

The challenge of translating nutrition research into public health nutrition, University College, Dublin, 18–20 June 2008

## Effective evaluation of small dense LDL

Wendy S. Jones<sup>a</sup>, Bruce A. Griffin<sup>b</sup> and Max Wong<sup>b</sup> and Ian G. Davies<sup>a</sup>

<sup>a</sup>Liverpool John Moores University, Liverpool L17 6BD, UK and <sup>b</sup>University of Surrey, Guildford, Surrey, UK

Small dense LDL (sdLDL) is a subtype of LDL that expresses greater atherogenicity than large buoyant LDL and is characteristic of the dyslipidaemia seen in metabolic syndrome, obesity and type 2 diabetes<sup>(1)</sup>. With a dramatic increase in these conditions in both adults and children worldwide, a rapid and reliable method of estimating sdLDL is of potential value in the identification and subsequent management of ‘at-risk’ individuals.

Separation of LDL subclasses has been achieved by methods including preparative ultracentrifugation or polyacrylamide gradient gel electrophoresis (PGGE); the former has been developed to allow quantification using iodixanol density-gradient media and pre-staining<sup>(2,3)</sup>. While this method is suitable for high-through-put analysis, the procedure is only semi-quantitative. Fully-quantitative ultracentrifugation is more time-consuming, and not therefore suitable for large-scale screening. A simple and rapid method for sdLDL quantification based on Mg–heparin precipitation has been described by Hirano *et al*<sup>(4)</sup>. The present report describes a comparison of the latter method for sdLDL quantification with the iodixanol gradient ultracentrifugation method<sup>(3)</sup>.

Blood sampled into a tripotassium citrate anticoagulant was obtained from nine adults. Plasma removed by centrifugation was separated into two portions that were used for sdLDL analysis by one of the two methods; the procedures were carried out blind by different operators. Ultracentrifugation of one portion in an iodixanol gradient was followed by fractionation and measurement of the cholesterol in twenty fractions, providing a complete lipoprotein profile for each individual from which the sdLDL could be estimated. The Mg–heparin method was performed as described<sup>(4)</sup>. Briefly, heparin–MgCl<sub>2</sub> was added to plasma to separate VLDL, IDL and large buoyant LDL. sdLDL and HDL remained in the infranant fraction and LDL-C was determined by the direct LDL-C method on an ILAB 650 autoanalyser (Randox Laboratories Ltd, UK).

On the small sample studied these two methods gave a reasonable correlation (Figure), as indicated by the similarity in fractionated cholesterol profiles and significant correlation between the cholesterol content of sdLDL. The Mg–heparin precipitation method may provide a suitable method for estimation of sdLDL in ‘at-risk’ individuals. This method may allow for quantitative high-through-put analysis for use in large-scale dietary interventions in populations who are dyslipidaemic.

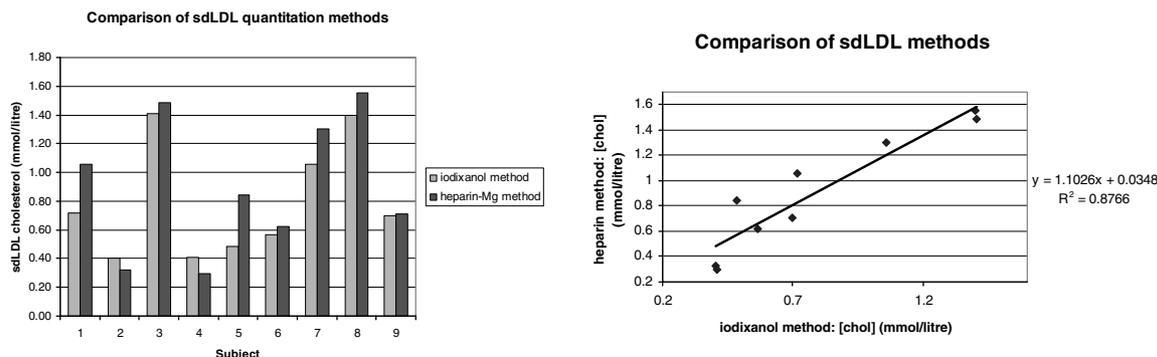


Figure Comparison of sdLDL measurements by alternative methods.

- Berneis KK & Krauss RM (2002) *J Lipid Res* **43**, 1363–1379.
- Davies IG, Graham JM & Griffin BA (2003) *Clin Chem* **49**, 415–418.
- Graham JM, Higgins JA, Gillott T, Taylor T, Wilkinson J, Ford T & Billington D (1996) *Atherosclerosis* **124**, 125–135.
- Hirano T, Ito Y, Saegusa J & Yoshino G (2003) *J Lipid Res* **44**, 2193–2201.