

Оценка диаметра лёгочной артерии при различной степени тяжести течения COVID-19 (по данным бесконтрастной компьютерной томографии лёгких)

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АННОТАЦИЯ

Обоснование. Компьютерная томография является методом выбора при оценке объёма поражения лёгких при вирусных пневмониях, в том числе ассоциированных с COVID-19. Помимо оценки объёма поражения лёгких, компьютерная томография позволяет определить размеры магистральных сосудов грудной клетки. Это позволило проанализировать связь между тяжестью течения COVID-19 и наличием изменения диаметров лёгочной артерии и восходящей аорты. Расширение лёгочной артерии является признаком лёгочной гипертензии. Изучение данных закономерностей может иметь клиническое значение в отношении определения тактики лечения и прогноза течения заболевания COVID-19.

Цель — оценить зависимость между диаметром лёгочной артерии и степенью тяжести течения COVID-19 у пациентов различного возраста.

Материалы и методы. Одноцентровое одномоментное сплошное неконтролируемое исследование выполнено в группе пациентов (*n*=511, 267 мужчин, медиана 59 лет, IQR 49,0–65,0, размах от 31 до 84 лет), проходивших лечение во временном госпитале для лечения пациентов с COVID-19. При госпитализации все пациенты прошли компьютерное томографическое исследование органов грудной клетки с помощью мобильной системы Airo TruCT (Stryker, CША). Степень поражения лёгочной ткани оценивалась по шкале КТ 1–4. Измерение диаметра лёгочной артерии и восходящей аорты проводилось стандартными инструментами рабочей станции врача-рентгенолога перпендикулярно длинной оси сосуда.

Результаты. Получены следующие статистически значимые закономерности: расширение лёгочной артерии и увеличение отношения лёгочной артерии/восходящей аорты было связано с увеличением степени поражения лёгких при COVID-19 (критерий Краскела–Уоллиса, *p* <0,001; медианный тест, *p* <0,001); диаметр восходящего отдела аорты достоверно увеличивается с возрастом пациента (критерий Краскела–Уоллиса, *p* <0,001; медианный тест, *p* <0,001). Показаны недостоверная связь между увеличением диаметра лёгочной артерии и возрастом пациента (критерий Краскела–Уоллиса, *p*=0,094; медианный тест, *p*=0,311) и недостоверная связь между изменением диаметра восходящей аорты и степенью поражения лёгких (критерий Краскела–Уоллиса, *p*=0,061; медианный тест, *p*=0,165). Во всех возрастных группах с тяжёлым течением заболевания и большим объёмом поражения лёгких (КТ-3 и КТ-4) показано достоверно большее количество пациентов с признаками лёгочной гипертензии (расширенная от 29 мм и более лёгочная артерия).

Заключение. Дилатация лёгочной артерии и увеличение отношения диаметров лёгочной артерии/восходящей аорты достоверно связано с увеличением объёма поражения лёгких при COVID-19 во всех возрастных группах.

Ключевые слова: COVID-19; лёгочная артерия; лёгочная гипертензия; аорта; компьютерная томография.

Как цитировать

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Changing of pulmonary artery diameter in accordance with severity of COVID-19 (assessment based on non-contrast computer tomography)

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ABSTRACT

BACKGROUND: Computed tomography is the method of choice for assessing the volume of lung damage in viral pneumonia, including those associated with COVID-19. In addition, computed tomography can determine the main vessels size of the thorax. This allowed us to analyze the relationship between the severity of COVID-19 and the changes in the diameters of the pulmonary artery (PA) and ascending aorta (Ao). Dilation of the PA is a sign of pulmonary hypertension. The study of these patterns may be of clinical significance in determining the treatment tactics and prognosis of the course of COVID-19 disease.

AIM: To evaluate the relationship between PA diameter and the severity of the COVID-19 course in patients of different ages.

