

Prognostic Value Of 12-Leads Admission Electrocardiogram In Low-Risk Patients Hospitalized For Covid-19

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Prognostic Value Of 12-Leads Admission Electrocardiogram In Low-Risk Patients Hospitalized For Covid-19.

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ECG predictors of adverse clinical course.

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ABBREVIATIONS LIST

- 1 ACE-I, angiotensin-converting-enzyme inhibitors
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3 AF, atrial fibrillation
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5 COVID-19, coronavirus disease 2019
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7 BMI, body mass index
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9 BZD, benzodiazepine
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11 CIs, confidence intervals
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13 Cr, creatinine
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15 CRP, C-reactive protein
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17 DM, diabetes mellitus
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19 ECG, electrocardiogram
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21 eGFR, estimated glomerular filtration rate
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23 HR, heart rate
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25 Hs-cTn-I, high-sensitivity cardiac troponin-I
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27 ICU, intensive care unit
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29 IL, interleukin
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31 IMV, invasive mechanical ventilation
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33 LAFB, left anterior fascicular block
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35 LBBB, left bundle branch block
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37 LPFB, left posterior fascicular block
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39 LVH, left ventricular hypertrophy.
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41 LDH, lactate dehydrogenase
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43 PCT, procalcitonin
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45 PRWP, poor R wave progression
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47 QTc, corrected QT interval
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49 RBBB, right bundle branch block
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51 RV, right ventricular
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53 SARS-CoV-2, severe acute respiratory syndrome coronavirus 2
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55 TIA, transient ischemic attack
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57 TG, triglyceride
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59 VPC, ventricular premature complex
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ABSTRACT

Background: Cardiac involvement significantly contributes to coronavirus disease 2019 (COVID-19) mortality. 12-lead electrocardiogram (ECG) represents a fast, cheap, and easy to perform exam with the adjunctive advantage of the remote reporting possibility. In this study, we sought to investigate if electrocardiographic parameters are able to identify patients, deemed at low-risk at admission, who will face in-hospital unfavourable course.

Methods: From March 1, 2020 through March 30, 2021, 384 consecutive patients with confirmed low-risk COVID-19 were hospitalized at the Azienda Ospedaliero Universitaria Policlinico di Bari (Italy). Criteria for low-risk were: admission to the division of Pneumology or Infectious Diseases, no need for immediate (within 24 hours from admission) transfer to Intensive Care Unit or for respiratory support with invasive mechanical ventilation (IMV) or for circulation support (either mechanical or pharmacological). Admission ECGs were reviewed and interpreted by two expert cardiologists. The primary outcomes were in-hospital death and the composite outcome of in-hospital death and IMV.

Results: In low-risk COVID-19 patients, atrial fibrillation (AF), poor R wave progression (PRWP), tachycardia, and right bundle branch block (RBBB) resulted as statistically significant and independent predictors of in-hospital all-cause mortality; AF, PRWP, Tachycardia, RBBB, and corrected QT interval showed to be statistically significant and independent risk factors for the occurrence of the composite endpoint of death and IMV.

Conclusions: Our study demonstrated for the first time that RBBB and PRWP, assessed upon admission with ECG, are associated with unfavourable clinical course in a baseline low-risk population hospitalized for COVID-19.

KEY WORDS

- COVID-19
- Electrocardiogram
- Atrial fibrillation
- Right bundle branch block

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INTRODUCTION

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3 The severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) has caused a worldwide
4 increase in hospitalizations for pneumonia and multi-organ disease. Overall, in-hospital mortality from
5 coronavirus disease 2019 (COVID-19) is approximately 15-20%, ranging from less than 5% among
6 patients younger than 40 years up to 60% for octogenarians and older patients.¹⁻⁵ When clinical
7 conditions require intensive care unit (ICU) admission, the reported mortality rate raises to
8 approximately 40%.^{6,7}

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11 There is increasing evidence that cardiac involvement significantly contributes to COVID-19
12 mortality.⁸⁻¹² Cardiovascular complications associated with COVID-19 comprise myocarditis, acute
13 myocardial infarction, acute heart failure, cardiomyopathy, and dysrhythmias. Cardiac damage
14 recognises different pathogenic mechanisms including direct myocardial injury, thromboembolic
15 events, and medication toxicity or adverse drug to drug interactions.¹³⁻¹⁵

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18 Despite cardiac involvement is frequent and potentially life-threatening, not all inpatients
19 currently undergo a complete cardiological evaluation at admission, especially during the climax of the
20 pandemic waves when high pressure is exerted on the healthcare systems. In this study, we sought to
21 investigate if admission 12-lead electrocardiogram (ECG) can provide relevant information to predict
22 the hospital prognosis of COVID-19 inpatients presenting with an initial low-risk clinical profile. ECG
23 represents indeed a fast, cheap, easy to perform, and widely used exam with the adjunctive advantage,
24 in the context of a highly infectious disease, of the remote reporting possibility.

