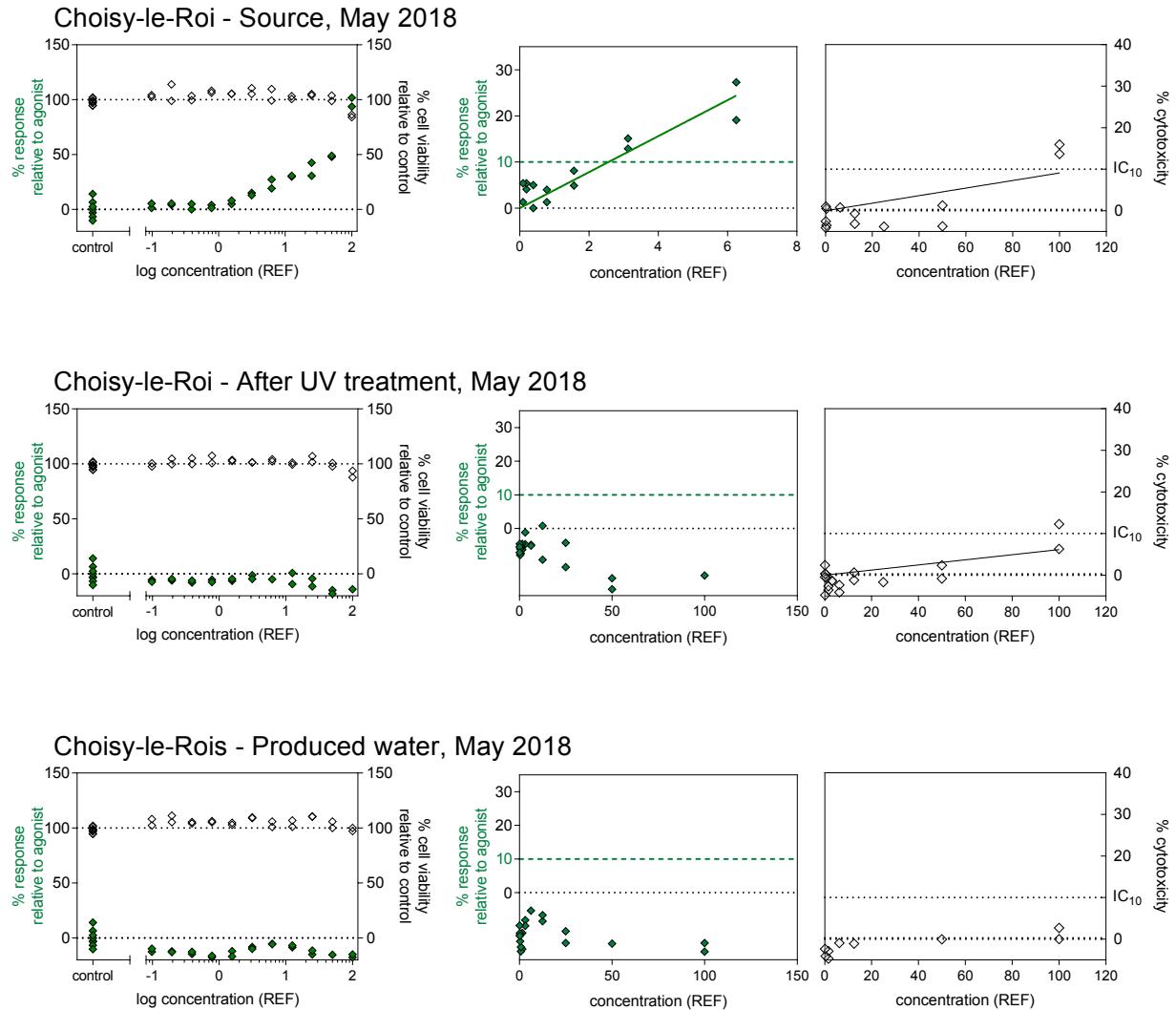


Table S4:  $IC_{01}$  and  $EC_{SR0.2}$  values for ER $\alpha$  GeneBLAzer (antagonist mode) in units of relative enrichment factor (REF). NB: samples were not run in October and December 2018.

WTP	Sample Name	May 2018		July 2018	
		Cytotoxicity $IC_{01}$ (REF)	Inhibition of ER $EC_{SR0.2}$ (REF)	Cytotoxicity $IC_{01}$ (REF)	Inhibition of ER $EC_{SR0.2}$ (REF)
Méry-sur-Oise	Source	N/A	N/A	7.33	Cytotoxic
	After nanofiltration	>100	>100	5.52	Cytotoxic
	After biological treatment	19.7	Cytotoxic	4.61	Cytotoxic
Choisy-le-Roi	Produced water	>100	>100	4.21	Cytotoxic
	Source	17.0	Cytotoxic	6.67	Cytotoxic
	After UV treatment	>100	>100	>100	>100
Neuilly-sur-Marne	Produced water	>100	>100	>100	>100
	Source	7.05	Cytotoxic	10.12	Cytotoxic
	After UV treatment	>100	>100	14.37	Cytotoxic
Control	Produced water	>100	>100	>100	>100
	Bottled water with sodium thiosulphate	99.6	Cytotoxic	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	0.37	Cytotoxic

Table S5: IC<sub>10</sub> and EC<sub>10</sub> values for AR GeneBLAzer (agonist mode) in units of relative enrichment factor (REF). NB: samples were not run in October and December 2018.

WTP	Sample Name	May 2018		July 2018	
		Cytotoxicity IC <sub>10</sub> (REF)	Activation of AR EC <sub>10</sub> (REF)	Cytotoxicity IC <sub>10</sub> (REF)	Activation of AR EC <sub>10</sub> (REF)
	Source	N/A	N/A	>100	>100
Méry-sur-Oise	After nanofiltration	>100	>100	90.3	Cytotoxic
	After biological treatment	>100	>100	>100	>100
	Produced water	>100	>100	>100	>100
Choisy-le-Roi	Source	>100	>100	61.2	Cytotoxic
	After UV treatment	>100	>100	>100	>100
	Produced water	>100	>100	>100	>100
Neuilly-sur-Marne	Source	>100	>100	99.0	Cytotoxic
	After UV treatment	>100	>100	>100	>100
	Produced water	56.1	Cytotoxic	>100	>100
Control	Bottled water with sodium thiosulphate	>100	>100	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	>100	>100

Table S6:  $IC_{01}$  and  $EC_{SR0.2}$  values for AR GeneBLAzer (antagonist mode) in units of relative enrichment factor (REF). NB: samples were not run in October and December 2018.

WTP	Sample Name	May 2018		July 2018	
		Cytotoxicity $IC_{01}$ (REF)	Inhibition of AR $EC_{SR0.2}$ (REF)	Cytotoxicity $IC_{01}$ (REF)	Inhibition of AR $EC_{SR0.2}$ (REF)
Méry-sur-Oise	Source	N/A	N/A	81.6	Cytotoxic
	After nanofiltration	>100	>100	13.7	Cytotoxic
	After biological treatment	>100	>100	>100	>100
Choisy-le-Roi	Produced water	>100	>100	>100	>100
	Source	>100	>100	5.57	Cytotoxic
	After UV treatment	53.5	Cytotoxic	10.5	Cytotoxic
Neuilly-sur-Marne	Produced water	>100	>100	>100	>100
	Source	>100	>100	8.38	Cytotoxic
	After UV treatment	32.6	Cytotoxic	>100	>100
Control	Produced water	5.09	Cytotoxic	>100	>100
	Bottled water with sodium thiosulphate	>100	>100	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	>100	>100

Table S7: IC<sub>10</sub> and EC<sub>10</sub> values for GR GeneBLAzer (agonist mode) in units of relative enrichment factor (REF). NB: samples were not run in October and December 2018.

