

Both typical and atypical radiological changes predict poor COVID-19 outcome in HIV-positive patients from a multinational observational study – data from Euroguidelines in Central and Eastern Europe Network Group

Justyna D. Kowalska<sup>1</sup>, Carlo Bieńkowski<sup>1</sup>, Lukáš Fleischhans<sup>2</sup>, Sergii Antoniuk<sup>3</sup>, Agata Skrzat-Klapaczyńska<sup>1</sup>, Magdalena Suchacz<sup>4</sup>, Nikolina Bogdanic<sup>5</sup>, Deniz Goengin<sup>6</sup>, Cristiana Oprea<sup>7</sup>, Igor Karpov<sup>8</sup>, Kerstin Kase<sup>9</sup>, Raimonda Matulionyte<sup>10</sup>, Antonios Papadopoulos<sup>11</sup>, Nino Rukhaze<sup>12</sup>, Arjan Harxi<sup>13</sup>, David Jilich<sup>2</sup>, Botond Lakatos<sup>14</sup>, Dalibor Sedlacek<sup>15</sup>, Gordana Dragovic<sup>16</sup>, Marta Vasylyev<sup>17</sup>, Antonia Verhaz<sup>18</sup>, Nina Yancheva<sup>19</sup>, Josip Begovac<sup>5</sup> and Andrzej Horban<sup>1</sup>

<sup>1</sup>Department of Adult's Infectious Diseases, Medical University of Warsaw (Poland) and Hospital for Infectious Diseases in Warsaw (Poland), <sup>2</sup>Department of Infectious Diseases, 1<sup>st</sup> Faculty of Medicine, Faculty Hospital Bulovka, Charles University in Prague (Czech Republic), <sup>3</sup>Viral Hepatitis and AIDS Department, Gromashevsky Institute of Epidemiology and Infectious Diseases, Kyiv (Ukraine), <sup>4</sup>Department of Infectious and Tropical Diseases and Hepatology, Medical University of Warsaw (Poland), <sup>5</sup>School of Medicine, University Hospital for Infectious Diseases, University of Zagreb (Croatia), <sup>6</sup>Department of Infectious Diseases and Clinical Microbiology, Faculty of Medicine, Ege University, Izmir (Turkey), <sup>7</sup>Victor Babes Clinical Hospital for Infectious and Tropical Diseases, Carol Davila University of Medicine and Pharmacy, Bucharest (Romania), <sup>8</sup>Department of Infectious Diseases, Belarusian State Medical University, Minsk (Belarus), <sup>9</sup>West Tallinn Central Hospital, Tallin (Estonia), <sup>10</sup>Faculty of Medicine, Vilnius University Hospital Santaros Klinikos, Vilnius University (Lithuania), <sup>11</sup>Medical School, University General Hospital Attikon, National and Kapodistrian University of Athens (Greece), <sup>12</sup>Infectious Diseases, AIDS and Clinical Immunology Center, Tbilisi (Georgia), <sup>13</sup>Infectious Disease Service, University Hospital Center of Tirana (Albania), <sup>14</sup>National Institute of Hematology and Infectious Diseases, South-Pest Central Hospital, National Center of HIV, Budapest (Hungary), <sup>15</sup>Faculty of Medicine in Plzen, University Hospital Plzen, Charles University, Plzen (Czech Republic), <sup>16</sup>Department of Pharmacology, Clinical Pharmacology and Toxicology, School of Medicine, University of Belgrade (Serbia), <sup>17</sup>Astar Medical Center, Lviv (Ukraine), <sup>18</sup>Department for Infectious Diseases, Faculty of Medicine, University of Banja Luka (Bosnia and Herzegovina), <sup>19</sup>Department for AIDS, Specialized Hospital for Active Treatment of Infectious and Paraistic Disease Sofia, Medical University of Sofia (Bulgaria)

Background

In countries with limited resources, people living with HIV (PLWH) may differently present lung infections hindering the differential diagnosis and the choice of treatment during coronavirus disease 2019 (COVID-19). This study aims to investigate the association between radiological changes and poor COVID-19 outcome in PLWH from Central and Eastern Europe.

