Journal of Pharmaceutical Research International



33(42A): 25-31, 2021; Article no.JPRI.71399 ISSN: 2456-9119 (Past name: British Journal of Pharmaceutical Research, Past ISSN: 2231-2919, NLM ID: 101631759)

Idiopathic Thrombocytopenic Purpura in Post COVID-19 Condition: Case Study

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Authors' contributions

This work was carried out in collaboration among all authors. Author CKVLSNAM did the concept design, critical revision and editing. Authors NVY, KV and VP did the case report gathering and drafting manuscript. Literature collection, verification done by authors AJA and TND. All authors Read and approved the final manuscript.

Article Information

DOI: 10.9734/JPRI/2021/v33i42A32380 <u>Editor(s)</u>: (1) Dr. Ana Cláudia Coelho, University of Trás-os-Montes and Alto Douro, Portugal. (2) Dr. Debarshi Kar Mahapatra, Rashtrasant Tukadoji Maharaj Nagpur University, India. (3) Dr. Sachin Kumar Jain, IPS Academy College of Pharmacy, India. (3) Dr. Sachin Kumar Jain, IPS Academy College of Pharmacy, India. (1) Dheeraj Lamba, , Jimma University, Ethiopia. (2) Papadatu Carmen-Penelopi, Dunarea de Jos University, Romania. Complete Peer review History: <u>https://www.sdiarticle4.com/review-history/71399</u>

Case Study

Received 10 June 2021 Accepted 15 August 2021 Published 25 August 2021

ABSTRACT

A 50 years old woman was referred to hospital with complaints of bluish black discoloration of skin or multiple ecchymosis since 6 days and skin rashes, pink changes or petechiae changes since 1 week, dark color stools since 3 days. She had a history of COVID-19 positive on past "3 months" back and she received the corticosteroids, antiviral drugs, broad spectrum antibiotics, anticoagulants, and vitamin B and C supplements. Now patient is admitted and investigated for further management. Her bone marrow examination reveals marrow cytological features are compatible with immune thrombocytopenic Purpura and peripheral examination reveals red cells are microcytic hypochromic with elongation forms, platelets are markedly reduced and elevation of CRP, reduction of the Hb, PCV, MCH, MCV, MCHC, APTT, Serum vitamin B12, Lymphocytes and

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her HRCT-Chest shows CORADS 5, Rapid antigen test shows positive. A review on introduction of the disease, etiology, pathogenesis, pathophysiology, mechanism of thrombocytopenia in COVID-19 patients and grading is explained in literature review.

Keywords: Idiopathic thrombocytopenia; petechiae; ecchymosis; purpura, corona virus disease; antiviral drugs.

1. INTRODUCTION

Wuhan is the capital city of Hubei province in China. In December 2019, a cluster of unknown acute respiratory illness occurred to many patients. This virus subsequently spread to other states in China and patients have been identified in other countries also [1,2,3]. The disease is known as corona virus disease (COVID-19) caused by severe acute respiratory syndrome corona virus 2 (SARS-CoV-2). It can be declared a pandemic by the World Health Organization on 11 March 2020 [1]. There are six types of corona viruses that can infect to humans have been identified those are HCoV-229E, HCoV-OC43, HCoV-NL63, HCoV-HKU1, SARS-CoV and MERS-CoV. The first four viruses mainly cause the common cold, where as the SARS-CoV and MERS-CoV viruses cause Severe Acute Respiratory Syndrome (SARS) and Middle East Respiratory Syndrome (MERS). COVID-19is known to cause serious respiratory symptoms and complications but, available data suggest that the symptoms of the disease can also result from involvement of other body systems such as hematopoietic, neurological and immune system [4,5]. The most common symptoms of the disease is febrile illness, fatigue associated with respiratory symptoms such as dry cough, dyspnea and anorexia but other atypical manifestations of the infection have been observed [6] and human to human transmission of COVID-19has been conformed [7].

