

Public perceptions of the association between drug effectiveness and drug novelty in France during the COVID-19 pandemic

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- 1 Public perceptions of the association between drug effectiveness and drug novelty in
- 2 France during the COVID-19 pandemic
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21 Abstract

- 22 <u>Objectives:</u>
- 23 During the COVID-19 pandemic, public debates overtly addressed the promises of new
- 24 innovative treatments. Many of these debates pitted those who advocated for the
- 25 development of new treatments by pharmaceutical companies against those who favored

- 26 the repositioning of existing drugs. Our study explored perceptions of the association
- 27 between drug novelty and effectiveness as well as perceptions of the role of the
- 28 pharmaceutical industry in drug development.
- 29 Methods
- 30 Data were collected in January 2021 from a quota sample of the French population aged
- 31 18–75 years (N = 1,000) during the second round of the "Health Literacy Survey 2019"
- 32 (HLS₁₉).
- 33 Results
- We tested the hypothesis that individuals with a high level of familiarity with the health care
- 35 system and those with a high level of trust in institutions are more likely to agree that new
- 36 drugs are more effective than old ones and that drug development should be driven by the
- 37 pharmaceutical industry. A quarter (25%) of respondents agreed that new drugs are always
- 38 more effective than old ones. Agreement with this statement was stronger among
- 39 respondents with a high level of familiarity with the health care system (as measured by the
- 40 Navigational Health Literacy score, OR 3.34 [2.13-5.24]) and among those with a high level
- 41 of trust in pharmaceutical companies or politicians. A high level of trust in pharmaceutical
- 42 companies was reported by 42% of respondents, and 43% agreed that drug development
- 43 should be driven by the pharmaceutical industry. Respondents who agreed that new drugs
- 44 are always more effective than old ones were almost four times more likely to agree that
- 45 drug development should be driven by the pharmaceutical industry (OR 3.85 [2.76-5.39]).
- 46 Conclusion
- 47 A better understanding of public attitudes towards new treatments is needed to elucidate
- 48 individual preferences in health care and their consequences on health behavior.
- 49 **Keywords:** drug innovation; COVID-19; public attitude; sociology

Introduction

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During the COVID-19 pandemic, the development of new treatments, including vaccines, became a global priority[1,2]. Innovative treatments were designed, manufactured, and tested at an accelerated pace. They were also overtly discussed in the media as they became available, sometimes even before their efficacy was robustly validated in clinical trials. Some of the innovations that garnered media attention include mRNA vaccines and the monoclonal antibodies administered to former president Donald Trump for his COVID-19 disease. Public debates on how to find a suitable cure pitted two camps against each other[2-4]. On the one side were those who advocated for the development of new treatments (such as Remdesivir [5]) by pharmaceutical companies. On the other were those who favored the repositioning of existing drugs on the grounds that they are massively available at low cost and that their safety has already been demonstrated. In the particular context of France, the controversy over the effectiveness of hydroxychloroquine against COVID-19 increased the demand for the drug, which carried the risk of adverse effects[6,7] and led to drug shortages for other indications[8]. The following question was thus brought to the forefront of public debates: Are new drugs more effective than old ones for the treatment of COVID-19? Insofar as public perceptions of the association between drug novelty and drug effectiveness affect health behavior and drive medical innovation – as illustrated by the issue of vaccine hesitancy[9,10] – it is important that we better understand these perceptions, especially in a social pharmacology perspective [11]. The success of modern medicine has largely hinged on the development of medical technologies and on their diffusion in the general population. As a result, medical innovation has become central to how modern medicine is perceived and discussed in the media[12,13]. Expectations for better and more effective treatments, which underpin health and innovation policies[14], are based on trust in scientific and technological

progress[15,16]. To fulfill those expectations, most modern democracies have adopted what can be described as the standard model of innovation in medicine. According to this model, academic and clinical researchers and actors of the pharmaceutical industry must collaborate to achieve new effective therapeutic solutions under the regulation of state agencies[17–20].

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The standard model of innovation, however, is increasingly being criticized, both within the medical profession and in the public space. Studies have demonstrated that novelty does not equal better effectiveness, with only 25% of tested drugs showing increased therapeutic effect[21]. Both the "me-too" marketing model (i.e. the commercialization of a drug that is similar to a pre-existing drug) and economic incentives to market new products appear to have affected the relationship between drug effectiveness and drug novelty[22]. The acceleration of drug approval by regulatory agencies, in particular through mechanisms like the Orphan Drug Designation and PRIME programs put in place by the European Medicines Agency, have weakened the requirement for proof of effectiveness[23]. In this context, physicians have begun publicizing the fact that new drugs are not necessarily better than old ones and that their higher prices are not always justified[24]. Biomedical research, especially when it is clearly driven by the pharmaceutical industry[25], is regularly criticized in the public space, with some pointing out the unknown adverse effects of new drugs or vaccines[26] and others highlighting potential conflicts of interest[27,28]. In particular, the rising cost of innovative oncological treatments has been criticized as unjustified in view of gains in effectiveness that could be considered as marginal or minor[29–31]. Unfortunately, while public criticisms can contribute to a better regulation of the pharmaceutical industry[32], they can also reinforce science-related populism[33].

