Table e-4. Comparison of time to treatment before and after awareness of marked delays in treatment administration for SE under different sensitivity analyses

SENSITIVITY ANALYSIS 1: 2011-2016 versus 2017-2019								
	2011-2016	2017-2019	Univariate analysis p-value	Multivariate analysis				
ALL PATIENTS (N=328)								
	N=237	N=91						
Time to the	17 (5-48)	15 (5-36)	0.3919	HR=0.92 (95% confidence				
first BZD	minutes	minutes		interval: 0.72-1.19), <i>p</i> =0.5210				
(N=328)								
Time to the	65 (35-150)	66 (31.5-	0.7328	HR=0.92 (95% confidence				
first non-BZD	minutes	151.5)		interval: 0.71-1.18), <i>p</i> =0.4915				
ASM (N=328) Time to the	190 (121 520)	minutes 140 (85-540)	0.2236	IID 0.05 (050)				
first	180 (121-539) minutes	140 (85-540) minutes	0.2236	HR=0.95 (95% confidence interval: 0.64-1.42), <i>p</i> =0.8149				
continuous	minutes	illillutes		mtervar. 0.04-1.42), p=0.0149				
infusion								
(N=152)								
/	SE O	NSET OUT OF	THE HOSPITAL (N=	=222)				
At least one	75/126	23/31 (74.2%)	0.1513	OR=2.60 (95% confidence				
BZD before	(59.5%)			interval: 0.98-7.51), <i>p</i> =0.0632				
hospital arrival								
(N=157)								
At least one	39/67 (58.2%)	17/18 (94.4%)	0.0042	OR=21.94 (95% confidence				
BZD before				interval: 3.19-459.88), <i>p</i> =0.0081				
hospital arrival among patients								
with a prior								
diagnosis of								
epilepsy								
(N=85)								
	N=168	N=54						
Time to the	20 (7.8-56.3)	20 (5-48.8)	0.5444	HR=1.00 (95% confidence				
first BZD	minutes	minutes		interval: 0.73-1.39), <i>p</i> =0.9792				
(N=222) Time to the	83 (45-165.2)	78.5 (51.3-	0.7463	HR=0.99 (95% confidence				
first non-BZD	minutes	206.8)	0.7403	interval: $0.72-1.37$), $p=0.951$				
ASM (N=222)	minutes	minutes		interval. 0.72-1.37), p=0.931				
Time to the	172 (123.5-	217.5 (67.5-	0.6875	HR=0.88 (95% confidence				
first	604) minutes	900) minutes		interval: $0.50-1.54$), $p=0.650$				
continuous								
infusion								
(N=101)								
			E HOSPITAL (N=10	6) 				
Time to the	N=69	N=37 9 (5-23)	0.5074	HR=0.88 (95% confidence				
Time to the first BZD	9 (4-26) minutes	9 (5-23) minutes	0.3074	interval: 0.56-1.40), p=0.597				
(N=106)	minutes	minutes		πιοιναι. 0.30-1.40 <i>j</i> , <i>μ</i> =0.39 <i>i</i>				
Time to the	42 (20-76)	31 (24-96)	0.6807	HR=1.08 (95% confidence				
first non-BZD	minutes	minutes		interval: 0.68-1.73), <i>p</i> =0.7501				
ASM (N=106)				2				
Time to the	186 (120-	135 (90.5-	0.3845	HR=1.37 (95% confidence				
first	487.5)	326.5)		interval: 0.67-2.79), <i>p</i> =0.388				
continuous	minutes	minutes						
infusion								
(N=51)	ANAT VOIC 2: C	ONGINED ON	V CENTEDO AL DE	ADV IN SCEDE DUDING 2011				
SENSITIVITY ANALYSIS 2: CONSIDER ONLY CENTERS ALREADY IN pSERG DURING 2011- 2013								
			4013					

	ORIGINA	L COMPARISO	ON 2011-2014 versus	2015-2019		
	2011-2014	2015-2019	Univariate analysis	Multivariate analysis		
		ALL PAT	ENTS (N=268)	•		
	N=150	N=118				
Time to the	20 (5-53.8)	19 (5-38)	0.3825	HR=1.07 (95% confidence		
first BZD (N=268)	minutes	minutes		interval: 0.84-1.38), <i>p</i> =0.5726		
Time to the	66.5 (34.3-	65.5 (38.5-	0.9475	HR=1.01 (95% confidence		
first non-BZD	160) minutes	149.8)		interval: 0.79-1.30), <i>p</i> =0.9113		
ASM (N=268)		minutes				
Time to the	180 (123.5-	212 (122-	0.7007	HR=0.97 (95% confidence		
first	539) minutes	660.5)		interval: 0.66-1.43), <i>p</i> =0.8922		
continuous		minutes				
infusion						
(N=119)						
			<u>THE HOSPITAL (N</u>			
At least one	40/80 (50%)	43/54 (79.6%)	0.0006	OR=5.22 (95% confidence		
BZD before				interval: 2.16-13.83), <i>p</i> =0.0005		
hospital arrival						
