# The most influential COVID-19 articles: A systematic review

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#### Abstract

**Background:** Since December 2019, the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2), causative pathogen of coronavirus disease 2019 (COVID-19), has triggered a pandemic with challenges for health care systems around the world. Researchers have studied and published on the subject of SARS-CoV-2 and the disease extensively. What is the significance of articles published, shared and cited in the early stages of such a pandemic?

Materials and methods: A systematic literature search in a time frame of 12 months and analysis rating using Principle Component Analysis (PCA) and Multiple Factor Analysis (MFA) were performed.

**Results:** The 100 most cited COVID-19 articles were identified. The majority of these articles were from China (n = 54), followed by United States of America (USA) (n = 21) and United Kingdom (UK) (n = 8). All articles were published in high-ranked, peer-reviewed journals, with research focusing on the the diagnosis, transmission and therapy of COVID-19. The level of evidence of the 100 most cited COVID-19 articles on average was low.

**Conclusion:** In the early stages of a pandemic, new and innovative research can emerge and be highly cited, regardless of the level of evidence.

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#### I. Introduction

The onset of the COVID-19 pandemic was not only a test of resilience for the human race, but it also put scientists through their paces. In being a novel virus there was initially a lack of literature to aid the medical workforce; it fast became a race for

scientists to contribute to the evidence-base to guide management of unwell patients accordingly. Newly proposed treatments based on anecdotal evidence were being used across the world, however policy-makers and those treating patients on the 'front-line' were unable to rely on such data alone for assurance that these novel treatments would be best for patient care. Some countries such as the UK with NICE guidelines, heavily rely on validated and peer-reviewed evidence in order to formulate treatment guidelines and regimens.

One of the largest barriers to clinical confidence in hastily published 'COVID-19' articles is the distinct lack of high hierarchical levels of evidence. Whilst this could largely be due to the lack of time alongside intense pressure to publish research, there may also be a general lack of understanding that results

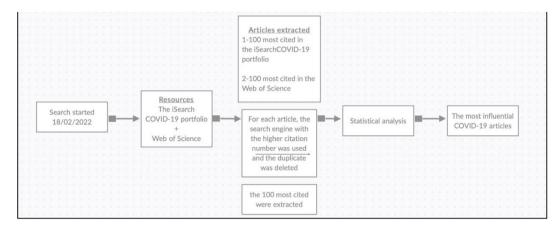


FIG. I. Flow diagram demonstrating the methodology and data extraction.

from case-studies of small sample sizes cannot be extrapolated to be true of entire populations.

This paper aims to highlight, understand and assess the top 100 most-cited articles published under the topic of COVID-19 through a systematic search using stringent inclusion and exclusion criteria. As shown in the results section, most papers originated from China (n = 54) and USA (n = 21). Difficulties with translations of Chinese papers were found to be an issue (although most were published in English), with their focus on diagnosis, mechanism, transmission and treatment, whilst Western papers focused only on transmission and treatment.

Using Principle Component Analysis (PCA) and Multiple Factor Analysis (MFA) of the filtered search results, this systematic review explores the possible correlations between objective metrics including: number of citations, density, article age, hierarchical evidence level and impact factors. Our findings suggest that pioneering evidence was published and subsequently heavily cited regardless of the level of evidence (mainly levels IV & V). We hope that this review will be of use to those contributing to the evidence base in future time-pressured scenarios such as subsequentnovel pathogen emergences.

# 2. Materials and methods

The Web of Science and the iSearch COVID-19 portfolio were utilised as effective tools for retrieval of citation information of published Covid-19 articles.

 The Web of Science provides comprehensive citation data for articles published in Medline, Web of Science Core Collection, BIOSIS Citation Index, KCI-Korean Journal Database, Russian Science Citation Index, and SciELo Citation index [1–3]. Topic fields of articles (title, abstract, author's keywords and keywords within a record) were searched for the following keywords:

"Wuhan Coronavirus" OR "Wuhan Seafood Market Pneumonia Virus" OR "COVID19" OR "COVID-19" OR "COVID-2019" OR "Coronavirus Disease 2019" OR "SARS-CoV-2" OR SARS2 OR "2019-nCoV" OR "2019 Novel Coronavirus" OR "Severe Acute Respiratory Syndrome Coronavirus 2" OR "2019 Novel Coronavirus Infection" OR "Coronavirus Disease 2019" OR "Coronavirus Disease-19" OR "Novel Coronavirus" OR "Coronavirus" OR "SARS-CoV-2019" OR "SARS-CoV-19".

 The iSearch COVID-19 portfolio is the National Institute of Health's comprehensive source for publications related to COVID-19. It demonstrates cutting-edge analytical capabilities and is updated daily.

Only COVID-related articles submitted after 31/12/2019 (first reported COVID-19 case) were included in the study and the 100 most cited articles were identified and evaluated by two independent reviewers (Fig. 1).

COVID-19 articles were classified and assigned a level of evidence.

The levels of evidence (I-V) were adapted from the National Health and Medical Research Council (NHMRC) and The Centre for Evidence-Based Medicine (CEBM) [4].

Articles were categorized, using LitCovid, by different research topics as following:

Clinical Features, Mechanism, Diagnosis, Treatment, Transmission, Prevention, Forecasting and General [5].

