

---

**"STUDY OF COAGULATION PROFILE DYNAMICS IN  
PATIENTS WITH COVID 19 INFECTION - A ONE YEAR  
CROSS SECTIONAL STUDY IN KLE'S DR.PRABHAKAR  
KORE HOSPITAL AND MEDICAL RESEARCH CENTRE,  
BELAGAVI."**

---

**BY**

**REG NO: BG0120021**

**Disse  
Dissertation**

*Submitted to*

*KAHER, Belagavi, Karnataka,*

*In partial fulfilment of the requirements for the degree of*

**M.D.**

**IN**

**GENERAL MEDICINE**

**JAWAHARLAL NEHRU MEDICAL COLLEGE,  
KAHER, BELAGAVI – 590010 KARNATAKA.**

---

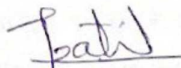
**JUNE/JULY – 2023**

---

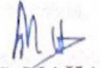
KLE ACADEMY OF HIGHER EDUCATION AND RESEARCH,  
BELAGAVI, KARNATAKA.

**ENDORSEMENT BY THE HOD/ PRINCIPAL/  
HEAD OF THE INSTITUTION**

This is to certify that the dissertation "STUDY OF COAGULATION PROFILE DYNAMICS IN PATIENTS WITH COVID 19 INFECTION - A ONE YEAR CROSS SECTIONAL STUDY IN KLE'S DR.PRABHAKAR KORE HOSPITAL AND MEDICAL RESEARCH CENTRE, BELAGAVI." is a bonafide research work done by REG NO. BG0120021.

  
**Dr. REKHA S. PATIL MD,**  
Professor & Head  
Department of General Medicine  
J.N. Medical College  
Nehru Nagar, Belagavi- 590010

Date:  
Place: Belagavi

  
**Dr. N. S. MAHANTASHETTI MD,**  
Principal  
J.N. Medical College  
Nehru Nagar, Belagavi- 590010

Date:  
Place: Belagavi

## UNDERTAKING

I, Reg No. BG0120021, hereby declare that the information and data mentioned in my dissertation entitled “STUDY OF COAGULATION PROFILE DYNAMICS IN PATIENTS WITH COVID 19 INFECTION - A ONE YEAR CROSS SECTIONAL STUDY IN KLE’S DR.PRABHAKAR KORE HOSPITAL AND MEDICAL RESEARCH CENTRE, BELAGAVI.” belongs to me and is original. I am aware of the definition of plagiarism as detailed below:

- AN ACT OR INSTANCE OF USING OR CLOSELY IMITATING THE LANGUAGE AND THOUGHTS OF ANOTHER AUTHOR, WITHOUT AUTHORIZATION AND REPRESENTATION OF THAT AUTHOR'S WORK AS ONE'S OWN, AS BY NOT CREDITING THE ORIGINAL AUTHOR.
- A PIECE OF WRITING OR OTHER WORK REFLECTING SEARCH UNAUTHORIZED USE OR IMITATION.
- THE DELIBERATE OR RECKLESS REPRESENTATION OF ANOTHER'S WORDS, THOUGHTS OR IDEAS AS ONE'S OWN WITHOUT ATTRIBUTION IN CONNECTION WITH SUBMISSION OF ACADEMY, WORK, WHETHER GRADED OR OTHERWISE.

I HEREBY DECLARE THAT THE DISSERTATION PREPARED BY ME IS ORIGINAL-ONE AND DOES NOT INVOLVE PLAGIARISM ANYWHERE. IN CASE AT A LATER STAGE, IT IS FOUND THAT I HAVE INDULGED IN PLAGIARISM, THEN I AM SOLELY RESPONSIBLE FOR THE SAME AND THE INSTITUTION IS AT LIBERTY TO TAKE ANY DISCIPLINARY ACTION AGAINST ME INCLUDING CANCELLATION OF MY DISSERTATION, OR ANY OTHER PENALTIES AS IMPOSED BY THE UNIVERSITY.



Date: 19/01/2023

Place: Belagavi

*Bathe*

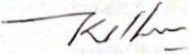
REG NO: BG0120021


# ANTI-PLAGIARISM CHECK – ACCEPTANCE LETTER


	<b>JAWAHARLAL NEHRU MEDICAL COLLEGE</b> <small>[Recognized by Medical Council of India, New Delhi]</small>	
Accredited 'A+' Grade by NAAC (3 <sup>rd</sup> Cycle)		Placed in Category 'A' by MHRD (Govt)
Nehru Nagar, Belagavi- 590 010, Karnataka, INDIA		
☎ 0831 - 2471350	☎ 0811 - 2470759	✉ principal@jnmc.edu
Ref No: MDC/PG/		Date: 15-12-2022

**ACCEPTANCE LETTER**

The softcopy of thesis entitled: "STUDY OF COAGULATION PROFILE DYNAMICS IN PATIENTS WITH COVID 19 INFECTION - A ONE YEAR CROSS SECTIONAL STUDY IN KLE'S DR. PRABHAKAR KORE HOSPITAL AND MEDICAL RESEARCH CENTRE, BELAGAVI" has been submitted for Anti-Plagiarism check through Turnitin software. The scan has been carried out and the scanned output reveals a match percentage of 10% which is within the acceptable limits of 10% as per the guidelines given by UGC.

  
Guide.



  
Dr. (Mrs.) N.S. Mahantashetti,  
Chairperson-Antiplagiarism Committee &  
Principal,  
J. N. Medical College, Belagavi.

To,  
Reg. No. BG0120021,  
Postgraduate Student,  
2020-21 Batch,  
Department of General Medicine,  
J. N. Medical College, Belagavi.

# ETHICAL CLEARANCE CERTIFICATE



K.I.E. ACADEMY OF HIGHER EDUCATION AND RESEARCH  
(Deemed to be University)

Accredited 'A' Grade by NAAC (2<sup>nd</sup> Cycle)

Placed in Category 'A' by MHRD (Govt)

**JAWAHARLAL NEHRU MEDICAL COLLEGE,**  
**NEHRU NAGAR, BELAGAVI-590010 (KARNATAKA-INDIA)**

Website: <http://www.jnmc.edu>  
E-Mail : [dome@jnmc.edu](mailto:dome@jnmc.edu)

Phone: (+ 91-0)831 Office : 2472550  
Principal: 2471701  
Fax No. +91 (0)831 - 2470759

Ref: MDC/DOME/ 07

Date: 25/01/2021

**REG NO: BG0120021**

PG student in Medicine,  
J.N.Medical College,  
BELAGAVI.

Sub: Institutional Ethical Clearance for the study.

With reference to the above, we wish to inform you that your proposed research project titled  
**"STUDY OF COAGULATION PROFILE DYNAMICS IN PATIENTS WITH COVID 19  
INFECTION – A ONE YEAR CROSS SECTIONAL STUDY IN KLE'S DR. PRABHAKAR  
KORE HOSPITAL AND MEDICAL RESEARCH CENTRE, BELAGAVI"**, is ethical and  
justifiable. The proposed research project has been cleared by the JNMC Institutional Ethics  
Committee on Human Subjects Research.

(Dr. Smita Sonoli)  
Member Secretary  
JNMC Institutional Ethics Committee  
on Human Subjects Research,  
J.N.Medical College, Belagavi.

(Dr. Harsha Hegde)  
Chairman,  
JNMC Institutional Ethics Committee  
on Human Subjects Research,  
J.N.Medical College, Belagavi.

## ABSTRACT

**Background:** Thrombotic events such deep vein thrombosis, pulmonary embolism, myocardial infarction, and ischemic strokes have become more common in critically ill patients.

**Material & Method:** The patients that were received hospital care with COVID-19 infection were the subjects of the current cross-sectional study. Following enrollment in the study and completion of a questionnaire and comprehensive clinical examination, all patients meeting the inclusion criteria will be evaluated. The patient's outcome was evaluated based on the length of their hospital stay, how much oxygen they needed, how they were feeling when they were discharged, and whether or not they died. The coagulation profile included the platelet count, INR, aPTT, serum D-dimer, fibrinogen, other inflammatory markers like LDH, hs-CRP, Ferritin and IL-6. The data were analysed and tabulated with the SPSS 21v, with  $p < 0.05$  was considered statistically significant

**Results:** After receiving informed consent, The study covered 150 patients in total. The mean age of the patients was found to be 55.45yrs, among them 80.7% were male patients and 19.3% were female with male preponderance in the study. On assessment of the parameters between the outcome of patients, the CT severity score was significantly higher in patient with worst outcome as death compared to the patients discharged. Similarly, there was significant higher mean level of the ferritin in patients with worst outcome as death. The mean level of Fibrinogen, Homocysteine, D-dimer was significantly higher among the patients with worst outcome as death compared to patients discharged. There was significant higher

incidence of death among the patients with ISTH score  $>5$  (75%) compared to patients discharged.

**Conclusion:** The present study documented the significant changes in the coagulation profile among the patients with worst outcome as death of the patients, also there is higher incidence of the inflammatory changes compared to the patients discharged from hospital. The ISTH score was found to be with good sensitivity and specificity to detect the worst outcome among the patients.

**KEY WORDS:** COVID 19, Coagulopathy, D-dimer, Fibrinogen, INR-APTT, Mortality

## **ABBREVIATIONS**

ACE-2	ANGIOTENSIN-CONVERTING ENZYME 2 RECEPTOR
APTT	ACTIVATED PARTIAL THROMBOPLASTIN CLOTTING TIME
AUC	AREA UNDER CURVE
CFR	CASE FATALITY RATE
CRP	C-REACTIVE PROTEIN
CT	COMPUTED TOMOGRAPHY
HIF	HYPOXIA-INDUCIBLE FACTOR
ICU	INTENSIVE CARE UNIT
IRE	IRON-RESPONSIVE ELEMENTS
MERS	MIDDLE EAST RESPIRATORY SYNDROME
MPV	MEAN PLATELET VOLUME
PDW	PLATELET DISTRIBUTION WIDTH
PLCC	PLATELET LARGE CELL COUNT
PLCR	PLATELET LARGE CELL COUNT & RATIO
PLT	PLATELET COUNT
TMPRS	TRANSMEMBRANE PROTEASE SERINE
UTR	UNTRANSLATED REGION
WHO	WORLD HEALTH ORGANISATION
ISTH	INTERNATIONAL SOCIETY ON THROMBOSIS & HEMOSTASIS

## TABLE OF CONTENTS

<b>SL.NO</b>	<b>CONTENTS</b>	<b>PAGE NO.</b>
<b>1</b>	<b>INTRODUCTION</b>	<b>1-2</b>
<b>2</b>	<b>OBJECTIVE OF STUDY</b>	<b>3</b>
<b>3</b>	<b>REVIEW OF LITERATURE</b>	<b>4-23</b>
<b>4</b>	<b>MATERIALS AND METHODS</b>	<b>24-29</b>
<b>5</b>	<b>RESULTS</b>	<b>30-49</b>
<b>6</b>	<b>DISCUSSION</b>	<b>50-54</b>
<b>7</b>	<b>CONCLUSION</b>	<b>55</b>
<b>8</b>	<b>SUMMARY</b>	<b>56-57</b>
<b>9</b>	<b>BIBLIOGRAPHY</b>	<b>58-66</b>
<b>10</b>	<b>ANNEXURES</b>	
	<b>Annexure I: Consent form</b>	<b>67-71</b>
	<b>Annexure II: Proforma</b>	<b>72-75</b>
	<b>Annexure III: Master Chart</b>	<b>76-77</b>

## **LIST OF FIGURES**

<b>SL NO.</b>	<b>FIGURE</b>	<b>PAGE NO.</b>
1	Timeline Of COVID-19	4
2	COVID-19 virus structure	6
3	Life cycle Of the Corona virus	8
4	Pathogenesis Of COVID-19 infection	11
5	Relation Of RAAS and SARS-CoV-2	12
6	Transmission and entry Of COVID-19	15
7	Pathophysiology Of COVID-19	16
8	Laboratory findings associated with severity Of COVID infection	20
9	Mean age of the patients included in the study	30
10	Gender distribution of study participants	32
11	Vaccination status of the patients included in the study	33
12	Outcome of the patients in the study	34
13	Comparison of CT severity score with outcome among study participants	35
14	Comparison of the mean level of blood parameters with outcome of patients	37
15	Comparison of the mean level of blood parameters with outcome of patients	38

16	Comparison of the biomarkers with outcome of the study participants	40
17	Comparison of the biomarkers with outcome of the study participants	41
18	Comparison of the renal and liver function test parameters with outcome of the patients	43
19	Comparison of the ISTH score with outcome of the patients	44
20	Diagnostic characteristics of ISTH score to predict the outcome	45

## LIST OF TABLES

<b>SL NO.</b>	<b>TABLE</b>	<b>PAGE NO.</b>
1	Mean age of the participants included in the study	30
2	Distribution of age (years) of the participants included in the study	31
3	Gender distribution of study participants	32
4	Vaccination status of the patients included in the study	33
5	Outcome of the patients in the study	34
6	Comparison of CT severity score with outcome among study participants	35
7	Comparison of the mean level of blood parameters with outcome of patients	36
8	Comparison of the biomarkers with outcome of the study participants	37
9	Comparison of the renal and liver function test parameters with outcome of the patients	42
10	Comparison of the ISTH score with outcome of the patients	44
11	Diagnostic characteristics of ISTH score to predict the outcome	45

12	Co-relation of CT severity score with ISTH score	46
13	Co-relation of day 3 Platelet Count and Mortality	46
14	Co-relation of day 3 D-dimer level and mortality	47
15	Co-relation of vaccination status to ISTH score	48
16	Co-relation of vaccination status to mortality	48
17	Co-relation of T2DM and mortality	49
18	Co-Relation of T2DM with ISTH score	49

## **INTRODUCTION**

In December 2019, the SARS-CoV-2 novel coronavirus disease spread around the globe. It first appeared in China before spreading fast elsewhere. The illness has been designated as COVID-19 by the International Committee on Virus Taxonomy (ICTV).

COVID-19's strong infectivity led in a quick surge in new cases, and the illness has spread internationally to 188 nations and six continents, making it a current global health burden. COVID 19 initially believed to be pneumonia has now believed to have a much more diverse spectrum of disease. "COVID is a coagulopathy" is now the new understanding as multiple studies worldwide have confirmed the vasculopathy associated with this infection.

The frequency of thrombotic events include deep vein thrombosis, pulmonary embolism, myocardial infarction, and ischemic strokes has increased in critically ill patients. Although clinical observations of thromboembolic events have been made, the precise mechanism is yet unknown.

Multiple studies done in China, United States, England and Spain have suggested a positive correlation between rise in inflammatory markers and deranged D-dimer Fibrinogen PT INR and APTT, Other- (coagulation profile markers) in patients and determined their relation to the predictable prognosis of the patients. The studies also suggest the use of effective Anti Coagulation Therapy in patients to minimise the risk of complications.

The novel SARS COV2 virus has a very significant effect on the vasculature of the human body reflecting as dynamic changes in the coagulation profile and possibly indicating the prognosis of the patient.

Because India has the world's second most affected population, a comprehensive analysis of the Coagulation profile dynamics in our population is required to understand clinical management of our patient population and provide adequate medication and prophylaxis. Thromboprophylaxis is a very dynamic intervention which needs an individualistic approach with a broad understanding of the vasculopathy, clinical presentation, associated co-morbidities and multiple factors.

COVID-19 thrombotic syndrome is an evolving spectra which needs in depth study and analysis in order to achieve optimum therapeutic goals, improve prognosis, minimise post infection complications and devise a good predictive scoring system to predict this elusive and complex coagulopathy.

## **OBJECTIVES**

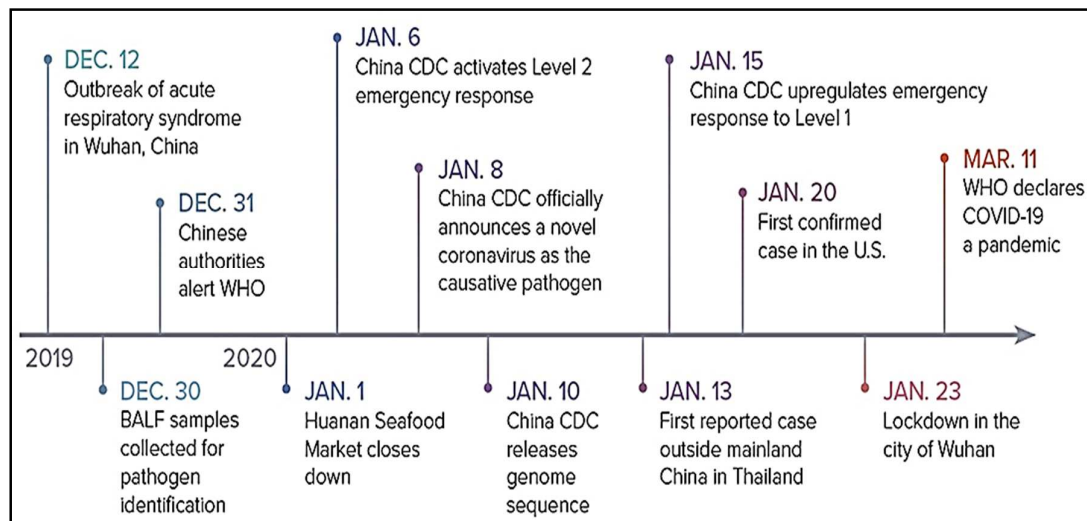
- To study the coagulation profile in patients with COVID 19 infection.
- Analyse the risk of thrombotic and thrombo-embolic events.
- Draw a comparison to the clinical outcome &
- Devise an effective thromboprophylaxis for patients to prevent further complications of the infection.

## REVIEW OF LITERATURE

Novel corona viruses are becoming important diseases for both humans and animals.<sup>1</sup> End of 2019 saw the discovery of a novel corona-virus as the cause of an outbreak of pneumonia cases in Wuhan, Hubei Province, China. As a result of its rapid growth, China saw an epidemic and a rise in cases worldwide.<sup>2</sup> The World Health Organization (WHO) designated the disorder as COV-19, which represents for coronavirus disease 2019. The SARS-CoV-2 virus is what causes COV-19 (2019-nCoV). On March 11, 2020, the WHO proclaimed COV-19 to be a pandemic.<sup>3</sup>

### Epidemiology

On December 31, 2019, Wuhan, Hubei Province, China reported the first case of SARS CoV-2 infection.



**Figure 1: Timeline of COVID-19<sup>4</sup>**

From this, it swiftly expanded throughout China and internationally, infecting over 185 countries as of April 23, 2020, culminating in the current global pandemic. COVID-19 was declared a Public Health Emergency of International Concern by the

World Health Organization on January 30, 2020, and on March 11, 2020, a pandemic status was given to it.

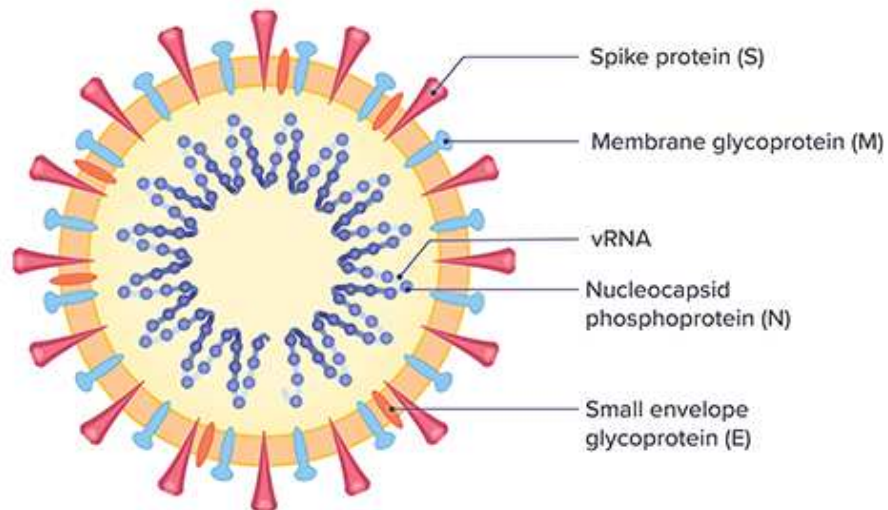
Its extremely contagious nature is shown by the fundamental replication number ( $R_0$ ) of SARS CoV-2, which is predicted to be between 1.4 and 3.9. In public gathering venues like cruise ships, religious, political, intellectual, and corporate congregations, as well as hospitals that do not adhere to personal privacy protections, the  $R_0$  may be deemed to be significantly greater. Expected are the incubation duration and serial interval to be 5–6 days and 8 days, respectively, in comparison to SARS CV and MERS CV.<sup>5</sup> The case-fatality rate (CFR) for the pandemic was predicted to range between 0.9 and 3 percent early on., which was lower than that of previous HCVs (SARS CV (6 percent -17 percent) and MERS CV). However, by the 24th of May 2020, the CFR had climbed exponentially in many countries.

