

# Dynamics of age characteristics and prevalence of concomitant cardiovascular and non-cardiovascular diseases in patients hospitalized with COVID-19 during epidemic wave: data from TARGET-VIP registry

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**Aim.** According to hospital-based registry, to evaluate the age characteristics and prevalence of concomitant cardiovascular and noncardiovascular diseases in patients hospitalized with COVID-19 during epidemic wave.

**Material and methods.** The TARGET-VIP register included 1130 patients aged 57,5±12,8 years (men, 51,2%) hospitalized at the Pirogov National Medical and Surgical Center from April 6, 2020 to June 22, 2020 with COVID-19. Cardiovascular diseases (CVDs) were diagnosed in 51,6% of patients, non-cardiovascular chronic diseases — in 48,6%, while CVDs and/or non-cardiovascular chronic diseases — in 65,8% of patients.

Results. The average age of patients significantly increased by an average of 0,77 years per week (p<0,001), while the difference between the 1<sup>st</sup> week (52,8 years) and 11<sup>th</sup> week (62,2 years) was 9,4 years; the proportion of men did not change significantly. The proportion of patients with CVDs increased significantly - from 34,2% to 66,7%, on average by 3,7% per week (p<0,001; Incidence Risk Ratio (IRR)=1,037; 95% confidence interval (CI), 1,017-1,058), with chronic non-cardiovascular diseases - from 32,5% to 43,2%, on average by 2,5% per week (p<0,001; IRR=1,025; 95% CI, 1,002-1,049), as well as those with CVDs and/or chronic non-cardiovascular diseases - from 47,5% to 75,3%, on average by 3,2% per week (p<0,001; IRR=1,032; 95% CI, 1,017-1,048). Over the entire period, the proportion of people with hypertension (HTN) was 47,0%, with coronary artery disease (CAD) - 15,4%, with heart failure (HF) - 4,0%, and with atrial fibrillation (AF) - 10,1%. The proportion of patients with HTN increased by 9,5% (p<0,001; OR=1,095; 95% CI, 1,047-1,144), with CAD - by 9,4% (p=0,01; OR=1,094; 95% CI, 1,022-1,172) and with AF - by 9,4% (p<0,001; OR=1,094; 95% CI, 1,023-1,170) per week. The proportion of patients with diabetes was 16,5%, with respiratory diseases -11,4%, with chronic kidney disease (CKD) - 12,6%, with digestive diseases -22,5%, with obesity -6,1%. During the epidemic wave, the most pronounced increase in the proportion of patients with CKD was by 6,2% (p=0,036; OR=1,062; 95% CI, 1,004-1,124) and with digestive diseases — by 6,0% (p=0,01; OR=1,060; 95% CI, 1,014-1,109) per week. Conclusion. According to the 11-week TARGET-VIP registry, the age of patients increased by 9,4 years, CVD cases - by 1,9 times (mainly HTN, CAD, AF), and chronic non-cardiovascular pathology - by 1,3 times (mainly CKD and digestive diseases). These trends in hospital practice corresponded to a weekly increase in the proportion of patients with a higher risk of fatal and non-fatal complications, which is the basis for further research in order to develop a system for a comprehensive prognostic assessment of the degree and rate of increase in the load on hospitals during COVID-19 epidemic wave.

**Keywords:** coronavirus disease 2019, COVID-19, hospital-based registry, age and sex characteristics, cardiovascular diseases, chronic non-cardiovascular diseases, comorbidity, weekly dynamics, epidemic wave.

#### Relationships and Activities: none.

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### Introduction

The pandemic of coronavirus disease 2019 (COVID-19) has claimed the lives of more than 5 million people and damaged the economic and social life of the entire world community. Every week, millions of new cases continue to be registered worldwide [1, 2]. At the same time, age, concomitant cardiovascular (CVD) and non-cardiovascular diseases have an important prognostic value in COVID-19 patients [3-10]. However, in this category of patients, the dynamics of age and sex characteristics, the detection rate of concomitant CVD and non-cardiovascular diseases during the epidemic wave of COVID-19, in particular, among hospitalized patients, have been little studied. The TARGET-VIP prospective in-hospital registry created in Russia makes it possible to study the clinical and anamnestic characteristics, structure and dynamics of comorbidity in patients with COVID-19 during the epidemic wave.

Based on the foregoing, the aim of the study was to evaluate the age characteristics and prevalence of concomitant cardiovascular and non-cardiovascular diseases in patients hospitalized with COVID-19 during epidemic wave.