WTP	Sample Name	May 2018		July 2018	
		Cytotoxicity IC <sub>10</sub> (REF)	Activation of GR EC <sub>10</sub> (REF)	Cytotoxicity IC <sub>10</sub> (REF)	Activation of GR EC <sub>10</sub> (REF)
	Source	N/A	N/A	50.0	Cytotoxic
Méry-sur-Oise	After nanofiltration	>100	>100	>100	>100
	After biological treatment	>100	>100	>100	>100
	Produced water	>100	>100	>100	>100
Choisy-le-Roi	Source	>100	>100	71.8	Cytotoxic
	After UV treatment	>100	>100	>100	>100
	Produced water	>100	>100	>100	>100
Neuilly-sur-Marne	Source	>100	>100	>100	>100
	After UV treatment	>100	>100	>100	>100
	Produced water	>100	>100	>100	>100
Control	Bottled water with sodium thiosulphate	>100	>100	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	>100	>100

Table S8:  $IC_{01}$  and  $EC_{SR0.2}$  values for GR GeneBLAzer (antagonist mode) in units of relative enrichment factor (REF). NB: samples were not run in October and December 2018.

WTP	Sample Name	May 2018		July 2018	
		Cytotoxicity $IC_{01}$ (REF)	Inhibition of GR $EC_{SR0.2}$ (REF)	Cytotoxicity $IC_{01}$ (REF)	Inhibition of GR $EC_{SR0.2}$ (REF)
Méry-sur-Oise	Source	N/A	N/A	4.08	Cytotoxic
	After nanofiltration	22.8	Cytotoxic	10.4	Cytotoxic
	After biological treatment	25.9	Cytotoxic	9.77	Cytotoxic
Choisy-le-Roi	Produced water	22.9	Cytotoxic	17.1	Cytotoxic
	Source	22.0	Cytotoxic	5.21	Cytotoxic
	After UV treatment	41.3	Cytotoxic	>100	>100
Neuilly-sur-Marne	Produced water	>100	>100		Cytotoxic
	Source	25.3	Cytotoxic	16.0	Cytotoxic
	After UV treatment	>100	>100	>100	>100
Control	Produced water	11.5	Cytotoxic	>100	>100
	Bottled water with sodium thiosulphate	15.3	Cytotoxic	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	8.53	Cytotoxic

Table S9: IC<sub>10</sub> and EC<sub>10</sub> values for PR GeneBLAzer (agonist mode) in units of relative enrichment factor (REF). NB: samples were not run in October and December 2018.

WTP	Sample Name	May 2018		July 2018	
		Cytotoxicity IC <sub>10</sub> (REF)	Activation of PR EC <sub>10</sub> (REF)	Cytotoxicity IC <sub>10</sub> (REF)	Activation of PR EC <sub>10</sub> (REF)
Méry-sur-Oise	Source	N/A	N/A	>100	>100
	After nanofiltration	>100	>100	>100	>100
	After biological treatment	>100	>100	>100	>100
Choisy-le-Roi	Produced water	>100	>100	>100	>100
	Source	>100	>100	>100	>100
	After UV treatment	>100	>100	>100	>100
Neuilly-sur-Marne	Produced water	>100	>100	>100	>100
	Source	>100	>100	>100	>100
	After UV treatment	>100	>100	>100	>100
Control	Produced water	>100	>100	>100	>100
	Bottled water with sodium thiosulphate	>100	>100	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	>100	>100

Table S10: IC<sub>01</sub> and EC<sub>SR0.2</sub> values for PR GeneBLAzer (antagonist mode) in units of relative enrichment factor (REF). NB: samples were not run in October and December 2018.

WTP	Sample Name	May 2018		July 2018	
		Cytotoxicity IC <sub>01</sub> (REF)	Inhibition of PR EC <sub>SR0.2</sub> (REF)	Cytotoxicity IC <sub>01</sub> (REF)	Inhibition of PR EC <sub>SR0.2</sub> (REF)
Méry-sur-Oise	Source	N/A	N/A	13.8	Cytotoxic
	After nanofiltration	38.2	Cytotoxic	>100	>100
	After biological treatment	>100	>100	87.4	Cytotoxic
Choisy-le-Roi	Produced water	>100	>100	>100	>100
	Source	>100	>100	13.2	Cytotoxic
	After UV treatment	>100	>100	>100	>100
Neuilly-sur-Marne	Produced water	>100	>100	>100	>100
	Source	>100	>100	>100	>100
	After UV treatment	>100	>100	>100	>100
Control	Produced water	>100	>100	>100	>100
	Bottled water with sodium thiosulphate	>100	>100	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	37.1	Cytotoxic

Figure S2: Example full concentration-effect curves for inhibition (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in ER $\alpha$  GeneBLAzer (antagonist mode) (left plot), along with linear concentration-effect curves for inhibition (centre plot) and cytotoxicity (right plot).