#### 2.1. Statistical analysis

Statistical analyses were conducted using the R programming language. Normality of data was checked using the Shapiro-

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# TABLE I. Overview of the top 100 cited COVID-19 articles (\* next to rank number indicates systematic review).

lank	Title	Citations(Density Citations/Age In months))	First Author	Last Author
	Clinical features of patients infected with 2019 novel coronavirus in Wuhan, China. Clinical Characteristics of Coronavirus Disease 2019 in China. Clinical Characteristics of Covid-19 in China	18958(790) 13699(652) 13699(652)	Chaolin Huang Wei-jie Guan Alexandre P.	Bin Cao Nan-Shan Zhong Diego R. Falci
	Clinical course and risk factors for mortality of adult inpatients with COVID-19 in	12819(537)	Zavascki Fei Zhou	Bin Cao
	Wuhan, China: a retrospective cohort study. A Novel Coronavirus from Patients with Pneumonia in China, 2019 Clinical Characteristics of 138 Hospitalized Patients With 2019 Novel Coronavirus-	2 82(530)  0228(445)	Na Zhu Dawei Wang	Wenjie Tan Zhiyong Peng
	Infected Pneumonia in Wuhan, China. Epidemiological and clinical characteristics of 99 cases	10013(417)	Nanshan Chen	Li Zhang
	of 2019 novel coronavirus pneumonia in Wuhan, China: a descriptive study A pneumonia outbreak associated with a new coronavirus of probable bat origin Characteristics of and Important Lessons From the Coronavirus Disease 2019 (COVID- 19) Outbreak in China: Summary of a Report of 72,314 Cases From the Chinese Center	9991 (434) 7855(457)	Peng Zhou Zunyou Wu	Zheng-Li Shi Jennifer M. McGoog
)	for Disease Control and Prevention. SARS-CoV-2 Cell Entry Depends on ACE2 and TMPRSS2 and Is Blocked by a Clinically Device Devices Institute for the second	7797(354)	Markus Hoffmann	Hannah Kleine-Web
	Proven Protease Inhibitor. Early Transmission Dynamics in Wuhan, China, of Novel Coronavirus-Infected	6343(288)	Qun Li	Zijian Feng
	Pneumonia. Genomic characterisation and epidemiology of 2019 novel coronavirus: implications for virus origins and receptor binding	5779(251)	Roujian Lu	Wenjie Tan
	Antibody Responses to SARS-CoV-2 in Patients With Novel Coronavirus Disease 2019	5122(320)	Juanjuan Zhao	Zheng Zheng
	Clinical course and outcomes of critically ill patients with SARS-CoV-2 pneumonia in		Xiaobo Yang	You Shang
		4651(194)	Jasper Fuk-Woo	Kwok-Yung Yuen
5	person-to-person transmission: a study of a family cluster The psychological impact of quarantine and how to reduce it: rapid review of the	4599(200)	Chan Samantha K Brooks	Gideon James Rub
	evidence Presumed Asymptomatic Carrier Transmission of COVID-19	4555(207)	Brooks Yan, Bai	Meiyun Wang
	Pathological findings of COVID-19 associated with acute respiratory distress syndrome		Zhe Xu	Fu-Sheng Wang
	COVID-19: consider cytokine storm syndromes and immunosuppression. Aerosol and Surface Stability of SARS-CoV-2 as Compared with SARS-CoV-1	4386(199) 4362(198)	Puja Mehta Neeltje Van Doremalen	Jessica J Manson Vincent J Munster
	A new coronavirus associated with human respiratory disease in China.	4309(187)	Fan Wu	Yong-Zhen Zhang
	Presenting Characteristics, Comorbidities, and Outcomes Among 5700 Patients Hospitalized With COVID-19 in the New York City Area.	4118(206)	Safiya Richardson	Karina W. Davids
	SARS-CoV-2 Viral Load in Upper Respiratory Specimens of Infected Patients	4079(185)	Lirong Zou	Jie Wu
	Cryo-EM structure of the 2019-nCoV spike in the prefusion conformation.	3912(170)	Daniel Wrapp	Jason S. McLellan
	Structure, Function, and Antigenicity of the SARS-CoV-2 Spike Glycoprotein	3750(170)	Alexadra C. Walls	