MATERIALS AND METHODS: This study is a single-centered, cross-section, continuous, uncontrolled study performed on a group of patients (*n*=511, 267 men, median 59 years, IQR 49.0–65.0, ages 31–84 years) who were treated in a temporary hospital to treat patient with COVID-19. During hospitalization all patients were examined by CT scan of the chest. All studies were carried out using a mobile CT scan system Airo TruCT (Stryker, USA). The degree of damage to the lung tissue was assessed using the CT volume scale 1–4. Measurement of the LA and Ao diameters was carried out using standard instruments of the radiologist's CT workstation perpendicular to the long axis of the vessel.

RESULTS: The following statistically significant regularities were obtained: the detection of a dilated pulmonary artery (PA) and an increased PA/Ao ratio correlated to an increase in the degree of lung damage in COVID-19 (Kruskal-Wallis test, K-W p < 0.001; median test, MT p < 0.001), the diameter of the ascending aorta (Ao) significantly increases with the patient's age (K-W p < 0.001; MT p < 0.001). An insignificant correlation between an increase in the diameter of the pulmonary artery (PA) and the patient's age (K-W p = 0.094; MT p = 0.311) and an insignificant correlation between detection of a change in aortic (Ao) diameter and the degree of lung damage (K-W p = 0.061; MT p = 0.165) were shown. In groups with a severe course of the disease and a large volume of lung lesions (CT-3 and CT-4), a significantly greater number of patients with signs of pulmonary hypertension (detection of the dilated pulmonary artery: 29 mm and more) was shown for all age groups.

CONCLUSION: The study showed that PA dilatation and increased PA/Ao diameter ratio were significantly associated with increased pulmonary lesion volume in COVID-19 in all age groups.

Keywords: COVID-19; pulmonary artery; pulmonary hypertension; thoracic aorta; X-ray computed tomography.

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评价当前COVID-19不同严重程度的光动脉直径 (根据肺的非敏感性计算机断层扫描)

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简评

论证。计算机断层扫描是评估病毒性肺炎(包括COVID-19相关肺炎)肺部损伤的一种选择方法。除了评估肺损伤外,计算机断层扫描还可以确定胸腔主要血管的大小。分析了COVID-19流动的严重程度与肺动脉和上主动脉直径变化的存在之间的关系。肺动脉扩张是肺高压的标志.研究这些模式可能对确定治疗策略和预测COVID-19疾病的趋势具有临床意义。

目标是估计不同年龄患者的光动脉直径和COVID-19流量的严重程度

材料与方法患者组中进行单核单级固体不受控制的研究 (n=511,267男性,中美洲59 岁,IQR 49.0-65.0,范围从31至84岁),在临时医院治疗COVID-19患者。 随着住院治疗, 所有患者都通过了使用IISRO Truct Mobile System (Stryker,美国)的胸部器官的计算机 断层研究。 CT 1-4规模上估计了轻微组织的病变程度。 通过垂直于容器的长轴的放射科医 生的工作站的标准工具来进行光动脉直径和上升主动脉的测量。

结果。获得以下统计学显着的模式:光动脉的膨胀和光动脉/升序姿势的升高与COVID-19 肺部病变程度的增加有关(克拉克尔-沃利斯准则,p<0.001;中值测试,p<0.001); 升 高主动脉的直径随着患者的年龄而显着增加(克拉克尔-沃利斯准则,p<0.001;中值测 试,p<0.001)。显示了光动脉直径和患者年龄的直径之间的不准确的关系(克拉克尔-沃 利斯准则,p=0.094;嗜源性测试,p=0.311)和腹腔直径的变化和肺部的病变程度之间的不 准确的联系(克拉克尔-沃利斯准则,p=0,061;中值测试,p=0.165)。所有年龄段的疾病 方面和大量的肺病灶(KT-3和KT-4)中,可靠地增加了患有轻质高血压症状的患者(从29毫 米和更多的光动脉膨胀)是显示。

结论。光动脉的扩张和光动脉/升序的直径比率的增加与在所有年龄组中COVID-19的肺病变的体积增加是可靠的。

关键词: COVID-19; 易动脉; 轻盈高血压; 主动脉; CT扫描。

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BACKGROUND

In the context of the coronavirus disease-2019 (CO-VID-19) pandemic, pulmonary computed tomography (CT) became one of the leading methods for COVID-19 severity diagnosis and assessment [1–3].

