

## ORIGINAL ARTICLE

# Electrocardiographic characteristics in patients with coronavirus infection: A single-center observational study

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## Abstract

**Background:** A global outbreak of coronavirus disease (COVID-19), caused by severe acute respiratory coronavirus 2 (SARS-CoV-2), has emerged since December 2019, in Wuhan, China. However, electrocardiographic (ECG) manifestations of patients with COVID-19 have not been fully described. We aim to investigate ECG characteristics in COVID-19 patients and risk factors of intensive care unit (ICU) admission.

**Methods:** This retrospective observational study included the patients with COVID-19 at the Wuhan Asia General hospital between February 10, and 26, 2020. Demographic, clinical, and ECG characteristics were collected, and comparisons were made between the ICU and non-ICU admission groups. Logistic regression was used to identify risk factors of ICU admission.

**Results:** Among 135 included patients (median age: 64 years [interquartile range: 48–72]), ST-T abnormalities (40%) were the most common ECG feature, followed by arrhythmias (38%). Cardiovascular disease (CVD) was presented in 48% of the patients. Six (4.4%) died during hospitalization, and 23 (17.0%) were admitted to the ICU. Compared with non-ICU group, the ICU group showed higher heart rate ( $p = .019$ ) and P-wave duration ( $p = .039$ ) and was more frequently associated with CVD ( $p < .001$ ), ST-T abnormalities ( $p = .007$ ), arrhythmias ( $p = .003$ ), QTc interval prolongation ( $p = .003$ ), and pathological Q waves ( $p < .001$ ). Twenty-seven patients were re-examined ECG during admission, and 17 of them presented new findings compared with their initial ECG presentations. ST-T abnormalities ( $p = .040$ ) and history of CVD ( $p = .0047$ ) were associated with increased risk of ICU hospitalization.

**Conclusions:** COVID-19 is frequently related to cardiovascular manifestations including ECG abnormalities and cardiovascular comorbidities. ST-T abnormalities and CVD at admission were associated with increased odds of ICU admission.

## KEYWORDS

arrhythmias, cardiovascular disease, coronavirus disease, electrocardiogram

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## 1 | INTRODUCTION

The novel coronavirus 2019 (COVID-19) refers to infection caused by SARS-CoV2. As of May 30, more than 5 million individuals have been infected worldwide, which has become a global pandemic. Comprehensive investigation for the performance of cardiac abnormalities should be assessed in patients with COVID-19 because the novel coronavirus has been proven to have definite effects on the cardiovascular system (Madjid et al., 2020). However, there is no specific research on initial electrocardiogram (ECG) abnormalities, which were not uncommon especially in COVID-19 patients with underlying CVDs (Lakkireddy et al., 2020). Here, we retrospectively investigated the patients with COVID-19 in Wuhan Asia General hospital, described the features of ECG and identified the risk factors of the admission to intensive care unit (ICU).

## 2 | METHODS

### 2.1 | Patients

We retrospectively included the patients with confirmed COVID-19 admitted to Wuhan Asia General Hospital in Wuhan, China, from February 10, to 26, 2020. Definition of a case was based on the COVID-19 guidelines of seventh edition established by the Chinese Centers for Disease Control and Prevention (CDC), requiring computed tomographic evidence of pulmonary involvement accompanying RNA detection of the novel coronavirus in nasopharyngeal swab samples. The baseline characteristics were recorded on admission, including age, gender, symptoms, medication history, cardiovascular comorbidities, blood pressure, and heart rate. Admission to the ICU was arranged if vital signs and oxygen saturation were difficult to maintain, or the pulmonary infection progressed rapidly.

The study complied with the edicts of the Declaration of Helsinki (World Medical Association, 2013) and was approved by the Research Ethics Commission of Wuhan Asia General Hospital (WAGHMEC-KY-2020005). Given the retrospective review of this study, informed consent was waived.