### Material and methods

The TARGET-VIP registry includes 1130 patients hospitalized at the Pirogov National Medical and Surgical Center from April 6, 2020 to June 22, 2020 due to COVID-19 (age, 57,5±12,8 years; men, 51,2%). Concomitant CVDs was diagnosed in 51,6% of patients, while chronic non-CVDs - in 48,6%, CVDs and/or non-CVDs - in 65,8%. At 11-week enrollment period, corresponding to the epidemic wave in Moscow, the number of new COVID-19 cases in Moscow on April 6, 2020 and June 22, 2020 was 691 and 662, respectively, and the maximum number (6703 cases) was registered on May 07, 2020. The weekly dynamics of average patient age and the proportion of following CVDs was assessed: hypertension (HTN); coronary artery disease (CAD); heart failure (HF); atrial fibrillation (AF); prior myocardial infarction (MI) and stroke. The dynamics of the proportion of following chronic non-CVDs was also assessed: diabetes, respiratory diseases (RDs), chronic kidney disease (CKD), digestive diseases (DDs), anemia, and obesity.

Descriptive statistics were used for statistical data processing. Numerical data are presented as M±SD or Me [Q25%; Q75%]. The statistical significance of differences in numerical data was assessed using the Student's test, while categorical data — using the  $\chi^2$  test. For assessing weekly dynamics, regression analysis was used. For binary variables (sex, presence of certain diseases), logistic regression was used with an odds ratio (OR) and 95% confidence interval (CI). For data on the number of certain diseases (number of CVDs, chronic non-CVDs or their sum), Poisson regression was used with an Incidence Risk Ratio (IRR) and corresponding 95% CI. Sex and age of patients were included as covariates in the regression models. Differences were considered significant at p<0,05. Statistical processing was performed using Statistica 7.0 and Stata 15.0 software.

### **Results**

The dynamics of age, sex characteristics and comorbidities in patients admitted is presented in Table 1. The mean age of patients over 11-week period increased significantly, on average, by 0,77 years per week, while this parameter for the first (52,8 years) and 11<sup>th</sup> weeks (62,2 years) differed by 9,4 years. The proportion of men did not change significantly during the followup, with a maximum of 56,3% in the first week and a minimum of 48,2% in the 11<sup>th</sup> week (p=0,145; OR=0,973; 95% CI: 0,938-1,009). From the 1<sup>st</sup> to the 11<sup>th</sup> week of the inclusion period, the proportion of people with CVDs increased by 1,9 times, with chronic non-CVDs — by 1,3 times, and with both types of pathology — by 1,6 times.

The proportion of patients with CVDs increased significantly, with the CVD number per patient increasing by an average of 3,7% per week (p<0,001; IRR=1,037; 95% CI: 1,017-1,058). The proportion of patients with CVD was the lowest in the 1<sup>st</sup> week (34,2%) and the highest during the 8<sup>th</sup> and 11<sup>th</sup> weeks (68,9 and 66,7%). The average number of chronic non-CVDs per patient significantly increased by 2,5% per week (p<0,001; IRR=1,025; 95% CI: 1,002-1,049). The proportion of patients with CVD and/or chronic non-CVDs increased from 47,5 to 75,3% during the analyzed period. The mean number of CVDs and/or chronic non-CVDs increased by 3,2% per week (p<0,001; IRR=1,032; 95% CI: 1,017-1,048).

Over the entire period of epidemic wave, the proportion of people with hypertension, on average, was 47,0%, with CAD without MI - 15,4%. At the same time, the proportion of patients with prior MI and stroke averaged 5,0 and 4,0%, while the proportion of patients with CAD and AF was 4,0 and 10,1%, respectively.

Dynamics of CVD detection rate in analyzed patients during the study period is presented in Table 2. During the epidemic wave from the 1<sup>st</sup> to the 11<sup>th</sup> week, the most pronounced increase was noted in patients HTN (from 30,0 to 64,2%, i.e. 2,1 times; p<0,001), CAD (from 13,2 to 37,3%, i.e. 2,8 times; p<0,001) and AF (from 7,1 to 21,0%, i.e. 3,0 times; p<0,001). The proportion of patients with HTN per week increased on average by 9,5% (p<0,001; OR=1,095; 95% CI: 1,047-1,144), with CAD without MI — by 9,4% (p=0,01; OR=1,094; 95% CI: 1,022-1,172) and with AF by 9,4% (p<0,001; OR=1,094; 95% CI: 1,023-1,170). There was no significant weekly increase in the proportion of patients with CAD who underwent MI and stroke (p>0,05).