The federal clinical guidelines for the prevention, diagnosis, and treatment defined four degrees of COVID-19 severity [2]. This division is based on several clinical and laboratory parameters, including the volume of pulmonary tissue lesions according to grades 1-4 of the CT classification [1, 3]. Patients with a mild degree of disease severity had no lung lesions (CT-0); moderate disease severity had up to 25% (CT-1) and 50%-75% (CT-2) lung involvement degree; 50%-75% (CT-3) in severe disease course; and over 70% (CT-4) in extremely severe disease. In addition, the degree of pneumonia severity according to the CT classification has prognostic value. Thus, a significantly increased proportion of patients who died depending on the volume of the affected pulmonary parenchyma according to the CT scale was shown, including a jump in the risk of lethality when transitioning from mild and moderate to severe and extremely severe disease course (from CT 1-2 to CT 3-4) [4].

The radiological sign analyses, which are typical for COVID-19-associated pneumonia, paid considerable attention to the pulmonary parenchyma, pleura, and bronchial tree assessment, and to a lesser extent, to the great vessel changes [5, 6]. The most characteristic and stable signs of COVID-19-associated pneumonia are segmental pulmonary artery (PA) dilation [7] and high risks of pulmonary embolism and acute respiratory distress syndrome, which is accompanied by gas exchange worsening, hypoxemia, and multiple organ failure [4, 7, 8].

A significantly increased PA diameter and PA and ascending aorta (Asc Ao) diameter ratio is shown in patients with lethal outcomes compared to those who recovered [9].

Thus, the lung lesion volume and the state of the pulmonary vessels (both segmental PAs, [10–12] and the pulmonary trunk [main PA]) are considered to assess the course of COVID-19, current therapy effectiveness, and prognosis using CT [9]. Increased pressure in the arterial vascular bed of the pulmonary circulation indicates the difficulty of blood transit through the lungs, which reduces the patient's compensatory abilities, leading to heart failure.

AIM: This study is aimed to determine the relationship between the COVID-19 severity and the PA and Asc Ao diameters in patients of different ages.

METHODS

Study design

A single-center, cross-sectional, continuous, and noncontrolled study based on chest CT was conducted (Fig. 1).

Eligibility criteria

Inclusion criteria are COVID-19 diagnosis that is confirmed by polymerase chain reaction, signs of viral pneumonia on CT scan, and voluntary informed consent for



Fig. 1. Study design

CT: computed tomography; CT 1-4: adapted scale for visual assessment of the relationship between the severity of the patient's general condition and the nature and intensity of radiological signs of pulmonary changes in COVID-19; PA: pulmonary artery.

medical intervention. Chest CT scans were performed for the first time in a reserve hospital during admission or within 4 days after admission, provided that chest CT scans were performed in another medical institution at the prehospital stage [3].

Exclusion criteria are the history of chronic lung disease diagnosis, surgical interventions on the chest organs, and pronounced artifacts on CT scans that prevent an adequate lung lesion assessment and measurement of the great vessel diameters.

The non-inclusion criterion is the absence of viral pneumonia signs on CT scans (CT-0) due to a limited number of such admissions.

Setting

The study was conducted with a group of patients who were treated at the temporary reserve hospital in Krylatskoye Ice Palace (Vorokhobov City Clinical Hospital No. 67) from October 08, 2020, to December 05, 2020.

Medical intervention

in a specific period, 2998 non-contrast chest CT scans were performed in the hospital for diagnosis or follow-up of patients with COVID-19 viral pneumonia.