MATERIALS AND METHODS

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27 Our observational single-center cohort study included all consecutive patients with confirmed
28 SARS-CoV-2 infection hospitalized at the Azienda Ospedaliero Universitaria Policlinico di Bari
29 (Italy) from March 1, 2020 through March 30, 2021 with a low clinical risk profile at admission.
30 All cases of COVID-19 were confirmed by real-time reverse-transcriptase polymerase chain reaction on
31 nasopharyngeal swabs. Patients were deemed at low-risk if admitted to the division of Pneumology or

1 Infectious Diseases and did not require immediate (within 24 hours from admission) transfer to ICU,
2 respiratory support with invasive mechanical ventilation (IMV), or circulation support either mechanical
3 or pharmacological. Admission ECG was available for all patients. ECGs were reviewed and
4 interpreted by two expert cardiologists blinded to each other and to the clinical course of the patients;
5 disagreements were resolved by consensus. ECG interpretation was standardized and focused on the
6 following data: rhythm, heart rate, QRS axis, QRS amplitude and morphology, ST segment and T wave
7 morphology, and QT interval. The ST segment level was measured at 80 ms after the J-point, and ST
8 depression or elevation was adjudged if ≥ 1 mm in ≥ 2 contiguous leads. Patients with a duration of the
9 QRS complex ≥ 120 ms were excluded from ST, T, and QT analysis. QT interval was corrected (QTc)
10 using the Bazett formula.

11 Laboratory data were extracted from the electronic health record and comprised, among others,
12 the following parameters: D-dimer, high-sensitivity cardiac troponin-I (Hs-cTn-I), complete blood
13 counts, C-reactive protein (CRP), interleukin-6 (IL-6), lactate dehydrogenase (LDH), and blood gas
14 test.

15 The primary outcomes were in-hospital death and the composite outcome of in-
16 hospital death and IMV.

17 Patients and/or the public were not involved in the design, conduct, reporting or dissemination
18 plans of this research.

19 Data availability: the data associated with the paper are not publicly available but are available
20 from the corresponding author on reasonable request.

21 STATISTICAL ANALYSIS

22 Statistical analysis was performed using SPSS version 22 software (IBM, Inc., Armonk, NY).
23 Data are presented as means \pm standard deviations for continuous variables and proportions for
24 categorical variables. Categorical variables were analyzed using the chi-square analysis and Fisher's
25 exact test for counts < 5 . Continuous variables were compared using paired Student's t-tests. P-value

1 <0.05 was determined to be significant. The relationship between outcomes and both ECG and clinical
2 characteristics was examined using univariate logistic regression analyses with odds ratio (OR) and 95%
3 confidence intervals (CIs). Statistically significant ($p < 0.05$) predictors of in-hospital mortality and of in-
4 hospital death and/or IMV were entered into multivariable logistic regression models.
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10 RESULTS

11 384 patients with low-risk COVID-19 at the time of hospitalization were included in the study.
12 Clinical characteristics of the population at admission are shown in **Table I-II** (home therapy,
13 laboratory and radiographic findings are depicted in **Supplementary Digital Material 1:**
14 **Supplementary Table I-II**). During hospital stay a total of 29 (7.5%) patients died and 44 (11.5%)
15 reached the composite outcome of death and IMV.
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23 Cardiovascular comorbidities such as diabetes mellitus (DM) type 2, arterial hypertension,
24 dyslipidemia, history of any heart disease and the use of any antiplatelet therapy or beta-blocker drugs
25 resulted more prevalent in the group of patients facing in-hospital death or the composite outcome of
26 in-hospital death and IMV.
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32 The electrocardiographic findings at admission are presented for the whole population and, in a
33 comparative fashion, by groups based on the occurrence of the considered clinical outcomes (in-
34 hospital all-cause mortality and the composite of death and IMV); significant correlations with main
35 outcomes are depicted in **Table III** (overall data are available in **Supplementary Digital Material 1:**
36 **Supplementary Table III**).
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43 In the univariate logistic regression analyses, electrocardiographic predictors of in-hospital death
44 resulted: atrial fibrillation (AF), poor R wave progression (PRWP) in V1-V6, tachycardia, low QRS
45 voltage in precordial leads, ST segment depression in any lead, lateral ST segment depression,
46 precordial ST segment depression, negative T wave, inferior negative T wave, lateral negative T wave,
47 QTc, right bundle branch block (RBBB), and left anterior fascicular block (LAFB) (**Supplementary**
48 **Digital Material 1: Supplementary Table IV**). The multivariate analysis confirmed that AF,
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tachycardia, PRWP, and RBBB were significantly associated with in-hospital mortality. (**figure 1, Table IV**)