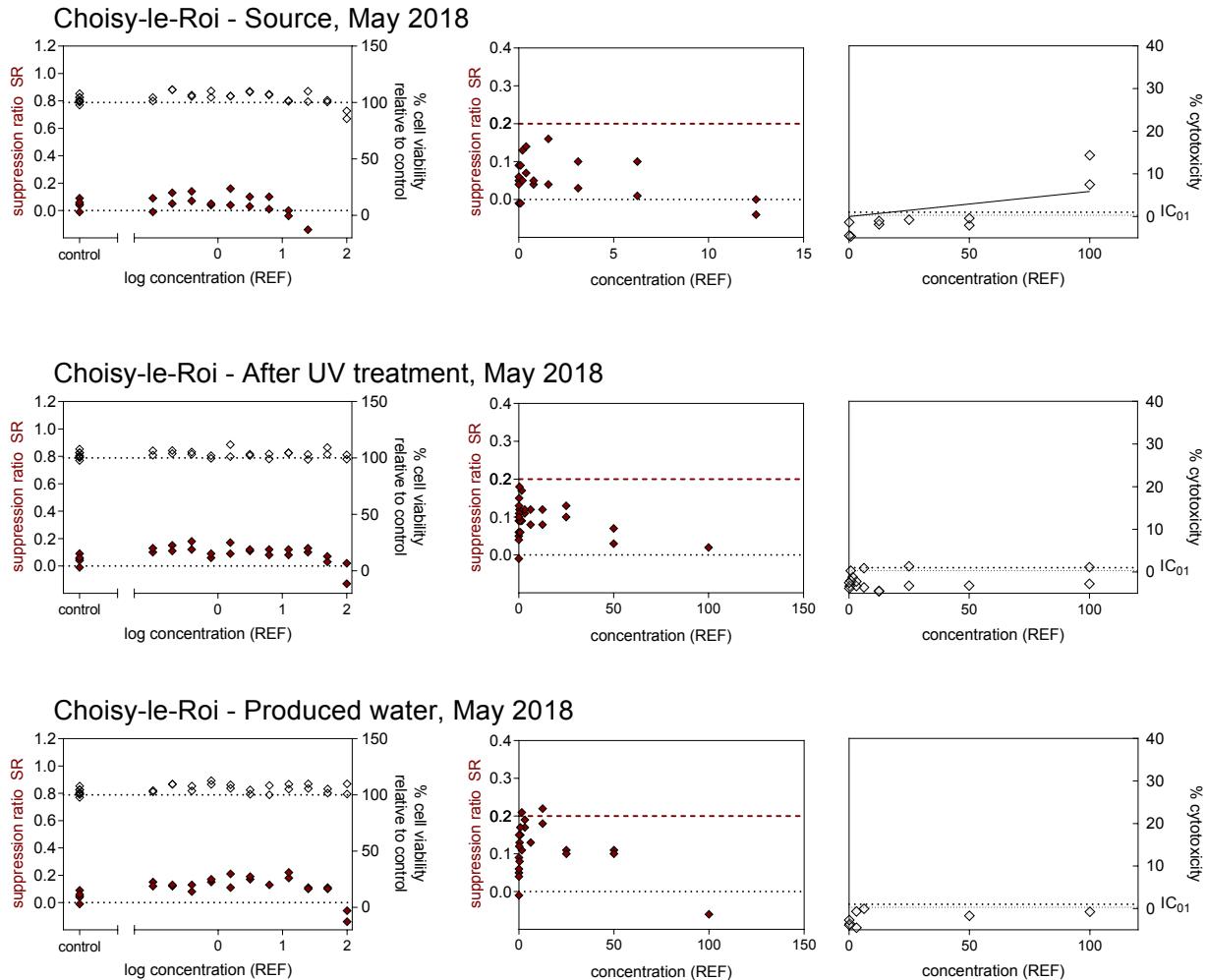


Figure S3: Example full concentration-effect curves for induction (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in AR GeneBLAzer (agonist mode) (left plot), along with linear concentration-effect curves for induction (centre plot) and cytotoxicity (right plot).

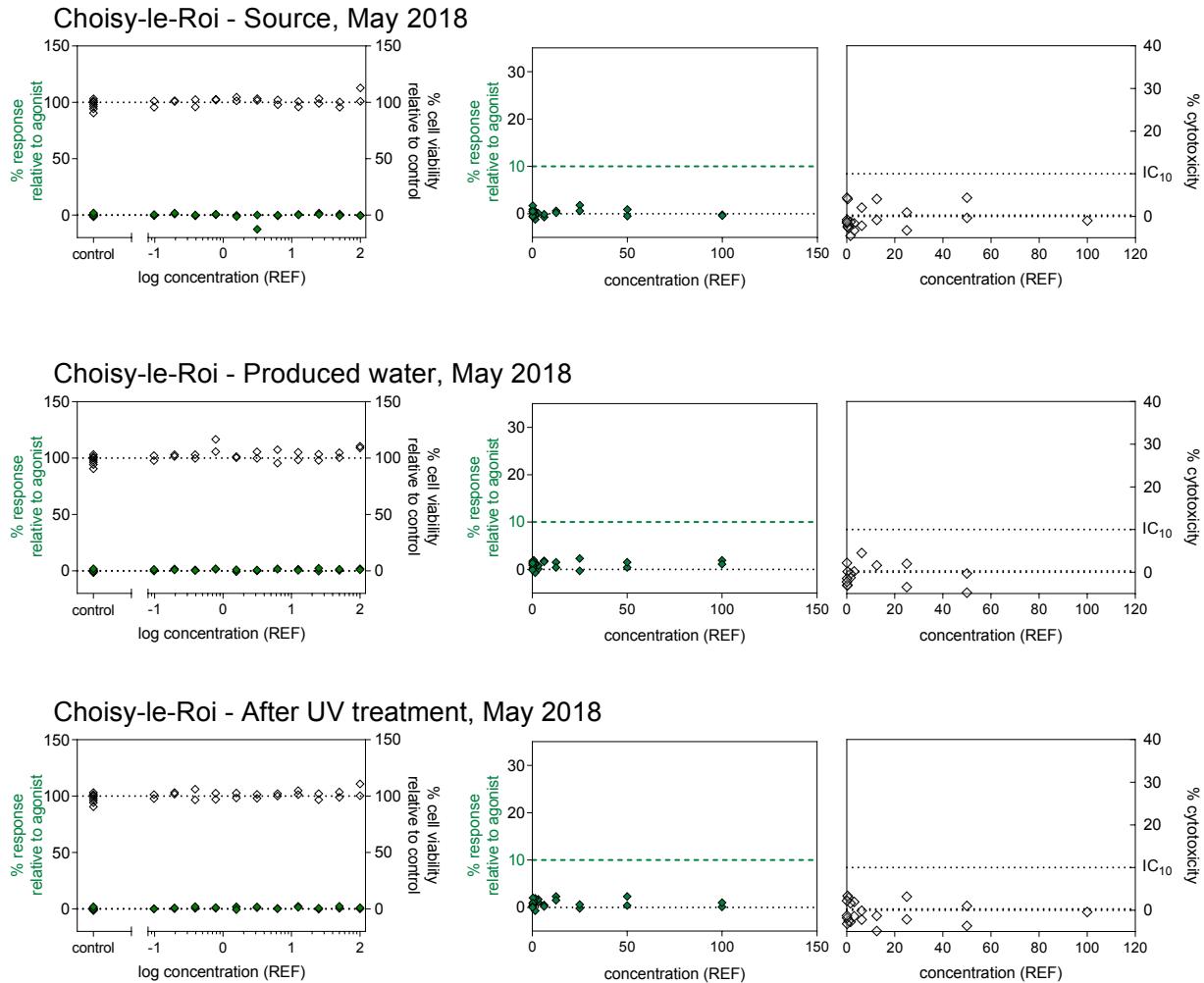


Figure S4: Example full concentration-effect curves for inhibition (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in AR GeneBLAzer (antagonist mode) (left plot), along with linear concentration-effect curves for inhibition (centre plot) and cytotoxicity (right plot).