	Dexamethasone in Hospitalized Patients with Covid-19 Abnormal coagulation parameters are associated with poor prognosis in patients	3746(163) 3737(170)	Peter Horby Ning Tang	Martin J. Landray Ziyong Sun
	with novel coronaviruspneumonia Risk Factors Associated With Acute Respiratory Distress Syndrome and Death in	3705(195)	Chaomin Wu	Yuanlin Song
	Patients With Coronavirus Disease 2019 Pneumonia in Wuhan, China. The species Severe acute respiratory syndrome-related coronavirus: classifying 2019-	3560(162)	Alexander E.	John Ziebuhr
	nCoV and naming it SARS-CoV-2 Remdesivir and chloroquine effectively inhibit the recently	3473(151)	Gorbalenya Manli Wang	Gengfu Xiao
	emerged novel coronavirus (2019-nCoV) in vitro Neurologic Manifestations of Hospitalized Patients With Coronavirus Disease 2019 in Wuhan, China	3467(173)	Ling Mao	Bo Hu
	An interactive web-based dashboard to track COVID-19 in real time	3460(173)	Ensheng Dong	Lauren Gardner
	Safety and Efficacy of the BNT162b2 mRNA Covid-19 Vaccine.	3413(263)	Fernando P. Plack	
	Detection of 2019 novel coronavirus (2019-nCoV) by real-time RT-PCR	3205(134)	Victor M. Corman	
	A novel coronavirus outbreak of global health concern Virological assessment of hospitalized patients with COVID-2019.	3190(133) 3100(155)	Chen Wang Roman Woelfel	George F. Gao Camilla Rothe
	Immediate Psychological Responses and Associated Factors during the Initial Stage of the 2019 Coronavirus Disease (COVID-19) Epidemic among the General Population in		Cuiyan Wang	Roger C. Ho
	China. A Trial of Lopinavir-Ritonavir in Adults Hospitalized with Severe Covid-19	2894(134)	Bin Cao	Chen Wang
	First Case of 2019 Novel Coronavirus in the United States.	2787(121)	Michelle, Holshue	Satish K Pillai
	Remdesivir for the Treatment of Covid-19 - Final Report.	2768(185)	John H. Beigel	H. Clifford Lane
	Detection of SARS-CoV-2 in Different Types of Clinical Specimens. Correlation of Chest CT and RT-PCR Testing for Coronavirus Disease 2019 (COVID- 19) in China: A Report of 1014 Cases	2732(130) 2721(146)	Wenling Wang Tao Ai	Wenjie Tan Xia Liming
	Hydroxychloroquine and azithromycin as a treatment of COVID-19: results of an open- label non-randomized clinical trial	2718(143)	Philippe Gautret	Didier Raoult
	Endothelial cell infection and endotheliitis in COVID-19.	2718(129)	Zsuzsanna Varga	Holger Moch
	Incidence of thrombotic complications in critically ill ICU patients with COVID-19 Factors Associated With Mental Health Outcomes Among Health Care Workers	2683(141) 2550(116)	Frederikus A. Klok Jianbo Lai	Henrick Endeman Shaohua Hu
	Exposed to CoronavirusDisease 2019 Baseline Characteristics and Outcomes of 1591 Patients Infected With SARS-CoV-2 Admitted to ICUs of the Lombardy Region, Italy.	2524(120)	Giacomo Grasselli	Antonio Pesenti
	Coronavirus Disease 2019-COVID-19	2508(157)	Kuldeep Dhama	Alfonso J. Rodrigu Morales
	Clinical predictors of mortality due to COVID-19 based on an analysis of data of 150 patients from Wuhan, China	2491(119)	Qiurong Ruan	Song JX
	The Incubation Period of Coronavirus Disease 2019 (COVID-19) From Publicly Reported Confirmed Cases: Estimation and Application	2442(116)	Stephen A. Lauer	Justin Lessler
	The Epidemiological Characteristics of an Outbreak of 2019 Novel Coronavirus Diseases (COVID-19) - China, 2020	2427(106)	Zijian Feng	Jennifer M. McGoo
	Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) and coronavirus disease-2019 (COVID-19): The epidemic and the challenges	2396(104)	Chih-Cheng Lai	Po-Ren Hsueh
		2366(131)	Chuan Qin	Dai-Shi Tian
	Dysregulation of Immune Response in Patients With Coronavirus 2019 (COVID-19) in Wuhan, China	()	Cinduan Quin	

