# ETTER TO THE EDITOR



# Patterns of HIV and SARS-CoV-2 co-infection in Wuhan, China

Wei Guo<sup>1,2,\*</sup> (b), Fangzhao Ming<sup>3,\*</sup>, Yong Feng<sup>4,\*</sup>, Qian Zhang<sup>5</sup>, Pingzhen Mo<sup>6</sup>, Lian Liu<sup>7</sup>, Ming Gao<sup>8</sup>, Weiming Tang<sup>9,†</sup> and Ke Liang<sup>6,§,†</sup>

<sup>§</sup>Corresponding author: Ke Liang, Department of Infectious Diseases Zhongnan Hospital in Wuhan University, Wuhan, 430071, China. Tel: +86-15327198964. (keliang@whu.edu.cn)

\*These authors equally contributed to this manuscript and are co-first authors. †These authors contributed equally.

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People living with HIV/AIDS (PLWHA) were initially considered to be at an increased risk of infection with SARS-CoV-2 due to their risk of immunosuppresion [1]. However, few studies have reported on co-infections between HIV and SARS-CoV-2 and treatment outcomes, and data to date do not support either increased acquisition or severity of COVID-19 [2,3]. Here we present the clinical and laboratory patterns in 14 co-infected cases, including three asymptomatic SARS-CoV-2 carriers in Wuhan, China, from 16 February to 30 April 2020.

The 14 cases were coded P1 to P14 in order of their infection severity, namely: critical (P1 and P2), severe (P3 to P5), mild (P6 to P11) and asymptomatic (P12 to P14) (Table 1). The case group consisted of 13 men who have sex with men (MSM) and one heterosexual female. Only P10 and P13 had smoking history, and none of them abused alcohol.

The median age of the COVID-19 patients was 56 (range 31 to 71 years old), whereas the three SARS-CoV-2 carriers were 24, 57 and 64 years old. Five patients had hypertension and were on anti-hypertensive agents.

Ten patients and two asymptomatic carriers showed undetectable HIV viral load (HIV-VL) and CD4<sup>+</sup> T lymphocyte count (CD4 count) between 141 to 817/µL prior to SARS-CoV-2 co-infection. Two HIV-VL unsuppressed individuals included patient (P5) who presented with a CD4 count of 34/ μL without having ever received antiretroviral therapy (ART), and an asymptomatic carrier (P12) who presented with a CD4 count of  $29/\mu$ L after two weeks of ART. Four co-morbidity cases presented with a CD4 count less than 200/µL, including two severe cases and two mild cases (All recovered at the end of the study). Although P12 presented with a low CD4 count, he did not even show any symptoms of COVID-19 or any abnormality in the radiograph. This may indicate that low CD4 count did not inevitably lead to death under HIV and SARS-CoV-2 co-infection. On the contrary, the two deaths occurred in cases who had a relatively high CD4 count (350/  $\mu$ L and 695/ $\mu$ L). This finding is consistent with a previous study, which reported one AIDS/COVID-19 patient who presented a low CD4 count  $(13/\mu L)$  and recovered finally [2]. Considering that SARS-CoV-2 might cause multi-organ injury through the inflammatory cytokine storm [4], we hypothesized that previous low CD4 count might act a protective role in preventing hyperimmune response. However, our sample size limits the deduction for a generalized conclusion, and hence large-scale investigation is needed to validate this hypothesis.

The proportion of severe cases (5/11, 45.5%) was higher among the case series than reported in the general population [5]. The COVID-19 mortality rate (18.2%) was also higher than the overall COVID-19 mortality (7.7%, 3869/50333) in Wuhan [6]. The median age among the co-infected cases was older than that of overall patients, and the higher proportion of comorbidities in our patients (8/11, 72.7%) coupled with the senior age might have contributed partially to the higher proportion of severe case and mortality. Living with HIV/AIDS also played a role in leading to comorbidities prior to admission, such as lymphoma and opportunistic infection (pneumocystis pneumonia, PCP).

