Innovation and Interdependence: The Case of Gene-Editing Technology*

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Abstract

Technological breakthroughs often disrupt social, economic, and political systems, posing unique challenges for governments. In this paper, we examine how breakthroughs shape patterns of interdependence between states. We argue that rapid technological advancement increases interdependence among national governments in two ways. First, it lowers barriers to entry and creates opportunities for forum-shopping by researchers, firms, and other actors. This facilitates arbitrage as actors relocate to more permissive jurisdictions, generating incentives for regulatory competition. Second, public unease about new technologies creates the potential for backlash against controversial applications. This backlash can spill across borders: accidents or misuse in one jurisdiction undermines support for research and commercial development elsewhere. Together, these processes create incentives for states to mismanage risk, generating inefficient cycles of accelerated progress disrupted by damaging controversies. We test these mechanisms in the case of gene editing, a field that has experienced rapid technological advancement in recent years. We find support for the theory in patterns of gene editing research, social media data, and a novel survey experiment. Our results demonstrate that technological disruption increases interdependence, undermining states’ ability to regulate in isolation and strengthening the case for international policy coordination.

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1 Introduction

Technological innovation is a defining feature of contemporary social and economic life. Recent advances in fields like robotics, biotechnology, digital finance, and artificial intelligence promise improved welfare through enhanced health, productivity, and economic growth. They also pose significant risks: many emerging technologies can be misused to cause harm or violate ethical norms. The recent revolution in gene editing technology, for example, has been celebrated for facilitating new medical therapies and also criticized for enabling controversial modifications of human DNA.

Governments navigate this tradeoff by regulating the development and application of emerging technologies. Regulations seek to guide the path and speed of technological progress, balancing the economic and social potential of technological change against the risk of harm. Countries often make different choices in this environment — imposing more restrictive or permissive rules on the use of a particular technology — as governments align regulations with public preferences and social norms.

We argue that, in addition to provoking a regulatory choice within countries, technological disruption also shapes patterns of interdependence between them. Technological breakthroughs lower barriers to entry, erode incumbent advantages, and allow more actors in more jurisdictions to compete for status and profit. As a technology diffuses, countries’ regulatory efforts become more closely linked in two ways.

First, the reduced entry barriers associated with new technologies create greater opportunities for forum-shopping by researchers and firms. This facilitates regulatory arbitrage as actors evade national regulations by relocating to more permissive jurisdictions. In some cases, governments will face pressure to weaken regulatory standards in order to stimulate domestic innovation and lure researchers, firms, and capital from other jurisdictions. While regulatory arbitrage and competition are well-established features of national governance
we argue that technological shocks often exacerbate the problem by lowering costs and increasing the cross-border mobility of production.

The second source of interdependence is rooted in public attitudes regarding emerging technologies. Because technological innovations involve risks of harm or misuse, they generate apprehension among citizens and potential consumers. When controversies occur, they often spur public backlash and undermine support for related research and commercial development. We argue that this backlash frequently spills across national boundaries, such that controversies in one state affect public attitudes in another. As a result, one government’s decision to weaken regulation can damage confidence in the technology around the world. Unlike regulatory arbitrage, we are not aware of existing scholarship that examines the potential for spillovers in public backlash. Nonetheless, we argue that it is an important source of interdependence in the governance of new technologies.

These patterns of interdependence combine to produce several problems for governments as they regulate emerging technologies. They create incentives for states to mismanage risk, generating inefficient cycles of accelerated progress disrupted by damaging controversies. Enhanced international arbitrage opportunities encourage states to compete by lowering regulatory barriers. The presence of weak regulation may temporarily speed technological progress, but it also increases the systemic risk of controversial accidents or misuse. These controversies arouse public anxiety, undermine support, and stall continued progress.

In addition, these dynamics weaken the power of national governments to regulate technology in isolation. Increased opportunities for arbitrage make it easier for targets of regulation to evade national rules. The potential for spillovers in public backlash mean that effective national governance cannot insulate a country from poor regulation in another jurisdiction. Both of these mechanisms therefore increase the need for international policy coordination to manage interdependence.

We examine these processes in the case of gene editing, a field in the midst of a techno-
logical revolution. Gene editing, or genetic engineering, refers to the targeted manipulation of an organism’s genetic material. The emergence of CRISPR and associated techniques in the last decade provides a dramatically more accurate, efficient, and economical method for editing genes. In awarding the 2020 Nobel Prize in Chemistry to CRISPR architects Emmanuelle Charpentier and Jennifer Doudna, the Royal Swedish Academy of Sciences celebrated the technology’s revolutionary capability for “rewriting the code of life” (Royal Swedish Academy of Sciences, 2020).

In response to these technological advances, scientists in academia and industry are applying gene editing to an increasingly diverse set of objectives. Laboratory researchers routinely “knock out” genes in mice or other animals to study gene function and expression. Therapeutic developers are creating new gene therapies to treat cancer and correct harmful genetic mutations (Khan et al., 2016). Agricultural producers are applying CRISPR to both produce and livestock. Research teams have successfully altered the DNA of mosquitos to prevent the transmission of malaria (Gantz et al., 2015). More recently, gene-editing technology has been used to develop diagnostic tests for COVID-19 (Straiton, 2020).

We argue that gene editing is an archetypal disruptive technology, and as a result it enhances the two forms of interdependence described above. First, it encourages regulatory arbitrage among actors in science and industry. As the capital and infrastructure needed to edit genes fall, countries with weaker regulatory environments become more attractive destinations for cutting-edge researchers. These countries can more easily capitalize on the scientific and economic potential of gene editing, in part by drawing human and financial capital away from countries with more stringent regulations.

Second, controversial applications of gene-editing technology generate public backlash.

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1 The acronym CRISPR stands for clustered regularly interspaced short palindromic repeats. The term was coined by Ishino et al. (1987), who first noted the appearance of repeating DNA sequences in bacteria.

2 See, for example, the Knockout Mouse Project: [https://www.komp.org](https://www.komp.org)
that spans national borders. These controversies can undermine public support, reduce funding for related research, and constrain even responsible scientific activity. As a result, the field of gene-editing research has progressed in fits and starts, with periods of promising technological advancements interrupted by crises of public confidence. As clinical gene-editing applications are brought to market, lack of public trust could reduce demand for potentially life-saving technologies.

We probe these arguments with two sets of empirical tests. We first analyze patterns of gene editing research and development to examine incentives for regulatory arbitrage. Specifically, we assess whether countries with weaker regulations outperform more stringent jurisdictions in the development and application of gene technologies. We leverage the 2012 introduction of CRISPR as a temporal shock to examine how national regulatory environments shape patterns of employment among gene scientists and the clinical development of gene therapies. Our results are consistent with theoretical expectations: the development of CRISPR technology enhanced the frequency of scientific forum-shopping and increased the economic returns to weak regulatory environments.

To test for spillovers in public backlash, we examine changes in Twitter sentiment in the wake of a high-profile gene-editing scandal in November 2018. Analysis of nearly one million tweets about gene-editing technology suggest that controversies in one country (China) depress public opinion in other jurisdictions. We then implement a novel survey experiment that probes American respondents’ reaction a scandalous application of gene-editing technology. The experiment varies information about the occurrence and location of inappropriate gene-editing activity. We find that both foreign and domestic gene-editing controversies negatively affect domestic public support for gene-editing research.

Our paper adds to a growing literature on international competition, cooperation, and

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Drezner (2008) argues that distrust of genetically-modified organisms in Europe, for example, led to increased regulations and low public support for GMO consumption despite its potential health and economic benefits.
technological advancement (Milner & Solstad, 2020; Perlman, 2020; Canfil, 2021). We develop a theory of technology and interdependence in a domain, biotechnology, that has been largely neglected by scholarship in international relations and political science. While international relations scholars have paid close attention to the security implications of technological advancements (Buchanan & Keohane, 2015; Kreps & Wallace, 2016; Ayoub & Payne, 2016), we know less about governance of scientific issues in non-security sectors. We expect biotechnology to increase in salience for political science as governments and their citizens grapple with the unprecedented technological progress in this field.

More broadly, we identify two theoretical mechanisms — regulatory arbitrage and spillovers in public backlash — that link countries’ fates as they govern emerging technologies. In doing so, we demonstrate how technological shocks interact with patterns of economic and political exchange to induce interdependence among countries. These mechanisms have clear implications for the design of international institutions, which are likely to be charged with managing these spillovers.

The rest of the paper is organized as follows. In section 2 we provide background on the case of gene editing, summarizing the emergence and governance of this rapidly advancing technology. Section 3 draws on this case to develop our theory of technological innovation and international interdependence. Section 4 describes our empirical strategy and presents our findings, and section 5 concludes.

2 Gene Editing: Technological and Political Landscape

While the ability to modify genetic material is not new, scientific advances have transformed the field over the past decade. The term “gene editing” (also referred to as genetic engineering, modification, or manipulation) refers to the direct alteration of an organism’s DNA. The goal is typically to suppress or alter naturally-occurring biological traits of the organism.
Historically, the field of gene editing evolved from splicing together naturally-occurring genetic material (producing “recombinant” DNA) in the 1970s to using cells’ own DNA-repair technology to selectively edit specific genes (using “programmable nucleases”) in the early 2000s.\footnote{For a general overview of scientific progress in gene-editing technology, see Gupta et al. (2014).}

The emergence of the CRISPR method in the 2010s represented a particularly significant breakthrough in gene-editing technology. The name CRISPR (clustered regularly interspaced short palindromic repeats) refers to a series of repeating DNA sequences originally found in bacteria. These sequences provide bacteria with adaptive immunity, allowing them to recognize and destroy the DNA of harmful viruses. Scientists adapted this technique for programmable gene editing (Jinek et al., 2012). CRISPR targets specific gene sequences and cleaves them with a nuclease, most commonly the Cas9 enzyme. This “CRISPR-Cas9” system is significantly more accurate, efficient, and economical than previous gene-editing methods.