Methods

Since November 2020 Euroguidelines in Central and Eastern Europe Network Group have started collecting data on HIV/COVID-19 co-infection. In total data was submitted from 16 countries (eCRF) on 557 HIV-positive patients.

Analysis included patients with radiological examination performed. Logistic regression models were used to identify factors associated with death (Figure 1), ICU admission or partial recovery (poor COVID-19 outcome). Factors significant in univariate models (p<0.1) were included in multivariate model. Figure 2, 3 and 4 presents examples of none, typical and atypical radiological changes.

Results

Radiological data were available for 224 (40.2%) patients, 108 (48.2%) had computed tomography and 116 (51.8%) chest X-ray. Of these 211 (94.2%) were diagnosed with RT-PCR, 212 (94.6%) were symptomatic, 123 (55.6%) were hospitalized,

137 (16.6%) required oxygen therapy and 28 (13.1%) either died, was admitted to ICU or only partially recovered. By radiologist's description 138 (61.6%) patients had typical, 18 (8.0%) atypical and 68 (30.4%) no radiological changes. Baseline characteristics are presented in Table.

In univariate models, CD4 count (OR=0.86 [95% CI: 0.76–0.98]), having a comorbidity (2.33 [1.43–3.80]), co-infection with HCV and/or HBV (3.17 [1.32–7.60]), being currently employed (0.31 [0.13–0.70]), being on antiretroviral therapy (0.22 [0.08–0.63]), and having typical (3.90 [1.12–13.65]) or atypical (10.8 [2.23–52.5]) radiological changes were significantly asso-

associated with poor COVID-19 outcome.

In the multivariate model being on antiretroviral therapy (OR=0.20 [95% CI: 0.05–0.80]) decreased the odds of poor COVID-19 outcome. Having a comorbidity (2.12 [1.20–3.72]), as wells as both typical (4.23 [1.05–17.0]) and atypical (6.39 [1.03–39.7]) radiological changes (vs. no changes) increased the odds of poor COVID-19 outcome.

Conclusions

Among HIV-positive patients diagnosed with symptomatic SARS-CoV-2 infections presence of both typical and atypical radiological COVID-19 changes independently predicted poorer COVID-19 outcome.

Table. Baseline characteristics of HIV-positive patients with COVID-19 regarding presence and type of radiological changes.

		Radiological changes			
Characteristic	All n=224	Typical n=138	Atypical n=18	None n=68	P value
Age in years, median (IQR)	45 (35.0-55.0)	47.0 (38.5-57.0)	45.5 (38.0-52.0)	40 (34.5-48.5)	0.0080
BMI in kg/m2, median (IQR)	24.6 (21.4-28.7)	24.6 (21.6-29.0)	20.9 (17.8-24.4)	24.0 (21.3-29.0)	0.0044
Female sex, n (%)	77 (34.7)	44 (32.1)	7 (41.2)	26 (38.2)	0.5788
Currently employed, n (%)	133 (59.4)	82 (59.4)	10 (55.6)	41 (55.6)	0.9358
Ever smoking cigarettes, n (%)	136 (60.7)	54 (61.4)	12 (66.7)	40 (58.8)	0.8308
One or more comorbidity, n (%)	83 (37.0)	53 (38.4)	9 (50.0)	21 (30.9)	0.2848
Nr of comorbidities, median (IQR)	0 (0-1)	0 (0-1)	1 (0-1)	0 (0-1)	0.2281
HCV and/or HBV co-infection, n (%)	39 (17.7)	24 (17.5)	8 (44.4)	7 (10.8)	0.0017
MSM mode of HIV infection, n (%)	64 (28.6)	40 (29.0)	5 (27.8)	19 (27.9)	0.1342
Time since HIV diagnosis in years, median (IQR)	9 (5-14)	10 (6-15)	11.5 (1-19)	7 (3-11)	0.0107
CD4 count in cells/ul, median (IQR)	539 (307-818)	545 (370-830)	344 (140-609)	521 (268-833)	0.1017
HIV VL <50 copies/ml, n (%)	174 (77.7)	109 (62.6)	9 (50.0)	56 (82.3)	0.0114
On antiretroviral therapy, n (%)	203 (90.6)	130 (94.2)	13 (72.2)	60 (88.2)	0.0078
InSTI as third drug in cART, n (%)	134 (65.4)	81 (62.3)	11 (73.3)	42 (70.0)	0.2546
TDF or TAF in backbone, n (%)	146 (65.2)	90 (65.2)	13 (72.2)	43 (63.2)	0.7762
Any COVID-19 symptoms , n (%)	212 (94.6)	135 (97.8)	17 (94.4)	60 (88.2)	0.0160
Hospitalized, n (%)	123 (55.6)	88 (63.8)	13 (72.2)	22 (32.8)	<0.0001
Requiring oxygen therapy, n (%)	37 (16.6)	34 (24.6)	3 (16.7)	0 (0.0)	<0.0001
Died, admitted to ICU or no improvement, n(%)	28 (13.1)	20 (15.3)	5 (33.3)	3 (4.4)	0.0054

