Change of HbA_{1c} reporting to the new SI units

Graham RD Jones, George Barker, Ian Goodall, Hans-Gerhard Schneider, Mark DS Shephard and Stephen M Twigg

aemoglobin $A_{\rm lc}$ (Hb $A_{\rm lc}$ — a term that is sometimes used interchangeably with "glycated haemoglobin") measurements are an indicator of time-averaged blood glucose levels (previous 2–3 months), and are used as the best marker of long-term diabetes control. A recent consensus statement on the worldwide standardisation of Hb $A_{\rm lc}$ measurement has updated previous international recommendations on the standardisation of Hb $A_{\rm lc}$ measurement and reporting. Here, we provide the rationale and guidance for implementation of Hb $A_{\rm lc}$ reporting in the new Système International (SI) units in Australia. This article represents the views of the Australasian Association of Clinical Biochemists, the Australian Diabetes Educators Association, the Australian Diabetes Society and the Royal College of Pathologists of Australasia, and was prepared by a working party of representatives of these organisations.

The International $\mathrm{HbA_{1c}}$ Consensus Committee recommends that all $\mathrm{HbA_{1c}}$ levels be reported in SI units (mmol/mol, no decimal places) — with results directly traceable to the International Federation of Clinical Chemistry and Laboratory Medicine (IFCC) reference method — and in the currently used, National Glycohemoglobin Standardization Program (NGSP) units (percentage, one decimal place). We recommend that dual reporting in Australia begins in July 2011, and that reporting of percentages ceases 2 years later. In New Zealand, dual reporting commenced in August 2009.

The key reasons for implementing this recommendation in Australia are that:

- the SI units relate to a scientifically valid measure of HbA_{1c};
- the SI units remove potential confusion between HbA_{1c} values as a percentage and blood glucose values in mmol/L;
- the change is in keeping with the international consensus statement;¹ and
- the change has already been initiated in New Zealand and a number of countries in the European Union.

Until now, all HbA_{1c} measurements performed in Australia have been reported as percentages (HbA_{1c} as a percentage of total haemoglobin) that are aligned with those produced in the Diabetes

Control and Complications Trial.³ These units and this standardisation have been promoted by the NGSP in the United States, and the activities of this organisation have produced marked improvement in the accuracy of HbA_{1c} results worldwide. More recently, the IFCC has developed a reference method that is more specific for HbA_{1c} and more analytically robust. 4 The IFCC method is now used as the reference system by the NGSP and for all routine methods for measurement of HbA_{1c}, although a numerical conversion is required during the calibration process. The changes recommended here will provide results that are directly aligned with the IFCC method. As the IFCC method is more specific for HbA_{1c}, not measuring several other haemoglobin-sugar complexes, the results are

• An interna

- An international consensus statement recommends that dual reporting of haemoglobin A_{1c} (HbA_{1c}) levels — in the current units (percentage) and Système International (SI) units (mmol/ mol) — be used as an interim measure for a 2-year transition period before progressing towards the use of SI units only.
- This recommendation is supported by the Australasian Association of Clinical Biochemists, the Australian Diabetes Educators Association, the Australian Diabetes Society and the Royal College of Pathologists of Australasia.
- The SI units are a true measure of HbA_{1c} and remove potential confusion between HbA_{1c} values and blood glucose values.

MJA 2011; 195: 45-46

10% to 40% lower than those from the NGSP system, depending on HbA_{1c} concentration. Because reporting these results as percentages may lead to confusion (eg, producing a result of 5.3% rather than 7.0%), the units are changed to mmol/mol (millimoles HbA_{1c} per mole of total haemoglobin [53 mmol/mol in the previous example]), which is consistent with the SI units recommended for use in Australia.

There is a linear relationship between results from the two methods, and the "master equation" is used to convert results between the two methods: HbA_{1c} SI unit (mmol/mol) = $10.93 \times HbA_{1c}$ NGSP unit (%) $-23.50.^5$

To make the conversion easier for clinicians, it is important to translate current treatment advice to the new units. A general conversion table for clinical use is provided in Box 1. The general HbA $_{\rm 1c}$ target of $\leq 7.0\%$ for patients with type 1 and type 2 diabetes mellitus equates to $\leq 53\, \rm mmol/mol$, although these values need to be individualised for patients. The recently updated diabetes treatment guidelines are shown with SI units in Box 2 and Box 3, and recommendations for reporting HbA $_{\rm 1c}$ levels in Australia are summarised in Box 4. In addition, supporting

material for doctors and patients will be presented in SI units in the future.

