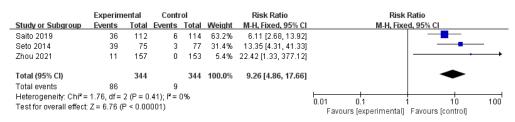
	Erlotinib plus bevaciz	umab	Erloti	nib		Risk Ratio		Risk	Ratio	
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl		M-H, Fixe	ed, 95% Cl	
Saito 2019	53	112	47	114	31.4%	1.15 [0.86, 1.54]		-	-	
Seto 2014	61	75	60	77	39.9%	1.04 [0.89, 1.23]			•	
Zhou 2021	62	157	42	153	28.7%	1.44 [1.04, 1.99]				
Total (95% CI)		344		344	100.0%	1.19 [1.03, 1.38]			•	
Total events	176		149							
Heterogeneity: Chi ² =	3.93, df = 2 (P = 0.14); l	²= 49%					0.01	0.1	 1 10	100
Test for overall effect:	Z = 2.29 (P = 0.02)								Favours [control]	100

FigureS1 Forest plot of AEs of diarrhea

	Experim	ental	Contr	ol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Zhou 2021	56	157	45	153	38.6%	1.21 [0.88, 1.68]	
Seto 2014	54	75	22	77	37.8%	2.52 [1.72, 3.69]	
Saito 2019	31	112	3	114	23.6%	10.52 [3.31, 33.42]	
Total (95% CI)		344		344	100.0%	2.66 [1.12, 6.31]	-
Total events	141		70				
Heterogeneity: Tau ² =	= 0.47; Chi ^a	² = 18.60	3, df = 2 (P < 0.0	001); I ^z =	89%	
Test for overall effect	: Z = 2.22 (F	P = 0.03)				0.01 0.1 1 10 100 Favours [experimental] Favours [control]

FigureS2 Forest plot of AEs of haemorrhagic event



FigureS3 Forest plot of AEs of proteinuria

	Experim	ental	Contr	rol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl
Saito 2019	52	112	11	114	42.2%	4.81 [2.65, 8.73]	_ _
Seto 2014	57	75	10	77	38.2%	5.85 [3.24, 10.58]	_ _
Zhou 2021	29	157	5	153	19.6%	5.65 [2.25, 14.22]	·
Total (95% CI)		344		344	100.0%	5.37 [3.66, 7.88]	•
Total events	138		26				
Heterogeneity: Chi ² =	0.22, df=	2 (P = 0	.89); I ² =	0%			
Test for overall effect	Z = 8.60 (f	P < 0.00	1001)				Favours [experimental] Favours [control]

FigureS4 Forest plot of AEs of hypertension

	Erlotinib plus bevaci	umab	Erlotii	nib		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
1.10.1 rash							
Saito 2019	98	112	99	114	30.6%	1.01 [0.91, 1.11]	•
Seto 2014	74	75	76	77	46.7%	1.00 [0.96, 1.04]	•
Zhou 2021	108	157	117	153	22.7%	0.90 [0.78, 1.03]	-
Subtotal (95% CI)		344		344	100.0%	0.98 [0.90, 1.07]	•
Total events	280		292				
Heterogeneity: Tau ² =	= 0.00; Chi ² = 6.02, df =	2 (P = 0.0	05); I ² = 61	7%			
Test for overall effect	Z = 0.49 (P = 0.63)						
Total (95% CI)		344		344	100.0%	0.98 [0.90, 1.07]	•
Total events	280		292				
Heterogeneity: Tau ² =	= 0.00; Chi ² = 6.02, df =	2 (P = 0.0	05); l ² = 6	7%			
Test for overall effect	Z = 0.49 (P = 0.63)	•					0.01 0.1 1 10 100
Test for subaroup dif	ferences: Not applicabl	е					Favours [experimental] Favours [control]

FigureS5 Forest plot of AEs of rash

	Experim	ental	Contr	ol		Risk Ratio	Risk Ratio	
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl	
Saito 2019	17	112	18	114	18.5%	0.96 [0.52, 1.77]		
Seto 2014	57	75	50	77	51.1%	1.17 [0.95, 1.44]		
Zhou 2021	28	157	29	153	30.4%	0.94 [0.59, 1.50]		
Total (95% CI)		344		344	100.0%	1.06 [0.87, 1.30]	•	
Total events	102		97					
Heterogeneity: Chi ² =	1.20, df=	2 (P = 0	.55); l² =	0%				100
Test for overall effect:	Z = 0.58 (F	^o = 0.57)				Favours [experimental] Favours [control]	100

FigureS6 Forest plot of AEs of paronychia

	Experim	ental	Contr	ol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Saito 2019	23	112	12	114	23.3%	1.95 [1.02, 3.73]	-
Seto 2014	47	75	46	77	49.8%	1.05 [0.81, 1.35]	÷
Zhou 2021	25	157	17	153	26.9%	1.43 [0.81, 2.55]	
Total (95% CI)		344		344	100.0%	1.32 [0.89, 1.96]	•
Total events	95		75				
Heterogeneity: Tau ² =	= 0.06; Chi ^a	= 4.17,	df = 2 (P	= 0.12)); I ² = 52%	6	
Test for overall effect	Z=1.37 (P = 0.17)				0.01 0.1 1 10 100 Favours [experimental] Favours [control]

