

1 *Online Resource 5 - Table 1. Sensitivity analysis on delirium resolution the following day in motor delirium*
 2 *subtypes using a Markov regression model.*

Motor delirium subtype	Exposure on day X	n	Adjusted OR (95%CI) ^a	p value
Hyperactive or mixed delirium day	No haloperidol or clonidine	633	Reference ^b	
	Haloperidol only	747	0.43 (0.32-0.59)	<.001
	Clonidine only	473	0.86 (0.63-1.17)	0.331
	Both haloperidol and clonidine	728	0.45 (0.34-0.61)	<.001
Hypoactive delirium day	No haloperidol or clonidine	1119	Reference ^b	
	Haloperidol only	560	0.65 (0.51-0.84)	<.001
	Clonidine only	256	0.72 (0.51-1.01)	0.055
	Both haloperidol and clonidine	175	0.64 (0.44-0.95)	0.029

3 CI = Confidence Interval; OR = Odds Ratio

4 ^a Adjusted for age, APACHE IV score, admission type (acute surgery, elective surgery, medical reason), patients'
 5 mental status on the prior day (day X), modified SOFA score, metabolic acidosis, use of ventilation and
 6 administration of any antipsychotic other than haloperidol, benzodiazepines, propofol and opioids.

7 ^b Reference is transition from delirium to delirium or unarousable state on the following day, with no haloperidol or
 8 clonidine administered on the delirium day. Mortality or discharge on day X + 1 was included as a competing event.

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10 *Online Resource 5 - Table 2. Sensitivity analysis on delirium resolution the following day based on inclusion period*
 11 *using a Markov regression model.*

Inclusion period	Exposure on day X	n	Adjusted OR (95%CI) ^a	p value
2011 - 2013	No haloperidol or clonidine	853	Reference ^b	
	Haloperidol only	772	0.49 (0.38-0.63)	<0.001
	Clonidine only	156	0.88 (0.58-1.33)	0.533
	Both haloperidol and clonidine	277	0.43 (0.29-0.63)	<0.001
2015 - 2019	No haloperidol or clonidine	913	Reference ^b	
	Haloperidol only	536	0.44 (0.33-0.59)	<0.001
	Clonidine only	575	0.75 (0.57-0.99)	0.041
	Both haloperidol and clonidine	626	0.47 (0.35-0.61)	<0.001

12 CI = Confidence Interval; OR = Odds Ratio

13 ^a Adjusted for age, APACHE IV score, admission type (acute surgery, elective surgery, medical reason), patients'
 14 mental status on the prior day (day X), modified SOFA score, metabolic acidosis, use of ventilation and
 15 administration of any antipsychotic other than haloperidol, benzodiazepines, propofol and opioids.

16 ^b Reference is transition from delirium to delirium or unarousable state on the following day, with no haloperidol or
 17 clonidine administered on the delirium day. Mortality or discharge on day X + 1 was included as a competing event.

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28 *Online Resource 5 - Table 3. Sensitivity analysis on delirium resolution on day X+2 using a Markov regression*
29 *model .^a*

Exposure on day X	Mental status on day X+2	n	Adjusted OR (95% CI)^b	p value
No haloperidol or clonidine	Delirious or unarousable	816	Reference	
Haloperidol only	Awake without delirium	260	0.50 (0.41-0.61)	<.001
Clonidine only	Awake without delirium	228	0.96 (0.77-1.20)	0.747
Both haloperidol and clonidine	Awake without delirium	187	0.56 (0.46-0.70)	<.001

30 CI = Confidence Interval; OR = Odds Ratio.

31 ^a Mortality or discharge on day X + 2 was included as a competing event.

32 ^b Adjusted for age, APACHE IV score, admission type (acute surgery, elective surgery, medical reason), patients'
33 mental status on day X, modified SOFA score, metabolic acidosis, use of ventilation, and administration of any
34 antipsychotic other than haloperidol, benzodiazepines, propofol and opioids.