

# CRISPR Patents: An ELSI Review

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# *CRISPR Patent Licenses*

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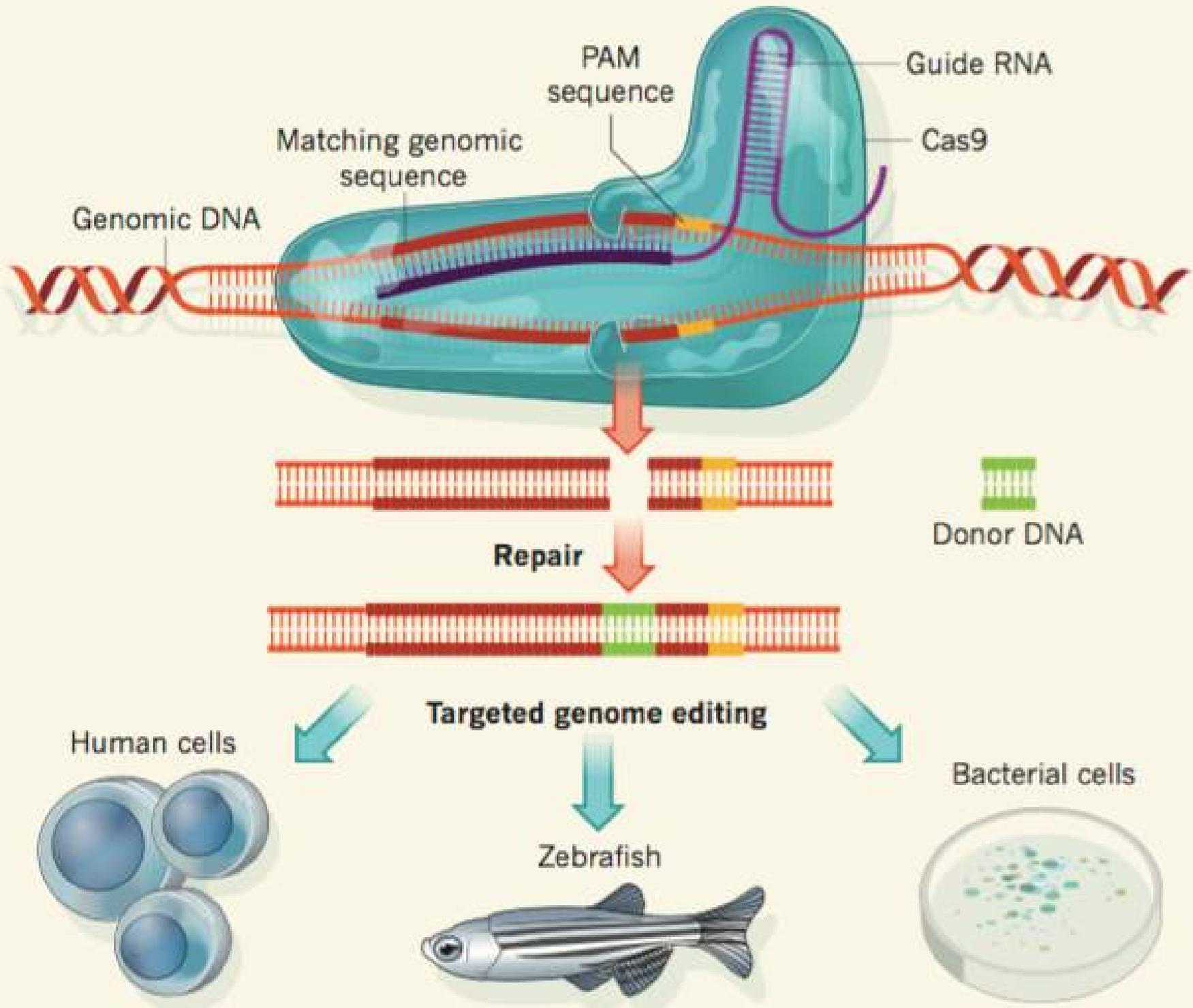


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# Multiplex Genome Engineering Using CRISPR/Cas Systems

