Monoclonal antibodies in prevention and early treatment of COVID-19 in lung transplant recipients: a systematic review and perspective on the role of monoclonal antibodies in the future

Online supplement

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1. Search strategy

Search terms used on 8th February 2023:

PubMed/MEDLINE

(("Lung Transplantation" [Mesh] OR lungtransplant*[tiab] OR ((lung OR pulmon*)AND transplant*[tiab]) OR ((lung OR pulmon*)AND allotransplant* [tiab]) OR (lung tissue AND transplant*[tiab])) OR ("Organ Transplantation" [Mesh] OR organ transplant* [tiab] OR organ graft* [tiab] OR (solid AND organ AND transplant* [tiab]) OR SOT[tiab] OR organ allotransplant* [tiab])

AND

(Bamlanivimab[tiab] OR (Bamlanivimab AND etesevimab[tiab]) OR Regdanvimab[tiab] OR Regkirona[tiab] OR Ronapreve[tiab] OR (Casirivimab AND imdevimab[tiab]) OR Sotrovimab[tiab] OR Xevudy[tiab] OR Bebtelovimab[tiab] OR Evusheld[tiab] OR (Tixagevimab AND cilgavimab[tiab]) OR Tocilizumab[tiab] OR Sarilumab[tiab] OR Kevzara[tiab])

Embase

(('lung transplantation'/exp OR 'lung transplant*':ti,ab,kw OR 'pulmonary transplant*':ti,ab,kw OR 'lung allotransplant*':ti,ab,kw OR 'pulmonary allotransplant*':ti,ab,kw OR 'pulmonary allograft*':ti,ab,kw) OR ('organ transplantation'/exp OR 'organ transplant*':ti,ab,kw OR 'organ graft*':ti,ab,kw OR 'solid organ transplant*':ti,ab,kw OR 'SOT':ti,ab,kw OR 'organ allotransplant*':ti,ab,kw))

AND

('Bamlanivimab':ti,ab,kw OR '(Bamlanivimab AND etesevimab)':ti,ab,kw OR 'Regdanvimab':ti,ab,kw OR 'Regkirona':ti,ab,kw OR 'Regdanvimab':ti,ab,kw OR 'Regkirona':ti,ab,kw OR 'Regdanvimab':ti,ab,kw OR 'Casirivimab AND imdevimab)':ti,ab,kw OR 'Sotrovimab':ti,ab,kw OR 'Evusheld':ti,ab,kw OR 'Tocilizumab':ti,ab,kw OR 'Sarilumab':ti,ab,kw OR 'Kevzara':ti,ab,kw)

Cochrane Controlled Trials Register (CENTRAL/CCTR)

(([mh "Lung Transplantation"] OR (lungtransplant* OR ((lung OR pulmon*) AND transplant*) OR ((lung OR pulmon*) allotransplant*) OR (lung tissue AND transplant*) OR ((lung OR pulmon*) allograft)):ti,ab,kw) OR ([mh "Organ Transplantation"] OR (organ transplant* OR organ graft* OR (solid AND organ AND transplant*) OR SOT OR organ allotransplant*):ti,ab,kw))

AND

(("Bamlanivimab" OR "(Bamlanivimab AND etesevimab)" OR "Regdanvimab" OR "Regkirona" OR "Ronapreve" OR "(Casirivimab AND imdevimab)" OR "Sotrovimab" OR "Xevudy" OR "Bebtelovimab" OR "Evusheld" OR "(Tixagevimab AND cilgavimab)" OR "Tocilizumab" OR "Sarilumab" OR "Kevzara"):ti,ab,kw)

Search results for tocilizumab (a non-COVID-specific monoclonal antibody) were not included in the final manuscript.

