

June 28, 2017

Via email: CRISPR@mpegla.com

Mr. Lawrence A. Horn President and CEO MPEG LA, LLC 4600 S. Ulster St. Suite 400 Denver, CO 80237 USA

Dear Mr. Horn:

The Broad Institute shares the goal of developing innovative technologies such as CRISPR genome editing tools and promoting its translation into genomic medicines to benefit patients.

We are committed to making these technologies widely available for research to help ensure that therapeutic development — bringing this technology to the clinic — has the best chance of success and are working with our joint owners to license our CRISPR technology consistently.

For four years, we have made CRISPR tools, knowledge, methods and other IP for genome-editing freely available to the academic and non-profit community, and will continue to do so. In addition, we license CRISPR IP non-exclusively to companies to use in their own commercial research, and non-exclusively to companies wishing to sell tools and reagents for genome editing. For human therapeutics, we have developed an "inclusive innovation" model, which limits exclusivity to ensure CRISPR technology will be appropriately available for human therapeutics.

We were pleased to learn that MPEG LA was considering the creation of a CRISPR-Cas9 joint licensing pool and are interested in exploring participation, if consistent with our practice of simplifying and expanding, rather than restricting, access to these important tools and methods.

The Broad Institute hereby submits for evaluation of eligibility to participate in discussions facilitated by MPEG LA regarding creation of a CRISPR Joint Licensing Platform the ten (10) patent families described on the attached list, which include, *inter alia*, the following CRISPR-Cas9 patents:

US Patent No. 8,697,359	US Patent No. 8,906,616	EP Patent No. EP2825654
US Patent No. 8,771,945	US Patent No. 8,932,814	EP Patent No. EP2840140
US Patent No. 8,795,965	US Patent No. 8,945,839	EP Patent No. EP2896697
US Patent No. 8,865,406	US Patent No. 8,993,233	EP Patent No. EP2898075
US Patent No. 8,871,445	US Patent No. 8,999,641	EP Patent No. EP2921557
US Patent No. 8,889,356	EP Patent No. EP2764103	EP Patent No. EP2931898
US Patent No. 8,889,418	EP Patent No. EP2771468	
US Patent No. 8,895,308	EP Patent No. EP2784162	

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The Broad Institute submits this application also on behalf of joint-owners of Broad filed CRISPR patents and patent applications including: Massachusetts Institute of Technology, President and Fellows of Harvard College, and The Rockefeller University each a joint-owner of one or more of the above referenced properties.

While there is no obligation or commitment from such submission, and only one patent asset needs to be submitted and determined eligible to qualify to participate in discussions, to underscore our commitment to making CRISPR technologies broadly available for research, we have included not only issued CRISPR-Cas9 patents, but more broadly relevant CRISPR patents and application filed by the Broad Institute that relate to this important technology.

Of course, these patent pools are only truly effective in streamlining the process and driving research forward if many parties participate. We know that our approach differs sharply from and is much more open than what we understand to be the case with licenses given by others.

We look forward to discussions with additional like-minded potential partners about opportunities to streamline licensing of CRISPR IP to ensure these technologies are freely available to the academic and non-profit community and reasonably and non-exclusively available to companies to use in their own commercial research or to sell tools and reagents, and, for human therapeutics, in a manner to encourage sufficient investment to develop fully CRISPR-gene editing technology to treat human diseases and to maximize patient benefit.

Please let us know when and where the meeting will take place, as representatives of Broad Institute and also of our current licensing partners are interested in joining this discussion.

Regards,

Ellen Law

Senior Advisor to the Director

The Broad Institute of MIT and Harvard

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Cambridge, MA 02142

CC: Dahlia Fetouh, Counsel, MIT Office of the General Counsel Lesley Millar-Nicholson, Director, MIT Technology Licensing Office Lauren Foster, Associate Director, MIT Technology Licensing Office Diane Lopez, Deputy General Counsel, Harvard Office of the General Counsel Vivian Berlin, Director of Business Development, Life Sciences, Harvard Office of Technology Development Karen Sinclair, Director of Intellectual Property, Harvard Office of Technology Development Deborah Yeoh, Vice President and General Counsel, The Rockefeller University Office of the General Counsel Kathleen Denis, Associate Vice President, The Rockefeller University Office of Technology Transfer