Similarly, in the univariate analysis electrocardiographic predictors of in-hospital death and/or IMV were: AF, PRWP, lateral ST depression, precordial ST depression, negative T wave in any lead, lateral negative T wave, tachycardia, QTc, RBBB, and LAFB (**Supplementary Digital Material 1: Supplementary Table IV**). These variables were afterward included into the multiple logistic regression: AF, PRWP, tachycardia, QTc, and RBBB resulted independent predictors of in-hospital death and/or IMV. (**figure 1, table V**)

DISCUSSION

This was a retrospective analysis of electrocardiographic data from all consecutive patients with confirmed COVID-19 with low clinical risk profile at admission.

The main findings of the present study are:

1. in a low-risk COVID-19 population specific electrocardiographic parameters resulted associated with a poor prognosis during hospital stay, proving the usefulness and reliability of this fast, cheap, and easy to perform exam in the prognostic stratification process;
2. tachycardia, AF, PRWP, and RBBB resulted independent predictors of in-hospital death; the same parameters, along with QTc, were also strongly associated with the composite outcome of in-hospital death and/or need for IMV;
3. among all, RBBB was the strongest independent predictor of poor outcome during hospital stay for low-risk COVID-19 patients.

There is growing evidence that cardiac involvement significantly contributes to COVID-19 related mortality.⁸⁻¹¹ Four main pathophysiologic pathways to explain cardiac involvement in COVID-19 have been proposed: coronary artery thrombosis related to the hypercoagulable state¹⁶, direct viral myocardial infiltration¹⁷, demand-supply mismatch due to hypoxemia, and pro-inflammatory cytokines storm possibly leading to fulminant myocarditis¹⁸. In spite of the relevant prognostic impact

1 of the cardiac involvement, during the pandemic peaks when high pressure is exerted on the healthcare
2 systems, the shortage of human and instrumental resources cannot guarantee a complete cardiological
3 evaluation to all COVID-19 patients. In this context the detection of electrocardiographic parameters
4 able to predict early unfavourable outcome in a baseline low risk-population seems of paramount
5 importance; ECG represents indeed a fast and easy instrument which could be used to address the
6 rationed healthcare resources where mainly needed.
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12 Being rapid and low cost, ECG is routinely performed to all patients hospitalized for COVID-19
13 in our center. Some other studies have already suggested the association between specific
14 electrocardiographic patterns and patients prognosis.¹⁹ Nevertheless, these previous analyses have
15 always included all COVID-19 patients admitted to the Emergency Department and hospitalized
16 thereafter. This "all-inclusive" approach significantly jeopardizes the prognostic validity of
17 electrocardiographic findings since the inclusion of severely diseased patients raises the interpretation
18 dilemma of considering baseline parameters as "simple markers" of already advanced disease rather
19 than real predictors of subsequent unfavourable clinical course.
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30 The uniqueness of our study is represented indeed by the restricted focus on low-risk COVID-19
31 patients; proof of this is the overall in-hospital mortality in our population of 7.5% which is lower than
32 most of the recent reports from unselected cohorts.^{20, 21} The detection of early markers of poor outcomes
33 in patients initially deemed at low risk is highly worthwhile to let physicians identify the patients who
34 deserve a closer monitoring, a more accurate cardiological evaluation, and a longer hospital stay.
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41 Our analysis demonstrated that tachycardia, AF, PRWP, and RBBB are independent predictors
42 of in-hospital death in a low-risk COVID-19 population; when the composite outcome of in-
43 hospital death and IMV is considered, also QTc should be added to the above mentioned parameters.
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47 AF in COVID-19 can result from multiple pathogenic pathways: enhanced inflammatory
48 signalling eventually leading to inflammatory cytokines storm, direct viral endothelial damage,
49 electrolytes and acid-base unbalances, and increased sympathetic activation. In a not negligible quote
50 of cases, AF is triggered by the COVID-19 in the context of pre-existing cardiovascular
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1 comorbidities.²²The AF related detrimental hemodynamic effects and its thrombotic risk, further
2 enhanced by the COVID-19 hypercoagulable state, could explain the increased mortality in this
3 population,²⁰ as already suggested by previous studies.²³⁻²⁵

