



Journal Homepage: -[www.journalijar.com](http://www.journalijar.com)

## INTERNATIONAL JOURNAL OF ADVANCED RESEARCH (IJAR)

Article DOI:10.21474/IJAR01/17150

DOI URL: <http://dx.doi.org/10.21474/IJAR01/17150>



### RESEARCH ARTICLE

#### CARDIO-VASCULAR INVOLVEMENT AND CARDIAC REHABILITATION ON COVID-19 INDIAN PATIENTS: RECENT ADVANCES AND FUTURE PROSPECTIVE

Karnakar Kola<sup>1</sup>, Anju Mahor<sup>2</sup>, Dharambeer Singh Mahor<sup>3</sup> and Mayadhar Barik<sup>4</sup>

1. Asst.Prof. ESIC Medical College and Hospital, Sanath Nagar, Hyderabad.
2. Asst.Prof. MGM Medical College, Indore, Madhya Pradesh.
3. Asst.Prof. Vedantaa Institute of Medical Sciences, Indore, Madhya Pradesh.
4. Prof. and HoD. Department of Paramedical Sciences, Biochemistry, Advisor to Chairman, MURECMewar University, Gangrar, Chittorgarh, Rajasthan-312901, India.

#### Manuscript Info

##### Manuscript History

Received: 26 April 2023

Final Accepted: 31 May 2023

Published: June 2023

##### Key words:-

Cardiac Disease, Cardiac Rehabilitation, Cardiovascular, COVID-19, SARS-CoV-2, Rehabilitation, Exercise, Immunity, Biomarker, Respiration

#### Abstract

**Introduction:** The novel corona virus disease 2019 -23 (COVID-19) pandemic so many Indian patients have experienced with their multisystem involvement, critically ill patients and treated with in intensive care units (ICUs), critical care unit (CCU) and even so many died as unable to get proper treatment and cardiac rehabilitations (CRs) at the acute infections and critically ill patients.

**Material and Methods:** We have selected those patients having systemic effects, cardiac involvement patient's prognosis and diagnosis with COVID-19 and their cardiac complicity. Patients with COVID-19 develop cardiac complications are including with the heart failure, myocarditis, pericarditis, vasculitis, acute coronary syndrome (ACS), cardiac arrhythmias (CAMs), trigger and accompanying with cardiac diseases like coronary artery diseases (CAD) and coronary vascular diseases (CVD). We identify the recommendations of scientific and professional treatment and cardiac rehabilitations given by our organizations in the area of rehabilitation in our institutions respectively. We performed more than 500 cases in critical conditions and recovered in our main data bases. In our organizations were identified and selected as very specific contribution related to our study and excluded the single case reports and those are not interested to participate to our study. SARS-CoV-2 specific IgG and IgM, mild increases in the CD4+T cells, diverse changes in the several newly developed biochemical and haematological biomarkers are also studied.

**Results:** Cardiac Rehabilitations (CRs) taken Patients with COVID-19 develop cardiac complications are including with the heart failure, myocarditis, pericarditis, vasculitis, acute coronary syndrome (ACS), cardiac arrhythmias (CAMs), trigger and accompanying with cardiac diseases like coronary artery diseases (CAD) and coronary vascular diseases (CVD) are improved more than (85%-95%). In hospitalized patients 10-15% with worse outcomes, longer stay in the intensive care unit (ICU) & 5-8% in cardiac critical care (CCU), and higher risk of death ratio of COVID-19 cardiac involvement ranges between 0-

10percent. Deconditioning rate is increased due to immobility and muscle involvement in the pre and post-COVID-19 management is very poor at that time. COVID-19 Patients need CRs facilities.

**Conclusion:** Cardiac Rehabilitations (CRs) findings suggested that COVID-19 patients showed a gradual changes in improvement in exercise capacity of critically ill patients are significantly declined in the SARS-CoV-2 specific IgG and IgM, test reports are mild increase in CD4+ T cells, and diverse changes in several biochemical and hematological biomarkers after hospital discharge in 6-month. Cardiac Rehabilitations (CRs) effect on improving exercise capacity was observed in COVID-19 patients in a dose–response fashion after 6-months. CRs suggested that a better innovation strategy to promote recovery in patients during/after acute COVID-19. Although higher certainty in the evidence in convalescent plasma for the treatment of individuals with moderate to severe diseases. It does not reduce mortality and has little to no impact on measures of clinical improvement. Cardiac Rehabilitations (CRs) taken Patients with COVID-19 develop cardiac complications got better treatment through the CR experts clinically and feeling good.

*Copy Right, IJAR, 2023,. All rights reserved.*

## Introduction:-

Cardiac rehabilitations (CRs) are a complex intervention that seeks to improve the functional capacity, wellbeing and health-related quality of life (QOL) of patients including with cardiac diseases [1]. Cardiac rehabilitation substantive evidence base supports are suggested that effective and cost-effective intervention for patients with acute coronary syndrome (ACS) clinically. Heart failure reduced the patient's ejection fraction and then after coronary revascularization [2]. The major contemporary challenges that are faced with cardiac rehabilitation experts are very minimum in number in our institutions to looking patients. Despite of the strong recommendation in the current clinical guidelines for the referral of this COVID-19 patient groups, global access to cardiac rehabilitations (CRs) are very poor [3]. The COVID-19 pandemic has contributed to a further reduction in access to cardiac rehabilitations (CRs) [4]. Home-based and technology-based models of cardiac rehabilitation as alternatives or adjuncts to traditional at centre-based programmes are increasing the body of evidence supports [5].