CT scans were performed on an Airo TruCT mobile CT scanner (Stryker, USA), which was installed at the reserve hospital for the treatment of patients with COVID-19 (Fig. 2). The mobile CT scanner has an extended aperture diameter of 105 cm and 32 detector rows. This CT system was designed for neurosurgical operating rooms. However, its compact size and low power requirements (possibility of connection to the 1.5 kW power grids) allows an effective usage of the device in a temporary hospital setting. In addition, the power supply unit of the tomograph allows short-term scanning with a power of up to 30 kV. Given the necessity of providing around-the-clock operation mode, the following scanning parameters were chosen: X-ray tube voltage: 120 kV, current: 38 mA, slice thickness:1 mm, matrix: 512×512, pitch: 1.415, rotation time: 1.92 sec, average scanning length: 30 cm (12 sec), and effective dose: 3.9 mSv. Maximum flow is up to 6 patients per hour.



Fig. 2. Airo TruCT mobile CT scanner was installed in the admission department of the temporary reserve hospital for the treatment of patients with COVID-19, which was organized in the Krylatskoe Ice Palace.

The anonymized CT scans were independently assessed by two radiologists with 3 and 9 years of experience, respectively. The software of the Unified Radiological Information Service of the Unified Medical Information and Analytical System of Moscow City was used on the Agfa Enterprise Imaging platform (Agfa HealthCare, Belgium) to view CT scans and perform measurements. Axial CT 1 mm slices with pulmonary and soft-tissue viewing window parameters, (window width and level of 1500 HU, -500 HU, 350 HU, and 50 HU, respectively) were used for lung lesion degree assessment and great vessel measurement.

With a large patient flow, an empirical visual scale was used to assess the lung lesions based on the visual assessment of the thickened lung tissue volume according to axial and reconstructed chest coronal and sagittal images. Computer-assisted methods of assessing the affected parenchyma were not used.

The lung lesions, according to grades 1–4 of the CT classification, were assessed according to the approved guidelines [1, 2] and the scheme for describing pathological changes, which was proposed by domestic authors, based on the indication of the pulmonary parenchymal lesion volume. Pulmonary tissue changes (frosted glass, reticular thickening, consolidation, "cobblestone" appearance, "air bronchogram," and inverse "halo") were determined with the assessment of the lesion volume and localization of changes by the lung lobes and segments.

The PA size was measured on axial sections at the widest point, perpendicular to the long axis of the vessel (Fig. 3, *a*, *c*, *e*, *g*). The Asc Ao size was measured at the level of the maximum diameter of the right PA [13]. The accuracy of measurements was limited by the absence of contrast enhancement, cardiac gating, and reduced signal-to-noise ratio. Results were obtained by averaging independent measurements taken by two radiologists. A significant difference in the results of parenchymal lesion assessment or great vessel measurement (>2 mm) considers the opinion of a third radiologist with 15 years of work experience.

Main study outcome

The main study outcome is an assessment of the correlation between various measured PA and Asc Ao parameters and their derivative value with pulmonary parenchymal lesions and age dependence.

Additional study outcomes

An additional study outcome is an assessment of the relationship between the Asc Ao diameter and the patient's age, which confirm the previously established patterns.

Subgroup analyses

In the course of the study, patients of both genders were divided into four groups depending on the lung lesion volume (CT 1-4) and three age ranges (up to 50, 51-69, and over 70 years).



Fig. 3. Examples of CT scans with different PA diameters and lung lesion degrees: a, b, non-dilated PA (27.0 mm) with CT-1 lung lesion (<25%); c, d, dilated PA (30.5 mm) with CT-2 lung lesion (25%–50%); e, f, dilated PA (32.6 mm) with CT-3 lung lesion (50%– 75%); g, h, dilated PA (34.8 mm) with CT-4 lung lesion (over 75%). CT: computed tomography; CT 1-4: adapted scale for visual assessment of the relationship between the patient's general condition severity and the nature and intensity of radiological signs of pulmonary changes in COVID-19; PA: pulmonary artery.