In the years since its development, CRISPR has become the dominant technology in the field of gene editing (Carroll, 2018)\footnote{Figure A1 in the appendix shows the number of patent applications utilizing three prominent gene-editing technologies: CRISPR, TALENs, and ZFNs. CRISPR has dominated technological development in the field, significantly outperforming other methods.}. A report in Stanford Medicine notes that while “other gene-editing tools have emerged in recent years...none seems to match the precision, low cost and usability of CRISPR” (Shwartz, 2018). Like many other breakthrough technologies, CRISPR dramatically reduces the costs associated with editing genes: a CRISPR-Cas9 RNA template may cost $65 to design compared to $1000 for the same template using other technologies (Shwartz, 2018)\footnote{However, as one gene researcher noted, there continue to be barriers to entry for many laboratories. In particular, innovation using CRISPR technology requires financial resources and investment in learning how to adapt CRISPR tools to specific projects (Interview by authors, 11.13.2019).}.

Lower costs have enabled the diffusion of CRISPR technology to laboratories around the
world. Diffusion is also facilitated by scientific norms regarding the accessibility of research materials. As a condition of publication, authors often must make their data and materials available to other researchers. Much of the biological material – including the plasmids used to edit genes – is handled by third-party distributors like AddGene, a repository that stores and disseminates genetic material used in published studies. Although CRISPR-related materials represent a small minority of AddGene’s repository, they are among the most commonly requested plasmids from researchers. Appendix Figure A2 displays the number of registered AddGene researchers by country of origin. American researchers are the largest group, followed by China, France, Japan, India, and Germany.

Many have cheered the spread of gene-editing technology, which has stimulated a “biotechnological revolution” in clinical care, diagnostics, agriculture, and other fields (Knott & Doudna, 2018). As with other technologies, however, rapid progress has also been accompanied by ethical concerns and fears of potential misuse. Anxiety over gene editing varies based on the type of genetic material researchers seek to alter. There is a strong norm against editing germline (heritable) cells in humans due to the unknown long-term effects of the changes, concerns about the ethics of editing heritable genes, and the difficulty of ensuring the safety of the procedure (Miller, 2015).

Concern about inappropriate genetic modification escalated sharply in 2018, when the Chinese scientist He Jiankui announced the birth of the world’s first gene-edited infants.

7 AddGene has served as a popular repository for CRISPR plasmids since Jinek et al. (2012) used the organization to store materials from their landmark paper. AddGene “is a global, nonprofit repository that was created to help scientists share plasmids;” the site allows scientists to register and deposit plasmids on the site and send a sample to the lab. Addgene then allows other scientists to request the material for their own research. Researchers who want to access these materials register on the AddGene website and pay a fee for the plasmid transfer. They are then free to replicate the parent study or alter the plasmids for their own research purposes. For more information, see http://addgene.org.

8 An AddGene employee estimated that 25% of researcher requests are for CRISPR-related plasmids (Interview by authors, 11.25.2019).

9 Researchers from the United States are overwhelmingly the most frequent depositors of CRISPR plasmids (see Appendix Figure A3).
He’s team used CRISPR to genetically alter twins in-vitro in order to render them immune to HIV (Cyranoski 2019). The revelation sparked international outcry, raising concerns about safety, consent of the participants, and the risks of modifying heritable germline cells that will pass alterations to subsequent generations. Many questioned whether China had the institutional capacity or will to reign in potential ethical violations by its scientists.

Other concerns are linked to the purposes that gene-editing technology can serve. The potential for genetic modifications to enhance socially-desirable traits in a human without conferring health benefits evoke the dark history of gene science\textsuperscript{10}. The increased accessibility and public profile of gene-editing technology has also encouraged amateur scientists to experiment in unsafe conditions. Communities of self-proclaimed “bio-hackers” use gene-editing tools on test animals, livestock, or even themselves (Keulartz & van den Belt 2016).

\subsection{Governance of Gene-Editing Technology}

Gene editing is governed by a fragmented patchwork of scientific norms, national laws, and international guidelines. When targeted gene editing first became feasible in the 1970s with the advent of recombinant DNA techniques, scientists attempted to construct self-governing arrangements with standards for appropriate gene-editing research. In 1973, leading geneticists announced a voluntary moratorium on gene-editing experiments involving certain viruses and toxins (Berg \textit{et al.} 1974). The moratorium was maintained for two years until it was replaced by formal guidelines adopted by the National Institutes of Health. Scientists involved in drafting the original guidelines argue that this decentralized approach was successful in constraining potentially inappropriate applications of gene-editing technology (Berg & Mertz 2010).

\footnotesize\begin{itemize}
\item Following 19th century advances in genetics, the British scientist Francis Galton popularized the idea of “improving the human stock” by encouraging procreation based on desirable traits (Goering 2014). The concept of “eugenics” took hold in the popular imagination and was weaponized to exclude immigrants, sterilize disabled individuals, and justify human rights abuses (Bouche & Rivard 2014).
\end{itemize}
In recent years, similar efforts have sought to establish new norms for the research community. A 2019 international conference of geneticists called for a global five-year ban on editing DNA in human eggs, sperm, or embryos that are brought to term (Lander et al., 2019). However, there is dissent about this approach even among the most prominent genetic researchers (Cohen, 2019). The lack of consensus creates uncertainty about appropriate applications of gene-editing technology, potentially contributing to misuse. In addition, there remain serious questions about whether voluntary, decentralized rules can succeed in an environment where gene-editing technology is more accessible and diffusely distributed than in the 1970s.

As early gene-editing technology progressed, national regulations began to supplement scientific norms. The United States created national guidelines that built upon the partial gene-editing moratorium of 1973-4 (Baskin et al., 2016). Other states followed suit as the practice became more widespread. Currently, there is significant variation in the structure and rigor of national rules. Some countries, for example, maintain a legal ban on the alteration of human germline cells. Some have less formal “guidelines” prohibiting germline editing, while others are more permissive in the regulatory constraints they place on the technology (Araki & Ishii, 2014; Ishii, 2017). Figure 1 displays a composite measure of national gene-editing regulations combining information from three recent surveys of regulatory policies. Countries are shaded according to regulatory rigor, with darker shades indicating more restrictive national rules.

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11 Baylis (2019) states that He Jiankui likely believed his controversial research was in-line with existing ethical standards due to a lack of a clear scientific moratorium and conflicting recommendations from experts: “He [Jiankui] believed he had checked all the boxes.”

12 For example, despite efforts by academic journals to ensure published research meets strict ethical guidelines, norm-violating research can often be published in second-tier journals (Baylis, 2019).

13 According to Ishii (2017), this group includes Canada, Brazil, Australia, and much of Western Europe.

14 Source data are from Isasi et al. (2016), Araki & Ishii (2014), and Baylis et al. (2020). For details on these measures and the construction of the composite measure, see Section 4.1.
Figure 1: National Regulation of Gene-editing Technology. Thirty-nine countries are rated by the permissiveness of national gene-editing technology regulations. Ratings combine data from Isasi et al. (2016), Araki & Ishii (2014), and Baylis et al. (2020). See Section 4.1 for details on the coding and source data.

Inconsistent rules across countries stem, in part, from different national historical experiences and cultural expectations regarding the appropriate use of gene-editing technology. For example, Germany’s experience with unethical human subjects experiments during the Nazi regime and their subsequent societal accounting with these human rights violations has conditioned the state’s regulation of human subjects research.\footnote{Bioethicist George Annas states, “Germany has been very reluctant to get involved with anything that could lead to a re-introduction of eugenic practices in their society” (Begley 2019).} South Korea developed relatively strict biological research guidelines in response to a high-profile controversy regarding the falsification of data in a cloning experiment by Seoul University investigator Woo Suk Hwang (Resnik et al. 2006). The United States developed comparatively weaker regulations for gene editing, consistent with a policy process that is more receptive to industry influence.\footnote{One biotech investor referred to the regulation of gene-editing technology in the US as “the Wild West” (Interview by authors, 9/21/2020).} As described above, China also has come under criticism for lax ethical guidelines...
In many cases, however, national regulations have simply not kept up with rapid advances in the field (Baylis 2019).

At the global level, there is growing interest in international coordination. Notably, the prospects for multilateral cooperation have not been not plagued by the strong political cleavages common to other issue areas (e.g., geopolitical rivalries or North-South divisions). A set of legacy international agreements, negotiated in the 1990s in reaction to concerns about cloning, provide a precedent for global governance of genetic research. The 1997 Oviedo Convention prohibits the misuse of innovations in biomedicine (Oviedo 1997). Its 34 signatories – all members of the Council of Europe – have agreed to ban the cloning of human beings and prohibit genetic screening for non-health purposes. In a similar declaration, members of the United Nations Educational, Scientific, and Cultural Organization (UNESCO) unanimously adopted the Universal Declaration on the Human Genome and Human Rights in 1997 (UNESCO 1997). Subsequent UNESCO declarations articulated norms about genetic data and trade in genetic resources (UNESCO 2003).

In recent years, formal international institutions have been slow to develop rules despite calls for new global standards. In 2016, environmental activists unsuccessfully pushed for the UN Convention on Biological Diversity to expand its mandate to regulate synthetic biology and gene drive organisms (Callaway 2016). The World Health Organization is among the few international organizations explicitly addressing the issue, establishing an Advisory Committee on Human Genome Editing that remains in preliminary stages. The Committee’s stated goal is to “advise and make recommendations on appropriate governance mechanisms for Human Genome editing.” More informally, a transnational network of National Academies of Science and Medicine are collaborating to evaluate social and ethical challenges in gene editing research and incorporate these issues into research protocols.

