

Supplemental Table 1: Summary of Study 2 treatment-related findings for animals sacrificed after 28 or 90 days of ring exposure. Microscopic findings were either reported as present (i.e., no adverse finding) or assigned a severity grade of minimal, slight, moderate, marked, or severe.

Treatment Duration	Site / Finding	Degree	Control	Placebo IVR	TFV IVR
28 Days	Vagina / Lymphocytic Infiltration	Minimal	2	1	0
		Slight	1	4	4
		Moderate	0	0	1
		Total	3	5	5
		Number of tissues examined	4	5	5
90 Days	Uterine Cervix / Lymphocytic Infiltration	Minimal	2	1	2
		Slight	0	0	2
		Moderate	0	1	0
		Total	2	2	4
		Number of tissues examined	5	5	5

Supplemental Table 2: Study 2 TFV vaginal fluid, vaginal tissue, and plasma concentrations from the TFV IVR.

Collection Time and Location	Animal Number					Mean	SD
	1	2	3	4	5		
Vaginal fluid (ng/g)							
Proximal, Day 28	5.8E+6	5.6E+6	9.6E+6	7.9E+6	4.8E+6	6.8E+6	2.0E+6
Distal, Day 28	1.8E+6	1.3E+7	1.4E+6	4.4E+6	1.9E+6	4.6E+6	5.0E+6
Proximal, Day 90	3.0E+6	1.7E+6	3.6E+5	1.6E+6	1.4E+6	1.6E+6	9.5E+5
Distal, Day 90	3.3E+6	1.5E+6	2.1E+5	1.5E+6	1.3E+6	1.6E+6	1.1E+6
Vaginal tissue (ng/g)							
Proximal, Day 28	8.1E+4	4.7E+3	6.1E+3	7.6E+3	4.0E+3	2.1E+4	3.4E+4
Distal, Day 28	6.7E+4	2.5E+4	9.2E+3	1.7E+4	4.5E+3	2.4E+4	2.5E+4
Proximal, Day 90	7.8E+3	2.0E+3	1.4E+3	1.3E+3	1.7E+3	2.9E+3	2.8E+3
Distal, Day 90	3.6E+3	1.4E+3	6.7E+2	5.5E+3	1.6E+3	2.5E+3	2.0E+3
Plasma (ng/mL)							
Day 28	23.8	18.0	29.4	24.0	31.7	25.4	5.4
Day 90	11.0	2.0	*	8.1	15.6	7.3	6.4

* BLQ<1.0 ng/mL