Author(s): Cruciani F, De Crescenzo F, Vecchi S, Saulle R, Mitrova Z, Amato L, Davoli M. **Question**: Should Interferon β -1a vs Standard treatment be used for COVID-19 patients? **Setting**: Inpatient

			Certainty asse	ssment			Nº of p	atients	Effect		
№ of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Interferon β-1a	Standard treatment	Relative (95% CI)	Absolute (95% CI)	Certainty
All-cause M	ortality										
4 1,2,3,4	randomised trials	serious a	very serious ^b	not serious	not serious	none	255/2164 (11.8%)	245/2166 (11.3%)	RR 0.63 (0.31 to 1.29)	42 fewer per 1.000 (from 78 fewer to 33 more)	⊕○○○ VERY LOW
All-cause M	ortality mild /	moderat	e ill patients								
1 ¹	randomised trials	serious c	not serious	not serious	not serious	none	10/482 (2.1%)	13/490 (2.7%)	RR 0.78 (0.35 to 1.77)	6 more per 1.000 (from 17 fewer to 20 more)	⊕⊕⊕⊖ MODERATE
All-cause M	ortality sever	e ill patie	nts								
1 ¹	randomised trials	serious c	not serious	not serious	not serious	none	178/1429 (12.5%)	163/1430 (11.4%)	RR 1.09 (0.90 to 1.33)	10 more per 1.000 (from 11 fewer to 38 more)	⊕⊕⊕⊖ MODERATE
All-cause M	ortality critica	ally ill pat	ients								
11	randomised trials	serious c	not serious	not serious	not serious	none	55/139 (39.6%)	40/130 (30.8%)	RR 1.29 (0.92 to 1.79)	89 more per 1.000 (from 25 fewer to 243 more)	⊕⊕⊕○ MODERATE

Number discharged

			Certainty asse	ssment			Nº of p	atients	Effect		
№ of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Interferon β-1a	Standard treatment	Relative (95% CI)	Absolute (95% CI)	
2 2,3	randomised trials	serious d	not serious	not serious	not serious	none	70/94 (74.5%)	59/96 (61.5%)	RR 1.19 (0.98 to 1.43)	117 more per 1.000 (from 12 more to 264 more)	⊕⊕⊕⊖ MODERATE
Number of p	atients with a	ny advers	se events								
2 2,3	randomised trials	serious d	very serious ^e	not serious	serious ^f	none	40/94 (42.6%)	31/96 (32.3%)	RR 3.14 (0.13 to 78.69)	691 more per 1.000 (from 281 fewer to 1.000 more)	⊕○○○ VERY LOW
Number of p	atients with s	evere ad	verse events								
1 ³	randomised trials	not serious	not serious	not serious	serious ^g	none	7/48 (14.6%)	14/50 (28.0%)	RR 0.52 (0.23 to 1.18)	134 fewer per 1.000 (from 216 fewer to 50 more)	⊕⊕⊕○ MODERATE
Length of st	ay in intensive	e care									
1 ²	randomised trials	very serious h	not serious	not serious	serious ^g	none	46	46	-	SMD 0.1 lower (0.51 lower to 0.31 higher)	⊕○○○ VERY LOW

Length of stay in hospital (mean days)

	Certainty assessment							№ of patients		fect	
№ of studies	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	Interferon β-1a	Standard treatment	145%	Absolute (95% CI)	Certainty
1 ²	randomised trials	very serious h	not serious	not serious	serious ^g	none	46	46	-	SMD 0.32 higher (0.09 lower to 0.73 higher)	⊕○○○ VERY LOW

Length of stay in hospital

1 ³	randomised trials	not serious	not serious	not serious	serious ^g	none	Tendency to a shorter duration of hospitalization in favour of Interferon beta 1a but not gaining statistical significance HR: 1.37 (0.85-2.20) p = 0.196	⊕⊕⊕○ MODERATE
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Progression of COVID-19 disease severity

1 ¹ rar	ndomised serious c	not serious	not serious	not serious	none	209/1911 (10.9%)	210/1920 (10.9%)	RR 1.00 (0.83 to 1.20)	0 fewer per 1.000 (from 19 fewer to 22 more)	⊕⊕⊕○ MODERATE
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Explanations

- a. Downgraded of one level for high risk of performance bias in two studies, high risk of attrition bias in one study, unclear risk of selection bias in two studies, and unclear risk of reporting bias in one study
- b. Downgraded of two levels for high heterogeneity: I²=71%
- c. Downgraded of one level for high risk of performance bias
- d. Downgraded of one level for high risk of performance and attrition bias and unclear risk of selection and reporting bias in one study
- e. Downgraded of two levels for high heterogeneity: I²=90%
- f. Downgraded of one level for wide confidence interval
- g. Downgraded of one level for small sample size
- h. Downgraded of two levels for high risk of performance and attrition bias and unclear risk of selection and reporting bias

References

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