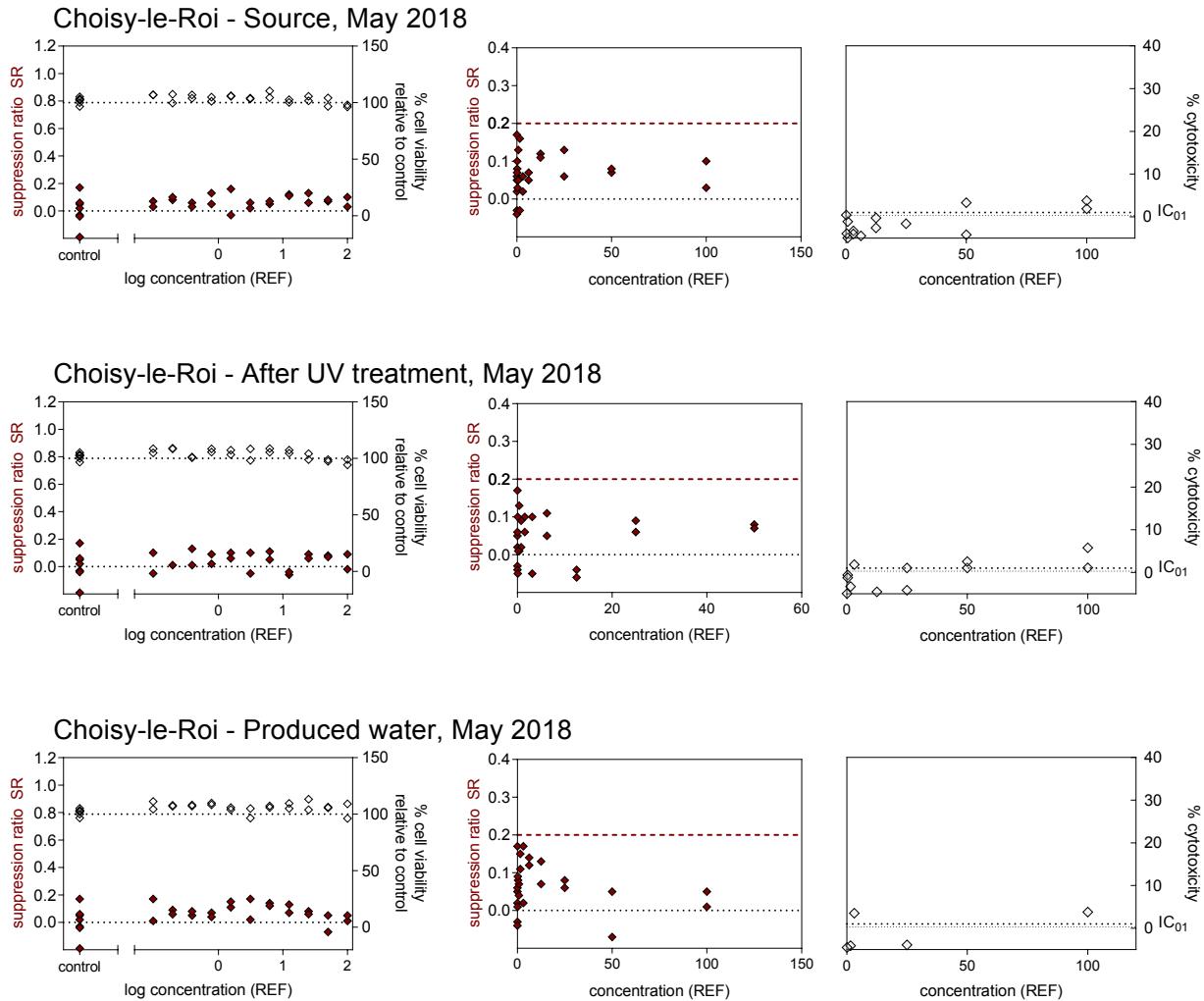


Figure S5: Example full concentration-effect curves for induction (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in GR GeneBLAzer (agonist mode) (left plot), along with linear concentration-effect curves for induction (centre plot) and cytotoxicity (right plot).

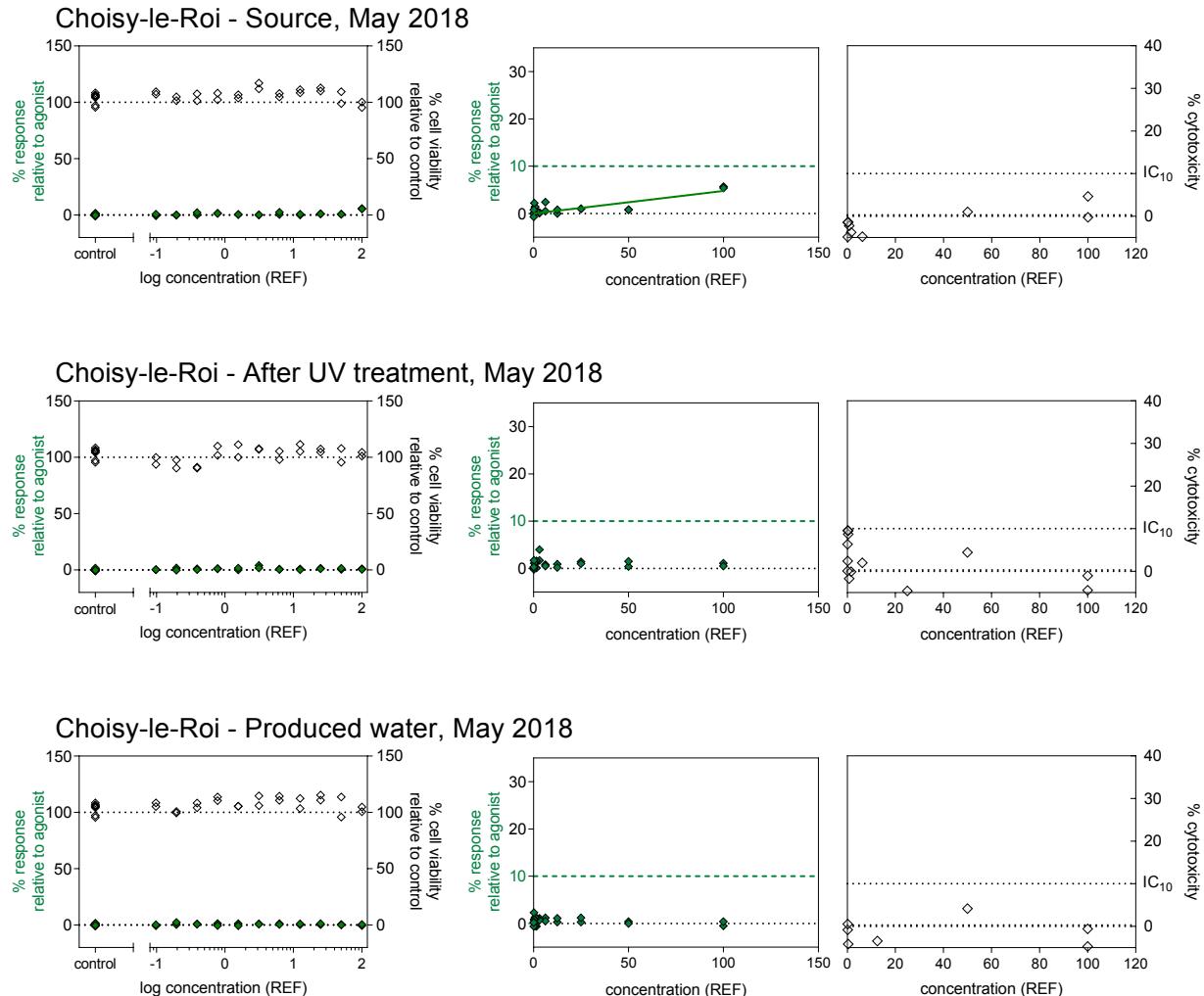


Figure S6: Example full concentration-effect curves for inhibition (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in GR GeneBLAzer (antagonist mode) (left plot), along with linear concentration-effect curves for inhibition (centre plot) and cytotoxicity (right plot).