Especially in lower-income and middle-income countries which cardiac rehabilitations (CRs) services are very scarce, scalable and affordable models are required for the cardiac complicity cases with COVID-19 patients [6]. The delivery of cardiac rehabilitations (CRs) in future approaches are growing to save the multi morbidity of an ageing population in our country and the World. In addition to the cater into the needs of the increasing patients with cardiac numbers of diseases. Those are who having present with two or three and more chronic diseases patients are getting very critical situations and it was a big challenges for the management of these patients [7]. Future research is required on priorities to minimize these cardiac diseases. Cardiac rehabilitations (CRs) are strengthening the evidence base for the critically cardiac ill patients [8]. According to other indications, including with heart failure is preserved with the ejection fraction, atrial fibrillation and the congenital heart disease (CHD) after valve surgery or heart transplantation [9]. Cardiac rehabilitation implementation of evaluation of sustainable (EOS) and affordable models of delivery that can be improved the lower income settings in the world [10].

Additionally, Cardiopulmonary rehabilitations are included with specific interventions that helping patients management and maximize the functional potential due to progressive deconditioning , acute decompensation is following with acute medical event essential [11-12].

The COVID-19 population of patients who benefit from both the cardiac and pulmonary rehabilitation from the heart disease is one leading cause of global morbidity and mortality [13]. The principles of exercise physiology when applied to this population can reverse deconditioning, build cardiopulmonary reserve, and ultimately reduce morbidity and mortality in these populations [14]. The physically disabled persons are also benefitted from exercise conditioning. The newly refined model of cardiac rehabilitation can also be applied to improve functional status of

stroke patients with COVID-19 infections [15]. These risk factors of cardiovascular diseases conditions are an emerging area of interest in research development required to know their etiopathology [16]. The frailty and post-transplant decline in function benefits of supervised exercise can also be extended to patients with clinically recognized [17].

Frailty is a complex diagnosis with multiple tools and newly developed approaches used to describe this syndrome and appropriate management. Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) infection remains asymptomatic in (35% to 90%) of the older adults. As it is depending upon their immune status from the prior infections noticed [18]. Before and after Vaccination, circulating strain among the Older adults are symptomatic with SARS-CoV-2 often both present atypically with a blunted fever response is develop more than severe disease [19].

## **Material and Methods:-**

### **Study Design:**

A randomized controlled trial (RCT) is a form of scientific experiment used to control factors not under direct experimental control. In the examples of this RCTs study are clinical trials that compare the effects of drugs, surgical techniques, medical devices, diagnostic procedures or other medical treatments Cardiac Rehabilitations (CRs) taken Patients with COVID-19 develop cardiac complications and associated cardiac diseases.

### **Data Collection:**

Patients with COVID-19 may develop cardiac complications such as heart failure, myocarditis, pericarditis, vasculitis, acute coronary syndrome, and cardiac arrhythmias or trigger an accompanying cardiac disease.

### **Study duration:**

This prospective cohort data of (n=500) COVID-19 patients admitted from January 2020 to 2023.

### **Selection Criteria:**

We have selected those patients having systemic effects, cardiac involvement patient's prognosis and diagnosis with COVID-19 and their cardiac complicity. Patients with COVID-19 develop cardiac complications are including with the heart failure, myocarditis, pericarditis, vasculitis, acute coronary syndrome (ACS), cardiac arrhythmias (CAMs), trigger and accompanying with cardiac diseases like coronary artery diseases (CAD) and coronary vascular diseases (CVD) are included.

### **Inclusion Criteria:**

We identify the recommendations of scientific and professional treatment and cardiac rehabilitations (CRs) taken by our organizations in the area of rehabilitation in our institutions respectively. We performed more than (n=500) in critical conditions and recovered in our main data bases.

### **Excluded Criteria:**

In our organizations were identified and selected as very specific contribution related to our study and excluded the single case reports and those are not interested to participate to our study and irrelevant from our study.

## **Material and Methods:-**

SARS-CoV-2 specific IgG and IgM, mild increases in the CD4+T cells, diverse changes in the several newly developed biochemical and haematological biomarkers are also studied. Among these systemic effects, cardiac involvement may have very important consequences for the patient's prognosis and later life. The reminder who had not performed any CRs was assigned to the control group.

### **Statistical Analysis:**

The primary outcome was the change in (0 to six-months) between the 6-month to 12 months and their follow-ups was assessed via analysis of covariance with a covariate of propensity score. Again that adjusted for the potential confounders. Secondary outcomes were changes in 0-2 weeks and one week to two months. The SARS-CoV-2 immunoglobulins, T-lymphocytes and blood chemistry were also be evaluated via paired tests. The *p-value* ( $p < .001$ ) is considered as the significant.