17 The Committee is described at https://www.who.int/ethics/topics/human-genome-editing/committee-members/en/.
In summary, regulation of gene-editing technology has been characterized largely by a decentralized national approach and inconsistency across jurisdictions. We now turn to our theoretical argument regarding the spillovers that afflict governance in this environment.

3 Interdependence & Technological Innovation

We argue that recent advances in gene editing, like many other disruptive technologies, increase international interdependence. We identify two specific sources of interdependence — regulatory arbitrage and spillovers from public controversies — through which policy decisions in one country affect outcomes in another. In each case, we specify the underlying conditions that give rise to interdependence, draw analogies to other issue areas, and specify observable implications that are tested in Section 4. While our primary focus is gene editing, we argue that these linkages are likely to recur in other emerging fields such as artificial intelligence and cybersecurity.

3.1 Regulatory Arbitrage

Gene-editing technologies are inputs to an array of commercial applications that are expected to grow substantially over the next decade. In 2019, the gene-editing market was estimated to be worth $3.8 billion and is projected to exceed $10 billion in the next five years. The most direct applications are in the pharmaceutical and healthcare industries, where firms are developing gene therapies to address a range of disorders and chronic illnesses. Among these are CRISPR Therapeutics, co-founded by Nobel laureate Emmanuelle Charpentier to develop gene-based medicines, which went public in 2016 and has since increased more

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18 Projections are from Ugalmugle & Swain (2020). McKinsey estimates the broader biotechnology sector to have direct economic impact $2-4 trillion annually in the next 10-20 years (Chui et al., 2020).
than tenfold in market value.\footnote{CRISPR Therapeutics (CRSP) closed at $13.82 on its first day of trading in October 2016. Its closing price on February 8, 2021 was $169.23.} Other sectors like agriculture, veterinary medicine, and industrial production processes also increasingly draw on gene-editing technology (Brinegar \textit{et al.} \citeyear{brinegar2017}).

The competition for these potential economic returns is fierce. Patent applications associated with gene-editing technologies have grown from less than 1,500 in 2000 to over 12,000 in 2019.\footnote{Data on patent applications are from \texttt{lens.org}, a publicly available patent database, and reflect searches for “gene editing.”} Firms are racing to develop commercial applications and successfully navigate regulatory hurdles to exploit the rapid market growth in gene editing. As in other emerging fields, pioneer firms may gain a first-mover advantage that endures even as competitors subsequently enter the market (Lieberman & Montgomery, \citeyear{lieberman1988} Agarwal & Gort, \citeyear{agarwal2001}).

Consequently, researchers and firms are highly motivated to accelerate the development and commercialization of gene editing technology. One strategy for doing so is to seek out more permissive regulatory jurisdictions. While there are longstanding concerns about regulatory arbitrage in genetic research,\footnote{Notably, the potential for regulatory arbitrage by gene-editing researchers was raised by the geneticist Irving P. Crawford when scientists first developed recombinant DNA technology in the 1970s. Crawford argued that researchers would simply evade strict regulations, citing several clinical trials that had moved to Europe and South America to sidestep burdensome rules in the United States (Baskin \textit{et al.} \citeyear{baskin2016}).} we argue that the recent revolution in gene-editing technology has substantially \textit{increased} the potential for arbitrage. Reduced costs have expanded access to a more diverse set of actors in more jurisdictions. The ability to apply gene editing more easily and cheaply increases cross-border mobility, allowing human and financial capital to select into countries with less rigorous rules.

The ability of actors to opportunistically select into other jurisdictions creates several problems for governments as they regulate gene editing. First, it reduces a government’s ability to restrict the use of the technology. Prior to recent technological breakthroughs – when altering genes was performed using difficult and expensive recombinant DNA tech-
niques, for example – gene editing was constrained to a handful of institutions in a handful of countries with the funding and infrastructure to support such methods. With few alternatives, researchers were largely forced to accept the regulations that governments imposed. The advent of programmable nucleases like CRISPR significantly increased the exit options available to scientists in academia and industry. This shift increased their ability evade unfavorable rules and weakened the hand of national governments.

A second problem is an increase in regulatory competition among national governments. The economic returns associated with biotechnology, coupled with the potential relocation of researchers, firms, and investment flows into more permissive environments, creates strong pressures to lower regulatory barriers. For example, European plant breeders have pressed the EU to relax restrictions on gene editing, arguing that existing rules put them at a competitive disadvantage. South Korea is reviewing its rules on gene therapy research in order to maintain its competitiveness in medical technology (Ji-young, 2017). In the United States, the agricultural industry successfully lobbied the government to weaken restrictions on gene-edited crops and livestock (Cancryn & Crampton, 2021; Stokstad, 2020). More broadly, commentators argue that China’s progress in biotechnology and America’s “eroding technological advantage” necessitate regulatory reforms and increased public investment (Darby & Sewall, 2021). Other countries have announced similar regulatory reviews or new public initiatives to capitalize on gene-editing technologies. Since lax regulations can reap economic returns, governments have an incentive to undercut each other’s rules. This can generate a deregulatory spiral in which governments collectively weaken standards.

In international relations scholarship, regulatory competition is most closely associated

\[\text{22See } \url{https://www.mpg.de/13748566/position-paper-crispr.pdf} \]

\[\text{23For example, policymakers in New Zealand are reviewing the country’s gene-editing regulations, which some claim are outdated (Morton, 2019). In Russia, the government recently announced a collaboration with Rosneft, the country’s largest oil producer, to develop gene-editing technology (Morton, 2020). For an overview of policy reforms regarding gene editing in agriculture, see Schmidt et al. (2020).}\]
with issues like tax policy, financial regulation, and environmental standards (Trachtman 1993; Bretschger & Hettich 2002; Angelini & Cetorelli 2003; Konisky 2007; Genschel & Schwarz 2011). These are policy domains in which nations differ in their regulatory approach, economic output is responsive to rules, and assets have high cross-border mobility. We argue that these conditions increasingly characterize the field of gene editing. More broadly, the conditions are likely to develop in other fields undergoing rapid technological change. Technological advancement often lowers entry barriers and encourages diffusion to a wider set of regulatory jurisdictions. It frequently occurs in industries with enormous growth potential and where initial economic advantages can yield substantial returns. As a result, we should expect emerging technologies to exhibit higher levels of regulatory arbitrage and the potential for more intense regulatory competition.

An implication of this argument is that gene editing will be governed by a weaker set of regulations than would exist with fewer arbitrage opportunities. If the CRISPR revolution enables firms and researchers to relocate gene-editing activity to more permissive jurisdictions — a hypotheses we test in Section 4 — we should expect an increase in the systemic risk of accidents or misuse. The repercussions of these controversies are discussed in the following section.

3.2 Spillovers in Public Controversies

The second source of interdependence is public attitudes about emerging technologies. Like other fields, continued progress in gene editing requires maintaining a high level of public confidence. Public attitudes affect the trajectory of the technology in at least three ways. First, beliefs about the safety and morality of gene editing shape consumer demand for gene

\[\text{Notably, other processes can sometimes counteract competitive pressures to generate a “race to the top” (Genschel & Plumper 1997; Prakash & Potoski 2006; Barkin 2015). In particular, if jurisdictions with sufficient market size adopt stringent rules, these can encourage higher standards elsewhere. Our argument is not that competitive pressures will dominate these countervailing forces, but that the pressure of regulatory competition increases in the face of rapid technological advancement.}\]
therapies and other products. Aiyegbusi et al. (2020), for example, identify public perceptions of gene therapies as “central to their uptake and use.” Second, public opinion affects the ability of firms to attract investors. Historically, controversies regarding one application of gene-editing technology have extinguished investor interest more broadly (Gardner 2020). Finally, public attitudes influence regulation, which determines the permissible ends to which the technology may be applied.

We argue that public attitudes about disruptive technologies are often fragile. Rapid technological advances challenge existing systems of practice and thought. The ramifications of disruptive technologies frequently do not nest neatly into existing ideological or political cleavages; instead, they create unexpected coalitions and give rise to a mix of emerging public narratives. As a result, we view emerging technologies as particularly vulnerable to backlash when controversies arise. If the technology intersects traditional political divides, elites may lack incentives to provide a narrative for individuals to anchor their own beliefs (Druckman et al. 2013). With no pre-existing reference frame to “fix” individuals’ views and moderate extreme reactions, high profile events can create quick and profound shifts in public opinion. Controversies can spark public backlash, lead to reductions in public funding, and engender knee-jerk regulatory responses that constrain even responsible scientific activity.

The recent history of gene therapy provides an example of such backlash. In 1999, 18-year old Jesse Gelsinger joined a clinical trial for a developmental gene therapy treatment run by the University of Pennsylvania. Unlike the previous seventeen participants, Gelsinger suffered an unexpected immune response that ultimately lead to his death. The tragic loss of the teenager led to an immediate and precipitous drop in public trust in clinical applications for gene-editing technology. As Jennifer Doudna recalls, the incident “made the whole field of gene therapy go away, mostly, for at least a decade. Even the term gene therapy became

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25In the case of artificial intelligence, an overwhelming number of Americans (81%) “believe that robots and/or AI should be carefully managed,” a statistic which Zhang & Dafoe (2019) suggest will shape governance outcomes for this and other technologies.
kind of a black label.” (Rinde, 2019).

This example illustrates how individuals update their beliefs in a highly uncertain environment with few consistent cues and even fewer data points. These features engender instability in public attitudes. As a result, emerging technologies that rely on public support often progress in fits and starts, with periods of promising technological advancements interrupted by crises of public confidence. There is evidence for this dynamic in the related field of genetically-modified organisms, where media exposure to controversies has been found to meaningfully affect public opinion (Prakash & Kollman, 2003; Drezner, 2008; Vigani et al., 2012). Ciocca et al. (2021) similarly note the potential for “hype-induced backsliding” in the field of artificial intelligence.

