

# High prevalence of primary dyslipidaemia in black South African patients at a tertiary hospital in northern Gauteng, South Africa

A A Khine,<sup>1</sup> MD, MMed, PG Dip (HSE); A D Marais,<sup>2</sup> MB ChB, FCP

<sup>1</sup> Department of Chemical Pathology, National Health Laboratory Service, Pretoria; and Department of Chemical Pathology, Sefako Makgatho Health Sciences University, Pretoria, South Africa

<sup>2</sup> Department of Chemical Pathology, National Health Laboratory Service, Cape Town; Lipidology Research Unit, Faculty of Health Sciences, University of Cape Town; and Lipid Clinic, Grootte Schuur Hospital, Cape Town, South Africa

Corresponding author: A A Khine ([ayeaye.khine@smu.ac.za](mailto:ayeaye.khine@smu.ac.za))

**Background.** The prevalence of cardiovascular disease is projected to be 38.7% for the USA in 2020, including coronary heart disease at 8.6% and stroke at 3.6%. In South Africa (SA), premature deaths due to heart and blood vessel diseases in people of working age (35 - 64 years) have been predicted to increase by 41% between 2007 and 2030, with enormous negative economic impact. Atherosclerosis underlies much of the pathogenesis, which involves risk factors including dyslipidaemia. Secondary dyslipidaemia associated with diabetes mellitus, hypothyroidism, chronic renal disease, cholestasis, nephrotic syndrome, alcohol excess, drugs such as thiazide diuretics and antiretroviral agents may respond to treatment of underlying causes, but residual dyslipidaemia may in such cases be due to primary disorders of metabolism. Primary dyslipidaemias are uncommon and to a large extent underdiagnosed, especially in the black population of SA, reflecting a lack of clinical and laboratory awareness or expertise. Specific diagnoses enable effective intervention in the patients as well as the families.

**Objective.** To assess the burden and prevalence of dyslipidaemia in the SA black population at Dr George Mukhari Hospital in the north region of Gauteng.

**Method.** A retrospective data analysis of 12-month lipid profiles comprising triglyceride (TG), total cholesterol (TC), high-density lipoprotein cholesterol and directly measured low-density lipoprotein (LDL) cholesterol (LDLC).

**Results.** There were 24 656 requests for 6 348 patients. The lipid cut-off levels were somewhat arbitrary but were based on the commonly used decision-making levels in the treatment guidelines. Severe hypercholesterolaemia (>7 mmol/L) was seen in 299 (4.7%) patients and extreme hypercholesterolaemia (>12 mmol/L) was seen in 30 (0.5%) patients. LDLC (>5 mmol/L) occurred in 80 (1.3%) patients and >10 mmol/L in 19 (0.3%) patients. A predominant triglyceride problem was seen in 578 (9.1%) patients with TG (>2 mmol/L) and TC (<5 mmol/L), whereas moderate hypertriglyceridaemia (>5 mmol/L) was present in 113 (1.8%) patients, and more severe hypertriglyceridaemia (>10 mmol/L) in 10 (0.2%). TC (>5 mmol/L) with LDL (>3 mmol/L) but TG in the normal range was seen in 369 (5.8%) patients, indicating a cholesterol-predominant problem. In contrast, LDLC (>3 mmol/L) and TG (>1.7 mmol/L) was seen in 249 (3.87%) representing mixed hyperlipidaemia. Paediatric patients with severe dyslipidaemia mostly suffered from nephrotic syndrome.

**Conclusion.** A significant burden and a high prevalence of dyslipidaemias were present in adults in whom a monogenic disorder should be considered. The extent and severity of dyslipidaemia justify a special clinic and laboratory to ensure accurate diagnosis with effective intervention for patients and their families.