The term Idiopathic Thrombocytopenic Purpura refers Unknown to an oriain or Thrombocytopenic state characterized bv decreased numbers of circulating platelets, normal or increased numbers of megakaryocytic in the bone marrow [8.9]. It is divided into 2 forms those are Acute and Chronic. The Acute form most commonly occurs in 2-7years of age although it is also present in adults but it has no preference for gender and it resolves within 6 months. Chronic form occurs more often in females between 20-40years of age with females to male ratio of 3:1. It is more persistent disease, lasting for more than 6 months, has an insidious onset and incidence is unknown [10].

2. ETIOLOGY

- 1.30% Idiopathic
- 2.30% of Drugs related

3.30% of underlying disease

- Connective tissue disorder,
- Lymphoma,
- Chronic Lymphocytic Leukemia.

4.10% Viral Infection

- HIV, Rubella, Rubeola etc.
- 5. Genetics: SLE

6. Immune factors

- Autoantibody mediated destruction of Platelets.
- Predisposing conditions and exposures

3. PATHOGENESIS

ITP appears to be immunologically mediated. An ant platelet IgG antibody reacts with host Platelets, causing rapid destruction by the Reticuloendothelial system (Fig.1).

They are two main reasons. Those are

A) The demonstration of increased level of Platelet associated IgG antibodies in more than 90% of patient with ITP.

B) Some studies show normal individuals Developed thrombocytopenia when injected with plasma from patients with ITP.

4. PATHOPHYSIOLOGY

Lymphocytes produce anti platelet antibodies directed at platelet surface glycoproteins.

A) Platelets are taken up and internally degraded by antigen presenting cells (APCs)

B) APCs present platelet antigen in association with major histocompatibility complex (MHC) class II to T helper cells, which become activated and secrete the Th1 cytokines interleukin-2 and IFN-g.

C) Th1 cytokines activate and drive auto reactive B cells to differentiate into autoantibody producing cells.

The IgG coated platelets are cleared by splenic macrophages, which results in thrombocytopenia.

The course of ITP is affected by pregnancy (relapse during pregnancy after remission worse if active).

Placental transfer of the IgG platelet antibodies can result in fetal or neonatal thrombocytopenia.

5. CASE REPORT

A 50 years old woman was referred to hospital with complaints of bluish black discoloration of skin or multiple ecchymosis since 6 days and skin rashes, pink changes or petechiae changes since 1 week, dark color stools since 3 days. She had a history of COVID-19positive on past "3 months" back in treatment.

6. PAST MEDICAL AND MEDICATION HISTORY

Chief Complaints: Cough, Shortness of breath since 4 days.

Diagnosis: COVID - 19 Positive, viral Pneumonia

7. PERIPHERAL BLOOD SMEAR

- Red cells are micro cystic hypochromic with elongated forms
- Platelets are markedly reduced

Table 1. Physical examinations

Vitals	
Pulse rate	84 beats/minute
Respiratory rate	20 cycles/min
Blood pressure	130/80 mm Hg
Temperature	98.6

Table 2. Systemic examinations

CVS	S1S2+
CNS	E4V5M6
RS	BAE+, Clear
P/A	Normal
SPO2	100%

8. BONE MARROW

Marrow cytological features are compatible with immune thrombocytopenic Purpura (Fig. 2).

Blood Group: AB Positive (blood transfusion was done)

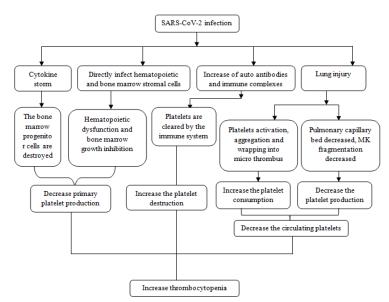


Fig 1. Mechanism of thrombocytopenia in COVID-19patients [11,12,13]

Parameters Abnormal values Normal values Haemoglobin 9.1 g/dl 14-16 g/dl PCV 30.6% 37-47% MCV 61.5fl 80-95 fl MCH 18.2pg 27-33 pg MCHC 29.6g/dl 31-36 g/dl RDW 19.0% 12.2-16.1% 1,50,000-4,00,000 Cells/cu.mm 15000Cells/cu.mm Platelets PΤ 11.9 seconds 11-13.5 seconds APTT 24.6 seconds 30-40 seconds Serum vitamin B12 139.6pg/mL 200-900 pg/m Serum folic acid 4.9ng/MI 2.7-17.0 ng/MI 326.7 mg/dL 200-400 mg/DI Fibrinogen levels Direct anti-globulin test Negative Indirect anti-globulin test Negative