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#### TABLE I. Continued

Rank	Title	Citations(Density Citations/Age In months))	First Author	Last Author
55 56	Efficacy and Safety of the mRNA-1273 SARS-CoV-2 Vaccine. Comorbidity and its impact on 1590 patients with COVID-19 in China: a nationwide analysis	2225(93) 2223(106)	Lindsey R. Baden Wei-jie Guan	Tal Zaks Jian-Xing He
57 58	Structural basis for the recognition of SARS-CoV-2 by full-length human ACE2. Pulmonary Vascular Endothelialitis, Thrombosis, and Angiogenesis in Covid-19.	2221(101) 2182(128)	Renhong Yan Maximilian	Qiang Zhou Danny Jonigk
59 60 *61	Transmission of 2019-nCoV Infection from an Asymptomatic Contact in Germany Case-Fatality Rate and Characteristics of Patients Dying in Relation to COVID-19 in Italy The proximal origin of SARS-CoV-2	2133(93) 2083(99) 2068(94)	Ackermann Camilla Rothe Graziano Onder Kristian G. Andersen	Michael Hoelscher Silvio Brusaferro Robert F. Garry
*62	The epidemiology and pathogenesis of coronavirus disease (COVID-19) outbreak	2062(98)	Hussin A. Rothan	Siddappa N. Byrareddy
63 64	Clinical and immunological features of severe and moderate coronavirus disease 2019. Structure of the SARS-CoV-2 spike receptor-binding domain bound to the ACE2 receptor.	2057(98) 2056(98)	Guang Chen jun Lan	Qin Ning Xinquan Wang
65	Receptor Recognition by the Novel Coronavirus from Wuhan: an Analysis Based on Decade-Long Structural Studies of SARS Coronavirus.	2048(93)	Yushun Wan	Fang Li
66	Nowcasting and forecasting the potential domestic and international spread of the 2019- nCoV outbreak originating in Wuhan, China: a modelling study	2003(87)	Joseph T. Wu	Gabriel M. Leung
67	Association of Cardiac Injury With Mortality in Hospitalized Patients With COVID-19 in Wuhan, China.	2000(105)	Shaobo Shi	Congxin Huang
68 *69	Epidemiology of COVID-19 Among Children in China World Health Organization declares global emergency: A review of the 2019 novel coronavirus (COVID-19)	1996(100) 1972(90)	Yuanyuan Dong Catrin Sohrabi	Shilu Tong Riaz
70	Clinical characteristics of 113 deceased patients with coronavirus disease 2019: retrospective study.	1972(90)	Tao Chen	Qin Ning
71	Critical Care Utilization for the COVID-19 Outbreak in Lombardy, Italy Early Experience and Forecast During an Emergency Response	1959(93)	Giacomo Grasselli	Maurizio Cecconi
72	Multidisciplinary research priorities for the COVID-19 pandemic: a call for action for mental health science	1948(97)	Emily A. Holmes	Ed Bullmore
*73	The origin, transmission and clinical therapies on coronavirus disease 2019 (COVID-19) outbreak - an update on the status	1927(84)	Yan-Rong Guo	Yan Yan
74		1924(101)	Tao Guo	Zhibing Lu
75	Clinical characteristics and intrauterine vertical transmission potential of COVID- 19 infection in nine pregnant women: a retrospective review of medical records	1904(83)	Huijun Chen	Yuanzhen Zhang
76	Factors associated with COVID-19-related death using OpenSAFELY	1886(105)	Elizabeth J. Williamson	Ben Goldacre
77 78	Temporal dynamics in viral shedding and transmissibility of COVID-19. Anticoagulant treatment is associated with decreased mortality in severe coronavirus disease 2019 patients with coagulopathy.	879(89)   837(87)	Xi He Ning Tang	Gabriel M. Leung Ziyong Sun
*79 *80 81 82	The socio-economic implications of the coronavirus pandemic (COVID-19): A review The trinity of COVID-19: immunity, inflammation and intervention Clinical characteristics of 140 patients infected with SARS-CoV-2 in Wuhan, China. Temporal profiles of viral load in posterior oropharyngeal saliva samples and serum	1738(87) 1719(86) 1718(90) 1715(82)	Maria Nicola Mathew Zirui Tay Jin-Jin Zhang Kelvin Kai-Wang	Riaz Agha Lisa F.P. NG Ya-dong Gao Kwok-Yung Yuen
83	antibody responses during infection by SARS-CoV-2: an observational cohort study. Remdesivir in adults with severe COVID-19: a randomised, double-blind, placebo-	1714(82)	To yeming Wang	Chen Wang
84	controlled, multicentre trial. Radiological findings from 81 patients with COVID-19 pneumonia in Wuhan, China: a description study.	1660(76)	Heshui Shi	Chuansheng Zheng
85	descriptive study. Imbalanced Host Response to SARS-CoV-2 Drives Development of COVID-19.	1648(82)	Daniel Blanco- Melo	Benjamin R. tenoever
86 87	A SARS-CoV-2 protein interaction map reveals targets for drug repurposing. Characteristics and Outcomes of 21 Critically III Patients With COVID-19 in Washington State	633(86)  632(78)	David E. Gordon Matt Arentz	Nevan J. Krogan Melissa Lee
88	Clinical management of severe acute respiratory infection (SARI) when COVID-19 disease is suspected. Interim guidance	1609(89)	Caroline Quach- Thanh	Titus Yeung
*89	Prevalence of comorbidities and its effects in patients infected with SARS-CoV-2: a systematic review and meta-analysis.	1597(76)	Jing Yang	Yonging Zhou
90		1592(72)	Jianyin Qiu	Yifeng Xu
91	Tracking Changes in SARS-CoV-2 Spike: Evidence that D614G Increases Infectivity of the COVID-19 Virus.	1560(92)	Bette Korber	David C. Montefiori
*92 *93 *94	COVID-19 and the cardiovascular system The psychological impact of the COVID-19 epidemic on college students in China WHO Declares COVID-19 a Pandemic.	1555(74) 1503(74) 1512(69)	Ying-Ying Zheng Wenjun Cao Domenico Cucinotta	Xiang Xie Jianzhong Zheng Maurizio Vanelli
95 96 97	Structural basis of receptor recognition by SARS-CoV-2. Estimates of the severity of coronavirus disease 2019: a model-based analysis. Targets of T Cell Responses to SARS-CoV-2 Coronavirus in Humans with COVID-19 Disease and Unexposed Individuals.	1491(65) 1473(74) 1462(77)	Jian Shang Robert Verity Alba Grifoni	Fang Li Neil M. Ferguson Alessandro Sette
98	Substantial undocumented infection facilitates the rapid dissemination of novel coronavirus (SARS-CoV-2).	1424(68)	Ruiyun Li	Jeffrey Shaman
99 100	Antibody responses to SARS-CoV-2 in patients with COVID-19. Compassionate Use of Remdesivir for Patients with Severe Covid-19.	1410(71) 1405(70)	Quan-Xin Lo Jonathan Grein	Ai-Long Hunag Timothy Flanigan

Wilk test. The distribution of a parameter was characterised by the median and interquartile range. The Kendall rank correlation coefficient was used to measure the ordinal association between two values. Multiple Factor Analysis (MFA) was used to analyse quantitative variables simultaneously. A *P*-value of <0.05 was considered statistically significant. Microsoft Excel software was used for descriptive statistical analysis.