### 2.2 | ECG collections

18-lead electrocardiogram (ECG) and laboratory tests are the main observation indicators. The first ECG on admission was recorded with GE MAC 5000 electrocardiograph (General Electric Company). Patients with worsening conditions or heart-related symptoms during admission rechecked for ECG after assessment. The criteria of parameter measurement and ECG diagnosis are based on the recommendation of American Heart Association (AHA) et al (AHA/ACCF/HRS, 2007–2009). ST-T abnormalities were defined as the following criteria: (1) Abnormalities in the ST segment: the ST

segment was measured at 80 ms after J point, and the meaningful change was described as ST-segment depression  $\geq 0.05$  mV, or ST-segment elevation  $\geq 0.10$  mV in limb leads and/or  $\geq 0.20$  mV in chest leads. (2) T-wave abnormalities: a. high and sharp T wave: the peak of T wave was  $>0.5$  mV in limb leads and/or  $>1.5$  mV in chest leads; b. low and flat T wave: the peak of T wave was  $<0.1$  mV in limb leads or  $<0.2$  mV in chest leads; c. bidirectional T wave; d. inverted T wave (inversion depth  $\geq 0.1$  mV). Secondary ST-T changes caused by ventricular pre-excitation, intraventricular conduction block, and ventricular pacing rhythm were not included. QT interval is determined between the earliest ventricular depolarization point and the latest T-wave ending point. QTc interval prolongation was defined as corrected QT (QTc) interval (Bazett formula) was  $\geq 450$  ms in male and  $\geq 460$  ms in female. The measurement and diagnosis of each case were independently analyzed by two experienced cardiologists.

**TABLE 1** Baseline characteristics of patients with COVID-19

	No. (%)			
	Total (n = 135)	ICU (n = 23)	Non-ICU (n = 112)	p value
Age, median (IQR), years	64 (48–72)	71 (55–76)	64 (47–69)	.037
Male	69 (51.1)	17 (73.9)	52 (46.4)	.016
Cardiovascular comorbidity				<.001
Hypertension	44 (32.6)	10 (43.5)	34 (30.4)	.221
Diabetes	20 (14.8)	5 (21.7)	15 (13.4)	.481
Hyperlipidemia	2 (1.5)	1 (4.3)	1 (0.9)	.313
Coronary heart disease	11 (8.1)	5 (21.7)	6 (5.4)	.028
Previous myocardial infarction	2 (1.5)	0	2 (1.8)	>.999
Stroke	7 (5.2)	4 (17.4)	3 (2.7)	.017
Previous PCI procedure	7 (5.2)	4 (17.4)	3 (2.7)	.017
Permanent pacemaker	2 (1.5)	0	2 (1.8)	>.999
Medication history				
$\beta$ receptor blocker	4 (3.0)	1 (4.3)	3 (2.7)	.531
Calcium channel blocker	20 (14.8)	6 (26.1)	14 (12.5)	.178
ACEIs/ARBs	13 (9.6)	2 (8.7)	11 (9.8)	>.999
Oral anti-diabetic agents	15 (11.1)	3 (13.0)	12 (10.7)	>.999
Antiplatelet agents	6 (4.4)	1 (4.3)	5 (4.5)	<.999

Abbreviations: ACEIs, angiotensin-converting enzyme inhibitors; ARBs, angiotensin receptor blockers; IQR, interquartile range; PCI, percutaneous coronary intervention.

