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DHB

Thursday, August 11, 2016

(Room 3)

8:30 - 10:30 WS #5: Starting Insulin in Primary Care

11:00 - 13:00 WS #11: Starting Insulin in Primary Care (Repeated)

Barriers to Insulin

Average HbA1c of $\geq 75\text{mmol/mol}$ (9%) for up to 2 years before starting...

Patient:

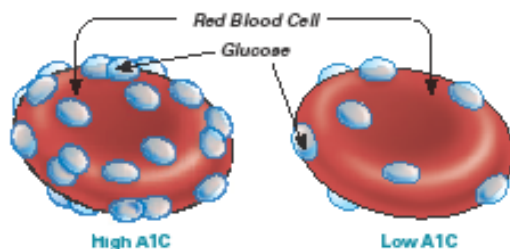
- Fear/ inconvenience of injections
- Social/ work issues
- Hypoglycaemia
- Fear of 'inevitable end stage of disease'
- Perception of failure
- Weight gain

Doctor:

- Benefit to patient unclear
- Doing harm (hypos/ weight gain)
- Time and resources
- Expertise/ confidence

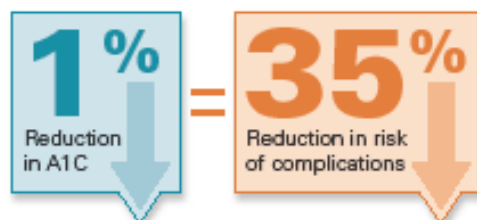
How to prevent complications?

What is A1C?



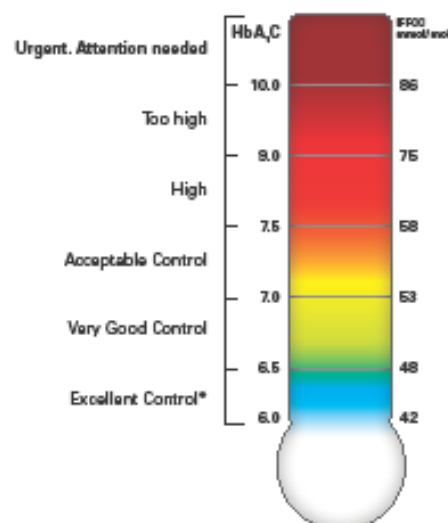
Work on your A1C

American Diabetes Association.
Implications of the United
Kingdom prospective diabetes
study. Diabetes Care 2003;
26(Suppl. 1): S28-S32



Diabetic Control

HbA_{1c} Thermometer

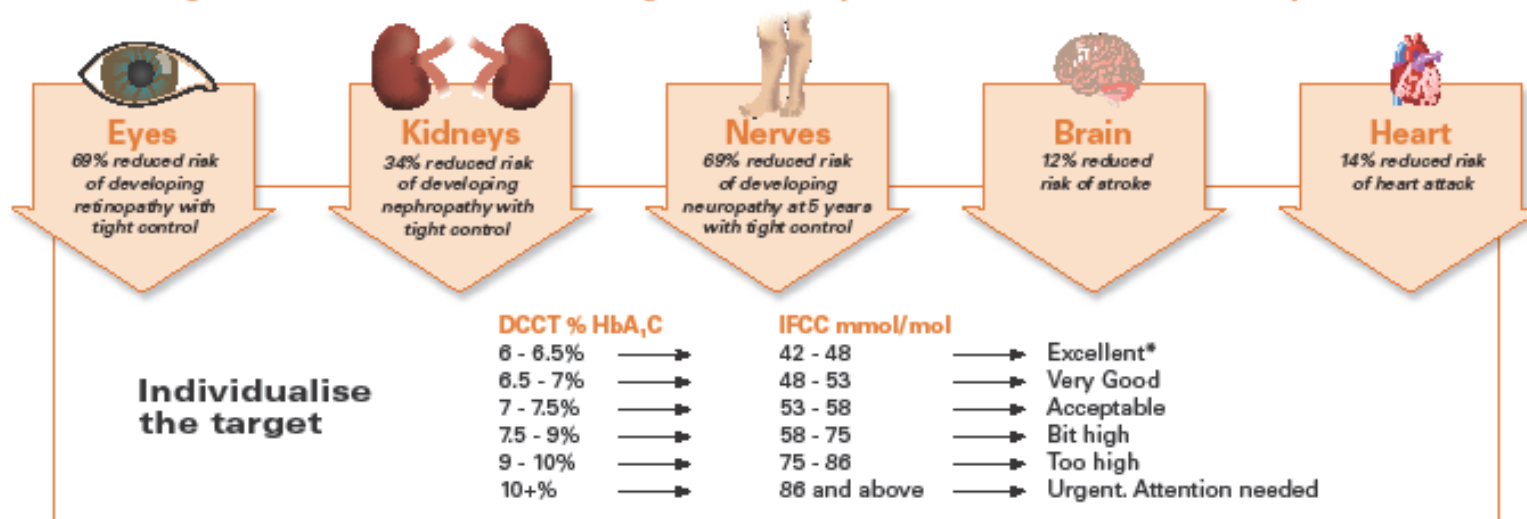


*The targets for everyone is less than 7% or 53 mmol/mol though targets should be individualised. **Caution should be taken with targets lower than 7 for those on insulin and sulphonylureas due to risk of hypoglycaemic events.**

HbA_{1c} is the best test of overall diabetic control.