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\*Any disagreements between the reviewers were resolved via consensus.

#### 3. Results

Table I gives an overview of the top 100 most cited articles on COVID-19. All articles were published in 2020 (100%). The highest number of citations was 18958 and the lowest number was 1410. The median age of the articles was 21 months (range 13-24). In terms of levels of evidence - 14 articles were evidence level I, 7 were level II, 12 were III, 45 were IV and 22 were V.

13 articles were basic science, 2 case control studies, 4 case reports, 32 case series, 3 clinical consensus articles, 12 cohort studies, 8 cross sectional studies, 5 expert opinions, 7 randomised controlled trials and 14 systematic reviews (Fig. 2).

The Top 100 cited COVID-19 originated from 13 countries (Fig. 3), of which more than half were from China (n = 54), followed by United States of America (USA) (n = 21); United Kingdom (UK) (n = 8); Germany (n = 4); Italy (n = 4); Netherlands (n = 2); Brazil (n = 1); Canada (n = 1); Colombia (n = 1); France (n = 1); Singapore (n = 1); Switzerland (n = 1) and Taiwan (n = 1).

Table 2 shows the journals in which the top 100 cited COVID-19 articles were published with accompanying journal metrics. Of the 100 cited COVID-19 articles, 13 were published in the New England Journal of Medicine (NEJM), followed by 21 in the Lancet or associated journals (The Lancet (n = 13), Lancet Infectious Disease (n = 4), Lancet Respiratory medicine (n = 2), Lancet Psychiatry (n = 1), Lancet Oncology (n = 1)); 13 in Nature or associated journal (Nature (n = 7), Nature Medicine (n = 3), Nature Microbiology (n = 1), Nature Reviews Cardiology (n = 1), Nature Reviews Immunology (n = 1)); 13 in

JAMA or associated journals (JAMA (n = 8), JAMA Neurology (n = 1), JAMA Network Open (n = 1), JAMA Cardiology (n = 2), JAMA Internal Medicine (n = 1)), 4 in Cell and 3 articles in Science.

Articles published in China concentrated on the diagnosis, mechanism, transmission and treatment of COVID-19. On the other hand, articles published in Europe and the USA mainly focused on the transmission and the treatment of the virus (Table 3).

Principle Component Analysis (PCA) revealed a strong correlation between the number of citations and the citation density of citations. Furthermore, there was also a strong correlation between the age of the article, the level of the evidence and the impact factor. There was a significant trend towards increased frequency of citations with age of the article (r = 0.26, P = 0.0004). The number of citations an article had was not significantly associated with the level of evidence (r = 0.152, p = 0.152) (Fig. 4).

### 4. Discussion

This systematic review identified the 100 most cited articles on Covid-19 and sought to identify trends within them by applying citation analysis techniques. In late 2019, the COVID-19 pandemic presented one of the greatest challenges of the modern scientific era. With an estimated 503,862 deaths worldwide reported within the first 6 months of 2020 [6], the gravity and urgency of the problem required rapid advancement in knowledge to a degree not previously seen. It is unsurprising that with the amount of funding and resource invested, great volumes of scientific literature were produced in a relatively short period of time. What is surprising is the degree to which this occurred. Despite the first case of COVID-19 being only

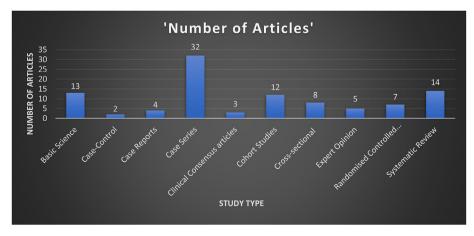


FIG. 2. The hundred most cited Covid-19 articles study types.

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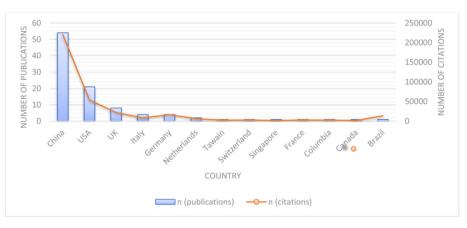


FIG. 3. Number of publications and citations per country.

reported in December 2019, by July 2020 over 27,000 COVID-19 related articles had been published [7], with Covid related articles accounting for more scientific publication then all other topics combined. This unprecedented level of publication provokes many questions around the quality of research and the readiness of article acceptance [8-12].