**TABLE 2** ECG features of patients with COVID-19

	No. (%)			p value
	Total (n = 135)	ICU (n = 23)	Non-ICU (n = 112)	
<b>Basic parameters</b>				
Heart rate, bpm	81.4 ± 17.8	93.2 ± 26.4	79.0 ± 14.6	.019
P-wave duration <sup>a</sup> , median (IQR), ms	110 (101–116)	110 (92–111)	110 (102–117)	.039
PR interval <sup>d</sup> , ms	157.3 ± 24.7	151.6 ± 23.7	158.4 ± 24.9	.274
QRS complex duration, median (IQR), ms	93 (86–102)	94 (85–105)	92 (86–102)	.572
Frontal plane QRS axis, degree	33.6 ± 44.6	32.2 ± 75.4	33.8 ± 35.7	.921
QTc interval, ms	435.0 ± 31.7	447.8 ± 52.9	432.4 ± 24.8	.184
<b>ECG diagnosis</b>				
Normal	30 (22.2)	1 (4.3)	29 (25.9)	.024
ST-T abnormalities <sup>b</sup>	54 (40.0)	15 (65.2)	39 (34.8)	.007
Conduction block	16 (11.9)	4 (17.4)	12 (10.7)	.584
First degree AVB	5 (3.7)	0	5 (4.5)	.588
Second-degree type I AVB	1 (0.7)	1 (4.3)	0	.170
Third-degree AVB	2 (1.5)	2 (8.7)	0	.028
Right bundle branch block	5 (3.7)	2 (8.7)	3 (2.7)	.201
Left anterior fascicular block	2 (1.5)	0	2 (1.8)	>.999
Intraventricular conduction delay	4 (3.0)	1 (4.3)	3 (2.7)	.531
Atrial and ventricular arrhythmias	20 (14.8)	9 (39.1)	11 (9.8)	.001
Atrial premature complex	5 (3.7)	3 (13.0)	2 (1.8)	.035
Atrial tachycardia	5 (3.7)	4 (17.4)	1 (0.9)	.002
Atrial fibrillation	8 (5.9)	1 (4.3)	7 (6.3)	>.999
Ventricular premature complex	6 (4.4)	4 (17.4)	2 (1.8)	.006
Sinus tachycardia	9 (6.7)	1 (4.3)	8 (7.1)	.976
Sinus bradycardia	11 (8.1)	1 (4.3)	10 (8.9)	.754
Arrhythmias (total) <sup>c</sup>	51 (37.8)	15 (65.2)	36 (32.1)	.003
QTc interval prolongation	18 (13.3)	8 (34.8)	10 (8.9)	.003
Abnormal Q wave	11 (8.1)	7 (30.4)	4 (3.6)	<.001
Extensive anterior wall (V1–V6, I, aVL)	1 (0.7)	—	—	—
Antero-septal wall (V1–V3)	3 (2.2)	—	—	—
Anterior wall (V3–V5)	2 (1.5)	—	—	—
Inferior wall (II, III, aVF)	5 (3.7)	—	—	—
Posterior wall (V7–V9)	1 (2.2)	—	—	—
Low voltage in limb leads	6 (4.4)	3 (13.0)	3 (2.7)	.101
Left ventricular hypertrophy	8 (5.9)	1 (4.3)	7 (6.3)	>.999
Left atrial abnormality	20 (14.8)	1 (4.3)	19 (17.0)	.219
Right atrial abnormality	2 (1.5)	0	2 (1.8)	>.999
Others <sup>d</sup>	4 (3.0)	—	—	—

Abbreviation: AVB, atrioventricular block.