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6 Moreover, PRWP is a common ECG finding that is often interpreted as suggestive, but not
7 conclusively diagnostic, of anterior myocardial infarction.²⁶ Lately, PRWP has been also proposed as
8 an early sign of left ventricle dysfunction in diabetic patients²⁷ and a predictor of overall and
9 cardiovascular mortality in healthy women.²⁸ Despite the relation between PRWP and mortality has
10 been already reported in smaller cohorts^{29, 30}, our data confirmed this finding in a larger population and
11 suggested PRWP to be also associated with the occurrence of the composite outcome of death and
12 IMV.
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21 Also baseline RBBB proved to be associated with both death and the composite of death and
22 IMV. Pneumonia and respiratory failure are the principal clinical conditions leading to hospitalization
23 in COVID-19 patients. Being RBBB a marker of right ventricular (RV) dysfunction, it can express the
24 increased afterload and over-distension of the RV resulting from endothelial damage of the pulmonary
25 circulation as a consequence of both the COVID-19 related thromboembolic complications and hypoxic
26 vasoconstriction. RV dysfunction assessed by 2-dimensional speckle-tracking echocardiography was
27 indeed lastly suggested to be associated with a higher mortality in COVID-19 patients³¹. Moreover a
28 recent study showed that RBBB at the time of hospital admission was associated with worse
29 survival²³. This study nevertheless did not discriminate between high and low-risk COVID-19
30 presentation as proved, relative to our investigation, by higher overall short-term mortality, a more
31 elevated rate of coronary heart disease (14.4% vs 8.4%) and, above all, of home O₂ therapy (20.5% vs
32 1.1%). The two latter conditions can both raise question on the identification beyond doubt of the RV
33 dysfunction as an outcome predictor rather than a simple marker of disease. Of note, no other ECG
34 patterns suggestive of pulmonary embolism (S1Q3T3, S1Q3, isolated negative T3 wave) were
35 associated with poor outcomes (**Supplementary Digital Material 1: Supplementary Table**
36 **IV**), probably because all signs of more advanced RV dysfunction and severe clinical deterioration. As
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1 proof, they are more frequently found in pulmonary embolism when complicated by cardiogenic
2 shock.³²
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4 QTc prolongation was previously proposed as a predictor of mortality²¹. Nevertheless, also
5 regarding QTc calculation the timing is crucial in hospitalized patients. QT interval is indeed
6 influenced by multiple factors which include pre-existing comorbidities (e.g. cardiovascular diseases),
7 extra cardiac organs failure (e.g. advanced kidney disease) which frequently complicates the course of
8 COVID patients, and medications such as hydroxychloroquine. In this view, a rigorous approach
9 was adopted by Rubin and co-authors who demonstrated that Sars-CoV-2 infection led to significant
10 mean QTc prolongation from baseline, independently of other causes³³. Since in our study QTc was
11 obtained upon admission and in a low-risk population, the established relation with the occurrence of
12 death and/or IMV did not suffer from the mentioned potentially confounding factors.
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15 Despite the topic, of great scientific interest, has been already explored by others, our study
16 presents several peculiar strengths. First, as previously described, we restricted the analysis to a low-risk
17 population affected by COVID-19 which allowed to identify "real" predictors of unfavourable clinical
18 course. Moreover, low risk patients represent the category which can benefit the most from the
19 modulation of the diagnostic and therapeutic choices during hospital stay. Second, we rigorously
20 considered only baseline ECG for avoiding disease and drug related confounders. Third, we conducted
21 a deeply detailed and highly standardized ECG analysis including numerous ECG parameters.
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24 On the other hand, some limitations needs to be recognized as well. First, the study design is
25 observational and all therapeutic choices were up to treating physicians; as a consequence data would
26 result in selection bias and the study should be considered as hypotheses generating. Second, for the
27 great majority of our patients previous ECGs were missing and comparison unfeasible. Third, some
28 anamnestic and laboratory data were missing.
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51 CONCLUSIONS

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1 In low-risk COVID-19 patients AF, PWPR, tachycardia, and RBBB resulted independent
2 predictors of in-hospital all-cause mortality; AF, PWPR, Tachycardia, RBBB and QTc showed to be
3 independent risk factors for the occurrence of the composite endpoint of in-hospital death and IMV. Our
4 study demonstrated for the first time that RBBB and PRWP, assessed upon admission, are associated
5 with unfavourable clinical course in a low-risk population with Sars-CoV-2 infection. Being ECG a
6 fast, easy, and low-cost exam, this finding appears of great interest since would guide physicians to
7 identify the patients who could benefit the most from a stricter clinical monitoring, more accurate
8 cardiological evaluation, and more aggressive pharmacological therapies.
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Conflicts of interest:

Prof. Biondi-Zoccai has consulted for InnovHeart and Replycare.