In contrast to SARS CoV, a significant portion of SARS CoV-2 carriers have asymptotic or pauci-symptomatic signs, making them easy to spot and prospective carriers.<sup>6</sup> It is crucial to remember that not all clinical contacts get infected, suggesting that genetic susceptibility varies among individuals.<sup>7,8</sup> Normally, the virus enters people through the upper respiratory tract. SARS CV-2 was recently isolated from patient faeces, indicating the possibility of fecal-oral transmission.<sup>9,10</sup>

Women who are pregnant and infected with SARS CV-2 are more likely to transmit the disease vertically. Vertical transmission was ruled out due to negative viral testing on swabs obtained from the six infected pregnant women's amniotic fluid, cord blood, neonate pharynx, and breast milk.<sup>11</sup> Long-distance airborne transmission is also postulated, which is reliant on the virus's flow dynamics as well as the ventilation state Of the location.<sup>12</sup> Furthermore, mapping techniques like as cartograms

can be used to depict the dissemination and expansion of COVID-19.<sup>13</sup> Understanding the SARS CoV-2 transmission routes will enable the implementation of suitable containment measures.

## SARS-CoV-2 Structure



**Figure 2: COVID-19 virus structure<sup>4</sup>**

**SPIKE PROTEIN:** The cascade of inflammation is started by the binding of spike protein with the ACE 2 receptor of alveolar cells. Receptor TMP2R2 cleaves the spike protein releasing the S2 component which invokes the immune response and enters the cell and replicates.

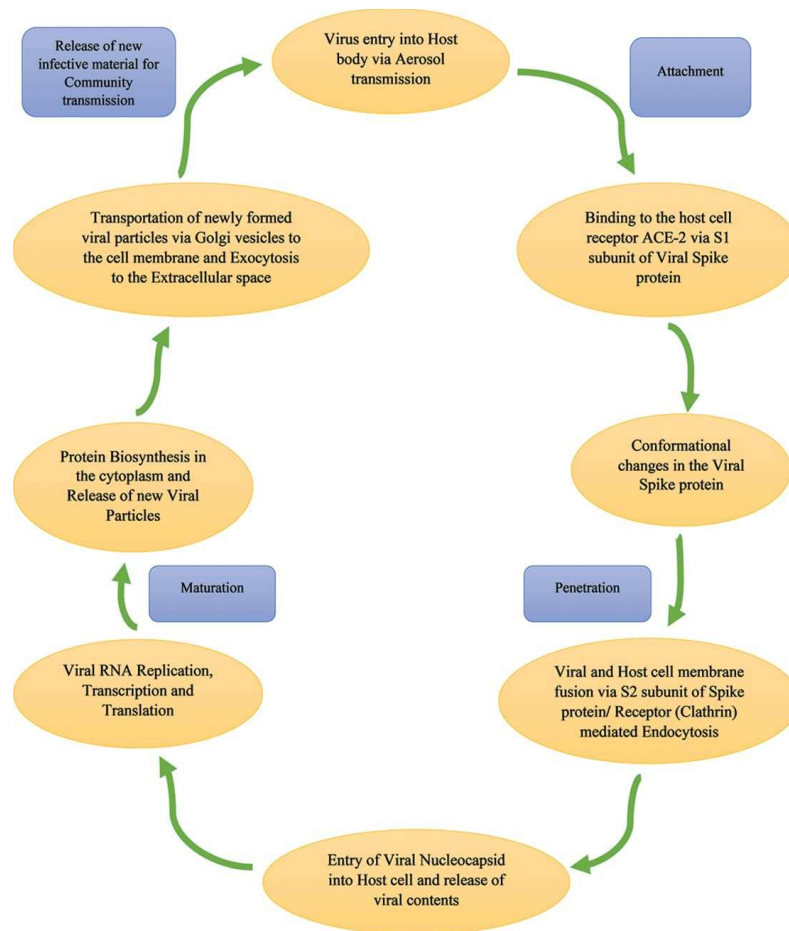
As the virus replicates there is more fusion of new virion and the inflammatory and immune response evoked by the S2 SPIKE PROTEIN magnifies and induces the cytokine storm. This exaggerated immune response marks the beginning of acute inflammatory state and provokes a disseminated intravascular coagulopathy. Antibodies of the COVID vaccine were designed to S2 spike protein

there by attenuating the initial fusion of virion and replication and in turn suppressing the inflammatory cascade.

Children had a lower incidence of serious illness and death from SARS CV-2 infection, despite the fact that all age groups are susceptible to infection.<sup>14</sup> Women often experience less adulthood-related effects than men because of higher plasma ACE2 levels.<sup>15</sup> The older population, especially those with co-morbidities such as diabetes, hypertension, stroke, and chronic lung, heart, or renal disease, are more prone to serious infections because of weak body defence against infection and underlying age-related organ system breakdown.<sup>16,17</sup> According to a recent study, group A of non-blood viruses, like SARS CoV, had greater infection and mortality rates from COVID-19 because there were insufficient protective anti-A IgM antibodies. Many aspects of the SARS CoV-2 epidemiology are still unknown, such as how the virus and human host interact, how susceptible humans are, and how the epidemic spreads.<sup>18,19</sup>

The two main categories of Corona viruses (CoVs) are  $\alpha$  and  $\beta$  (found in both humans and other animals);  $\alpha$  and  $\beta$  (seen in avian species). Due to the presence of spike protein in their envelop, they resemble to a crown in electron microscopy, hence their Latin name "coronam." With respiratory, gastrointestinal, hepatic, and central nervous system infections, they are able to infect avian and mammalian species, including humans. According to estimates, 2% to 4% of the population has Corona viruses in a healthy state, and 5% to 10% of acute respiratory infections are brought on by these viruses.

It is hypothesised that these viruses undergo rapid mutation and recombination as they move from natural host tissues to humans via an intermediary amplifying host, leading to the creation of more virulent CoVs that cause human sickness. Novel CoVs have resulted in pandemics with pulmonary and extra-pulmonary symptoms, including severe acute respiratory syndrome (SARS - SARS CoV) in China in 2002–2003 and Middle East respiratory syndrome (MERS - MERS CoV) in Saudi Arabia in 2012. The most common way that CoVs are discovered in children is as a co-infection with other respiratory viral infections, but they can also be the major pathogen causing infection in kids with underlying chronic conditions. Because of this, CoVs have become crucial pathogens in rising outbreaks of respiratory disease.



**Figure 3: Life cycle Of the Corona virus<sup>20</sup>**

### **Pathogenesis and Coagulopathy of COVID -19 infection**

**COVID 19 infection** is a multi systemic illness multiple pathological processes involving host immune responses, inflammation and disseminated coagulation. Important factors are over activation of immune cells, cytokine storm and excessive oxidative stress which are common pathophysiological mechanisms of ARDS, sepsis, DIC and multi organ failure. Hypercoagulable state of Covid-19 infection is multifactorial and causes micro and macro thrombi in arteries and veins predisposing to increased risk of deep vein thrombosis ,pulmonary embolism, strokes ,peripheral vascular disease, myocardial infarctions and disseminated intravascular coagulopathy.

### **4 CHEIF MECHANISMS OF VASCULOPATHY AND COAGULOPATHY IN COVID-19**

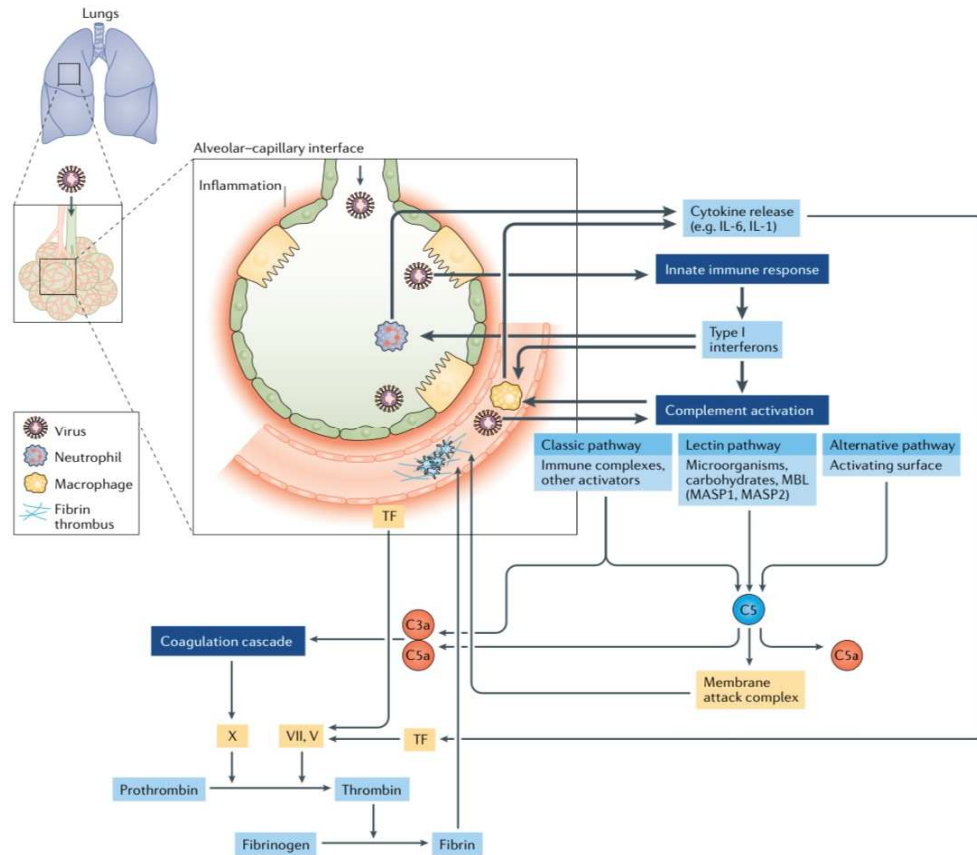
Cytokine Storm-acute inflammatory response and complement pathway activation

ACE 2 deactivation by the virion particles

Severe endothelial dysfunction, persistent hypoxia and vascular stasis

Postulated mechanisms like Lupus anticoagulant activation, Fibrinogen activation, on Willebrand factor activation and plasminogen activator inhibitor.

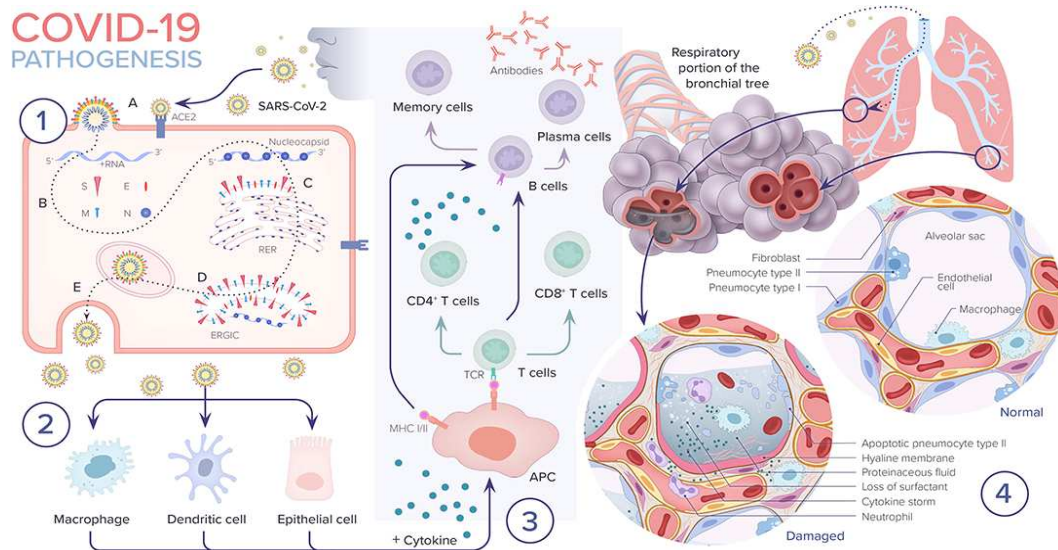
**VIRUS ENTRY PORT AND INITIATION OF CYTOKINE STORM:**



The highly glycosylated spike (S) protein S1 fragment with receptor binding domain (RBD), which binds to the angiotensin-converting enzyme 2 receptor (ACE-2 R) with 10–20 fold greater affinity than SARS CoV, is thought to be the mechanism by which SARS CoV-2 would infect human cells.<sup>21</sup> The majority of ACE-2 R is found in type II rather than type I human alveolar epithelial cells (type II > type I).<sup>22–24</sup> Endothelial cells, gastro-intestinal (esophageal and intestinal) epithelium, and cardiac myocytes all express the ACE-2 R.<sup>25</sup> Transmembrane proteases serine (TMPRSSs) expressed in host cells recognise and cleave the SARS CoV-2's unique polybasic S1/S2 protease cleavage site on the spike protein in order to release the fusinoid protein (S2 fraction), causing the membranes of the virus and host cell to fuse. It has been established that ACE-2 R and TMPRSSs are strongly exhibited in

upper esophageal epithelium, absorptive enterocytes, and alveolar type 2 pneumocytes, demonstrating that SARS CoV-2 can enter the human body through these epithelial types as well as alveolar epithelium. 5

As a result, TMPRSSs and ACE-2 R should be present in potential SARS CoV-2 target tissues. This membrane fusion enables viral RNA to be taken up and translated to produce new viral proteins in the cytoplasm of the host cell. Viral assembly, which involves nucleocapsid (N) proteins adhering to RNA molecules and being surrounded by envelope and membrane proteins to form whole viruses capable of infecting other cells, is the penultimate stage before viruses are discharged from infected cells.



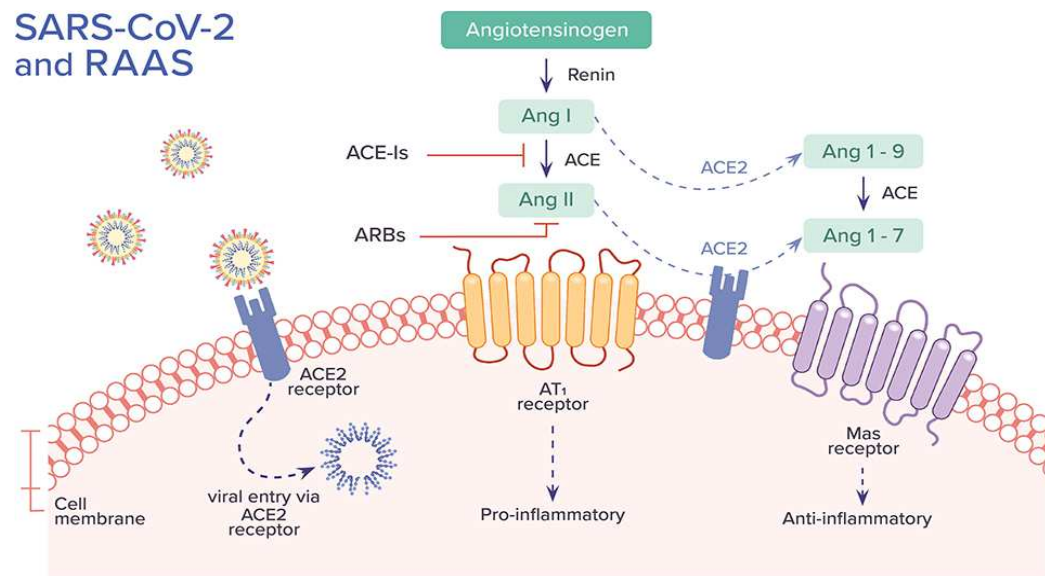
**Figure 4: Pathogenesis Of COVID-19 infection<sup>4</sup>**

Type II pneumocytes, which are responsible for tissue healing and surfactant production, are killed as a result of SARS CoV-2 infection, leading to an increase in surface tension and dyspnea. Due to an excess of cytokine production and release (cytokine storm) by activated inflammatory cells as a result of an accumulation of

angiotensin II, these injured type II pneumocytes impair the alveolar immune system's ability to operate.

Renin angiotensin system - COVID-19 & pro-coagulant state

The renin-angiotensin-aldosterone system (RAAS) is influenced by SARS-CoV-2 and SARS-CoV-2 via the enzyme ACE2, which also serves as a receptor for both viruses and normally lowers RAAS activity. In the RAAS, ACE converts angiotensin I to angiotensin II. The Angiotensin II type 1 receptor is how Angiotensin II exerts its vasoconstrictory and pro-inflammatory actions (AT<sub>1</sub>R). A multitude of functions, including vasodilation and anti-inflammatory properties, are stimulated when angiotensin II is converted into angiotensin I-7 by ACE2. Additionally, ACE2 converts angiotensin I into angiotensin I-9, which is then converted by ACE into angiotensin I-7. By removing Angiotensin II and stimulating the synthesis of Angiotensin I-7, ACE2 reduces the harmful vasoconstriction and pro-inflammatory effects of Angiotensin II.<sup>4</sup>



**Figure 5: Relation Of RAAS and SARS-CoV-2<sup>4</sup>**

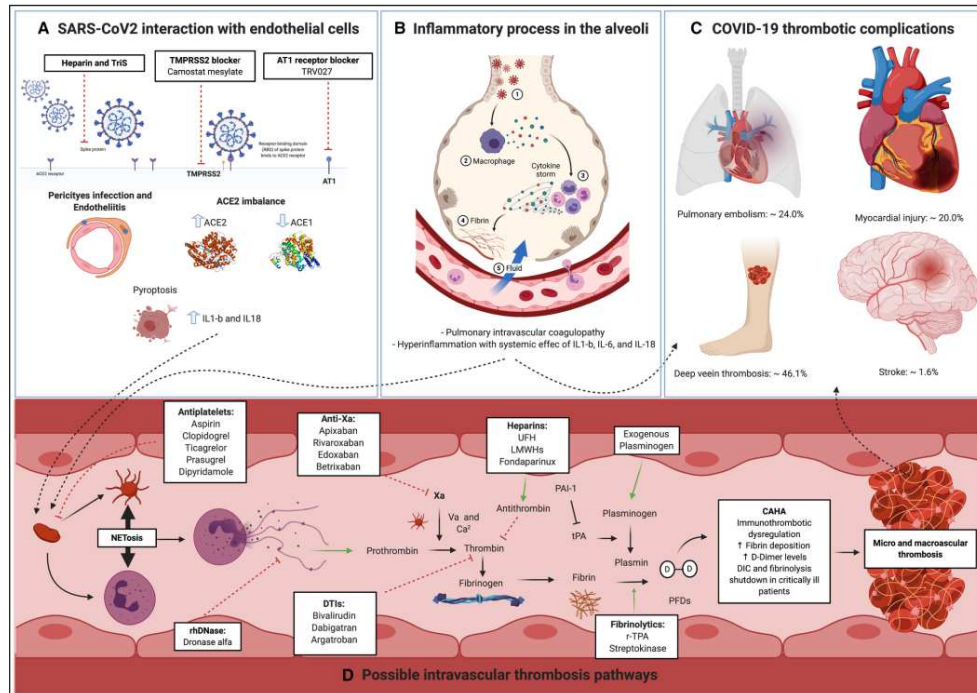
SEVERE ENDOTHELIAL DYSFUNCTION: Due to the potent pro inflammatory state there is a endothelial dysfunction “endothelitis” which in turn increases the risk of coagulation by triggering the cascade. As COVID -19 in its initial stages behaves like a acute respiratory illness it predisposes to a relative hypoxic state which further increases the risk of coagulation’s per the pathophysiology of this coagulopathy it appears that the virus in turn disturbs the micro environment of haemostasis and causes macro and microangiopathy.

With a backround of co-morbidities like diabetes, hypertension, dyslipidemia vascular disorders which potentially worsen endothelial function, predispose to this vasculopathy and can be fatal.

COAGULOPATHY HYPOTHESIS-Multiple other mechanisms like direct activation of lupus anticoagulant activity and direct fibrinogen activation were postulated which would induce a pro thrombotic state, however the trial studies had insufficient data and were were inconclusive.