During 11-week period, the proportion of patients with diabetes, on average, was 16,5%, with respiratory diseases -11,4%, with CKD -12,6%, with DDs -22,5%, with anemia -3,5%, and with obesity -6,1%. The dynamics of the detection rate of chronic non-CVDs is presented in Table 3. Of the analyzed chronic non-CVDs, to the greatest extent for 11-week

	Dyní to the ce	amics of a nter for C	tge, sex chi 30VID-19	aracteristi and com	ics and no munity-ae	oncommur cquired pr	nicable di neumonia	seases in t during th	patients ac he enrollm	lmitted tent perio	þ			Table 1
Parameter	Weeks of	enrollment 1	period			4						OR/B/	95% CI	a
	1 n=240	2 n=86	3 n=140	4 n=127	5 n=93	6 n=127	7 n=65	8 n=45	9 n=84	10 n=42	11 n=81	IRR		-
Age (years)	$52,8\pm 16,$	0 54,3±15,	5 57,3±14,4	59,6±16,(	) 56,9±15,2	2 60,0±15,7	57,2±15,2	), 62,7±17,0	) 59,3±15,3	57,4±14,2	62,2±16,2	0,77	0, 49; 1, 06	<0,001
Proportion of men (%)	56,3	48,8	49,3	52,8	52,7	50,4	50,8	48,9	41,7	57,1	48,2	0,973	0,938; 1,009	0,145
Proportion of patients with CVD (%)	34,2	43,0	52,1	52,0	50,5	58,3	53,9	68,9	61,9	54,8	66,7	1,037	1,017; 1,058	<0,001
Proportion of patients with non-CVD (%)	32,5	37,2	38,6	44,9	43,0	52,0	41,5	44,4	48,8	45,2	43,2	1,025	1,002; 1,049	<0,001
Proportion of patients with CVD and/or non-CVD (%)	47,5	60,5	63,6	66,9	59,1	76,4	64,6	77,8	77,4	73,8	75,3	1,032	1,017; 1,048	<0,001
Note: OR (for "proportion of men"), $\beta$ – confidence interval, OR – odds ratio, CVD	«coefficien — cardiovae	t β» (for "a <sub>ξ</sub> cular disease	ge"), IRR (fo es.	r "proportic	on of patient	s with CVD'	, "proporti	on of patien	ts with non-	CVD" and 1	for a combine	ation of t	hese paramete	rs). CI —
														Table 2
	Ц	bynamics and	of CVD de l communi	stection r ty-acquir	ate in pati ed pneun	ients admi 10nia duri	tted to th ng the en	e center f rollment	or COVII period	0-19				
Parameter, %	Weeks of	enrollment 1	period									OR	95% CI	d
		2	3	4	5	9	7	8	6	10	11	1		4
	n=240	n=86	n=140	n=127	n=93	n=127	n=65	n=45	n=84	n=42	n=81			
Proportion of patients with HTN	30,0	39,5	50,0	49,6	46,2	54,3	49,2	62,2	57,1	47,6	64,2	1,095	1,047; 1,144	<0,001
Proportion of patients with CAD without MI	10,7	9,9	10,4	18,3	11,4	14,1	20,6	31,0	19,7	19,5	27,4	1,094	1,022; 1,172	0,010
Proportion of patients with MI	2,5	5,8	3,6	5,5	5,4	4,7	3,1	6,7	9,5	2,4	9,9	1,060	0,970; 1,158	0,197
Proportion of patients with HF	5,4	11,6	9,3	9,5	7,5	11,8	10,8	13,3	10,7	4,8	18,5	1,022	0,952; 1,097	0,538
Proportion of patients with AF	7,1	4,7	5,0	8,7	11,8	13,4	9,2	20,0	14,3	7,1	21,0	1,094	1,023; 1,170	<0,001
Proportion of patients with CVA	2,1	2,3	3,6	7,9	5,4	5,5	4,6	2,2	2,4	4,8	3,7	1,003	0,912; 1,104	0,945
Note: CVA – all cases of MI and transient i CI – confidence interval, CAD – coronary	ischemic att artery disea	ack. MI and se, MI – m	CVA is taken yocardial infa	into accour ction, CVA	it on the date 	e of hospitaliz scular accider	zation; MI a nt, OR – od	nd CVA devi Ids ratio, AF	eloped in the - atrial fibri	hospital are llation, HF	not taken int – heart failur	o account e.	t. HTN – hyp	ertension,
														Table 3
	fo	Dyna r COVID	mics of no	n-CVD c mmunity	letection 1 y-acquired	rate in pat	ients adm nia durinį	nitted to tl g the enrc	he center ollment pe	riod				
Parameter, %	Weeks of	enrollment l	period									OR	95% CI	d
	1 n=740	2 n=86	3 n=140	4 n=127	5 n=03	6 n=177	7 n=65	8 n=45	9 n=84	10 n=47	11 n=81			
Proportion of patients with diabetes	12.5	11.6	18.6	15.0	14.0	21.3	15.4	26.7	17.9	11.9	23.5	1.037	0.983: 1.091	0.154
Proportion of patients with RD	13,3	11,6	12,9	8,7	14,0	8,7	9,2	11,1	13,1	7,1	11,1	0,946	0,890; 1,006	0,079
Proportion of patients with CKD	6,7	14,0	13,6	11,8	15,1	10,2	15,4	17,8	11,9	21,4	19,8	1,062	1,004; 1,124	0,036
Proportion of patients with anemia	1,7	2,3	3,6	4,7	3,2	3,9	3,1	2,2	3,6	7,1	6,2	1,062	0,960; 1,174	0,243
Proportion of patients with DD	14,6	18,6	17,9	25,2	22,6	33,9	21,5	26,7	28,6	28,6	24,7	1,060	1,014; 1,109	0,011
Proportion of patients with obesity	4,2	3,5	5,0	8,7	8,6	7,1	9,2	4,4	7,1	9,5	3,7	1,041	0,964; 1,124	0,304
Note: RD — respiratory diseases. DD — dig	estive diseas	es. CI — cor	afidence inter	val. OR – o	dds ratio. CI	CD = chronic	e kidnev dise	9366						