The publicity given to medical controversies is not without consequences, as was made clear during the COVID-19 pandemic. The violent debate over hydroxychloroquine had a marked impact on health behaviors and attitudes[34,35] and on the production of scientific knowledge. Traditional strategies for conducting clinical research were called into question,

slowing down enrollment in clinical trials such as Discovery (NCT04315948) and Recovery (NCT04381936). The debate also had lasting consequences on public health policy in countries such as Brazil, precluding the adoption of much-needed prevention strategies[36]. As such, it raised broader questions regarding the role of science and research in society[37].

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Although medical innovation is central to promises of progress in health, few public health or social sciences studies have explored public perceptions of the association between drug novelty and drug effectiveness. The few available data on the topic appear contradictory. Some studies have shown that branded drugs are perceived more favorably than generic ones[38], with a specific effect of labelling a treatment as 'new' [22]. Others have found that older drugs tend to be perceived as safer and more effective than new ones, except in the case of some innovative oncological treatments[39]. On a different note, a study on patient preferences has suggested that perceptions of novelty are associated with specific psychological characteristics[40]. Studies focusing on the food industry have shown that trust in institutions has a positive impact on perceptions of new food products[41]. However, given the specificities of the health sector, public perceptions of innovation in other domains can hardly be extrapolated to it[42]. Most social science studies examining public perceptions of innovation have focused on attitudes towards science in general and not towards the medical sector. Nevertheless, these studies do provide context for perceptions of medical innovation, with some stressing the effect of overall trust in institutions and others highlighting the pervasiveness of suspicion of corporate influence in medicine[27,43]. Again, vaccine hesitancy again provides a perfect illustration of these issues[10].

Our study aimed to explore perceptions of the association between drug novelty and effectiveness from a social science perspective. Data were collected from a representative sample of the French population via an online survey in January 2021. More specifically, we tested the hypothesis that positive perceptions of medical innovation correlate with greater acceptance of the standard model of innovation underlying medical institutions (H1).

Following our main hypothesis H1, individuals with a high level of familiarity with the health care system and those with a high level of trust in institutions (doctors, scientists, politicians, and/or pharmaceutical companies) are more likely to agree that new drugs are more effective than old ones and that drug development should be driven by the pharmaceutical industry. Two subsidiary hypotheses were tested, namely that individuals with a more direct and practical experience of the health care system due to illness or ageing (H2a) and those with low socio-economic status (H2b) are less likely to believe that drug novelty and drug effectiveness are associated. Given the ongoing pandemic and massive media exposure of biomedical research[3], we also tested the hypothesis that COVID-19 has had a specific effect on perceptions of the association between drug effectiveness and drug novelty (H3).

Methodology

Design and sample

Data were collected the first week of January 2021 from a sample of the French population aged 18–75 years (N = 1000) as part of the second round of the cross-sectional online "Health Literacy Survey 2019" (HLS₁₉ - International Health Literacy Population Survey 2019-2021 from M-POHL). The questionnaire administered during the second round of the survey contained questions – on household income and on perceptions of the role of the pharmaceutical industry in drug development (see *infra*) – that were not in the questionnaire administered during the first round (in June 2020). We made sure that there were no significant differences in responses to the questions that were common to both questionnaires.

Participants were selected from a nationally representative French household research panel developed by the survey research firm IPSOS (Paris, France). A total of 23,813 individuals were initially invited by email to fill out the questionnaire. Quota sampling was used to match the French general population with regards to gender, age, area of residence (rural *vs.* urban) and population density in the region of residence (as per official census

data). The largest difference between theoretical quotas and our sample was -1.3% for rural area of residence (21.1% of the effective sample compared to 22.4% of the theoretical sample). Collected data were weighed according to participants' demographic profile to match the French general population – which explains why the number of respondents is sometimes written with decimals. The study was approved by the Ethics Committee of the French national biomedical research Institute INSERM (CEEI, IRB 00003888, 2020/04/04).

Data collection

Participants responded to the online self-administered questionnaire after providing informed consent. The questionnaire focused on five main topics: (1) demographic characteristics (gender, age, level of education, region of residence, household income, financial difficulties, occupation, current health condition); (2) health literacy, health information-seeking behavior, and ability to navigate the health care system; (3) perception of and familiarity with clinical research; (4) trust in institutions (doctors, scientists, politicians, and/or pharmaceutical companies); and (5) knowledge and concerns about the COVID-19 epidemic. The first two topics were shared with HLS₁₉ European survey when the last three were specific to the French survey.

The present study focuses specifically on responses to the following two statements: "New drugs are always more effective than old ones" and "Drug development should be driven by the pharmaceutical industry." Response options were "Completely agree," "Somewhat agree," "Somewhat disagree," and "Completely disagree."

Statistical Analysis

Several variables were recoded to ensure comparability and interpretability: "Educational level" was recoded into three groups and "age" was recoded into four groups. Four-point

scales were dichotomized into "yes" vs. "no" for the following variables: "trust in doctors," "trust in politicians," "trust in scientists," "trust in pharmaceutical companies," and "fear of

180 COVID-19."

To measure familiarity with the health care system, we used the pre-existing Navigational Health Literacy (Navigational HL, or NHL in this article) index proposed by Griese et al.[44]. This self-assessed measure draws on the 12-item version of the European Health Literacy Scale (HLS-EU). In our study, each item of the NHL index was dichotomized into two responses: "easy" (value 1) and "difficult" (value 0). The final score was calculated by summing up the values obtained for each item. Participants were divided into quartiles ranging from Q1 (low navigating skills) to Q4 (high navigating skills). The consistency of our results was checked using the Health Literacy scale[45].