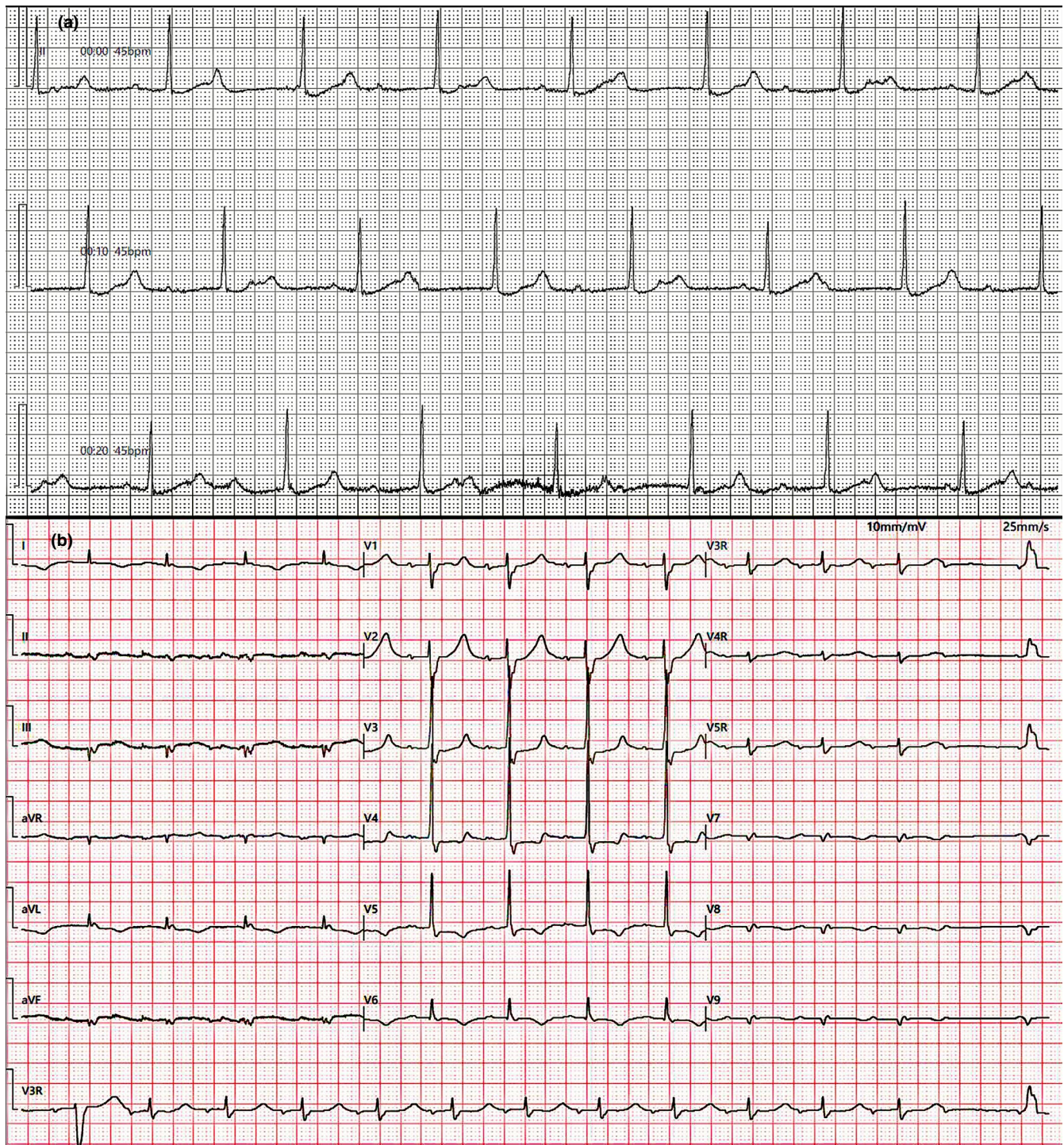
<sup>a</sup>P-wave duration and PR interval were not measured in a total of 11 patients (4 in ICU group and 7 in non-ICU group) with atrial fibrillation, the third-degree AVB or ventricular pacing rhythm.

<sup>b</sup>ST-T abnormality: including 5 cases of acute myocardial infarction.

<sup>c</sup>Arrhythmias: including sinus tachycardia, bradycardia, conduction block, atrial and ventricular arrhythmias. Multiple arrhythmias from one patient were counted once.

<sup>d</sup>Others: including short PR interval and ventricular pacing rhythm.