Broad CRISPR-Cas9 Patent Portfolio, MPEG LA discussion

OwnerName	ApplicationStatus	AppNumber	PubNumber	AppTitle	Applaventors
Broad, MIT, Harvard & Rockefeller	Granted and/or Pending in US, Europe, UAE, Australia, Brazil, Canada, China, Japan, Republic of Korea, Qatar, Russian Federation and Hong Kong.	PCTAUS2013/074611	WO 2014/093595	CRISPR-CAS COMPONENT SYSTEMS, METHODS AND COMPOSITIONS FOR SEQUENCE MANIPULATION	Zhang, Feng, Bikard, David Olivier, Cong. Le, Cox, David Benjamin Turitz, Itsu, Patrick; Jiang, Wenyan; Lin, Shuailiang; Marraffini, Luciano, Platt, Randall Jeffiey; Ran, Fei; Sanjana, Neville Espi
Broad, MIT & Harvard	Granted and/or Pending in US, Europe, UAE, Australia, Brazil, Canada, China, Egypt, India, Israel, Japan, Republic of Korea, Mexico, Qatar, Russian Federation, Sinapore, South Africa and Hong Kong.	PCT/US2013/074667	WO 2014/093622	DELIVERY, ENGINEERING, AND OPTIMIZATION OF SYSTEMS, METHODS AND COMPOSITIONS FOR SEQUENCE MANIPULATION AND THERAPEUTIC APPLICATIONS	Zhang, Feng, Heidenreich, Matthias, Ran, Fer, Swiech, Lukasz
Broad, MIT & Harvard	Granted and/or Pending in US, Europe, Australia, Canada, China, India, Israel, Japan, Republic of Korea, Sinapore and Hong Kong.	PCT/US2013/074691	WO 2014/093635	ENGINEERING AND OPTIMIZATION OF IMPROVED SYSTEMS, METHODS AND ENZYME COMPOSITIONS FOR SEQUENCE MANIPULATION	Zhang, Feng; Ran, Fei; Shalem, Ophir
Broad, MIT & Harvard	Granted and/or Pending in US and Europe.	PCT/US2013/074736	WO 2014/093655	ENGINEERING AND OPTIMIZATION OF SYSTEMS, METHODS AND COMPOSITIONS FOR SEQUENCE MANIPULATION WITH FUNCTIONAL DOMAINS	Zhang, Feng, Cong, Le; Platt, Randall Jeffrey; Ran, Fei; Sanjana, Neville Espi
Broad & MIT	Granted and/or Pending in US, Europe, UAE, Australia, Brazil, Canada, China, Japan, Republic of Korea, Qatar, Russian Federation and Hong Kong.	PCT/US2013/074743	WO2014/093661	CRISPR-CAS SYSTEMS AND METHODS FOR ALTERING EXPRESSION OF GENE PRODUCTS	Zhang, Feng
Broad, MIT & Harvard	Granted and/or Pending in US.	PCT/US2013/074790	WO 2014/093694	CRISPR-CAS NICKASE SYSTEMS, METIIODS AND COMPOSITIONS FOR SEQUENCE MANIPULATION IN EUKARYOTES	Zhang, Feng, Cong, Le; Ran, Fei
Broad, MIT & Harvard	Granted and/or Pending in US, Europe, Australia, Canada, China, India, Israel, Japan, Republic of Korea, Mexico, Russian Federation and Hong Kong.	PCT/US2013/074819	WO 2014/093712	ENGINEERING OF SYSTEMS, METHODS AND OPTIMIZED GUIDE COMPOSITIONS FOR SEQUENCE MANIPULATION	Zhang, Feng, Cong, Le; Hsu, Patrick; Ran, Fei
Broad & MIT	Pending in US and Europe.	PCT/US2014/041806	WO2014/204727	FUNCTIONAL GENOMICS USING CRISPR-CAS SYSTEMS, COMPOSITIONS, METHODS, SCREENS AND APPLICATIONS THEREOF	Zhang, Feng; Shalem, Ophir, Sanjana, Neville Espi; Doench, John; Root, David E., Biewener Hartenian, Ella Nicole
Broad, MIT & Rockefeller	Pending in US, Europe, UAE, Australia, PCT/US2014/070135 Brazil, Canada, China Egypt, India, Israel, Japan, Republic of Korea, Mexico, Qatar, Russian Federation, Singapore, South Africa,	PCT/US2014/070135	WO2015/089465	DELIVERY, USE AND THERAPEUTIC APPLICATIONS OF THE CRISPR-CAS SYSTEMS AND COMPOSITIONS FOR HBY AND VIRAL DISEASES AND DISORDERS	Bhatia, Sangecta; Rice, Charles; Zhang, Feng; Cox, David Benjamin Turitz; Ramanan, Vyas; Schwartz, Robert; Shlomai, Amir
Broad & MIT	Pending as PCT filing (unexpired) and in US, Europe, Argentina, Taiwan and Hong Kong.	PCT/US2016/038034	WO2016/205613	CRISPR ENZYME MUTATIONS REDUCING OFF- TARGET EFFECTS	Zhang, Feng, Gao, Linyi, Zetsche, Bernd, Slaymaker, Ian