The association between variables was measured using Pearson's correlation coefficient for numeric variables and the chi-square test for categorical variables. Estimated proportions were interpreted based on the margins of error provided by IPSOS, which ranged from 1.4 to 3.1 points.

Two binomial logistic regression analyses were performed to identify the (adjusted) factors that influenced perceptions of the association between drug novelty and drug effectiveness and perceptions of the role of the pharmaceutical industry in drug development. The analyses were carried out using a main effect model with purposeful selection of variables[46]. First, we selected variables showing univariate association with a p=0.20 and entered them in the binomial regression model. Second, we removed non-significant variables from the model (threshold value of p=0.05). Lastly, we re-entered each removed variable in the model and tested it for significance. Statistical analyses were performed using Python (Pandas 1.3.4 – Scipy 1.7.2 – Statsmodel 0.13.1).

Results

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203 Perceptions of the association between drug novelty and drug effectiveness 204 A quarter (25%) of respondents agreed that new drugs are always more effective than old 205 ones, with only 3% agreeing completely. 206 Perceptions of the association between drug novelty and drug effectiveness were correlated 207 with familiarity with the health care system (Table 1). Indeed, 45% of respondents with a 208 high NHL score agreed that new drugs are always more effective than old ones, compared 209 to 16% of those with a low NHL score. Agreement was stronger among younger 210 respondents (33%) and among respondents situated at the two ends of the income scale 211 (34% for low household income and 37% for high household income). It was also stronger 212 among respondents with a high level of trust in politicians or pharmaceutical companies 213 (45% and 34%). 214 By contrast, respondents with poor health condition were less likely to agree that new drugs 215 are always more effective than old ones. A correlation was also found between perception of 216 risk and perception of the association between drug novelty and drug effectiveness. Thus, 217 16% of respondents who did not fear COVID-19 agreed that new drugs are always more 218 effective than old ones compared to 35% of those who did fear the disease. 219 The binary logistic regression analysis (**Table 2**, R²=0.11) found a significant effect of 220 familiarity with the health care system on perceptions of the association between drug 221 novelty and drug effectiveness. Indeed, respondents with a high NHL score were more likely 222 to agree that new drugs are always more effective than old ones (OR 3.34 [2.13-5.24])

compared to respondents with a low NHL score. Perception of risk also appeared to have a

strong effect, as respondents who did not fear COVID-19 were more than two times less likely to agree with this statement than those who did (OR 0.38 [0.18-0.82]).

Trust in doctors and scientists had a non-significant effect on perceptions of the association between drug novelty and drug effectiveness. Respondents with a low level of trust in pharmaceutical companies or politicians were two times less likely to agree that new drugs are always more effective than old ones compared to those with a high level of trust in these institutions (OR 0.63 [0.42-0.95] and OR 0.68 [0.49-0.94], respectively).

Finally, older respondents were two times less likely than younger ones to agree with this statement (OR 0.57 [0.31-1.03]). Neither gender nor health information-seeking behavior had a statistically significant effect in the final model.

Perceptions of the role of the pharmaceutical industry in drug development

Respondents had a high level of trust in doctors (92%) and scientists (87%) but a low level of trust in politicians (only 15%). They were much more split concerning pharmaceutical companies (42% trusted them).

Most respondents (86%) agreed that doctors need to collaborate with the pharmaceutical industry to develop new treatments. Less than half (43%) agreed that drug development should be driven by the pharmaceutical industry.

Different factors were associated with perceptions of the role of the pharmaceutical industry in drug development. Thus, 71% of respondents who agreed that new drugs are always more effective than old ones also agreed that drug development should be driven by the pharmaceutical industry. Likewise, 67% of respondents with a high NHL score agreed with the latter statement, compared to only 31% of respondents with a low NHL score. Low

educational level (51%), high household income (60%), low health information-seeking behavior (54%), trust in pharmaceutical companies (66%) or politicians (60%), and good health condition (48%) were all associated with the belief that drug development should be driven by the pharmaceutical industry.

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However, this belief did not exclude positive attitudes towards public intervention. Almost all respondents felt that the state should be directly involved in drug development when the disease is serious (86%). Even those who agreed that the pharmaceutical industry should drive drug development considered that direct state action was sometimes necessary (41%), with only 2% of respondents stating that drugs should be developed by the pharmaceutical industry alone.

The logistic regression analysis (**Table 3**, R²=0.15) showed that perceptions of the association between drug novelty and drug effectiveness had an important effect on perceptions of the role of the pharmaceutical industry in drug development. Respondents who agreed that new drugs are always more effective than old ones were almost four times more likely to agree that drug development should be driven by the pharmaceutical industry (OR 3.85 [2.76-5.39] compared to respondents who disagreed). The NHL score had an independent effect, as respondents with a high NHL score were more than twice as likely to agree that drug development should be driven by the pharmaceutical industry (OR 2.62 [1.73-3.97] compared to respondents with a low NHL score). By contrast, respondents with a low level of trust in pharmaceutical industry were less likely to agree with that statement (OR 0.38 [0.28-0.50] compared to respondents with a high level of trust). Likewise, respondents with poor health condition were less likely to agree that drug development should be driven by the pharmaceutical industry (OR 0.60 [0.44-0.82] for respondents with average health condition compared to those with good health condition; low statistical significance for the difference between respondents with poor health condition and those with good health condition, p=0.35).