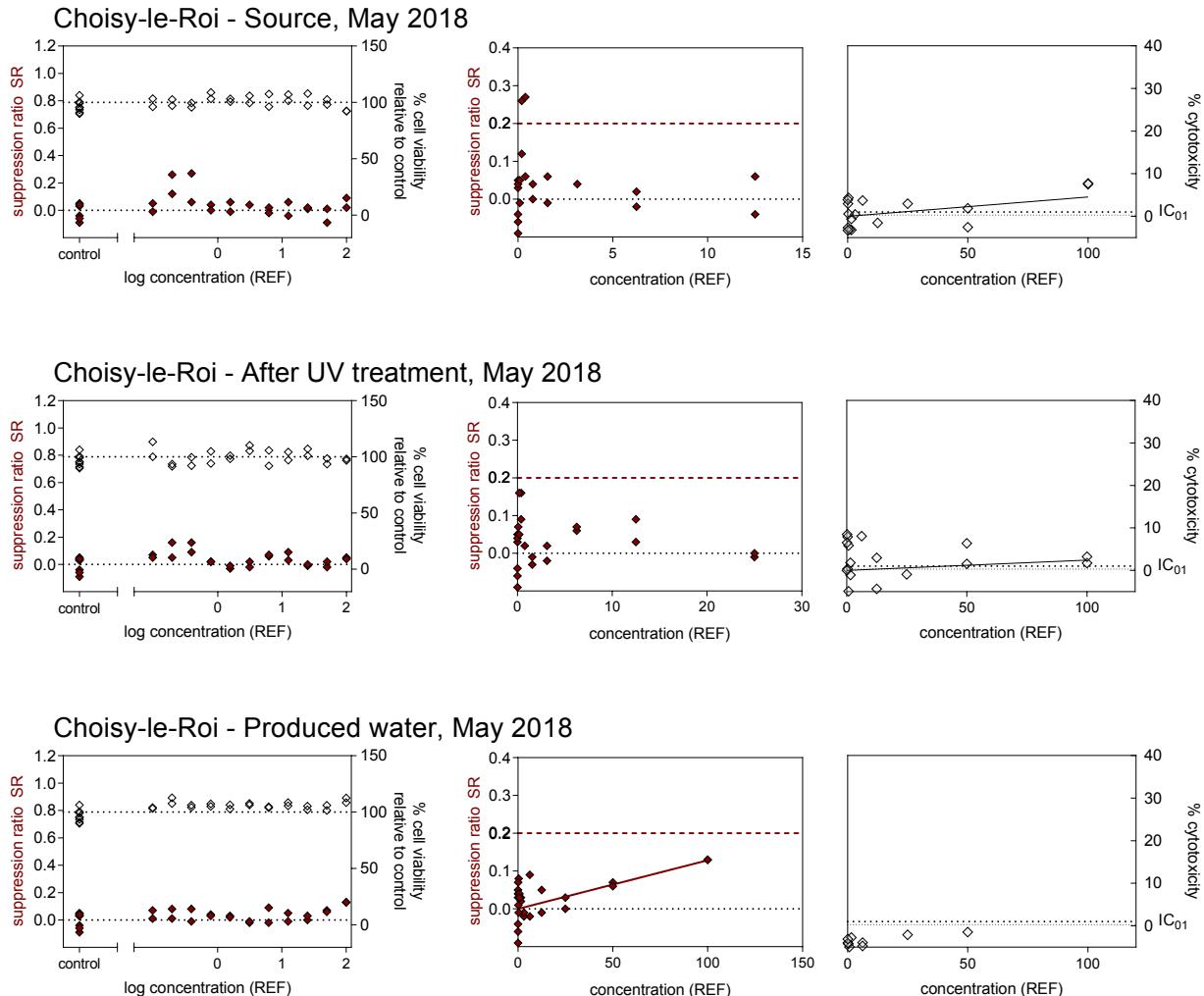


Figure S7: Example full concentration-effect curves for induction (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in PR GeneBLAzer (agonist mode) (left plot), along with linear concentration-effect curves for induction (centre plot) and cytotoxicity (right plot).

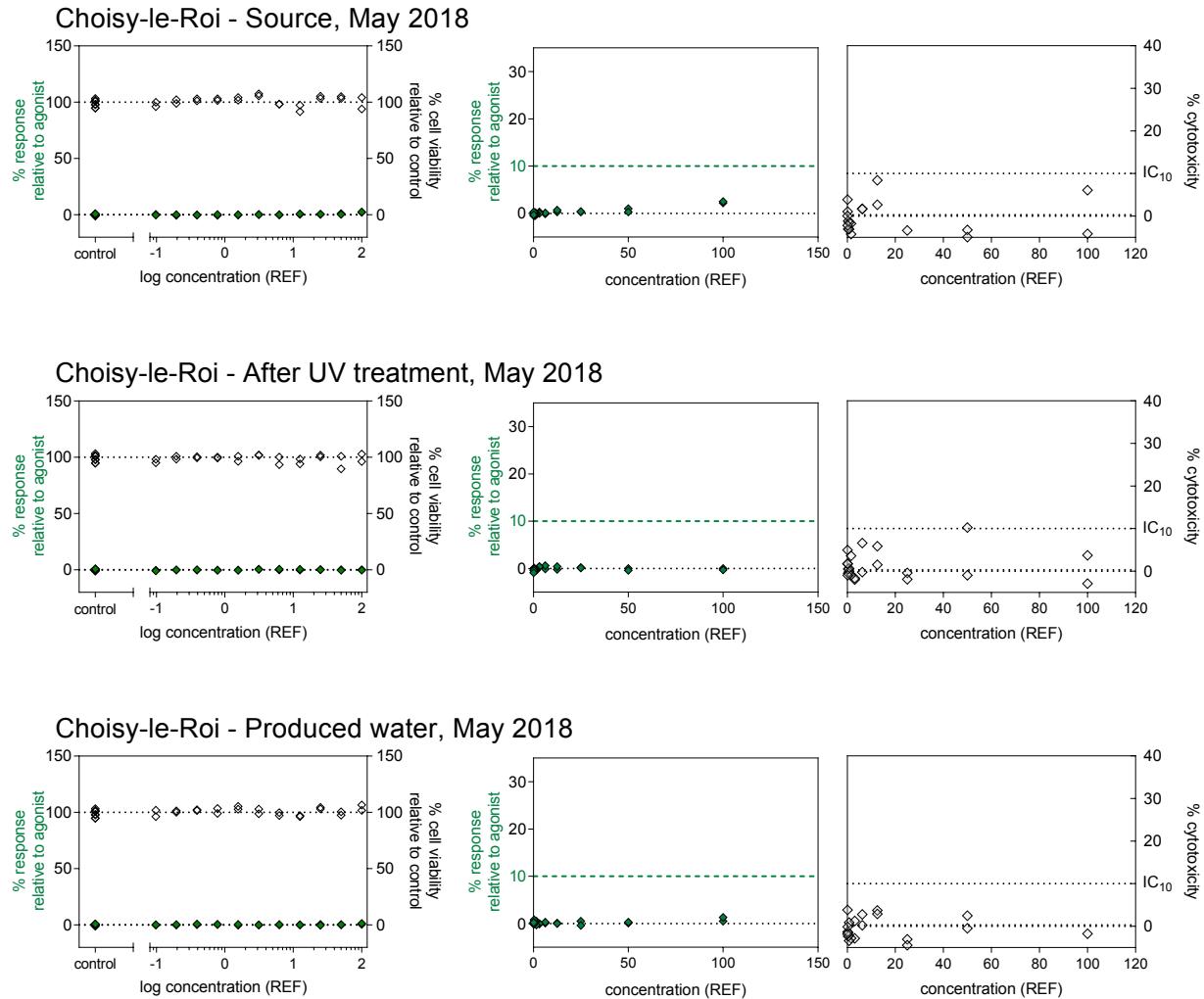


Figure S8: Example full concentration-effect curves for inhibition (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in PR GeneBLAzer (antagonist mode) (left plot), along with linear concentration-effect curves for inhibition (centre plot) and cytotoxicity (right plot).