Ethical review

The study was conducted under the Declaration of Helsinki, which was developed by the World Medical Association.

The study was approved by the Independent Ethics Committee of the Moscow Regional Branch of the Russian Society of Roentgenologists and Radiologists, No. 2/2021, dated February 18, 2021.

Informed consent for medical intervention (CT examination) and the use of their medical data for scientific purposes was obtained from all participants.

Statistical analysis

Statistical data processing was conducted using StatSoft Statistica 12.0 software. The required sample size was determined using Altman's nomogram (with a power of 0.8 and a significance level of 0.05). Examination of over 500 patients obtained the minimum sufficient sample size and statistically significant results. The quantitative data distribution normality was checked using the Kolmogorov-Smirnov criterion and adjusted for Lilliefors and Shapiro-Wilk tests. Given the non-normal distribution of PA and Asc Ao parameters in several studied groups and different numbers of patients divided into groups, the reliability of differences was assessed using the nonparametric Kruskal-Wallis (K-W) test, nonparametric ANOVA analog, and median test (MT). The nonparametric Spearman's correlation coefficient (ρ) was used to further assess the relationship of several studied parameters of great vessel sizes, whereas 2×2 tables and nonparametric Fisher F-criterion were used to analyze differences between the groups. The significance level (p) in all tests was <0.05.

RESULTS Study subjects/participants

CT scans of 511 patients both male (52.2%) and female (47.8%), with a mean age of 57 ± 12 years, the median age of 59 years, IQR of 49.0-65.0 years, min. of 31 years, and max. of 84 years, were analyzed. The distribution of patients by gender, lung lesion degree, and age intervals is shown in Table 1. Patient CT scans with different pulmonary parenchymal lesions and corresponding PA measurements are

Main study results

The following significant patterns were obtained: an increased PA diameter with lung lesion severity (K-W, p < 0.001; MT, p < 0.001), increase Asc Ao diameter with the patient's age (K-W, p < 0.001; MT, p < 0.001), increased PA/Asc Ao ratio with lung lesion severity (K-W, p < 0.001; MT, p = 0.008), and decreased parameter with age (K-W, p < 0.001; MT, p < 0.001). Both the correlation of a small increase in the pulmonary trunk diameter with age (K-W, p = 0.094; MT, p = 0.311) and a small increase in the aortic diameter depending on lung lesion degree (K-W, p = 0.061; MT, p = 0.165) were not significant.

Results presented in Fig. 4 determined the following Spearman's correlation coefficients (between the PA diameter and lesion severity): for the entire sample, $\rho = 0.268$, p < 0.005; up to 50 years, $\rho = 0.245$, p < 0.005; 50–70 years, $\rho = 0.229$, p < 0.005; and over 70 years, $\rho = 0.374$, p < 0.005, thus a weak positive relationship was noted. The analysis of differences by Fisher's one-sided F-criterion showed reliable PA dilation (>29 mm) in patients with severe pneumonia (CT 3-4) (Table 2).

Table 1. Distribution of	f patients in groups	s by gender (M: males),	age, and severity of	pneumonia accordin	g to the CT 1-4 scale
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Parameters	CT-1	CT-2	CT-3	CT-4	Total, <i>n</i>
Group I (up to 50 years old)	47 (34 M)	56 (38 M)	19 (14 M)	9 (9 M)	131 (95 M)
Group II (50–69 years old)	84 (36 M)	127 (56 M)	78 (39 M)	24 (19 M)	313 (150 M)
Group III (over 70 years old)	24 (8 M)	18 (4 M)	16 (4 M)	9 (6 M)	67 (22 M)
Total, <i>n</i>	155 (78 M)	201 (98 M)	113 (57 M)	42 (34 M)	511 (267 M)



Fig. 4. Dependence of the PA diameter on age and lung lesion severity in COVID-19.