272 Discussion

The notion of techno-scientific progress has been intertwined with biomedical advances[47] and the organization of modern medicine[48] at least since the eighteenth century. Health innovations are now ubiquitous and range from targeted drugs in genomic medicine[49] to imaging technologies based on Artificial Intelligence[50]. Innovations are introduced in medical practices at such a high pace that some of them remain on the market for short periods of time, before they are replaced with the next wave. With the COVID-19 pandemic, medical debates surrounding innovative treatments have come to the attention of the general public as never before[3,4]. The main opposition is observed between those who advocate for the development of new drugs to treat COVID-19 and those who favor the repositioning of old drugs with proven efficacy. Indeed, despite being a major achievement, new vaccines are being criticized due to their purported uncertainty[10]. Research on treatments for COVID-19 is ongoing and continues to impact both health behavior and public perceptions of the disease.

The paternalistic approach that prevailed in medicine up to the end of the twentieth century left little room for patients and their families to intervene in medical decisions. While the therapeutic decision is still up to doctors, shared decision-making is gaining ground, with patients and patients' advocates increasingly demanding that patients be actively involved in medical decisions and evaluations[51]. The advent of patient participation in their own care was an important change, but it also created a space for the pharmaceutical industry to exert direct influence on consumers and end-users (in particular through patients' associations). Patients can now pressure their doctors to develop or prescribe treatments that they believe to be "useful." Moreover, the greater emphasis on individual responsibility for health decisions has led many patients to engage in what some refer to as "doctor shopping" or "medical tourism." Indeed, some people may now choose to refuse a novel treatment (e.g., a

new vaccine), while others may favor a specific medical center or may decide to travel abroad to receive innovative treatment[52]. However, in the context of clinical trial recruitment, the narrative that stresses the value of drug novelty can be misleading given the uncertain benefits of the tested treatments[53]. Instances of unproven therapies and unfulfilled promises have led some people to criticize the pharmaceutical lobby, with frequent denunciations of its economic power, influence on health policy, and nefarious effect on citizen health. In view of this, a better understanding of public attitudes towards new treatments, and more generally towards medical innovation, is needed to elucidate individual preferences in health care.

Our findings support our hypothesis that respondents with a high level of trust in institutions and those with a high level of familiarity with the health care system are more likely to believe that new drugs are more effective than old ones (H1). Familiarity with the health care system was measured using the NHL index. This indicator was developed to evaluate people's difficulties in accessing, understanding, appraising, and applying information for navigating the health care system[44]. Compared to standard health literacy indexes, which focus more broadly on health and illness, the NHL index measures respondents' familiarity with their own national health care system. From a sociological perspective, it captures familiarity with specific health practices and health contexts. Such familiarity is often described as a specific form of what sociologists, following Pierre Bourdieu, have referred to as cultural capital[54]. In the context of health care, the possession of cultural capital has been shown to affect health choices and the way that people navigate the health care system[55]. The strong effect of the NHL index observed in our study indicates that it is also correlated with attitudes towards medical innovation.

In addition, the strong effect of the NHL index reflects the value attributed to innovation in French medical culture. Indeed, the value of medical innovation is not only promoted by professionals and health policies in France, but is also embodied by individuals with a high

level of familiarity with the health care system and by the beneficiaries of new treatments. As such, our finding is consistent with earlier studies on public understandings of science, which found institutional proximity to affect attitudes towards science and technology[56].

Moreover, our study supports the claim that attitudes towards new treatments reflect broader attitudes towards the health care system and towards institutions in general. Positive perceptions of medical innovation were also found to correlate with greater acceptance of the standard model of innovation, as the belief that new drugs are more effective than old ones was associated with a high level of trust in politicians, and/or pharmaceutical companies.

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While pandemics have historically prompted the intervention of public actors in biomedical research[57], pharmaceutical companies remain the main actors of medical innovation today. However, public perceptions of the role of the pharmaceutical industry are increasingly polarized, with criticisms generally focusing on the industry's excessive role in the medical innovation process and on potential conflicts of interest[28]. This is a major concern, as the association between public research and the pharmaceutical industry has consequences for the legitimacy of science[25]. As several studies have shown, overall trust in pharmaceutical companies is especially low in France[43,58]. There is currently little explanation for this mistrust, though a series of public scandals, including the Mediator case[59], may have contributed to this. Interestingly, in our study, respondents who believed that new drugs are more effective than old ones were more likely to consider that drug development should be driven by the pharmaceutical industry. Likewise, respondents with a high level of familiarity with the health care system had a high level of trust in pharmaceutical companies. In a context where industrial R&D is both central to the innovation process and negatively depicted in the news, a better understanding of the factors associated with trust in the pharmaceutical industry is needed to reduce polarization and, consequently, to ensure adequate public vigilance of biomedical research.