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The American Heart Association predicted the rise in the prevalence of cardiovascular disease as 38.7% by 2020, where the coronary heart disease would rise for 8.6% and stroke for 3.6%.<sup>[1]</sup> In South Africa (SA), Liberty Life Insurance group has forecast the premature deaths due to heart and blood vessel diseases in people of working age (35 - 64 years) to increase by 41% between 2007 and 2030, with a warning of enormous negative economic impact.<sup>[2]</sup> Locally, an increased awareness of cardiovascular disease at the Dr George Mukhari referral hospital has led to more requests for plasma lipid profiles comprising fasting triglyceride (TG), total cholesterol (TC), high-density lipoprotein (HDL) cholesterol (HDL) and low-density lipoprotein (LDL) cholesterol (LDLC). In 2013, 18 234 lipid profiles from 7 255 patients were analysed compared with 24 656 requests for 6 348 patients in 2014. The 35% increase occurred despite opening a new antiretroviral (ARV) clinic in the neighbourhood, which reduced patient flow at the tertiary hospital, and the introduction of electronic gatekeeping. Requests were in response to patients' history of early death among first-degree relatives due to heart disease and stroke,<sup>[3]</sup> and cardiovascular risk suggested by obesity, metabolic

syndrome, smoking, sedentary lifestyle, lack of exercise and stressful living. Genetic disorders in lipoprotein metabolic pathways generally have a greater impact on risk and typically display more severe and/or persistent lipid abnormalities after treating the secondary factors.<sup>[4,5]</sup> Cascade testing of the family is advised.<sup>[6]</sup> Admixture of primary and secondary causes is especially evident in Fredrickson hyperlipidaemia types III, IV and V, where predisposing variant genes are stressed by diet, obesity with insulin resistance, hormonal derangements and alcohol intake. In SA, at least three low-density lipoprotein receptor (LDLR) mutations are known in Afrikaans-speaking whites, one in Jewish people from Eastern Europe and one in Indians of Gujerati descent.<sup>[7,8]</sup> Familial hypercholesterolaemia (FH) is mostly due to a pathogenic mutation in one of three genes (LDLR, apolipoprotein B (apoB) or proprotein convertase subtilisin/kexin type 9). FH is relatively common (prevalence 1:200 - 1:500) and is therefore the most important monogenic disorder of lipoprotein metabolism. It exists as either heterozygous, where only one allele of the gene involved is mutated and the other allele is normal, or homozygous, where both alleles are abnormal. Persons with untreated

homozygous FH are at an approximately 20-fold increased risk for coronary heart disease (CHD), which usually presents in their early 20s. Heterozygous FH is more common (no data for SA yet) and untreated men have a 50% risk of having a coronary event by age 50 years; untreated women are at a 30% risk by age 60 years.<sup>[9]</sup> FH has not been fully investigated in all SA population groups. The commonly held view that disorders of lipoprotein metabolism are absent in the SA black population needs to be dispelled, as it detracts from the recognition of disorders imparting high, but remediable, risk; especially if the previous (rural) lifestyle was protective against atherosclerosis as urbanisation increases cardiovascular diseases. A founder effect may exist in the black population for a six-nucleotide deletion.<sup>[9]</sup> The homozygous FH phenotype has been described in a patient with LDLR mutations and autosomal recessive hypercholesterolaemia.<sup>[10]</sup> Dysbetalipoproteinaemia seems common in black patients at an SA lipid clinic.<sup>[11]</sup> Other than apolipoprotein E (apoE2) allele 2/2 homozygosity accounting for dysbetalipoproteinaemia, De Villiers *et al.*<sup>[12]</sup> found that the apoE Arg145Cys mutation was present in patients of mixed ancestry, whites and more commonly in blacks in Cape Town. The mode of inheritance was autosomal dominant with delayed and incomplete penetrance.<sup>[12]</sup>

Most dyslipidaemias can be diagnosed by the lipid profile, but subtler dyslipidaemias in which triglyceride-rich particles such as very-low-density lipoprotein (VLDL) and intermediate-density lipoprotein accumulate, or the more atherogenic small dense LDL, are found to require more sophisticated investigations. A non-denaturing gradient gel electrophoresis method was shown to be practical, cost-effective and a good diagnostic test for dysbetalipoproteinaemia.<sup>[13,14]</sup>

The Dr George Mukhari Hospital is a tertiary referral hospital drawing patients from northern Gauteng, North West and Limpopo provinces. It is affiliated to Sefako Makgatho Health Sciences University (previously Medunsa). Most patients were indigenous black Africans. To study the prevalence of lipid abnormalities and recognition of monogenic disorders of lipoprotein metabolism in this population, an audit of the lipid requests was performed at this hospital.