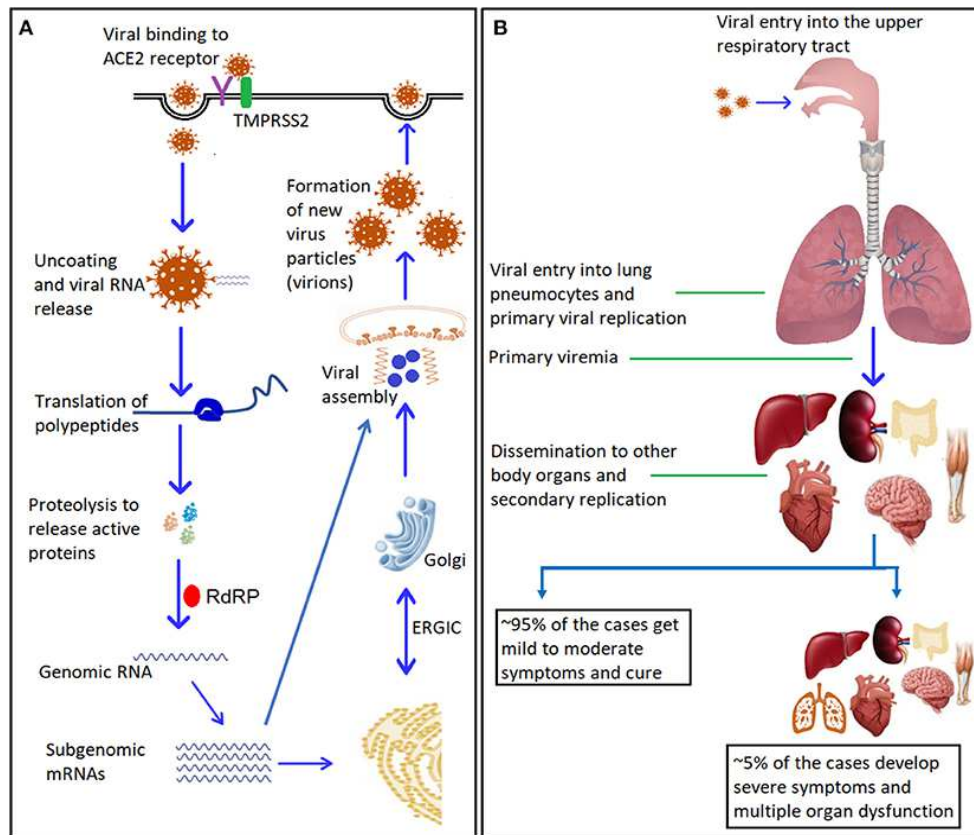
Plasminogen activator inhibitor plays a key role in regulating fibrinolytic activity. Impaired fibrinolysis or fibrinolysis shutdown has been demonstrated by thromboelastography studies in covid 19 patients postulating that the virus can inhibit Plasminogen activity.

Von willibrand factor is a multimeric protein involved in platelet adhesion and hemostasis was found to be elevated in patients with COVID 19 predisposing to thrombosis. Therefore the conclusion of this pathogenesis was that the exact mechanism remains unknown. The coagulopathy is complex and multifactorial and almost similar to the pathophysiology of sepsis associated disseminated coagulopathy.



## Transmission

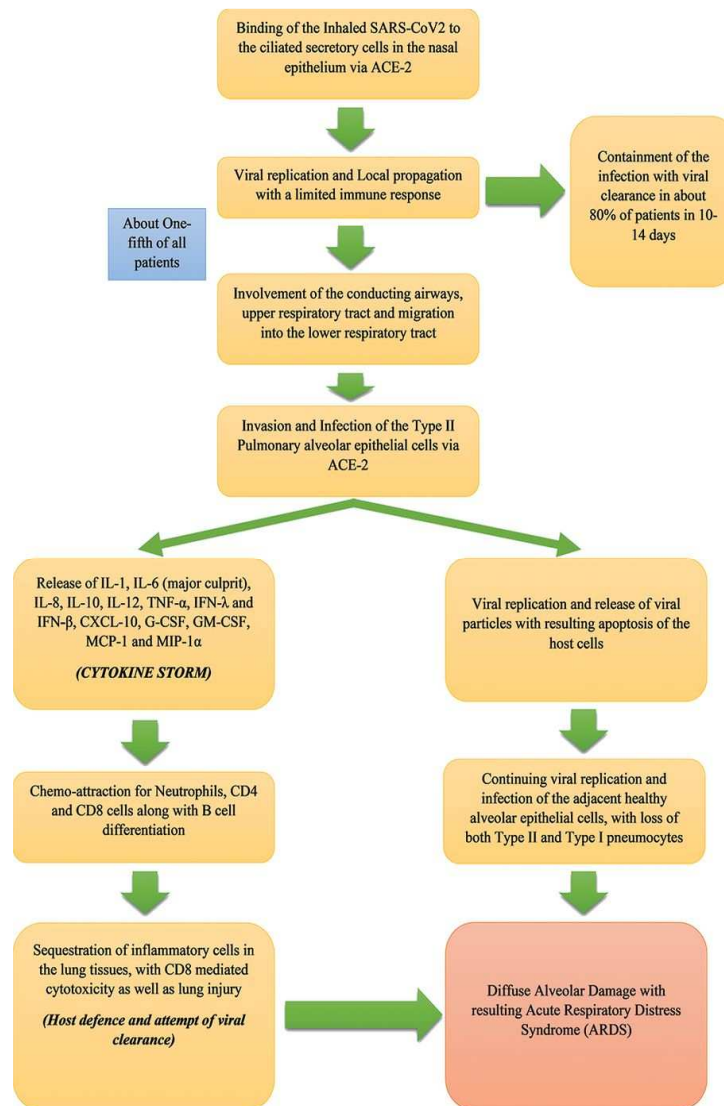
In observational studies of home transmission before the development of variants, the secondary attack rate ranged from 4 to 57 percent.<sup>26–31</sup> A meta-analysis of 87 home transmission studies including 1,249,163 household links from diverse nations showed that the secondary attack rate among adult contacts was 30%. Variations that are more transmissible boost household transmission rates. The secondary attack rate in adults was 90% in a study of family clusters during the dominant circulation of the Alpha (B.1.1.7 lineage variant).<sup>32,33</sup>



**Figure 6: Transmission and entry of COVID-19<sup>35</sup>**

There have also been reports of epidemics involving the provision of healthcare, as well as instances of potential transmission from educators to students and children in educational settings. 36–39 According to a case-control study, irregular mask use at school was linked to SARS-CoV-2 infection but not to attendance. 34

Although nothing is known about SARS-CoV-2 transmission to households in contact with confirmed asymptomatic SARS-CoV-2, transmission to asymptomatic (as opposed to presymptomatic) persons has been documented. 40 Additionally, reports of familial clusters with asymptomatic persons as well as possible transmission from asymptomatic individuals to family members have been made. 40–43 According to these data, asymptomatic patients may play a part in transmission.44,45



**Figure 7: Pathophysiology Of COVID-19<sup>20</sup>**

### **Clinical presentation**

The most common representative symptoms among people with symptomatic COVID-19 are cough, myalgias, and headache. Other characteristics include the presence of diarrhoea, severe vomiting, abnormal smell, or abnormal taste. The most common type of infection, pneumonia, is characterised by fever, coughing, dyspnea, and bilateral infiltrates on chest imaging.

---

<b>Symptoms</b>	<b>Percentage (%)</b>
Fever	88.7
Cough	67.8
Fatigue	38.1
Sputum production	33.7
Shortness of breath	18.7
Myalgia or arthralgia	14.9
Sore throat	13.9
Headache	13.6
Chills	11.5
Nausea or vomiting	5
Nasal congestion	4.8
Diarrhea	3.8

---

**Severe complications:**

The complications described as

Respiratory failure

Cardiac and cardiovascular complications

Thromboembolic complications

Neurologic complications

Inflammatory complications

Secondary infections

### **Infection fatality rate**

Only the mortality rate in reported instances is represented by the case fatality rate. Because many severe acute respiratory syndrome infections are asymptomatic and many mild infections go undiagnosed, the SARS-CoV-2 fatality rate (i.e., the estimated mortality rate among all infected individuals) has been estimated in some analyses of unvaccinated individuals to be between 0.15 and 1%, with significant heterogeneity by location and risk group. <sup>46</sup>

Using prevalence surveys or extensive tracking programmes, <sup>27</sup> research from resource-rich environments that determined the total number of community infections up to September 2020 were systematically reviewed and meta-analyzed. that the infection fatality rate would rise exponentially by age (0.002 percent at age 10, 0.01 percent at age 25, 0.4 percent at age 55, 1.4 percent at age 65, 4.6 percent at age 75, 15 percent at age 85, and >25 percent at age 90 years). The majority of the geographic variation in infection mortality rates appears to be explained by these age-related variations (i.e., locales with higher median population ages reported greater fatality rates). <sup>47</sup>

### **Risk factors for severe illness**

Although it can affect otherwise healthy people of any age, severe illness is more common in those over the age of 65 or in those who have specific underlying medical conditions. Severe illness has also been connected to certain demographic traits and aberrant test results.

Numerous predictor tests have been designed to identify patients who are more likely to have severe illness based on epidemiological, clinical, and laboratory

features; however, the majority of studies evaluating these tests are constrained by the possibility of bias and they haven't been sufficiently validated for clinical management. Increasing age

- Comorbidities
- Socioeconomic background and gender

### **Laboratory findings**

The findings Of laboratory are variable. In meta-analysis, the data Of following laboratory abnormalities were noted down

- Elevated serum ferritin: 47%
- Elevated C-reactive protein: 54%
- Elevated D-dimers: 35%
- Elevated lactate dehydrogenase: 37%
- Elevated procalcitonin: 21%
- Elevated leukocytes: 20%
- Elevated erythrocyte sedimentation rate: 19%
- Lymphocytosis: 8%
- Lymphocytopenia: 19%
- Elevated creatine kinase myocardial bands: 25%
- Elevated serum aminotransferases: 30%

Abnormality	Possible threshold
<b>Elevations in:</b>	
D-dimer	>1000 ng/mL (normal range: <500 ng/mL)
CRP	>100 mg/L (normal range: <8.0 mg/L)
LDH	>245 units/L (normal range: 110 to 210 units/L)
Troponin	>2× the upper limit of normal (normal range for troponin T high sensitivity: females 0 to 9 ng/L; males 0 to 14 ng/L)
Ferritin	>500 mcg/L (normal range: females 10 to 200 mcg/L; males 30 to 300 mcg/L)
CPK	>2× the upper limit of normal (normal range: 40 to 150 units/L)
<b>Decrease in:</b>	
Absolute lymphocyte count	<800/microL (normal range for age =21 years: 1800 to 7700/microL)

**Figure 8: Laboratory findings associated with severity of COVID infection<sup>16,17,48,49</sup>**

**Various studies with Emerging evidence of COVID 19 thrombotic syndrome has treatment implications**

To evaluate the prevalence of anticoagulant, coagulant factors, and antiphospholipid antibody in critically sick patients, Zhang Y et al. conducted a study. There were anti-phospholipid antibodies in ten of the people. Raised D-Dimer (100%), a prolonged prothrombin time (73.7%), and hyperfibrinogenemia (73.7%) were routine coagulation test abnormalities. The median activity of natural anticoagulants, including protein C, protein S, and antithrombin, was below the normal range. All patients displayed significantly above-normal factor VIII activity (median value 307%, interquartile range 198–441). The activity of factors V and VII

was significantly lower in those who were nearing the end of their lives. Unexpectedly, four incidences of cerebral infarcts were linked to individuals who had anti-phospholipid antibodies of various subtypes. Sustained hypercoagulability and thrombotic events were common in extremely ill COVID-19 patients. High Factor VIII levels, the presence of antiphospholipid antibodies, and natural anticoagulant activity may all contribute to the clogging of the arteries in people with COVID-19.<sup>50</sup> In order to determine the clinical importance of the dynamic neutrophil to lymphocyte ratio and the d-dimer in people with COVID-19 infection, Jianhong F et al. conducted a study. When compared to the mild/moderate group, the severe group's WBC, neutrophil to lymphocyte ratio (NLR), D-dimer, and fibrinogen levels were all considerably greater ( $P < 0.05$ ), but lymphocyte levels were lower. According to the RCT, the AUC for hematological parameters was higher than for inflammatory variables. Between the two groups on day one, there was a significant difference in lymphocyte and fibrinogen levels ( $P < 0.05$ ). However, on days 1, 4, and 14 both NLR and D-dimer levels were higher in the severe group compared to the mild/moderate group ( $P < 0.01$ ) ( $P < 0.05$ ). Hyperfibrinogenemia, lymphopenia, D-dimer elevation, and leukopenia were the most common COVID-19 abysmal hematological parameters upon arrival and varied considerably across the mild/moderate and severe COVID-19 groups. Additionally, it is possible to discriminate between severe and mild COVID-19 individuals based on the dynamic shift in NLR and D-dimer levels.<sup>51</sup>

To evaluate the haematological features and risk factors for pregnancies in COVID patients, Liao D et al. conducted a study. Rapid blood tests can help doctors assess the severity and prognosis of COVID-19 patients. These tests include platelet count, prothrombin time, D-dimer, and neutrophil to lymphocyte ratio. The sepsis-

induced coagulopathy score system can be used to evaluate and treat people who are critically unwell at an early stage.<sup>52</sup>

Researchers Tang N et al. conducted a study to see if reduced coagulation parameters are related to pregnancies in patients with coronavirus pneumonia. The D-dimer and fibrin degradation product (FDP) levels in non-survivors' admissible samples were significantly higher, and their prothrombin time and activated partial thromboplastin time were longer ( $P < .05$ ); 71.4% of non-survivors and 0.6% of survivors met the criteria for disseminated intravascular coagulation disease. According to the results of the current investigation, abnormal coagulation readings—notably noticeably high D-dimer and FDP—are associated with NCP fatalities.<sup>53</sup>

Garcia-Olive I et al. conducted a research to analyse the D-dimer in COVID-19 infected individuals and its result. Patients with higher D-dimer levels and those who needed intubation were more likely to develop PE. With odds ratios of 1.7, 2.0, 2.4, and 2.4, respectively, higher D-dimer levels were linked to a higher likelihood of PE three, six, nine, and twelve days after D-dimer levels were measured. Finally, people infected with COVID-19 who require OTI and have higher D-dimer levels are more likely to suffer PE.<sup>54</sup>

In a study by Ranucci M et al., fibrin lysis biomarkers and thrombin generation were used to evaluate the COVID-19-associated gastrointestinal disease. A thrombin burst and subsequent coagulation activation define severe COVID-19 ARDS patterns. The mechanisms of fibrinolysis control appear to be skewed toward inhibiting fibrinolysis. This tendency improves in survivors while worsening in non-survivors.<sup>55</sup>

Samkari HA et al. conducted a research to examine COVID-19 and coagulation abnormalities in patients. ESR, CRP, fibrinogen, ferritin, and procalcitonin levels were higher in patients with thrombotic issues than in those without. DIC, clinically significant thrombocytopenia, and low fibrinogen levels were all unusual and were associated with severe bleeding symptoms. Given the reported bleeding rates, randomised studies are necessary to establish whether enhanced anticoagulant prophylaxis may help COVID-19 patients.<sup>56</sup>

Luo HC et al. carried out a study to learn more about the characteristics of coagulation change in COVID-19 patients. A multivariate logistic analysis revealed an independent relationship between mortality and PT-act 75%. To predict death at the time of admissibility, one can use the area under the PT-act, D-dimer, and FDP receiver operating characteristic curves. The AUCs for PT-act, D-dimer, and FDP did not differ statistically significantly from one another. After taking the drug for two weeks, the coagulation levels of the survivors improved. Typically, a hormonal imbalance is linked to COVID-19. PT-act levels, D-dimer levels, and FDP levels can all be used to predict mortality in COVID-19 patients. PT-act 75% is associated with death on its own. 57

## MATERIALS AND METHODS

### MATERIAL:

Source of study: COVID 19 patients admitted in the isolation wards and intensive care units at KLE'S Dr. Prabhakar Kore Hospital and Medical Research Centre, Belagavi.

Study Design: A Cross sectional study.

Study Period: 1 year.

Sample Size: 150

Sample Size Calculation:

. The prevalence rate-based formula for the minimal sample size is

$$n = \frac{z_{\alpha}^2 P(1-P)}{d^2}$$

where d is the expected difference in prevalence and P is the prevalence percentage.

$z_{\alpha}$  is linked with the level of significance. For 5% level of the significance  $z_{\alpha} = 1.96$ .

Ref:

With  $P = 56.3\%$  and  $d = 20\%$  of  $P = 11.26\%$ , the minimum sample size is 150.

### Inclusion criteria

- Age  $\geq 18$  years.
- COVID -19 positive patients confirmed by RT-PCR/Rapid Antigen Test (RAT).

**Exclusion criteria**

- Pregnant Female patients
- End stage liver disease
- Long term Anti coagulation therapy
- Anti Folate Medication

**METHODOLOGY**

- Informed consent was obtained and then patient was enrolled for the study.
- All patients fulfilling the inclusion criteria was enrolled in the study then subjected to a questionnaire and thorough clinical examination.
- Only a confirmed case of COVID-19 that is a positive result on RT-PCR /RAT of nasal and pharyngeal swab specimens were taken into the study.
- At admission the respiratory rate, saturation at room air and requirements of oxygen delivery device or ventilation were noted.
- qSOFA score is a mortality predictor scoring system calculated by considering respiratory rate , blood pressure and mental status was calculated at the time of admission.
- All laboratory investigations including routine, coagulation profile, inflammatory markers were sent on the day of admission were taken into the study data.

- D-dimer and platelet count was repeated at 72 hrs of admission to study their variation during the course of treatment and hospital stay.
- CT thorax severity score (measured out of 25) on the day of admission was noted.
- Vaccination status with COVISHEILD VACCINE only were also noted.
- The patient's outcome was evaluated based on how long they stayed in the hospital, oxygen requirement and mode of ventilation, supportive medications and requirement of thromboprophylaxis ,steroids, immunosuppressives and fibrinolytics.
- During the course of hospital stay clinical outcome was also assessed on the basis of evidence of any thrombotic or thromboembolic event like deep vein thrombosis, pulmonary embolism ,stroke, myocardial infarction , disseminated intravascular coagulopathy or critical illness ARDS, sepsis or acute kidney injury.
- ISTH (International Society of Thrombosis & Hemostasis ) scoring system is a score to assess coagulopathy which considers platelet count, fibrinogen level ,PTINR and D-dimer and is expressed as <5 or >5 .This score was calculated from the study data to co-relate with the clinical outcome of the patients included in the study

---

Variable	Value	Points
Platelets, K/ $\mu$ L	>100	0
	50-100	1
	<50	2
INR	<1.3	0
	1.3-1.7	1
	>1.7	2
D-Dimer, ng/mL	<400	0
	400-4000	2
	>4000	3
Fibrinogen, mg/dL	>100	0
	<100	1

---

Abbreviations: DIC, disseminated intravascular coagulation; INR, international normalized ratio; ISTH, International Society on Thrombosis and Haemostasis.

**SCORE OF >5 is suggestive of overt DIC**

Investigations for the study purpose

Following Laboratory investigations will be considered-

A) **Basic Investigations-**

1. Complete blood count
2. Random blood sugar
3. Blood urea
4. Serum Creatinine
5. Liver function test

B) **COVID Profile**

COAGULATION PROFILE

PT INR

APTT

Serum D-dimer

Serum Fibrinogen

Platelet count

**INFLAMMATORY MARKERS**

- 1) LDH.
- 2) Sr. Ferritin,
- 3) hs -CRP.
- 4) IL-6.
- 5) Sr. Procalcitonin

**RADIOLOGICAL INVESTIGATION:**

HRCT-THORAX: CORADS SCORE out of 25

## STATISTICAL ANALYSIS

This was an observational research, the analytic plan was as follows.

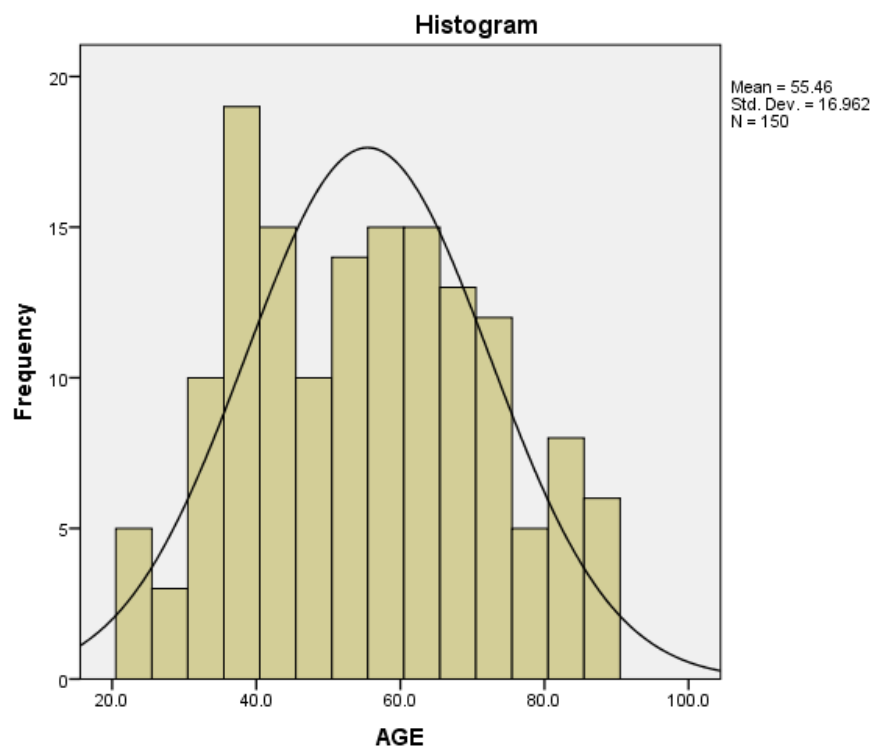
The mean and standard deviation of the continuous quantitative variables were computed. If the data is separated into two groups based on a certain qualitative trait, the continuous variables were compared using appropriate statistical methods such as the student's unpaired t test. Rates, ratios, and percentages were used to convey categorical data. The association between the result, clinical, and demographic factors was evaluated using the Chi-square test, test of proportion, or Fisher's exact test. For all tests, a p value of less than 5% (0.05) was regarded as significant.

