

**Investigation of Risk Factors Affecting the Development of Nephrotoxicity in Patients Receiving Colistin Therapy:** 

## BEZMÍÂLEM VARIF ONIVERSITESI

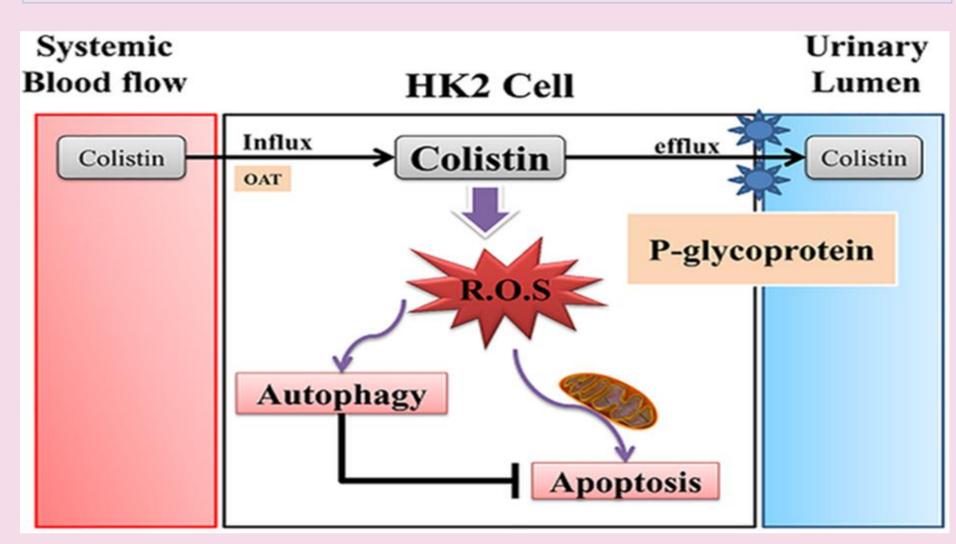
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**Introduction:**Colistin (COL) has become the backbone of the treatment of infections due to extensively drug-resistant (XDR) Gramnegative bacteria. The most common restriction to its use is acute kidney injury (AKI).Colistin is an antibiotic that was introduced many years ago and was withdrawn because of its nephrotoxicity. Nowadays, reemergence of this antibiotic for multi-drug resistant Gram-negative infections, and a new high dosing regimen recommendation increases concern about itsnephrotoxicity. Studies reported that approximately one third of colistin used patients have developed nephrotoxicity In experimental studies, when nephrotoxicity caused by colistin is reduced by oxidative release and use of antioxidants. However, the risk is increased with the presence of a different risks, such as age, hypertension, use of contrast media, and the use of other nephrotoxic drugs.

## Inclusion<br/>criteriaExclusion<br/>criteriaPatients over 18 years old<br/>admitted to anesthesia and<br/>reanimation<br/>ICUThose with a prior history of<br/>Renal disease history

At least 48 hours IV Colistin Pregnant women



Picture 1 Show the pathophysiological mechanisms of the colistin in nepfrons

**Methods:** We reached datas by using our Bezmialem Vakıf University' patients data system called 'Bizmed' retrospectively (July 2020 -June 2021). When we were arranging our patients groups, we took care exclusion and inclusion criteria. We didn't include patients that have chronic renal disease .In addition to above-mentioned exclusion criteria , we also determined inclusion criteria such as: Patients should be over 18 years old and admitted to anesthesia and reanimation ICU. Also they should take at least 48 hours IV colistin. Colistin treatments recieved ones "Patients with no pre-existing kidney dysfunction were compared in terms of risk factors( age, hypertension, use of contrast media, and the use of other nephrotoxic drugs) and outcomes of AKI graded according to the KDIGO criteria

treatmens recieved ones

Patients Hospitalized with Acute Hypotensive Shock

## Table 1 Show which groups are covered by our study

**Results:**6 out of 14 patients developed colistin nephrotoxicity.The average age of the patient group was %56.5 New onset AKI developed in 42,8% of the patients.Patients who developed nephrotoxicity ,the rate of sepsis was % 33,3, while in those who do not developed it, it was % 25. We didnt reach the targeted patient number on the ethical form.Statistical analysis could not be performed.

**Conclusion :**COL-induced nephrotoxicity occurred significantly more often in patients older than 60y of age and was related to low initial GFR estimations and high CCI scores, which were basicall determined by age.