## Objectives

To assess the burden of lipid profile abnormalities and possible primary or familial dyslipidaemias at Dr George Mukhari Hospital, northern Gauteng, SA.

## Methods

Analyses of plasma or serum were done in an accredited laboratory with commercially available enzymatic assays for TG and TC; homogenous immunoassay assays for HDLC and LDLC (direct) were performed with Synchron series reagent kits using a DXc automated analyser (Beckman Coulter, USA). The laboratory practised quality control to ensure that data were consistent and comparable with other laboratories. Samples were analysed in the fasted state, except in children.

A cross-sectional audit was done of 12 months of data (1 January - 31 December 2014) extracted from the laboratory information system. Clinical files of patients with severe lipid abnormalities were viewed for clinical details and treatment and data were captured on Microsoft Excel (Microsoft Corporation, USA) without revealing identity. Adults mostly had only one test whereas in paediatric patients lipid profiles were often done 3-monthly. The numbers of patients with various dyslipidaemias were derived from the master database. Hypercholesterolaemia was taken as >5 mmol/L, severe as >7 mmol/L and extreme as >12 mmol/L; the commensurate LDLC levels were 3 mmol/L, 4 mmol/L, 5 mmol/L and 6 mmol/L taking a stepwise approach of classifying the severity. Hypertriglyceridaemia was taken as mild (>2 mmol/L), moderate (>5 mmol/L) and

severe (>10 mmol/L). These are convenient for clinical decision-making limits.

Patient identities were kept confidential. Relevant patients' files were reviewed with the permission from the hospital manager.

## Results

There were 24 656 lipid profiles for 6 348 patients over 12 months. Table 1 summarises the mean, standard deviation (SD), median and range (minimum; maximum) for each lipid parameter extracted. The distribution of each component of the lipid profile is shown in Figs 1 - 4.

The data were analysed separately for adult and paediatric patients. In addition to the information on the request forms and patients' files, locations (outpatient departments, clinics and wards) at the hospital from which the patient was referred also indicated chronic diseases, such as diabetes, HIV treated with ARVs, endocrine conditions such as hypothyroidism, and paediatric high care (nephrotic syndrome) and cardiac high care as well as cardiology outpatients.

Table 2 summarises significantly deranged lipid levels and percentage of patients affected out of the total of 6 348 with clinical diagnoses indicated on the laboratory request forms. TC (<5 mmol/L) with TG (<1.7 mmol/L) (better lipid levels) was present in 3 574 (56.3%). The desired target of LDLC for primary prevention

**Table 1. Distribution of lipid parameters in the study population**

	TC (mmol/L)	LDLC (mmol/L)	TG (mmol/L)	HDLC (mmol/L)
Mean (SD)	4.42 (1.66)	2.32 (1.18)	1.50 (1.17)	1.13 (0.49)
Median	4.28	2.27	1.18	1.14
Minimum	0.38	0.07	0.10	0.13
Maximum	23.00	12.77	13.47	4.50

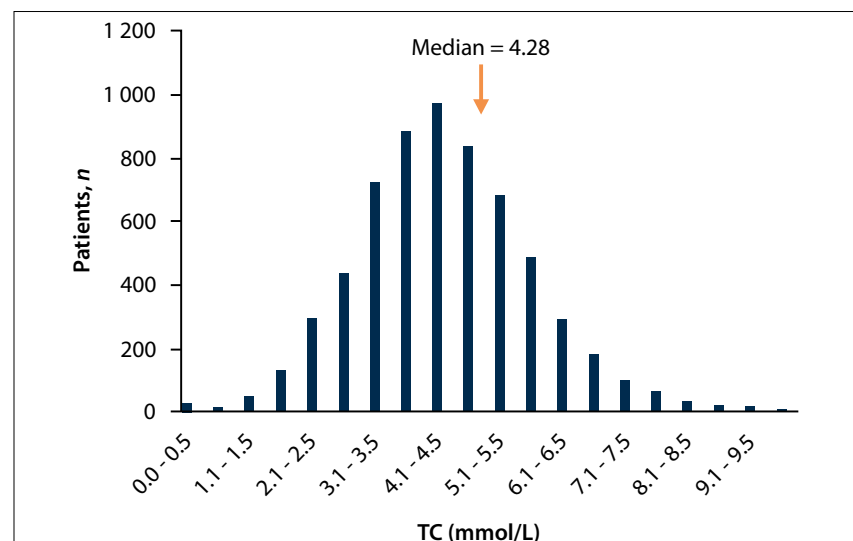


Fig. 1. Total cholesterol distribution (results abridged).