Le Cong,<sup>1,2\*</sup> F. Ann Ran,<sup>1,4\*</sup> David Cox,<sup>1,3</sup> Shuailiang Lin,<sup>1,5</sup> Robert Barretto,<sup>6</sup> Naomi Habib,<sup>1</sup> Patrick D. Hsu,<sup>1,4</sup> Xuebing Wu,<sup>7</sup> Wenyan Jiang,<sup>8</sup> Luciano A. Marraffini,<sup>8</sup> Feng Zhang<sup>1†</sup>

Functional elucidation of causal genetic variants and elements requires precise genome editing technologies. The type II prokaryotic CRISPR (clustered regularly interspaced short palindromic repeats)/Cas adaptive immune system has been shown to facilitate RNA-guided site-specific DNA cleavage. We engineered two different type II CRISPR/Cas systems and demonstrate that Cas9 nucleases can be directed by short RNAs to induce precise cleavage at endogenous genomic loci in human and mouse cells. Cas9 can also be converted into a nicking enzyme to facilitate homology-directed repair with minimal mutagenic activity. Lastly, multiple guide sequences can be encoded into a single CRISPR array to enable simultaneous editing of several sites within the mammalian genome, demonstrating easy programmability and wide applicability of the RNA-guided nuclease technology.

**P**recise and efficient genome-targeting technologies are needed to enable systematic reverse engineering of causal genetic variations by allowing selective perturbation of individual genetic elements. Although genome-editing technologies such as designer zinc fingers (ZFs) (1–4), transcription activator–like effectors (TALEs) (4–10), and homing meganucleases (11) have been

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gun to enable targeted genome modifications, there remains a need for new technologies that are scalable, affordable, and easy to engineer. Here, we report the development of a class of precision genome-engineering tools based on the RNA-guided Cas9 nuclease (12–14) from the type II prokaryotic clustered regularly interspaced short palindromic repeats (CRISPR) adaptive immune system (15–18).

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**UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION**  
Washington, D.C. 20549

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**FORM S-1  
REGISTRATION STATEMENT**  
*Under  
The Securities Act of 1933*

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**EDITAS MEDICINE, INC.**  
(Exact name of registrant as specified in its charter)

**Delaware**  
(State or other jurisdiction of  
incorporation or  
organization)

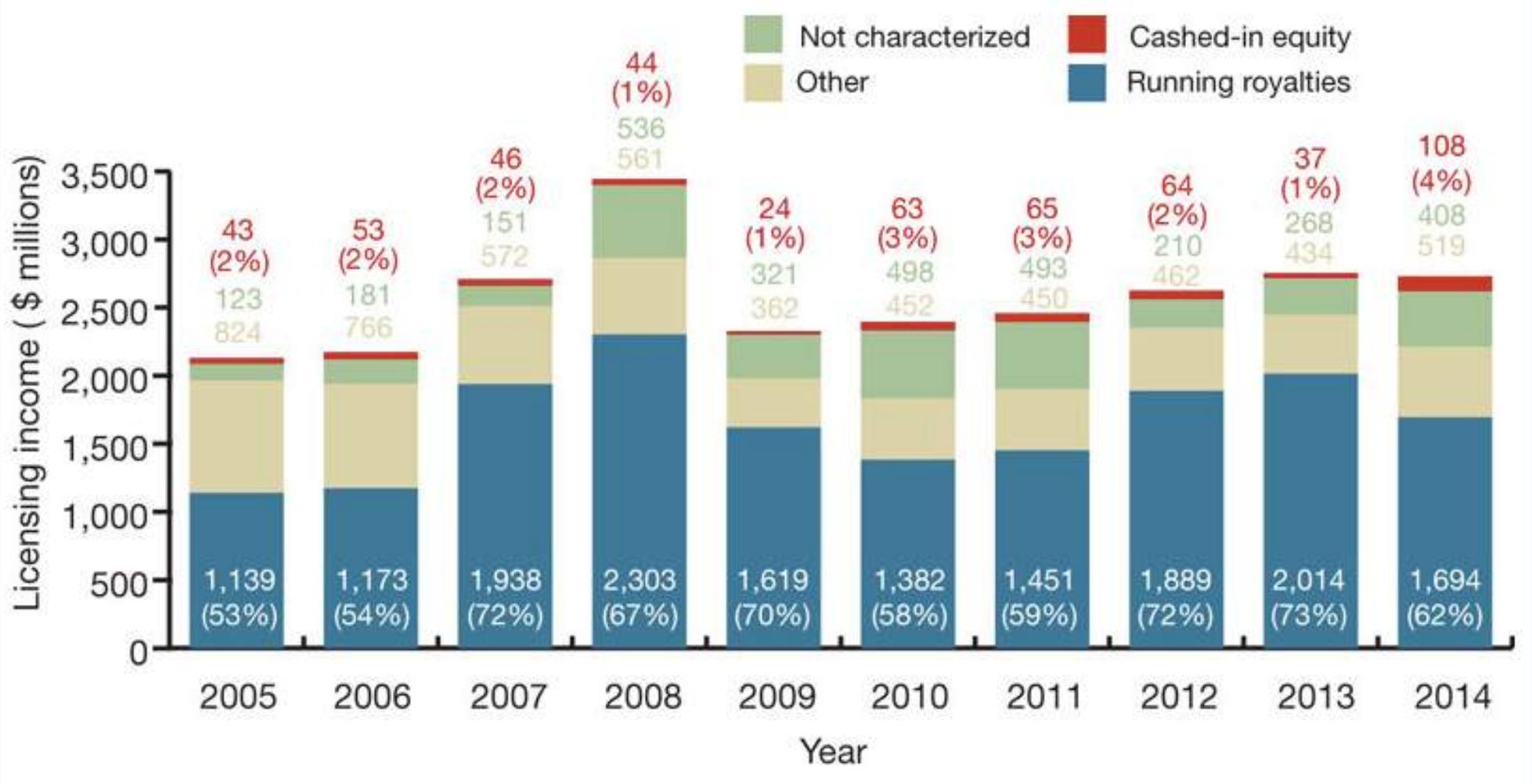
**2836**  
(Primary Standard Industrial  
Classification Code Number)