Table 3. Lab tests

Table 4. Drug chart

Drugs	Doses	Frequency	
Inj. Pan	40mg	OD	
Inj. Vitamin B12	1000mcg	OD	
Tab. Fovite	5mg	OD	
Cap. Orofer-XT	1capsule	OD	
Inj. Methylprednisoslone	1gm	OD	
Tab. Primolut-N	4mg	TID	
Tab. Zolpidolom	5mg	HS	

Table 5. Lab investigations

Tests	Result	
HRCT-Chest	CORADS 5	
Rapid antigen test	Positive	
C-Reactive protein	250mg/L	
Serum ferritin	48ng/ml	
Blood group	AB Positive	

Table 6. Hematology

Parameters	Abnormal values	Normal Values	
Hb	7.5 g/dl	11-14g/dl	
PCV	26%	35-47%	
Lymphocytes	15%	26-46%	
Polymorphs	77%	40-65%	
ESŔ	50mm/hr	0-30mm/hr	

Table 7. HRCT-Chest severity

CT-Severity	% of involvement	Score
Right upper lobe	5-25%	2
Right middle lobe	5%	1
Right lower lobe	5-25%	2
Left upper lobe	5-25%	2
Left lower lobe	25-50%	3
	Total	10
	Severity	Moderate

Table 8. Hospital treatment

Drugs	Doses	
Tab. Augmentin	6 doses	
Tab. Pantocid-DSR	6 doses	
Syp. Ascoril	18 doses	
Tab. Prednisolone	12 doses	
Tab. Dexamethasone	6 doses	
Inj. Clexane	12 doses	
Tab. Limcee	6 doses	
Cap. Becosules	6 doses	
Inj. Remdesiver	6 doses	

Table 9. Discharge medications

Drugs	Dose	Frequency	
Tab. Pantocid-DSR	40/30mg	OD	
Tab. HCQ	200mg	BD	
Tab. Limcee	500mg	OD	
Tab. Omnacortil	20mg	BD	
Tab. Dabiclot	110mg	BD	
Cap.Becosules	1capsule	HS	



Fig. 2. This is the picture to the patient. The patient who is suffered with COVID-19can developed the Idiopathic Thrombocytopenic Purpura

9. DISCUSSION

Idiopathic Thrombocytopenic Purpura is one of the rare diseases. The incidence of the disease is more in females when compared to males. The most common cause of the disease is unknown. The standard treatment given to the patient is corticosteroids, intravenous immunoglobulins (IVIG) and Anti-D immune globulin the patients who having the Rh-positive blood group. In this condition patient have the Marrow cytological features are compatible with immune thrombocytopenic Purpura, Red cells are micro cystic hypochromic with elongated forms, Platelets are markedly reduced and elevation of C-Reactive protein, reduction of the Hemoglobin, packed cell volume, mean corpuscular hemoglobin, mean corpuscular volume, mean corpuscular hemoglobin concentration, APTT, Serum vitamin B12, Lymphocytes and her HRCT-Chest shows CORADS 5, Rapid antigen test shows positive.

We observed the decreasing level of platelets due to an immunologic reaction that can be seen in viral infections is the most common cause as it has been seen in dengue, influenza and HIV infections [14]. Like the SARS-CoV and MERS-CoV viruses [11], SARS-CoV may induce the thrombocytopenia and the mechanism involves the abnormal immune function, impairing bone marrow hemopoiesis directly [15,16] and lung damage [17]. Previous studies have reported thrombocytopenia in patients with COVID-19during the infection [18,19,20]. This is commonly represented by significant lymphopenia [21].