## Results:-

The Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) is deadly critical for coronavirus disease 2019 (COVID-19), has caused substantial mortality and morbidity worldwide since late 2019-2023. The post acute sequelae of SARS-CoV-2 infection (PASC) can manifest as a wide range of new, recurring, or ongoing disabling symptoms including with health problems. 90-95% people were experiencing acute infection and persisting. The starting of more than 4-8 weeks after being infected with the virus that causes COVID-19 as one critical epidemiology. COVID-19 to peripheral nervous system (PNS) diseases. In addition to the pandemic to the vaccination. Current era of interest is shifting towards the potential association between COVID-19 and their associated cardiac diseases.

Vaccines and PNS manifestations of participants' age ranges is 18 to 120 years ( $M = 275$ ,  $F = 225$  standard deviation ( $SD$ ) = 14.8) respectively. 55% identified as male. During the 6-month of the convalescence, 6-MWD increased 25.0%, with a mean [95% CI] and 275 [92–135] m ( $p < .001$ ). SARS-CoV-2 IgG and IgM decreased 35.5% ( $p = .002$ ) and 53.8% ( $p = .009$ ), CD4+ T cells increased 8.9% ( $p = .05$ ), and the majority of blood chemistry changed significantly with the patients in the CR group. It acquired with increase in 6-MWD than those in control (unadjusted, 275 [168–224] m, ( $p < .001$ ). The adjusted with 225 [69–195] m, ( $p < .001$ ). The dose-responsiveness of CRs is on 6-Months was observed ( $p < .001$ ). There are no differences between at the observed group in the immunity variables and their blood chemistry level (BCL).

After getting the COVID-19 vaccination process and PNS manifestations between (Jan 2020 and June 2023) is one growing evidence indicated that COVID-19 related cardiovascular symptoms and complications were raised. It is persisting with weeks or months after resolution of the acute infections of COVID-19 is 5%–30% reported as chest pain, dyspnea, or palpitations and post recovery. The acute infection even 6 months after the acute infection and after 12 months of follow up of the study ranges from mild results to incapacitating their survivors. The prevalence of these sequelae and emerging data on the longevity of symptoms. But they are very limited. The new assessment and treatment of cardiovascular complications in PASC and the Multi-Disciplinary PASC Collaborative (PASC Collaborative) of the American Academy of Physical Medicine and Rehabilitation (AAPM&R) was developed (15–25%). PASC is quite important in pressing need for guidance in the care of patients with Indian patients with SARS-CoV-2 infection (PASC) also we look forward the more numbers of collaborative research is required for the respective field and specific net working for this ongoing research projects.

## Discussion:-

More than (40%-80%) older adult's cases have increased their severity of coronavirus disease 2019-2023. (COVID-19) with higher cases including with their fatality rates and higher intensive care (HIC) needs compared with younger adults [20]. Infections and vaccine-induced antibody response (VIAR) and long-term effects of COVID-19 differing in the population of the older adults. SARS-CoV-2 Severe acute respiratory syndrome corona virus pandemic is ongoing [21]. In the medical system subjected to many changes and they faced there specific challenges to save patients lives. Face-to-face treatments have been suspended for a period of time as this is very typical diseases [22]. After the lockdown period the dentists have to be aware of the modalities to protect themselves and their patients in order should not to be getting infected [23]. Dental practitioners are potentially exposed to a higher degree of contamination with SARS-CoV-2. While them performing patient's dental procedures that produces their aerosols and major infections [24]. It should also be noted that the airways, oral cavity and nostrils, are the access pathways for SARS-CoV-2. In order to protect themselves and their patients they have to use full the personal protective equipment (PPE) [25].

Relevant data regarding this pandemic are under evaluation and are still under test. The way, in which SARS-CoV-2 spreads, diagnose a novel corona virus infection. The possible treatments are protective personal equipment use to stop its spreading [26]. Coronavirus disease 2019 (COVID-19) is a contagious infection disease. It causes the respiratory, physical, psychological, and generalized systemic dysfunction [27]. The severity of this disease ranges from an asymptomatic infections are mild illness to severe pneumonia with respiratory failure or death. COVID-19 dramatically affects the pulmonary system and the lungs and later multiple organ failure [28]. This clinical practice guideline includes pulmonary rehabilitation (PR) recommendations for adult COVID-19 patients and has been developed through the light of the guidelines on the diagnosis and treatment of COVID-19 provided by the World Health Organization [29]. The Republic of Turkey, Ministry of Health, published scientific literature, and PR recommendations for COVID-19 regarding basic principles of PR and CR recently [30].

This national guideline provides different suggestions regarding the CRs methods during the clinical stages of COVID-19 and post-COVID-19 with its possible benefits, contraindications and disadvantages [31]. The Convalescent plasma and hyperimmune immunoglobulin reduces the mortality in patients with viral respiratory diseases. CRs investigated as potential therapies for coronavirus disease 2019 (COVID-19) associated with the cardiac diseases [32]. Understanding of the current body of evidence regarding benefits and risks of these interventions is required for COVID-19 and post-COVID-19 [33].