Journals	Articles	Median Citation	Median Citation density	Country	Quartile (Impact factor)	Categor Rank
Lancet Respiratory Medicine	2 Articles (2 level IV)	4665	217	UK	30.7	QI
The Lancet	13 Articles (1 level I, 2 level II, 1 level III, 6 level IV, 3 level V)	4599	199	England	79.323	QI
Clinical Infection Diseases	2 Articles (I level III, I level IV)	3744	226	USA	9.079	QI
AMA Internal Medicine	I Article (I level III)	3705	195	USA	21.873	QΪ
New England Journal of Medicine	16 articles (5 level II, 9 level IV, 2 Level V)	3580	185	USA	91.253	οÌ
Nature Microbiology	I Article (I level V)	3560	162	UK	17.745	οÌ
Cell Research	I Article (I level V)	3473	151	UK	25.617	õi
IAMA Neurology	I Article (I level IV)	3467	173	USA	18.302	Qi
Eurosurveillance	I Article (I level V)	3205	134	Sweden	6.307	Qi
International Journal of Enviromental Research and Public Health		2829	123	Switzerland	3.39	QI
Journal of Thrombosis and Haemostasis	2 Articles (1 level III, 1 level IV)	2787	129	UK	5.824	QI
Radiology	I Article (I level IV)	2721	151	USA	11.105	õi
Thrombosis Research	I Article (I level IV)	2683	141	USA	3.944	õi
Journal of the American Medical Association	8 Articles (6 level IV, 2 level 5	2628	125	USA	56.274	õi
International Journal of Antimicrobial Agents	2 Articles (1 level I, 1 level II)	2557	125	Netherlands		Qi
AMA Network Open	I Article (I level IV)	2550	126	USA	8.485	Qi
Clinical Microbiology Reviews	I Article (I level I)	2508	157	USA	26.132	õi
Intensive Care Medicine	I Article (I level IV)	2491	119	USA	17.44	õi
Annals of Internal Medicine	I Article (I level IV)	2442	116	USA	25.391	õi
China CDC Weekly	I Article (I level IV)	2427	106	China	23.371 Na	Na
		2263	98		41.316	
Lancet Oncology	I Article (I level III)	2223	106	England		QI
The European Respiratory Journal Science	Article (Llevel III)	2223	106	England	16.671 47.728	QI
	3 Articles (1 level IV, 2 level V)			USA		QI
Journal of Autoimmunity	I Article (I level I)	2062	98	UK	7.094	QI
Journal of Clinical Investigation	I Article (I level IV)	2057	98	USA	14.808	QI
Nature	7 Articles (1 level III, 3 level IV, 3 level V)	2056	105	UK	49.962	QI
Journal of Virology	I Article (I level IV)	2048	93	USA	5.103	QI
Paediatrics	I Article (I level IV)	1996	100	USA	7.125	QI
British Medical Journal	I Article (I level IV)	1972	90	UK	14.093	QI
JAMA Cardiology	2 Articles (I level III, I level IV)	1962	103	USA	14.676	QI
Lancet Psychiatry	I Article (I level V)	1948	97	UK	26.481	QI
Military Medical Research	Article (  level  )	1927	84	UK	3.329	Q2
Nature Medicine	3 Articles (I level I, I level III, I Level IV)	1879	89	USA	53.44	QI
International Journal of Surgery	2 Articles (2 level II)	1855	88	UK	6.071	QI
Nature Reviews Immunology	Article (  level  )	1719	86	UK	53.106	QI
Allergy	I Article (I level IV)	1718	90	Denmark	13.146	QI
Lancet Infectious Diseases	4 articles (2 level III, 1 level IV, 1 level V)	1689	79	USA	25.071	QI
Pediatria i Medycyna Rodzinna	I Article (I level V)	1609	89	Poland	0.07	Q4
Cell	4 articles (1 level III, 3 level V)	1604	87	USA	41.584	QI
International journal of Infectious Diseases	Article (  level  )	1597	76	Canada	3.623	Q2
General Psychiatry	I Article (I level IV)	1592	72	UK	2	Q3
Nature Reviews Cardiology	I Article (I level I)	1555	74	UK	32.43	QI
Psychiatry research	l Article (l level l)	1553	74	UK	3.222	Q2
Acta Biomedica	I Article (I level I)	1512	69	Italy	1.35	03

TABLE 2. Journals in which top 100 cited COVID-19 articles were published with accompanying journal metrics.

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#### TABLE 3. Articles topic field.

Field	Country	Number of Articles
Clinical Features	Brazil	1
Diagnosis	China	6
Diagnosis	Germany	1
Diagnosis	Italy	1
Diagnosis	USÁ	1
Diagnosis AND Mechanism	China	7
Diagnosis AND Mechanism	USA	1
Diagnosis AND Prevention	China	1
Diagnosis AND Transmission	China	2
Diagnosis AND Transmission	USA	1
Diagnosis AND Treatment	China	2
Diagnosis AND Treatment	UK	1
Diagnosis AND Treatment	USA	1
Forecasting	China	1
Forecasting	Italy	1
Forecasting	UK	1
Forecasting AND Prevention AND Transmission	China	1
Forecasting AND Transmission	USA	1
General	China	6
General	UK	3
General	Italy	2
General	Netherlands	1
General	USA	1
Mechanism	China	8
Mechanism	USA	2
Mechanism	China	1
Mechanism	Switzerland	1
Mechanism AND Diagnosis	Germany	1
Mechanism AND Diagnosis AND Treatment AND	Canada	1
Transmission AND Prevention Mechanism AND Diagnosis AND Treatment AND Transmission AND Prevention	Columbia	I
Mechanism AND Diagnosis AND Treatment AND	UK	1
Transmission AND Prevention Mechanism AND Diagnosis AND Treatment AND Transmission AND Prevention	USA	I
Mechanism AND Prevention	China	1
Mechanism AND Transmission	China	2
Mechanism AND Treatment	USA	7
Mechanism AND Treatment	China	4
Mechanism AND Treatment	Germany	i
Mechanism AND Treatment	Singapore	i
Mechanism AND Treatment	UK	1
Mechanism AND Treatment AND Transmission	China	1
Transmission	China	2
Transmission	Germany	1
Transmission	USA	1
Transmission AND Prevention	China	1
Treatment	China	8
Treatment	USA	3
Treatment	France	i.
Treatment	Netherlands	1
Treatment	UK	1
Treatment	USA	1
Treatment AND Transmission AND Prevention	Taiwan	1