Lymphocytes and their sub types play a vital role in maintenance of the immune system [22]. The decreasing the lymphocytes and their subtypes, especially the CD4+ T cells, CD8+ T cells and NK (CD56+) cells was significantly indicates an impairment of the immune system caused by the SARS-CoV-2 infection, which might be trigger an autoimmune response [23]. CD4+ T cells help B cells to produce the virus specific antibodies and CD8+ T cells can kill the virus infected cells by using the cytotoxicity mechanism. Therefore the depletion and dysfunction of the lymphocytes and their subtypes induced the immune system become abnormal. This results the thrombocytopenia.

10. CONCLUSION

A 50 years old female patient was admitted in hospital with complaints of bluish black discoloration of skin or multiple ecchymosis since 6 days and skin rashes, pink changes or petechiae changes since 1 week, dark color stools since 3 days. She had a history of COVID-19positive on past "3 months" back in treatment. Her peripheral examination reveals Red cells are micro cystic hypochromic with elongated forms, Platelets are markedly reduced and bone marrow test reveals that Marrow cytological features are compatible with immune thrombocytopenic Purpura. Her lab tests reveal that elevation of C reactive protein and reduction of the Hemoglobin, cell volume, mean corpuscular packed hemoglobin, mean corpuscular volume, and mean corpuscular hemoglobin concentration, APTT, Serum vitamin B12, Lymphocytes and her HRCT-Chest shows CORADS 5, Rapid antigen test shows positive. The novel corona virus can also cause the systemic involvement those are Respiratory and Hematopoietic complications.

SUPPLEMENTARY MATERIAL

Supplementary material available in this following link:

https://www.journaljpri.com/index.php/JPRI/librar yFiles/downloadPublic/17

CONSENT

As per international standard or university standard, patient's consent has been collected and preserved by the authors.

ETHICAL APPROVAL

As per international standard or university standard written ethical approval has been collected and preserved by the author(s).

ACKNOWLEDGEMENT

We thank the manipal hospital and Nirmala College of pharmacy for carrying this information and giving support for publishing this case study.

COMPETING INTERESTS

Authors have declared that no competing interests exist.

REFERENCES

- Phan LT, Nguyen TV, Luong QC, Nguyen TV, Nguyen HT, Le HQ, Nguyen TT, Cao TM, Pham QD. Importation and human-tohuman transmission of a novel corona virus in Vietnam. New England Journal of Medicine. 2020;382(9):872-4.
- Rothe C, Schunk M, Sothmann P, Bretzel G, Froeschl G, Wallrauch C, Zimmer T, Thiel V, Janke C, Guggemos W, Seilmaier M. Transmission of 2019-nCoV infection from an asymptomatic contact in Germany. New England journal of medicine. 2020; 382(10):970-1.
- Holshue ML, DeBolt C, Lindquist S, Lofy KH, Wiesman J, Bruce H, Spitters C, Ericson K, Wilkerson S, Tural A, Diaz G. First case of 2019 novel corona virus in the United States. New England Journal of Medicine; 2020.
- Mehta P, Mc Auley DF, Brown M, Sanchez E, Tatter sall RS, Manson JJ. COVID-19: consider cytokine storm syndromes and immunosuppression. The lancet. 2020; 395(10229):1033-4.
- Driggin E, Madhavan MV, Bikdeli B, Chuich T, Laracy J, Biondi-Zoccai G, Brown TS, Der Nigoghossian C, Zidar DA, Haythe J, Brodie D. Cardiovascular considerations for patients, health care workers, and health systems during the COVID-19pandemic. Journal of the American College of Cardiology. 2020; 75(18):2352-71.
- 6. Zhang JJ, Dong X, Cao YY, Yuan YD, Yang YB, Yan YQ, Akdis CA, Gao YD. Clinical characteristics of 140 patients infected with SARS–CoV-2 in Wuhan, China. Allergy. 2020;75(7):1730-41.
- Li Q, Guan X, Wu P, Wang X, Zhou L, Tong Y, Ren R, Leung KS, Lau EH, Wong JY, Xing X. Early transmission dynamics in Wuhan, China, of novel corona virus– infected pneumonia. New England journal of medicine; 2020.