CT 1-4: adapted scale for visual assessment of the relationship between the patient's general condition severity and the nature and intensity of radiological signs of pulmonary changes in COVID-19; PA: pulmonary artery.

Additional study results

The following additional study results were obtained: a more frequently (37%) observed severe COVID-19 (CT 3–4) in the older patient group (>70 years); less frequently (21%) observed in the second age group (<50 years); and intermediately (32%) observed in the middle age group (50–69 years old).

In addition, an increased PA diameter from CT-1 to CT-4 was 12.9% (calculated as the ratio of the difference in values to half of the sum), whereas the ratio of the standard

deviation to the mean PA values was 13.5%. The relative increase of the PA/Asc Ao ratio was 8.3% and the ratio of the standard deviation to the mean was 11.9%. Therefore, an increased PA diameter was more pronouncedly associated with an increased CT degree compared to the PA/Asc Ao ratio with a smaller difference in the coefficient of variation. In addition, a separate PA changes assessment by this parameter in contrast to the PA/Asc Ao ratio is more preferable.

Adverse events

No adverse events were observed in patients when performing the non-contrast chest CT. Scans were obtained according to the standard non-contrast protocol for chest disease diagnoses.

No adverse events associated with the use of the Airo TruCT mobile CT scanner were registered. The radiation exposure did not exceed the typical values for stationary computed tomographs.

DISCUSSION

Main study result summary

The relationships of PA and Asc Ao diameters and their ratios in patients with different COVID-19-associated pneumonia severity depending on age were studied. A significant positive correlation between the pulmonary trunk diameter and the pulmonary parenchymal lesion in COVID-19 was revealed. A non-significant increased pulmonary trunk diameter with age was shown. A significant aortic dilatation was observed in older age groups, without relation to the increasing disease severity.

Table 2. Distribution of examined	patients by age interval,	pneumonia severity, and	pulmonary artery dilation
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Age interval, years	Parameters				
	CT severity	PA < 29 mm	PA ≥ 29 mm	PA ≥ 29 mm, %	
	CT 3-4	10	18	64,3	
Up to 50	CT 1-2	80	23	22,3	
F0 (0**	CT 3-4	47	51	52,0	
JU-09	CT 1-2	145	62	29,9	
Over 70***	CT 3-4	10	19	65,5	
	CT 1–2	30	16	34,8	

Note. The differences are significant: *p < 0.0001, **p = 0.0001, and ***p = 0.0089 (Fisher's F-criterion). CT: computed tomography; CT 1–4: adapted scale for visual assessment of the relationship between the patient's general condition severity and the nature and intensity of radiological signs of pulmonary changes in COVID-19; PA: pulmonary artery.

Main study result discussion

An equal but not significant increased pulmonary trunk diameter with age (see Fig. 4) was shown. Values for age intervals up to 50, 50-69, and over 70 years amounted for 27.0 ± 3.6, 27.6 ± 3.8, and 28.2 ± 4.1 mm (mean ± standard deviation), respectively. However, a significantly increased PA diameter was shown about pulmonary parenchymal lesion severity in COVID-19-associated viral pneumonia (see Fig. 4). CT grades 1–4 determined the following mean PA values: 26.7 ± 3.4, 26.9 ± 3.4, 28.8 ± 4.1, and 30.4 ± 4.4 mm, respectively. Reverse patterns were shown for the aortic diameter. In addition, a significant change (increase) in the aortic diameter with age $(31.2 \pm 3.7, 34.6 \pm 3.9,$ and 34.8 ± 5.6 mm) for age intervals defined above was observed. No significant change (increase) was noted in the aortic diameter with increasing lung lesion severity (CT 1–4), that is, 33.3 ± 4.9 , 33.7 ± 4.2 , 34.2 ± 3.9 , and 34.6 ± 4.3 mm, respectively. The derived parameter (PA/Asc Ao ratio) showed a significant change (decrease) with age $(0.87 \pm 0.11, 0.80 \pm 0.12, \text{ and } 0.82 \pm 0.15 \text{ RU}, \text{ respectively})$ due to a pronounced increased Asc Ao diameter with age and a slight increased PA diameter. A significant change (increase) in the PA/Asc Ao ratio with increasing degree of the pulmonary tissue lesion (CT 1-4) was shown, that is 0.81 ± 0.13 , 0.81 ± 0.11 , 0.85 ± 0.14 , and 0.88 ± 0.12 RU, respectively.