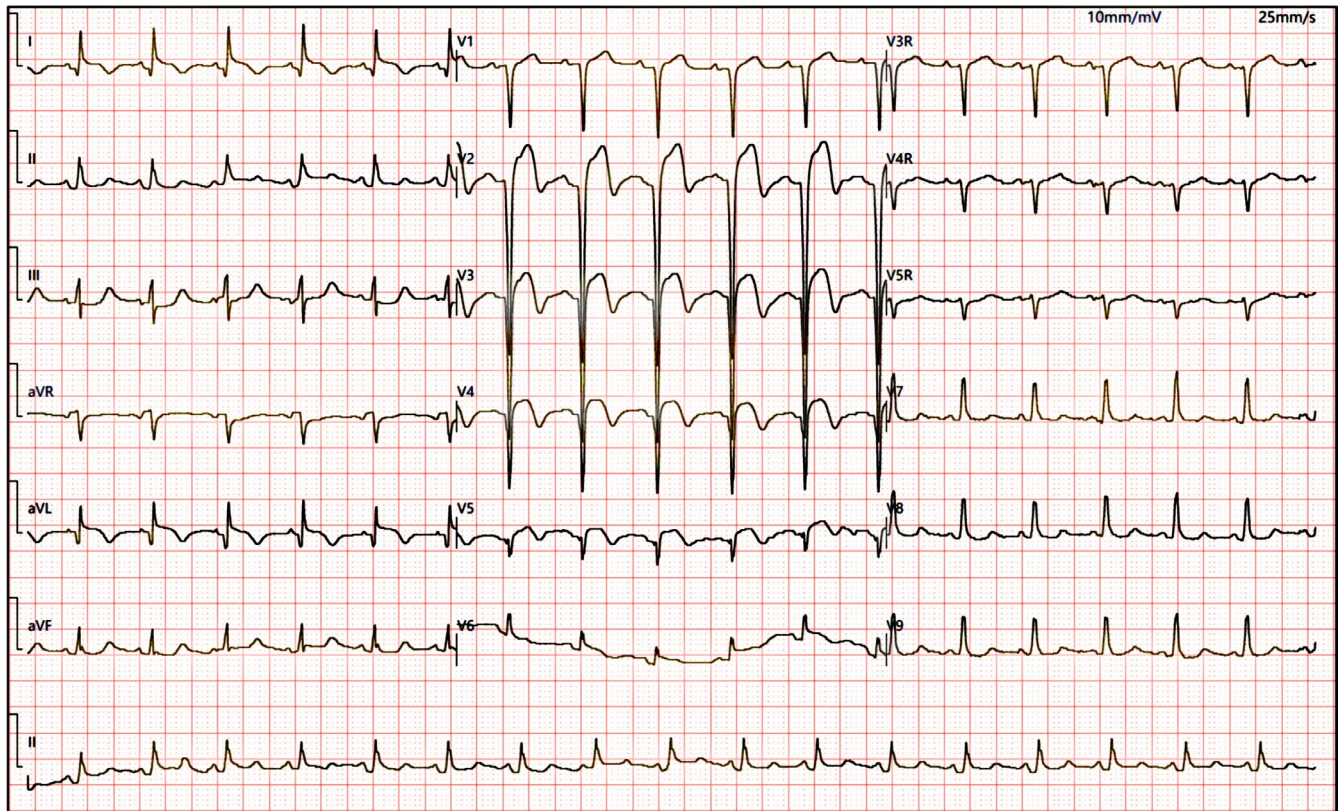
**FIGURE 1** (a) ECG of the third-degree atrioventricular block in a patient with COVID-19 combined with acute myocarditis. (b) ECG of the second-degree type I atrioventricular block in a patient with COVID-19 combined with coronary heart disease

### 2.3 | Statistical analysis

Categorical variables were depicted as frequency rates with percentages, and continuous variables were described using median and interquartile range (IQR) values. After the normal distribution test, Mann-Whitney  $U$  test, chi-square test, or Fisher's exact test were used to compare differences between the ICU group and non-ICU group where appropriate.  $p < .05$  was considered statistically

significant. The inclusion criteria of independent variables in binary regression analysis were as follows: a. dichotomous variables with statistically significant differences; b. variables with no potentially collinearity (such as CVDs and coronary heart disease, arrhythmias, and atrial tachycardia); c. variables associated with adverse cardiac events and may aggravate the condition in patients with COVID-19. All statistical analyses were calculated by Statistical Package for the Social Sciences (SPSS) version 21.0 software.





**FIGURE 2** ECG of acute extensive anterior myocardial infarction in a patient with COVID-19

### 3 | RESULTS

#### 3.1 | Baseline characteristics

A total of 135 hospitalized patients with COVID-19 were included in this analysis. Twenty-three (17.0%) patients were admitted or transferred to the ICU since the aggravation of infection and multiple organs injury. Six (4.4%) patients died during hospitalization. The median age of 135 patients was 64 years (IQR, 48–72), and over half of patients were male (Table 1). About 45.2% of patients had cardiovascular diseases (CVDs). Hypertension (32.6%), diabetes (14.8%), and coronary heart disease (8.1%) were the top three most common comorbidities. The most common cardiovascular medication prescribed was calcium channel blockers, followed by oral antidiabetic agents and angiotensin-converting enzyme inhibitors/angiotensin receptor blocker (ACEI/ARB). Compared with patients who were not admitted to ICU ( $n = 112$ ), patients who received ICU care ( $n = 23$ ) are significantly older (71 years [55–76] vs. 64 years [47–69];  $p = .037$ ) and were more likely to complicate with underlying CVDs (18 [78.3%] vs. 43 [38.4%];  $p < .001$ ).