Actors in both academic and industry are keenly aware that continued research depends on managing public anxiety about gene editing. Participants at a 2015 conference on gene editing, for example, explicitly called for slowing down the more controversial germline gene-editing research “in order to create a safe political space” (Isaacson, 2021, 288). Public recommendations for scientific and national regulation of gene editing frequently note “increasing legitimacy and trust” in the technology as a primary goal (Kuzma et al., 2018, 23).

Despite these calls for cautious progress, both firms and researchers have strong incentives to push the scientific frontier. The recent controversy surrounding He Jiankui’s alteration of human germline cells triggered a new round of public concern regarding the safety and propriety of gene-editing research. Unlike the Gelsinger tragedy, however, it unleashed a response that spilled across national borders. Calls for a global moratorium on some avenues of gene-editing research swiftly followed the revelation of He’s experiment (Lander et al., 2019). Recognizing the potential for public backlash, leading scientists were quick to condemn the

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26The legacy of Jesse Gelsinger’s death weighed heavily on top researchers in the field and depressed research output. Doudna remarked that, after the tragedy, “You didn’t want that in your grants. You didn’t want to say, ‘I’m working on gene therapy.’ It sounded terrible” (Isaacson, 2021, 279).
At an international conference, a senior colleague accused He of “jeopardizing the entire field of genetic engineering” (Isaacson, 2021, 306).

We conceptualize backlash to gene-editing controversies as a negative externality that readily spills across national borders. Countries receive several benefits from scientists who push the boundaries of gene-editing research. Successful innovation brings economic rewards in the form of marketable and profitable new technology and also enhances the prestige of the nation’s scientific establishment. There are also clear costs associated with lax regulation, including the potential for domestic public backlash, but these costs are not fully internalized by the home country. Like the technology itself, backlash diffuses across borders. Controversies may damage support for gene-editing research even in jurisdictions that are comparatively well-regulated.

While regulatory arbitrage has been studied and documented in other contexts, we are not aware of existing scholarship that examines the potential for spillovers in public backlash. Nonetheless, we expect that it is an important source of interdependence that occurs whenever a technology is associated with safety risks or ethical concerns. We argue that these are common traits of emerging technologies – from nuclear energy in the 1950s to artificial intelligence today – and expect public backlash to pose a recurring challenge for technology governance.

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27 Given this legacy, the He Jiankui case was remarkable not only for the ethical misconduct it displayed, but, as one participant stated, “the boldness of presenting this at a conference” (Author’s interview with biologist, 8/21/2020).

28 After initially heralding the achievement, China sentenced He and two colleagues to three years in prison for “illegal medical practice” (Cyranoski, 2020). Chinese scientists working in the field of gene editing expressed concern after the sentencing “that the international condemnation that followed He’s explosive announcement in 2018 might have a wider chilling effect on CRISPR work in China” (Cyranoski, 2020).
4 Empirical Tests

We present two sets of empirical tests of the theoretical logics described above. First, we leverage data on clinical trials and scientific employment patterns to examine patterns of international arbitrage. Because we expect the recent transformation of gene-editing technology to lower entry barriers and exacerbate forum shopping, we use the year in which CRISPR was introduced (2012) as a cutpoint in the analysis. We test whether countries with weaker gene-editing regulations benefited from this technological shock more than countries with strict regulations.

For the second test, we identify the presence of public backlash using observational data from Twitter and an original survey experiment on American respondents. We first track public sentiment toward gene editing within approximately one million tweets across several dozen countries before and after the He Jiankui controversy. We then experimentally test the effect of information about a hypothetical gene-editing controversy on public support for gene-editing research and policy. The experiment varies whether the controversy occurs domestically or in a foreign country, allowing us to test whether foreign misuse of gene-editing technology affects public attitudes towards gene-editing use and policy in the United States.

4.1 Regulatory Arbitrage

We argued above that actors can evade strict regulations by relocating to jurisdictions with weaker rules, and that this behavior should increase in the wake of significant technological breakthroughs. To test this claim, we first examine patterns of clinical trials devoted to the development of gene therapies. While this test cannot definitively establish forum shopping behavior, it can test whether the recent breakthrough in gene-editing technology benefited countries with more permissive regulations, compared to their more stringent counterparts.
To directly test whether researchers relocate to weaker jurisdictions, we also analyze employment patterns of over 65,000 gene researchers.

4.1.1 Data on National Gene-Editing Regulations

The independent variable in our analysis is the rigor of national regulations governing the use of gene-editing technology. To measure regulation of gene editing and its applications, we develop a composite national regulatory score drawn from three sources. First, Isasi et al. (2016) classify national regulations on a range of gene editing-related issues, including gene therapy, human germline editing, and genetic diagnosis. Each country is rated as “permissive”, “intermediate,” or “restrictive” on each issue; we transform these into a 1-3 scale of increasing regulatory rigor and average across the fields to generate a single national regulatory score. Second, Araki & Ishii (2014) provide a separate classification of countries based on the regulation of heritable genetic editing. Finally, Baylis et al. (2020) examine national rules regarding the use of genetically modified in vitro embryos in laboratory research.

These three measures are positive correlated (all greater than 0.4) but prioritize different applications of gene-editing technology. To get a broad measure of each country’s regulatory environment, we combine them via principal components analysis. This provides a continuous, cross-national measure of gene-editing regulation for 39 countries that engage in

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29No existing data measures a country’s broad approach to gene editing regulation. As a result, we draw from a variety of sources that touch on particular aspects of gene-editing research and clinical use.

30The categories include “ban based on legislation,” “ban based on guidelines,” “restrictive,” and “ambiguous.” We transform these into a 1-4 scale of increasing regulatory rigor.

31Baylis et al. (2020) provide a similar measures of gene-editing regulations regarding assisted human reproduction and those that focus on other applications. Since we are interested in rules regarding gene therapy development, we use the “not for reproduction” measure. The authors categorize countries’ regulatory approach as permissive, indeterminate, prohibitive with exceptions, or prohibitive based on a review of national legislation, guidelines, and codes of conduct.
research and clinical development in the field. Cross-national variation in these regulations is visualized in Figure 1. The most restrictive regulatory environments include Germany, Sweden, Switzerland, and Brazil. The most permissive are China, Ireland, and the United States. The regulatory are centered at zero and range from -2.1 to 3.3.

### 4.1.2 Gene Therapy Clinical Trials

Our first outcome is gene therapy clinical trials, which test the safety and efficacy of clinical applications of gene-editing technology. They are a necessary step to gain market authorization for gene therapies, a growing industry with high consumer demand and significant profit potential (Macpherson & Rasko, 2014; Hirakawa et al., 2020). Several high-profile gene therapies have been approved following successful clinical trials (June et al., 2018; Gong et al., 2018), and the race to develop new treatments has intensified in recent years.

We collect data on clinical trials from the Journal of Gene Medicine’s “Gene Therapy Clinical Trials Worldwide” database. They capture all registered trials that perform gene therapy clinical interventions on human subjects. We structure the data at the level of the country-year, such that the outcome variable represents the number of new gene therapy trials registered in each country in each year. We restrict the sample to the 39 countries for which we have data on national regulations. The data include a total of 3,535 clinical trials.

Figure 2 displays temporal trends in gene therapy trials located in countries with different levels of regulatory rigor. Each line represents the number of clinical trials registered in a particular country from 1989-2019. The dotted red line reflects the introduction of CRISPR in 2012. The top panel features countries with the most restrictive gene-editing regulations, followed by countries with intermediate regulatory environments (middle panel), and permissive jurisdictions (bottom panel). The figure provides some suggestive evidence that

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32 The three source measures vary widely in geographic coverage. Thirty-nine countries are classified by at least two sources. For these countries, we impute the missing scores before estimating the principal components.
regulatory rigor is positively associated with clinical trial development in the post-CRISPR era. Countries with weak regulations (e.g., the United States and Russia) experience both higher overall numbers of clinical trials and a sharper increase in later years compared to countries with strict policies (e.g., Germany).

To test the relationship between regulation and clinical trials more systematically, we employ a difference-in-differences specification. The analysis leverages the 2012 introduction of CRISPR as a technological shock that should interact with countries’ national regulations. We expect low-regulation countries to benefit more from this technology than countries with
more exacting rules. After 2012, technological innovation lowers the human and financial
capital needed to engage in gene-editing research. Cost and expertise recede as barriers to
commercial development, while regulatory restrictions are more likely to become binding
constraints. We therefore expect countries with weaker regulatory jurisdictions to attract a
larger share of gene therapy trials after 2012 than those in more rigorous jurisdictions. We
estimate the following linear model to determine how the regulatory environment affects the
quantity of a country’s gene therapy trials.

\[
\text{Trials}_{it} = \alpha + \beta_1 \text{Regulation}_i + \beta_2 \text{CRISPR}_t + \beta_3 (\text{Regulation}_i \times \text{CRISPR}_t) + \beta_4 X_{it} + \epsilon_{it} \quad (1)
\]

where \(\text{Trials}_{it}\) are the number of gene therapy clinical trials conducted in country \(i\) in year \(t\), \(\text{Regulation}_i\) is the composite measure of regulatory rigor, \(\text{CRISPR}_t\) is an indicator for the post-2012 period when CRISPR was introduced, and \(X_{it}\) is a vector of control variables. We
expect \(\beta_3\), the difference-in-differences coefficient, to be negative and statistically significant.
This would signify that CRISPR led to the acceleration of clinical trial development in weak
regulatory environments, compared to countries with more restrictive rules.

We include a set of control variables that reflect a country’s national economic output
and human capital, which are likely to shape gene therapy development. We include GDP to
account for each country’s economic capacity and GDP per capita for its level of develop-
ment. We proxy scientific capital with a count of the number of patent applications in each
country by residents Patent Residents and non-residents Patent Non-Residents. Data
for all control variables are drawn from the World Bank’s World Development Indicators
(WDI) for the years 1989-2020. Each is measured at the country-year level and logged to
account for skewed distributions.