2. Summary of findings tables

Tixagevimab-cilgavimab

Lung transplant-specific outcomes

Bibliography:

- a) Jurdi AA, Morena L, Cote M, et al. Tixagevimab/Cilgavimab pre-exposure prophylaxis is associated with lower breakthrough infection risk in vaccinated solid organ transplant recipients during the Omicron Wave. 2022;22(12):3130-36.
- b) Nguyen Y, Flahault A, Chavarot N, et al. Pre-exposure prophylaxis with tixagevimab and cilgavimab (Evusheld) for covid-19 among 1112 severely immunocompromised patients. Clinical Microbiology and Infection. 2022;28(12).

Quality assessment			N° of patients			Effect		
N° of studies	Study design	Risk of bias	Intervention	No intervention	Relative	Absolute		
Overall mor	rtality			intervention				
2 ^{a,b}	<i>-</i>	1	442	70*	h.a			
24,5	Observational	Intermediate	113	70*	Mortality was 0%	in both groups in the matched control study (0/77 vs. 0/70) and also 0% (0/36) in the other study.		
COVID-19-re	elated mortality	,						
2 ^{a,b}	Observational	Intermediate	113	70*	COVID-19-related	mortality was 0% in both groups in the matched control study (0/77 vs. 0/70) and also 0% (0/36) in		
					the other study.			
Breakthroug	gh COVID-19 inf	ection	•					
2 ^{a,b}	Observational	Low	113	70*	Breakthrough COVID-19 infection was 8% (3/36) in the study by Nguyen et al. and significantly lower in the patients who received tixagevimab-cilgavimab in the matched control study (6/77 or 8% vs. 16/70 or 23%, p=0.010). Lower dose tixagevimab-cilgavimab was associated with higher breakthrough infection for SOTR (150/150 mg vs 300/300 mg) (p=0.025).			
Hospital adı	mission							
1 ^a	Observational	Intermediate	77	70*	1/77 (1%) treated	with 150/150 mg single dose vs. 1/70 (1%) control LTR.		
ICU admissi	on							
-	-	-		-		•		
Additional r	espiratory supp	ort (HFNC, NIV,	MV)					
1 ^b	Observational	Intermediate	36		0% (0/36) received non-invasive ventilation.			
Incidence ve	enous thromboo	embolism						
-	-	-		-		-		
Incidence se	econdary infecti	ons						
-	-	-		-		-		

Incidence re	Incidence renal impairment										
-	-	-	-	-							
Long-term lu	Long-term lung function data										
-	-	-	-	-							

- a) Retrospective multicenter matched cohort study including 222 SOTR (77 LTR) who received tixagevimab-cilgavimab as pre-exposure prophylaxis compared to 222 matched controls (with 70 LTR). COVID-19 variant: Omicron B.1.1.529, BA.2 and BA.2.12.1.
- b) Retrospective, multicenter study of 1112 immunocompromised patients, including 36 LTR, received tixagevimab-cilgavimab as pre-exposure prophylaxis. COVID-19 variant: Omicron BA.1 and BA.2.

- a) Alejo JL, Kim JD, Chiang TP, et al. Patient-reported outcomes after Tixagevimab and Cilgavimab Pre-exposure prophylaxis among solid organ transplant recipients: Safety, effectiveness, and perceptions of risk. Clinical Transplantation. 2023;37(4).
- b) Cochran W, Salto-Alejandre S, Barker L, et al. Covid-19 outcomes in solid organ transplant recipients who received Tixagevimab-cilgavimab prophylaxis and/or Bebtelovimab treatment in a nurse-driven monoclonal antibody program during the Omicron Surge. Transplantation. 2022;107(2).