FigureS7 Forest plot of AEs of stomatitis

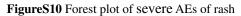
	Experim	ental	Contr	rol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl
Saito 2019	8	112	1	114	40.0%	8.14 [1.04, 64.04]	
Seto 2014	6	75	0	77	19.9%	13.34 [0.76, 232.75]	
Stinchcombe 2019	5	43	0	45	19.7%	11.50 [0.66, 201.90]	
Zhou 2021	11	157	0	153	20.4%	22.42 [1.33, 377.12]	
Total (95% CI)		387		389	100.0%	12.75 [3.54, 45.97]	
Total events	30		1				
Heterogeneity: Chi ² =	0.34, df=	3 (P = 0	.95); I ^z =	0%			
Test for overall effect	: Z = 3.89 (I	P < 0.00	01)				0.01 0.1 1 10 100 Favours [experimental] Favours [control]

FigureS8 Forest plot of severe AEs of proteinuria

	Experim	ental	Contr	ol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Saito 2019	26	112	1	114	12.4%	26.46 [3.65, 191.71]	
Seto 2014	45	75	8	77	30.7%	5.78 [2.92, 11.42]	
Stinchcombe 2019	17	43	9	45	30.5%	1.98 [0.99, 3.95]	
Zhou 2021	29	157	5	153	26.4%	5.65 [2.25, 14.22]	
Total (95% Cl)		387		389	100.0%	5.00 [2.14, 11.68]	-
Total events	117		23				
Heterogeneity: Tau ² =	= 0.49; Chi ^a	² = 10.20), df = 3 (l	• = 0.0	2); I ² = 71	%	
Test for overall effect	Z = 3.72 (P = 0.00	02)				0.01 0.1 1 10 100 Favours [experimental] Favours [control]

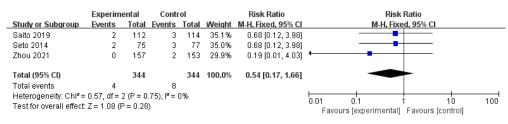
FigureS9 Forest plot of severe AEs of hypertension

	Erlotinib plus bevac	izumab	Erlotir	nib		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl
Saito 2019	23	112	24	114	48.0%	0.98 [0.59, 1.62]	
Seto 2014	19	75	15	77	29.9%	1.30 [0.72, 2.36]	
Stinchcombe 2019	4	43	6	45	11.8%	0.70 [0.21, 2.30]	
Zhou 2021	8	157	5	153	10.2%	1.56 [0.52, 4.66]	
Total (95% CI)		387		389	100.0%	1.10 [0.78, 1.56]	•
Total events	54		50				
Heterogeneity: Chi ² =	= 1.46, df = 3 (P = 0.69);	I ² = 0%					
Test for overall effect	Z = 0.53 (P = 0.59)						0.01 0.1 1 10 100 Favours [experimental] Favours [control]



	Erlotinib plus bevaciz	umab	Erlotir	nib		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl
Saito 2019	6	112	2	114	21.2%	3.05 [0.63, 14.81]	
Seto 2014	1	75	1	77	10.6%	1.03 [0.07, 16.12]	
Stinchcombe 2019	4	43	6	45	62.8%	0.70 [0.21, 2.30]	
Zhou 2021	4	157	0	153	5.4%	8.77 [0.48, 161.56]	
Total (95% CI)		387		389	100.0%	1.67 [0.76, 3.68]	-
Total events	15		9				
Heterogeneity: Chi ² =	3.98, df = 3 (P = 0.26); l	² = 25%					
Test for overall effect:	Z = 1.27 (P = 0.20)						Favours [experimental] Favours [control]

FigureS11 Forest plot of severe AEs of diarrhea



FigureS12 Forest plot of severe AEs of paronychia

	Experim	ental	Cont	rol		Risk Ratio	Risk Ratio	
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl	
Saito 2019	1	112	1	114	24.9%	1.02 [0.06, 16.07]	ŧ	
Seto 2014	1	75	2	77	49.6%	0.51 [0.05, 5.54]		
Zhou 2021	1	157	1	153	25.5%	0.97 [0.06, 15.44]		
Total (95% CI)		344		344	100.0%	0.76 [0.17, 3.36]		
Total events	3		4					
Heterogeneity: Chi ² =	0.18, df=	2 (P = 0	.91); I² =	0%				100
Test for overall effect	: Z = 0.37 (P = 0.71)				0.01 0.1 1 10 1 Favours [experimental] Favours [control]	00

FigureS13 Forest plot of severe AEs of stomatitis

	Experim	ental	Contr	rol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl
Saito 2019	2	112	1	114	49.8%	2.04 [0.19, 22.13]	
Seto 2014	2	75	0	77	24.8%	5.13 [0.25, 105.14]	
Zhou 2021	2	157	0	153	25.4%	4.87 [0.24, 100.69]	
Total (95% CI)		344		344	100.0%	3.52 [0.74, 16.87]	
Total events	6		1				
Heterogeneity: Chi ² =	= 0.31, df =	2 (P = 0	.86); I ^z =	0%			
Test for overall effect	: Z = 1.58 (I	P = 0.11)				0.01 0.1 1 10 100 Favours [experimental] Favours [control]

FigureS14 Forest plot of severe AEs of haemorrhagic event