of atherosclerosis (>3 mmol/L) combined with a normal HDL (>2 mmol/L) was seen in 85 (1.2%) patients, indicating isolated LDL problem, while 99 (1.6%) patients had additional higher risk owing to low HDLC (<0.8 mmol/L) over and above their LDL being >3 mmol/L, indicating both LDL and HDL problems. Highlighting the cholesterol-predominant problem, TC (>6 mmol/L) with LDL (>4 mmol/L) with TG in the normal range (<1.7 mmol/L) was seen in 176 (2.8%) patients. To indicate a triglyceride-predominant problem, TC (<5 mmol/L) with TG (>2.0 mmol/L) was seen in 602 (9.5%) patients. In another group, TC (>6.0 mmol/L), LDL (>4 mmol/L) and TG (>2.0 mmol/L) were seen in 322 (5.1%), representing mixed hyperlipidaemia and imposing cardiovascular risk. Hypercholesterolaemia from 5 mmol/L to 7 mmol/L was seen in 1 659 (26.1%) patients. Severe hypercholesterolaemia was seen in 315 patients (5.0%) and extreme hypercholesterolaemia was seen in 33 (0.5%) patients. These cases may indicate underlying monogenic disorders as well as secondary causes. While LDLC (>5 mmol/L) is frequently taken as a cut-point for FH, a stricter cut-off of >6 mmol/L was present in 78 (1.2%) patients and >8 mmol/L in 19 (0.3%) patients. Hypertriglyceridaemia

(>5 mmol/L) was present in 113 (1.8%) patients and >10 mmol/L in 10 (0.2%), signalling a risk for pancreatitis. HDLC (<0.8 mmol/L) was seen in 1 341 (21.1%) patients and hyperalipoproteinaemia (HDLC >2 mmol/L) was seen in 248 (3.9%) patients.

Table 3 provides the information on patients in whom secondary dyslipidaemia was excluded, as well as those on thiazides.

The thiazides probably increase LDLC by 1 - 2% and therefore any TC >7 mmol/L should still be counted as pathological.<sup>[15,16]</sup> Patients treated with simvastatin without initial lipid profiles in the laboratory information system had estimated levels by the calculation based on the LDLC lowering efficacy in African Americans. Adjusted LDLC = current LDLC + 26% (±3%) for simvastatin 20 mg/day dose and + 21% (±1%) for simvastatin 10 mg once

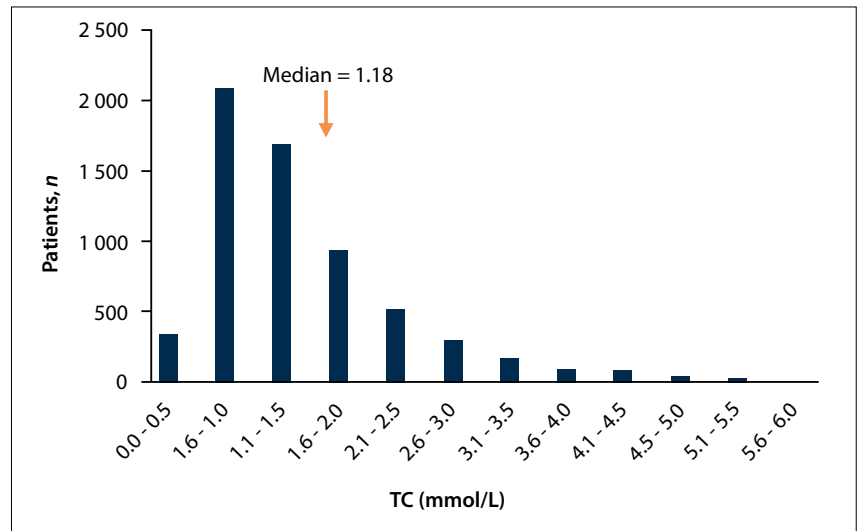


Fig. 2. Triglyceride distribution (results abridged).

