



DIVISION OF
CORPORATION FINANCE

UNITED STATES
SECURITIES AND EXCHANGE COMMISSION
WASHINGTON, D.C. 20549

November 10, 2020

David Kirn
Chief Executive Officer
4D Molecular Therapeutics Inc.
5858 Horton Street #455
Emeryville, CA 94608

Re: 4D Molecular Therapeutics Inc.
Draft Registration Statement on Form S-1
Submitted October 14, 2020
CIK No. 0001650648

Dear Dr. Kirn:

We have reviewed your draft registration statement and have the following comments. In some of our comments, we may ask you to provide us with information so we may better understand your disclosure.

Please respond to this letter by providing the requested information and either submitting an amended draft registration statement or publicly filing your registration statement on EDGAR. If you do not believe our comments apply to your facts and circumstances or do not believe an amendment is appropriate, please tell us why in your response.

After reviewing the information you provide in response to these comments and your amended draft registration statement or filed registration statement, we may have additional comments.

Draft Registration Statement on Form S-1

Prospectus Summary

Overview, page 1

1. Please balance the disclosure in the summary with disclosure that your product candidates are based on a novel AAV gene therapy technology with which there is limited regulatory and clinical experience and that few gene therapy products have been approved by the FDA or comparable foreign regulatory agencies. Therefore, it is difficult to predict how long it will take to develop your product candidates and obtaining regulatory approval. Also disclose in the summary that the regulatory approval process for novel product candidates can be more expensive and take longer than for other, better known or extensively studied therapeutic modalities. Finally, revise your disclosure on page 5 and

page 117 that you will "apply [y]our modular product design and engineering approach to accelerate the pace of product development" to remove any implication that you will be able to accelerate the development of your product candidates as such statements are speculative.

2. Please delete the statements that an evolved vector increases the likelihood of success and could accelerate the pace of product development and that your competitive advantages and experience uniquely position you to successfully create, develop and manufacture targeted gene therapies. Given the number of product candidates that never receive FDA approval, the time required to obtain approval and the number of companies currently developing product candidates, your statements that you are in a unique position, have an increased likelihood of success and can develop products more quickly is not appropriate.

Our Therapeutic Vector Evolution Platform, page 2

3. Please substantiate your statement that you have developed an "industry-leading" collection of synthetic capsid sequences.
4. Please revise your disclosure here and in the Business section to provide appropriate context for various conclusions and predictions as to the performance of your product candidates and revise and/or remove any statements that imply safety or efficacy as safety and efficacy are determinations that are solely within the authority of the FDA or similar foreign regulators. For example, we note statements that your vectors "achieved enhanced delivery, increased transgene expression, reduced immunogenicity and/or improved antibody resistance when compared to conventional AAV vectors," that you expect to demonstrate improved safety and efficacy of your product candidates versus conventional AAV vectors and various other claims regarding the superiority of your product candidates to those using conventional AAV vectors, including certain statements in the sections regarding competition and differentiation of your various product candidates. Please revise your disclosure to remove any suggestion that there is an expectation that your product candidates will be safe and effective or will have improved safety and efficacy over conventional AAV vectors and instead refer to the relevant objective data from your preclinical trials or studies that relate to your product candidate's performance.
5. We note your disclosure here and in the Business section regarding preclinical head-to-head comparisons between your targeted and evolved vectors with relevant conventional AAV vectors. If you have not conducted actual head-to-head trials, please revise your disclosure to clearly state this fact and disclose why you believe these comparisons are appropriate. If you provide disclosure regarding results from other trials, expand your disclosure to provide the other information regarding these trials that would help an investor make a meaningful comparison (e.g, number of subjects, dosage, how the baseline was measured in each study, etc.).

Our Product Candidate Pipeline, page 3

6. We note your disclosure on page 166 that planning is underway for an IND-enabling GLP toxicology and biodistribution study of 4D-710 in NHP. Please revise the pipeline table to shorten the line for 4D-710 and revise the anticipated milestone accordingly. We also note that you have included in your pipeline table 4D-710, 4D-135, 4D-3XX and 4D-7XX, all of which appear to be in the discovery phase. Given the early-stage development of these programs, please explain why each program is sufficiently material to your business to warrant inclusion in your pipeline table.
7. Please explain what is involved in "lead-optimization" and why you believe this is a separate and distinct development phase, as opposed to part of vector discovery and/or IND-Enabling studies.

Implications of Being an Emerging Growth Company, page 7

8. Please supplementally provide us with copies of all written communications, as defined in Rule 405 under the Securities Act, that you, or anyone authorized to do so on your behalf, present to potential investors in reliance on Section 5(d) of the Securities Act, whether or not they retain copies of the communications.

Risk Factors

Our success depends on our ability to protect our intellectual property and our proprietary technologies, page 51

9. Please revise this risk factor to disclose the patent rights or technologies subject to march-in rights.

Use of Proceeds, page 86

10. Please revise to clarify whether you expect that you will be able to complete the IND-enabling studies for 4D-150 and 4D-710 and the Phase 1/2 clinical trial for 4D-125 using the proceeds from this offering.

Business

Our Proprietary Therapeutic Vector Evolution Platform, page 140

11. Please substantiate your statement that you have the "largest and most diverse portfolio of targeted and evolved vectors in the field of gene therapy."

Competition and Differentiation: AAV Gene Therapy for wet AMD and Diabetic Retinopathy, page 150

12. We note your disclosure that, to your knowledge, 4D-150 would be the only AAV gene therapy asset in wet AMD and DR that has shown superior transduction on human retinal cells *ex vivo* versus conventional AAV vectors such as AAV2 and is the first gene therapy product candidate for the eye to directly inhibit three different angiogenic growth factor targets, including VEGF and PlGF. Please revise these statements to eliminate any

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implication that 4D-150 is effective and to provide the data that support these claims.

Strategic Collaborations, page 172

13. Please provide the current expiration date for the last-to-expire licensed patent right under the Roche Agreement, the uniQure agreements and the UC Agreements.

You may contact Jenn Do at 202-551-3743 or Jeanne Baker at 202-551-3691 if you have questions regarding comments on the financial statements and related matters. Please contact Ada Sarmiento at 202-551-3798 or Suzanne Hayes at 202-551-3675 with any other questions.

Sincerely,

Division of Corporation Finance
Office of Life Sciences