

**eTable 4** Geometric means sVEGF and mean and SD  $\log_e(s\text{VEGF})$  across study visits by SSAE group

	IME (n=25)		ATE (n=53)		OTH (n=104)		No SSAEs (n=449)	
	Mean (SD)		Mean (SD)		Mean (SD)		Mean (SD)	
	GM	$\log_e(s\text{VEGF})$	GM	$\log_e(s\text{VEGF})$	GM	$\log_e(s\text{VEGF})$	GM	$\log_e(s\text{VEGF})$
<b>Visit 0</b>								
sVEGF (including imputed)	173	5.16 (0.83)	210	5.35 (0.91)	164	5.10 (1.02)	166	5.11 (0.79)
Below detectable limit (n, %)	1/24	4.2%	1/47	2.1%	6/83	7.2%	14/427	3.3%
<b>Visit 1</b>								
sVEGF (including imputed)	90	4.50 (0.71)	115	4.75 (1.11)	96	4.57 (1.06)	90	4.50 (1.13)
Below detectable limit (n, %)	1/21	4.8%	4/47	8.5%	13/83	15.7%	59/428	13.8%
<b>Visit 11</b>								
sVEGF (including imputed)	132	4.88 (0.92)	98	4.58 (1.88)	73	4.29 (1.48)	74	4.31 (1.21)
Below detectable limit (n, %)	2/20	10.0%	8/34	23.5%	19/66	28.8%	91/385	23.6%
<b>Visit 12</b>								
sVEGF (including imputed)	62	4.13 (1.79)	152	5.02 (1.24)	83	4.42 (1.24)	80	4.38 (1.27)
Below detectable limit (n, %)	6/17	35.3%	5/36	13.9%	17/71	23.9%	81/387	20.9%
<b>Visit 23</b>								
sVEGF (including imputed)	80	4.38 (0.90)	88	4.47 (1.21)	78	4.36 (1.34)	61	4.12 (1.32)
Below detectable limit (n, %)	3/14	21.4%	5/24	20.8%	16/62	25.8%	113/374	30.2%
<b>Visit 24</b>								
sVEGF (including imputed)	77	4.35 (1.40)	91	4.51 (1.02)	78	4.36 (1.26)	61	4.11 (1.42)
Below detectable limit (n, %)	4/17	23.5%	5/27	18.5%	15/63	23.8%	118/398	29.6%

GM=geometric mean. SD=standard deviation. sVEGF= serum vascular endothelial growth factor. SSAE=systemic serious adverse event. There are 20 patients who appear in more than one SSAE category above; see Figure 4 for details. sVEGF values below 32 could not be detected; therefore values below the detectable limit were imputed from a uniform distribution between 0 and 32. SDs on the  $\log_e$  scale can be added or subtracted from the mean on the  $\log_e$  scale and exponentiated to calculate the approximate range within which 68% (1SD) or 95% (2SD) of the data points lie.