All the articles were published in 2020 with a mean article age of 21 months (range 13–24 months). There was a weak but significant association between age and citation number; as citations take time to accumulate and consequently more recently published articles may not yet have achieved sufficient citations to have entered the review. The weakness of this relationship is likely a result of the short time-frame over which the articles have accrued the citations. The most highly cited article has been cited 18958 times and had a citation density of 790, the median citations was 2434.5 (IQR 1989.5–3749.0) and median citation density was 117.5 (IQR 89.5–185.2). This is particularly impressive as a variety of other citation classics have reported significantly lower median citations despite covering time periods of many years [2,13–16]. On average a

journal article will peak in citation density approximately 3 years after publication [17] which presents a potential problem in applying citation analysis to a novel and rapidly evolving field. The strong correlation between density and citation number combined suggests that highly cited articles continue to be cited and may be establishing 'authority' status. Given the ongoing expansion in literature there is a risk that articles, considered powerful by traditional metrics, may already be scientifically out of date but not yet past their peak in terms of citation accrual.

54% of the articles originated from China which is unusual for citation classics reviews. Similar reviews on other topics tend to draw most of their articles from the USA [1,2,13-15]. This is likely explained by early geographic distribution of cases which would have granted a significant advantage for Chinese-based labs, resulting in earlier publication and thus citation accrual. Interestingly the USA provided almost half of the remaining articles, which allowing for the above explanation is in keeping with what would be expected. The early geographical distribution of cases may also explain diagnosis playing a significant role in articles from China but not from the rest of the world.

Articles representing level IV and V levels of evidence account for 67% of those identified. Whilst citation classics often demonstrate the inclusion of the lowest levels of evidence, it is seldom to this degree. For example, a review into general medical articles found 38% of articles were drawn from the lowest two levels [1] and another review into GI surgery 44% [2]. Only 7 RCTs were identified which is significantly lower than what would have been expected. It must be considered that higher levels of evidence such as RCTs (and systematic reviews of these) can take many months to conduct. It is probable that the lack of high-level evidence, and the overrepresentation of lower levels of evidence, is partially a result of the literature not yet reaching maturity. Another interesting finding of this review is the degree to which high impact factor journals are publishing low levels of evidence. It has been previously shown that in the top three general medical journals (The Lancet, New England Journal of Medicine and Journal of the American Medical Association) the level of evidence represented by an article regarding COVID-19 was significantly lower when compared to both contemporary and historic controls [9].

The main limitation of this review is the time at which it was conducted; this makes comparisons to similar reviews of different topics difficult. Due to the short publication span of the papers the definition of citation density had to be modified, using a reference period of a month rather than a year. It is likely that as the literature around COVID-19 matures trends in publications will change. It is possible that in the early stages of an emerging topic traditional citation metrics may not be the

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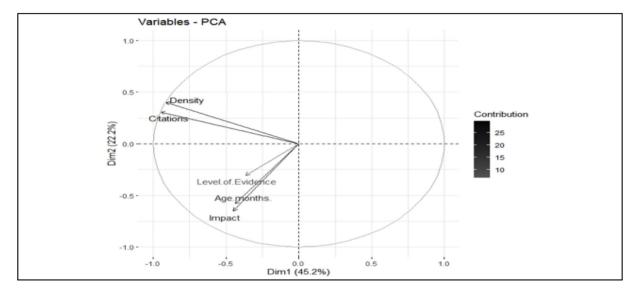


FIG. 4. Principal component analysis of the relationship between citation number, citation density, level of evidence and Impact.

most reliable way of identifying the most influential research in the longer term. Presence on social media may play an important role in identification of future influential articles; number of tweets within the first 7 days of a publication are shown to correlate with high levels of citation [18]. The simple and easily repeatable methods of this review, however, allow for later comparative review to examine how these trends have changed.

# 5. Conclusion

This review has collated the 100 most influential COVID-19 papers and assessed trends within them. We have established that in the early phases of a pandemic new and ground-breaking research surfaces regardless of the evidence level and can gain high levels of citation.

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#### **Declaration of competing interest**

The authors declared no conflict of interest.