- Mahévas M, Moulis G, Andres E, Riviere E, Garzaro M, Crickx E, Guillotin V, Malphettes M, Galicier L, Noel N, Darnige L. Clinical characteristics, management and outcome of COVID-19-associated immune thrombocytopenia: a French multicentre series. British journal of haematology. 2020;190(4):e224-9.
- Vermeiren C, Marchand-Senécal X, Sheldrake E, Bulir D, Smieja M, Chong S, Forbes JD, Katz K. Comparison of Copan Eswab and FLOQ swab for COVID-19PCR diagnosis: working around a supply shortage. Journal of clinical microbiology; 2020.
- Helms RA, Quan DJ, editors. Textbook of therapeutics: drug and disease management. Lippincott Williams & Wilkins; 2006.
- 11. Xu P, Zhou Q, Xu J. Mechanism of thrombocytopenia in COVID-19patients. Annals of hematology. 2020;99(6):1205-8.
- 12. Yang M, Ng MH, Li CK. Thrombocytopenia in patients with severe acute respiratory syndrome. Hematology. 2005;10(2): 101-5.
- Chan JF, Kok KH, Zhu Z, Chu H, To KK, Yuan S, Yuen KY. Genomic characterization of the 2019 novel humanpathogenic corona virus isolated from a patient with atypical pneumonia after visiting Wuhan. Emerging microbes & infections. 2020;9(1):221-36.
- 14. Hottz ED, Bozza FA, Bozza PT. Platelets in immune response to virus and immunopathology of viral infections. Frontiers in medicine. 2018;5:121.
- The L. Emerging understandings of 2019nCoV. Lancet (London, England). 2020; 395(10221):311.
- 16. Eickmann M, Gravemann U, Handke W, Tolksdorf F, Reichenberg S, Müller TH, Seltsam A. Inactivation of three emerging viruses-severe acute respiratory syndrome corona virus, Crimean-Congo haemorrhagic fever virus and Nipah virusin platelet concentrates by ultraviolet C light and in plasma by methylene blue plus

visible light. Voxsanguinis. 2020;115(3): 146-51.

- Poon TC, Pang RT, Chan KA, Lee NL, Chiu RW, Tong YK, Chim SS, Ngai SM, Sung JJ, Lo YD. Proteomic analysis reveals platelet factor 4 and beta thromboglobulin as prognostic markers in severe acute respiratory syndrome. Electrophoresis. 2012;33(12): 1894-900.
- Chang D, Lin M, Wei L, Xie L, Zhu G, Cruz CS, Sharma L. Epidemiologic and clinical characteristics of novel corona virus infections involving 13 patients outside Wuhan, China. Jama. 2020;323(11):1092-3.
- 19. Chan JF, Yuan S, Kok KH, To KK, Chu H, Yang J, Xing F, Liu J, Yip CC, Poon RW, Tsoi HW. A familial cluster of pneumonia associated with the 2019 novel corona virus indicating person-to-person transmission: a study of a family cluster. The lancet. 2020;395(10223):514-23.
- Huang C, Wang Y, Li X, Ren L, Zhao J, Hu Y, Zhang L, Fan G, Xu J, Gu X, et al. Clinical features of patients infected with 2019 Novel corona virus in Wuhan, China. Lancet (London, England). 2020; 395(10223):497–506.
- Terpos É, Ntanasis□Stathopoulos I, Elalamy I, Kastritis E, Sergentanis TN, Politou M, Psaltopoulou T, Gerotziafas G, Dimopoulos MA. Haematological findings and complications of COVID□19. American journal of hematology. 2020 ;95(7):834-47.
- Chen Y, Xie Y, Ruan M, Shi J. The levels of T lymphocyte subsets in immune thrombocytopenia associated with anti-GPIIb/IIIa and/or anti-GPIbα-mediated responses are differentially sensitive to dexamethasone. Act a haematological. 2018;140(1): 60-6.
- Audia S, Mahévas M, Samson M, Godeau B, Bonnotte B. Pathogenesis of immune thrombocytopenia. Autoimmunity reviews. 2017;16(6):620-32.

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