**RESULTS**

In present study, total of 150 patients included after obtaining the informed consent. The mean age of the patients was found to be 55.45yrs.

**Table 1: Mean age of the participants included in the study**

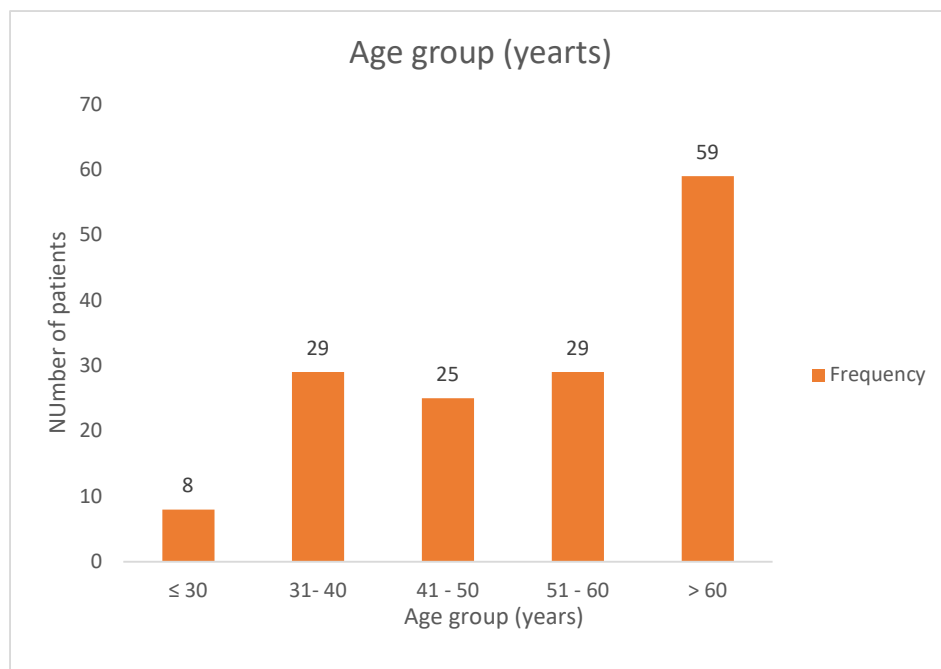
	N	Minimum	Maximum	Mean	SD
AGE	150	23.0	90.0	55.460	16.96



**Figure 9: Mean age of the patients included in the study**

**Table 2: Distribution of age (years) of the participants included in the study**

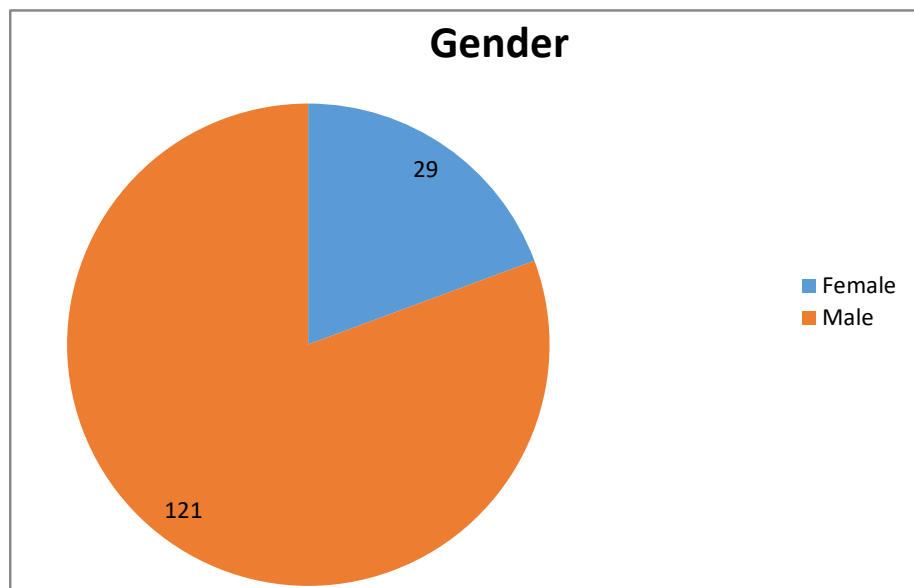
Age group					
	Age group	Frequency	Percent	Valid Percent	Cumulative Percent
Valid	≤ 30	8	5.3	5.3	5.3
	31- 40	29	19.3	19.3	24.7
	41 - 50	25	16.7	16.7	41.3
	51 - 60	29	19.3	19.3	60.7
	> 60	59	39.3	39.3	100
	Total	150	100	100	



**Table 3: Gender distribution of study participants**

		Frequency	Percent
Gender	Female	29	19.3
	Male	121	80.7
	Total	150	100.0

Among them 80.7% were male patients and 19.3% were female with male preponderance in the study.

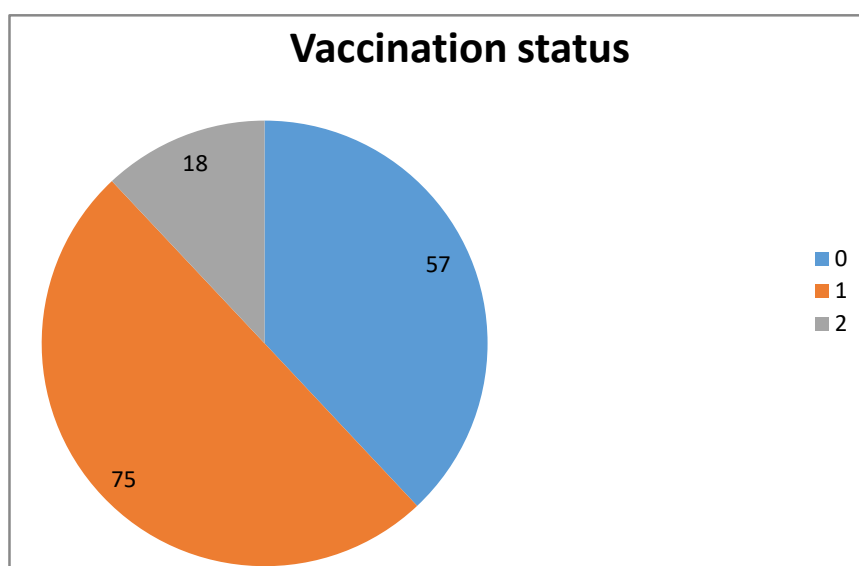


**Figure 10: Gender distribution of study participants**

**Table 4: Vaccination status of the patients included in the study**

		Frequency	Percent
<b>Vaccination status</b>	Not vaccinated (0)	57	38.0
	1.0	75	50.0
	2.0	18	12.0
	Total	150	100.0

Vaccination status was found to be 38% without vaccination, 50% with first dose and 12% with second dose.

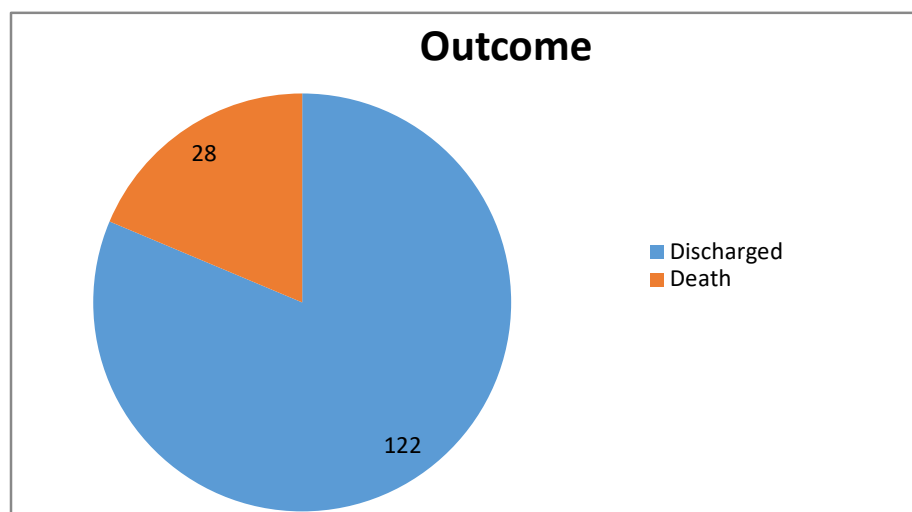


**Figure 11: Vaccination status of the patients included in the study**

**Table 5: Outcome of the patients in the study**

		Frequency	Percent
<b>Outcome</b>	Discharged	122	81.3
	Death	28	18.7
	Total	150	100.0

On assessment of outcome, 18.7% had the worst outcome as death, 81.3% were discharges. 14 patients discharged were worsened at the time of discharged and sent to higher centre.



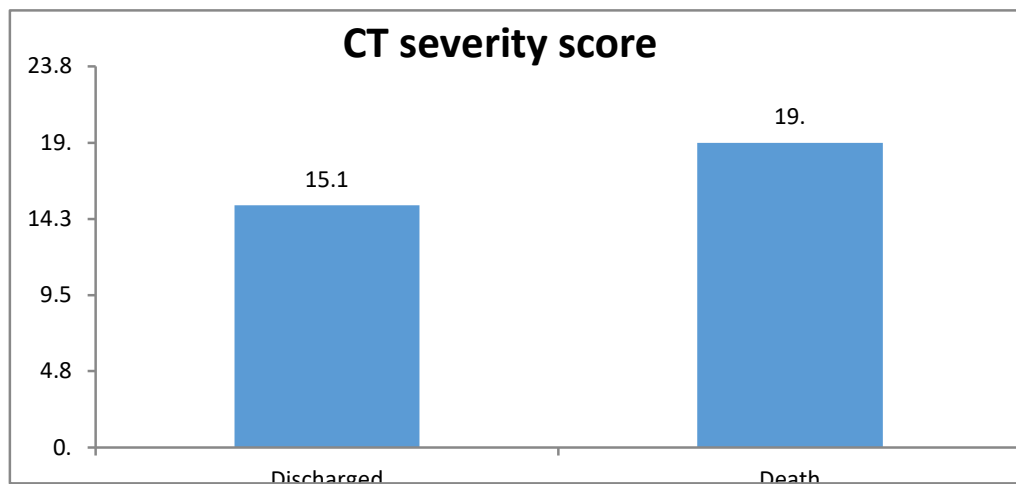
**Figure 12: Outcome of the patients in the study**

**Table 6: Comparison of CT severity score with outcome among study participants**

	Outcome				p-value
	Discharged		Death		
	Mean	SD	Mean	SD	
CT severity score	15.08	5.38	19.04	5.03	< 0.001*

\*Significant (p-value < 0.05) Unpaired t-test used

On assessment of the parameters between the outcome of patients, the CT severity score was significantly higher in patient with worst outcome as death compared to the patients discharged.



**Figure 13: Comparison of CT severity score with outcome among study participants**

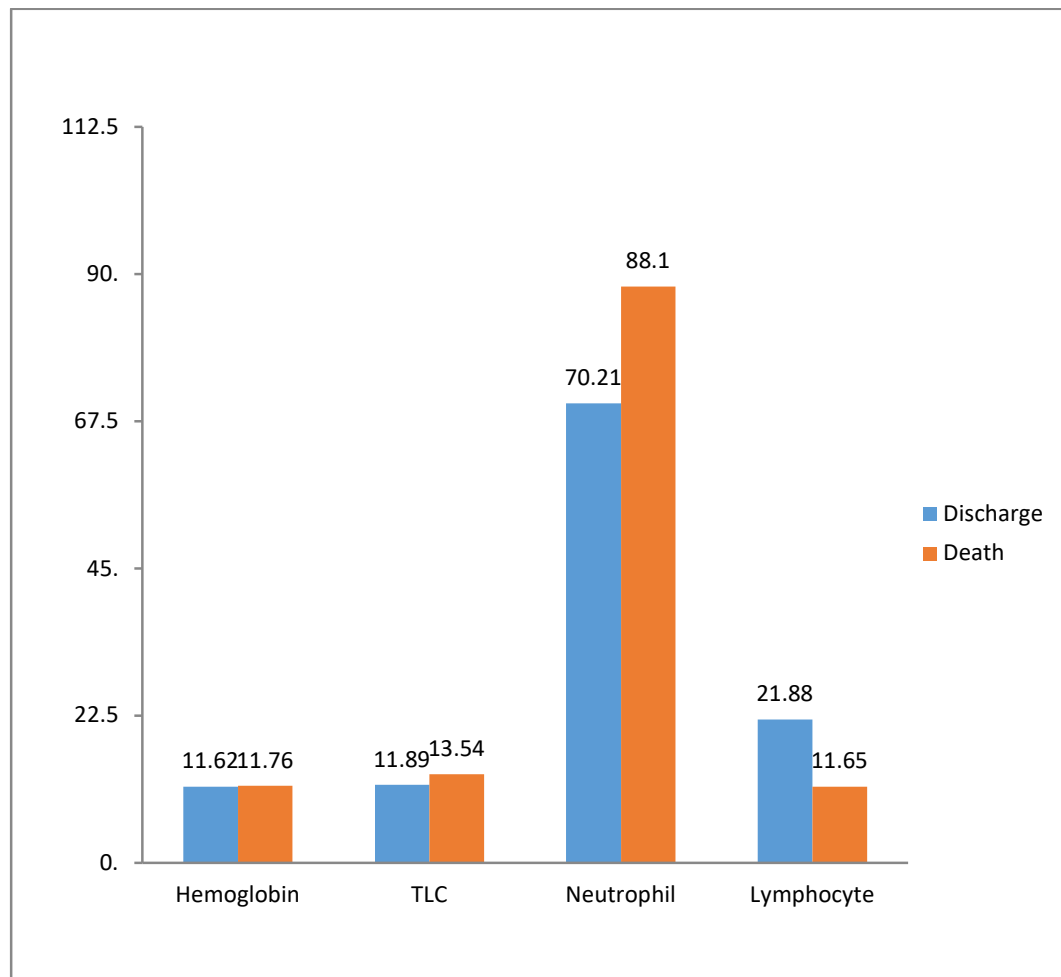
**Table 7: Comparison of the mean level of blood parameters with outcome of patients**

Parameter	Discharge (n=122)		Death (n=28)		p-value
	Mean	SD	Mean	SD	
Hemoglobin	12.85	2.16	13.10	2.53	0.619
Total leucocyte count	10.87	5.95	11.04	8.16	0.920
Neutrophils	82.96	10.74	78.75	10.94	0.073
Lymphocyte	12.64	11.26	14.18	9.98	0.477
INR	1.24	0.26	1.22	0.21	0.627
APTT	1.02	0.28	1.05	0.27	0.599

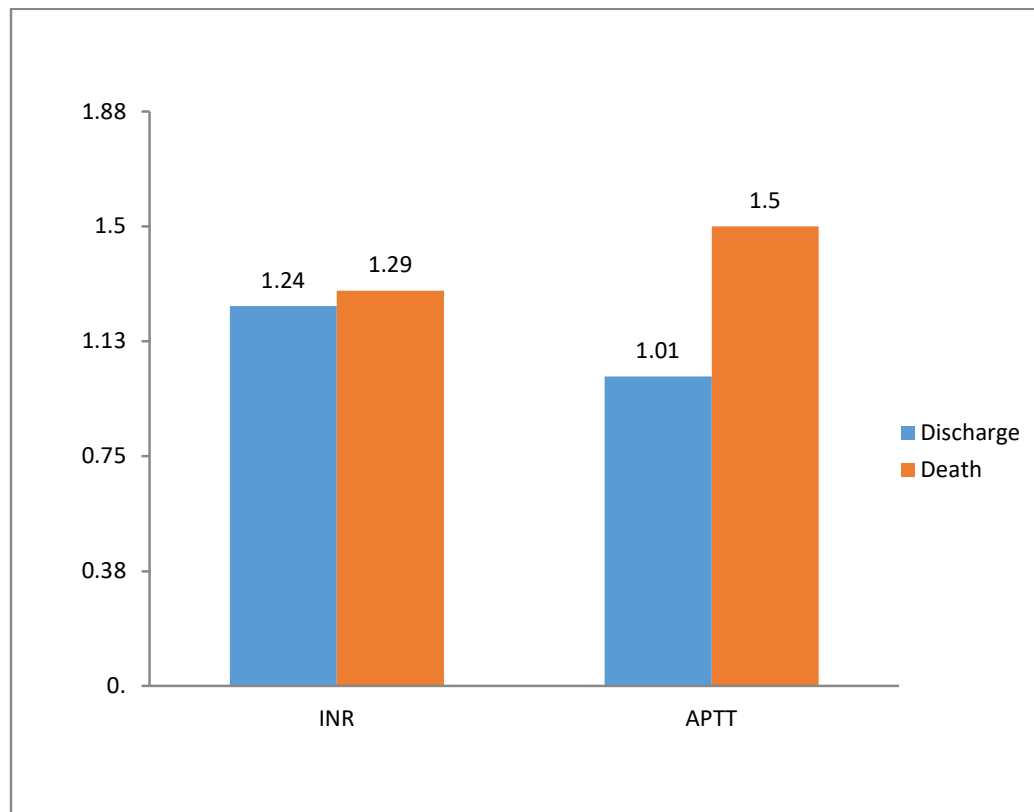
p-value > 0.05 (Not Significant) Unpaired t-test used

Hemoglobin level, total leucocyte count, neutrophil percent, lymphocyte percent

APTT, PT-INR level did not have any statistical significance with outcome of the patient.



**Figure 14: Comparison of the mean level of blood parameters with outcome of patients**



**Figure 15: Comparison of the mean level of blood parameters with outcome of patients**

**Table 8: Comparison of the biomarkers with outcome of the study participants**

Parameter	Discharge (n=122)		Death (n=28)		p-value
	Mean	SD	Mean	SD	
Ferritin	858.88	1418.53	933.07	915.50	0.439
Fibrinogen	426.93	245.13	391.46	198.58	0.652
LDH	537.48	370.67	648.32	251.12	0.007*
Homocysteine	26.45	59.26	46.15	144.17	0.723
HsCRP	98.52	115.12	146.60	93.65	0.002*
IL-6	84.11	147.68	71.70	116.84	0.791
D-Dimer	715.74	597.95	3180.12	2038.27	< 0.001*

\*Significant (p-value < 0.05) Mann-Whitney U test used

There was significant higher mean level of the ferritin in patients with worst outcome as death. The mean level of LDH, Ferritin, Fibrinogen, Homocysteine, D-dimer and CRP, IL6 was significantly higher among the patients with worst outcome as death compared to patients discharged. LDH, HsCRP, and D-dimer levels were significantly higher and directly related to mortality in the given study.

