Fully-Automated Ultrasensitive Differential Scanning Calorimeter for High-Throughput Studies of Biopharmaceutical Formulations and Drug Discovery

Lung-Nan Lin, John F. Brandts, J. Michael Brandts, Valerian Plotnikov, Verna Frasca MicroCal, LLC, 22 Industrial Drive East, Northampton, Massachusetts, 01060

Ultrasensitive DSC is used to characterize the structure and stability of proteins down to concentrations of 0.1 mgm/ml and below. With existing instruments, sample throughput can be a limiting factor for certain applications, including: biopharmaceutical formulation stability studies; characterization of mutant protein stability; ligand binding studies. The VP-Capillary DSC has been developed for applications requiring higher throughput. The instrument scans effectively at rates up to 250 °C/hr, about 3 times faster than conventional ultrasensitive DSCs. The instrument is mated to a fully-integrated autosampler which allows for unattended operation. VPViewer software controls all cell cleaning, cell filling, and scanning operations, and Origin® software is used for post-run data analysis. Maximum throughput is up to 50 samples during 24 hours of continuous, unattended operation. The sample and reference cells have working volumes of ~130 microliters which permit rapid equilibration consistent with the fastest scan rates. Instrument response time is software-selectable to maximize performance at all scan rates. An improved adiabatic mode is used for upscanning over the temperature range –10 to +130 °C.