Reports from laboratory examinations at admission showed leukopenia in six cases and lymphopenia in discrepant six cases. Although the exact immune-pathogenesis of the decrease in lymphocytes in peripheral blood remains unclear, lymphopenia is also reported as a frequent presentation in HIV negative COVID-19 patients, especially in severe cases [5,7]. The pattern of immune cells in AIDS/COVID-19 patients showed minor differences from that of overall COVID-19 patients [8]. However, patients' CD4 count, HIV-VL and inflammatory cytokines were not tested on the admission, which prevented us from knowing more about the immune status at the early stage of SARS-CoV-2 infection in PLHWA. Another limitation should be mentioned that only one ART naïve COVID-19 case was enrolled in our investigation, although knowing the COVID-19 prognosis in ART naïve patients is critical. However, together with the case series

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	Patient 1 <sup>a</sup>	Patient 2 <sup>a</sup>	Patient 3	Patient 4	Patient 5	Patient 6	Patient 7	Patient 8	Patient 9	Patient 10	Patient 11	Patient 12 <sup>b</sup>	Patient 13 <sup>b</sup>	Patient 14 <sup>b</sup>
Demographic and baseline features of the patients	features of the	patients												
Ages (years)	56	68	60	58	38	31	71	63	44	51	44	24	57	64
sex HIV-risk factor	MSM	MSM	MSM	Heterosexual	MSM	MSM	MSM	MSM	MSM	MSM	MSM	MSM	MSM	MSM
Comorbidities	COPD	Hypertension	Diabetes	Hypertension	PCP	None	Hypertension	Hypertension	None	Hypertension	None	Kaposi's	Bronchiectasia	None
			Lymphoma	Atrial fibrillation				Cerebral infarction				Sarcoma	Pulmonary tuberculosis	
ART regimen	+ TZA	AZT +	TDF +	RPV +	None	AZT +	+ TZA	AZT +	EFV +	AZT +	Lpv/r +	EVG/c +	Anemia AZT + 3TC+EFV	TDF +
)	3TC+EFV	3TC+ NVP	3TC+ FFV	TDF+ FTC		3TC+ NVP	3TC+NVP	3TC+ FFV	3TC+ TDF	3TC+ NVP	3TC+	FTC+TAF		3TC+EFV
Latest CD4 count	695	350	141	722	34	420	586	169	506	817	198	29	750	302
(cells/μL) Latest HIV-VL	<20	<20	<20	<20	35600	<20	<20	<20	<20	<20	<20	I	<20	452
(copies/mL)														
Clinical features on admission Diagnosis symptoms	uo													I
Fever	+	+	+	+	+	+	I	+	+	+	+	I	I	I
Cough	+ (dry)	+(dry)	+ (dry)	+ (dry)	1	+ (dry)	+ (dry)	I	I	+ (dry)	1	I	I	I
Lyspnoea Eatimue	+ +	+ +	+ +	+ +	+ +	+ +	I	4	1 4	1 4	+ +	I	1 4	I
raugue Blood nressure (mmHø)	98/45	133/90	117/69	1.39/79	145/93	139/87	- 109/95	153/97	130/80	135/80	116/84	- 124/80	102/75	130/86
Body temperature (°C)	38.6	38.5	39.3	38.9	39.5	40.0	36.8	38.7	38.6	37.5	39.8	Normal	Normal	Normal
Oximetry saturation	76%	82%	90%	87%	91%	94%	86%	97%	896	88%	%06	%66	98%	100%
Chest CT scan	GGO	GGO	660	Bilateral patchy	GGO	Bilateral patchy	660	660	660	660	660	Normal	Bronchiectasia Bilateral	Normal
				shadowing		shadowing							pleural	
WBC (x 10 <sup>9</sup> cells/L) (normal range 3.5	4.36	6.7	3.41	2.29	4.2	1.83	4.51	1.65	3.1	4.44	5.2	4.32	2.16	7.04
LYM (×10 <sup>9</sup> cells/L) (normal range 1.1	1.1	0.35	0.9	0.62	1.55	0.95	1.45	0.41	1.6	1.86	0.68	1.22	1.13	1.44
to 3.2)														
PLT (×10 <sup>9</sup> cells/L) (normal range 125 to 350)	360	65	95	132	267	120	168	118	234	165	232	383	111	210
L-6 (pg/mL) (normal	I	18.62	I	I	9.87	14.08	I	I	I	10.21	I	I	3.25	I
Clinical classification	Critical	Critical	Severe	Severe	Severe	Mild	Mild	Mild	Mild	Mild	Mild	Asymptomatic	Asymptomatic	Asymptomatic
Treatment and clinical outcomes	omes											carrier	carrier	carrier
Admitted to ICU	None	Yes	None	None	None	None	None	None	None	None	None	None	None	None
Mechanical ventilation	None	Yes	None	None	None	None	None	None	None	None	None	None	None	None
Anti-virus Therapy	None	None	Useltamivir	Abidol	Abidol	Useltamivir	None	None	None	None	None	None	None	Abidol
Corticosteroids	None	Yes	None	Yes	Yes	None	None	None	None	None	None	None	None	None
Prognosis	Death	Death	Cure	Cure	Cure	Cure	Cure	Cure	Cure	Cure	Cure	Cure	Cure	Cure
Length of hospital	1	6	18	21	47	11	00	6	24	14	23	10	14	7
buration of positive	I	I	12	17	10	9	9	00	21	10	15	7	4	Ś
nucleic acid														