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Table I. Baseline characteristics.

	Overall (n=384)	In hospital death and/or IMV treatment (n=44)	Alive without IMV (n= 340)	p	In hospital death (n=29)	Alive (n=355)	p
Male, <i>n</i> (%)	206 (53.6%)	24 (54.5%)	182 (53.5%)	0.899	13 (44.8%)	193 (54.4%)	0.322
Female, <i>n</i> (%)	178 (46.4%)	20 (45.5%)	158 (46.5%)	0.899	16 (55.2%)	162 (45.6%)	0.322
Weight, <i>kg</i>	76.6 ±20.3	68.4 ±12.8	77.7 ±20.9	0.009	63 ±11.2	77 ±20.5	0.001
Height, <i>cm</i>	165.3 ±15.9	164.6 ±7.4	165.4 ±16.7	0.783	162 ±8.1	165 ±16.3	0.406
Hypertension, <i>n</i> (%)	176 (45.8%)	26 (59.1%)	150 (44.1%)	0.061	20 (69.0%)	156 (43.9%)	0.008
Dyslipidaemia, <i>n</i> (%)	80 (20.8%)	17 (38.6%)	63 (18.5%)	0.002	14 (48.3%)	66 (18.6%)	<0.001
DM 1, <i>n</i> (%)	4 (1.0%)	0 (0.0%)	4 (1.2%)	0.613	0 (0.0%)	4 (1.1%)	0.730
DM 2, <i>n</i> (%)	56 (14.6%)	13 (29.5%)	43 (12.6%)	0.003	9 (31.0%)	47 (13.2%)	0.015
BMI > 30, <i>n</i> (%)	61 (15.9%)	7 (15.9%)	54 (15.9%)	0.996	1 (3.4%)	60 (16.9%)	0.038
Current smoker, <i>n</i> (%)	13 (3.4%)	2 (4.5%)	11 (3.2%)	0.451	2 (6.9%)	11 (3.1%)	0.256
Previous smoker, <i>n</i> (%)	36 (9.4%)	5 (11.4%)	31 (9.1%)	0.397	3 (10.3%)	33 (9.3%)	0.526
eGFR < 30 ml/min, <i>n</i> (%)	16 (4.2%)	4 (9.1%)	12 (3.5%)	0.098	4 (13.8%)	12 (3.4%)	0.025
Any lung disease, <i>n</i> (%)	49 (12.8%)	10 (22.7%)	39 (11.5%)	0.052	9 (31.0%)	40 (11.3%)	0.006
Allergies, <i>n</i> (%)	59 (15.4%)	2 (4.5%)	57 (16.8%)	0.034	1 (3.4%)	58 (16.3%)	0.044
Any Heart disease, <i>n</i> (%)	66 (17.2%)	18 (40.9%)	48 (14.1%)	<0.001	15 (51.7%)	51 (14.4%)	<0.001
Coronary heart diseases, <i>n</i> (%)	33 (8.6%)	7 (15.9%)	26 (7.6%)	0.068	4 (13.8%)	29 (8.2%)	0.230
Valvular heart diseases, <i>n</i> (%)	9 (2.3%)	2 (4.5%)	7 (2.1%)	0.276	2 (6.9%)	7 (2.0%)	0.142
Previous Stroke / TIA, <i>n</i> (%)	16 (4.2%)	6 (13.6%)	10 (2.9%)	0.005	4 (13.8%)	12 (3.4%)	0.025
Connectivitis, <i>n</i> (%)	7 (1.8%)	0 (0.0%)	7 (2.1%)	0.424	0 (0.0%)	7 (2.0%)	0.575
Liver diseases, <i>n</i> (%)	7 (1.8%)	1 (2.3%)	6 (1.8%)	0.576	1 (3.4%)	6 (1.7%)	0.425
Cirrhosis, <i>n</i> (%)	1 (0.3%)	0 (0.0%)	1 (0.3%)	0.885	0 (0.0%)	1 (0.3%)	0.924
Cancer, <i>n</i> (%)	39 (10.2%)	8 (18.2%)	31 (9.1%)	0.061	7 (24.1%)	32 (9.0%)	0.019
Immunodepression, <i>n</i> (%)	38 (9.9%)	6 (13.6%)	32 (9.4%)	0.258	6 (20.7%)	32 (9.0%)	0.054

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35 IMV, invasive mechanical ventilation; DM, diabetes mellitus; BMI, body mass index; eGFR, estimated glomerular filtration rate; TIA, transient

36 ischemic attack.