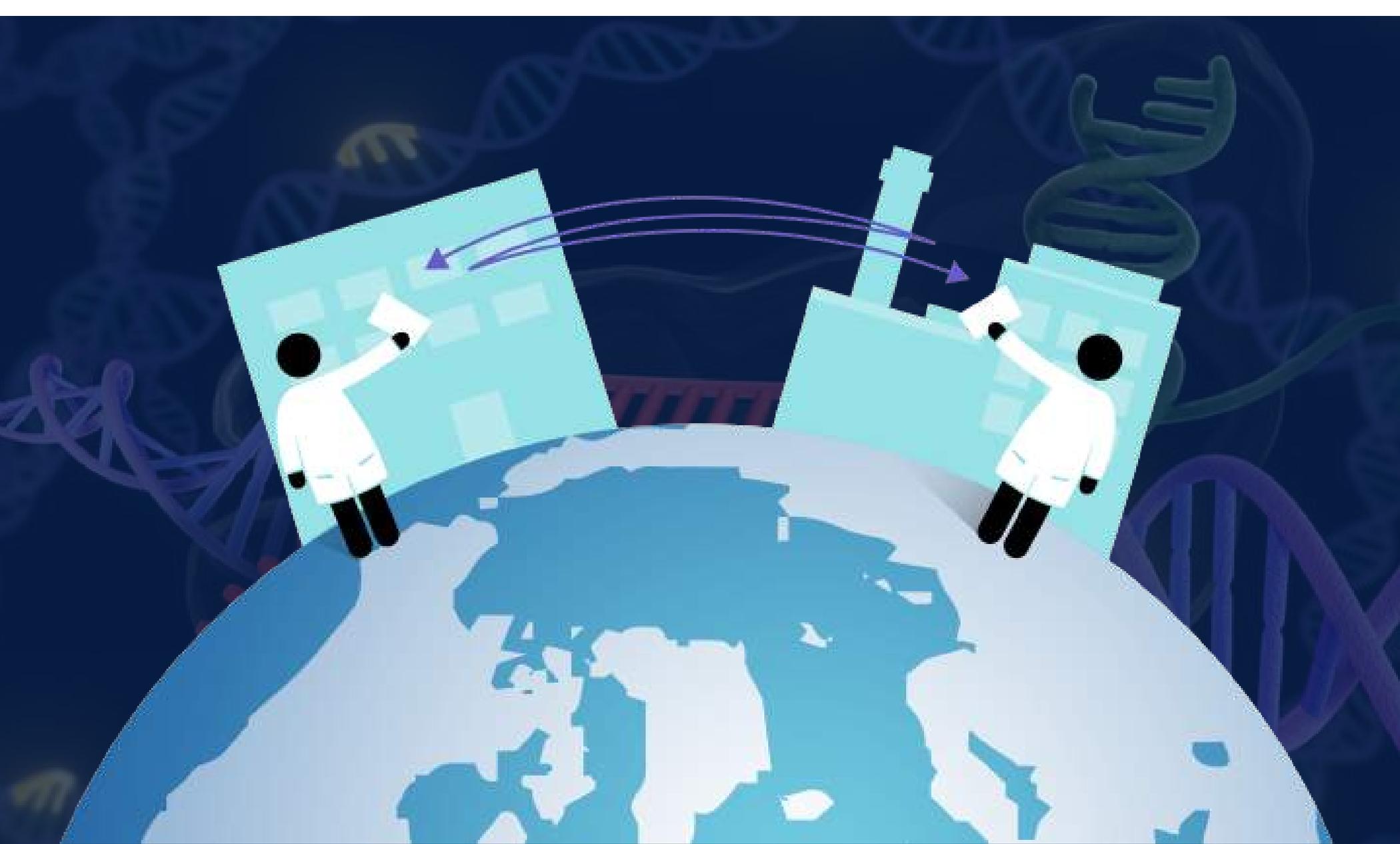
**46-4097528**  
(I.R.S. Employer  
Identification No.)