	Quality assessment		N° of patients			Effect		
N° of	Study design	Risk of bias	Intervention	No	Relative	Absolute		
studies				intervention				
Overall mor	rtality							
1 ^{a,b}	Observational	Intermediate			a) 0/392 (0%). b) 2/205 (1%) (7% (1/14) in SOTR treated with 150/150 mg, 0.6% (1/156) in SOTR with 300/300 mg and none (0/35) in 150/150 mg double dose).			
COVID-19-re	elated mortality							
1 ^{a,b}	Observational	Intermediate	597		0% (0/392) and 2,	/205 (1%).		
Breakthrou	gh COVID-19 inf	ection						
1 ^{a,b}	Observational	Low	597			reakthrough COVID-19 infection was 29% (4/14) in SOTR treated with single dose 150/150 mg, 8% with 300/300 mg and none (0/35) in the cohort of double 150/150 mg dose).		
Hospital ad	mission	•						
1 ^{a,b}	Observational	Intermediate	597		a) 2/392 (0.5%). b) 3/205 (1%) (tixagevimab-cilgavimab 150/150mg: 1 (7%), double 150/150mg: 0 (0%), single 300/300 mg: 2 (1%)).			
ICU admissi 1 ^a	Observational	Intermediate	392		0/392 (0%).			

^{*} Study of Jurdy et al. matched cohort with 222 control SOTR including 70 LTR.

Additional r	Additional respiratory support (HFNC, NIV, MV)									
1 ^a	Observational	Intermediate	392	0/392 (0%) (NIV).						
Incidence ve	nous thromboe	mbolism								
-	-	-	-	-						
Incidence se	condary infection	ons								
-	-	-	-	-						
Incidence re	nal impairment									
-	-	-	-	-						
Long-term lu	Long-term lung function data									
-	-	-	-	-						

a) Nationwide prospective cohort study of 392 SOTR, including 54 LTR. COVID-19 variant: Omicron BA.2 and BA.4/5.

b) Retrospective single-center study of 350 SOTR, of whom 205 SOTR were treated with tixagevimab-cilgavimab. COVID-19 variant: Omicron BA.2 and BA.5. COVID-19: coronavirus 2019 disease, HFNC: high-flow nasale canula, MV: mechanical ventilation, NIV: non-invasive ventilation.

Sotrovimab

Lung transplant-specific outcomes

Bibliography:

a) Malahe SR, Hoek RA, Dalm VA, Broers AE, et al. Clinical characteristics and outcomes of immunocompromised patients with coronavirus disease 2019 caused by the Omicron variant: A prospective, observational study. Clinical Infectious Diseases. 2022;76(3).

	Quality assessment		N° of patients		Effect		
N° of	Study design	Risk of bias	Intervention	No	Relative	Absolute	
studies				intervention			
Overall me	ortality						
1 ^a	Observational	Intermediate	14	16	Before sotrovimab outpatient treatment, outpatient treatment.	1/16* (6%) LTR died versus 0/14 (0%) after implementation of sotrovimab	
COVID-19	related mortalit	у					
1 ^a	Observational	Intermediate	14	16	Before sotrovimab outpatient treatment, outpatient treatment.	1/16* (6%) LTR died versus 0/14 (0%) after implementation of sotrovimab	
Hospital a	dmission						
1 ^a	Observational	Intermediate	14	16	Before sotrovimab administration in outp outpatient sotrovimab with 1 LTR requiring	atient therapy, 69% of LTR (11/16) required hospital admission. 14 LTR received ng hospital admission (7%) (p= <0.001).	
ICU admis	sion						
-	-	-	-	-		-	
Additiona	respiratory sup	port (HFNC, N	IV, MV)				
1 ^a	Observational	Intermediate	NA	16	Before sotrovimab outpatient therapy, 4/ or high-flow nasal oxygen therapy. 0% ned	11 (36%) who were hospitalized (25% of all 16 LTR*) required at least 15 L/min eded mechanical ventilation.	
Incidence	venous thrombo	embolism					
-	-	-	-	-		-	
Incidence	secondary infec	tions					
-	-	-	-	-		-	
Incidence	renal impairmer	nt					
-	-	-	-	-		-	
Long-tern	n lung function o	lata					
-	-	-	-	-		-	

a) Prospective single-center cohort study with 114 immunocompromised patients including 16 LTR. Sotrovimab was initially used for patients requiring hospital admission and later in 14 outpatient LTR immediately upon COVID-19 diagnosis. * It is unclear how many of the 16 LTR, of whom 11 required hospitalization, were treated with sotrovimab upon hospitalization. COVID variant: 46% Omicron (mainly BA.1).