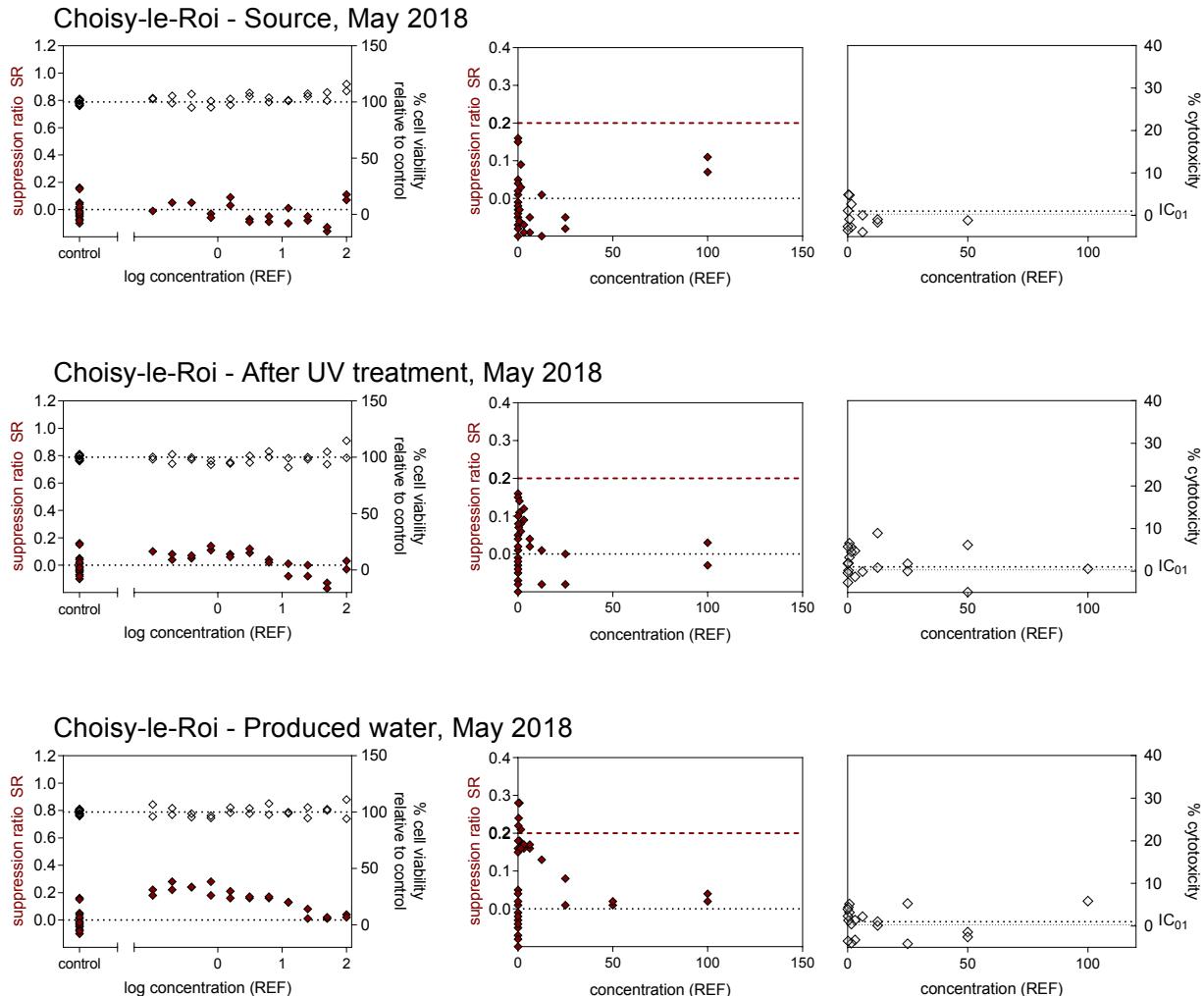


Figure S9: Example full concentration-effect curves for induction (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in AREc32 (agonist mode) (left plot), along with linear concentration-effect curves for induction (centre plot) and cytotoxicity (right plot).

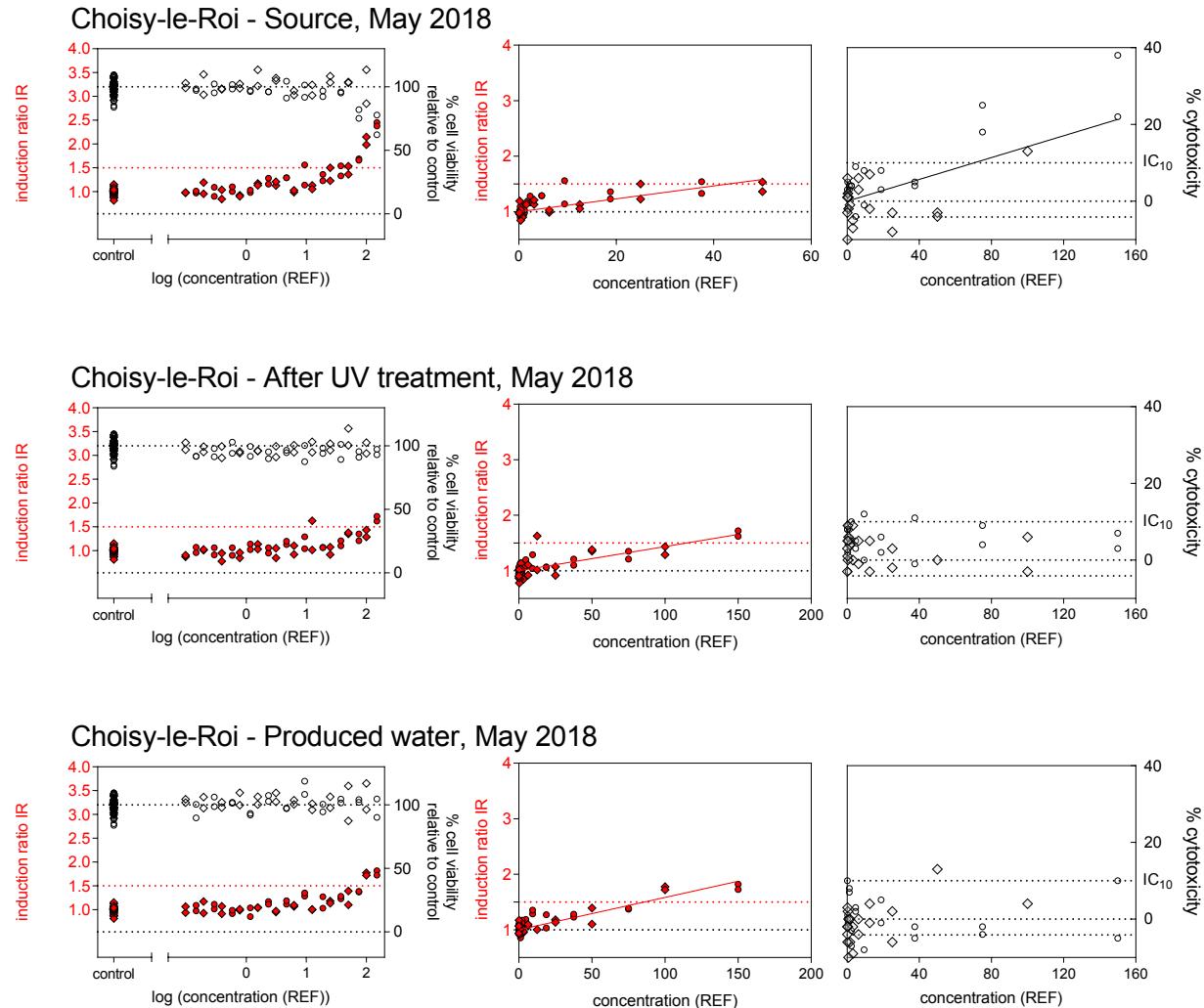


Figure S10: Example full concentration-effect curves for induction (filled symbols) for Choisy-le-Roi (May 2018) in NF-κB GeneBLAzer (agonist mode) (left plot), along with linear concentration-effect curves for induction (right plot).