#### 3.2 | ECG characteristics

ECG characteristics showed a variety of difference between the two groups. Patients in the ICU presented higher heart rate

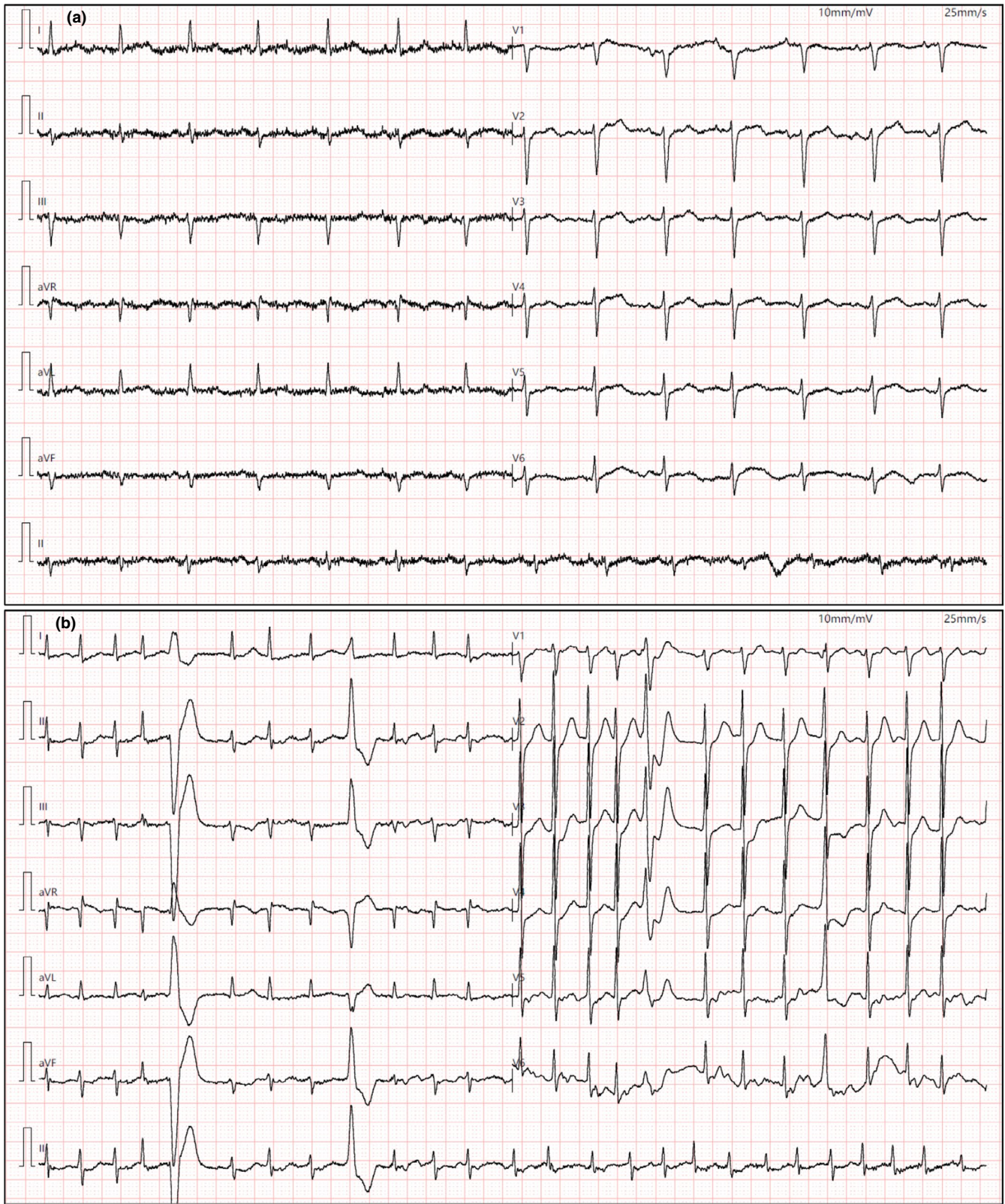
**TABLE 3** New electrographic findings in patients with COVID-19 during admission

New ECG findings <sup>a</sup>	No.
Pathological Q wave	2
ST-T abnormality	5
Atrial premature complex	1
Ectopic atrial rhythm	1
Atrial fibrillation	3
Ventricular premature complex	1
Limb lead low voltage	1
Intermittent bundle branch block	1
QTc prolongation	1
Left ventricle hypertrophy	3
Resolved ECG abnormalities	
Sinus tachycardia resolved	1
Atrial tachycardia resolved	1
ST-T abnormalities resolved	1
QTc prolongation resolved	1

<sup>a</sup>A total of 27 patients re-examined ECG during admission, and 10 of them presented no significant changes compared with their initial ECG results.