Our findings appear also to support our hypothesis of a specific effect of health condition and age on perceptions of the association between drug novelty and drug effectiveness (H2a). This effect, however, was statistically significant in bivariate analysis only. The lack of statistical significance in multivariate analysis may be explained by the limited statistical power of our survey, but also by the fact that statistical models do not account for individual trajectories and experiences. Some studies have shown that the experience of chronic disease leads to specific navigating skills and favors specific attitudes towards medical innovation[52]. One could suppose that people with chronic diseases, who tend to have a high level of familiarity with the health care system, are more likely to develop a critical view of this system. This is especially likely if the disease is deadly or debilitating, with little hope for improvement or recovery. By contrast, our findings did not support our hypothesis of a specific effect of socio-economic status (H2b). Beyond the limited statistical power of the survey, this absence of effect may be explained by the fact that the NHL index already accounts for health condition or socio-economic status. In other words, familiarity with the health care system as measured by the NHL index likely acts as a mediating factor between socio-economic status and perceptions of the association between drug novelty and effectiveness. By contrast, our hypothesis H3 was supported by our findings, as fear of COVID-19 appeared to correlate with the belief that new drugs are more effective than old ones. This finding suggests that in an epidemic context, people who are more anxious about the disease are more likely to support medical innovation. More generally, one could argue that perceptions of the association between drug novelty and drug effectiveness are influenced by health policy, the availability of existing treatments, the actual and perceived severity of the disease, and the global or restricted nature of the health threat.

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This study extends the analysis initiated in June 2020 with the first round of HLS₁₉[43]. Our findings indicate that trust in institutions and perceptions of clinical research in France have remained stable during the COVID-19 pandemic (**Table 1**). They also go beyond the original findings by showing that the role of the pharmaceutical industry in drug development is

perceived as normal, even necessary, with almost half of respondents agreeing that drug development should be driven by this industry. This finding is especially interesting given that the pharmaceutical lobby is regularly criticized, both in France and beyond, in terms of regulation failures and potential conflicts of interests[60]. Our findings also suggest that perceptions of the association between drug novelty and drug effectiveness vary depending on the terms used in public debates to refer to pharmaceutical actors. Thus, the questionnaire used in the first round of HLS₁₉ measured respondents' level of trust in "*l'industrie pharmaceutique*" (the pharmaceutical industry), while that used in the second round measured respondents' level of trust in "*les laboratoires pharmaceutique*" (pharmaceutical companies). Interestingly, the level of trust rose from 25% in the first round to 42% in the second one, which may be explained by the fact that the term "laboratory" evokes the scientific field while that of "industry" is associated with the economic domain. It should be noted, however, that the high level of trust reported in the second round may also reflect the increased presence of pharmaceutical actors in the media during the COVID-19 pandemic – and especially the success of the new vaccines.

Today, public acceptance of the standard model of innovation in medicine is a major political issue. The economic influence exerted by the pharmaceutical industry on the orientation and supervision of biomedical research, and especially on the conduct of clinical trials[61], has been well documented[17,18]. In France, public funding of biomedical research by international private companies has been regularly criticized. For instance, a major controversy erupted over the important public funds allocated to the pharmaceutical company Sanofi (via the *Crédit d'Impôt Recherche*), which laid off many workers and failed to develop a "French" vaccine. Moreover, the economic domination of the pharmaceutical industry was loudly denounced when Pfizer decided to raise the cost of vaccines. In a recent book, Fierlbeck et al. argued that "there is plenty of controversy surrounding pharmaceuticals, but it cannot be denied that the pharmaceutical industry is both socially beneficial and profitable." However, they also recognized that "the fact that citizens believe

medicines are reasonably safe and that the medications work is based more on trust than on the scientific evidence (p.4)"[32]. Because the innovation process is an important component of health care, the way it is framed in public debates can have a lasting impact on trust in the medical system, in regulatory agencies, and even in science in general.

Limitations

This study has several limitations. First, our study was an exploratory analysis of an understudied topic that aimed to test a series of hypotheses. Our findings are therefore tentative and will need to be confirmed in future studies. Second, only two questions concerning the topic of interest were analyzed, which precluded the development of an explanatory model. Political orientation, knowledge of specific aspects of the health care system, and perceptions of the cost of innovations will also need to be investigated in the future. Third, our sample was not probabilistic. However, it was representative of the French population since we used quota sampling. Lastly, our questionnaire could not fully capture the diversity of attitudes towards medical innovation, suggesting a need for qualitative studies on the topic. Such studies could determine the extent to which the COVID-19 pandemic changed people's perceptions of the association between drug novelty and drug effectiveness.

Conclusion

Today, medical research and innovation are framed both as a solution and as a problem in public debates. While drugs and vaccines with proven efficacy allow for sound health policy, pharmaceutical regulation failures and the overwhelming role of the pharmaceutical industry in drug development have created legitimate suspicion in the general public. Because the positive value assigned to medical innovation is directly linked to the belief in scientific progress that underpins medical institutions, public criticism of medical research has an

impact on the process of drug development and, consequently, on the level of trust in experts and scientists. In fact, public attitudes towards medical innovation can be ambivalent, as novelty increases both the perceived value of a given product and the perceived risk associated with it[41]. This ambivalence is not limited to drugs and can be extended to other innovations – for instance, the contact tracing apps developed during the COVID-19 pandemic[62,63]. In a global context where trust in institutions, and especially in science, has a major impact on health behavior and democratic life, there is a growing need for a better understanding of public perceptions of medical innovation.