- a) Cochran W, Langlee J, Barker L, et al. Short-term outcomes in a nurse coordinator—led and nurse practitioner—led Sotrovimab initiative for solid organ transplant recipients during the omicron surge. Transplantation. 2022;106(9).
- b) Hedvat J, Lange NW, Salerno DM, et al. COVID-19 therapeutics and outcomes among solid organ transplant recipients during the Omicron BA.1 era. Am J Transplant. 2022;22:2682-88.
- c) Rasmussen LD, Lebech A-M, Øvrehus A, et al. Experience with sotrovimab treatment of SARS-CoV-2-infected patients in Denmark. Br J Clin Pharmacol. 2023;89(6):1820-33.
- d) Solera JT, Árbol BG, Alshahrani A, et al. Impact of vaccination and early monoclonal antibody therapy on coronavirus disease 2019 outcomes in organ transplant recipients during the Omicron Wave. Clinical Infectious Diseases. 2022;75(12):2193-200.
- e) Yetmar ZA, Beam E, O'Horo JC, et al. Outcomes of bebtelovimab and sotrovimab treatment of solid organ transplant recipients with mild-to-moderate Coronavirus disease 2019 during the omicron epoch. Transplant Infectious Disease. 2022;24(4).

	Quality assessm	ent	N° of patients		Effect		
N° of studies	Study design	Risk of bias	Intervention	No intervention	Relative	Absolute	
Overall mo	ortality		•	•	1		
5ª-e	Observational	Intermediate	1314	262*	Two studies with control groups reported reduced mortality in the sotrovimab group (0/51 (0%) vs. 3/75 (4%) and 0/106 vs. 12/187 (6%)). The other studies also had a low mortality rate (0/88 (0%), 9/800 (1%) and 2/269 (0.7%)). Hedvat et al. reported an association with lower risk of 30-day hospitalization and death and sotrovimab (aRR 0.15, CI95% 0.05–0.47). Rasmussen et al. reported that a delay in time of sotrovimab admission (≤ 3 vs > 3 days after testing positive) was associated with risk of death (multivariate HR: 4.88, CI95%: 1.27-18.73).		
COVID-19-	related mortality	1	•	•	<u>, , , , , , , , , , , , , , , , , , , </u>		
3 ^{a,b,d}	Observational	Intermediate	245	75**	-	in the sotrovimab group (0/51, 0%) than the control group (3/75, 4%) in the study by Hedvat other two studies, no patients treated with sotrovimab died due to COVID-19 (0/88 and	
Hospital ac	lmission		•	•	,		
5 ^{a-e}		Intermediate	1314	262*	(17/106 (16%) v. day hospitalizati hospitalization o lower risk of 30- Hospital admissi	the control groups reported a lower incidence of hospitalization in the sotrovimab group s. 52/187 (28%), RR 0.58 (95%CI 0.35-0.94, p=0.33), and 6/51 (12%) vs. 25/75 (33%) for 30-on or death from any cause (p=0.009) or 5/51 (10%) vs. 23/75 (31%) for 30-day or death from COVID-19 (p=0.007)). In the latter study, sotrovimab was associated with a day hospitalization or death (aRR 0.15, CI95% 0.05–0.47). on ranged from 3% to 23% in the other studies (8/269, 9/88 and 183/800). or ted median stay of 4 days (IQR 3-7) vs. 7 days in control SOTR (IQR 5-16) (p=0.002).	
ICU admiss	sion	•	•	•			