( $93.2 \pm 26.4$  vs.  $79.0 \pm 14.6$  bpm;  $p = .019$ ) and P-wave duration (110 [92–111] vs. 110 [102–117] ms;  $p = .039$ ) (Table 2). ST-T abnormalities (40%) were the most common ECG changes in patients





**FIGURE 3** (a) A normal ECG of a patient with COVID-19 at admission. (b) The ECG recorded 6 days after admission indicated new atrial fibrillation with rapid ventricular response and ventricular premature complex

with COVID-19, followed by total arrhythmias (37.8%) and left atrial abnormality (14.8%). Two patients with acute myocarditis developed third-degree atrioventricular block (Figure 1a), and

one patient with coronary heart disease presented second-degree type I atrioventricular block (Figure 1b). Five patients were admitted with ST-segment elevation, which was later diagnosed as acute

**TABLE 4** Risk factors of the admission to ICU

	B	SE	Wal	Sig.	Exp (B)	95% CI
Cardiovascular disease	1.188	0.598	3.940	0.047	3.280	1.015–10.598
Male	1.127	0.599	3.545	0.060	3.087	0.955–9.983
Arrhythmias	0.594	0.582	1.042	0.307	1.811	0.579–5.662
ST-T abnormalities	1.141	0.555	4.221	0.040	3.130	1.054–9.298
QTc interval prolongation	0.727	0.640	1.287	0.257	2.068	0.589–7.256

<sup>a</sup>Binary logistic regression: enter method.

ST-segment elevation myocardial infarction by coronary angiography (Figure 2). Compared with the non-ICU group, patients in ICU were more prone to manifest ST-T abnormalities (15 [65.2%] vs. 39 [34.8%];  $p = .007$ ), QTc interval prolongation (8 [34.8%] vs. 10 [8.9%];  $p = .003$ ), and pathological Q wave (7 [30.4%] vs. 4 [3.6%];  $p < .001$ ), as well as arrhythmias (Table 2). During hospitalization, 27 patients were re-examined ECG and 17 of them presented new findings compared with their initial ECG results (Table 3). ST-T abnormalities (5/17), atrial fibrillation (3/17) and left ventricular hypertrophy (3/17) were the three most common emerging ECG changes (Figure 3). The initial ECG abnormalities of 4 patients were resolved in re-examination.

### 3.3 | Risk factors for admission to ICU

After the procedure elaborated earlier, a history of CVD, cardiac arrhythmias, gender (male), ST-T abnormalities, and QTc prolongation were included into our final binary logistic regression. We found that ST-T abnormalities ( $p = .040$ , C.I. [1.054–9.298]) and CVD history ( $p = .047$ , C.I. [1.015–10.598]) at admission were associated with increased odds of ICU admission (Table 4).

## 4 | DISCUSSION

This retrospective study revealed the ECG characteristics and risk factors of ICU admission in patients with COVID-19. Previous studies have confirmed that SARS-CoV can cause the ECG abnormalities in both animals and humans (Alexander et al., 1999; Yu et al., 2006). As a virus homologous to SARS-CoV, SARS-CoV-2 was also found to be potentially arrhythmogenic. Wang et al. showed 16.7% (23/138) of COVID-19 patients combined with arrhythmias and were more common in the ICU group (44.4% vs. 6.9%,  $p < .001$ ) (Wang et al., 2020). However, comprehensive investigations on the ECG characteristics in patients with COVID-19 are still sparse.

In our study, we found that only 25% of COVID-19 patients presented generally normal ECG and ST-T abnormalities were the most prominent ECG manifestation among the patients with COVID-19. The ST-segment elevation in COVID-19 patients is mainly due to

obstructive/nonobstructive coronary artery diseases, myocarditis, pericarditis, or stress cardiomyopathy (Dehghani et al., 2020; Inciardi et al., 2020). However, the exact reasons for the other nonspecific ST-T abnormalities remain unclear. Infection-mediated myocardial injury and prior CVD comorbidities may contribute.