Figure 1. Results of the logistic regression model where unadjusted and adjusted odds ratios of death are presented for patients with HIV/COVID-19 co-infection.

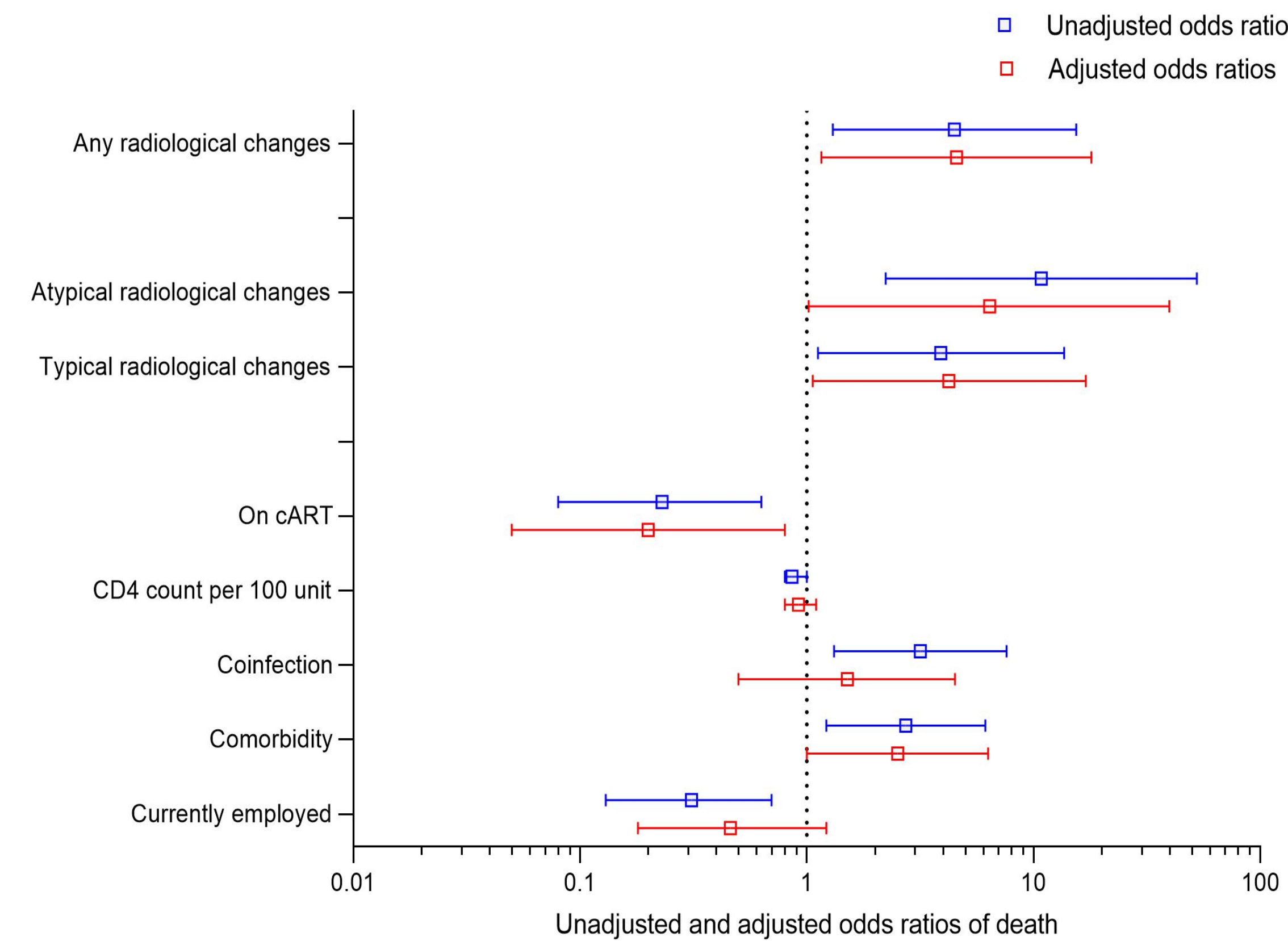