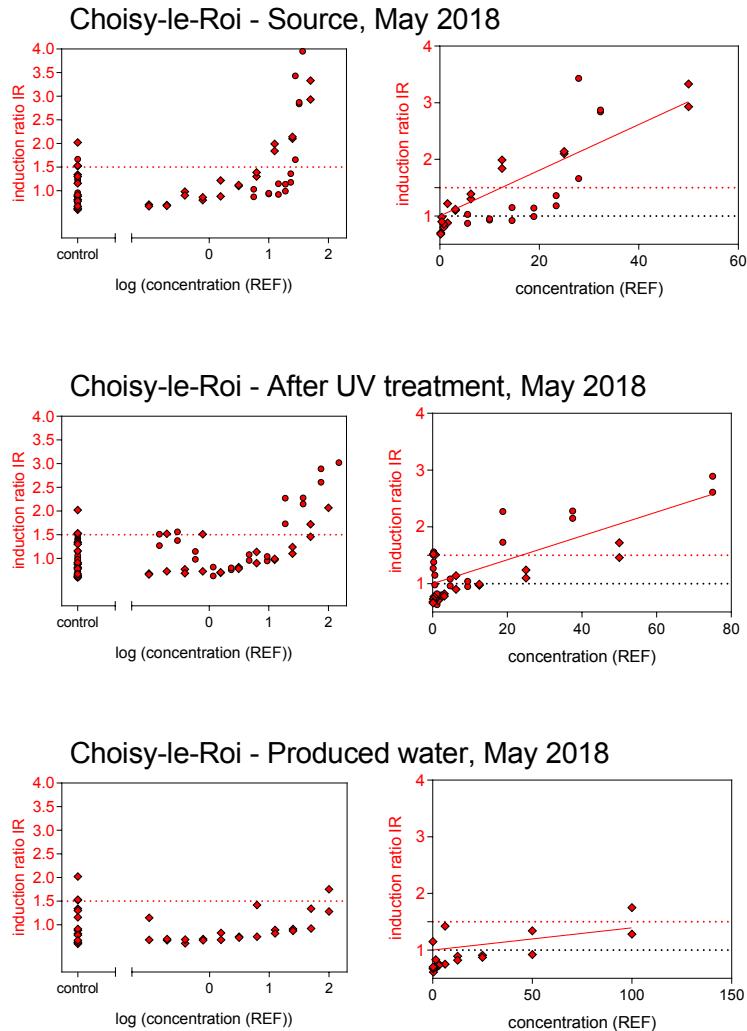


Figure S11: Example full concentration-effect curves for induction (filled symbols) and cell viability (empty symbols) for Choisy-le-Roi (May 2018) in p53RE GeneBLAzer (agonist mode) (left plot), along with linear concentration-effect curves for induction (centre plot) and cytotoxicity (right plot).

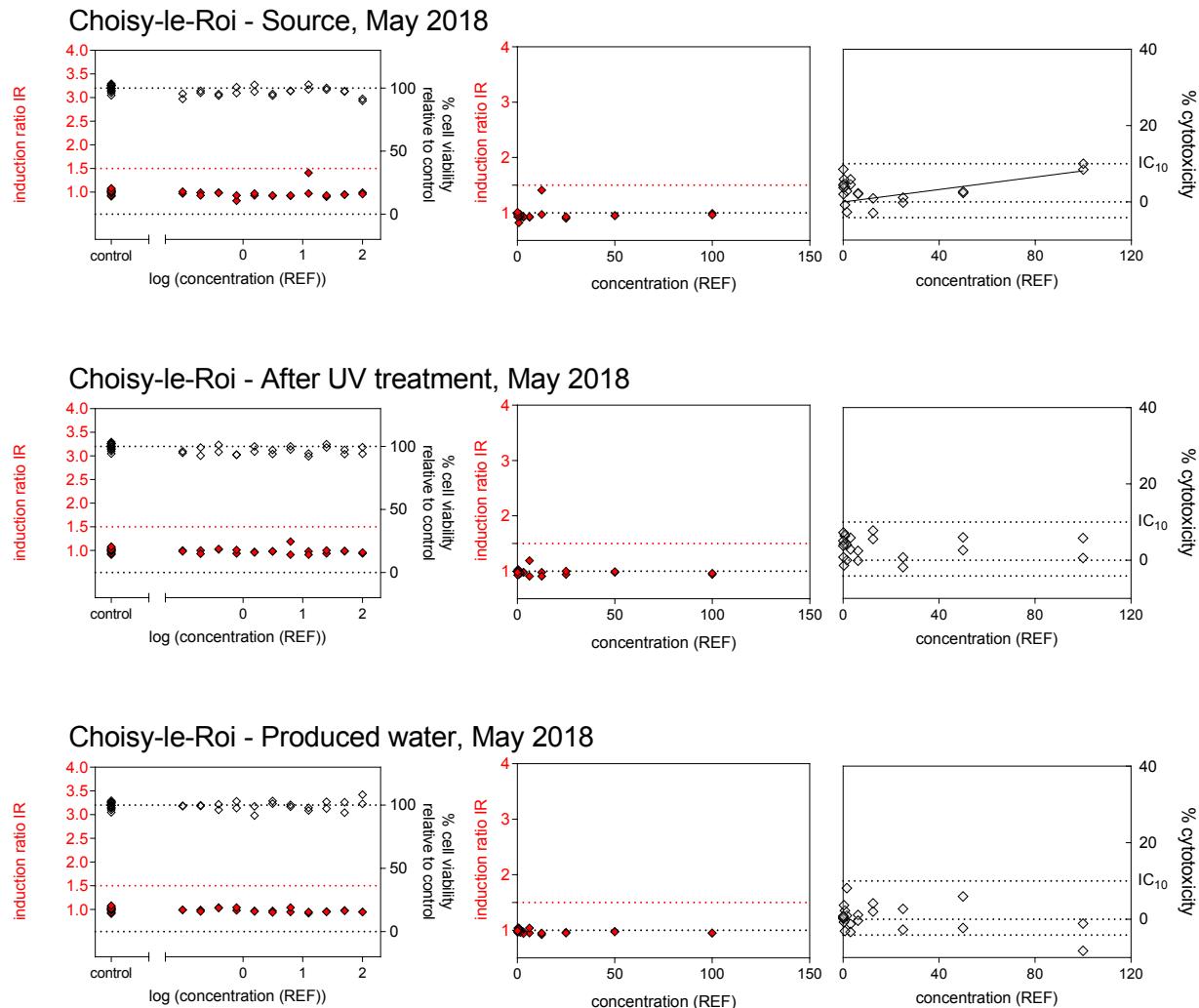


Table S11:  $IC_{10}$  and  $EC_{IRI.5}$  values for AREc32 in units of relative enrichment factor (REF).

WTP	Sample Name	May 2018		July 2018	
		Cytotoxicity $IC_{10}$ (REF)	Oxidative stress response $EC_{IRI.5}$ (REF)	Cytotoxicity $IC_{10}$ (REF)	Oxidative stress response $EC_{IRI.5}$ (REF)
		N/A	N/A	37.5	Cytotoxic
Méry-sur-Oise	Source	N/A	N/A	37.5	Cytotoxic
	After nanofiltration	>100	114 ± 7.6	>100	>100
	After biological treatment	>100	>100	>100	>100
Choisy-le-Roi	Produced water	>100	>100	>100	>100
	Source	70.3	43.0 ± 5.1	36.4	Cytotoxic
	After UV treatment	>150	115 ± 12	>100	>100
Neuilly-sur-Marne	Produced water	>150	85.8 ± 5.5	>100	>100
	Source	88.9	61.5 ± 4.9	>100	>100
	After UV treatment	>150	127 ± 12	>100	>100
Control	Produced water	>150	86.1 ± 6.3	>150	103 ± 7.1
	Bottled water with sodium thiosulphate	>100	>100	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	>100	>100
October 2018					
WTP	Sample Name	Cytotoxicity $IC_{10}$ (REF))	Oxidative stress response $EC_{IRI.5}$ (REF)	Cytotoxicity $IC_{10}$ (REF)	Oxidative stress response $EC_{IRI.5}$ (REF)
		45.5	Cytotoxic	40.6	Cytotoxic
		>150	136 ± 7.4	>150	>150
Oise	After nanofiltration	>150	82.3 ± 3.1	N/A	N/A
	After biological treatment	>150	102 ± 6.1	>150	>150
	Produced water	>150	>150	>150	>150
Choisy-le-Roi	Source	59.0	Cytotoxic	49.7	Cytotoxic
	After UV treatment	>150	>150	>150	114 ± 8.4
	Produced water	>150	>150	>150	77.5 ± 3.6
Neuilly-sur-Marne	Source	49.2	Cytotoxic	36.0	Cytotoxic
	After UV treatment	>150	88.0 ± 4.4	>150	>150
	Produced water	>150	87.2 ± 4.2	>150	>150
Control	Bottled water with sodium thiosulphate	>150	>150	N/A	N/A
	Bottled water without sodium thiosulphate	N/A	N/A	>150	>150

Table S12: EC<sub>IR1.5</sub> values for NF-κB GeneBLAzer in units of relative enrichment factor (REF).