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Bibliography

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439 440 441	[1]	McCreary EK, Meyer NJ. Covid-19 controversies: the tocilizumab chapter. BMJ [Internet]. 2021 Jan 27;372:n244. Available from: http://www.ncbi.nlm.nih.gov/pubmed/33472855
442 443 444	[2]	Sourimant J, Aggarwal M, Plemper RK. Progress and pitfalls of a year of drug repurposing screens against COVID-19. Curr Opin Virol [Internet]. 2021 Jun; Available from: https://doi.org/10.1016/j.coviro.2021.06.004
445 446 447	[3]	Schultz É, Ward JK. Science under Covid-19's magnifying glass: Lessons from the first months of the chloroquine debate in the French press. J Sociol [Internet]. 2021 Mar 10; Available from: http://journals.sagepub.com/doi/10.1177/1440783321999453
448 449 450 451	[4]	Berlivet L, Löwy I. Hydroxychloroquine Controversies: Clinical Trials, Epistemology, and the Democratization of Science. Med Anthropol Q [Internet]. 2020 Dec 18;34(4):525–41. Available from: https://onlinelibrary.wiley.com/doi/10.1111/maq.12622
452 453 454	[5]	Moynihan R, MacDonald H, Bero L, Godlee F. Commercial influence and covid-19: Greater independence from commercial interests is more important than ever. BMJ. 2020;369:10–1.

455 456 457	[6]	Romain Lutaud, Ward J, Gentile G, Verger P. Between an Ethics of Care and Scientific Uncertainty: Dilemmas of General Practitioners in Marseille. In: The COVID-19 Crisis: Social Perspectives. 2021. p. 144–55.
458 459 460 461	[7]	Lutaud R, Scronias D, Ward J, Verger P. The hydroxychloroquine debate: a therapeutic dilemma for general practitioners. Eur J Public Health [Internet]. 2021 Apr 24;31(2):283–5. Available from: https://academic.oup.com/eurpub/article/31/2/283/6135355
462 463 464 465	[8]	Tuccori M, Convertino I, Ferraro S, Cappello E, Valdiserra G, Focosi D, et al. The Impact of the COVID-19 "Infodemic" on Drug-Utilization Behaviors: Implications for Pharmacovigilance. Drug Saf [Internet]. 2020;43(8):699–709. Available from: https://doi.org/10.1007/s40264-020-00965-w
466 467	[9]	Goldenberg MJ. Vaccine Hesitancy: Public Trust, Expertise, and the War on Science. University of Pittsburgh Press; 2021.
468 469 470 471	[10]	Ward JK, Alleaume C, Peretti-Watel P, COCONEL group. The French public's attitudes to a future COVID-19 vaccine: The politicization of a public health issue. Soc Sci Med [Internet]. 2020 Nov;265:113414. Available from: https://linkinghub.elsevier.com/retrieve/pii/S027795362030633X
472 473 474 475	[11]	Roustit M, Guilhaumou R, Molimard M, Drici M-D, Laporte S, Montastruc J-L. Chloroquine and hydroxychloroquine in the management of COVID-19: Much kerfuffle but little evidence. Therapies [Internet]. 2020 Jul;75(4):363–70. Available from: https://linkinghub.elsevier.com/retrieve/pii/S0040595720301001
476	[12]	Briggs C, Hallin D. Making health public. London: Routledge; 2016.
477 478 479	[13]	Marchetti D. Quand la santé devient médiatique. Les logiques de production de l'information dans la presse. Grenoble: Presses universitaires de Grenoble; 2010. 191 p.
480 481 482 483	[14]	Stiller I, van Witteloostuijn A, Cambré B. Determinants of radical drug innovation: a systematic literature review [Internet]. Management Review Quarterly. Springer International Publishing; 2021. Available from: https://doi.org/10.1007/s11301-021-00218-9
484 485 486 487	[15]	Neresini F, Crabu S, Di Buccio E. Tracking biomedicalization in the media: Public discourses on health and medicine in the UK and Italy, 1984–2017. Soc Sci Med [Internet]. 2019;243(October):112621. Available from: https://doi.org/10.1016/j.socscimed.2019.112621
488 489 490	[16]	Borup M, Brown N, Konrad K, Van Lente H. The sociology of expectations in science and technology. Technol Anal Strateg Manag [Internet]. 2006;18(September):285–98. Available from: http://dx.doi.org/10.1080/09537320600777002
491 492	[17]	Sismondo S. Ghost-managed medicine: Big pharma's invisible hands. Mattering Press; 2018.