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	А	В	С	D	E	F	G	Н	1	J	K	L
		BÖBREK Y <mark>ETMEZLİĞİ GELİŞEN (</mark> ÖLÜM (VA COMORBIDITIES										
1	NUMBE R		GENDE	AGE		DIABET ES	HYPER TENSIO N	COPD	NEOPL ASIA	HEART FAILUR E	CORO NARY ARTER DISOR DER	NEURO LOGIC AL DISOR DER
3	32734155954	0,0	1	50	1	0	1	0	0	0	1	0
4	13049747522	1,0	1	20	0	0	0	0	0	0	0	0
5	25549939208	0,0	0	69	1	0	0	0	1	0	0	1
6	18722251266	1,0	0	57	1	1	0	0	0	0	0	0
7	16916347412	1,0	0	83	1	0	0	0	1	0	0	0
8	26102004982	1,0	1	68	1	0	0	0	1	0	0	0
9	18941230660	0,0	0	85	0	0	0	0	0	0	0	0
10	34081320764	0,0	1	31	0	0	0	0	0	0	0	1
11	43816425834	0,0	0	63	0	0	0	1	0	1	1	0
12	35105005278	1,0	1	76	1	0	0	1	0	1	0	0
13	29311561536	0,0		46	0	0	0	0	0	0	0	0
14	34876358408	0,0		38	1	0	0	0	0	0	0	0
15	32791825524	0,0		62	1	0	0	0	1	0	0	0
16	61150418466	1,0		43	0	0	0	0	0	0	0	0

		Urine output		
	RIFLE	AKIN	KDIGO	
Definition	SCr increase $\geq$ 50% within 7 days	SCr increase $\geq$ 50% or $\geq$ 0.3 mg/dL within 48 h	SCr increase $\geq 0.3 \text{ mg/dL}$ within 48 h or $\geq 50\%$ within 7 days	UO $<$ 0.5 mL/kg/l for 6 h
Staging	RIFLE	AKIN	KDIGO	
RIFLE-Risk	SCr increase $\geq$ 50% or	SCr increase $\geq$ 50%	SCr increase ≥0.3 mg/dL	$\rm UO < 0.5~mL/kg/k$
AKIN stage 1 KDIGO stage 1	GFR decrease $>25\%$	or $\geq$ 0.3 mg/dL	within 48 h or $\geq$ 50% within 7 days	for 6 h
RIFLE-Injury AKIN stage 2 RIFLE stage 2	SCr increase $\geq 100\%$ or GFR decrease $>50\%$	SCr increase $\geq 100\%$	SCr increase ≧ 100%	UO < 0.5 mL/kg/ for 12 h
RIFLE-Failure AKIN stage 3 KDIGO stage 3	SCr increase $\geq 200\%$ or GFR decrease $>75\%$ or SCr $\geq 4 \text{ mg/dL}$ (with an acute rise $\geq 0.5 \text{ mg/dL}$ )	SCr increase $\geq 200\%$ or SCr $\geq 4 \text{ mg/dL}$ (with an acute rise $\geq 0.5 \text{ mg/dL}$ ) or need RRT	SCr increase $\geq$ 200% or SCr $\geq$ 4 mg/dL or need RRT	UO < 0.3 mL/kg/ł for 24 h or anuria for 12 h
RIFLE-Loss RIFLE-End stage	Need RRT for >4 weeks Need RRT for >3 months	<b>u</b> ,		

Abbreviation: RIFLE, risk of renal failure, injury to the kidney, failure of kidney function, loss of kidney function, and end-stage renal failure; AKIN, Acute Kidney Injury Network; KDIGO, Kidney Disease Improving Global Outcome; SCr, serum creatinine; UO, urine output; GFR, glomerular filtration rate; RRT, renal replacement therapy.

Picture2: show rating AKI according to different parameters(urine output,creatine,GFR)

Table 2 Show which parient developed AKI during antibiotic regimes recieving

**References:** 1Deryke, C. A., Crawford, A. J., Uddin, N., & Wallace, M. R. (2010). Colistin Dosing and Nephrotoxicity in a Large Community Teaching Hospital. Antimicrobial Agents and Chemotherapy, 54(10), 4503–4505. doi: 10.1128/aac.01707-09 2 Li, J., K. Coulthard, R. Milne, R. L. Nation, S. Conway, D. Peckham, C. Etherington, and J. Turnidge. 2003. Steady-state pharmacokinetics of intravenous colistin methanesulphonate in patients with cystic fibrosis. J. Antimicrob. Chemother. 52:987-992. 3Doshi NM, Mount KL, Murphy CV (2011) Nephrotoxicity associated with intravenous colistin in critically ill patients. Pharmacother J Human Pharmacol Drug Ther 31(12):1257–1264

4Balkan, I. I., Dogan, M., Durdu, B., Batirel, A., Hakyemez, I. N., Cetin, B., ... Tabak, F. (2014). Colistin nephrotoxicity increases with age. Scandinavian Journal of Infectious Diseases, 46(10), 678–685. doi: 10.3109/00365548.2014.926021
5 Yousef, J. M., Chen, G., Hill, P. A., Nation, R. L., & Li, J. (2011). Ascorbic acid protects against the nephrotoxicity and apoptosis caused by colistin and affects its pharmacokinetics. Journal of Antimicrobial Chemotherapy, 67(2), 452–459. doi: 10.1093/jac/dkr483