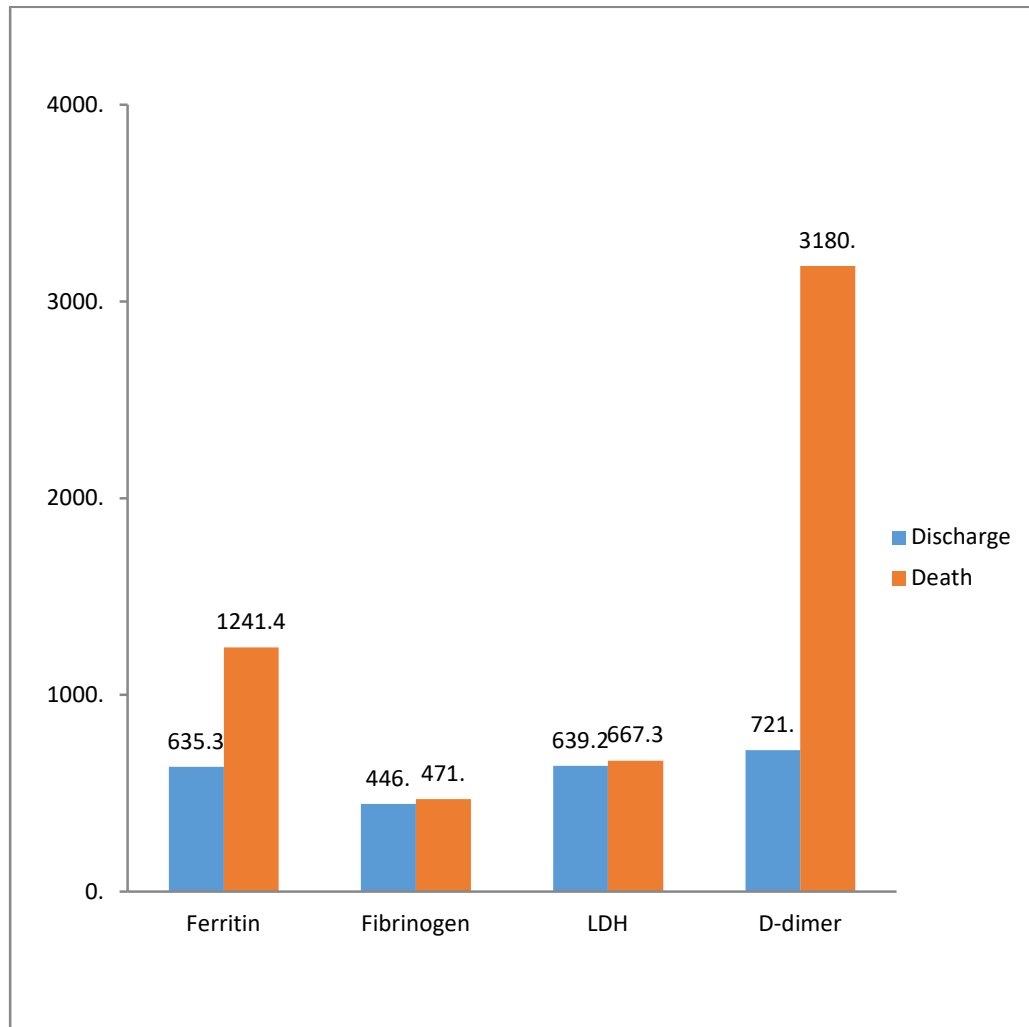
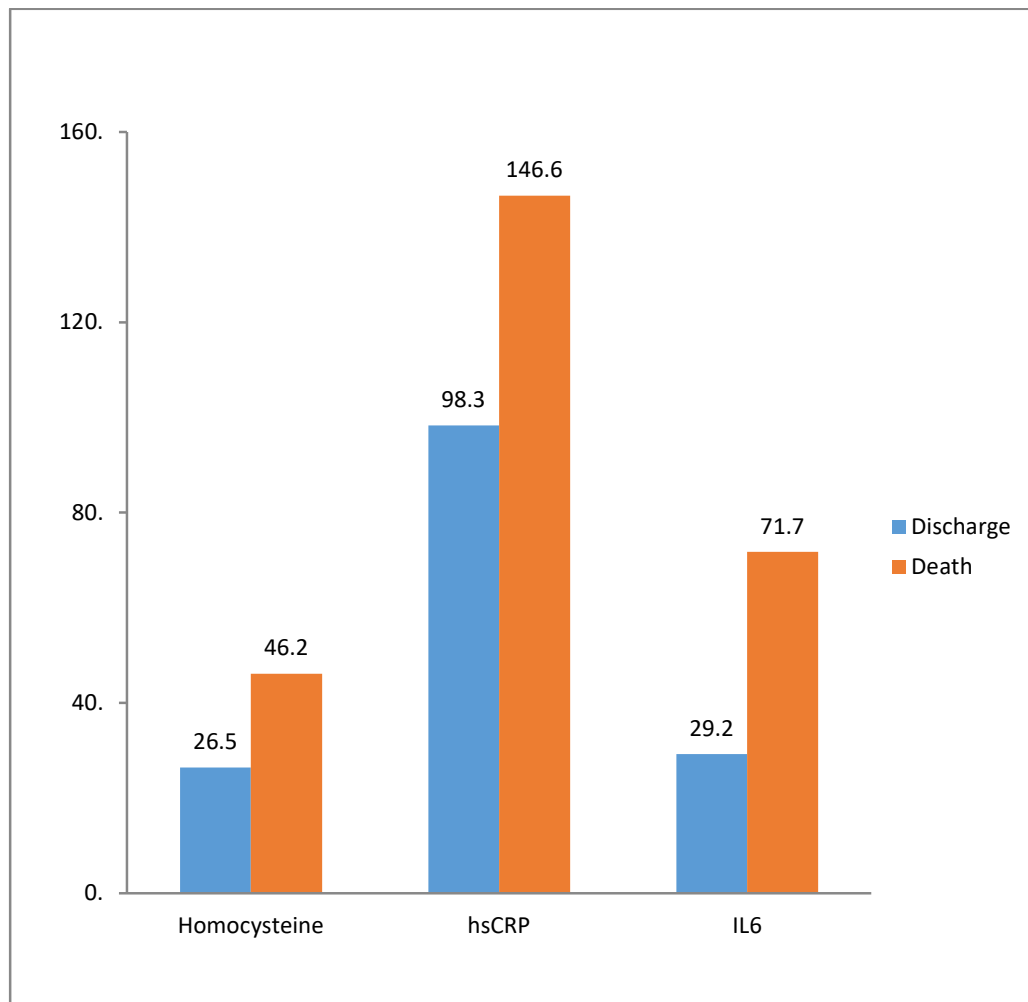


Figure 16: Comparison of the biomarkers with outcome of the study participants



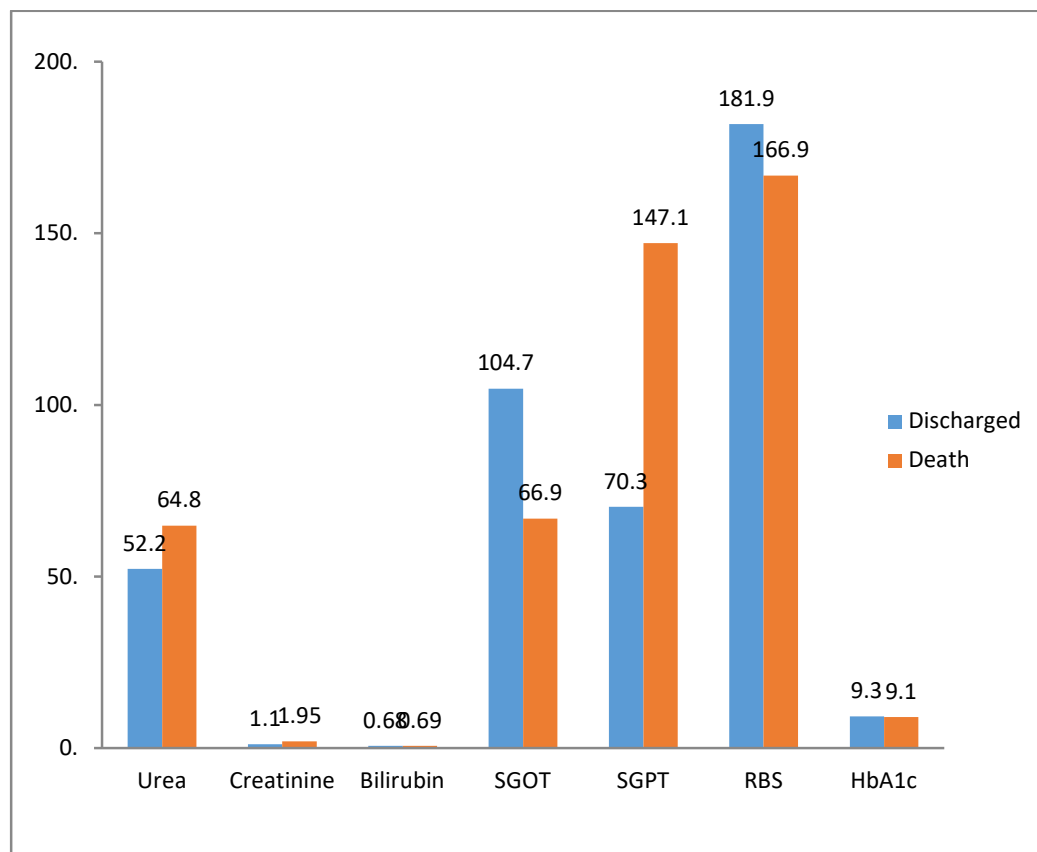
**Figure 17: Comparison of the biomarkers with outcome of the study participants**

**Table 9: Comparison of the renal and liver function test parameters with outcome of the patients**

Parameter	Discharge (n=122)		Death (n=28)		p-value
	Mean	SD	Mean	SD	
Urea	52.27	45.37	64.75	54.72	0.037*#
Creatinine	1.47	1.87	1.46	1.71	0.980 \$
Bilirubin	0.68	0.97	0.69	0.53	0.892 \$
SGOT	104.43	670.14	66.86	64.55	0.100 #
SGPT	40.23	44.77	68.86	92.65	0.325 #
RBS	181.86	101.79	166.86	71.55	0.794 #
HbA1c	2.16	3.67	2.09	5.40	0.962 \$

\*Significant (p-value < 0.05), # Mann-Whitney U test used,  
\$ Unpaired t-test used

In the study there is no significant difference in the mean level of the blood urea, creatinine and the liver profile.



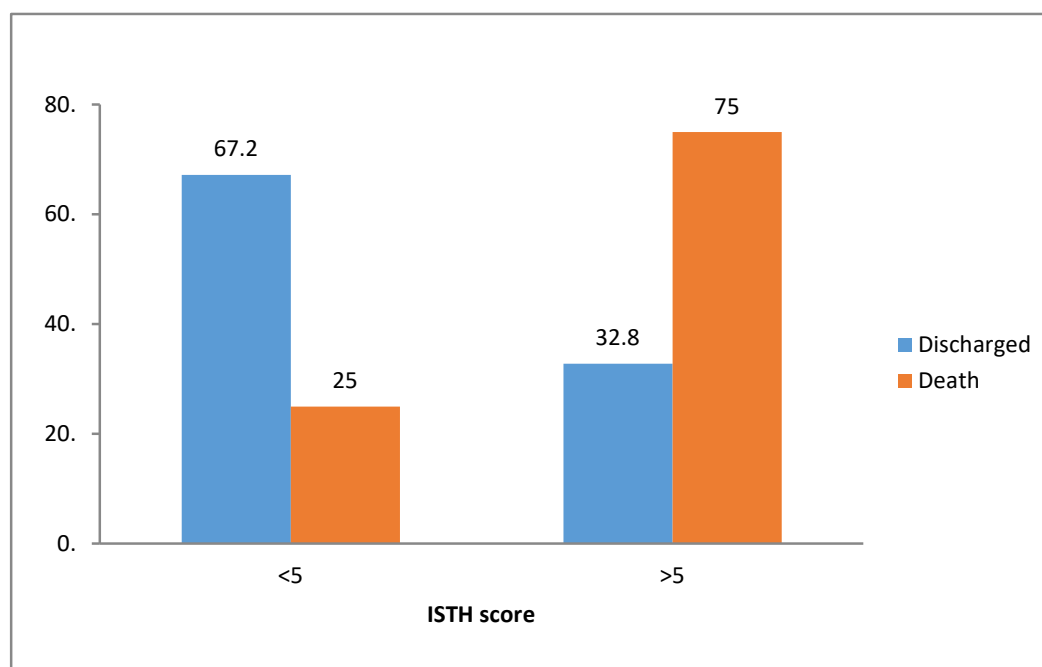
**Figure 18: Comparison of the renal and liver function test parameters with outcome of the patients**

**Table 10: Comparison of the ISTH score with outcome of the patients**

		Outcome				p-value
		Death		Discharged		
		Count	%	Count	%	
ISTH score	> 5	21	75.0%	40	32.8%	< 0.001*
	< 5	7	25.0%	82	67.2%	
Total		28	100.0%	122	100.0%	

\*Significant (p-value < 0.05) Chi-square test used

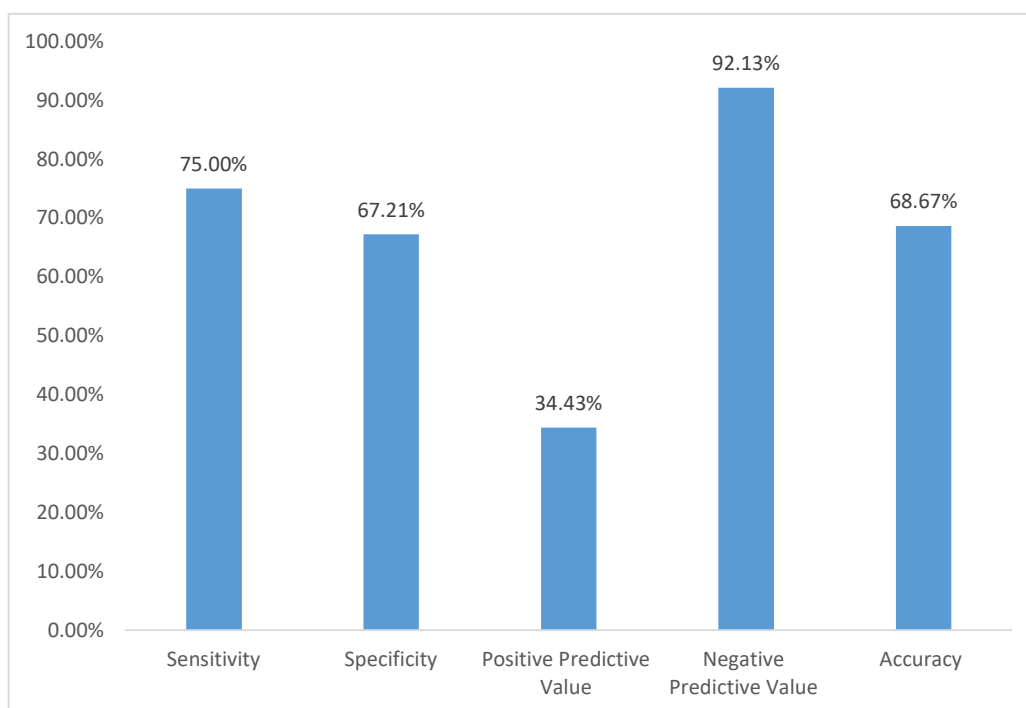
There was significant higher incidence of death among the patients with ISTH score >50 (75%) compared to patients discharged.



**Figure 19: Comparison of the ISTH score with outcome of the patients**

**Table 11: Diagnostic characteristics of ISTH score to predict the outcome**

Statistic	Value	95% CI
Sensitivity	75.0%	55.13% to 89.31%
Specificity	67.21%	58.13% to 75.44%
Positive Predictive Value	34.43%	27.36% to 42.26%
Negative Predictive Value	92.13%	85.90% to 95.75%
Accuracy	68.67%	60.59% to 75.98%

**Figure 20: Diagnostic characteristics of ISTH score to predict the outcome**

<b>Table 12: Co-relation of CT SEVERITY SCORE WITH ISTH SCORE</b>					
	ISTH SCORE RE DIC _Group	N	Mean	SD	p-value
CT severity score	< 5	89	15.3	5.382	0.172
	> 5	61	16.57	5.682	

**There is no direct co relation with CT severity and ISTH SCORE with  
p value of 0.172**

<b>Table 13: Co-relation of DAY 3 PLATELET COUNT AND MORTALITY</b>					
	Mortality	N	Mean	SD	p-value
Plateletsday 1	Yes	28	221.89	122.37	0.818
	No	122	227.64	97.67	
Platelet day 3	Yes	28	168.36	108.26	0.009
	No	122	229.44	92.62	

**It was noted that the mean platelet count at 72 hrs was 168.36 with a drop of 53.5 from platelet count at admission and was statistically significant with mortality of patient.**

<b>Table 14: Co-relation of DAY 3 D-dimer level AND MORTALITY</b>					
	Mortality	N	Mean	SD	p-value
d_Dimer_1	Yes	28	3180.12	2038.27	< 0.001
	No	120	715.74	597.95	
d_Dimer_3	Yes	28	3344.39	2083.14	< 0.001
	No	122	1256.32	1283.89	

**It was noted that the mean platelet count at 72 hrs was 3344.39 with a was statistically significant with mortality of patient.**

<b>Table 15: Co-relation of vaccination status to ISTH SCORE</b>					
Count					
		ISTH SCORE DIC _Group		Total	p-value
		< 5	> 5		
Vaccination status	0	27	30	57	0.057
	1	49	26	75	0.034
	2	13	5	18	0.027
Total		89	61	150	

<b>Table 16: Co-relation of vaccination status to mortality</b>					
Count					
		Mortality		Total	p-value
		Yes	No		
Vaccination status	0	13	44	57	0.495
	1	13	62	75	0.461
	2	2	16	18	0.472
Total		28	122	150	

**It was noted that vaccination status of 1 and 2 had a significantly low ISTH score at admission. There was no significant relation of vaccination status and mortality.**

<b>Table 17: Co-relation of T2DM AND MORTALITY</b>					
Count					
		Mortality		Total	p-value
		Yes	No		
DM	Present	18	61	79	0.172
	Absent	10	61	71	
Total		28	122	150	

<b>Table 18: Co-Relation of T2DM with ISTH SCORE</b>					
Count					
		ISTH SCORE DIC _Group		Total	p-value
		< 5	> 5		
DM	Present	47	32	79	0.966
	Absent	42	29	71	
Total		89	61	150	

**There was no statistically significant co relation between Diabetic patients and mortality.**

## **DISCUSSION**

Thrombotic events such deep vein thrombosis, pulmonary embolism, myocardial infarction, and ischemic strokes have become more common in critically ill patients. Contrary to popular belief ,thromboembolic events have been clinically observed, the specific mechanism is unknown. The new SARS COV2 virus has a considerable influence on the human vasculature, manifesting as dynamic changes in the coagulation profile and perhaps suggesting the patient's prognosis.

COVID-19 thrombotic syndrome is an expanding spectrum that requires further research and analysis to accomplish optimal treatment goals, enhance prognosis, reduce post-infection consequences, and develop a robust prediction score system to anticipate this elusive and complicated coagulopathy. There is no question that DIC diagnosis and risk categorization are essential in the triage, prognosis, and treatment of COV-19 patients, particularly in cases of urgency, given the dismal prognosis of DIC. ISTH offers flexible, highly valid, laboratory-based scoring methods for identifying patients with overt DIC and septic persons who are prone to DIC regardless of clinical symptoms. 58

After receiving informed consent, a total of 150 patients were included in the study. The mean age of the patients was found to be 55.45yrs, among them 80.7% were male patients and 19.3% were female with male preponderance in the study. In the study 28.7% required oxygen mask and 28% were on room air. Vaccination status was found to be 38% without vaccination, 50% with first dose and 12% with second dose.

In study by Bauer W et al., documented the mean age of the patients as 70.0yrs with 48% were female.<sup>59</sup> In another study, the severe cases with mean age of 65yrs with male preponderance (61%).<sup>60</sup> Men and women's immune systems differ biologically, which may affect our capacity to fight viruses like SARS-2-CoV-2. Females are often more resistant to infections than men, which may be due to a combination of variables such as sex hormones and high expression of coronavirus receptors (ACE 2) in men, as well as lifestyle factors such as greater levels of smoking and drinking in men compared to women.