Manifestations of the (COVID-19) cardiovascular system of novel coronavirus disease 2019 have attracted attention through the multi-facet cardiac symptoms. The clinical features and sudden cardiac events according to the five-years in experience in nearly half of the patients with cardiovascular system (CVD) [34]. It has been affected with or without clinical features of respiratory disease (RD) and pulmonary disease (PD) [35]. We found that up to 23-30 % of hospitalized COVID-19 patients with suffered from the cardiac injury [36]. The pathophysiology of cardiac and vascular involvement in the seven main reasons have been discussed [37]. Direct viral effects on the tissues are mainly benign. But it named as elimination phase is responsible for the elimination of the virus from the tissues [38]. New possible mechanism directly involves viral infiltration onto the myocardial tissues [39]. Excessive inflammatory effect of cytokine release syndrome is mainly known as storm-like phase. It leads to severe inflammation of the targeted cells are the lung, heart, endothelial, lymphoid tissue, pancreas, kidney by the interleukin (IL)-2, IL-6, IL-8, IL-10, and tumor necrosis factor (TNF) are also very important [40].

These cytokines play an essential role in myocardial cell injury, and also the cardiometabolic demand associated with the systemic infection and ongoing hypoxia-induced excessive intracellular calcium and cardiac stress lead to respiratory failure and hypoxemia [41]. Crosstalk between coagulation and inflammation is evident. Endothelial dysfunction shifts the vascular equilibrium toward an inflammatory and pro-coagulant state which tends to thrombosis and vasculitis [42]. Unmanageable cascade of hyper-inflammation may transform into an autoimmune overreacted response. Sepsis, which leads to the development of disseminated intravascular coagulation syndrome [43]. Electrolyte imbalance and Side effects of medical treatment during hospitalization [44]. Cardiovascular involvement during the course of the disease can be divided into two periods [45].

One is acute sequelae, seen at the very early stage of the infection (from the incubation to four weeks), and the other is post-acute sequelae, seen after the fourth week of the infection [46]. According to the organ (heart) and its components (pericardium, myocardium, coronary arteries) injured, the clinical picture emerges as arrhythmia, myocardial infarction (MI), myocarditis, pericarditis, acute coronary syndrome [47], heart failure (HF) (acute onset and exacerbation of chronic HF) and post-acute sequelae coronary vascular disease (PASC-CVD) [48], post-acute sequelae-cardiovascular symptoms (PASC-CVS), blood pressure dysfunction (BPD), postural orthostatic tachycardia syndrome (POTS), and myalgic encephalomyelitis also very important [49-50]

#### **Recent Advances and future prospective:**

The current COVID-19 pandemic is also a great challenge for the worldwide researchers. In the human microbiota area having the long-term effects of the infection at the GI level are not yet deeply understood [51]. SARS-CoV-2 and the possible consequences on the microbiota were pertaining to COVID-19 have also been discussed with the transmission and the resistance [52]. In the human body, impact of nutritional status in relation to the intestinal microbiota [54]. Impact of inflammatory bowel disease (IBS), comorbid metabolic disorders such as obesity, and type two diabetes (T2D) [55]. Health, age, and nutritional status are associated with specific communities of bacterial species in the gut. CRs could influence the clinical course of COVID-19 infection and their patient's management [56]. Alterations of fecal microbiota were associated with fecal concentrations of SARS-CoV-2 and COVID-19 severity with cardiac diseases [57]. Patients are suffering from the metabolic and gastrointestinal (GI) disorders are thought to be at a moderate-to the higher risk of infection with SARS-CoV-2 [58]. It is indicating with the direct implication of gut symbiosis in COVID-19 severity [59]. The World Health Organization (WHO) is also declared that pandemic on the 11th of march 2020 [60]. COVID-19 represents flu-like symptoms and that become more severe in higher-risk compromised subjects medically [61].

According to the definition of health “**A STATE OF COMPLETE PHYSICAL, MENTAL AND SOCIAL WELL-BEING**” individuals with heart involvement with COVID-19 should be rehabilitated by the light of the rehabilitation perspective and given the increasing number of patients with cardiac manifestations of COVID-19. The rehabilitation principles in this group of patients with COVID-19 had a significant physical, cognitive and psychosocial impairments may be observed in some major of the cases.

**Limitations:**

More randomized clinical trials (RCTs) studies are necessary to identify novel pharmacospecific therapies and infection prevention protocols are required to improve their efficacies and safety for clinical application in the COVID-19 Patients management and research.