Observational	Intermediate	375?	187***	No SOTR treated with sotrovimab (0/106) were admitted to ICU vs. 10% (18/187) in the control group in
				Solera et al. ICU admission was 1% (3/269) in Yetmar et al.
I respiratory supp	ort (HFNC, NI	V, MV)	•	
Observational	Intermediate	245	262*	In the studies with control groups, no SOTR treated with sotrovimab required mechanical ventilation vs. 5-8% in the controls (0/51 (0%) vs. 4/75 (5%) and 0/106 vs. 14/187 (8%)). In the other study, 0/88 (0%) required high-flow oxygen therapy, non-invasive ventilation or mechanical ventilation. Solera et al. reported a RR 0.24 (95% CI, 0.1 to 0.59) of sotrovimab for supplemental oxygen treatment. NNT preventing supplemental oxygen treatment of 6.64 (95% CI, 4.56 to 13.66). Significant after adjusting age/transplant type/number of vaccines and comorbidities (p=0.026).
venous thrombo	embolism			
Observational	Intermediate	106	187***	1% (2/187) SOTR without sotrovimab suffered a pulmonary embolism.
secondary infect	ions	•	•	
Observational	Intermediate	106	187***	8/106 (8%) in sotrovimab group vs. 28/187 (15%) in control group (7% of SOTR with sotrovimab had a bacterial pneumonia, 1% a COVID-associated pulmonary aspergillosis and 0% a bloodstream infection vs. 9% (p=0.51), 2% (p=1) and 4%, respectively, in control SOTR).
renal impairmen	t			
Observational	Intermediate	157	262	10% of SOTR with sotrovimab suffered from acute kidney injury (AKI) in Hedvat et al. compared to 28% in SOTR without sotrovimab treatment (p=0.17). Solera et al. reported 13% of AKI in the sotrovimab group vs 21% in controls (p=0.12).
m lung function d	ata			
-	-	-	-	-
	Observational venous thrombo Observational secondary infect Observational renal impairmen Observational	Observational Intermediate venous thromboembolism Observational Intermediate secondary infections Observational Intermediate renal impairment	Observational Intermediate 106 secondary infections Observational Intermediate 106 renal impairment Observational Intermediate 157	Al respiratory support (HFNC, NIV, MV) Observational Intermediate 245 262* Evenous thromboembolism Observational Intermediate 106 187*** Evecondary infections Observational Intermediate 106 187*** Observational Intermediate 106 262*

- a) Retrospective single-center cohort study with 154 SOTR, of whom 24 LTR. 51 SOTR were treated with sotrovimab (including 4 LTR) upon COVID-19 diagnosis, 28 SOTR with nirmatrelvir/ritonavir and 75 SOTR received no SARS-CoV-2-specific treatment. COVID-19 variant: Omicron BA.1.
- b) Retrospective single-center cohort-study with 88 SOTR including 18 LTR. COVID-19 variant: Omicron BA.1.
- c) Prospective single-center cohort study with 300 SOTR, of whom 62 LTR. 106 SOTR were treated with sotrovimab (including 34 LTR). COVID-19 variant: Omicron BA.1.
- d) Nationwide, population-based cohort study with 2933 high-risk patients including 800 SOTR (with 49 LTR and 2 heart-lung transplants). Sotrovimab was mainly given as outpatient treatment (88% of SOTR). COVID-19 variant: majority of all patients had Omicron BA.2 (54%), followed by Omicron BA.1 (13%) and Delta (10%).
- e) Retrospective multicenter cohort study with 361 SOTR, including 21 LTR. 269 SOTR (17 LTR) were treated with sotrovimab. COVID-19 variant: Omicron B.1.1.527 and BA.2.

^{*} Studies of Hedvat et al. (n=75) and Solera et al. (n=187)

^{**} Study of Hedvat et al. including 75 SOTR without additional treatment

^{***} Study of Solera et al. including cohort of 187 SOTR

Casirivimab-imdevimab

- a) Sarrell BA, Bloch K, El Chediak A, et al. Monoclonal antibody treatment for Covid-19 in solid organ transplant recipients. Transplant Infectious Disease. 2021;24(1).
- b) Yetmar ZA, Beam E, O'Horo JC, et al. Monoclonal antibody therapy for COVID-19 in solid organ transplant recipients. Open Forum Infectious Diseases. 2021;8(6).