On assessment of outcome, 18.7% had the worst outcome as death, 81.3% were discharges. Fourteen patients discharged were worsened at the time of discharged and sent to higher centre. On assessment of the parameters between the outcome of patients, the CT severity score was significantly higher in patient with worst outcome as death compared to the patients discharged. Similarly, there was significant higher mean level of the ferritin in patients with worst outcome as death. The mean level of Fibrinogen, Homocysteine, D-dimer was significantly higher among the patients with worst outcome as death compared to patients discharged. There was significant higher incidence of death among the patients with ISTH score >5 (75%) compared to patients discharged. The mechanisms of fibrinolysis control appear to be skewed toward inhibiting fibrinolysis. This tendency improves in survivors while worsening in non-survivors.<sup>55</sup>

In addition to thrombosis and pulmonary embolism, D-dimer can indicate a significant viral infection. A viral infection can produce sepsis and coagulation disruption, which is common in the evolution of severe disease. In addition, a rise in D-dimer levels may be an indirect symptom of an inflammatory response, since

inflammatory cytokines may produce an imbalance of coagulation and fibrinolysis in the alveoli, stimulating the fibrinolysis system and so elevating D-dimer levels.<sup>61,62</sup>

In a study by Luo HC et al., the PT-act, D-dimer, and FDP may be used to predict death in a manner similar to the current study. There was no statistically significant difference between the AUCs for PT-act and those for D-dimer and FDP. After two weeks of medication, the coagulation parameters of the patients who survived improved. Uneven coagulation is usually linked to COV-19. In COV-19 patients, PT-act levels upon admission, D-dimer levels, and FDP values can all predict death. <75% of PT-acts are connected to death on their own. 57

Yu HH et al., documented that Patients with severe COVID-19 had higher D-dimer levels than those with non-severe illness, and D-dimer levels more than 0.5 g/ml are related with severe infection in COVID-19 patients.<sup>63</sup>

In study by Garcia-Olive et al., documented Patients with greater D-dimer values and those requiring intubation were more likely to develop PE. Three, six, nine, and twelve days after D-dimer levels were measured, with ORs of 1.7, 2.0, 2.4, and 2.4, respectively, higher D-dimer levels were linked to a higher risk of PE. Last but not least, individuals with COVID-19 infection who need OTI and have higher D-dimer levels are more likely to experience PE. 54 The levels of D-dimer and fibrin degradation product (FDP), as well as the prothrombin time and activated partial thromboplastin time, were significantly higher in non-survivors at the time of admission (P.05); 71.4% of non-survivors and 0.6% of survivors met the criteria for disseminated intravascular coagulation. The current investigation found that NCP deaths frequently have aberrant coagulation values, particularly noticeably high D-dimer and FDP. 53

As demonstrated by Liao D et al., rapid blood tests can assist doctors in determining the severity and prognosis of COV-19 patients. These tests include platelet count, prothrombin time, D-dimer, and neutrophil to lymphocyte ratio. Early evaluation and management of patients with severe illness can be done using the sepsis-induced coagulopathy scoring system. 52 In another study, NLR and D-dimer levels were higher on days 1, 4, and 14 in the severe group compared to the mild/moderate group ( $P<0.01$ ), and D-dimer levels were higher on days 1, 7, and 14 ( $P<0.05$ ). Hyperfibrinogenemia, lymphopenia, D-dimer elevation, and leukopenia were the most prevalent COV-19 aberrant haematological indices upon arrival, and their prevalence varied considerably across the mild/moderate and severe COV-19 groups. Additionally, the dynamic variation of NLR and D-dimer levels can differentiate between patients with severe and mild COVID-19. 51

According to Samkari HA et al., DIC, clinically significant thrombocytopenia, and low fibrinogen levels are unusual and are connected with signs of extreme bleeding. Given the reported bleeding rates, randomised studies are necessary to establish whether enhanced anticoagulant prophylaxis may help COVID-19 patients.<sup>56</sup>

Higher ISTH and SIC scores are linked to increased sickness severity and death, according to research by Hosseini S. et al., which is similar to the current study. Additionally, more nonsurvivors and patients with severe illnesses met the ISTH and SIC DIC classifications. Given the high prevalence of coagulopathy associated with COVID-19 infection, it is essential to continuously monitor important indicators, such as the ISTH and SIC scoring systems, in order to identify DIC in suspect people as soon as possible. 58

Zhang Y et al. found that thrombotic events and persistent hypercoagulability were frequent in the critically ill COVID-19 patients. The presence of

antiphospholipid antibodies, high factor VIII levels, and natural anticoagulant activity could all contribute to the coagulopathy etiology in COVID-19 patients. Additionally, standard coagulation tests revealed elevated D-Dimer (100%), an extended prothrombin time (73.7%), and hyperfibrinogenemia (73.7%). Protein C, protein S, and antithrombin all had median activities of natural anticoagulants that were below the normal range. 50

Another study by Liu Y et al. found that the PT, DD, and FDP values were positively associated to the traditional APACHE II, SOFA, and qSOFA scores, but adversely related to the ATIII ( $p < 0.001$ ). High levels of PT, FDP, and DD were linked to in-hospital mortality ( $p < 0.001$ ). Finally, blood coagulation issue was prevalent in COVID-19 ICU patients and was linked to a variety of inflammatory variables. Abnormalities in blood coagulation parameters may be a risk factor for COVID-19 ICU patients.<sup>64</sup>

Thrombocytopenia was connected to coagulation function, DIC rate, and survival in a research by Bao C et al. Six of the seven fatal patients experienced thrombocytopenia while in the hospital, and their platelet count fell until they died. Within one week of being hospitalised, six recovered patients developed thrombocytopenia. Higher platelet levels were followed by positive SARS-CoV-2 IgM/IgG and negative coronavirus nucleic acid tests in 8 recovered individuals. A low platelet count is associated with poor coagulation function, an increased risk of DIC, severe disease presentation, and mortality in COVID-19 patients.<sup>65</sup>

The present study documented the significant changes in coagulation profile in patients with COVID-19 infection.

## CONCLUSION

The present study documented the significant changes in the coagulation profile among the patients with worst outcome as death of the patients.

D-dimer level at admission and at 72 hours was significantly higher in patients with the worst outcome.

There is a higher incidence of the inflammatory markers and deranged coagulation profile among critical patients as compared to the patients discharged from hospital.

Vaccinated patients had a lower ISTH score at admission as compared to non-vaccinated patients.

The ISTH score was found to have sensitivity and specificity to detect the worst outcome among the patients. This knowledge is helpful to assess and prevent the further complications among the patients.

## SUMMARY

The patients who were admitted to the hospital with COVID-19 infection were the subjects of the current cross-sectional study. All patients fulfilling the inclusion criteria will be enrolled in the study then subjected to a questionnaire and thorough clinical examination. Outcome of the patient were assessed in terms of duration of stay in hospital, oxygen requirement, and condition of the patient at the time of discharge and mortality. The coagulation profile included the platelet count, INR, aPTT, serum D-dimer, fibrinogen, other inflammatory markers like LDH, hs-CRP, Ferritin and IL-6. The data were analysed and tabulated with the SPSS 21v, with  $p < 0.05$  was considered statistically significant.

- After receiving informed consent, a total of 150 patients were included in the study.
- The mean age of the patients was found to be 55.45yrs.
- Among them 80.7% were male patients and 19.3% were female with male preponderance in the study.
- In the study 28.7% required oxygen mask and 28% were on room air.
- Vaccination status was found to be 38% without vaccination, 50% with first dose and 12% with second dose.
- On assessment of outcome, 18.7% had the worst outcome as death, 81.3% were discharges.
- 14 patients discharged were worsened at the time of discharged and sent to higher centre.
- On assessment of the parameters between the outcome of patients, the CT severity score was significantly higher in patient with worst outcome as death compared to the patients discharged.

- Similarly, there was significant higher mean level of the D-DIMER in patients with worst outcome as death.
- The mean level of Fibrinogen, Homocysteine, D-dimer was significantly higher among the patients with worst outcome as death compared to patients discharged.
- LDH, Ferritin and D-DIMER Levels at admission were significantly higher at admission in patients with the worst outcome.
- Platelet level at 72 hrs were significantly lower (Mean:168.36) in patients with the worst outcome compared to those discharged.
- D-dimer level at admission and at 72 hrs was significantly higher (Mean:3344.39) in patients with the worst outcome.
- Vaccination status of 1 or 2 doses had a significantly lower ISTH score at admission. However there was no statistical significance of vaccination status with outcome of the patients.
- Patients with diabetes mellitus did not show any statistical significant relation with ISTH score or mortality in this study.

When compared to patients who had been discharged, patients with an ISTH score of  $>5$  had a significantly greater likelihood of death (75%).

**BIBLIOGRAPHY**

1. Weiss SR, Navas-Martin S. Coronavirus pathogenesis and the emerging pathogen severe acute respiratory syndrome coronavirus. *Microbiol Mol Biol Rev.* 2005;69(4):635–64.
2. World Health Organization. Director-General’s remarks at the media briefing on 2019-nCoV on 11 February 2020 [Internet]. 2020. Available from: <http://www.who.int/dg/speeches/detail/who-director-general-s-remarks-at-the-media-briefing-on-2019-ncov-on-11-february-2020>
3. World Health Organization (WHO). WHO Director-General’s opening remarks at the media briefing on COVID-19 -- 11 March 2020. 2020.
4. Chams N, Chams S, Badran R, Shams A, Araji A, Raad M, et al. COVID-19: A Multidisciplinary Review. *Front Public Heal* [Internet]. 2020;8:1–15. Available from: <https://www.frontiersin.org/articles/10.3389/fpubh.2020.00383>
5. Shanmugam C, Mohammed AR, Ravuri S, Luthra V, Rajagopal N, Karre S. COVID-2019 - A comprehensive pathology insight. *Pathol Res Pract.* 2020;216:153222.
6. Rothe C, Schunk M, Sothmann P, Bretzel G, Froeschl G, Wallrauch C, et al. Transmission of 2019-nCoV Infection from an Asymptomatic Contact in Germany. *N Engl J Med.* 2020;382(10):970–1.
7. Tsang KW, Ho PL, Ooi GC, Yee WK, Wang T, Chan-Yeung M, et al. A Cluster of Cases of Severe Acute Respiratory Syndrome in Hong Kong. *N Engl J Med.* 2003;348(20):1977–85.
8. Kim JY, Ko J-H, Kim Y, Kim Y-J, Kim J-M, Chung Y-S, et al. Viral Load Kinetics of SARS-CoV-2 Infection in First Two Patients in Korea. *J Korean Med Sci.* 2020;35(7).

9. Holshue ML, DeBolt C, Lindquist S, Lofy KH, Wiesman J, Bruce H, et al. First Case of 2019 Novel Coronavirus in the United States. *N Engl J Med.* 2020;382(10):929–36.
10. Poutanen SM, Low DE, Henry B, Finkelstein S, Rose D, Green K, et al. Identification of Severe Acute Respiratory Syndrome in Canada. *N Engl J Med.* 2003;348(20):1995–2005.
11. Chen H, Guo J, Wang C, Luo F, Yu X, Zhang W, et al. Clinical characteristics and intrauterine vertical transmission potential of COVID-19 infection in nine pregnant women: a retrospective review of medical records. *Lancet.* 2020;395(10226):809–15.
12. Morawska L, Tang JW, Bahnfleth W, Bluyssen PM, Boerstra A, Buonanno G, et al. How can airborne transmission of COVID-19 indoors be minimised? *Environ Int.* 2020;142:105832.
13. Gao P, Zhang H, Wu Z, Wang J. Visualising the expansion and spread of coronavirus disease 2019 by cartograms. *Environ Plan A Econ Sp.* 2020;52(4):698–701.
14. Zimmermann P, Curtis N. Coronavirus Infections in Children Including COVID-19: An Overview of the Epidemiology, Clinical Features, Diagnosis, Treatment and Prevention Options in Children. *Pediatr Infect Dis J.* 2020;39(5).
15. Ciaglia E, Vecchione C, Puca AA. COVID-19 Infection and Circulating ACE2 Levels: Protective Role in Women and Children. *Front Pediatr.* 2020;8.
16. Guan W, Ni Z, Hu Y, Liang W, Ou C, He J, et al. Clinical Characteristics of Coronavirus Disease 2019 in China. *N Engl J Med.* 2020;382(18):1708–20.
17. Wang D, Hu B, Hu C, Zhu F, Liu X, Zhang J, et al. Clinical Characteristics of

- 138 Hospitalized Patients With 2019 Novel Coronavirus–Infected Pneumonia in Wuhan, China. *JAMA*. 2020;323(11):1061–9.
18. Cheng Y, Cheng G, Chui CH, Lau FY, Chan PKS, Ng MHL, et al. ABO blood group and susceptibility to severe acute respiratory syndrome. Vol. 293, *JAMA*. United States; 2005. p. 1450–1.
19. Zhao J, Yang Y, Huang H, Li D, Gu D, Lu X, et al. Relationship between the ABO Blood Group and the COVID-19 Susceptibility. *medRxiv*. 2020;2020.03.11.20031096.
20. Parasher A. COVID-19: Current understanding of its pathophysiology, clinical presentation and treatment. *Postgrad Med J*. 2021;97(1147):312–20.
21. Wrapp D, Wang N, Corbett KS, Goldsmith JA, Hsieh C-L, Abiona O, et al. Cryo-EM structure of the 2019-nCoV spike in the prefusion conformation. *Science*. 2020;367(6483):1260–3.
22. Gui M, Song W, Zhou H, Xu J, Chen S, Xiang Y, et al. Cryo-electron microscopy structures of the SARS-CoV spike glycoprotein reveal a prerequisite conformational state for receptor binding. *Cell Res*. 2017;27(1):119–29.
23. Tian X, Li C, Huang A, Xia S, Lu S, Shi Z, et al. Potent binding of 2019 novel coronavirus spike protein by a SARS coronavirus-specific human monoclonal antibody. Vol. 9, *Emerging microbes & infections*. 2020. p. 382–5.
24. Zhou P, Yang X-L, Wang X-G, Hu B, Zhang L, Zhang W, et al. A pneumonia outbreak associated with a new coronavirus of probable bat origin. *Nature*. 2020;579(7798):270–3.
25. Xiong T-Y, Redwood S, Prendergast B, Chen M. Coronaviruses and the cardiovascular system: acute and long-term implications. *Eur Heart J*.

- 2020;41(19):1798–800.
26. Somekh E, Gleyzer A, Heller E, Lopian M, Kashani-Ligumski L, Czeiger S, et al. The Role of Children in the Dynamics of Intra Family Coronavirus 2019 Spread in Densely Populated Area. *Pediatr Infect Dis J.* 2020;39(8):e202–4.
  27. Laws RL, Chancey RJ, Rabold EM, Chu VT, Lewis NM, Fajans M, et al. Symptoms and Transmission of SARS-CoV-2 Among Children - Utah and Wisconsin, March-May 2020. *Pediatrics.* 2021;147(1).
  28. Rosenberg ES, Dufort EM, Blog DS, Hall EW, Hofer D, Backenson BP, et al. COVID-19 Testing, Epidemic Features, Hospital Outcomes, and Household Prevalence, New York State-March 2020. *Clin Infect Dis.* 2020;71(8):1953–9.
  29. Bi Q, Wu Y, Mei S, Ye C, Zou X, Zhang Z, et al. Epidemiology and transmission of COVID-19 in 391 cases and 1286 of their close contacts in Shenzhen, China: a retrospective cohort study. *Lancet Infect Dis.* 2020;20(8):911–9.
  30. Li W, Zhang B, Lu J, Liu S, Chang Z, Peng C, et al. Characteristics of Household Transmission of COVID-19. *Clin Infect Dis.* 2020;71(8):1943–6.
  31. Grijalva CG, Rolfes MA, Zhu Y, McLean HQ, Hanson KE, Belongia EA, et al. Transmission of SARS-COV-2 Infections in Households - Tennessee and Wisconsin, April-September 2020. *MMWR Morb Mortal Wkly Rep.* 2020;69(44):1631–4.
  32. Madewell ZJ, Yang Y, Longini IMJ, Halloran ME, Dean NE. Factors Associated With Household Transmission of SARS-CoV-2: An Updated Systematic Review and Meta-analysis. *JAMA Netw open.* 2021;4(8):e2122240.
  33. Somekh I, Sharabi A, Dory Y, Simões EAF, Somekh E. Intrafamilial Spread

- and Altered Symptomatology of SARS-CoV-2, During Predominant Circulation of Lineage B.1.1.7 Variant in Israel. *Pediatr Infect Dis J*. 2021;40(8):e310–1.
34. Hobbs C V, Martin LM, Kim SS, Kirmse BM, Haynie L, McGraw S, et al. Factors Associated with Positive SARS-CoV-2 Test Results in Outpatient Health Facilities and Emergency Departments Among Children and Adolescents Aged <18 Years - Mississippi, September-November 2020. *MMWR Morb Mortal Wkly Rep*. 2020;69(50):1925–9.
35. Triggler CR, Bansal D, Ding H, Islam MM, Farag EABA, Hadi HA, et al. A Comprehensive Review of Viral Characteristics, Transmission, Pathophysiology, Immune Response, and Management of SARS-CoV-2 and COVID-19 as a Basis for Controlling the Pandemic. *Front Immunol*. 2021; 12:1–15.
36. Brown NE, Bryant-Genevier J, Bandy U, Browning CA, Berns AL, Dott M, et al. Antibody Responses after Classroom Exposure to Teacher with Coronavirus Disease, March 2020. Vol. 26, *Emerging infectious diseases*. 2020. p. 2263–5.
37. Hains DS, Schwaderer AL, Carroll AE, Starr MC, Wilson AC, Amanat F, et al. Asymptomatic Seroconversion of Immunoglobulins to SARS-CoV-2 in a Pediatric Dialysis Unit. *JAMA*. 2020;323(23):2424–5.
38. Krass P, Zimbrick-Rogers C, Iheagwara C, Ford CA, Calderoni M. COVID-19 Outbreak Among Adolescents at an Inpatient Behavioral Health Hospital. *J Adolesc Health*. 2020;67(4):612–4.
39. Macartney K, Quinn HE, Pillsbury AJ, Koirala A, Deng L, Winkler N, et al. Transmission of SARS-CoV-2 in Australian educational settings: a prospective cohort study. *Lancet Child Adolesc Heal*. 2020;4(11):807–16.

40. Lopez AS, Hill M, Antezano J, Vilven D, Rutner T, Bogdanow L, et al. Transmission Dynamics of COVID-19 Outbreaks Associated with Child Care Facilities - Salt Lake City, Utah, April-July 2020. *MMWR Morb Mortal Wkly Rep.* 2020;69(37):1319–23.
41. Wong J, Jamaludin SA, Alikhan MF, Chaw L. Asymptomatic transmission of SARS-CoV-2 and implications for mass gatherings. *Vol. 14, Influenza and other respiratory viruses.* 2020. p. 596–8.
42. Jung J, Hong MJ, Kim EO, Lee J, Kim M-N, Kim S-H. Investigation of a nosocomial outbreak of coronavirus disease 2019 in a paediatric ward in South Korea: successful control by early detection and extensive contact tracing with testing. *Vol. 26, Clinical microbiology and infection: the official publication of the European Society of Clinical Microbiology and Infectious Diseases.* 2020. p. 1574–5.
43. Qiu H, Wu J, Hong L, Luo Y, Song Q, Chen D. Clinical and epidemiological features of 36 children with coronavirus disease 2019 (COVID-19) in Zhejiang, China: an observational cohort study. *Lancet Infect Dis.* 2020;20(6):689–96.
44. Huff H V, Singh A. Asymptomatic Transmission During the Coronavirus Disease 2019 Pandemic and Implications for Public Health Strategies. *Clin Infect Dis.* 2020;71(10):2752–6.
45. Kelvin AA, Halperin S. COVID-19 in children: the link in the transmission chain. *Lancet Infect Dis.* 2020;20(6):633–4.
46. Meyerowitz-Katz G, Merone L. A systematic review and meta-analysis of published research data on COVID-19 infection fatality rates. *Int J Infect Dis.* 2020; 101:138–48.
47. Levin AT, Hanage WP, Owusu-Boaitey N, Cochran KB, Walsh SP,

- Meyerowitz-Katz G. Assessing the age specificity of infection fatality rates for COVID-19: systematic review, meta-analysis, and public policy implications. *Eur J Epidemiol.* 2020;35(12):1123–38.
48. Zhou F, Yu T, Du R, Fan G, Liu Y, Liu Z, et al. Clinical course and risk factors for mortality of adult inpatients with COVID-19 in Wuhan, China: a retrospective cohort study. *Lancet (London, England).* 2020;395(10229):1054–62.
49. Xie J, Tong Z, Guan X, Du B, Qiu H. Clinical characteristics of patients who died of coronavirus disease 2019 in China. *JAMA Netw open.* 2020;3(4):e205619–e205619.
50. Zhang Y, Cao W, Jiang W, Xiao M, Li Y, Tang N, et al. Profile of natural anticoagulant, coagulant factor and anti-phospholipid antibody in critically ill COVID-19 patients. *J Thromb Thrombolysis.* 2020;50(3):580–6.
51. Fu J, Kong J, Wang W, Wu M, Yao L, Wang Z, et al. The clinical implication of dynamic neutrophil to lymphocyte ratio and D-dimer in COVID-19: A retrospective study in Suzhou China. *Thromb Res.* 2020; 192:3–8.
52. Liao D, Zhou F, Luo L, Xu M, Wang H, Xia J, et al. Haematological characteristics and risk factors in the classification and prognosis evaluation of COVID-19: a retrospective cohort study. *Lancet Haematol.* 2020 Sep;7(9): e671–8.
53. Tang N, Li D, Wang X, Sun Z. Abnormal coagulation parameters are associated with poor prognosis in patients with novel coronavirus pneumonia. *J Thromb Haemost.* 2020;18(4):844–7.
54. Garcia-Olivé I, Sintes H, Radua J, Abad Capa J, Rosell A. D-dimer in patients infected with COVID-19 and suspected pulmonary embolism. *Respir Med.*

- 2020 Aug; 169:106023.
55. Ranucci M, Sitzia C, Baryshnikova E, Di Dedda U, Cardani R, Martelli F, et al. Covid-19-Associated Coagulopathy: Biomarkers of Thrombin Generation and Fibrinolysis Leading the Outcome. *J Clin Med.* 2020;9(11):3487–9.
  56. Al-Samkari H, Karp Leaf RS, Dzik WH, Carlson JCT, Fogerty AE, Waheed A, et al. COVID-19 and coagulation: bleeding and thrombotic manifestations of SARS-CoV-2 infection. *Blood.* 2020 Jul;136(4):489–500.
  57. Luo H, You C, Lu S, Fu Y. Characteristics of coagulation alteration in patients with COVID-19. *Ann Hematol.* 2021;100(1):45–52.
  58. Hosseini SF, Behnam-Roudsari S, Alavinia G, Emami A, Toghyani A, Moradi S, et al. Diagnostic and prognostic value of Sepsis-Induced coagulopathy and International Society on Thrombosis and Hemostasis scoring systems in COVID-19-associated disseminated intravascular coagulopathy. *J Res Med Sci Off J Isfahan Univ Med Sci.* 2021;26:102.
  59. Bauer W, Galtung N, Neuwinger N, Kaufner L, Langer E, Somasundaram R, et al. A Matter of Caution: Coagulation Parameters in COVID-19 Do Not Differ from Patients with Ruled-Out SARS-CoV-2 Infection in the Emergency Department. *TH open companion J to Thromb Haemost.* 2021 Jan;5(1):e43–55.
  60. Xu W, Fei L, Huang CL, Li WX, Xie XD, Li Q, et al. Dynamic changes in coagulation parameters and correlation with disease severity and mortality in patients with COVID-19. *Aging (Albany NY).* 2021 May;13(10):13393–404.
  61. Li XY, Du B, Wang YS, Kang HYJ, Wang F, Sun B, et al. The keypoints in treatment of the critical coronavirus disease 2019 patient (2). *Zhonghua jie he he hu xi za zhi= Zhonghua jiehe he huxi zazhi= Chinese J Tuberc Respir Dis.* 2020;43(4):277–81.