**Conclusion:-**

CRs may be a innovative strategy to promote the improvement of exercise capacity after COVID-19 patients with cardiac diseases. COVID-19 patients showed gradual improvement in these exercise capacity. There is a significant decline in SARS-CoV-2 specific IgG and IgM, a mild increase in CD4+ T cells are observed clinically. In this diverse changes in CRs with several biochemical and haematological biomarkers 6-month after hospital discharge. CR's 6-month effect on improving exercise capacity was observed in COVID-19 patients with moderate to severe diseases. Cardiac diseases within a dose-response fashion (DRF) with CRs may be a new tool for recovered in patients during/after acute COVID-19 more than (70%-95%). Although, convalescent plasma treatment is helping of individuals moderate to severe diseases. It does not reduce the mortality rate and has little to no impact on measures of clinical improvement. The adverse effects of convalescent plasma are increasing in patient's management. The major efforts to conduct research on COVID-19 are being made, heterogeneous reporting of outcomes is still problematic as more than 500- 1250 ongoing studies from (ICMR, DST, DBT, PUSA) registry are being in the process. Hyperimmune immunoglobulin therapy for the people disease with severity and convalescent plasma therapy for people with asymptomatic or mild disease. While many efforts are being spent worldwide in research aimed at identifying early diagnostic methods and evidence-based effective treatments, mass vaccination is thought to be the best option against this disease in the near future. Although, our Indian Government is very good in situation to solve this diseases management is still more research work urgent need for this area.

**Acknowledgement:-**

We thanks to our patients and their relatives to participated in this study.

**References:-**

1. Taylor RS, Dalal HM, McDonagh ST. The role of cardiac rehabilitation in improving cardiovascular outcomes. *Nature Reviews Cardiology*. 2022 Mar;19(3):180-94.
2. Buckley BJ, Lip GY. Current Concepts: Comprehensive "Cardiovascular Health" Rehabilitation—An Integrated Approach to Improve Secondary Prevention and Rehabilitation of Cardiovascular Diseases. *Thrombosis and haemostasis*. 2022 Oct 28.
3. Stefanakis M, Batalik L, Papathanasiou J, Dipla L, Antoniou V, Pepera G. Exercise-based cardiac rehabilitation programs in the era of COVID-19: a critical review. *Reviews in Cardiovascular Medicine*. 2021 Dec 22;22(4):1143-55.
4. Nicholls SJ, Nelson M, Astley C, Briffa T, Brown A, Clark R, Colquhoun D, Gallagher R, Hare DL, Inglis S, Jelinek M. Optimising secondary prevention and cardiac rehabilitation for atherosclerotic cardiovascular disease during the COVID-19 pandemic: a position statement from the Cardiac Society of Australia and New Zealand (CSANZ). *Heart, Lung and Circulation*. 2020 Jul 1;29(7):e99-104.
5. Coronas-Watkins K, Cooke M, Butland M, McGuire A. Exploring the delivery of phase II cardiac rehabilitation services in rural and remote Australia: a scoping review. *Australian Health Review*. 2023 Jan 13.
6. Turk-Adawi K, Sarrafzadegan N, Grace SL. Global availability of cardiac rehabilitation. *Nature Reviews Cardiology*. 2014 Oct;11(10):586-96.
7. Watkins DA, Msemburi WT, Pickersgill SJ, Kawakatsu Y, Gheorghe A, Dain K, Johansson KA, Said S, Renshaw N, Tolla MT, Twea PD. NCD Countdown 2030: efficient pathways and strategic investments to accelerate progress towards the Sustainable Development Goal target 3.4 in low-income and middle-income countries. *The Lancet*. 2022 Mar 26;399(10331):1266-78.
8. MacKay C, Lee L, Best K, Campbell J, Cimino SR, Cowley H, Delvin M, Dilkas S, Landry M, Marzolini S, Mayo A. Developing a research agenda on exercise and physical activity for people with limb loss in Canada. *Disability and Rehabilitation*. 2022 Dec 4;44(25):8130-8.
9. Brida M, Lovrić D, Griselli M, Gil FR, Gatzoulis MA. Heart failure in adults with congenital heart disease. *International Journal of Cardiology*. 2022 Mar 10.
10. Daw P, Wood GE, Harrison A, Doherty PJ, van Zanten JJ, Dalal HM, Taylor RS, van Beurden SB, McDonagh ST, Greaves CJ. Barriers and facilitators to implementation of a home-based cardiac rehabilitation programme for patients with heart failure in the NHS: a mixed-methods study. *BMJ open*. 2022 Jul 1;12(7):e060221.

