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UNITED STATES DEPARTMENT OF COMMERCE
United States Patent and Trademark Office

February 12, 2014

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By Authority of the

**Under Secretary of Commerce for Intellectual Property** and Director of the United States Patent and Trademark Office

M. TARVER

**Certifying Officer** 

Mylan Exhibit 1003, Page 1



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## 233752

PATENT APPLICATION SERIAL NO.

U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE FEE RECORD SHEET

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OBLON, FISHER, SPIVAK, McCLE LAND AND MAIER, P.C.

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### QUINOLINE TYPE MEVALONOLACTONES

The present invention relates to novel mevalonolactones having a quinoline ring, processes for their production, pharmaceutical compositions containing them and their pharmaceutical uses particularly as anti-hyperlipidemic, hypolipoproteinemic and anti-atherosclerotic agents, and intermediates useful for their production and processes for the production of such intermediates.

Some fermentation metabolic products such as compactine, CS-514, Mevinolin or semi-synthetic derivatives or fully synthetic derivatives thereof are known to be inhibitors against HMG-CoA reductase which is a rate limiting enzyme for cholesterol biosynthesis. (A. Endo J. Med Chem., 28(4) 401 (1985))

CS-514 and Mevinolin have been clinically proved to be potentially useful anti-hyperlipoproteinemic agents, and they are considered to be effective for curing or preventing diseases of coronary artery sclerosis or atherosclerosis. (IXth Int. Symp. Drugs Affect. Lipid



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