	Quality assessm	ent	N° of p	atients		Effect
N° of	Study design	Risk of bias	Intervention	No	Relative Absolute	
studies				intervention		
Overall mo	ortality					
2 ^{a-b}	Observational	Intermediate	40	72*	No patients who in Sarrell et al.	received casirivimab-imdevimab died (0/18 and 0/22), versus 2/72 (3%) in the control group
COVID-19-	related mortality	•		•	<u> </u>	
2 ^{a-b}	Observational	Intermediate	40	72	No patients who r	received casirivimab-imdevimab died due to COVID-19 (0/18 and 0/22), versus 1/72 (1%) in in Sarrell et al.
Hospital ac	dmission		•			
2 ^{a-b}		Intermediate	40	72*		22 treated with casirivimab-imdevimab were admitted for COVID-19-directed therapy vs. e control group (with a total admission rate of 14/72 or 19%). In the other study, 1/18 (6%)
ICU admiss	sion		•	•		
2 ^{a-b}	Observational	Intermediate	40	72*	No patients treate Sarrell et al.	ed with casirivimab-imdevimab were admitted to ICU vs. 1/72 (1%) in the control group in
Additional	respiratory supp	ort (HFNC, NIV,	MV)	1	1	
2 ^{a-b}	Observational	1	40	72*	a) 0/18 (0%). b) 0/22 (0%) vs. 0	/72 (0%) in controls required mechanical ventilation.
Incidence v	venous thromboe	embolism				
-	-	-		-		-
Incidence s	secondary infection	ons				
-	-	-		-		-
Incidence r	renal impairment					
1 ^a	Observational	Intermediate	22	72	0/22 (0%) vs. 1/1:	1 (9%) hospitalized (or 1/72 (1%) overall) controls needed renal replacement therapy.
Long-term	lung function da	ta				
-	-	-		-		-

- a) Retrospective single-center cohort study with 165 patients, including 13 LTR and 72 controls, of whom 22 were treated with casirivimab-imdevimab. COVID-19 variant most likely alpha variant (B1.1.7).
- b) Retrospective single-center cohort study with 73 SOTR, including 2 LTR, of whom 18 were treated with casirivimab-imdevimab. COVID-19 variant most likely alpha variant (B1.1.7).

^{*} Study of Sarrell et al. with 72 SOTR who did not receive monoclonal antibody therapy.

Bamlanivimab

Solid organ transplant outcomes from studies that included lung transplant recipients

Bibliography:

- a) Sarrell BA, Bloch K, El Chediak A, et al. Monoclonal antibody treatment for Covid-19 in solid organ transplant recipients. Transplant Infectious Disease. 2021;24(1).
- b) Yetmar ZA, Beam E, O'Horo JC, et al. Monoclonal antibody therapy for COVID-19 in solid organ transplant recipients. Open Forum Infectious Diseases. 2021;8(6).

