

October 8, 2024

United States Securities and Exchange Commission  
Division of Corporation Finance  
100 F Street, N.E.  
Washington, DC 20549

Attention: Chris Edwards, Jimmy McNamara, Kevin Kuhar and Franklin Wyman

**Re: Upstream Bio, Inc. (the "Registrant")**  
**Registration Statement on Form S-1 (File No. 333-282197)**  
**Request for Acceleration**

Ladies and Gentlemen:

Pursuant to Rule 460 of the General Rules and Regulations under the Securities Act of 1933, as amended, we wish to advise that between October 7, 2024 and the date hereof, approximately 668 copies of the Preliminary Prospectus, dated October 7, 2024, were distributed to prospective underwriters, institutional investors and prospective dealers in connection with the above-captioned Registration Statement, as amended.

We wish to advise you that the participating underwriters have informed us that they have complied and will continue to comply with the requirements of Rule 15c2-8 under the Securities Exchange Act of 1934, as amended.

We hereby join in the request of the Registrant that the effectiveness of the above-captioned Registration Statement, as amended, be accelerated to 4:01 p.m. Eastern Time, on Thursday, October 10, 2024 or as soon thereafter as practicable.

*[signature page follows]*

Very truly yours,

**J.P. MORGAN SECURITIES LLC**  
**TD SECURITIES (USA) LLC**  
**PIPER SANDLER & CO.**  
**WILLIAM BLAIR & COMPANY, L.L.C.**  
As representatives of the Underwriters

**J.P. MORGAN SECURITIES LLC**

By: /s/ Benjamin Burdett  
Name: Benjamin Burdett  
Title: Managing Director, Head of Healthcare ECM

**TD SECURITIES (USA) LLC**

By: /s/ Mariel Healy  
Name: Mariel Healy  
Title: Managing Director

**PIPER SANDLER & CO.**

By: /s/ Chad Huber  
Name: Chad Huber  
Title: Managing Director

**WILLIAM BLAIR & COMPANY, L.L.C.**

By: /s/ Kevin Eisele  
Name: Kevin Eisele  
Title: Managing Director

*[Signature Page to Acceleration Request Letter]*