493 494 495	[18]	Davis C, Abraham J. Unhealthy Pharmaceutical Regulation [Internet]. Unhealthy Pharmaceutical Regulation. London: Palgrave Macmillan UK; 2013. Available from: http://link.springer.com/10.1057/9781137349477
496 497	[19]	Keating P, Cambrosio A. Cancer on Trial. Oncology as a New Style of Practice. Chicago: Chicago Press; 2011. 476 p.
498 499 500	[20]	Dalgalarrondo S. Quelle place pour les associations de malades dans le processus d'innovation médicamenteuse? Rev Fr Aff Soc [Internet]. 2007;3–4(3):171–91. Available from: http://www.cairn.info/article.php?ID_ARTICLE=RFAS_073_0171
501 502 503	[21]	Wieseler B, McGauran N, Kaiser T. New drugs: Where did we go wrong and what can we do better? BMJ [Internet]. 2019;366(July):1–8. Available from: http://dx.doi.org/doi:10.1136/bmj.l4340
504 505	[22]	Harrison M, Marra CA, Bansback N. Preferences for 'New' Treatments Diminish in the Face of Ambiguity. Heal Econ (United Kingdom). 2017;26(6):743–52.
506 507 508	[23]	Mahase E. FDA allows drugs without proven clinical benefit to languish for years on accelerated pathway. BMJ [Internet]. 2021 Jul 29;(July):n1898. Available from: https://www.bmj.com/lookup/doi/10.1136/bmj.n1898
509 510 511	[24]	Theodorou M, Tsiantou V, Pavlakis A, Maniadakis N, Fragoulakis V, Pavi E, et al. Factors influencing prescribing behaviour of physicians in Greece and Cyprus: Results from a questionnaire based survey. BMC Health Serv Res. 2009;9:1–9.
512 513	[25]	Besley JC, McCright AM, Zahry NR, Elliott KC, Kaminski NE, Martin JD. Perceived conflict of interest in health science Partnerships. PLoS One. 2017;12(4):1–20.
514 515 516	[26]	Dubé È, Ward JK, Verger P, Macdonald NE. Vaccine Hesitancy, Acceptance, and Anti-Vaccination: Trends and Future Prospects for Public Health. Annu Rev Public Health. 2020;42:175–91.
517	[27]	Eyal G. The Crisis of Expertise. Cambridge: Polity Press; 2019. 208 p.
518 519	[28]	Hauray B, Boullier H, Gaudillère J, Michel H. Conflict of interest and medicine. Routledge, editor. 2021.
520 521 522	[29]	Davis C. Drugs , cancer and end-of-life care : A case study of pharmaceuticalization ? Soc Sci Med [Internet]. 2015;131:207–14. Available from: http://dx.doi.org/10.1016/j.socscimed.2014.12.007
523 524 525 526	[30]	Salas-Vega S, Shearer E, Mossialos E. Relationship between costs and clinical benefits of new cancer medicines in Australia, France, the UK, and the US. Soc Sci Med [Internet]. 2020;258(April):113042. Available from: https://doi.org/10.1016/j.socscimed.2020.113042
527 528	[31]	Chabannon C. Thérapies cellulaires et géniques : la guérison à l'horizon ? Cancer(s) et psy(s). 2020;n°5(1):122.
529 530 531	[32]	Fierlbeck K, Graham J, Herder M. Transparency, Power, and Influence in the Pharmaceutical Industry: Policy Gain Or Confidence Game? University of Toronto Press; 2021.

532 533	[33]	Mede NG, Schäfer MS. Science-related populism: Conceptualizing populist demands toward science. Public Underst Sci. 2020;29(5):473–91.
534 535 536	[34]	Gould S, Norris SL. Contested effects and chaotic policies: the 2020 story of (hydroxy) chloroquine for treating COVID-19. Vol. 3, The Cochrane database of systematic reviews. 2021.
537 538 539 540	[35]	Perez J, Roustit M, Lepelley M, Revol B, Cracowski J-L, Khouri C. Reported Adverse Drug Reactions Associated With the Use of Hydroxychloroquine and Chloroquine During the COVID-19 Pandemic. Ann Intern Med [Internet]. 2021 Jun;174(6):878–80. Available from: https://www.acpjournals.org/doi/10.7326/M20-7918
541 542 543	[36]	Boschiero MN, Capasso Palamim CV, Ortega MM, Mauch RM, Lima Marson FA. One year of coronavirus disease 2019 (Covid-19) in brazil: A political and social overview. Ann Glob Heal. 2021;87(1):1–27.
544 545 546	[37]	Hilgartner S, Hurlbut JB, Jasanoff S. Was "science" on the ballot? Science (80-) [Internet]. 2021 Feb 26;371(6532):893–4. Available from: https://www.sciencemag.org/lookup/doi/10.1126/science.abf8762
547 548 549	[38]	Colgan S, Faasse K, Martin LR, Stephens MH, Grey A, Petrie KJ. Perceptions of generic medication in the general population, doctors and pharmacists: A systematic review. BMJ Open. 2015;5(12).
550 551 552	[39]	Jie Y. Older is better: Consumers prefer older drugs. Psychol Mark [Internet]. 2020 Nov 13;37(11):1498–510. Available from: https://onlinelibrary.wiley.com/doi/10.1002/mar.21395
553 554 555 556	[40]	Russo S, Jongerius C, Faccio F, Pizzoli SFM, Pinto CA, Veldwijk J, et al. Understanding Patients' Preferences: A Systematic Review of Psychological Instruments Used in Patients' Preference and Decision Studies. Value Heal [Internet]. 2019;22(4):491–501. Available from: https://doi.org/10.1016/j.jval.2018.12.007
557 558 559 560	[41]	Albertsen L, Wiedmann KP, Schmidt S. The impact of innovation-related perception on consumer acceptance of food innovations – Development of an integrated framework of the consumer acceptance process. Food Qual Prefer [Internet]. 2020;84(April):103958. Available from: https://doi.org/10.1016/j.foodqual.2020.103958
561 562 563	[42]	Connor M, Siegrist M. Factors Influencing People's Acceptance of Gene Technology: The Role of Knowledge, Health Expectations, Naturalness, and Social Trust. Sci Commun. 2010;32(4):514–38.
564 565 566 567	[43]	Schultz É, Ward JK, Atlani-Duault L, Holmes SM, Mancini J. French Public Familiarity and Attitudes toward Clinical Research during the COVID-19 Pandemic. Int J Environ Res Public Health [Internet]. 2021 Mar 5;18(5):2611. Available from: https://www.mdpi.com/1660-4601/18/5/2611
568 569 570	[44]	Griese L, Berens EM, Nowak P, Pelikan JM, Schaeffer D. Challenges in navigating the health care system: Development of an instrument measuring navigation health literacy. Int J Environ Res Public Health. 2020;17(16):1–17.