(N=134)	22/42 (52.49)	04/20 (75%)	0.0562	OD 2.07 (050) C.1		
At least one	22/42 (52.4%)	24/32 (75%)	0.0562	OR=3.07 (95% confidence		
BZD before				interval: 0.92-11.59), <i>p</i> =0.0783		
hospital arrival among patients						
with a prior						
diagnosis of						
epilepsy						
(N=74)						
,	N=104	N=80				
Time to the	26.5 (9.3-	20 (7-48.5)	0.3272	HR=1.02 (95% confidence		
first BZD	62.5) minutes	minutes		interval: 0.75-1.39), <i>p</i> =0.8882		
(N=184)						
Time to the	81 (45-182.5)	89.5 (59-	0.5638	HR=0.92 (95% confidence		
first non-BZD	minutes	162.8)		interval: 0.68-1.25), <i>p</i> =0.594		
ASM (N=184)		minutes				
Time to the	175 (134-	212 (121.5-	0.9893	HR=1.19 (95% confidence		
first	634.5)	660.5)		interval: 0.74-1.93), <i>p</i> =0.4771		
continuous	minutes	minutes				
infusion (N=84)						
(11-04)	C .	F ONSET IN TE	L IE HOSPITAL (N=8	24)		
	N=46	N=38	I HOSTITAL (N-C	/*/ 		
Time to the	8 (4.3-25.8)	11 (4-23.8)	0.9427	HR=1.23 (95% confidence		
first BZD	minutes	minutes		interval: $0.77-1.96$), $p=0.3972$		
(N=84)						
Time to the	44.5 (20.3-	32 (23-73.3)	0.6210	HR=1.20 (95% confidence		
first non-BZD	86.8) minutes	minutes		interval: 0.76-1.90), <i>p</i> =0.4431		
ASM (N=84)						
Time to the	210 (121-462)	205 (131.8-	0.5615	HR=0.88 (95% confidence		
first	minutes	535.5)		interval: 0.38-2.03), <i>p</i> =0.7604		
continuous		minutes				
infusion (N=35)						
(N=35)	1	<u> </u>	1	l		
SENSITIVITY ANALYSIS 3: CONSIDER ONLY CENTERS ALREADY IN pSERG DURING 2011- 2013						
			ersus 2017-2019			
	2011-2016	2017-2019	Univariate	Multivariate analysis		
	1		a alaia	1		
			analysis IENTS (N=268)			

	N=220	N=48					
Time to the	20 (5-49.3)	20 (5-33.5)	0.3825	HR=1.08 (95% confidence			
first BZD	minutes	minutes	0.3023	interval: 0.78-1.49), p=0.6462			
(N=268)	iiiiiides	iiiiiides		meer var. 0.70 1.45), p=0.0402			
Time to the	66.5 (37.8-	64.5 (31.8-	0.9475	HR=0.93 (95% confidence			
first non-BZD	150.8)	159.3)	0.5475	interval: $0.68-1.28$), $p=0.6454$			
ASM (N=268)	minutes	minutes		mtervar. 0.00-1.20), p=0.0434			
Time to the	186 (122-	420 (135-	0.7007	HR=0.82 (95% confidence			
first	574.8)	1000) minutes	0.7007	interval: 0.44-1.5), <i>p</i> =0.5141			
continuous	minutes	1000) illilutes		interval. 0.44-1.5), p=0.5141			
infusion	illitutes						
(N=119) SE ONSET OUT OF THE HOSPITAL (N=184)							
At least one	70/119	13/15 (86.7%)	0.0474	OR=6.70 (95% confidence			
BZD before	(58.8%)	13/13 (80.7%)	0.0474	interval: 1.50-48.48), p=0.0252			
hospital arrival	(36.6%)			interval. 1.30-48.48), p=0.0232			
(N=134)							
At least one	36/63 (57.1%)	10/11 (90.9%)	0.0437	OR=15.44 (95% confidence			
BZD before	30/03 (37.170)	10/11 (50.570)	0.0437	interval: 1.83-358.12), <i>p</i> =0.0288			
hospital arrival				mtervar. 1.03-330.12), p=0.0200			
among patients							
with a prior							
diagnosis of							
epilepsy							
(N=74)							
(11-7-1)	N=154	N=30					
Time to the			0.3272	HR=0.96 (95% confidence			
Time to the	21.5 (7.3-	25 (6.3-44.3)	0.3272	HR=0.96 (95% confidence interval: 0.64-1.46), n=0.8610			
first BZD			0.3272	HR=0.96 (95% confidence interval: 0.64-1.46), <i>p</i> =0.8610			
first BZD (N=184)	21.5 (7.3- 58.8) minutes	25 (6.3-44.3) minutes		interval: 0.64-1.46), <i>p</i> =0.8610			
first BZD (N=184) Time to the	21.5 (7.3- 58.8) minutes 88.5 (47.3-	25 (6.3-44.3) minutes 77 (58.3-	0.3272	interval: 0.64-1.46), <i>p</i> =0.8610 HR=0.88 (95% confidence			
first BZD (N=184) Time to the first non-BZD	21.5 (7.3- 58.8) minutes	25 (6.3-44.3) minutes 77 (58.3- 206.8)		interval: 0.64-1.46), <i>p</i> =0.8610			