	Quality assessi	ment	N° of	patients		Effect			
N° of	Study design	Risk of bias	Intervention	No	Relative	Absolute			
studies	, ,			intervention					
Overall m	ortality			I					
2 ^{a,b}	· ·	Intermediate	126	72*	No patients who received bamlanivimab died (0/55 al.	5 and 0/71), versus 2/72 (3%) in the control group in Sarell et			
COVID-19	related mortal	ity	-	1					
1 ^a	Observational	Intermediate	126	72	No patients who had bamlanivimab died due to CO group in Sarell et al.	VID-19 (0/55 and 0/71), versus 1/72 (1%) in the control			
Hospital a	admission		-	1	, -				
2 ^{a,b}	Observational	Intermediate	126		,	1% (8/71) - 13% (7/55) for SOTR treated with bamlanivimab. thout monoclonal antibody therapy were hospitalized,11/72			
ICU admi	ssion		•						
2 ^{a,b}	Observational	Intermediate	126	72*	In the study of Sarell et al., 2/71 (3%) were admitted the other study, no patients (0/55) were admitted t	d to ICU compared to 1/72 (1%) patient in control cohort. In to ICU due to COVID-19.			
Additiona	al respiratory su	pport (HFNC/N	IV/MV)						
2 ^{a,b}	Observational	1	126	72*	In Yetmar et al. noted no high-flow oxygen therapy, (1%) SOTR treated with bamlanivimab was mechani	, non-invasive or mechanical ventilation. In Sarrell et al., 1/71 ically ventilated vs. 0/72 (0%) in the controls.			
Incidence	venous thromb	oembolism	•	•		. , ,			
-	-	-	-	-		-			
Incidence	secondary infe	ctions		•					
-	-	-	-	-		-			
Incidence	renal impairme	ent							
1 ^a	Observational	Intermediate	71) treated with bamlanivimab had acute kidney injury with 0% ed to $4/11$ (36% or 6% overall) in the control group with 1%			
Long-ter	m lung function	data							

- a) Retrospective single-center cohort study with 165 patients, including 13 LTR and 72 controls, of whom 71 were treated with bamlanivimab. COVID-19 variant most likely alpha variant (B1.1.7).
- b) Retrospective single-center cohort study with 73 SOTR, including 2 LTR, of whom 55 were treated with bamlanivimab. COVID-19 variant most likely alpha variant (B1.1.7).

 * Study of Sarrell et al. with 72 SOTR who did not receive monoclonal antibody therapy.

Bebtelovimab

- a) Cochran W, Salto-Alejandre S, Barker L, et al. Covid-19 outcomes in solid organ transplant recipients who received Tixagevimab-cilgavimab prophylaxis and/or Bebtelovimab treatment in a nurse-driven monoclonal antibody program during the Omicron Surge. Transplantation. 2022;107(2).
- b) Yetmar ZA, Beam E, O'Horo JC, et al. Outcomes of bebtelovimab and sotrovimab treatment of solid organ transplant recipients with mild-to-moderate Coronavirus disease 2019 during the omicron epoch. Transplant Infectious Disease. 2022;24(4).

	O.,		NI9 - f	-4:4-		Fife as
	Quality assessr	1	•	atients		Effect
N° of	Study design	Risk of bias	Intervention	No	Relative	Absolute
studies				intervention		
Overall m	ortality					
2 ^{a,b}	Observational	Intermediate	237	-	0.7% (1/145) - 1	2% (2/92).
COVID-19	-related mortali	ty	•	•		
1 ^b	Observational	Intermediate	92	-	At least 1 SOTR	(1/2 who died, 1% of overall cohort) died due to COVID-19.
Hospital a	dmission		1	•	1	
2 ^{a,b}	Observational	Intermediate	237	-	3% (3/92) – 12%	(18/145).
ICU admis	sion					
1 ^b	Observational	Intermediate	92	-	0% (0/92).	
Additiona	l respiratory su	port (HFNC, N	IV, MV)			
1 ^a	Observational	Intermediate	145	-	0.7% (1/145) wa	as mechanically ventilated.
Incidence	venous thromb	oembolism				
-	-	-	-			-
Incidence	secondary infe	tions				
-	-	-	-			-
Incidence	renal impairme	nt				
-	-	-	-			-
Long-teri	n lung function	data				
-	-	-	-			-

a) Retrospective single-center study of 350 SOTR, of whom 145 were treated with bebtelovimab (including 18 LTR). COVID-19 variant: Omicron BA.2 and BA.5.

b) Retrospective multicenter cohort study with 361 SOTR, 92 SOTR (including 4 LTR) were treated with Bebtelovimab. COVID-19 variant: Omicron B.1.1.527 and BA.2. COVID-19: coronavirus 2019 disease, HFNC: high-flow nasale canula, MV: mechanical ventilation, NIV: non-invasive ventilation.