Table 2. Significantly deranged lipid levels and percentage of patients affected out of the total of 6 348 with clinical diagnoses (Dx) indicated on the laboratory request forms\*

Lipid level (mmol/L)	Patients, n (%)
TC >5	1 961 (30.9)
TC 5 - 7	1 659 (26.1)
TC >7	315 (5.0)
TC >12	33 (0.5)
TG >5	113 (1.8)
TG >10	10 (0.2)
LDLC >6	78 (1.2)
LDLC >8	19 (0.3)
HDLC <0.8	1 341 (21.1)
HDLC >2	254 (4.0)

Lipid level (mmol/L)	Patients, n (%)	Diabetes (%)	CKD (%)	Liver Ds (%)	Hypertension (%)	IHD/ CABG (%)	NS (%)	ARV (%)	No Dx indicated (%)
TC <5, TG <1.7	3 574 (56.3)	Diagnosis not reviewed because these lipid levels are generally acceptable							
TC >5, LDL >3, TG <1.7	777 (12.2)	30	5	0	30	10	10	10	5
TC >6, LDL >3, TG >2.0	322 (5.1)	45	10	1	30	10	4	0	0
TC <5, TG >2.0	602 (9.5)	20	20	0	5	0	15	5	35
LDLC >3, HDLC <0.8	99 (1.6)	25	12	3	20	20	20	0	0
LDLC >3, HDLC >2	85 (1.3)	These patients overlapped with the group of TC >5 mmol/L, LDL >3 mmol/L and TG <1.7 mmol/L, so their diagnoses are not mentioned separately here							

CKD = chronic kidney disease; Liver Ds = liver disease; IHD = ischaemic heart disease; CABG = coronary artery bypass graft; NS = nephrotic syndrome. \*Information such as chronic alcoholism and drugs are not usually indicated on the request forms.

**Table 3. Demography and clinical profile with the lipid levels\* in significantly affected (n=19) patients**

Patient No.	Age (yr)	Gender	Ethnicity	TC	LDLC	Estimated initial	Non-			Diagnosis	Medication
						LDLC, range	TG	HDLC	HDLC		
1	55	F	SW	8.4	6.5	7.94 - 8.44	1.27	1.66	6.74	IHD	Sim 20 mg
2	57	F	S	8.6	7.0	8.59 - 9.08	1.63	1.36	7.24	CABG	Sim 20 mg
3	61	M	SW	6.3	4.7	6.93 - 7.36	1.22	1.55	4.75	AS	Sim 20 mg + AHT
4	65	F	SW	6.9	4.9	5.99 - 6.36	0.71	2.17	4.73	AMI	Sim 20 mg
5	26	F	V	6.6	4.8	No statins	1.12	1.73	4.87	CCF/PTB	None
6	64	M	S	7.4	5.9	8.7 - 9.24	1.54	1.23	6.17	AMI	Sim 20 mg
7	59	F	S	6.1	4.1	5.01 - 5.32	1.37	1.61	4.49	HLP	Sim 20 mg
8	50	F	P	7.1	5.6	6.84 - 7.27	1.31	1.20	5.90	AS & IHD	Sim 20 mg
9	70	F	P	9.6	7.3	No statins	1.29	1.45	8.15	IHD/HHF	AHT
10	37	M	S	6.2	4.6	5.62 - 5.97	0.87	1.28	4.92	AP	Sim 20 mg
11	67	F	SW	9.1	7.0	No statins	1.67	1.84	7.26	AP/HPT	AHT
12	84	F	SW	7.6	4.4	No statins	1.31	1.19	6.41	CCF/LVD	Amiloride 5 mg
13	50	F	SW	6.6	5.0	6.11 - 6.49	1.21	1.34	5.26	HLP	Sim 20 mg
14	21	M	SW	6.8	5.4	No statins	0.66	1.52	5.28	PHT + smoking	Digoxin
15	68	F	SW	6.5	4.8	5.87 - 6.23	1.28	1.39	5.11	ACS	Sim 20 mg
16	50	F	P	6.47	4.8	5.87 - 6.23	0.68	1.66	4.81	AMI	Sim 20 mg
17	60	M	P	6.16	5.66	No statins	0.96	1.23	4.93	CCF	Amiloride, digoxin
18	51	F	S	6.86	5.12	6.26 - 6.64	0.86	2.59	4.27	AP	Sim 20 mg
19	55	F	S	6.21	4.88	5.96 - 6.33	1.52	0.86	5.35	IHD	Sim 20 mg