The obtained mean PA diameter and PA/Asc Ao ratio in the CT–1 group with the smallest lesion volume (26.7 \pm 3.4 and 0.81 \pm 0.13) were slightly higher compared with the mean values defined as normal in the Framingham Heart Study [14], where the mean PA size was 25.1 \pm 2.8 mm and the mean PA/Asc Ao ratio was 0.77 \pm 0.09. This is due to the meth-odological peculiarities, as measurements were performed without cardiac gating, which resulted in increased measured vessel diameters and disproportionate division of the studied groups by gender. The mean measured aortic diameter in the group of patients up to 50 years old (31.2 \pm 3.7 mm) was well comparable with the review results [15], which is devoted to age-related aortic measurements (31.1 \pm 4 mm). In addition, this review noted an increased aortic diameter with age due to decreased vessel wall elasticity.

The groups with severe COVID-2019 (CT 3–4) (see Fig. 2, *b*, *d*, *f*, *h*) demonstrated a statistically significant increased proportion of patients with PA diameter dilation of >29 mm for all ages (Table 2). On average, this group of patients was 60.6%, which was twice the average number of patients with PA dilation of >29 mm in the groups with mild COVID-19 (CT 1–2).

Our data agree well with the previously obtained results [9], where PA dilation in patients with COVID-19-associated pneumonia compared to the data before the disease showed a significantly increased PA/Asc Ao ratio. However, the relationship of these parameters with the pneumonia severity was not pronounced.

The threshold PA diameter values, above which the pulmonary hypertension is reliably stated, were different. For example, the upper limits of the PA diameter, above which patients had dyspnea, were 29 mm in males and 27 mm in females, whereas the critical PA/Asc Ao ratio was 0.9 [13, 16]. Another study proposed PA values of 29 mm and PA/Asc Ao ratio as criteria above which pulmonary hypertension is diagnosed. However, these values showed high sensitivity and insufficient specificity [14]. Alternatively, another proposed threshold of 31.5 mm, above which pulmonary hypertension is registered, showed low specificity [14]. The guidelines for pulmonary hypertension diagnosis and treatment noted a threshold of 29 mm [17].

Thus, the chosen threshold of 29 mm is reasonable. However, this value will probably be adjusted for the additional PA pressure assessment data, which is obtained by ultrasound measurements [18]. The statistical data on a significantly higher number of patients with a PA diameter of >29 mm in groups with a severe course (see Table 2) indicate a possible contribution of pulmonary hypertension to a more severe disease course.

The PA/Asc Ao ratio did not demonstrate threshold values exceeding 0.9 and 1 [9] and reached maximum values of 0.88 \pm 0.12 in the group with a CT-4 degree of lung lesions. A significant increase of this index corresponds to the results reported by P. Spagnolo et al. [9]. While emphasizing the clinical significance of PA dilation, the authors provided reliable data on increased PA diameter and PA diameter/Asc Ao ratio in patients with lethal outcomes compared to those who recovered. This study revealed a weak positive correlation between the lung lesion degree (also divided into severity grades 1–4) and an increased PA diameter. A significant PA diameter dilation was shown in the examined patients compared to the state before the development of pneumonia; however, the sample size was small (45 patients). The necessity of PA dilation correction with PA/Asc Ao ratio determination is confirmed by a reliably smaller, but rather high number of patients with PA dilation in the cohort with small lesion volume (up to 50%) (Table 2). The use of this criterion is reasonable for constitutionally excluding conditioned wide PA. However, this auxiliary comparative criterion is inefficient in patients over 50 years old with hypertension since the aorta also expands due to the vessel wall connective tissue damage [15].