first BZD (N=184) Time to the first non-BZD ASM (N=184)	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes	25 (6.3-44.3) minutes 77 (58.3- 206.8) minutes	0.5638	interval: 0.64-1.46), <i>p</i> =0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), <i>p</i> =0.545			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626)	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-		interval: 0.64-1.46), <i>p</i> =0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), <i>p</i> =0.545 HR=0.76 (95% confidence			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes	25 (6.3-44.3) minutes 77 (58.3- 206.8) minutes	0.5638	interval: 0.64-1.46), <i>p</i> =0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), <i>p</i> =0.545			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626)	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-	0.5638	interval: 0.64-1.46), <i>p</i> =0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), <i>p</i> =0.545 HR=0.76 (95% confidence			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626)	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-	0.5638	interval: 0.64-1.46), <i>p</i> =0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), <i>p</i> =0.545 HR=0.76 (95% confidence			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-1188) minutes	0.5638	interval: 0.64-1.46), <i>p</i> =0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), <i>p</i> =0.545 HR=0.76 (95% confidence interval: 0.33-1.74), <i>p</i> =0.5091			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-1188) minutes	0.5638	interval: 0.64-1.46), <i>p</i> =0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), <i>p</i> =0.545 HR=0.76 (95% confidence interval: 0.33-1.74), <i>p</i> =0.5091			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes	25 (6.3-44.3) minutes 77 (58.3- 206.8) minutes 600 (330- 1188) minutes	0.5638	interval: 0.64-1.46), <i>p</i> =0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), <i>p</i> =0.545 HR=0.76 (95% confidence interval: 0.33-1.74), <i>p</i> =0.5091			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84)	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-1188) minutes E ONSET IN TH N=18 12.5 (4.3-	0.5638 0.9893 E HOSPITAL (N=84	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84) Time to the	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes S N=66 9.5 (4-27.5)	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-1188) minutes E ONSET IN TH	0.5638 0.9893 E HOSPITAL (N=84	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091 HR=1.21 (95% confidence			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84) Time to the first BZD	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes Sinutes N=66 9.5 (4-27.5) minutes	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-1188) minutes E ONSET IN TH N=18 12.5 (4.3-20.8) minutes	0.5638 0.9893 E HOSPITAL (N=84	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091 HR=1.21 (95% confidence interval: 0.68-2.15), p=0.5194			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84) Time to the first BZD (N=84)	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes S N=66 9.5 (4-27.5)	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-1188) minutes E ONSET IN TH N=18 12.5 (4.3-	0.5638 0.9893 E HOSPITAL (N=84 0.9427	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091 HR=1.21 (95% confidence interval: 0.68-2.15), p=0.5194 HR=1.22 (95% confidence			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84) Time to the first BZD (N=84) Time to the first non-BZD	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes SEN=66 9.5 (4-27.5) minutes	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-1188) minutes