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NUVELO INC Form 424B3 October 13, 2006 PROSPECTUS SUPPLEMENT NO. THREE

This filing is made pursuant to Rule 424(b)(3)

(TO PROSPECTUS DATED October 13, 2005)

under the Securities Act of 1933 in connection with

Registration No. 333-128316

NUVELO, INC.

Common Stock

This Prospectus Supplement No. Three supplements and amends the prospectus dated October 13, 2005 relating to the resale by certain securityholders of up to 8,425,000 shares of our common stock.

This prospectus supplement should be read in conjunction with the prospectus dated October 13, 2005, as supplemented by Prospectus Supplement No. One dated November 18, 2005 and by Prospectus Supplement No. Two dated December 8, 2005 all of which are to be delivered with this prospectus supplement. This prospectus supplement is not complete without, and may not be delivered or utilized except in connection with, the prospectus, including any amendments or supplements to it.

We are filing this prospectus supplement to reflect a draw down by us pursuant to the common stock purchase agreement by and between us and Kingsbridge Capital Limited, dated August 4, 2005.

The table appearing under the caption Selling Stockholder on pages 24-25 of the prospectus is hereby supplemented by adding the following to the end of that table, immediately after the paragraph and bullet point added by Prospectus Supplement No. Two:

On October 2, 2006 we delivered notice to Kingsbridge to effect a draw down of up to \$10,000,000. The first trading day of the eight day pricing period for this draw down was October 3, 2006 and, in connection with this draw down, on October 13, 2006 we issued an aggregate of 568.247 shares of our common stock to Kingsbridge at an aggregate purchase price of \$10,000,000.

Investing in our common stock involves a high degree of risk. Please carefully consider the Risk Factors beginning on page 4 of the prospectus, as well as the section entitled Risk Factors included in our recent quarterly and annual reports filed with the Securities and Exchange Commission.

Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities or determined if this prospectus supplement is truthful or complete. Any representation to the contrary is a criminal offense.

The date of this Prospectus Supplement is October 13, 2006.