INTRODUCTION

Rapid and accurate determination of direct oral anticoagulants (DOACs) is still a major medical need in urgent clinical situations.

ICARESS JLY 9-13

AIM

The aim of this study was to identify the feasibility of algorithm for rapid exclusion of clinically significant concentration of DOACs (>30 ng/mL) by data from inpatients from cardiology and neurology departments.

METHOD

128 paired plasma and urine samples at trough levels from patients treated for non-valvular atrial fibrillation or venous thromboembolism factor Xa inhibitor (DXI) (apixaban, n=31, rivaroxaban, n=53) direct thrombin inhibitor (DTI) (dabigatran, n=44) Plasma DOACs concentrations - Innovance anti-FXa assay for rivaroxaban and apixaban and Innovance DTI assay for dabigatran on BCSXP analyzer (Siemens Healthineers, Germany). Urine DOAC qualitative test - DOAC Dipstick test (DOASENSE GmbH, Heidelberg, Germany) Statistics: receiver operating curve (ROC) analysis – diagnostic accuracy kappa statistics - comparison between DOAC Dipstick test results and plasma concentrations of DOACs

Exclusion of relevant concentrations of direct oral anticoagulants in blood by DOAC Dipstick – proposal of a diagnostic algorithm for improvement of clinical decision-making in emergencies

I. <u>ĆELAP¹</u>, S. MARGETIĆ^{1,2}, J. PERIŠA¹, M. RAZUM¹, S. ŠUPRAHA GORETA³, A. ČAJEVIĆ-GLOJNARIĆ¹ 1 Department of Clinical Chemistry, Sestre milosrdnice University Hospital Center, Zagreb, Croatia 2 Catholic University of Croatia, Zagreb, Croatia 3 Department of Biochemistry and Molecular Biology, Faculty of Pharmacy and Biochemistry, University of Zagreb, Zagreb, Croatia

RESULTS

ROC analysis revealed threshold values of plasma concentrations for DXI \geq 14 ng/mL and \geq 19 ng/mL for DTI for detection of DXI and DTI by DOAC Dipstick, respectively. At cut-off value ≥19 ng/mL for DTI there was no false negative or false positive results (kappa value=1.0). For DXI at cut-off ≥14 ng/mL one false positive result was obtained, κ=0.92 (95 % CI 0.74-1.00).

Table 1. Parameters of diagnostic accuracy obtained by receiver operating curve analysis (Margetic et al, 2022)

DOAC	<u>Threshold</u>	AUC	Sensitivity (%)	Specificity (%)	PPV (%)	NPV (%)
	<u>(ng/mL)</u>	(95 % CI)	(95 % CI)	(95 % CI)	(95 % CI)	(95 % CI)
DXI	<u>></u> 14	0.994	100	85.7	98.7	100
n = 84		(0.981–1.0)	(95.3–100)	(42.1–99.6)	(93.1–100)	(54.1–100)
DTI	<u>></u> 19	NA	100	100	100	100
n = 44			(91.8–100)	(2.5–100)	(91.8–100)	(2.5–100)

AUC = area under the curve, CI = confidence interval, PPV = positive predictive value, NPV = negative predictive value, NA = not applicable



Figure 1. Suggested algorithm for emergency care of patients treated with DOACs for exclusion clinically relevant concentration of DOACs (>30 ng/mL)

CONCLUSIONS

Application of an algorithm for exclusion of relevant plasma concentrations by DOAC Dipstick can be proposed by testing first DOACs in urine and if positive followed by quantitative DOAC assays if clinically required.

ACKNOWLEDGEMENTS

The study was funded as a part of Croatian Science Foundation research project IP-2016-06-8208.

REFERENCES

Margetic et al. DOAC Dipstick testing can reliably exclude the presence of clinically relevant DOAC concentrations in circulation. Thromb Haemost 2022. in press.

CONTACT INFORMATION

margeticsandra@gmail.com

ivana.celap@gmail.com