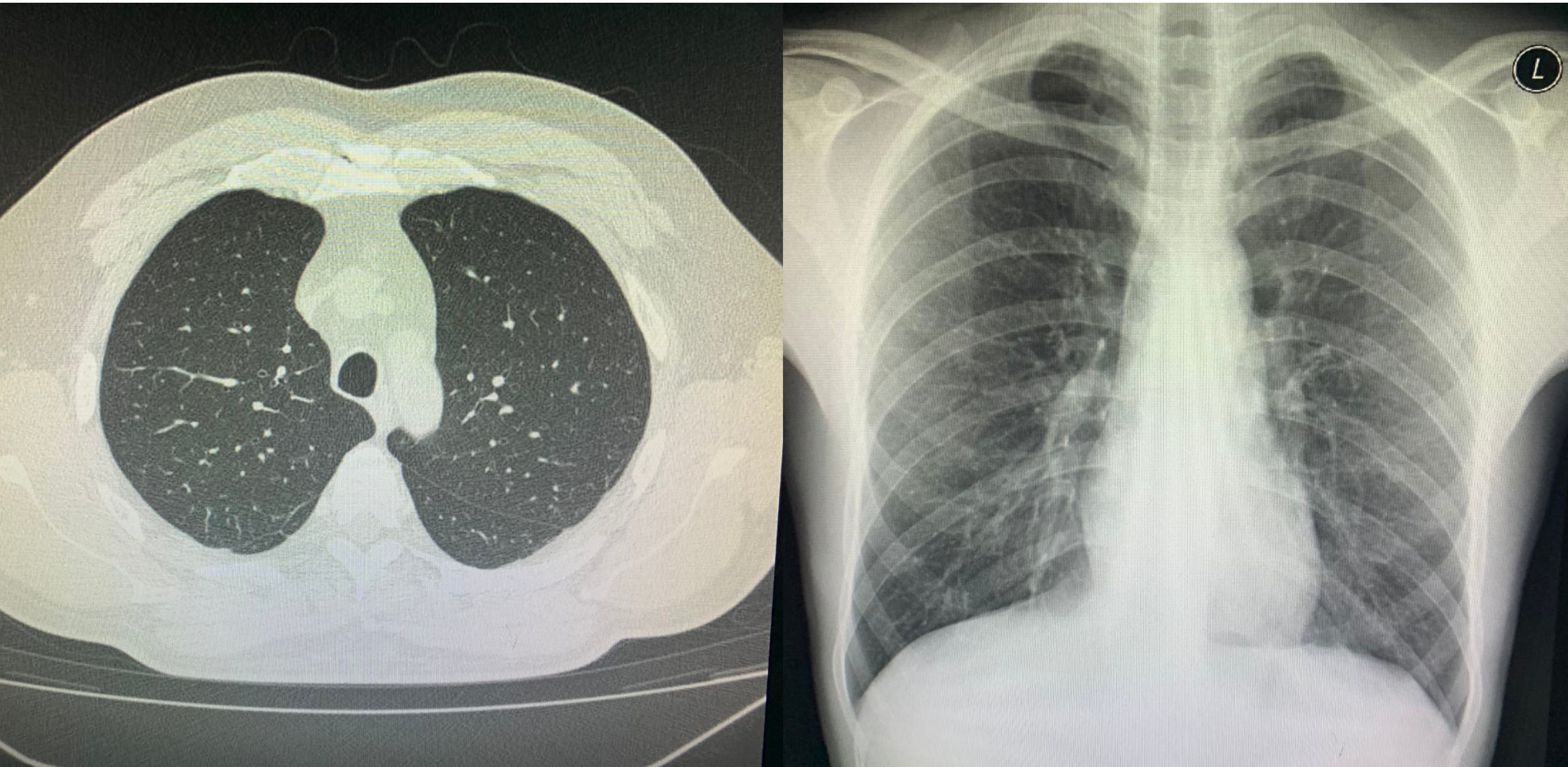
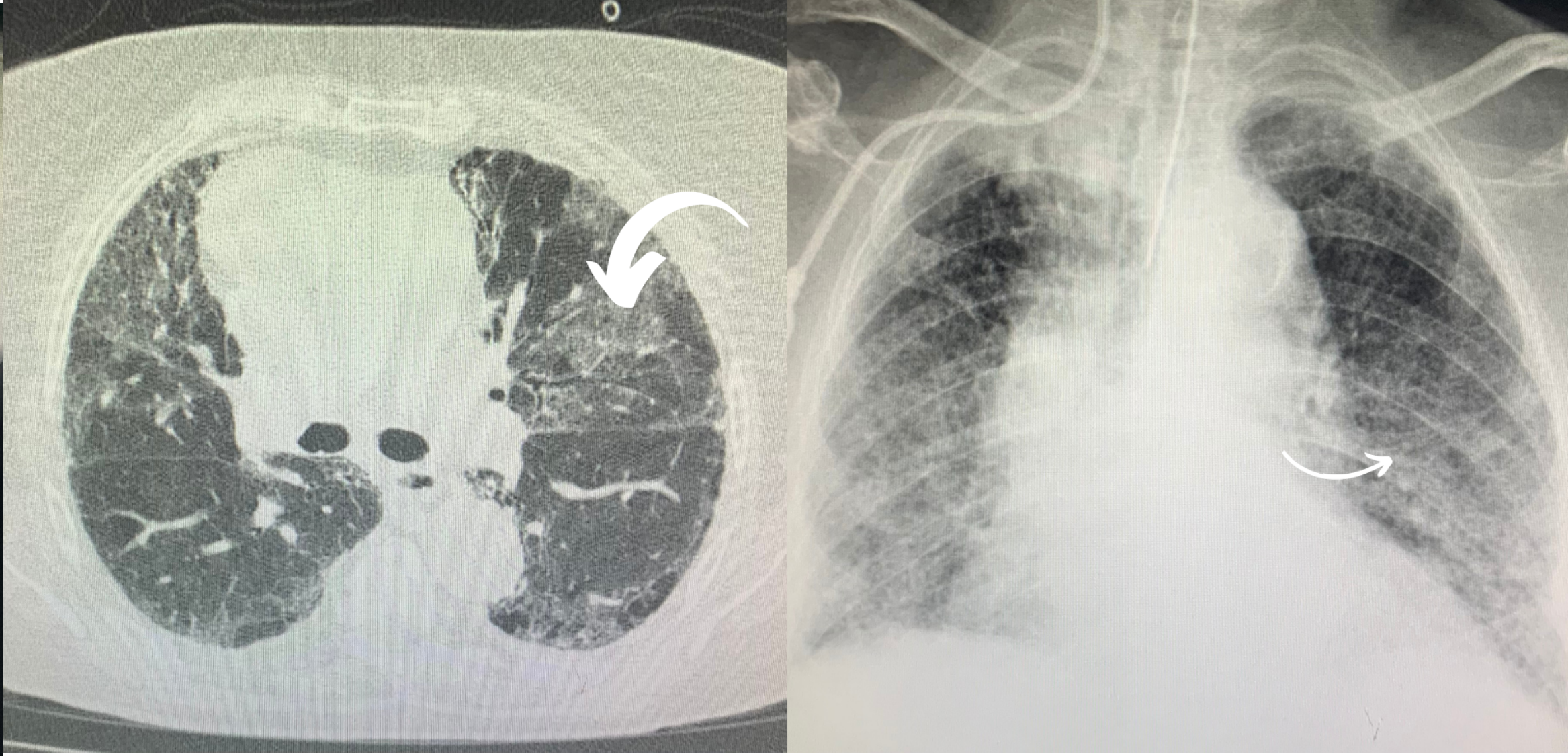