62. Tang N, Bai H, Chen X, Gong J, Li D, Sun Z. Anticoagulant treatment is associated with decreased mortality in severe coronavirus disease 2019 patients with coagulopathy. *J Thromb Haemost.* 2020;18(5):1094–9.
63. Yu H-H, Qin C, Chen M, Wang W, Tian D-S. D-dimer level is associated with the severity of COVID-19. *Thromb Res.* 2020 Nov; 195:219–25.
64. Liu Y, Gao W, Guo W, Guo Y, Shi M, Dong G, et al. Prominent coagulation disorder is closely related to inflammatory response and could be as a prognostic indicator for ICU patients with COVID-19. *J Thromb Thrombolysis.* 2020;50(4):825–32.
65. Bao C, Tao X, Cui W, Yi B, Pan T, Young KH, et al. SARS-CoV-2 induced thrombocytopenia as an important biomarker significantly correlated with abnormal coagulation function, increased intravascular blood clot risk and mortality in COVID-19 patients. *Exp Hematol Oncol.* 2020;9(1):1–8.

## **ANNEXURES - I- INFORMED CONSENT**

Dear Mr./Mrs./Dr. \_\_\_\_\_, you are kindly requested to enroll yourself in a research study titled: A STUDY OF COAGULATION PROFILE DYNAMICS IN PATIENTS INFECTED WITH COVID-19 ADMITTED TO KLES PRABHAKAR KORE HOSPITAL AND MEDICAL RESEARCH CENTRE. -A CROSS SECTIONAL STUDY DONE IN KLE'S DR.PRABHAKAR KORE HOSPITAL AND MEDICAL RESEARCH CENTRE, BELAGAVI being conducted by

REG NO: BG0120021 a post graduate student in M.D. General Medicine and the study will be carried out under the direct supervision and guidance of

Dr. \_\_\_\_\_

Professor, Department of General Medicine,

Registrar KAHER

Jawaharlal Nehru Medical College, Belagavi.

You have been requested to participate in this as you fit into the laid out criteria for a study 'subject'/ participant.

Your participation in study is voluntary. During the study you will be asked some questions and you are supposed to answer to the best of your knowledge. Your decision whether or not to participate in the study will not affect your treatment in any form. If you decide to participate you are free to withdraw at any time.

TITLE OF THE STUDY:

A STUDY OF COAGULATION PROFILE DYNAMICS IN PATIENTS INFECTED WITH COVID-19 ADMITTED TO KLES PRABHAKAR KORE HOSPITAL AND MEDICAL RESEARCH CENTRE.

A CROSS SECTIONAL STUDY DONE IN KLE'S DR.PRABHAKAR KORE HOSPITAL AND MEDICAL RESEARCH CENTRE, BELAGAVI

PURPOSE OF THE STUDY: To Study the dynamic changes in the coagulation profile and co relate it to the outcome of patients affected by COVID 19.

PROCEDURES INVOLVED: If you agree to enroll yourself in my study, you will be interviewed regarding your present, past and family history then you will be clinically examined in detail and investigated accordingly.

Then you will be subjected to a few blood investigations done routinely namely complete blood count, random blood glucose, blood urea, Sr.electrolyte, liver function test, urine routine and microscopy And coagulation profile which includes D-dimer, Serum Fibrinogen, PT INR,APTT along with other inflammatory markers like LDH,Sr.ferritin,IL-6,hs-CRP.

RISKS AND BENEFITS: There are no potential risks involved in this study.

BENEFITS OF TAKING PART IN THIS RESEARCH: By taking part in this study, the coagulation profile and prognosis of every patient will be studied and the therapeutic management will hence be refined which will benefit the forthcoming patients infected with covid-19.

VOLUNTARY PARTICIPATION / WITHDRAWAL FROM THE STUDY: Taking part in the study is voluntary. You may choose not to enroll yourself in this study and may choose to leave the study anytime in between.

ALTERNATIVES: Your decision regarding participation in study will not change present or future health care services offered to you at KLES Dr. Prabhakar Kore Hospital and Medical Research Centre, Belgaum. You would simply be excluded from the study if you wish to, and all your details shall be kept confidential and you will get the routine line of management.

**PRIVACY AND CONFIDENTIALITY:** All data collected or disclosed by you during the course of participation of study, will be kept fully confidential. If however during the course it becomes necessary for the progress of the course to disclose the identity, it would be done so only after your informed & written consent. The only people to know that you are a research subject are members of the research team. No information about you will be disclosed to other without your written permission except:

In emergency to protect your rights AND welfare.

If required by law.

**AUTHORIZATION TO PUBLISH RESULT:** The results of the study may be used to publish an article. When the results of research published or discussed, in a conference, no information will be displayed that would disclose your identity. Any information obtained in connection with this study and that can be identified with you will remain confidential.

**FINANCIAL INCENTIVES FOR PARTICIPATION:** No additional costs shall be incurred upon you for the purpose of this study. It is purely being done with the idea of research and all the cost of study will be borne by the investigator.

**COMPENSATION:** In the event that you become injured as a result of taking part in this study, treatment will be offered to you at KLES Dr. Prabhakar Kore Hospital and Medical Research Centre, Belgaum, or you will be given information about where to receive medical care. However, no reimbursement, compensation or free medical care will be given.

**QUESTIONS/CONTACT DETAILS:** You shall be free to contact the below mentioned name & addresses anytime during the study period for any clarification or help as you may desire for.

PRINCIPAL INVESTIGATOR:

REG NO: BG0120021

MD (Post Graduate Student), Department of General Medicine, Jawaharlal Nehru Medical College, Nehru Nagar, Belagavi 590010.

GUIDE:

Dr. \_\_\_\_\_

Department of General Medicine,  
Jawaharlal Nehru Medical College  
Nehru Nagar, Belagavi 590010.

Dr. Roopa M Bellad, M.D. D.C.H(PROFESSOR OF PAEDIATRICS Chairman,  
J.N.M.C Ethical Committee for Human Research,9448113403

**CONSENT FORM**

I voluntarily agree to take part in this study by signing below. I may withdraw at any time. I am not giving up any of my legal rights by signing this form. My signature below indicates that I have read this consent form, or it has been read to me, this consent form and have had all the questions answered.

Signature / Left Thumb print of the Participant or legally authorized representative

Participant's name :.....

Signature / Left thumb impression: .....

of the participant

Name of the legally authorized :.....

representative / guardian

Signature / Left thumb impression :.....

Witness' name :.....

Signature / Left thumb impression :.....

Investigator's name and signature :.....

Date:

Place:

**ANNEXURES – II - PROFORMA**

**IP No** :

**Date of Admission** :

**Age** :

**Sex** :

**Clinical Parameters At Admission**

*Symptoms:*

Tick the applicable Box					
Cough		Fever		Other Symptoms	
Myalgia		Breathlessness			
Time to hospital from symptom onset (Days)					

<b>Oxygenation Parameters at admission</b>	
Admission Respiratory rate (RR/min)	
Admission SPO2(%)	
Admission O2 modality (RBM,HFO,NIV, Ventilator)	
Admission O2 rate (in Litres per minute)	
Admission O2 FiO2 (%)	
Admission PO2 (Look at admission ABG)	

**Co-morbidities**

DM		HTN		IHD		CKD		CLD		CVA		Malignancy	
Others													

**Lab Parameters At Admission**

Hb		TLC		Lymphocyte%		Platelets	
Ferritin		LDH		hsCRP		PTINR	
d-Dimer		CRP		S Fibrinogen		Urea	
Creatinine		Bilirubin		SGOT		SGPT	
RBS		HbA1c				APTT	

<b>CT Severity score (Out of 25)</b>	
--------------------------------------	--

<b>Maximum OXYGENATION Support required during entire stay</b>	
Mode of O2 (RBM,HFO,NIV, Ventilator)	
FiO2%	
SpO2 %	
PO2% (Look at ABG)	

<b>Worst Lab Parameters during entire hospital stay</b>	
Lowest Absolute Lymphocyte count	
Maximum TLC	
Maximum IL-6	
Maximum D-dimer APTT PT INR S.Fibrinogen	
Maximum Creatinine	
Maximum SGOT	
Maximum SGOT	

**TREATMENT**

Remdesivir (No. of Doses)	
Tocilizumab (No. of Doses)	
LMWH (Max Dose)	
Un-fractionated Heparin (Max Dose)	
Type of Steroid used (methyle pred or dexamethasone)	
Max dose of steroid per day used	
Thymosin 1 Alpha (No of doses)	
Convalescent Plasma (Cycles)	
PO2/FiO2 at the time of convalescent plasma	
Mode of O2 at the time of plasma therapy	

---

---

Cytosob (Cycles)	
Plasma Exchange (Cycles)	
Alteplase (Dose)	
Alteplase (tPA)(Number of times Given)	
Pre tPA PO2/FiO2	
Post tPA pO2/FiO2	
Pre tPA Fibrinogen	
Post tPA Fibrinogen	
Pre tPA d-Dimer	
Post tPA D-dimer	

**OUTCOME**

Outcome	Tick	Date
Improved and discharged		
Died		
Worsened and went AMA		
Improved and went AMA		

0					CLINICAL PARAMETERS AT Admission													INFAMMATORY PARAMETERS AT Admission													Other Labs at Admission					
Sl. No.	IP No	Date of Admission	AGE	Sex	Symptoms (C=cough; F=Fever; B=breathlessness; M=Myalgia, Loss of smell=LS; Loss of taste=L,T)	Any other symptoms V: Vomiting, LO=loose stools, W: weakness, AP: Abd pain, AD: Abd distn, J: Jaundice; Hematuria, LAPP: Decreased Appetite, SoT: Sore throat, CP: Chest pain, AllSen: Altered sensorium, Ed:Edema, LoOP: Low urine, LWK: limb weakness, Hemoptysis	Time to hospital from symptom onset (Days)	Admission Respiratory rate (RR/min)	Admission SPO2	Admission O2 modality (RBM,HFO, NIV, Ventilator)	Admission O2 rate (L/min)	Admission FIO2	qSOFA	Co-morbidities	Vaccination status	CT severity score(-/25)	Hb 1	TLC1	NEUTROPHIL %1	Lymphocyte % 1	Platelets	Ferritin	FIBRINOGEN	LDH	HOMOCYSTEINE	hsCRP	IL-6	d-Dimer	d-Dimer	INR	APTT	ISTH SCORE	Urea	Creatinine	Bilirubin	SGOT
1	1032968	29.12.20	57	M	FB		7	28	96	RBM	10L/MIN	NA	1	DM	0	12	10.6	11	92	5	199	151	351	452	33	50	50	237		1.66	1.08	<50	24	0.9	0.7	30
2	1029460	28.11.20	59	M	FBC		6	26	98	RBM	8L/MIN	NA	1	HTN,IHD	0	16	11.8	10	78	14	262	119	392	232	19.5	53.7	17.06	1008	1624	1.28	0.9	<50	17	0.83	0.39	49
3	1029718	1.12.2020	25	M	CB		3	22	97	RBM	5L/MIN	NA	2	HTN	0	22	10.5	12	85	12	57	193	640	439	20.5	6.3	49	1381	514	2.42	0.81	<50	87	1.03	0.5	62
4	1049200	15.4.2021	40	M	CFM		4	24	94	O2 MASK	2L/MIN	NA	3	HTN,CKD	0	18	17.9	11.2	84	13	248	408.3	104	854	11.51	142	600	323	1352	1.22	0.97	>50	25	1.02	1.13	23
5	1050175	.21.4.21	23	M	CFMB		10	34	95	HFO	40L/MIN	NA	3	DM	0	24	15.6	5.2	46	44	208	143.6	237	223	12	4.2	13.6	166		1.1	1	<50	15	0.85	0.27	26
6	1049472	16.4.2021	51	M	B		4	28	94	O2 MASK	5L/MIN	NA	3	DM,HTN	0	22	13.6	14	82	10	561	235.9	436	634	7	89	13	5000	317	1.1	1	>50	44	0.61	0.32	18
7	1050328	23.4.21	46	M	FMB		7	28	94	RBM	8L/MIN	NA	3	DM, HTN	2	18	13.8	3	83	15	112	2086	108	255	25	57.9	17	474		1.7	1.2	<50	29	1.01	0.79	80
8	1050175	21.4.21	23	M	FB		3	26	92	RBM	15L/MIN	NA	2	DM	2	8	15.6	5.2	46	44	209	143	237	223	12.4	4.2	13.6	166		1.2	1.1	>50	15	0.85	0.27	26
9	1049472	16.4.21	51	M	M		3	24	90	O2 MASK	5L/MIN	NA	2	DM,CKD	2	19	13.6	5.2	46	44	208	143.6	237	223	12.43	4.2	13.69	166	202	1	1.08	<50	44	0.61	0.32	18
10	1051902	30.5.21	88	M	CB		5	28	88	RBM	10L/MIN	NA	2	IHD	2	24	9.5	6.5	86	13	152	8957	454	468	18	36	16	990		1.2	1.1	>50	41	1.32	0.39	103
11	1051907	10.5.21	65	F	CFB		3	28	90	O2 MASK	5L/MIN	NA	2	DM	2	22	9.2	10.3	92	6	283	325	778	545	16.7	307	9	872		1.18	0.93	<50	62	1.62	0.43	23
12	1051952	10.5.21	36	M	CFM		5	29	93	RBM	10L/MIN	NA	2	HTN/DM	1	25	11.5	14	88	6	287	1402	493	622	13	76	84	633	1	1	0.9	<50	42	0.94	2.2	33
13	1051195	1.5.21	42	M	BF		2	42	37	NIV		100	1	DM, HTN	1	24	13.7	5	75	18	159	748	399	352	14.6	195	64	160		1.12	1.06	<50	38	1.3	0.5	28
14	1051698	7.5.21	60	M	CB		3	26	80	NIV		100	1	HTN/OBESITY	1	22	15.6	8.7	87	12	90	2000	1103	416	13	156	48	758		1.27	0.89	<50	54	1.01	0.5	70
15	1051068	10.5.21	45	M	FB	V, LO	3	45	35	NIV		100	1	DM,HTN,CKD	1	25	13.3	5.1	83	12	150	3235	460	481	5.9	66	63	876		1.05	0.86	<50	31	0.6	0.2	58
16	1051728	10.5.21	40	M	CMB		2	26	92	O2 MASK	6L/MIN		1	NONE	1	15	13.7	14	87	7	170	605	283	961	8.6	5.6	21	1437	1541	1.16	0.6	<50	91	0.73	0.8	37
17	1051693	10.5.21	45	F	CF		5	28	100	NIV		100	3	DM,HTN,IHD	1	18	12.6	15.4	71	14	183	263	268	667	16.7	79	21	4682		1.16	0.6	>50	41	0.76	0.2	41
18	1051196	7.5.21	87	M	B	CP	4	22	96	O2 MASK	5L/MIN	NA	1	HTN,IHD	1	20	13.3	11.7	93	2	224	475	1149	296	19	759	654	690		1.3	1.5	>50	74	1.17	0.5	107
19	1051064	4.5.21	87	M	CFB		2	20	88	RBM	4L/MIN	NA	1	DM,HTN	1	22	13.6	5.5	82	13	147	569	1149	339	139	6	650	690		1.2	1	>50	46	1.45	0.57	26
20	1051262	10.5.21	60	M	FB	V	4	22	100	NIV		100	1	IHD	1	22	15.6	9.1	75	16	260	794	552	552	12.5	13.3	1.08	555		1.08	0.85	<50	20	0.6	0.42	20
21	1051304	10.5.21	62	M	CBM		3	30	88	RBM	15L/MIN	NA	1	HTN	0	18	15	6	84	10	160	1285	708	521	8.4	130	40	1057		1.28	0.9	>50	63	0.7	0.8	77
22	1051738	7.5.21	73	M	CFB		3	26	87	RBM	15L/MIN	NA	2	DM,HTN	0	24	11.3	6.3	74	21	164	191	191	323	11	117	7	891		1.19	0.7	<50	66	0.9	0.4	14
23	1051798	10.5.21	33	F	FB		7	22	94	RBM	15L/MIN	NA	3	DM,HTN	0	20	11.8	10	83	12	445	131	328	426	14.2	7	5.79	396		0.9	0.7	<50	35	1.25	0.5	15
24	1051477	4.5.21	75	M	CFB		4	36	82	RBM	10L/MIN	NA	3	ASTHMA	0	23	12.2	3.8	87	6	113	809	467	876	8.9	135	32	1352		0.9	0.7	<50	108	1.68	0.6	36
25	1051255	2.5.21	35	F	CB		4	34	70	NIV		100	3		0	6	8.7	21.6	93	5	143	93	470	1184	6.2	135	86.3	1232		1.7	0.05	>50	64	0.8	0.49	32
26	1051522	5.5.21	33	M	CFB		3	28	64	NIV		100	3	DM, HTN	0	24	14	13.1	67	9	160	749	934	555	7.4	274	45	1440	686	1.16	0.93	>50	45	1.71	0.62	41
27	1051068	10.5.21	45	M	CB		3	22	93	O2 MASK	15L/MIN	NA	1	DM	0	22	13.3	5.1	83	12	150	156	960	678	5.9	54	39	836		1.05	0.8	>50	26	0.48	1.06	70
28	1051728	7.5.21	40	M	CBF		4	24	93	RBM	8L/MIN	NA	1	DM,AKI	0	13	13.4	14	87	7	170	605	283	961	12	5.6	21	1437		1.16	0.66	>50	37	0.37	0.8	27
29	1054327	3.6.21	41	F	CFMB	LoOP, Hematu	2	18	95	NIV		100	2	DM	0	18	12.5	10.3	89	9	459000	626	333	773	13	431	254	1482		1.1	1.3	<50	18	1.1	0.3	42
30	1054254	3.6.21	88	M	CFB		3	28	87	RBM	15L/MIN	70	1	DM	1	22	14.8	12.5	95	3	150	1988	389	638	10	164.8	235	511		1.29	1.28	>50	86	1.64	0.33	68
31	1020598	3.6.21	49	M	FM		8	30	86	RBM	10L/MIN	NA	1	DM	1	16	14.1	4.7	81	16	147000	1349	433	826	5	96	35	937	7748	1.1	1.09	>50	34	1.01	0.5	55
32	1022740	3.6.21	74	M	B		1	32	70	NIV		70	3	DM,HTN	1	20	12.8	16.7	97	2	232000	499	345	3650	67	54	120	>5000		1.2	1.08	>50	139	3.52	1.2	7437
33	5783029	3.6.21	73	M	CFB		4	26	76	RBM	10L/MIN	NA	3	DM,HTN	0	22	10.8	14.1	89	6	1,84,000	1791	456	502	12	46	18.8	1205	273	1.1	1.02	>50	39	1.01	0.43	36
34	1023296	3.6.21	66	M	CFB		7	36	96	RBM	5L/MIN	NA	3	DM,HTN	0	24	14.3	2.6	75	20	169000	847.6	123	217	13	54	222	573		1.1	1.5	>50	23	0.78	0.78	50
35	1023272	3.6.21	52	M	CFB		2	35	66	RBM	15L/MIN	NA	3	HTN	0	22	13.9	10.9	76	21	140000	516.7	122	819	1	218.6	22.65	1217	5000	1	1.6	>50	52	1.02	0.68	83
36	1021459	3.6.21	56	M	M		4	32	84	RBM	15L/MIN	NA	1	DM	1	20	13	14.7	88	9	234000	923	122	842	12	140	23	765		1.9	1.7	<50	41	1.1	0.32	24
37	1021997	4.6.21	71	M	CFB		7	30	93	NIV		100	1	IHD	1	14	13.5	11.3	95	4	132000	271	128	854	14	14	19.57	660		1	1.8	<50	72	0.81	0.54	43
38	1021081	4.6.21	66	F	CFB		2	26	94	RBM	10L/MIN	NA	1	DM	1	20	12	9.7	86	11	2610000	82.91	178	280	12	28	164	328		1	1.8	<50	32	0.85	0.33	23
39	1021843	4.6.21	84	M	B		11	24	96	O2 MASK	8L/MIN	NA	1	DM,HTN,IHD	1	16	13.3	13	90	4	236000	385.3	189	489	10	29	247	898		1.4	1.2	<50	37	1.06	1.03	31
40	1021421	8.6.21	64	M	CB		5	24	94	NIV		100	2	DM,HTN	2	22	16.7	29.7	95	2	208000	1507	333	502	15	123	25	7028		1.4	1.1	>50	140	1.09	1.22	68
41	1021050	8.6.21	32	M	CFB		20	36	86	RBM	15L/MIN	NA	2	NONE	2	18	15.7	18	71	19	419000	759	555	579	16	158	29	871		1.2	1.1	>50	36	1.01	0.93	51
42	1022051	8.6.21	65	M	CFB		5	28	56	NIV		100	2	DM,HTN	0	22	10.1	17	93	5	107000	2000	567													