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Table II. Symptoms on admission.

Symptoms on admission	Overall (n=384)	In hospital death and/or IMV treatment (n=44)	Alive without IMV (n= 340)	p	In hospital death (n=29)	Alive (n=355)	p
Dyspnoea, <i>n</i> (%)	168 (43.8%)	33 (75.0%)	135 (39.7%)	<0.001	19 (65.5%)	149 (42.0%)	0.014
Tachypnea (>22 rpm), <i>n</i> (%)	87 (22.7%)	24 (54.5%)	63 (18.5%)	<0.001	12 (41.4%)	75 (21.1%)	0.012
Fatigue, <i>n</i> (%)	105 (27.3%)	14 (31.8%)	91 (26.8%)	0.479	9 (31.0%)	96 (27.0%)	0.643
Hypo/Anosmia, <i>n</i> (%)	29 (7.6%)	3 (6.8%)	26 (7.6%)	0.569	1 (3.4%)	28 (7.9%)	0.334
Disgeusia, <i>n</i> (%)	40 (10.4%)	2 (4.5%)	38 (11.2%)	0.134	0 (0.0%)	40 (11.3%)	0.036
Sore throat, <i>n</i> (%)	21 (5.5%)	2 (4.5%)	19 (5.6%)	0.559	2 (6.9%)	19 (5.4%)	0.483
Fever, <i>n</i> (%)	311 (81.0%)	36 (81.8%)	275 (80.9%)	0.882	22 (75.9%)	289 (81.4%)	0.463
Cough, <i>n</i> (%)	164 (42.7%)	21 (47.7%)	143 (42.1%)	0.474	13 (44.8%)	151 (42.5%)	0.810
Vomiting / Nausea, <i>n</i> (%)	17 (4.4%)	2 (4.5%)	15 (4.4%)	0.601	2 (6.9%)	15 (4.2%)	0.373
Diarrhoea, <i>n</i> (%)	46 (12.0%)	4 (9.1%)	42 (12.4%)	0.531	2 (6.9%)	44 (12.4%)	0.298
Myalgia / Arthralgia, <i>n</i> (%)	48 (12.5%)	2 (4.5%)	46 (13.5%)	0.063	0 (0.0%)	48 (13.5%)	0.018
Glasgow Coma Score, mean(±SD)	14.6 ±1.7	12.6 ±3.5	14.8 ±1.0	<0.001	11.7 ±3.5	14.8 ±1.2	<0.001
SBP <90 mmHg and/or DBP <60 mmHg, <i>n</i> (%)	22 (5.7%)	13 (29.5%)	9 (2.6%)	<0.001	10 (34.5%)	12 (3.4%)	<0.001

IMV, invasive mechanical ventilation; SBP, systolic blood pressure; DBP, diastolic blood pressure.

Table III. ECG characteristics upon admission.