**Editas Medicine, Inc.  
Notes to Financial Statements (Continued)**

(Information as of September 30, 2015 and for the nine months ended  
September 30, 2014 and 2015 is unaudited)

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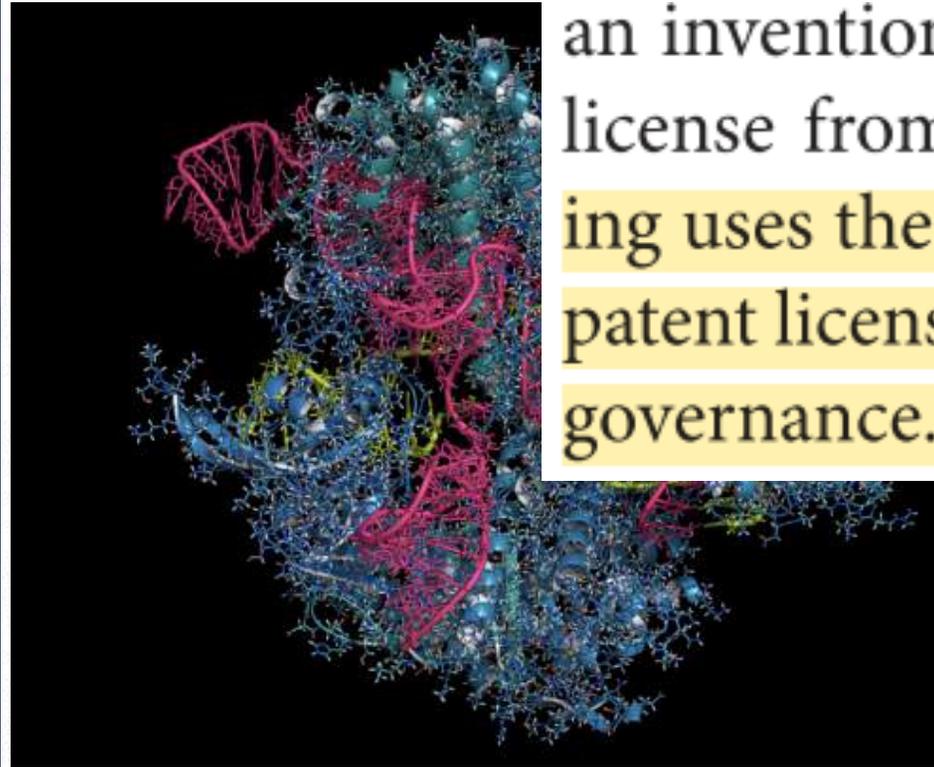
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# The rise of the ethical license