Table 1 summarizes the results. Column 1 is a simple baseline model with national
regulations, an indicator for the CRISPR era, and an interaction between them. Column
2 includes the covariates, and column 3 adds country and year fixed effects. Our core findings are consistent with theoretical expectations. In all models, the negative coefficient for Regulation reflects a general pattern of fewer gene therapy trials among countries with strict regulatory standards. The large and positive coefficient for CRISPR is consistent with an increase in the use of gene-editing technologies after 2012. Most importantly, the interaction term Regulation × CRISPR is negative and statistically significant. This result confirms that the effect of Regulation on gene therapy development is greater following the technological shock of CRISPR. Substantively, a one-point reduction in regulatory rigor (e.g., moving from Finland to Italy) increases the number of gene therapy trials by approximately 2.4 per year in the pre-CRISPR era and 3.7 per year afterwards. This shift is quite meaningful given the mean value of 3.1 trials per country-year.

Other covariates perform largely as expected. Countries with higher levels of economic development conduct greater numbers of gene therapy trials. The two patent measures are generally associated with increased clinical trial development, though patent applications by residents has an insignificant effect in the specification in Column 2.

We summarize the trends for low- and high-regulation countries in Figure 3. To simplify the visualization, we classify countries as permissive (below the median regulation score) and restrictive (median or above). The 2012 technological shock is represented by the vertical dotted line. The figure provides some evidence in favor of the parallel trends assumption needed to causally identify the difference-in-differences test. In the pre-CRISPR period, the two groups move roughly in parallel over time. They begin to diverge after 2012, when low-regulation countries experience increasing numbers of gene therapy trials compared to their high-regulation peers.

Our analysis provides strong evidence that weaker regulatory environments “outperform” expectations in the conduct of gene therapy trials. This evidence is consistent with regu-

33 These estimates are from the model estimated in Column 2.
<table>
<thead>
<tr>
<th></th>
<th>(1)</th>
<th>(2)</th>
<th>(3)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Regulation</td>
<td>-1.207***</td>
<td>-2.381***</td>
<td>-144.500***</td>
</tr>
<tr>
<td></td>
<td>(0.365)</td>
<td>(0.297)</td>
<td>(15.500)</td>
</tr>
<tr>
<td>CRISPR</td>
<td>2.766**</td>
<td>1.390*</td>
<td>3.687*</td>
</tr>
<tr>
<td></td>
<td>(0.729)</td>
<td>(0.732)</td>
<td>(2.060)</td>
</tr>
<tr>
<td>Regulation × CRISPR</td>
<td>-1.106*</td>
<td>-1.338**</td>
<td>-1.711***</td>
</tr>
<tr>
<td></td>
<td>(0.579)</td>
<td>(0.564)</td>
<td>(0.376)</td>
</tr>
<tr>
<td>GDP (log)</td>
<td>2.817***</td>
<td>-6.645***</td>
<td></td>
</tr>
<tr>
<td></td>
<td>(0.431)</td>
<td>(1.589)</td>
<td></td>
</tr>
<tr>
<td>GDP per capita</td>
<td>0.002***</td>
<td>0.0004***</td>
<td></td>
</tr>
<tr>
<td></td>
<td>(0.0002)</td>
<td>(0.0001)</td>
<td></td>
</tr>
<tr>
<td>Patent Resident (log)</td>
<td>-0.441</td>
<td>2.875***</td>
<td></td>
</tr>
<tr>
<td></td>
<td>(0.442)</td>
<td>(0.493)</td>
<td></td>
</tr>
<tr>
<td>Patent Non-Resident (log)</td>
<td>0.601***</td>
<td>1.104***</td>
<td></td>
</tr>
<tr>
<td></td>
<td>(0.198)</td>
<td>(0.267)</td>
<td></td>
</tr>
<tr>
<td>Country FE</td>
<td>✓</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Year FE</td>
<td></td>
<td>✓</td>
<td></td>
</tr>
<tr>
<td>Observations</td>
<td>1,248</td>
<td>1,138</td>
<td>1,138</td>
</tr>
<tr>
<td>Adjusted R²</td>
<td>0.038</td>
<td>0.270</td>
<td>0.711</td>
</tr>
</tbody>
</table>

Table 1: Effect of National Regulations on Gene Therapy Clinical Trials. The table displays coefficient estimates and standard errors from a linear difference-in-differences model. Statistical significance is denoted by: *p<0.1; **p<0.05; ***p<0.01.

Laboratory arbitrage, since after 2012 trials are increasingly concentrated in countries with lax national rules. However, it does not definitively establish arbitrage behavior. The same result could be obtained if low-regulation countries experienced increases in domestic clinical trial development, rather than attracting firms and researchers from other jurisdictions. We now examine arbitrage behavior directly by analyzing employment patterns among gene
Figure 3: Trends in Gene Therapy Trials, Low vs. High Regulation Countries. The graph depicts the average number of clinical trials for high (black line) regulation countries and low (red) regulation countries. The vertical dashed line represents the introduction of CRISPR in 2012. Points on the graph are the (logged) number of gene-editing clinical trials in a given country in a given year.

4.1.3 Employment of Gene Scientists

To test for forum-shopping behavior, we examine whether gene scientists relocate to academic institutions in low-regulation countries at higher rates after the invention of CRISPR. If our theory is correct, institutions located in countries with more permissive regulatory standards will be more attractive destinations for scientists in the post-CRISPR era. States with the capacity to generate strict regulations for advanced technology may also be locations with high levels of scientific production and the know-how to develop clear rules. However, CRISPR should lower the barriers to entry for conducting high-quality research on an advanced topic even outside of established scientific hubs. If scientists choose to move
to institutions in low-regulation countries at higher rates in the post-CRISPR period, this would constitute evidence of forum-shopping.

To test this claim, we examine patterns of employment among researchers who have published scientific papers in the field of genetic engineering. We collect these data from PubMed, a database of life sciences and biomedical publications maintained by the US National Library of Medicine at the National Institutes of Health. To obtain the sample of gene researchers, we extract the names and institutional affiliations of all authors who have published papers on the topic “genetic engineering” over the past 20 years. The search yields approximately 150,000 papers and more than 400,000 unique gene researchers for the period 2002-2021.  

We are interested in the employment choices made by gene-editing researchers, once they have decided to leave their home institution. Rather than model the choice to exit an institution, we condition on exit and examine the decision about where to relocate. To do so, we first subset to the 65,955 researchers who transfer institutions at least once during the time period. We structure our data at the level of the “potential transfer” dyad: a pairing of the researcher’s original country and each of the 39 potential destination countries for which we have data on national gene-editing regulations. If the researcher Jane Smith leaves Yale University for McGill University in 2012, for example, we add 39 observations to our dataset: each lists the United States as the home country and one of the regulatory jurisdictions (e.g., United States, Israel, Canada) as the destination country. The outcome, Observed Transfer, equals one for the US-Canada pairing and zero for the other observations. The resulting dataset includes approximately 6.3 million observations.

This data structure allows us to examine how differences in national regulatory environments shape employment patterns over time. We measure these differences by subtracting

34 We begin in the year 2002 because this is the first year PubMed records full author names for each publication.
the destination regulatory score from the researcher’s country of current employment, creating the variable \textit{Regulatory Difference}. As before, we interact this variable with an indicator for the post-2012 era when CRISPR technology enhanced forum-shopping opportunities. We also include the time-varying covariates from above: GDP, GDP per capita, and patents for both the origin and destination countries. Finally, we add an indicator for same-country pairings — observations in which a researcher finds employment in the same country as the institution he or she departed — to account for the propensity to remain in the same country.

Our theoretical expectations about forum shopping imply that transfers with lower \textit{Regulatory Difference} scores (representing movement to weaker jurisdictions) should become more attractive in the post-2012 period, compared to the prior era. This should manifest as a negative coefficient on the \textit{Regulatory Difference} \times \text{CRISPR} interaction term.

We report the results of linear probability models with and without the inclusion of covariates in Table 2. In both models, the interaction term \textit{Regulatory Difference} \times \text{CRISPR} is negative and statistically significant. This result confirms that the attractiveness of employment in permissive regulatory environments is greater following the technological shock of CRISPR. This effect is small but not insubstantial: a one-unit increase in the destination regulation score decreases the likelihood of researcher employment relocation by 0.4% in the pre-CRISPR era to 0.6% after 2012 (Column 1). These estimates are relative to a baseline rate of relocation of 1.4%.
Table 2: Employment Relocation of Gene Researchers: Linear probability model estimates for the likelihood that a gene-editing researcher who exits his or her home institution will move to an institution in another country. Column 2 includes the following controls (not shown): GDP origin country, GDP destination country, GDP per capita origin country, GDP per capita destination country, Patent Applications origin country, and Patent Applications destination country. Statistical significance is denoted by: *p<0.1; **p<0.05; ***p<0.01.

Together, these tests provide evidence consistent with forum-shopping behavior induced by low-cost, easy-to-use, disruptive technology. We now turn to the second hypothesized mechanism related to spillovers in public backlash.

4.2 Public Backlash

We first establish the presence of international public opinion backlash against the controversial use of technology using social media data. Using tweet content and location data, we
test for the effects of controversy on negative sentiment and moral outrage in tweets of users outside of the controversy’s jurisdiction (China). We then confirm that international controversy has a negative effect on domestic policy attitudes towards gene-editing technology using a survey experiment on an American population sample.