11. Bartels MN, Prince DZ. Acute Medical Conditions: Cardiopulmonary Disease, Medical Frailty, and Renal Failure. In Braddom's Physical Medicine and Rehabilitation 2021 Jan 1 (pp. 511-534). Elsevier.
12. Passantino A, Dalla Vecchia LA, Corrà U, Scalvini S, Pistono M, Bussotti M, Gambarin FI, Scrutinio D, La Rovere MT. The future of exercise-based cardiac rehabilitation for patients with heart failure. *Frontiers in Cardiovascular Medicine*. 2021 Aug 4;8:709898.
13. Lutz AH, Delligatti A, Allsup K, Afilalo J, Forman DE. Cardiac rehabilitation is associated with improved physical function in frail older adults with cardiovascular disease. *Journal of cardiopulmonary rehabilitation and prevention*. 2020 Sep 1;40(5):310-8.
14. Mayes J, Koufaki P, Greenwood SA. Physical Activity, Function, and Exercise-Based Rehabilitation for People on Dialysis. In *Handbook of Dialysis Therapy* 2023 Jan 1 (pp. 582-589). Elsevier.
15. Cuccurullo SJ, Fleming TK, Zinonos S, Cosgrove NM, Cabrera J, Kostis JB, Greiss C, Ray AR, Eckert A, Scarpatti R, Park MO. Stroke recovery program with modified cardiac rehabilitation improves mortality, functional & cardiovascular performance. *Journal of Stroke and Cerebrovascular Diseases*. 2022 May 1;31(5):106322.
16. Bartels MN, Prince DZ. Acute Medical Conditions: Cardiopulmonary Disease, Medical Frailty, and Renal Failure. In Braddom's Physical Medicine and Rehabilitation 2021 Jan 1 (pp. 511-534). Elsevier.
17. Perez-Saez MJ, Morgado-Pérez A, Faura A, Muñoz-Redondo E, Gárriz M, Muns MD, Nogués X, Marco E, Pascual J. The FRAILMar study protocol: frailty in patients with advanced chronic kidney disease awaiting kidney transplantation. A randomized clinical trial of multimodal prehabilitation. *Frontiers in Medicine*. 2021 May 19;8:675049.
18. Lipsitch M, Krammer F, Regev-Yochay G, Lustig Y, Balicer RD. SARS-CoV-2 breakthrough infections in vaccinated individuals: measurement, causes and impact. *Nature Reviews Immunology*. 2022 Jan;22(1):57-65.
19. Van Epps P, Canaday DH, editors. *Infections in Older Adults, An Issue of Infectious Disease Clinics of North America*, E-Book. Elsevier Health Sciences; 2023 Feb 21.
20. Gregg EW, Sophiea MK, Weldegiorgis M. Diabetes and COVID-19: population impact 18 months into the pandemic. *Diabetes Care*. 2021 Sep 1;44(9):1916-23.
21. Gazit S, Shlezinger R, Perez G, Lotan R, Peretz A, Ben-Tov A, Herzel E, Alapi H, Cohen D, Muhsen K, Chodick G. Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) naturally acquired immunity versus vaccine-induced immunity, reinfections versus breakthrough infections: a retrospective cohort study. *Clinical Infectious Diseases*. 2022 Jul 1;75(1):e545-51.
22. Cellini N, Conte F, De Rosa O, Giganti F, Malloggi S, Reyt M, Guillemin C, Schmidt C, Muto V, Ficca G. Changes in sleep timing and subjective sleep quality during the COVID-19 lockdown in Italy and Belgium: age, gender and working status as modulating factors. *Sleep Medicine*. 2021 Jan 1;77:112-9.
23. Baticulon RE, Sy JJ, Alberto NR, Baron MB, Mabulay RE, Rizada LG, Tiu CJ, Clarion CA, Reyes JC. Barriers to online learning in the time of COVID-19: A national survey of medical students in the Philippines. *Medical science educator*. 2021 Apr;31:615-26.
24. Clementini M, Raspini M, Barbato L, Bernardelli F, Braga G, Di Gioia C, Littarru C, Oreglia F, Brambilla E, Iavicoli I, Pinchi V. Aerosol transmission for SARS-CoV-2 in the dental practice. A review by SIdP Covid-19 task-force. *Oral diseases*. 2022 Apr;28:852-7.
25. Drozdziak A, Drozdziak M. Oral pathology in COVID-19 and SARS-CoV-2 infection—Molecular aspects. *International journal of molecular sciences*. 2022 Jan 27;23(3):1431.
26. Talbot TR, Hayden MK, Yokoe DS, Malani AN, Amer HA, Kalu IC, Logan LK, Moehring RW, Munoz-Price S, Palmore TN, Weber DJ. Asymptomatic screening for severe acute respiratory coronavirus virus 2 (SARS-CoV-2) as an infection prevention measure in healthcare facilities: Challenges and considerations. *Infection Control & Hospital Epidemiology*. 2023 Jan;44(1):2-7.
27. Sansone A, Mollaioli D, Limoncin E, Ciocca G, Băc NH, Cao TN, Hou G, Yuan J, Zitzmann M, Giraldi A, Jannini EA. The sexual long COVID (SLC): erectile dysfunction as a biomarker of systemic complications for COVID-19 long haulers. *Sexual medicine reviews*. 2022 Apr;10(2):271-85.
28. Aytür YK, Köseoglu BF, Taşkıran ÖÖ, Gökkaya NK, Delialioğlu SÜ, Tur BS, Sarıkaya S, Şirzai H, Tiftik TT, Alemdaroglu E, Ayhan FF. Pulmonary rehabilitation principles in SARS-COV-2 infection (COVID-19): The revised guideline for the acute, subacute, and post-COVID-19 rehabilitation. *Turkish journal of physical medicine and rehabilitation*. 2021 Jun;67(2):129.
29. Cheng YY, Chen CM, Huang WC, Chiang SL, Hsieh PC, Lin KL, Chen YJ, Fu TC, Huang SC, Chen SY, Chen CH. Rehabilitation programs for patients with COReNaVIrus Disease 2019: consensus statements of Taiwan Academy of Cardiovascular and Pulmonary Rehabilitation. *Journal of the Formosan Medical Association*. 2021 Jan 1;120(1):83-92.