The effectiveness of the PA measurement as a prognostic sign of pulmonary hypertension and an additional severity marker is confirmed by the preferential bilateral process of pulmonary tissue lesions [9]. The PA accumulates increasing hemodynamic distress in the small circle as an additional predictor of COVID-19-associated viral pneumonia severity. PA dilation, which was a reliable relationship with parenchymal lesions, resulted from increasing pulmonary hypertension.

Pulmonary autoimmune diseases show signs of interstitial pneumonia, and pulmonary hypertension develops,

for example, with progressing idiopathic pulmonary fibrosis due to massive interstitial lung lesions [19]. The normal architectonics of the pulmonary vessels is deformed due to the massive pulmonary interstitium inflammation. Capillary constriction and endothelium integrity failure lead to small vessel microthrombosis and sclerosis and, consequently, to an increased hydraulic resistance, which results in pulmonary hypertension [20]. The development of COVID-19-associated viral pneumonia is due to high levels of expression of angiotensin-converting enzyme type 2 (ACE2) in type II alveolar pulmonary cells, which are targets of the spike antigen of severe acute respiratory syndrome coronavirus 2 virus that leads to interstitial lung lesions [11, 12]. The affecting factor that leads to microthrombosis is both a direct vascular endothelium lesion due to ACE2 receptor expression [15, 20, 21] and a possible secondary vascular bed immune lesion and the development of edema and exudation. This is facilitated by the extensive accumulation of complement components in the alveolar walls, which results in endothelial cell damage to the pulmonary capillaries and subsequent coagulation activation [20]. ACE2 receptor system depletion in the pulmonary vascular bed contributes to endothelial dysfunction and inflammation and aggravates the course of atherosclerosis and diabetic angiopathy [15, 20]. In COVID-19, this process leads to delayed adverse reactions in the form of progressing atherosclerotic vascular lesions.

Moreover, the phenomenon of vascular enlargement sign that is a sign of dilated vessels in COVID-19 pneumonia was reported. However, this phenomenon was not deeply studied, and several available studies showed its inconsistent prevalence. This sign occurs in 76.9% of patients with viral pneumonia [7, 10].

Pulmonary hypertension, regardless of its causes, has a secondary negative effect on the course of viral pneumonia, especially with massive lung lesions.

The pulmonary trunk dilation, depending on the severity of pulmonary parenchymal lesions, allows pulmonary hypertension consideration as one of the important signs of COVID-19-associated pneumonia severity. The established pattern allows a more differentiated approach in predicting outcomes of this infectious disease, developing effective therapy methods, and identifying population risk groups for a severe disease course that require increased attention from clinicians.

Study limitations

The study does not answer the question of whether pulmonary hypertension, which is noted as PA dilation, is an initial condition or whether it develops within the course of coronavirus infection. This study did not measure the PA diameter in patients of different ages without signs of viral pneumonia to assess the population prevalence of pulmonary hypertension (PA dilation). Data on the population prevalence of pulmonary hypertension and PA dilation are presented in the mentioned study [20].

CONCLUSIONS

The study showed a statistically significant positive relationship between PA dilation and lung lesion severity in COVID-19-associated pneumonia in all age groups. A higher number of patients with PA dilation of \geq 29 mm (signs of pulmonary hypertension) and a severe degree of pulmonary parenchymal lesions (CT 3-4) is statistically significant. The obtained patterns require further study, which allows a more differentiated approach in predicting outcomes of this infectious disease, developing effective therapy methods, and identifying population risk groups for a severe disease course that require increased clinician attention.

ADDITIONAL INFORMATION

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