4.2.1 Observational social media data

We follow Müll er et al. (2020) in scraping and analyzing tweets with the keyword CRISPR. Using Twitter’s API through Barrie & Ho (2021)’s R package, academictwitter, we pulled approximately 50,000 tweets that contain the word “CRISPR” in the 50 days prior to and after the He Jiankui controversy. The bottom panel of Figure 4 shows a histogram of the appearance of CRISPR in tweets over this time period. Using a bag-of-words procedure, we take the average sentiment of each tweet by identifying the proportion of positive words in the tweet. Higher proportions indicate higher levels of positive sentiment. Only English-language tweets are included in the sentiment analysis. Only tweets with at least one word with a positive or negative valence are included in this sample. The top panel of Figure 4 displays change in average sentiment over time.

While sentiment in tweets does capture overall public opinion of Twitter users towards CRISPR technology, new research suggests that certain forms of expression on social media are more likely to drive conversations (Brady et al. 2021). In particular, tweets expressing moral outrage receive higher levels of positive feedback online and are therefore more likely to be seen and to influence online sentiment (Brady et al. 2021). We use Brady et al. (2021)’s measure of moral outrage to understand whether controversial events in CRISPR technology influence not only sentiment, but the type of language that drives greater engagement with CRISPR.

35We exclude replies and retweets in our analysis.
### Table 3: Pre-post analysis

<table>
<thead>
<tr>
<th>Dependent variable:</th>
<th>Sentiment (1)</th>
<th>Moral Outrage (2)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Post-He Jiankui</td>
<td>$-0.110^{***}$</td>
<td>$0.062^{***}$</td>
</tr>
<tr>
<td></td>
<td>(0.006)</td>
<td>(0.004)</td>
</tr>
<tr>
<td>Constant</td>
<td>$0.609^{***}$</td>
<td>$0.158^{***}$</td>
</tr>
<tr>
<td></td>
<td>(0.005)</td>
<td>(0.003)</td>
</tr>
<tr>
<td>Observations</td>
<td>21,856</td>
<td>50,839</td>
</tr>
</tbody>
</table>

The table reports the pre-post change in sentiment and outrage expressed in tweets.

We note that moral outrage is a form of negative sentiment. We measure the number of words stems associated with moral outrage in each tweet. The middle panel of Figure 4 displays change in moral outrage over time. Table 3 reports the pre-post change in sentiment and outrage expressed in tweets.

Subsetting the data to only tweets with identifiable geolocation data for the associated Twitter user, we replicate the main analysis by user country. We limit our analysis to English-language tweets, which likely affects the composition of countries in this sample. Figure 5 displays the results for countries with more than 10,000 unique tweets that mention CRISPR. As Figure 5 shows, the revelation of the gene-editing controversy produced negative sentiment in the days afterwards for every country in the sample. Importantly, the scandal

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36 Indeed, as confirmation of this concept, we find that CRISPR tweets with more moral outrage were retweeted and liked at higher rates. See Appendix A.5.2.

37 For a full glossary of moral outrage terms, see Appendix A.5.1.
Figure 4: **CRISPR tweet sentiment**: Top panel: sentiment analysis of tweets including the word “CRISPR” from September 2018 - January 2019. Higher values indicate more positive sentiment. Middle panel: moral outrage in same sample of tweets. Higher values indicate more moral outrage. Bottom panel: histogram of number of tweets per day. Dashed black line on November 26, 2018, the day the He Jiankui controversy became public. Data collected by authors.
Figure 5: *CRISPR tweets by location*: Sentiment analysis of tweets including the word “CRISPR” from September 2018 to January 2019 by Twitter user location. Only countries with over 10,000 CRISPR-related tweets included. Higher values indicate more positive sentiment. Blue line is a fitted loess model; grey bar indicates 95% confidence interval. Dashed black line on November 26, 2018, the day the He Jiankui controversy became public. Data collected by authors.

...did not occur in any of these countries. (China is not included in the sample as the country blocks access to Twitter for regular users).[^38]

4.2.2 Survey Experiment on Public Backlash

The observational data provides suggestive evidence that the controversial application of gene-editing technology decreases the public’s trust in the technology. We next examine the causal effect of information about controversial use of gene editing on public opinions about scientific policy via a survey experiment. The survey examines backlash among the general public in response to a hypothetical, norm-violating application of gene-editing technology. To gauge the spillover effect, we examine both the effect of controversial activity in one’s own country as well as activity in a foreign country.

The online survey was conducted in July 2020 on the survey platform Lucid on a sample of 1200 Americans quota-sampled to US census margins. Table A1 in the appendix displays summary statistics of our survey sample. For our primary results, we use the full sample of survey respondents. To address concerns that changes in Lucid recruitment methods may depress respondent quality, we ensure our findings hold after removing those who do not successfully pass attention checks (see Appendix Table A4).

We embed an experiment in the survey designed to address two questions. First, do controversies over the use of gene editing reduce public support for the technology and its potential applications? Second, does public backlash spill across national jurisdictions?

In the experiment, all respondents receive a basic summary of gene-editing technology. It reads:

All organisms, from bacteria to lizards to humans, have molecules called DNA, or deoxyribonucleic acid. These DNA molecules contain the genetic code for each organism. DNA provides the instructions that determine an organism’s physical characteristics and control how it develops, functions, and reproduces.

In recent years, scientists have developed new gene-editing technologies that can

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39In Appendix A.3, we list the full survey text and discuss its alignment with APSA Principles and Guidance for Human Subjects Research. Our pre-registration plan can be found under EGAP 20200505AA.
permanently alter an organism’s DNA. These technologies allow scientists to make targeted changes to DNA molecules in plants and animals, modifying their biological traits. For example, scientists have edited the genes of wheat plants to make them easier to grow.

Following this introduction, some respondents are randomly assigned to a treatment condition where they are given additional information about a gene-editing controversy. Among treated respondents, we randomize whether the controversy occurs in the US, UK, or China. Appendix Table A2 summarizes covariate balance across the randomized treatment conditions. In the survey, the treatments are presented as a hypothetical news article set in the year 2021. To increase external validity, we model the experimental intervention on the real-world controversy surrounding He Jiankui. We present the text for the UK treatment condition here:

**Birth of Genetically Altered Babies in the UK Provokes Outcry**

January 25, 2021

[LONDON]— A British research team announced that they have used a new gene-editing technology to alter the DNA of a group of infants. In an unprecedented intervention, scientists on the research team deleted a set of genes believed to be linked to breast and prostate cancer. The deleted genes are not considered essential to basic biological functions in humans, but the long-term effects of their removal are unclear. The research team plans to periodically examine the infants throughout their lives to assess any side effects of the genetic alteration.

The disclosure this week of the research — carried out in the UK — has sparked urgent debate about the ethics of genetic alteration. The infants’ birth represents a significant and controversial leap in the use of gene-editing technology. The British study has also increased concerns about a future in which parents produce
“designer babies” with selectively improved traits, such as height or intelligence.

After treatment assignment, respondents are asked to rate their agreement with four statements on a scale of 0 (no agreement) to 10 (complete agreement). The statements read as follows:

- Research in the US involving gene editing should be more strictly regulated.
- US patients should have access to medical treatments that involve gene editing.
- The US government should provide funding for gene editing research.
- Most US scientists conduct their research in a safe and responsible manner.

The statements estimate the extent of public confidence in the safety of gene-editing technology and support for continued development. Respondents’ answers constitute our dependent variables in the analyses below.

Our theory of public backlash against emerging technologies implies two patterns of response. First, we expect that respondents who read about a controversy in their own country will be less supportive of gene-editing research. This “domestic public backlash” should heighten demand for strict regulation, depress calls for patient access to gene therapy, decrease support for funding gene-editing research, and decrease confidence in the safety of scientific research. Second, we expect foreign controversies to similarly reduce public support for gene editing among US respondents. A “public backlash spillover” occurs if the controversial use of gene editing generates a domestic backlash even when the scandal occurs in another country.

We report treatment effects for each outcome of interest in Figure 6. Coefficients in the figure represent the treatment effect of exposure to a gene-editing controversy, compared to the control (no controversy) condition.\footnote{See Table A3 in the appendix for the full set of point estimates and standard errors.}

Within each panel, we display the estimated treat-
Figure 6: Public Response to Gene-Editing Controversy. The figure shows the treatment effect of learning about hypothetical, controversial gene-editing research in the United States, United Kingdom, and China. The four panels report effects on the four dimensions of public support listed above. 95% confidence intervals are reported.

We find evidence of a domestic public backlash in three of four outcomes. When American respondents read about a hypothetical misuse of gene-editing technology by American
researchers, they significantly reduce support for patient access to gene therapies and public funding for gene-editing research. They also have diminished perceptions of the safety and responsibility of scientific research in the United States. On average, the domestic controversy treatment shifts opinion on each of these outcomes by approximately 0.5 points. Contrary to expectations, respondents do not increase demand for strict regulations in reaction to domestic gene-editing scandals. This null finding may reflect a ceiling effect, as even respondents in the control condition call for strict regulations in high numbers (see Appendix Figure A3 for the distribution of responses across conditions).

There is clear evidence that backlash is not limited by national jurisdiction. As in the domestic scenario, neither of the foreign scandals significantly affects attitudes about gene-editing regulation. However, support for public funding of gene-editing research significantly decreases in response to foreign controversies in the UK and China. Additionally, both confidence in the responsibility of US scientists and support for expanded patient access to gene therapies decrease in the China and UK condition, though the UK treatment does not obtain conventional levels of statistical significance. Notably, the effects of domestic and foreign controversies are statistically indistinguishable across all four outcomes.

Together, these results suggest that the public does not discriminate between domestic and foreign research controversies. We find clear evidence that, for some public policy outcomes, the spillover effect of controversial research in one national jurisdiction negatively impacts domestic support for gene editing in another jurisdiction.

4.3 Discussion

Our findings provide evidence for two sources of interdependence that afflict national governance of gene-editing technology. We show that weaker national regulations boost gene therapy development, creating pressures for governments to engage in regulatory competition. We also demonstrate that the risks of weak national rules are not fully internalized
by the home country. If a government’s lax regulation increases the risk of inappropriate behavior, the resulting backlash spills across national boundaries.