30. Mengu S, Mengu M, Gunay K. Value-based communication during COVID-19 pandemic: a study on the twitter messages of Turkish Ministry of Health. *Essays on COVID-19 Research*. 2021:147.
31. Aytür YK, Köseoğlu BF, Taşkıran ÖÖ, Gökkaya NK, Delialioğlu SÜ, Tur BS, Sarıkaya S, Şirzai H, Tiftik TT, Alemdaroğlu E, Ayhan FF. Pulmonary rehabilitation principles in SARS-CoV-2 infection (COVID-19): The revised guideline for the acute, subacute, and post-COVID-19 rehabilitation. *Turkish journal of physical medicine and rehabilitation*. 2021 Jun;67(2):129.
32. Xie M, Chen Q. Insight into 2019 novel coronavirus—An updated interim review and lessons from SARS-CoV and MERS-CoV. *International Journal of Infectious Diseases*. 2020 May 1;94:119-24.
33. Chen L, Xiong J, Bao L, Shi Y. Convalescent plasma as a potential therapy for COVID-19. *The Lancet infectious diseases*. 2020 Apr 1;20(4):398-400.
34. Yuet-Wong C. 9th Asian Preventive Cardiology and Cardiac Rehabilitation Conference. *Journal of the Hong Kong College of Cardiology*. 2023;30(1):1-38.
35. Yuet-Wong C. 9th Asian Preventive Cardiology and Cardiac Rehabilitation Conference. *Journal of the Hong Kong College of Cardiology*. 2023;30(1):1-38.
36. Xu H, Hou K, Xu R, Li Z, Fu H, Wen L, Xie L, Liu H, Selvanayagam JB, Zhang N, Yang Z. Clinical characteristics and risk factors of cardiac involvement in COVID-19. *Journal of the American Heart Association*. 2020 Sep 15;9(18):e016807.
37. Inciardi RM, Lupi L, Zacccone G, Italia L, Raffo M, Tomasoni D, Cani DS, Cerini M, Farina D, Gavazzi E, Maroldi R. Cardiac involvement in a patient with coronavirus disease 2019 (COVID-19). *JAMA cardiology*. 2020 Jul 1;5(7):819-24.
38. Azkur AK, Akdis M, Azkur D, Sokolowska M, van de Veen W, Brüggen MC, O'Mahony L, Gao Y, Nadeau K, Akdis CA. Immune response to SARS-CoV-2 and mechanisms of immunopathological changes in COVID-19. *Allergy*. 2020 Jul;75(7):1564-81.
39. Tschöpe C, Ammirati E, Bozkurt B, Caforio AL, Cooper LT, Felix SB, Hare JM, Heidecker B, Heymans S, Hübner N, Kelle S. Myocarditis and inflammatory cardiomyopathy: current evidence and future directions. *Nature reviews cardiology*. 2021 Mar;18(3):169-93.
40. Tur BS, Köseoğlu BF, Gökkaya NK, Aytür YK, Taşkıran ÖÖ, Kabayel DD, Kesiktaş N, Tıkız C, Özdemir H, Alemdaroğlu E, Kaya BB. COVID-19, cardiac involvement and cardiac rehabilitation: Insights from a rehabilitation perspective-State of the Art. *Turkish Journal of Physical Medicine and Rehabilitation*. 2022 Sep;68(3):317.
41. EMRALINO FL. MDA5-mediated type I interferonopathy mouse model displays lethal response to immune stimulation.
42. Tur BS, Köseoğlu BF, Gökkaya NK, Aytür YK, Taşkıran ÖÖ, Kabayel DD, Kesiktaş N, Tıkız C, Özdemir H, Alemdaroğlu E, Kaya BB. COVID-19, cardiac involvement and cardiac rehabilitation: Insights from a rehabilitation perspective-State of the Art. *Turkish Journal of Physical Medicine and Rehabilitation*. 2022 Sep;68(3):317.
43. Chang JC. Sepsis and septic shock: endothelial molecular pathogenesis associated with vascular microthrombotic disease. *Thrombosis journal*. 2019 Dec;17(1):1-9.
44. Pickenhan L, Rungg C, Schiefermeier-Mach N. Electrolyte imbalances in nursing home residents: A review of prevalence, management and considerations. *The Journal of Nursing Home Research Sciences*. 2020;6:14-9.
45. Kikut J, Komorniak N, Ziętek M, Palma J, Szczuko M. Inflammation with the participation of arachidonic (AA) and linoleic acid (LA) derivatives (HETEs and HODEs) is necessary in the course of a normal reproductive cycle and pregnancy. *Journal of Reproductive Immunology*. 2020 Sep 1;141:103177.
46. Patterson BK, Francisco EB, Yogendra R, Long E, Pise A, Rodrigues H, Hall E, Herrera M, Parikh P, Guevara-Coto J, Triche TJ. Persistence of SARS CoV-2 S1 protein in CD16+ monocytes in post-acute sequelae of COVID-19 (PASC) up to 15 months post-infection. *Frontiers in immunology*. 2022 Jan 10;12:5526.
47. Tur BS, Köseoğlu BF, Gökkaya NK, Aytür YK, Taşkıran ÖÖ, Kabayel DD, Kesiktaş N, Tıkız C, Özdemir H, Alemdaroğlu E, Kaya BB. COVID-19, cardiac involvement and cardiac rehabilitation: Insights from a rehabilitation perspective-State of the Art. *Turkish Journal of Physical Medicine and Rehabilitation*. 2022 Sep;68(3):317.
48. Writing Committee, Gluckman TJ, Bhave NM, Allen LA, Chung EH, Spatz ES, Ammirati E, Baggish AL, Bozkurt B, Cornwell III WK, Harmon KG. 2022 ACC expert consensus decision pathway on cardiovascular sequelae of COVID-19 in adults: myocarditis and other myocardial involvement, post-acute sequelae of SARS-CoV-2 infection, and return to play: a report of the American College of Cardiology Solution Set Oversight Committee. *Journal of the American College of Cardiology*. 2022 May 3;79(17):1717-56.
49. Tur BS, Köseoğlu BF, Gökkaya NK, Aytür YK, Taşkıran ÖÖ, Kabayel DD, Kesiktaş N, Tıkız C, Özdemir H, Alemdaroğlu E, Kaya BB. COVID-19, cardiac involvement and cardiac rehabilitation: Insights from a rehabilitation perspective-State of the Art. *Turkish Journal of Physical Medicine and Rehabilitation*. 2022 Sep;68(3):317.