WTP	Sample Name	May 2018	July 2018	October	December
		NF-κB	NF-κB	NF-κB	NF-κB
		<i>response</i>	<i>response</i>	<i>response</i>	<i>response</i>
		EC <sub>IR1.5</sub> (REF)	EC <sub>IR1.5</sub> (REF)	EC <sub>IR1.5</sub> (REF))	EC <sub>IR1.5</sub> (REF)
Méry-sur-Oise	Source	N/A	>100	17.0 ± 3.7	6.23 ± 0.6
	After nanofiltration	>100	>100	>150	117 ± 32
	After biological treatment	32.4 ± 8.3	>100	56.8 ± 5.5	N/A
Choisy-le-Roi	Produced water	80.4 ± 18	>100	65.2 ± 8.9	49.4 ± 5.3
	Source	12.4 ± 1.2	39.1 ± 3.6	16.9 ± 1.1	5.11 ± 0.6
	After UV treatment	23.8 ± 2.7	40.8 ± 5.4	122 ± 20	49.6 ± 5.3
Neuilly-sur-Marne	Produced water	>100	>100	25.8 ± 3.1	118 ± 23
	Source	6.71 ± 0.5	38.1 ± 2.7	6.37 ± 0.6	7.71 ± 0.7
	After UV treatment	42.0 ± 5.2	>100	37.5 ± 2.9	51.0 ± 4.4
Control	Produced water	52.4 ± 9.3	>100	15.6 ± 1.4	68.4 ± 9.1
	Bottled water with sodium thiosulphate	>100	N/A	106 ± 26	N/A
	Bottled water without sodium thiosulphate	N/A	>100	N/A	>150

Table S13: IC<sub>10</sub> values for p53RE GeneBLAzer in units of relative enrichment factor (REF). None of the samples were active in the assay.

WTP	Sample Name	May 2018	July 2018	October 2018	December 2018
		Cytotoxicity IC <sub>10</sub> (REF)	Cytotoxicity IC <sub>10</sub> (REF)	Cytotoxicity IC <sub>10</sub> (REF)	Cytotoxicity IC <sub>10</sub> (REF)
Méry-sur-Oise	Source	N/A	34.5	55.5	40.3
	After nanofiltration	>100	>100	>100	>100
	After biological treatment	>100	>100	>100	>100
Choisy-le-Roi	Produced water	>100	>100	>100	>100
	Source	>100	50.2	57.2	37.6
	After UV treatment	>100	>100	>100	>100
Neuilly-sur-Marne	Produced water	>100	>100	>100	>100
	Source	86.6	62.2	57.6	43.8
	After UV treatment	>100	>100	>100	>100
Control	Produced water	>100	>100	>100	>100
	Bottled water with sodium thiosulphate	>100	N/A	>100	N/A
	Bottled water without sodium thiosulphate	N/A	>100	N/A	>100

Figure S12: Concentration-effect curves for growth inhibition in *Salmonella typhimurium* TA98.

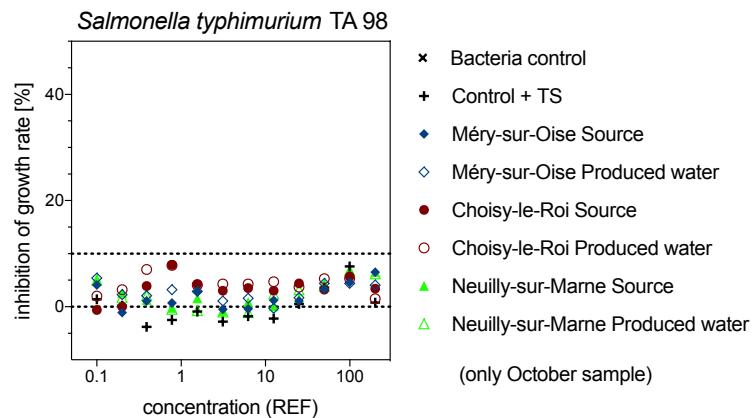


Figure S13: Concentration-effect curves of positive controls in *Salmonella typhimurium* TA98, TA100 and YG7108.

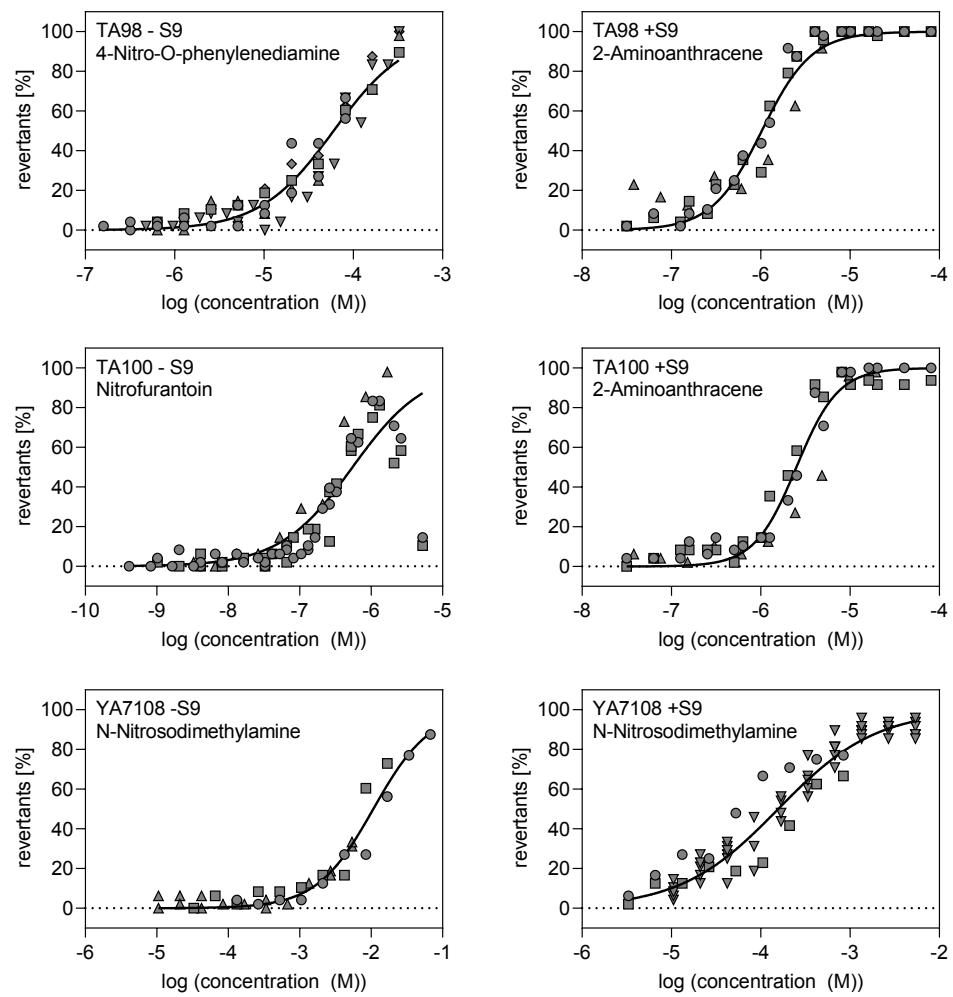


Figure S14: Concentration-effect curves of water extracts in *Salmonella typhimurium* TA98, TA100 and YG7108 with and without 0.15 mg<sub>protein</sub>/mL S9.