Cardiac arrhythmias (37.8%) were the second most common ECG change and should be taken seriously because some arrhythmias can be life-threatening. Two patients in our cohort occurred complete AVB on admission, and one of them had undertaken a permanent pacemaker implantation due to syncope. Recent studies have shown fatal arrhythmias including complete AVB, polymorphic ventricular tachycardia, and ventricular fibrillation in patients with COVID-19 (Kochav et al., 2020; Kir et al., 2020; Guo et al., 2020). The high prevalence of arrhythmias may be attributed to metabolic disorders, hypoxia, neurohormonal imbalance, or inflammatory reaction in patients with or without prior CVD (Driggin et al., 2020).

Although QTc prolongation was more common in the ICU group (34.8% vs. 8.9%), there was no significant difference in values between the two groups in our cohort ( $p = .184$ ). Jain S et al. reported that 19.7% (103/524) of patients with COVID-19 had prolonged QTc interval and were more likely to be admitted to the ICU (Jain et al., 2020). Drugs for treating COVID-19 were regarded as the main cause of QTc prolongation, and this effect may be amplified when multiple QT-prolonging drugs were used in combination. Hydroxychloroquine (HCQ), lopinavir/ritonavir, and azithromycin have been confirmed to prolong the QTc interval in patients with COVID-19 (Sacher et al., 2020). Moreover, some inflammatory factors (TNF- $\alpha$ , IL-6) and electrolyte disturbance may potentially contribute to this process (Cipriani et al., 2020). Abnormal Q wave often indicated myocardial necrosis or stunned, which may be caused by myocarditis and were associated with a poor prognosis (Buttà et al., 2020). This partially explained why the Q waves are more common in the ICU group.

As for risk factors, ST-T abnormalities and history of CVD were identified to increase the odds of ICU admission in our cohort. ST-T changes were not uncommon in patients with COVID-19 and often indicated ventricular repolarization abnormalities, which may be a potential cause of severe arrhythmias (Angeli et al., 2020). Comprehensive assessment on drug interactions, electrolyte abnormalities, and ECG changes was imperative for the patients with CVD history on admission, because they were



more sensitive to factors that could potentially lead heart injury (Kochi et al., 2020).

Finally, ECG changes during admission cannot be ignored or may be even more important. Although we did not re-examine the ECG for all the patients because of minimizing exposure of healthcare workers, ECG changes were observed in more than 60% of patients (17/27). New or worsening ECG abnormalities in patients with pneumonia often indicated the degree of cardiac involvement (Corrales-Medina et al., 2013; Stein et al., 2012). Hui et al. (Hui et al., 2020) reported two critically ill patients with COVID-19 developed new AF during hospitalization and eventually died.

To the best of our knowledge, this is the largest retrospective study regarding initial ECG manifestations among patients with COVID-19. The novel findings of this study are the CVD comorbidities and ECG abnormalities were not uncommon in patients with COVID-19 and some of which could be life-threatening. History of CVD and ST = T abnormalities were associated with increased odds of ICU admission. The ECG is a simple and useful screening tool, which can help us promptly assess the cardiac complications while avoiding adverse cardiac events during admission.

#### 4.1 | Study limitations

The following limitations of our study should be noted. First, not all patients had serial ECG recordings, and therefore, some transient ECG changes might have been missed. Second, because of the difficulty in obtaining the prior medical data due to the COVID-19 pandemic, there is a lack comparison of patient's previous ECG. Finally, as the logistic regression analysis in this study did not include laboratory indicators and some confirmed risk indicators, there may be selection bias in the process.

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#### CONFLICT OF INTEREST

The authors declared that they have no conflicts of interest to this work. We declare that we do not have any commercial or associative interest that represents a conflict of interest in connection with the work submitted.

#### AUTHOR CONTRIBUTIONS

Yi Li: Conceptualization, Software, Formal analysis, Resources, Writing original draft. Tong Liu: Methodology, Writing and Revision. Gary Tse: Investigation, Revision. Mingxiang Wu: Supervision. Jingjing Jiang: Data collection and curation. Ming Liu: Correction. Liang Tao: Validation.

#### ETHICS

The study complied with the edicts of the Declaration of Helsinki (World Medical Association, 2013) and was approved by the

Research Ethics Commission of Wuhan Asia General Hospital (WAGHMEC-KY-2020005). Given the retrospective review of this study, informed consent was waived.

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