M = male; F = female; SW = Tswana; P = Pedi; V = Venda; S = Sotho; TS = Tsonga; Sim = simvastatin daily intake; AHT = antihypertensive therapy with no hydrochlorothiazide; ACS = acute coronary syndrome; AMI = acute myocardial infarction; AP = angina pectoris; HPT = hypertension; VHD = valvular heart disease; AS = aortic stenosis; BC = bradycardia; MS = mitral stenosis; CCF = congestive heart failure; PTB = pulmonary thrombo-embolism or tuberculosis; PHT = pulmonary hypertension; HLP = hyperlipidaemia; HHF = hypertensive heart failure.  
\*All lipid levels are reported in mmol/L. Non-HDL cholesterol >4 mmol/L is a risk factor for atherosclerotic cardiovascular disease.

a day (od) (taken from pravastatin 10 mg od dose from the reference cited, as there was no adjustment for simvastatin 10 mg od).<sup>[17]</sup> The ratio of TC/HDLC or LDLC/HDLC may be used to assess risk when dealing with moderate dyslipidaemia.<sup>[18]</sup> In addition, non-HDLc was also calculated as a guide to atherogenicity.<sup>[19]</sup>

A total of 555 requests came from 148 patients in the paediatric cohort (average 3 - 4 tests per patient in 12 months), almost all from inpatients. Table 4 presents the proportion of paediatric patients with lipid profiles significantly elevated. Eighteen out of 148 (12%) had TC >6 mmol/L and 49 (33.1%) patients had LDLC >3 mmol/L, while 10 (7%) patients had LDLC >4 mmol/L and 31 (21%) patients had LDLC >5 mmol/L. Hypertriglyceridaemia of >1.7 mmol/L was present in 66 patients (44.6%), >2 mmol/L in 59 (40%) and >5 mmol/L in 15 (10.1%).

Clinical files of 46 paediatric patients (31 with LDLC >5 mmol/L and TG >5 mmol/L) were reviewed (Table 4). Of these, 23 (50%) were diabetic with glycated haemoglobin ranging from 7% to 10%, 15 (32%) patients had nephrotic syndrome, of whom 5 had severe hypercholesterolaemia (7.5 - 14.4 mmol/L). Eight (17%) were HIV patients on protease inhibitor treatment.

## Discussion

This study revealed an awareness of dyslipidaemia and cardiovascular risk with comorbidities in adult and paediatric patients. While secondary causes were prominent in children, a large portion of adults had moderate dyslipidaemias associated with other risk factors such as hypertension and diabetes, and drugs such as thiazides could have contributed. Severe hyperlipidaemia suggestive of monogenic disorders of lipoprotein metabolism was likely to be present in a

**Table 4. Numbers and percentages of paediatric patients affected, according to various lipid cut-off levels\***

Patients	Lipid level (mmol/L)	n (%)
With high TC	>6	18 (12.0)
With high LDL	>3	49 (33.1)
	>4	10 (7.0)
	>5	31 (21.0)
With high TG	>1.7	66 (44.6)
	>2	59 (40.0)
	>5	15 (10.1)

\*Diagnoses are described in the last paragraph of the 'Results' section.

small proportion but was not specifically considered in the notes. Patients with abnormal lipid levels were not worked up using lipid electrophoresis on agarose gel or polyacrylamide gel in order to look for specific types of hyperlipidaemias. Moreover, additional tests such as lipoprotein (a), which poses independent risk for cardiovascular complications, were not evaluated. This has a negative impact on patient management as the significant hyperlipidaemias could, to some extent, be affected or underplayed by genetic mutations or polymorphisms. If these were detected by proper work-up, both patient and family could benefit.

In the distribution graphs (Figs 1 - 4), the HDLC displayed a normal distribution, and TC and LDLC had some patients with strikingly low values that detracted from a normal distribution.