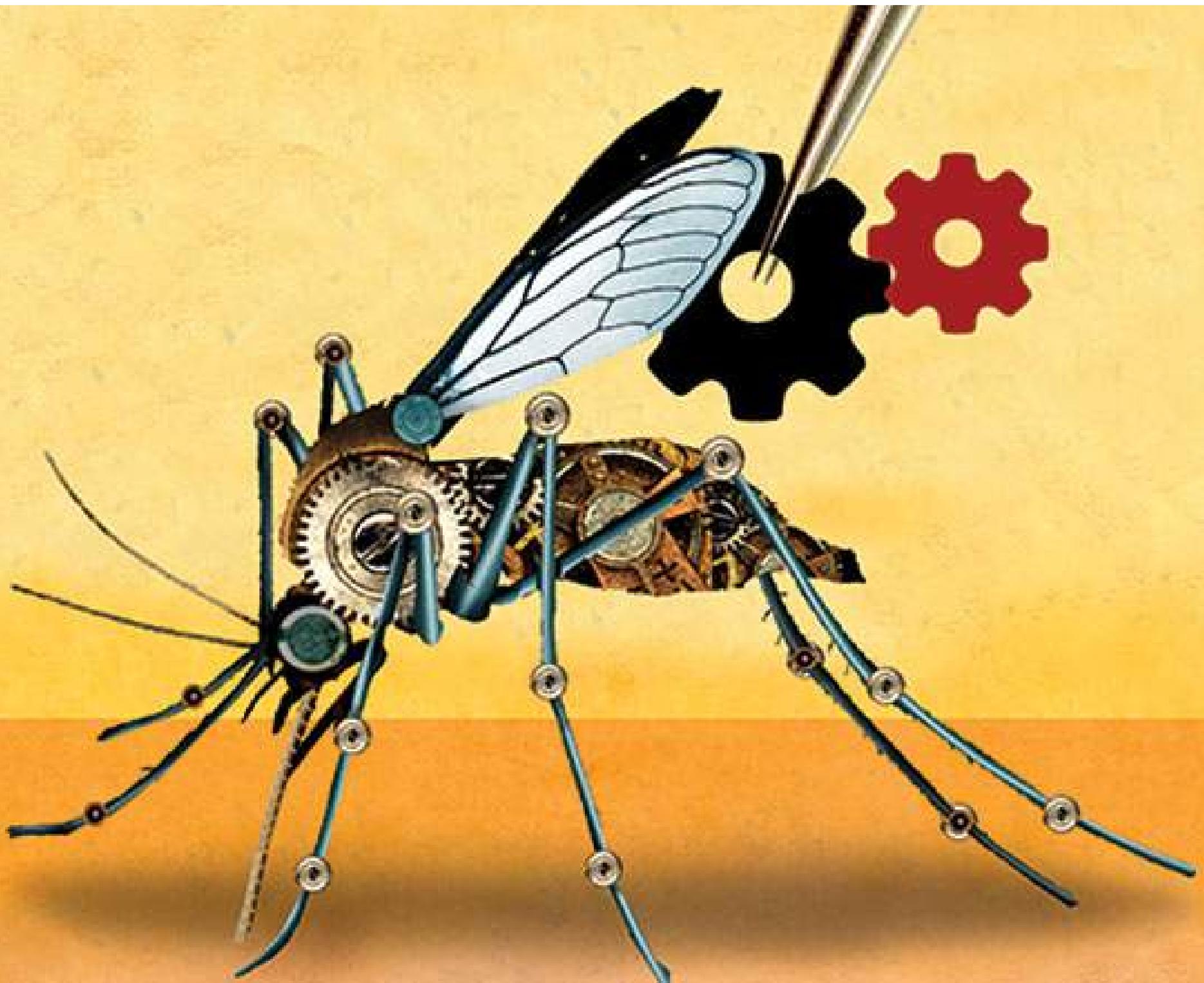
Christi J Guerrini, Margaret A Curnutte, Jacob S Sherkow & Christopher T Scott

The Broad Institute's recent licensing of its gene editing patent portfolio demonstrates how licenses can be used to restrict controversial applications of emerging technologies while society deliberates their implications.



an invention claimed in the patent without a license from the patent holder. By prohibiting uses the patent holder deems unethical, a patent license can function as a tool of private governance. And because the patent right is

ing and standard setting. First, this private solution is more efficient than formal policy making because it does not require consensus among many stakeholders but only the commitment of a single entity: the patent owner. And because the patent owner is frequently the original developer of the technology, it can be in the best position to anticipate controversial applications. Second, unlike most



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