The routine reporting of an estimated average glucose (eAG) value may be useful for consultations with individual patients. However, the working party strongly agrees with the revised consensus statement that routine reporting of eAG with all requests for HbA_{1c} analysis is not appropriate.1 The reasons for this include variability in the methods used to measure eAG, the risk of confusing a measure of long-term glycaemia (eAG) with a measure of short-term blood glucose control (actual blood glucose level), and its lack of applicability in the majority of patients with type 2 diabetes (in whom blood glucose levels are not measured at frequent intervals).7 Nonetheless, eAG values will be used as an educa-

1	Conversion table	for
	haemoglobin A _{1c}	(HbA_{1c}) values

HbA _{1c} as percentage (old units)	HbA _{1c} in mmol/mol (new units)
5.0	31
6.0	42
6.5	48
7.0	53
8.0	64
9.0	75
10.0	86
11.0	97
12.0	108

likely to limit life expectancy

2 Recommended haemoglobin A_{1c} (Hb A_{1c}) target ranges for patients with type 1 diabetes⁶

	HbA _{1c} target
General target	≤ 53 mmol/mol, ≤ 7.0%*
Specific clinical situations	
Pregnancy or planning pregnancy	\leq 53 mmol/mol, \leq 7.0%*†
Children and adolescents	≤ 58 mmol/mol, ≤ 7.5%*
Recurrent severe hypoglycaemia or hypoglycaemia unawareness	≤ 64 mmol/mol, ≤ 8.0%
Patients with major comorbidities	Symptomatic therapy of

^{*} Achievement of ${\rm HbA}_{1c}$ targets must be balanced against risk of severe hypoglycaemia. † An ${\rm HbA}_{1c}$ level of \leqslant 42 mmol/mol (\leqslant 6.0%) is desirable if it can be achieved safely. ‡ Where practical, suggest blood glucose target level < 15 mmol/L to help minimise risk of infection.

hyperglycaemia[‡] and

avoidance of ketosis

⊔ h ∧ + ~ ~ ~ ~ +

3 Recommended haemoglobin A_{1c} (HbA_{1c}) target ranges for patients with type 2 diabetes⁶

	nbA _{1c} target
General target	≤ 53 mmol/mol, ≤ 7.0%*
Specific clinical situations	
Diabetes of short duration [†] and no clinical cardiovascular disease	
Requiring lifestyle modification ± metformin	≤ 42 mmol/mol, ≤ 6.0%*

Requiring any antidiabetic agents other $\leq 48 \text{ mmol/mol}, \leq 6.5\%^*$ than metformin or insulin $\leq 53 \text{ mmol/mol}, \leq 7.0\%^*$

Pregnancy or planning pregnancy \leq 42 mmol/mol, \leq 6.0%* Children and adolescents \leq 53 mmol/mol, \leq 7.0%*

Diabetes of longer duration \uparrow or clinical cardiovascular disease (any therapy) $\leqslant 53 \, \text{mmol/mol}, \leqslant 7.0\%^*$ Recurrent severe hypoglycaemia or $\leqslant 64 \, \text{mmol/mol}, \leqslant 8.0\%$

hypoglycaemia unawareness (any therapy)

Patients with major comorbidities likely to Symptomatic therapy of limit life expectancy[‡] (any therapy)

hyperglycaemia[§]

4 Recommendations for reporting haemoglobin A_{1c} (HbA_{1c}) levels in Australia

- From July 2011, HbA_{1c} levels should be reported in both National Glycohemoglobin Standardization Program units (percentage) and the Système International (SI) units (mmol/mol) by all pathology laboratories and, where possible, from point-of-care devices
- The period of dual reporting will be 2 years, after which only the SI units will be used.
- These recommendations are consistent with international

tional tool at the discretion of individual clinicians, who can assist patients to understand the significance and limitations of the result.