Figure 2. Chest X-ray and CT without any radiological changes



No radiological changes

Figure 3. Chest X-ray and CT of typical radiological changes



Typical radiological changes (bilateral and peripheral ground glass and consolidated opacities)

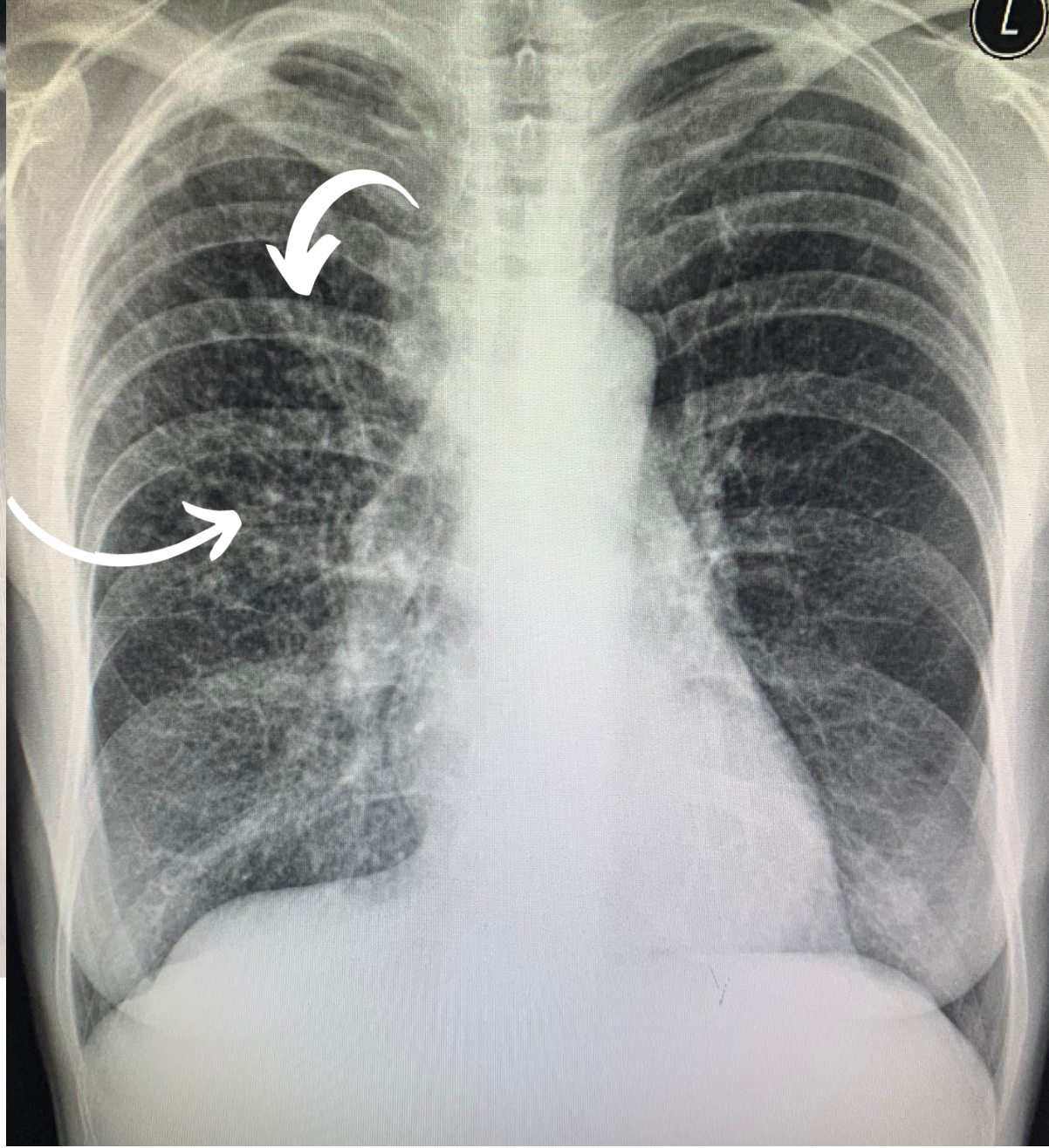


Figure 4. Chest X-ray of atypical radiological changes

Atypical radiological changes (diffuse nodular changes)