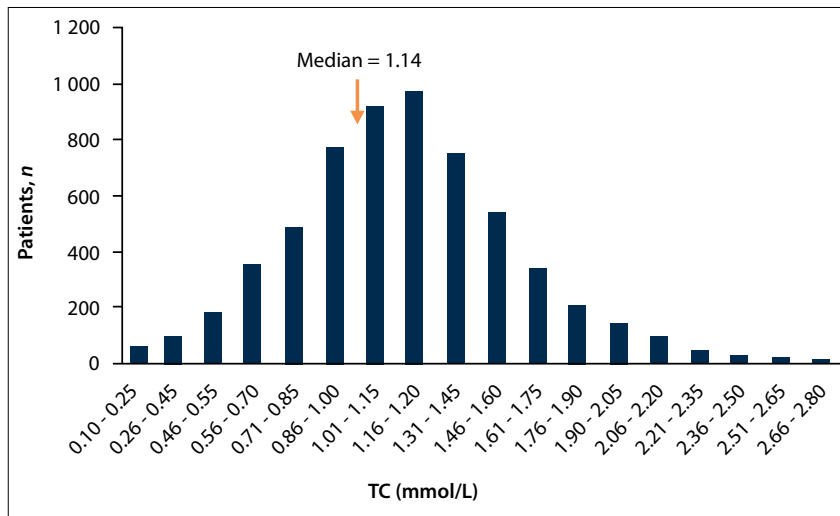


Fig. 3. HDL distribution (results abridged).

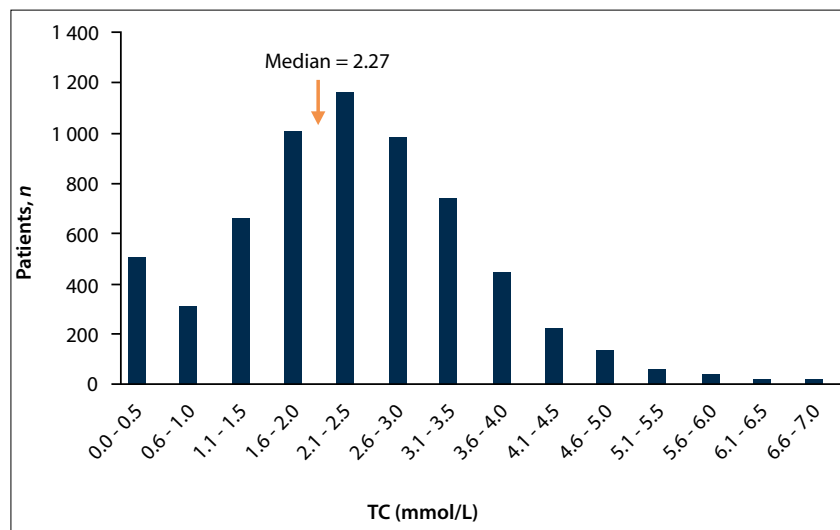


Fig. 4. LDL distribution (results abridged).

Thirty randomly selected patient files with TC <3 mmol/L and LDL <2 mmol/L were assessed and it was found that the bloods were taken for lipid levels during acute illness, such as acute myocardial infarction, which could represent apolipoproteins affected by negative acute phase reaction. Ten patients with lower levels of TC and TG (taken as a sample from TC <3 mmol/L and TG <1 mmol/L) were also found to have suffered from chronic cardiac failure with possible cardiac cirrhosis, which could have contributed to lower secretion of apoB 100. Despite these, TG concentrations, not corrected for background glycerol, also showed the typical skewed distribution towards higher values that could be attributed to patients with uncontrolled diabetes, chronic kidney disease and nephrotic syndrome. Upon closer examination of the combination of various lipid parameters at the relevant medical decision limits

(Table 2), there was a large number of patients with TC >5 mmol/L, the level at which cholesterol-lowering therapy should have been initiated. Patients whose TC >7.0 mmol/L, which could be suggestive of underlying heterozygous FH, are at a much higher risk of coronary complications; this category justifies medication as well as lifestyle changes.<sup>[5]</sup> TC >5 mmol/L with LDL >3 mmol/L and TG <1.7 mmol/L was seen in 777 (12.2%), and this may represent polygenic hypercholesterolaemia, while TC >6 mmol/L, LDL >3 mmol/L and TG >2.0 mmol/L may reflect type IIb hyperlipidaemia as seen in 322 (5.1%) patients. TC <5 mmol/L with TG >2 mmol/L was seen in 602 (9.5%), and could indicate chronic disorders of lifestyle, including obesity, metabolic syndrome or medications. Not surprisingly, the patients with patterns of moderate dyslipidaemia had a relatively high prevalence of monogenic disorders.

