Supplementary Information No 6 Meta-analyses of Anakinra versus control

	Anaki	nra	Cont	ol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% CI	M-H, Random, 95% CI
Declercq 2021	10	43	9	72	100.0%	1.86 [0.82, 4.21]	
Total (95% CI)		43		72	100.0%	1.86 [0.82, 4.21]	
Total events	10		9				
Heterogeneity: Not ap Test for overall effect	•	(P = 0.1	4)				0.2 0.5 1 2 5 Favours Anakinra Favours control

Fig. 1 Forest plot describing the difference between Anakinra compared to placebo or standard care alone regarding 60-day mortality. M-H = Mantel-Haenszel.

Study or Subgroup	log[Hazard Ratio]	SE	Anakinra Total		Weight	Hazard Ratio IV, Random, 95% CI	Hazard Ratio
Kyriazopoulou 2021	-0.7985		405	189	55.3%	0.45 [0.21, 0.96]	
Tharaux 2021	-0.2614	0.4323	59	55	44.7%	0.77 [0.33, 1.80]	i — -
Total (95% CI)			464	244	100.0%	0.57 [0.32, 1.01]	
Heterogeneity: Tau ² = Test for overall effect:		= 1 (P = 0).36); I² = 0%	6			0.1 0.2 0.5 1 2 5 10 Favours Anakinra Favours control

Fig. 2 Forest plot describing the difference between Anakinra compared to placebo or standard care alone regarding 28-day mortality, time-to-event. M-H = Mantel-Haenszel.

Study or Subgroup	log[Hazard Ratio]	SE	Anakinra Total		Woight	Hazard Ratio IV, Random, 95% Cl	Hazard Ratio IV. Random, 95% Cl
study of Subgroup	ισχίμαζαι η καιοί	3E	TULAI	TULAI	weight	IV, Rahuom, 95% Ci	IV, Kaliuolii, 95% Ci
Derde 2021	0	0	0	0		Not estimable	
Tharaux 2021	-0.0305	0.3807	59	55	100.0%	0.97 [0.46, 2.05]	
Total (95% CI)			59	55	100.0%	0.97 [0.46, 2.05]	
Heterogeneity: Not a	pplicable						
Test for overall effect	Z = 0.08 (P = 0.94)						0.5 0.7 1 1.5 2 Favours Anakinra Favours control

Fig. 3 Forest plot describing the difference between Anakinra compared to placebo or standard care alone regarding 90-day mortality, time-to-event. M-H = Mantel-Haenszel.

	Anakii	nra	Contr	ol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% CI	M-H, Random, 95% Cl
Declercq 2021	10	44	9	74	22.9%	1.87 [0.82, 4.24]	
Derde 2021	145	365	150	406	77.1%	1.08 [0.90, 1.29]	
Total (95% CI)		409		480	100.0%	1.22 [0.77, 1.92]	-
Total events	155		159				
Heterogeneity: Tau ² =	= 0.06; Chi	i ^z = 1.6	7, df = 1 (P = 0.2	0); I ^z = 40	%	
Test for overall effect	: Z = 0.86 ((P = 0.3	9)				Favours Anakinra Favours control

Fig. 4 Forest plot describing the difference between Anakinra compared to placebo or standard care alone regarding mortality at hospital discharge. M-H = Mantel-Haenszel.

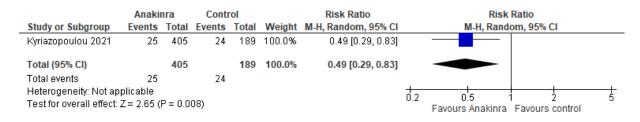


Fig. 5 Forest plot describing the difference between Anakinra compared to placebo or standard care alone regarding worsening of clinical status in patients with suPAR ≥ 6 ng/ml. M-H = Mantel-Haenszel.

	Anaki	nra	Cont	rol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Declercq 2021	12	43	14	72	22.7%	1.44 [0.73, 2.81]	
Kyriazopoulou 2021	65	405	41	189	41.7%	0.74 [0.52, 1.05]	— —— — —
Tharaux 2021	27	59	21	55	35.6%	1.20 [0.77, 1.85]	
Total (95% CI)		507		316	100.0%	1.02 [0.68, 1.53]	-
Total events	104		76				
Heterogeneity: Tau ² =	0.07; Chi ^a	²= 4.52	2, df = 2 (l	P = 0.10	0); I ² = 56'	%	
Test for overall effect:	Z = 0.10 (P = 0.9	2)				0.2 0.5 1 2 5 Favours Anakinra Favours control

Fig. 6 Forest plot describing the difference between Anakinra compared to placebo or standard care alone regarding serious adverse events. M-H = Mantel-Haenszel.

	Anaki	nra	Contr	rol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% Cl	M-H, Random, 95% Cl
Declercq 2021	26	43	43	72	63.5%	1.01 [0.74, 1.38]	
Tharaux 2021	29	59	23	55	36.5%	1.18 [0.78, 1.76]	
Total (95% CI)		102		127	100.0%	1.07 [0.84, 1.37]	
Total events	55		66				
Heterogeneity: Tau ² =	: 0.00; Ch	i ² = 0.3	4, df = 1 ((P = 0.5	6); I ² = 09	6 –	0.7 0.85 1 1.2 1.5
Test for overall effect:	Z = 0.53 ((P = 0.5	i9)				0.7 0.85 1 1.2 1.5 Favours Anakinra Favours control

Fig. 7 Forest plot describing the difference between Anakinra compared to placebo or standard care alone regarding any adverse events. M-H = Mantel-Haenszel.

	Anakii	nra	Contr	ol		Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% CI	M-H, Random, 95% Cl
Declercq 2021	16	43	24	72	100.0%	1.12 [0.67, 1.85]	
Total (95% CI)		43		72	100.0%	1.12 [0.67, 1.85]	
Total events	16		24				
Heterogeneity: Not a	pplicable						
Test for overall effect	: Z = 0.42 ((P = 0.6	67)				Favours Anakinra Favours control

Fig. 8 Forest plot describing the difference between Anakinra compared to placebo or standard care alone regarding adverse events, grade 3-4. M-H = Mantel-Haenszel.