E ONSET IN TH N=18 12.5 (4.3-20.8) minutes 30 (24-81)	0.5638 0.9893 E HOSPITAL (N=84 0.9427	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091 HR=1.21 (95% confidence interval: 0.68-2.15), p=0.5194			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84) Time to the first BZD (N=84) Time to the first non-BZD ASM (N=84)	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes SEN=66 9.5 (4-27.5) minutes 41.5 (20-75.5) minutes	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-1188) minutes E ONSET IN TH N=18 12.5 (4.3-20.8) minutes 30 (24-81) minutes	0.5638 0.9893 E HOSPITAL (N=84 0.9427	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091 HR=1.21 (95% confidence interval: 0.68-2.15), p=0.5194 HR=1.22 (95% confidence interval: 0.69-2.17), p=0.4921			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84) Time to the first BZD (N=84) Time to the first non-BZD	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes SEN=66 9.5 (4-27.5) minutes	25 (6.3-44.3) minutes 77 (58.3- 206.8) minutes 600 (330- 1188) minutes E ONSET IN TH N=18 12.5 (4.3- 20.8) minutes 30 (24-81) minutes 204.5 (146-	0.5638 0.9893 E HOSPITAL (N=84 0.9427	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091 HR=1.21 (95% confidence interval: 0.68-2.15), p=0.5194 HR=1.22 (95% confidence interval: 0.69-2.17), p=0.4921 HR=1.48 (95% confidence			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84) Time to the first BZD (N=84) Time to the first non-BZD ASM (N=84) Time to the	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes Sinutes N=66 9.5 (4-27.5) minutes 41.5 (20-75.5) minutes	25 (6.3-44.3) minutes 77 (58.3-206.8) minutes 600 (330-1188) minutes E ONSET IN TH N=18 12.5 (4.3-20.8) minutes 30 (24-81) minutes	0.5638 0.9893 E HOSPITAL (N=84 0.9427	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091 HR=1.21 (95% confidence interval: 0.68-2.15), p=0.5194 HR=1.22 (95% confidence interval: 0.69-2.17), p=0.4921			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84) Time to the first BZD (N=84) Time to the first non-BZD ASM (N=84) Time to the first non-BZD ASM (N=84)	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes Sinutes N=66 9.5 (4-27.5) minutes 41.5 (20-75.5) minutes	25 (6.3-44.3) minutes 77 (58.3- 206.8) minutes 600 (330- 1188) minutes E ONSET IN TH N=18 12.5 (4.3- 20.8) minutes 30 (24-81) minutes 204.5 (146- 372.5)	0.5638 0.9893 E HOSPITAL (N=84 0.9427	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091 HR=1.21 (95% confidence interval: 0.68-2.15), p=0.5194 HR=1.22 (95% confidence interval: 0.69-2.17), p=0.4921 HR=1.48 (95% confidence			
first BZD (N=184) Time to the first non-BZD ASM (N=184) Time to the first continuous infusion (N=84) Time to the first BZD (N=84) Time to the first non-BZD ASM (N=84) Time to the first non-BZD ASM (N=84)	21.5 (7.3- 58.8) minutes 88.5 (47.3- 169) minutes 180 (125-626) minutes Sinutes N=66 9.5 (4-27.5) minutes 41.5 (20-75.5) minutes	25 (6.3-44.3) minutes 77 (58.3- 206.8) minutes 600 (330- 1188) minutes E ONSET IN TH N=18 12.5 (4.3- 20.8) minutes 30 (24-81) minutes 204.5 (146- 372.5)	0.5638 0.9893 E HOSPITAL (N=84 0.9427	interval: 0.64-1.46), p=0.8610 HR=0.88 (95% confidence interval: 0.58-1.33), p=0.545 HR=0.76 (95% confidence interval: 0.33-1.74), p=0.5091 HR=1.21 (95% confidence interval: 0.68-2.15), p=0.5194 HR=1.22 (95% confidence interval: 0.69-2.17), p=0.4921 HR=1.48 (95% confidence			

Legend: BZD: Benzodiazepine. **N:** Number of patients. **NA:** Not applicable. **Non-BZD ASM:** Non-benzodiazepine antiseizure medication. **RMST:** Restricted mean survival times. **SE:** Status epilepticus.