Taken together, our results suggest that governments have compelling incentives to lower regulatory barriers beyond the level they would otherwise prefer. Each country can obtain individual economic benefits from weakening rules, while the risks of doing so are diffusely spread across multiple jurisdictions. If governments respond rationally to these incentives, effective regulation will be under-produced and the systemic risk of misuse will rise. However, the resulting public backlash in domestic constituencies could reduce demand for the technology or create a swell of public support for higher regulation of the technology, stymieing future progress. While public backlash could correct for under-regulation, the high volatility of this cycle creates an unstable atmosphere for technological innovation.

By illuminating patterns of interdependence among states, our arguments examine the functional case for global governance of gene-editing technology. Functionalists argue that states build international institutions to manage transnational spillovers, resolve information problems, and reduce transaction costs (Keohane 1982, 1984; Haggard & Simmons 1987). They view global governance bodies as a rational response to “political market failures” among states with fixed preferences (Keohane 1984). Though we adopt a functional framework in this paper, we acknowledge that other perspectives may generate valid arguments for or against a global gene-editing regime.

There are at least two functions an international institution could provide to improve governance of gene editing. First, it can encourage partial harmonization of the disparate national and subnational rules governing the technology. While it is politically-infeasible to expect countries to delegate regulatory power to a supranational body, a global gene-editing

\[\text{Keohane 1982, 1984; Haggard & Simmons 1987.}\]

41For example, Buchanan & Keohane (2015) adopt a primarily normative lens in arguing for a drone accountability regime. Johnston (2001) argues that international institutions can be used as vehicles for persuasion and social influence, changing state preferences. Bioethicists working in the field of gene editing have explicitly called for ethically-based global standards informed by public discussion and attention to local cultures and environments (Kofler et al. 2018).
regime could establish a floor of basic ethical and safety protections that all countries are expected to follow. This would limit the scope for regulatory competition and reduce the risk of scandalous applications of gene-editing technology.

Second, an international institution could assuage public anxiety by establishing clear norms of appropriate use and monitoring gene-editing applications to human subjects. As we argued above, one reason technological scandals generate wild fluctuations in public support is that citizens lack a coherent frame of reference for understanding the risks and benefits associated with disruptive technologies. International institutions can help fill this void by articulating norms that reflect a broad-based global consensus.

If public anxiety is rooted in the belief that existing governance efforts are insufficient, rigorous monitoring by international bodies may increase public confidence and support. To probe this idea, we added a corollary condition to our public backlash experiment in which the hypothetical news article alerts readers to monitoring by the World Health Organization (WHO). We found that this treatment failed to improve respondent confidence, suggesting the mere existence of monitoring does not substantially reduce public concern.

Finally, our findings have institutional design implications for a potential global governance regime. Scholars argue that international regimes are designed to address the unique cooperation problems states confront in an issue area (Martin, 1992; Koremenos et al., 2001). By theorizing and testing two patterns of state interdependence distinctive to gene-editing technology, we specify a set of important issues that international institutions should address. Given the widespread diffusion of gene-editing technology, an effective regime would need to be broad-based in membership and participation. This could be achieved by nesting

42 In these conditions, we add the following language to the news article: “Gene-editing research is closely monitored by the World Health Organization (WHO), which has established guidelines to ensure scientific activities are conducted safely and carefully. The study is currently under review by the WHO to assess its compliance with these guidelines.”

43 See Appendix A.4 for full results from this treatment arm.
the institution within an existing global body like the WHO. Our analysis of public back-
lash also suggests the regime should prioritize building resilience in public confidence. This
requires investing in features that increase the legitimacy of the institution in the eyes of the
public, including transparency and political impartiality.

5 Conclusion

The pace of innovation is sped up by international participation in research processes. With
competitive international systems providing incentives for states to invest in the adoption of
new technology (Milner & Solstad, 2020), the diffusion of novel, disruptive technology is a
key feature of the twenty-first century (Bloom et al., 2021). Scientific competition exacer-
bates the pace of technological advancement as scientists seek knowledge, status, and market
opportunities through innovative discoveries. As the pace of regulation struggles to keep up
with the speed of innovation, controversial applications and misuse of new technology are
likely to increase as more scientists engage in the field of research. However, the cost of
scientific scandals on public support for the technology is high. Controversial applications
of technology in one country can cause public backlash to spill over to other countries. The
opportunities for scientific risk created by new technology and enabled by low regulation
of this technology can lead to public outrage that can dampen technological progress. Cy-
cles of hyper progress and regressive backlash generate an inefficient environment for stable
technological progress.

In this paper, we provide evidence for international interdependence created by techno-
logical advancement. We demonstrate first that states are subject to regulatory arbitrage by
scientists and practitioners in the field of gene-editing. The accessibility of cheap, powerful
technology creates opportunities and incentives for regulatory arbitrage among industries
and scientists. To measure regulation, we construct a new index of gene-editing restrictions
across three gene-editing issues: gene therapy, germline editing, and in-vitro embryonic modification. Using two novel sources of data on gene therapy trials and scientific employment, we find strong support for the theory of arbitrage in gene therapy clinical trial data in the post-CRISPR era.

We then define a new form of interdependence, public backlash spillovers, which we identify using observational social media data and test with an online survey experiment. Data from Twitter suggest that real-life scientific controversies in the realm of gene-editing produce backlash from users across the globe. Results from our survey experiment on Americans show that this same information about foreign gene-editing scandals reduces trust in and support for funding for domestic gene-editing research.

While our tests focus on the field of gene editing, we argue these dynamics recur in the governance of disruptive technological innovations more generally. Future work can assess the generalizability of our theory by expanding empirical tests to other fields. In addition to contemporary emerging technologies like artificial intelligence, historical disruptions such as the nuclear energy and computer revolutions may shed light on whether international coordination was motivated in part by these mechanisms. We also hope our paper will inspire studies of other relatively “ungoverned” issue areas in world politics. While examining long-standing issues of global governance like trade and arms control are undoubtedly important, scholars of international cooperation should not neglect important policy domains that are not yet subject to multilateral regimes.
References


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*Available at SSRN 3312874.*
## Appendix

### A.1: Tables

<table>
<thead>
<tr>
<th>Variable</th>
<th>Sample Proportion</th>
</tr>
</thead>
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<td><strong>Party ID</strong></td>
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<td>Democrat</td>
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</tr>
<tr>
<td>Republican</td>
<td>0.36</td>
</tr>
<tr>
<td>Independent</td>
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</tr>
<tr>
<td><strong>Age</strong></td>
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</tr>
<tr>
<td>18-30</td>
<td>0.15</td>
</tr>
<tr>
<td>31-45</td>
<td>0.36</td>
</tr>
<tr>
<td>46-60</td>
<td>0.30</td>
</tr>
<tr>
<td>over 60</td>
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</tr>
<tr>
<td><strong>Education</strong></td>
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<tr>
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<tr>
<td>Some College</td>
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<tr>
<td>Bachelor’s Degree</td>
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<td>Post-Graduate</td>
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</tr>
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<td>Male</td>
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</tr>
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<tr>
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<tr>
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</tr>
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</tr>
<tr>
<td>No</td>
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<tr>
<td>$50-75,000</td>
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</tr>
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<td>&gt; $75,000</td>
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<td><strong>Region</strong></td>
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<tr>
<td>Northeast</td>
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<tr>
<td>Midwest</td>
<td>0.20</td>
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<tr>
<td>South</td>
<td>0.38</td>
</tr>
<tr>
<td>West</td>
<td>0.22</td>
</tr>
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</table>

Table A1: Survey sample statistics. For each category, we report the proportion of respondents who fit into the category among those that answered the relevant question.
Table A2: Balance Table of Covariates in Survey Experiment

<table>
<thead>
<tr>
<th></th>
<th>control (N=331)</th>
<th></th>
<th>uk (N=285)</th>
<th></th>
<th>china (N=292)</th>
<th></th>
<th>us (N=299)</th>
<th></th>
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</thead>
<tbody>
<tr>
<td></td>
<td>Mean</td>
<td>Std. Dev.</td>
<td>Mean</td>
<td>Std. Dev.</td>
<td>Mean</td>
<td>Std. Dev.</td>
<td>Mean</td>
<td>Std. Dev.</td>
</tr>
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<td>Income (1-24)</td>
<td>9.3</td>
<td>7.4</td>
<td>8.9</td>
<td>7.3</td>
<td>9.3</td>
<td>7.4</td>
<td>9.0</td>
<td>7.2</td>
</tr>
<tr>
<td>Education (1-8)</td>
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<td>2.0</td>
<td>4.4</td>
<td>2.0</td>
<td>4.3</td>
<td>1.9</td>
<td>4.5</td>
<td>2.0</td>
</tr>
<tr>
<td>Female</td>
<td>0.5</td>
<td>0.5</td>
<td>0.5</td>
<td>0.5</td>
<td>0.5</td>
<td>0.5</td>
<td>0.5</td>
<td>0.5</td>
</tr>
<tr>
<td>Age</td>
<td>43.5</td>
<td>17.0</td>
<td>44.4</td>
<td>16.2</td>
<td>46.1</td>
<td>17.3</td>
<td>45.0</td>
<td>17.1</td>
</tr>
<tr>
<td>Party (1-10)</td>
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<td>3.5</td>
<td>5.4</td>
<td>3.5</td>
<td>5.3</td>
<td>3.6</td>
<td>5.1</td>
<td>3.6</td>
</tr>
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<td>Political Attention (1-5)</td>
<td>3.6</td>
<td>1.2</td>
<td>3.5</td>
<td>1.2</td>
<td>3.5</td>
<td>1.2</td>
<td>3.6</td>
<td>1.2</td>
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Table A3: Survey Results. Estimated treatment effects and robust standard errors for the survey experiment. Effects are relative to the control condition (no additional informatton). Statistical significance is denoted by: *p<0.1; **p<0.05; ***p<0.01.
<table>
<thead>
<tr>
<th></th>
<th>Regulations</th>
<th>Access</th>
<th>Funding</th>
<th>Safety</th>
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</thead>
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<tr>
<td></td>
<td>(1)</td>
<td>(2)</td>
<td>(3)</td>
<td>(4)</td>
</tr>
<tr>
<td>US Controversy</td>
<td>-0.109</td>
<td>-0.644***</td>
<td>-0.572**</td>
<td>-0.416**</td>
</tr>
<tr>
<td></td>
<td>(0.197)</td>
<td>(0.223)</td>
<td>(0.231)</td>
<td>(0.199)</td>
</tr>
<tr>
<td>UK Controversy</td>
<td>-0.531**</td>
<td>-0.742***</td>
<td>-1.165***</td>
<td>0.080</td>
</tr>
<tr>
<td></td>
<td>(0.237)</td>
<td>(0.269)</td>
<td>(0.286)</td>
<td>(0.219)</td>
</tr>
<tr>
<td>China Controversy</td>
<td>-0.104</td>
<td>-1.170***</td>
<td>-1.347***</td>
<td>-0.447**</td>
</tr>
<tr>
<td></td>
<td>(0.223)</td>
<td>(0.240)</td>
<td>(0.254)</td>
<td>(0.224)</td>
</tr>
<tr>
<td>Observations</td>
<td>955</td>
<td>952</td>
<td>954</td>
<td>955</td>
</tr>
<tr>
<td>Adjusted R²</td>
<td>0.002</td>
<td>0.023</td>
<td>0.032</td>
<td>0.006</td>
</tr>
</tbody>
</table>