# **CRediT** authorship contribution statement

Suhaib JS. Ahmad: Conceptualization, Methodology, Literature search, Data curation, Project administration, Writing original draft, Writing - review & editing, Validation, Final approval. Konstantinos Degiannis: Methodology, Writing original draft, Writing - review & editing, Validation, Final approval. Joseph Borucki: Methodology, Writing - original draft, Writing - review & editing, Validation, Final approval. Sjaak Pouwels: Conceptualization, Methodology, Writing original draft, Writing - review & editing, Validation, Final approval. David Laith Rawaf: Methodology, Writing - original draft, Writing - review & editing, Validation, Final approval. Marion Head: Methodology, Writing - original draft, Writing - review & editing, Validation, Final approval. Chun Hei Li: Methodology, Data curation, Visualization, Writing - original draft, Writing - review & editing, Validation, Final approval. Rami Archid: Methodology, Writing - original draft, Writing - review & editing, Validation, Final approval. Ahmed R. Ahmed: Methodology, Writing - original draft, Writing review & editing, Validation, Final approval. Anil Lala: Methodology, Writing - original draft, Writing - review & editing, Validation, Final approval. Wasif Raza: Writing - original draft, Writing - review & editing, Validation, Final approval. Katie Mellor: Methodology, Writing - original draft, Writing - review & editing, Validation, Final approval. Doerte Wichmann: Writing - original draft, Writing - review & editing, Validation, Final approval. Aristomenis Exadaktylos: Conceptualization, Methodology, Supervision, Project

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#### References

- [1] Ahmad SJ, Ahmed AR, Kowalewski KF, Nickel F, Rostami K, Stocker CJ, Hakky SM, Archid R, McWhinnie D, Mohajer-Bastami A, Seimenis DS, Ahmad S, Mansour S, Ahmed MH, Mital D, Exadaktylos AK. Citation classics in general medical journals: assessing the quality of evidence; a systematic review. Gastroenterol Hepatol Bed Bench 2020;13(2):101–14. Spring.
- [2] Ahmad SJS, Ahmed AR, Exadaktylos AK, McWhinnie D, Nickel F, Hakky SM, Ramdin A, Müller PC. Systematic review on citation classics in minimally invasive gastrointestinal surgery. J Minimal Access Surg 2018 Oct-Dec;14(4):265–72.
- [3] Ahmad SS, Ahmad SS, Kohl S, Ahmad S, Ahmed AR. The hundred most cited articles in bariatric surgery. Obes Surg 2015 May;25(5):900-9. https://doi.org/10.1007/s11695-014-1542-1.
- [4] Straus S, Glasziou W, Richardson S, Haynes RB. Evidence based medicine: how to practice and teach it. fourth ed. Edinburgh, UK: Churchill Livingstone; 2011.
- [5] Chen Q, Allot A, Lu Z. LitCovid: an open database of COVID-19 literature. Nucleic Acids Res 2021;49(D1):D1534-40.
- [6] Organisation WH. Coronavirus disease (COVID-19) situation report - 162 (30th june 2020). 2020.
- [7] Wang J, Hong N. The COVID-19 research landscape: measuring topics and collaborations using scientific literature. Medicine 2020;99(43): e22849-e.
- [8] London Alex J, Kimmelman J. Against pandemic research exceptionalism. Science 2020;368(6490):476–7.

- [9] Zdravkovic M, Berger-Estilita J, Zdravkovic B, Berger D. Scientific quality of COVID-19 and SARS CoV-2 publications in the highest impact medical journals during the early phase of the pandemic: a case control study. PLoS One 2020;15(11):e0241826.
- [10] Jung RG, Di Santo P, Clifford C, Prosperi-Porta G, Skanes S, Hung A, et al. Methodological quality of COVID-19 clinical research. Nat Commun 2021;12(1):943.
- [11] Locher C, Moher D, Cristea IA, Naudet F. Publication by association: how the COVID-19 pandemic has shown relationships between authors and editorial board members in the field of infectious diseases. BMJ Evid Based Med 2022 Jun;27(3):133–6.
- [12] Horbach SPJM. Pandemic publishing: medical journals strongly speed up their publication process for COVID-19. Quantitative Sci Stud 2020;1(3):1056-67.
- [13] Brandt JS, Hadaya O, Schuster M, Rosen T, Sauer MV, Ananth CV. A bibliometric analysis of top-cited journal articles in obstetrics and gynecology. JAMA Netw Open 2019;2(12):e1918007-e.
- [14] Hennessey K, Afshar K, Macneily AE. The top 100 cited articles in urology. Canadian Urological Association journal = Journal de l'Association des urologues du Canada 2009;3(4):293–302.
- [15] Loomes DE, van Zanten SV. Bibliometrics of the top 100 clinical articles in digestive disease. Gastroenterology 2013;144(4). 673-6.e5.
- [16] Yoon DY, Yun EJ, Ku YJ, Baek S, Lim KJ, Seo YL, et al. Citation classics in radiology journals: the 100 top-cited articles, 1945-2012. AJR Am J Roentgenol 2013;201(3):471–81.
- [17] Margolis J. Citation Indexing and Evaluation of Scientific Papers: the spread of influence in populations of scientific papers may become a subject for quantitative analysis. Science 1967;155(3767):1213-9.
- [18] Eysenbach G. Can tweets predict citations? Metrics of social impact based on Twitter and correlation with traditional metrics of scientific impact. J Med Internet Res 2011;13(4):e123-.