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Rotational thromboelastometry (ROTEM) profiling of early stage COVID-19 patients and its prognostic value: a prospective observational study



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ROTATIONAL THROMBOELASTOMETRY (ROTEM) PROFILING OF EARLY STAGE COVID-19 PATIENTS AND ITS PROGNOSTIC VALUE: A PROSPECTIVE OBSERVATIONAL STUDY

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Background

- There is emerging evidence for hypercoagulability and thrombosis in coronavirus disease 2019 (COVID-19) patients, contributing to mortality
- The mechanisms underlying this pro-thrombotic state is still uncompletely understood, and optimal criteria for early identification of patients at high risk for thrombosis and death remain unclear



Almskog LM, Wikman A, Svensson J, et al. Rotational thromboelastometry results are associated with care level in COVID-19. J Thromb Thrombolysis. 2021;51(2):437-45. Kong R, Hutchinson N, Görlinger K. Hyper- and hypocoagulability in COVID-19 as assessed by thromboelastometry. Two case reports. Korean J Anesthesiol 2020. Published online on Aug 10, 2020. doi: 10.4097/kja.20327.



Scope

The aim of this study was to assess:

- whether rotational thromboelastometry (ROTEM) would indicate hypercoagulability and fibrinolysis shutdown in an early stage of severe COVID-19 patients
- if ROTEM variables in combination with D-dimer would predict mortality



Kruse JM, Magomedov A, Kurreck A, et al. Thromboembolic complications in critically ill COVID-19 patients are associated with impaired fibrinolysis. Crit Care. 2020;24(1):676. Schenk B, Görlinger K, Treml B, et al. A comparison of the new ROTEM sigma with its predecessor, the ROTEM delta. Anaesthesia. 2019;74(3):348-356.



Methods

- ✓ The study was designed as a prospective, observational study
- ✓ We used ROTEM in a cohort of 35 adult critically ill COVID-19 patients admitted to an intermediate ward
- Conventional coagulation assays and D-dimer levels were also analyzed





Rotational thromboelastometry (ROTEM)



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Görlinger K, Almutawah H, Almutawaa F, et al. The Role of Rotational Thromboelastometry during the COVID-19 Pandemic: A Narrative Review. Korean J Anesthesiol. 2021;74(2):91-102



ROTEM PROFILING OF COVID-19 PATIENTS Results (1)

- ✓ The median age of the study population was 74 years and 74% were males; the median number of comorbidities and SOFA score were 3 and 3, respectively; the median duration of symptoms before admission was 5 days
- ✓ The median D-dimer value was 1.08 mg/L
- ✓ The median EXTEM Maximum Clot Firmness (MCF) was 71.0mm and the median maximum lysis (ML) value was 3%





Results (2)

The overall 30 day-mortality was 40% (14/35)

Predicting factors	N (%)	Mortality	р*	VTE	р*
None	8 (23)	0 (0)	/	0 (0)	/
Fibrinolysis shutdown	1				
ML < 3.5%	18 (51)	9 (50)	0.01	5 (28)	0.06
D—ML > 3.7	7 (39)	4 (57)	0.01	2 (29)	0.1
D-dimer > 1 mg/L	19 (54)	11 (58)	0.005	6 (32)	0.07
CT > 79 sec	5 (14)	4 (75)	0.002	2 (40)	0.05
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Relation between presence of predicting factors, 30-day mortality and thromboembolic events. *vs group without risk factors. VTE: venous-thromboembolism ML: Maximum Lysis D - ML: maximum D-dimer — EXTEM ML CT: Clotting Time



Results (3)

5 patients with a specific profile characterized by a high D-Dimer value, ROTEM signs of fibrinolysis shutdown and high EXTEM clotting time (CT) presented an early death after admission (n=4) and one patient an hemodinamically unstable PE at admission

	Outcomes				
ROTEM characteristics	Early death or TVE events N=5	No early death or TVE events N=30	р		
D-Dimer median mg/dl	15.04 (12.52-33.35)	0.78 (0.24-17.97)	<0.0001		
CT mean sec.	87.5 (71.5-147.5)	63.5 (58.3-73.5)	0.001		
MCF mean mm	63.0 (50.0-65.0)	71.5 (68.0-73.0)	0.001		
ML %	0 (0-2.0)	3.5 (1.0-5.0)	0.15		

T: Clotting Time CF: Maximum Clot Firmness L: Maximum Lysis



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Discussion

- ✓ Our results suggest that the altered ROTEM values in combination with D-dimer could be used to better identify the presence of fibrinolysis shutdown in those patients
- ✓ Our findings highlight that the mechanisms to explain the coagulability state of patients infected with SARS-CoV-2 is very complex: for example, the significance of high value of EXTEM clotting time (CT) is not yet known since now



Ibanez C, Perdomo J, Calvo A, et al. High D dimers and low global fibrinolysis coexist in COVID19 patients: what is going on in there? J Thromb Thrombolysis. 2020;15:1-5. Iba T, Connors JM, Levy JH. The coagulopathy, endotheliopathy, and vasculitis of COVID-19. Inflamm Res 2020; 69:1181-9.



Conclusions

- ✓ Our results confirm altered ROTEM values in early stage of patients with severe COVID-19
- ✓ The combination of D-dimer concentrations, fibrinolysis shutdown and ROTEM CT prolongation may provide valuable in identifying patients at high risk of early onset of thrombosis and death which potentially requiring specific therapies



Moore HB, Barrett CD, Moore EE, Jhunjhnuwala R, McIntyre RC, Moore PK, et al. STudy of Alteplase for Respiratory failure in SARS-Cov2/COVID-19: Study Design of the Phase IIa STARS Trial. Res Pract Thromb Haemost 2020; 4: 984-96.





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