Table A4: *Survey Experiment Results on Attentive Sample*. Results of the survey experiment on the sample of respondents who successfully pass an attention check. We check for attention by asking treated individuals in which country gene-editing occurred. Statistical significance is denoted by: *p<0.1; **p<0.05; ***p<0.01
A.2: Figures

Figure A1: Gene-editing Patent Applications, 2012-2018: The figure displays annual patent applications related to CRISPR (red), TALENS (green), and ZFN (blue) technologies. Data from Oribit Intelligence.
Figure A2: *AddGene Registered Researchers by Country of Origin*: The figure shows the number of researchers registered on the AddGene website, by country of origin. Data collected by authors.

Figure A3: *AddGene depositors by country*
Figure A4: Distribution of Responses for Outcome Variables. The figure displays the distribution of responses by treatment condition for each of four outcomes.
A.3: Survey Experiment Consent and Text

In line with the APSA Principles and Guidance for Human Subjects Research, the author who provided the funding for this experiment submitted the survey protocol to the relevant Institutional Review Board (IRB) Human Subjects Committee prior to launching the survey experiment. The IRB reviewed this survey experiment and granted an exemption under federal regulation 45 CFR 46.104 (2)(ii) (IRB Protocol ID 2000027424). The survey does not contain deceptive material, intervene in political processes, or collect sensitive and/or personally identifiable information.

Respondents were recruited through Lucid, an automated marketplace that connects researchers with online research participants. The authors compensated Lucid $1 per completed interview. Lucid contracts with suppliers who provide financial incentives to survey respondents in the form of cash, gift cards, or loyalty reward points. All respondents are voluntary participants based in the United States. For further details, see https://luc.id/wp-content/uploads/2019/10/Lucid-IRB-Methodology.pdf.

Before beginning, potential respondents are informed that the study is voluntary and assured that their responses will be kept confidential. We then ask for their informed consent:

You are invited to participate in a research study that will take approximately 15 minutes to complete. You will be asked to answer some questions about yourself and your preferences.

There are no known or anticipated risks to you for participating.

Participation in this study is completely voluntary. You are free to decline to participate, to end participation at any time for any reason, or to refuse to answer any individual question without penalty or loss of compensation. The researcher will not know your name, and no identifying information will be connected to your survey answers in any way. The survey is therefore anonymous.
If at any time you have questions or concerns about the survey or your rights or welfare as a research subject, contact [Author name] at [Author email].

If you would like to talk with someone other than the researchers to discuss problems or concerns, to discuss situations in the event that a member of the research team is not available, or to discuss your rights as a research participant, you may contact the [Author’s university] Human Subjects Committee, [phone number], [email]. Additional information is available at [Link to statement of research participant’s rights at Author’s university].

If you would like to participate, simply click the ‘I agree to participate’ box below, then click the >> button to start the survey.

After a set of demographic questions, all respondents are provided the following information:

Now you will read some information related to recent advances in biotechnology.

All organisms, from bacteria to lizards to humans, have molecules called DNA, or deoxyribonucleic acid. These DNA molecules contain the genetic code for each organism. DNA provides the instructions that determine an organism’s physical characteristics and control how it develops, functions, and reproduces.

In recent years, scientists have developed new gene-editing technologies that can permanently alter an organism’s DNA. These technologies allow scientists to make targeted changes to DNA molecules in plants and animals, modifying their biological traits. For example, scientists have edited the genes of wheat plants to make them easier to grow.

Respondents are then randomly assigned to one of four conditions:

1. Control - no additional information
2. Domestic Controversy
3. Foreign Controversy (UK)

4. Foreign Controversy (China)

Those assigned to conditions 2-4 additionally read a hypothetical news article regarding a gene-editing controversy. We show the text for the Foreign Controversy (UK) here.

Below you will read a hypothetical news article about the use of gene-editing technology. The article describes events that could take place in the future. After you have read about the situation, we will ask for your opinions.

**Birth of Genetically Altered Babies in the UK Provokes Outcry**

January 25, 2021

[LONDON]— A British research team announced that they have used a new gene-editing technology to alter the DNA of a group of infants. In an unprecedented intervention, scientists on the research team deleted a set of genes believed to be linked to breast and prostate cancer. The deleted genes are not considered essential to basic biological functions in humans, but the long-term effects of their removal are unclear. The research team plans to periodically examine the infants throughout their lives to assess any side effects of the genetic alteration.

The disclosure this week of the research — carried out in the UK — has sparked urgent debate about the ethics of genetic alteration. The infants’ birth represents a significant and controversial leap in the use of gene-editing technology. The British study has also increased concerns about a future in which parents produce “designer babies” with selectively improved traits, such as height or intelligence.

Finally, we ask respondents to rate their agreement with four statements on a scale from zero to ten.
Please indicate your level of agreement with the following statements, with ”0” representing complete disagreement and ”10” representing complete agreement.

- Research involving gene editing should be more strictly regulated in the US
- US patients should have access to medical treatments that involve gene editing
- The US government should provide funding for gene editing research
- Most US scientists conduct their research in a safe and responsible manner
A.5: Moral outrage

A.5.1: Glossary of moral outrage terms from Brady et al. (2021)
<table>
<thead>
<tr>
<th>Word Stems</th>
</tr>
</thead>
<tbody>
<tr>
<td>abhorr</td>
</tr>
<tr>
<td>absolut</td>
</tr>
<tr>
<td>agit</td>
</tr>
<tr>
<td>anger</td>
</tr>
<tr>
<td>angi</td>
</tr>
<tr>
<td>annoy</td>
</tr>
<tr>
<td>antagonist</td>
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<td>appal</td>
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<td>arrog</td>
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<td>arsenol</td>
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<td>ass</td>
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<td>asshol</td>
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<td>attack</td>
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<tr>
<td>avoid</td>
</tr>
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<td>awful</td>
</tr>
<tr>
<td>baffil</td>
</tr>
<tr>
<td>barbar</td>
</tr>
<tr>
<td>bastard</td>
</tr>
<tr>
<td>bird</td>
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</tbody>
</table>
A.5.2: Moral outrage virality

<table>
<thead>
<tr>
<th></th>
<th>Retweets</th>
<th>Likes</th>
<th>Retweets</th>
<th>Likes</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Dependent variable:</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Moral Outrage</td>
<td>0.715***</td>
<td>1.373***</td>
<td>(0.185)</td>
<td>(0.349)</td>
</tr>
<tr>
<td>Sentiment</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td></td>
<td>−0.022</td>
<td>0.877***</td>
<td>(0.205)</td>
<td>(0.402)</td>
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<tr>
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<td>50,839</td>
<td>21,856</td>
<td>21,856</td>
</tr>
<tr>
<td></td>
<td>(5)</td>
<td>(6)</td>
<td>(7)</td>
<td>(8)</td>
</tr>
<tr>
<td>Moral Outrage</td>
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<td>2.332**</td>
<td>(0.444)</td>
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<tr>
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<td></td>
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<tr>
<td></td>
<td>0.556</td>
<td>2.035**</td>
<td>(0.385)</td>
<td>(0.944)</td>
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<tr>
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<tr>
<td></td>
<td>0.352</td>
<td>0.058</td>
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<tr>
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<tr>
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<td></td>
</tr>
<tr>
<td></td>
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<td>−1.816*</td>
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<tr>
<td>Observations</td>
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<td>50,839</td>
<td>21,856</td>
<td>21,856</td>
</tr>
</tbody>
</table>

Table 4: Virality: Correlation between average tweet sentiment and number of expressions of moral outrage with tweet virality (likes and retweets). Sentiment refers to expressions of positive sentiment and is only measured for tweets with at least one word that expresses positive or negative sentiment; positive values indicate more positive sentiment. Moral outrage refers to words that are categorized as expressing moral outrage and indexes the number of words per tweet that reflect this sentiment; positive values indicate increased outrage. Robust standard errors in parentheses. Sample is all tweets mentioning "CRISPR" from 9-25-2018 to 11-25-2018. Bottom panel displays pre-post results on virality.