SGPT	RBS	PCT	HbA1c	70										TREATMENT										OUTCOME																	
				Mode of O2 (RBM,HFO,NIV, Ventilator) 1	2	3	FIO2 1	2	3	SpO2 1	2	3	PAO2 ABG-1	2	3	Remdesivir (No. of Doses)	Tocilizumab (No. of Doses)	LMWH (Max Dose)	TOTAL DOSES	NOAC	Unfractionated Heparin (Max Dose)	Type of Steroid used (MethylePred or dexa)	Max dose per day used	TOTAL NO: OF DOSES	Alteplase (Doses)	Alteplase (Number of times Given)	TOFACITINIB	IVERMECTIN	ZINC+ VIT C	THROMBOTIC COMPLICATIONS ML,PE,DVT,CVA,DIC	CRITICAL ILLNESS ARDS SEPSIS AKI	ICU/HDU /WARD	HOSPITAL STAY	Improved and discharged	EXPIRED	Worsend and AMA	Improved and AMA	Outcome Date			
19	111	0.02		O2 MASK	NRBM	NRBM	10L	60	60	92	94	94	82	-	-	2		40MG	4		Methyl Pred	40MG	6		4								WARD	4	YES						
20	220	0.57		NIV	NIV	NIV	60	80		97%	97%	90%	77%	89%		9	1	40mg	8		MethylPred	40mg	8															AMA			
44	173	2.48		RBM	HFO	NIV	12L/MIN	60L/MIN	100%	92%	93%	91%	58%	77%	77	9	2	40MG	12		Methyl pred	40mg	12																		
27	120	0.01		RBM	NIV	NIV	5L/MIN	100%	100%	96%	94	96	77	72	69	6	1	40MG	4		Methyl pred	80mg	16														YES				
23	290	0.05		HFO	NIV	NIV	40L/MIN	80L/MIN	100%	92%	99%	90%	67%	69%	76	7	1	40MG	6		Methy pred	40mg	8															AMA			
28	233	0.05		HFO	NIV	NIV	60L/MIN	100%	100	80	80	86	68	69	77	7	2	40mg	8		Ivepred	80mg	16														YES				
66	133	0.01		NIV	INTUBATION		100%	100%	100	98	99%	90	45	44	67	8	2	80MG	10		MethylP	1G	3	30MG	1																
23	111	1		HFO	NIV	HFO	60L	100%	60L/min	94%	97%	93%	77%	78%	89	8	1	40MG	12		Methyl Pred	1 g	3	30	2																
28	322	0.05		HFO	HFO	INTUBATION	60L/MIN	70L/MIN	100%	97	95%	91%	66%	67%	77	7	1	40MG	8		Methylpred	500mg	3															YES			
13	44	3.4		NRBM	INTUBATION		10L	100%	100%	88%	67	87	56	68	66	8	3	40MG	8		MethylP	80MG	9															AMA			
13	133	1.33	10.1	HFO	NIV	HFO	60L/MIN	100%	60	96	90%	90	88	89	90	8	1	80MG	12		Methyl Pred	80mg	16																		
22	222	0.09		HFO	NIV	NIV	50L	60	80	75%	98%	96%	77%	77%	78	9	2	80mg	14		MethylP	500MG	5																		
16	123	0.29		NIV	NIV	HFO	100	100	80	95	92	90	89	-	-	7		80MG	8		MethylP	80	9																		
42	83	0.83		NIV	NIV	HFO	100	100	80	98	92%	90	88	-	-	6		40MG	9		DEXA	12MG	9																		
39	134	0.1	6.5	INTUBATED			100			92						1		80MG	1		METHYPRED	500MG	1		1												YES				
43	145	0.05	8.2	NIV	NIV	HFO	100	80	80	94	90	90	90	-	-	10	2	40mg	8		Methylpred	1gm	STAT+MD																		
18	122	0.05		INTUBATED			100			98						4		40MG	4		METHYPRED	120MG	4																		
329	300	0.05		HFO	NIV	INTUBATED	80L/MIN	100%	100%	92	96%	98%	90%	77%	89	7	3	40MG	10		METHYPRED	1000GM	5																		
305	199	0.23	10.7	HFO	HFO	HFO	60L/MIN	60	50	86	90	90	77	80	87	3	3	40MG	4		DEXA	12MG	9																AMA		
43	144	0.33		NIV	INTUBATED		100			68						9	3	40mg	3		Methylpred	1g	3														YES				
44	333	0.5		NIV	HFO	HFO	100	80	80	89	86	89	67	87	88	5	1	40 MG	7		METHYPRED	1000MG	2																		
28	145	0.03	9.8	NIV	INTUBATED		100	100		86	91%	77	76	66	56	5	2	40mg	4		Methylpred	500mg	5																		
20	233	0.03		RBM	HFO	NIV	15L/MIN	60L	100	89	92	89	78	80	87	4	1	40MG	5		Methylpred	500mg	2																		
26	68	0.7	7.6	NIV	HFO	HFO	100	60	70	78	84	87	78	88	88	5	3	40MG	5		Methyl Pred	40MG	5																		
65	111	0.05		NIV	NRBM	NRBM	100	60	60	70	82	82	82	88	89	6	1	80mg	6		Methylpred	80	6																		
55	123	0.17		HFO	RBM	NIV	30L/MIN	15L/MIN	100	93	94	95	90	90	77	6	2	80	6		Methyl Pred	1G	3	50MG	1																
80	133	0.02		NIV	NIV	NRBM	100	100	60	92	95	90	90	88	87	6	1	40	6		Methylpred	1000 mg	1																		
43	145	0.05		RBM	RBM	HFO	10L/MIN	15L/MIN	100	94	97	88%	66%	88%	90	5	1	40MG	6		Methylpred	500	2																		
10	333	16.21		NIV	NRBM	NRBM	100	60	60	90	90	90	90	84	83	6	2	40	7		Methylpred	40	9																AMA		
34	222	1.2		NIV	NIV	HFO	100	70	70	90	92	92	88	86	84	6	1	40	6		Methylpred	40	16																		
15	123	1		O2 MASK	O2	O2	10L	10L	8L	93	92	90	90	96	94	4	2	80	8		Methylpred	500	1+MD		6																
21	123	0.03		NIV	NIV	NIV	100	100	100	97	90	86	88	82	78	5	2	40	8		Methylpred	1g	1+MD																	AMA	
21	145	0.04		RBM	NIV	INTUBATED	10L/MIN	100	100	88	92	96	77	77	80	5	3	40	3		Methylpred	80MG	4	30MG	1	4													YES		
18	134	0.1		NIV	NIV	INTUBATED	100	100	100	92	84	80	66	62	64	11	2	80	6		Methylpred	500	1+MD																		
20	134	0.12		NIV	INTUBATED		100	100		76	84	77	78	66	62	5	2	80	4		Methylpred	1g	4	50mg	1																
29	146	0.23		RBM	NIV	NRBM	60	90	60	84	92	90	-	-	-	2	-	80	5		Methylpred	1g	1+MD		5																
19	500	0.44	6.9	NIV	NIV	NIV	100	80	100	93	96	84	89	-	-	5	2	80	4		Methylpred	1g	1+MD		5																
21	234	0.56		RBM	RBM	HFO	10L/MIN	15L/MIN	60L/MIN	94	89	70	80	-	-	4	1	80	6		Methylpred	500	1+MD		6																
14	67	0.65		NIV	NIV	HFO	100	100	60L/min	94	94	89	88	-	-	5		40	6		Methylpred	1g	1+MD		7																
28	77	0.78		NIV	INTUBATED		100	100		94	96	77	77	71	70	2	1	80	3		Methylpred	1g	3																		
86	567	1.2		HFO	NIV	HFO	60	90	60	80	90%	90	90	90	90	3	1	40	6		Methylpred	1g	1+MD		6																
35	420	1.7		NIV	HFO	HFO	100	70	70	94	89	89	90	90	-	5	1	60	5		Dexa	12MG	6																		
32	124	1		NIV	NIV	INTUBATED	100	100	100	92	60	70	66	66	61	7	1	40	6		Methylpred	1g	4																		
33	189	1.02		HFO	INTUBATED		60	100	90	90	77	77	71	56	-	3	2	80	3		Methylpred	120	6																		
26	200	1.03		RBM	RBM	NIV	15L/MIN	15L/MIN		100	100	92	88	88	89	4	2	40	6		DEXA	12	9																		
13	101	1.03		NIV	NIV	NIV	60	100	50	94	96	92	78	88	-	9	1	40	8		Methylpred	120	5																		
30	266	1.8		HFO	NIV	NIV	60	100	100	91	92	88	77	-	-	2	1	40	3		Methylpred	1GM	2																	AMA	
36	123	5.6		NIV	NIV	NRBM	100	100	60	90	90	100	90	80	-	3	2	40	7		Methylpred	80	7																		
117	146	4.8		NIV	NIV	INTUBATED	100	100	100	92	90	88	87	80	-	3	2	120	8		Methylpred	1G	1+MD																		
47	177	2.33		INTUBATED			100			88						1		40	1		Dexa	12MG	STAT																		
14	189	12.1		HFO	NIV	INTUBATED	60	100	100	90	99	92	80	78	-	5	1	40	3		Methylpred	120	4																		

Sl. No.	IP No	Date of Admission	Age	Sex	Symptoms (Cough, Fever, Rash, Shortness of Breath, Loss of consciousness, Myalgia, Loss of smell, Loss of taste, etc.)	CLINICAL PARAMETERS AT Admission															Other Labs at Admission																		
						Any other symptoms	Time to hospital from symptom onset (Days)	Admission SPO2	Admission O2 modality (O2 mask, RBM/HFO, NIV, Ventilator)	Admission O2 rate (L/min)	Admission FIO2	qSOFA	Co-morbidities	Vaccination status	CT severity score(-25)	Ib 1	TLC1	NEUTROPHIL %1	Lymphocyte % 1	Platelets 1	Ferritin 1	Fibrinogen	LDH 1	Homocysteine	hsCRP (value)	IL-6 1	d-Dimer 1	d-Dimer 2	INR	APTT	ISTH SCORE DIC	Urea	Creatinine 1	Bilirubin	SGOT	SGPT	RBS	HBsAg	RCT
1	1026582	10-29-20	37	M	CF		5	26	85%	O2 MASK	5L/min	NA	2%	ASTHMA	0	9	15.1	9	65	26	202	148	282	573	19	6.4	44.58	193	1.09	0.9	<5	16	0.8	0.45	17	12	98	-	0.1
2	1026610	27-10-2020	80	M	CFMB		5	32	91%	RA	NA	3%	CKD	0	12	7.5	9.8	79	15	174	267	288	240	15	12	31.6	938	1.31	1.04	>5	126	4.2	0.35	20	10	128	9.1	0.2	
3	1026721	31-10-20	68	F	FB		4	18	96%	RA	NA	NA	0	NONE	0	15	10.7	11.7	83	12	262	287	286	5	13.7	1.5	389	1.01	0.86	>5	24	0.9	0.24	18	17	99	9.1	0.1	
4	1026613	29-10-2020	60	F	CFMB		2	22	88%	O2 MASK	5L	NA	0	DM	0	12	13.2	9.8	84	13	361	1115	334	388	5.4	148	9.97	408	1.2	0.9	>5	27	0.63	0.53	35	28	174	0.63	
5	1026783	1-11-20	25	M	FB		1	16	99%	RA	NA	NA	0	NONE	0	8	13.8	6.2	60	27	291	64	395	227	14.6	2.5	1.5	111	1.4	1.03	<5	20	0.9	0.7	18	17	144	0.01	
6	1053241	4-6-21	44	M	CFB		5	14	96%	O2 MASK	2L O2	NA	1	HTN	1	15	10.1	10.5	92	6	375	404	468	846	138	17.2	767	1.13	1.07	<5	28	0.85	0.93	44	40	480	0.03		
7	1054134	03-06-21	65	F	CB		2	22	90%	O2 MASK	6L O2	NA	1	DM/HTN	1	17	12.1	10.5	86	9	159	1625	362	496	57	9.1	1115	1.23	0.9	<5	79	1.08	0.4	38	11	168	0.08		
8	1053539	4-6-21	44	M	CFB		2	26	90%	O2 MASK	5L O2	NA	3	DM/HTN	0	12	16	9.2	88	5	170	1131	660	453	16.6	180	1731	1.26	0.9	>5	44	0.8	0.76	30	20	85	0.13		
9	1053531	4-6-21	69	M	CFB		4	18	95%	O2 MASK	4L O2	NA	2	DM/HTN	1	14	13.7	14.7	91	5	268	900	324	811	72.6	26	1184	1.39	0.96	>5	46	1.36	0.32	42	36	52	0.09		
10	1053540	4-6-21	37	M	CF		5	12	96%	RA	RA	NA	1	DM/HTN, HHD	1	18	13.3	19.3	86	11	286	367	474	997	18	7.15	311	1.28	0.76	<5	31	0.6	1.06	38	42	156	0.06		
11	1053200	4-6-21	39	M	C		3	20	96%	RA	RA	NA	0	NONE	2	9	13.5	24.8	92	3	403	591	423	490	7.9	1.5	218	1.31	0.71	<5	40	0.82	0.45	27	66	191	0.04		
12	1053204	4-6-21	62	M	CMF		1	24	85%	O2 MASK	15L	NA	1	DM/HTN	0	14	11.1	8.8	86	10	170	176	423	620	413	424	737	1.31	0.89	<5	22	0.82	0.48	32	10	227	0.38		
13	1053971	4-6-21	40	M	F	WEAKNESS	2	14	99%	RA	RA	NA	2	NONE	1	12	13.8	3.8	74	24	162	613	904	105	16.9	5000	1.11	0.9	>5	27	0.68	0.69	266	329	137	0.25			
14	1054049	4-6-21	85	M	FM	WEAKNESS	4	18	96%	RA	RA	NA	1	NONE	1	14	8.8	11.7	74	14	253	257	275	8.8	81.8	1.55	5000	1.46	0.85	>5	96	1.29	0.42	21	14	86	10.23		
15	1053667	4-6-21	55	M	CFB		3	26	92%	O2 MASK	3L O2	NA	3	NONE	1	15	13.9	19.6	96	2	368	1294	496	725	339	38.6	699	1.2	0.86	<5	44	0.68	0.68	40	41	108	0.06		
16	1054303	4-6-21	31	M	CFBM	WEAKNESS	4	22	89%	O2 MASK	5L O2	NA	2	DM/HTN, HHD	1	24	12.6	9.2	82	14	228	800	781	500	171	122	333	1.19	0.9	<5	23	0.86	0.36	51	36	198	0.16		
17	1052330	4-6-21	54	M	CFB		3	21	90%	O2 MASK	6L	NA	1	DM	1	22	13.3	10.8	87	10	214	1024	435	583	264	84	278	1.06	0.91	<5	38	1.05	0.89	50	48	112	6.3	0.08	
18	1052979	4-6-21	80	F	CM	WEAKNESS	2	14	99%	RA	RA	NA	1	DM	1	19	11.6	13.8	92	5	184	619	376	630	43.4	15.7	377	0.91	0.83	>5	97	1.95	0.17	39	12	124	0.25		
19	1053907	4-6-21	72	M	CFMB	DIARRHOEA	2	12	96%	RA	RA	NA	0	NONE	1	14	13.1	17.4	95	3	304	438	614	257	13.4	45.12	253	1.28	0.95	<5	88	1.65	0.63	27	27	204	0.15		
20	1054129	1-6-21	45	M	CFB		5	22	89%	O2 MASK	RA	NA	3	TB, ASTHMA	1	20	11.2	13	70	14	278	741	468	303	13.5	41	1317	1.34	0.9	>5	36	0.89	0.2	20	14	470	0.12		
21	1053872	2-6-21	43	M	CFMB	WEAKNESS	2	16	97%	RA	RA	NA	2	DM/HTN	2	14	12.7	8.0	16	21	211	733	417	229	16	1.5	131	1.04	0.85	<5	24	1.41	0.17	30	21	104	0.12		
22	1054030	2-6-21	38	M	CFMB		4	12	96%	RA	RA	NA	1	NONE	1	12	16.9	13.2	77	6	236	186	555	363	51.7	31.3	459	1.16	0.83	<5	34	0.7	0.76	28	49	102	0.05		
23	1053968	2-6-21	38	F	CFB		3	10	99%	RA	RA	NA	0	NONE	0	6	15.1	5.9	86	11	311	190	287	190	10.2	2.7	96	1.2	0.9	<5	18	0.69	0.33	25	19	100	0.65		
24	1053215	20-5-21	52	M	CFB		5	20	93%	RA	2L MIN	NA	1	HTN	0	5	18.9	38	92	5	480	290	321	513	10.1	17.5	1385	1.23	0.85	>5	75	1.14	0.7	28	28	358	0.2		
25	1054158	2-6-21	35	M	CF		3	9	97%	RA	NA	NA	1	NONE	0	9	16.9	5.4	72	28	153	638	401	230	17.6	52	826	2.3	0.8	<5	21	1.12	0.4	28	28	114	0.09		
26	1053453	1-6-21	67	F	CM		3	15	90%	O2 MASK	2L MIN	NA	1	HTN, DM	1	12	13.2	8.3	81	15	226	127	423	232	111	33.4	485	1.25	0.94	<5	18	0.65	0.33	33	28	107	0.09		
27	1054285	6-6-21	40	F	CB		3	12	94%	O2 MASK	2L MIN	NA	1	NONE	1	15	10.9	4.1	64	30	48	10	256	173	1.9	13.3	189	1.14	0.96	<5	13	0.7	0.2	17	14	200	0.04		
28	1054277	1-6-21	44	M	CFB		8	16	90%	O2 MASK	2L MIN	NA	1	DM/HTN	2	13	11.9	9.8	70	21	149	355	417	264	12.5	24.4	121	1.22	0.87	<5	29	0.7	0.26	31	39	237	0.1		
29	1054278	3-6-21	36	M	CMB		3	9	99%	RA	NA	NA	1	NONE	1	15	13.6	8.8	80	12	267	79	400	199	65.7	43	236	1.1	0.86	<5	20	1.26	0.27	23	23	192	0.23		
30	1054191	2-6-21	41	M	CMB		3	12	94%	O2 MASK	1L MIN	NA	1	HTN	1	9	13.5	6.2	24	24	244	709	537	371	15.1	120	283	1.2	0.9	<5	25	0.8	0.25	30	27	90	0.09		
31	1053452	3-6-21	83	M	B	chest pain	1	16	95%	O2 MASK	1L MIN	NA	2	DM	1	11	13.5	8.2	66	24	132	129	229	376	1.9	55.9	409	1.17	0.94	<5	36	1.35	0.61	44	37	54	0.09		
32	1053013	10-6-21	70	M	CMF	chest pain	2	13	97%	RA	NA	NA	1	DM/HTN	1	13	13.8	3.1	78	15	109	1763	285	461	70	75	289	1.13	0.92	<5	22	0.88	0.6	53	29	109	0.08		
33	1054223	3-6-21	45	M	CF		4	15	97%	RA	NA	NA	1	NONE	1	14	11.4	4.6	79	16	149	813	233	313	37.5	50.6	240	1.13	0.92	<5	20	1.11	0.32	31	33	149	0.12		
34	1054154	2-6-21	85	M	CBF		2	26	88%	O2 MASK	4L MIN	NA	2	DM, ASTHMA	1	22	295	14.7	8.2	305	44	482	305	50.5	78.3	5000	1.1	0.67	<5	43	1.39	0.39	66	86	265	0.13			
35	1052456	15-5-21	52	F	CBF		3	22	91%	O2 MASK	2L MIN	NA	1	DM/HTN	1	16	13.7	2.3	73	19	171	298	461	504	29	8.1	314	1.19	0.4	<5	24	0.6	0.4	34	26	128	0.01		
36	1054263	3-6-21	49	F	CMF		3	16	99%	RA	NA	NA	1	NONE	2	9	10.1	5.4																					