## COVID-19 and diseases of the circulatory system

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period the proportion of patients with CKD (from 6,7 to 19,8%, i.e. 3,0 times; p<0,001) and DDs (from 14,6 to 24,7%, i.e. in 1,7 times; p=0,037) increased. During the epidemic wave, the most pronounced increase in the proportion of patients with DDS was noted. The proportion of CKD patients increased by an average of 6,2% per week (p=0,036; OR=1,062; 95% CI: 1,004-1,124), and those with CKD increased by 6,0% (p=0,01; OR=1,060; 95% CI: 1,014-1,109). There was no significant weekly increase in the proportion of patients with diabetes, RDs, anemia and obesity (p>0,05).

### Discussion

Monitoring of COVID-19 patients in a pandemic, according to published data, revealed that the severity of its clinical course increases with older patients, and the prognosis in the studied cohorts of patients worsens not only with an increase in the proportion of elderly patients, but also with an increase in the proportion of people with CVD and/or chronic non-CVDs (in particular, with diabetes, respiratory diseases, DDs, cancer, obesity) [6-9, 11, 12]. In addition, as with other respiratory tract infections, COVID-19 can not only worsen the course of CVD and non-CVDs, but also lead to cardiovascular events [6, 13-17].

TARGET-VIP registry revealed a significant weekly increase in the age and detection rate of concomitant CVDs and non-CVDs over 11-week inclusion period, which largely corresponded to the dates of the most pronounced part of COVID-19 wave. Thus, with each subsequent week of the inclusion period during the epidemic wave among hospitalized patients, the risk of fatal and non-fatal complications increased. This is an unfavorable factor, since severe patients are not only more difficult to treat [2, 17, 18], but they also have a longer length of hospital stay [4, 5, 7-9]. The above together leads to a progressive increase in the hospital

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load during the epidemic wave of COVID-19 [2, 17, 18]. Perhaps one of the reasons for this is that at the beginning of epidemic, predominantly more mobile and young people were infected, from which older members of their families and labor collectives subsequently became infected, significant part of whom had higher prevalence of noncommunicable diseases.

A more detailed analysis of the severity and rates of increase in the risk of fatal and non-fatal complications in patients hospitalized with COVID-19 will, in the future, make it possible to predict an increase in hospital load with each subsequent week of the epidemic wave. It seems very important to compare data over the epidemic wave period on the dynamics of age and detection rate of significant comorbidities in specific hospitals in certain cities and regions as a whole, in particular for Moscow. If a significant correlation is found in a particular hospital and in a corresponding region, there will be an additional opportunity to predict the hospital load in the context of subsequent epidemic waves in order to optimize inpatient care. This is the subject of further research.

### Conclusion

According to the 11-week TARGET-VIP registry, the age of patients increased by 9,4 years, CVD cases — by 1,9 times (mainly HTN, CAD, AF), and chronic non-cardiovascular pathology — by 1,3 times (mainly CKD and digestive diseases). These trends in hospital practice corresponded to a weekly increase in the proportion of patients with a higher risk of fatal and non-fatal complications, which is the basis for further research in order to develop a system for a comprehensive prognostic assessment of hospital load during COVID-19 epidemic wave.

### Relationships and Activities: none.

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