50. Writing Committee, Gluckman TJ, Bhavne NM, Allen LA, Chung EH, Spatz ES, et al. 2022 ACC expert consensus decision pathway on cardiovascular sequelae of COVID-19 in adults: Myocarditis and other myocardial involvement, post-acute sequelae of SARS-CoV-2 infection, and return to play: A report of the American College of Cardiology solution set oversight committee. *J Am Coll Cardiol* 2022;79:1717-56.
51. Naveen KV, Saravanakumar K, Sathiyaseelan A, MubarakAli D, Wang MH. Human fungal infection, immune response, and clinical challenge—a perspective during COVID-19 pandemic. *Applied Biochemistry and Biotechnology*. 2022 Sep;194(9):4244-57.
52. Burchill E, Lymberopoulos E, Menozzi E, Budhdeo S, McIlroy JR, Macnaughtan J, Sharma N. The unique impact of COVID-19 on human gut microbiome research. *Frontiers in Medicine*. 2021:267.
53. Liu Y, Kuang D, Li D, Yang J, Yan J, Xia Y, Zhang F, Cao H. Roles of the gut microbiota in severe SARS-CoV-2 infection. *Cytokine & growth factor reviews*. 2022 Jan 31.
54. Tkacheva ON, Klimenko NS, Kashtanova DA, Tyakht AV, Maytesyan LV, Akopyan AA, Koshechkin SI, Strazhesko ID. Gut Microbiome in Post-COVID-19 Patients Is Linked to Immune and Cardiovascular Health Status but Not COVID-19 Severity. *Microorganisms*. 2023 Apr 15;11(4):1036.
55. Abdel-Gawad M, Ali S, Azab M, Shawky R, Emara M. Relationship between COVID-19 and Human Gut Microbiome, Nutritional Factors, Type 2 Diabetes, and Obesity. *Journal of Advanced Pharmacy Research*. 2022 Apr 1;6(2):84-93.
56. Chatterjee P, Nirgude A, Chatterjee PK. Healthy eating—a modifiable contributor to optimize healthy living in the COVID-19 pandemic: a review. *Journal of the Science of Food and Agriculture*. 2022 Mar 30;102(5):1751-8.
57. Grenga L, Pible O, Miotello G, Culotta K, Ruat S, Roncato MA, Gas F, Bellanger L, Claret PG, Dunyach-Remy C, Laureillard D. Taxonomical and functional changes in COVID-19 faecal microbiome could be related to SARS-CoV-2 faecal load. *Environmental Microbiology*. 2022 Sep;24(9):4299-316.
58. Abdel-Gawad M, Ali S, Azab M, Shawky R, Emara M. Relationship between COVID-19 and Human Gut Microbiome, Nutritional Factors, Type 2 Diabetes, and Obesity. *Journal of Advanced Pharmacy Research*. 2022 Apr 1;6(2):84-93.
59. Fagni F, Simon D, Tascilar K, Schoenau V, Sticherling M, Neurath MF, Schett G. COVID-19 and immune-mediated inflammatory diseases: effect of disease and treatment on COVID-19 outcomes and vaccine responses. *The Lancet Rheumatology*. 2021 Oct 1;3(10):e724-36.
60. Onyeaka H, Anumudu CK, Al-Sharify ZT, Egele-Godswill E, Mbaegbu P. COVID-19 pandemic: A review of the global lockdown and its far-reaching effects. *Science progress*. 2021 May;104(2):00368504211019854.
61. Plassmeyer M, Alpan O, Corley MJ, Premeaux TA, Lillard K, Coatney P, Vaziri T, Michalsky S, Pang AP, Bukhari Z, Yeung ST. Caspases and therapeutic potential of caspase inhibitors in moderate–severe SARS-CoV-2 infection and long COVID. *Allergy*. 2022 Jan;77(1):118-29.