3TC. Lamivudine: ART, antiretroviral treatment: AZT, Zidovudine: COPD, chronic obstructive pulmonary disease; EFV, Efavirenz; EVG/c, Elvitegravir/cobi; FTC, Emtricitabine: GGO, ground glass opacity; HIV-VL, HIV viral load; ICU, intensive care unit; LPV/r, Lopinavir/Ritonavir; LYM, hymphocytes; MSM, men who have sex with men; NVP, Nevirapine; PCP, pneumooria; PLT, platelet; RPV, Rilpivirine; TDF, Tenofovir; WBC, white blood cells. The two patients died: "Asymptomatic carrier."

published earlier [3], the previous immune status and the clinical course of all the patients in this study still help to sum up some experiences in treating COVID-19 in PLWHA.

The COVID-19 pandemic has had a significant impact on people around the world, including PLWHA, and continues to grow. Health facilities should endeavour to maintain undisputed antiretroviral supply to PLWHA during the pandemic phase to ensure retention in care and treatment success. Co-infected patients who presented with low CD4 count and high HIV-VL, as well as opportunistic infections, should be considered for active treatments as they still have a chance to recover by proper treatment.

#### AUTHORS' AFFILIATIONS

<sup>1</sup>Department of Pathology, Zhongnan Hospital of Wuhan University, Wuhan, China; <sup>2</sup>Department of Pathology, School of Basic Medical Sciences, Wuhan University, Wuhan, China; <sup>3</sup>Wuchang District Center for Disease Control and Prevention, Wuhan, China; <sup>4</sup>State Key Laboratory of Virology/Department of Medical Microbiology, School of Basic Medical Sciences, Wuhan University, Wuhan, China; <sup>5</sup>Qingshan District Center for Disease Control and Prevention, Wuhan, China; <sup>6</sup>Department of Infectious Disease, Zhongnan Hospital of Wuhan University, Wuhan University, Hubei, China; <sup>7</sup>Caidian District Center for Disease Control and Prevention, Wuhan, China; <sup>8</sup>Xinzhou District Center for Disease Control and Prevention, Wuhan, China; <sup>9</sup>Dermatology Hospital, Southern Medical University, and the University of North Carolina at Chapel Hill Project-China, Guangzhou, China

#### COMPETING INTERESTS

None declared.

# AUTHORS' CONTRIBUTIONS

WT and KL both have full access to all of the data. WG, YF, WT and KL conceived the research. FM, QZ, LL, PM and MG collected all the data. WG, YF, and KL analysed the data and drafted the manuscript. All authors have read and approved the final manuscript.

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