After evaluating the patient files, patients with severe dyslipidaemias diagnosed with ischaemic heart disease, hypertension and coronary heart disease were analysed. The adjusted initial LDL levels of patients already on statins indicated possible underlying genetic disorders of LDL clearance. This subset of patients had no contributions to dyslipidaemias from secondary causes, and as mentioned before, patients with hypertension and on thiazide treatment are using a standard dose of 12.5 mg od, and at this level its effect on hyperlipidaemia is modest if not minimal.<sup>[18]</sup> The absence of causes of secondary dyslipidaemia in this severely hypercholesterolaemic group suggests either polygenic disorder, in which multiple polymorphisms of genes are involved in increased production of LDL, or reduced clearance of LDL due to underlying heterozygous FH.<sup>[10]</sup>

This study also revealed 637 (9.98%) patients who presented with cardiac ischaemia and coronary heart disease – not shown in the tables, but obtained from the original data extract by filtering based on the hospital locations where such patients are exclusively cared for. In this population, striking dyslipidaemias without comorbidities were seen and may suggest primary dyslipidaemia, including FH. The ratio of TC/HDL or LDL/HDL may be used to assess risk when dealing with moderate dyslipidaemia.<sup>[18]</sup> A ratio of TC/HDL >4 was seen in 2 475 patients, and a ratio of LDL/HDL >2 was seen in 2 912 patients. Non-HDL >4 mmol/L (indicative of cardiovascular risk) was seen in 1 567 patients.<sup>[19]</sup> A ratio of TC/TG >2.0 was observed in 4 219 patients, and in the study of Blom *et al.*,<sup>[20]</sup> such a ratio was seen in patients with dysbetalipoproteinaemia, although the ratio can be highly variable. The Friedewald calculation permits derivation of LDL when TG is <4.5 mmol/L, except in dysbetalipoproteinaemia. In 6 187 (97.5%) patients whose TG was <4.5 mmol/L (not shown in the tables), the calculated LDL exceeded the directly measured LDL by >1 mmol/L in 2 737 (43.11%) patients; this may suggest dysbetalipoproteinaemia (type III hyperlipidaemia).<sup>[21]</sup> Severe hypertriglyceridaemia compatible with Fredrickson type V was also noted, albeit in small numbers.

## Conclusion

These findings illustrate that various types of primary and secondary dyslipidaemias are seen in adult black patients at this hospital. Regarding paediatric patients, the majority presented with severe dyslipidaemia



owing to underlying nephrotic syndrome, type I diabetes and ARV therapy, although unfavourable underlying genetic variations could not be excluded. The high prevalence of dyslipidaemias may herald cardiovascular disease in a population known to be undergoing lifestyle changes with urbanisation. Unlike the other comorbidities, hypertension does not affect the lipid profile per se but promotes atherosclerosis in a possible positive feedback loop. Unfavourable changes of TC, VLDLC, LDLC/HDLC and HDLC/TC during thiazide treatment, especially at the low dosage of 12.5 mg od may only be a small contributor, but to obtain a more favourable lipid profile, consideration may need to be given to lipid-neutral antihypertensive agents such as angiotensin-converting enzyme inhibitors or calcium channel blockers.<sup>[18]</sup>

Ideally, further investigation should be undertaken to determine the cause(s) of the dyslipidaemias, by characterising the lipoprotein subsets (electrophoresis) and proceeding to appropriate genetic testing, to expedite cascade testing as well as to provide more accurate counselling about heritability. The information and interpretation of data in this study justify the establishment of a lipid clinic dedicated to work-up dyslipidaemias along with a laboratory that can perform more specialised tests.

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