ECG	Overall (n=384)	In hospital death and/or IMV treatment (n=44)	Alive without IMV (n= 340)	p	In hospital death (n=29)	Alive (n=355)	p
Sinus Rhythm, <i>n</i> (%)	361 (94.0%)	34 (77.3%)	327 (96.2%)	<0.001	19 (65.5%)	342 (96.3%)	<0.001
Sinus Arrhythmia, <i>n</i> (%)	2 (0.5%)	2 (4.5%)	0 (0.0%)	0.013	2 (6.9%)	0 (0.0%)	0.006
AF, <i>n</i> (%)	21 (5.5%)	8 (18.2%)	13 (3.8%)	0.001	8 (27.6%)	13 (3.7%)	<0.001
Tachycardia, <i>n</i> (%)	50 (13.0%)	17 (38.6%)	33 (9.7%)	<0.001	14 (48.3%)	36 (10.1%)	<0.001
Low QRS voltage (precordial), <i>n</i> (%)	50 (13.8%)	9 (20.5%)	41 (12.1%)	0.119	8 (27.6%)	42 (11.9%)	0.023
RBBB, <i>n</i> (%)	26 (6.8%)	7 (15.9%)	19 (5.6%)	0.02	5 (17.2%)	21 (5.9%)	0.037
LAFB, <i>n</i> (%)	39 (10.2%)	11 (25.0%)	28 (8.2%)	0.002	10 (34.5%)	29 (8.2%)	<0.001
RBBB and tachycardia, <i>n</i> (%)	5 (1.3%)	4 (9.1%)	1 (0.3%)	0.001	3 (10.3%)	2 (0.6%)	0.004
Absent / poor R wave progression (V1-V6), <i>n</i> (%)	32 (8.3%)	10 (22.7%)	22 (6.5%)	0.001	7 (24.1%)	25 (7.1%)	0.006
QTc (msec)	410.2 ±32.7	425.0 ±34.6	407.0 ±31.6	0.001	433.0 ±35.8	408.0 ±31.8	<0.001
ST depression, <i>n</i> (%)	9 (2.3%)	3 (6.8%)	6 (1.8%)	0.072	3 (10.3%)	6 (1.7%)	0.024
Lateral ST depression (V5-V6-DI-aVL), <i>n</i> (%)	3 (0.8%)	2 (4.5%)	1 (2.7%)	0.036	2 (6.9%)	1 (0.3%)	0.016
T wave inversion, <i>n</i> (%)	36 (9.4%)	9 (20.5%)	27 (7.9%)	0.013	7 (24.1%)	29 (8.2%)	0.012
Lateral T wave inversion (V5-V6-DI-aVL), <i>n</i> (%)	19 (4.9%)	7 (15.9%)	12 (3.5%)	0.003	7 (24.1%)	12 (3.4%)	<0.001
Inferior T wave inversion (DII-DIII-aVF), <i>n</i> (%)	9 (2.3%)	3 (6.8%)	6 (1.8%)	0.072	3 (10.3%)	6 (1.7%)	0.024

AF, atrial fibrillation; RBBB, right bundle branch block; LAFB, left anterior fascicular block; QTc, corrected QT interval; AV, atrio-ventricular.

Table IV. Univariate and multivariate logistic regression: ECG predictors of all-cause mortality

ECG findings	Univariate logistic regression analysis			Multivariate logistic regression analysis		
	OR	CI 95%	p	OR	CI 95%	p
AF	10.022	3.743-26.832	<0.001	7.401	1.623-33.751	0.010
Tachycardia	8.270	3.695-18.512	<0.001	4.263	1.430-12.714	0.009
absent/poor R progression V1-V6	4.200	1.636-10.781	0.003	4.381	1.087-17.665	0.038
Low QRS voltage (precordial)	2.839	1.183-6.815	0.020	2.996		0.064
ST depression	6.712	1.587-28.385	0.010	1.790		0.760
ST depression lateral	26.222	2.304-294.494	0.008	1.954		0.808
ST depression precordial	13.074	1.772-96.475	0.012	1.525		0.824
T wave inversion	3.577	1.409-9.079	0.007	0.436		0.499
T wave inversion inferior	6.712	1.587-28.385	0.010	4.851		0.212
T wave inversion lateral	9.095	3.257-25.398	<0.001	3.296		0.360
QTc	1.020	1.009-1.031	0.001	1.008		0.241
RBBB	3.313	1.148-9.560	0.027	8.039	1.229-52.603	0.030
LAFB	5.917	2.517-13.908	<0.001	2.337		0.217

AF, atrial fibrillation; QTc, corrected QT interval; RBBB, right bundle branch block; LAFB, left anterior fascicular block; OR, odds ratio; CI, confidence interval.

Table V. Univariate and multivariate logistic regression: ECG predictors of in-hospital death and/or invasive mechanical ventilation.

ECG findings	Univariate logistic regression analysis			Multivariate logistic regression analysis		
	OR	IC 95%	p	OR	IC 95%	p
AF	5.59	2.171-14.390	<0.001	3.878	1.029-14.611	0.045
absent/poor R progression V1-V6	4.251	1.859-9.720	0.001	5.313	1.847-15.282	0.002
ST depression laterale	16.143	1.433-181.862	0.024	3.060		0.481
ST depression precordial	8.048	1.104-58.639	0.040	3.266		0.345
T wave inversion	2.981	1.298-6.845	0.010	1.812		0.416
T wave inversion lateral	5.171	1.917-13.947	0.001	0.981		0.984
Tachycardia	5.857	2.894-11.856	<0.001	3.294	1.387-7.822	0.007
QTc	1.018	1.008-1.028	<0.001	1.011	1-1.023	0.048
RBBB	3.196	1.260-8.109	0.014	9.196	1.600-52.852	0.013
LAFB	3.714	1.695-8.138	0.001	0.883		0.844

AF, atrial fibrillation; QTc, corrected QT interval; RBBB, right bundle branch block; LAFB, left anterior fascicular block; OR, odds ratio; CI, confidence interval.