571 572 573 574	[45]	Rouquette A, Nadot T, Labitrie P, Van den Broucke S, Mancini J, Rigal L, et al. Validity and measurement invariance across sex, age, and education level of the French short versions of the European Health Literacy Survey Questionnaire. PLoS One. 2018;13(12):1–15.
575 576 577	[46]	Hosmer DW, Lemeshow S, Sturdivant RX. Applied Logistic Regression [Internet]. Wiley; 2013. 510 p. (Wiley Series in Probability and Statistics). Available from: https://onlinelibrary.wiley.com/doi/book/10.1002/9781118548387
578 579 580	[47]	Gaudillière J-P, Hess V. Ways of Regulating Drugs in the 19th and 20th Centuries [Internet]. Gaudillière J-P, Hess V, editors. London: Palgrave Macmillan UK; 2013. Available from: http://link.springer.com/10.1057/9781137291523
581 582	[48]	Pickstone J. Ways of Knowing: Towards a Historical Sociology of Science , Technology and Medicine. Br J Hist Sci. 1993;26(4):433–58.
583 584 585	[49]	Au L. Recent Scientific/Intellectual Movements in Biomedicine. Soc Sci Med [Internet]. 2021;278(April):113950. Available from: https://doi.org/10.1016/j.socscimed.2021.113950
586 587	[50]	Topol E. Deep medicine: how artificial intelligence can make healthcare human again. New York: Hachette; 2019.
588 589 590	[51]	Elwyn G, Frosch D, Thomson R, Joseph-Williams N, Lloyd A, Kinnersley P, et al. Shared decision making: A model for clinical practice. J Gen Intern Med. 2012;27(10):1361–7.
591 592 593	[52]	Besle S, Sarradon-Eck A. Choisir le risque : l'autonomie du malade en situation d'échec thérapeutique. Anthropol Santé [Internet]. 2019 Nov 3;(19). Available from: http://journals.openedition.org/anthropologiesante/4817
594 595 596 597	[53]	Mancini J. The impact of patient education on consideration of enrollment in clinical trials. J Community Support Oncol [Internet]. 2018 Mar 1;16(2):e81–8. Available from: https://www.mdedge.com/jcso/article/164500/patient-survivor-care/impact-patient-education-consideration-enrollment-clinical
598 599 600	[54]	Willis K, Collyer F, Lewis S, Gabe J, Flaherty I, Calnan M. Knowledge matters: producing and using knowledge to navigate healthcare systems. Heal Sociol Rev. 2016;25(2):202–16.
601 602 603	[55]	Paccoud I, Nazroo J, Leist A. A Bourdieusian approach to class-related inequalities: the role of capitals and capital structure in the utilisation of healthcare services in later life. Sociol Heal Illn. 2020;42(3):510–25.
604 605 606	[56]	Sturgis P, Allum N. Science in Society: Re-Evaluating the Deficit Model of Public Attitudes. Public Underst Sci [Internet]. 2004;13(1):55–74. Available from: http://journals.sagepub.com/doi/10.1177/0963662504042690
607 608 609	[57]	Beall RF, Hollis A. Global clinical trial mobilization for COVID-19: higher, faster, stronger. Drug Discov Today [Internet]. 2020;25(10):1801–6. Available from: https://doi.org/10.1016/j.drudis.2020.08.001
610	[58]	LEEM. Observatoire sociétal du médicament. 2019.

611 612	[59]	Gaudillière J-P. « Mediator n'est pas une exception, mais le révélateur de problèmes structurels dans le monde du médicament ». Mouvements. 2019;n°98(2):67.
613 614	[60]	Maraninchi D. Actuality of conflicts of interests in medicine. From financial interests to the run for scientific publication. Bull Cancer. 2021;108(7–8):677–85.
615 616	[61]	Jureidini J, McHenry LB. The Illusion of Evidence-Based Medicine: Exposing the crisis of credibility in clinical research. Wakefield Press; 2020.
617 618 619	[62]	Williams SN, Armitage CJ, Tampe T, Dienes K. Public attitudes towards COVID-19 contact tracing apps: A UK-based focus group study. Heal Expect. 2020;(November 2020):377–85.

- 620 [63] Touzani R, Schultz E, Holmes SM, Vandentorren S, Arwidson P, Guillemin F, et al.
- 621 Early Acceptability of a Mobile App for Contact Tracing During the COVID-19 Pandemic in
- 622 France: National Web-Based Survey. JMIR mHealth uHealth [Internet]. 2021 Jul
- 623 19;9(7):e27768. Available from: https://mhealth.jmir.org/2021/7/e27768