Competing interests

Graham Jones has received travel expenses from Roche Diagnostics and Bio-Rad, and his institution has received research support (not related to this article) from Roche Diagnostics. Stephen Twigg has received payment for advisory board membership from Sanofi-Aventis, Novartis and Novo Nordisk, and has received speaker fees from GlaxoSmithKline and Eli Lilly. His institution has received a grant from the Novo Nordisk Regional Diabetes Support Scheme. Hans-Gerhard Schneider has received payment, paid to his institution, for providing consultancy to Amgen and for providing expert testimony relating to legal cases, and has also received grants, paid to his institution, from Janssen-Cilag and Biosite. Mark Shephard has received a research grant from Siemens and payment to cover airfare and accommodation to attend a Siemens conference, all paid to his institution.

Author details

Graham R D Jones, MB BS, DPhil, FRCPA, Chemical Pathologist, ¹ and Conjoint Associate Professor²

George Barker, BHSc, CDE-RN, NP, Diabetes Educator³ Ian Goodall, BSc, FAACB, FFRCPA, Scientist in Charge, Complex Chemistry⁴

Hans-Gerhard Schneider, MD, FRACP, FRCPA, Head of Clinical Biochemistry and Director of Pathology⁵

Mark DS Shephard, MAACB, FFRCPA, PhD, Director of Community Point-of-Care Services⁶

Stephen M Twigg, MB BS, PhD, FRACP, Head of Endocrinology Research Laboratories and Deputy Head of Department of Endocrinology⁷

- 1 Department of Chemical Pathology, St Vincent's Hospital, Sydney, NSW.
- 2 Faculty of Medicine, University of New South Wales, Sydney, NSW.
- 3 Diabetes Education Service, Hornsby Ku-ring-gai Health Service, Sydney, NSW.
- 4 Austin Health, Melbourne, VIC.
- 5 Alfred Health, Melbourne, VIC.
- 6 Flinders University Rural Clinical School, Adelaide, SA.
- 7 Royal Prince Alfred Hospital, Sydney, NSW.

Correspondence: gjones@stvincents.com.au

References

- 1 Hanas R, John G; on behalf of the International HbA_{1c} Consensus Committee. 2010 consensus statement on the worldwide standardization of the hemoglobin A_{1c} measurement. *Clin Chem* 2010; 56: 1362-1364.
- 2 Consensus Committee. Consensus statement on the worldwide standardization of the hemoglobin A1C measurement: the American Diabetes Association, European Association for the Study of Diabetes, International Federation of Clinical Chemistry and Laboratory Medicine, and the International Diabetes Federation. *Diabetes Care* 2007; 30: 2399-2400.
- 3 NGSP. Background. http://www.ngsp.org/bground.asp (accessed Mar 2011).
- 4 Weykamp C, John WG, Mosca A, et al. The IFCC Reference Measurement System for HbA_{1c}: a 6-year progress report. *Clin Chem* 2008; 54: 240-248.
- 5 Hoelzel W, Weykamp C, Jeppsson JO, et al; on behalf of the IFCC Working Group on HbA_{1c} Standardization. IFCC reference system for measurement of hemoglobin A_{1c} in human blood and the national standardization schemes in the United States, Japan, and Sweden: a method-comparison study. *Clin Chem* 2004; 50: 166-174.
- 6 Cheung NW, Conn JJ, d'Emden MC, et al. Position statement of the Australian Diabetes Society: individualisation of glycated haemoglobin targets for adults with diabetes mellitus. Med J Aust 2009; 191: 339-344.
- 7 Goodall I, Shephard M, Tate J. Recommended changes in HbA_{1c} reporting units for Australian laboratories. Position statement of the Australasian Association of Clinical Biochemists. http://www.aacb.asn.au/admin/?getfile=2802 (accessed Jun 2011).

Provenance: Not commissioned; externally peer reviewed.

(Received 21 Feb 2011, accepted 30 May 2011)

See page 7.

^{*} Achievement of HbA_{1c} targets must be balanced against risk of severe hypoglycaemia, especially among older people. † In an older adult, long duration might be considered to be > 10–20 years, but for a person who develops type 2 diabetes at a young age, it might be considerably longer. ‡ Examples of major comorbidities include chronic medical conditions, such as chronic kidney disease stages 4 or 5; heart failure stages III or IV (New York Heart Association grading); incurable malignancy; and moderate to severe dementia. § Where practical, suggest blood glucose target level < 15 mmol/L to help minimise risk of infection.