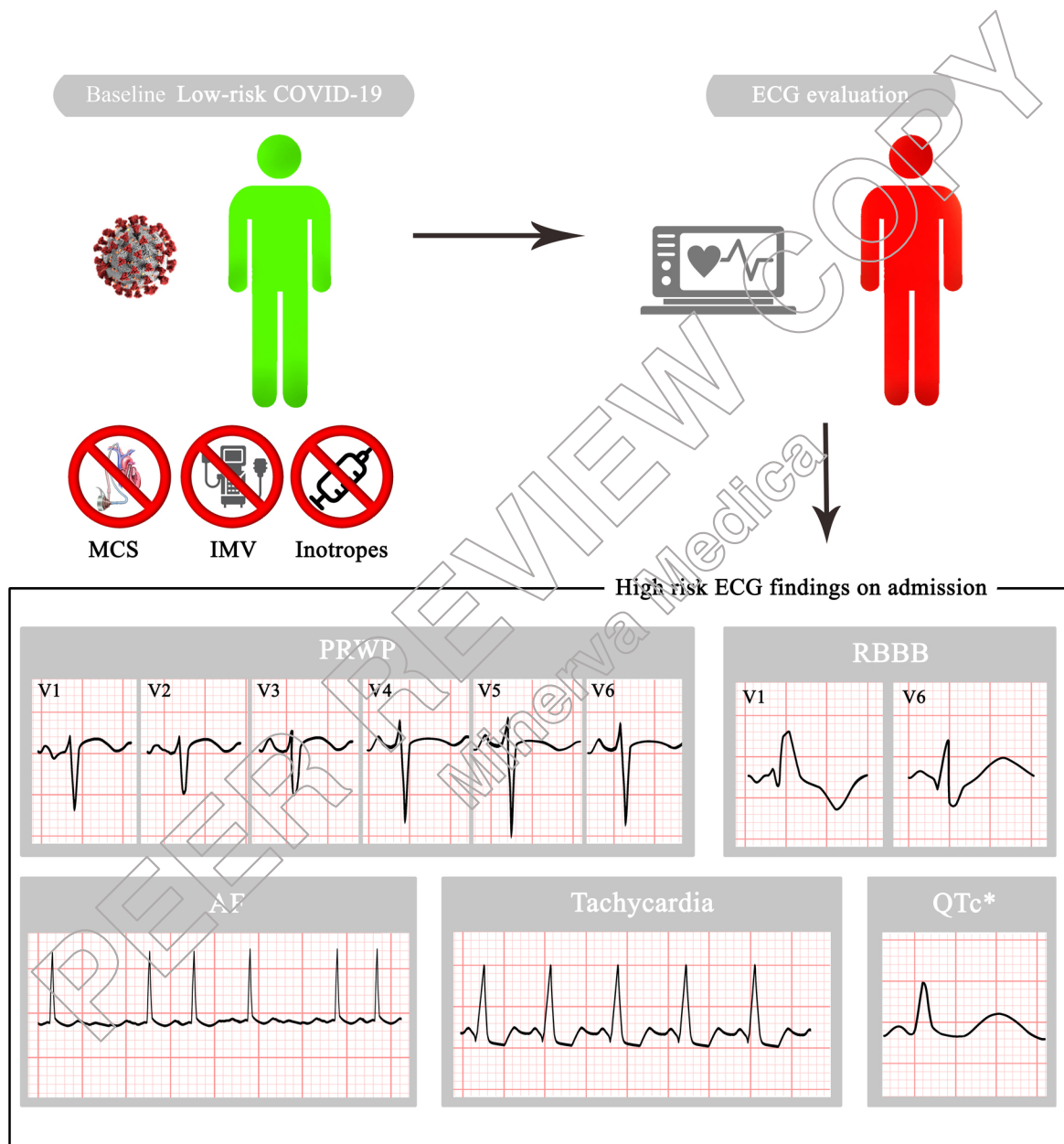
Figure 1.

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3 Electrocardiographic predictors of in-hospital death and/or IMV in low-risk patients hospitalized for COVID-19.

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9 In low-risk COVID-19 patients, AF, PRWP, tachycardia, and RBBB are independent predictors of in-hospital all-cause mortality; AF, PRWP,
10 Tachycardia, RBBB, and corrected QT interval (*) are independent risk factors for the occurrence of the composite endpoint of death and IMV. MCS,
11 mechanical circulation support; IMV, invasive mechanical ventilation; PRWP, poor R wave progression; RBBB, right bundle branch block; AF, atrial
12 fibrillation; QTc, QT corrected interval.
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