If your HbA_{1c} is 8% or 64 mmol/mol or more, please contact your nurse or doctor.

Being in control or intensive management* can prevent diabetes-related complications



Developed by:
Rob Burton,
Diabetes Nurse
Specialist

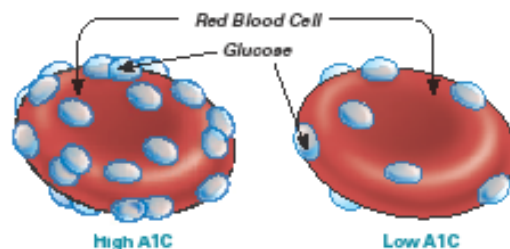
Overcoming the Barriers

You've got to be a good salesperson!

- Introduce concept of insulin deficiency early
- Tell the patient they are not “failures” for needing insulin
 - *(Explain UKPDS study showing , declining beta cells function over time ,50% initially at diagnosis then 4% decline every year therefore 50% of pts need Insulin after about 7 yrs)*
- Newer pens - “easy”, 4mm needles – “painless”
- Not complicated !!!!!Disposable pens !!!
- Use simple regimens to start *with (just one shot before bed helps to sleep better.....)*
- Emphasise the benefits of improved glycaemic control,
including *“feeling better/ more energy, less infections, less trips to toilets , less thirsty, benefits of or 11 mmols lowering Hba1c by 1% reduces complications by35 % UKPDS study ”*

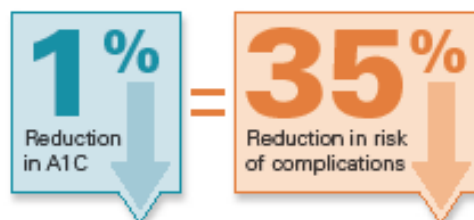
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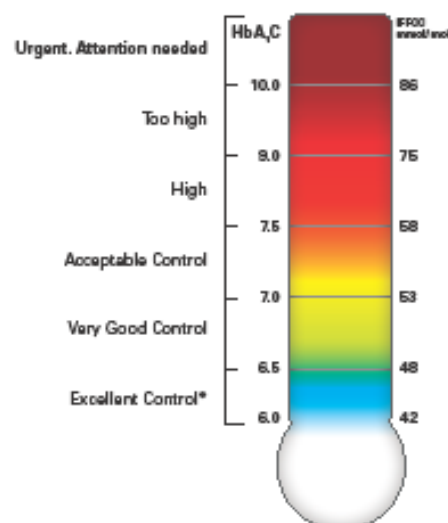
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Diabetic Control

HbA_{1c} Thermometer

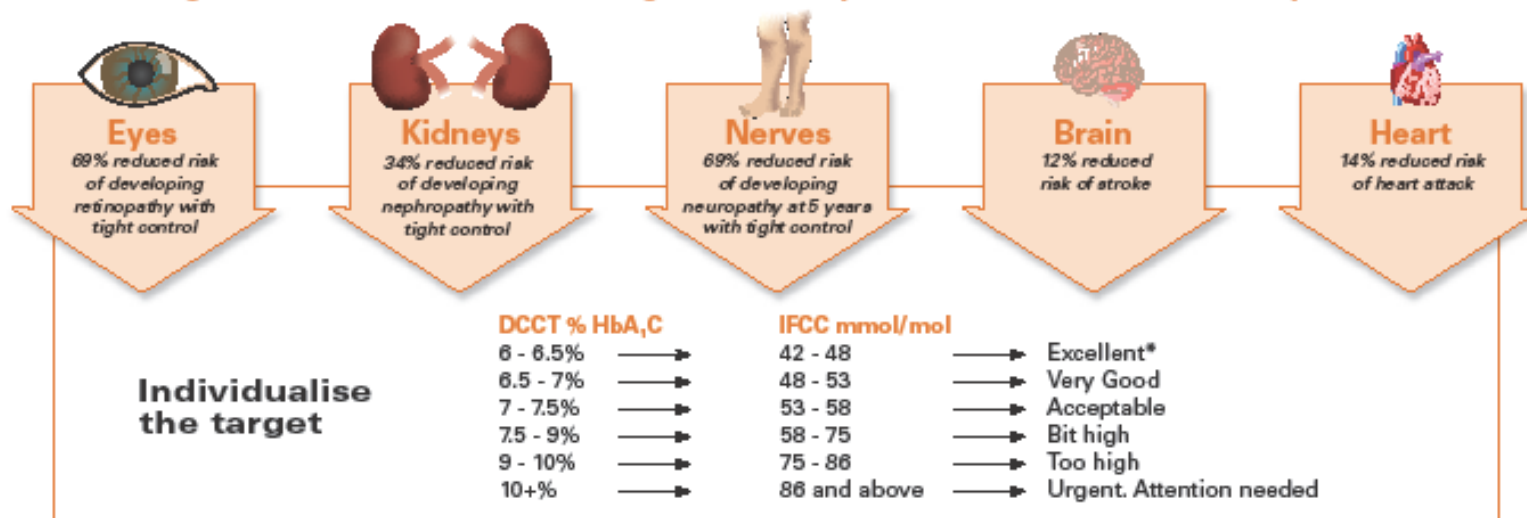


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When to start insulin?

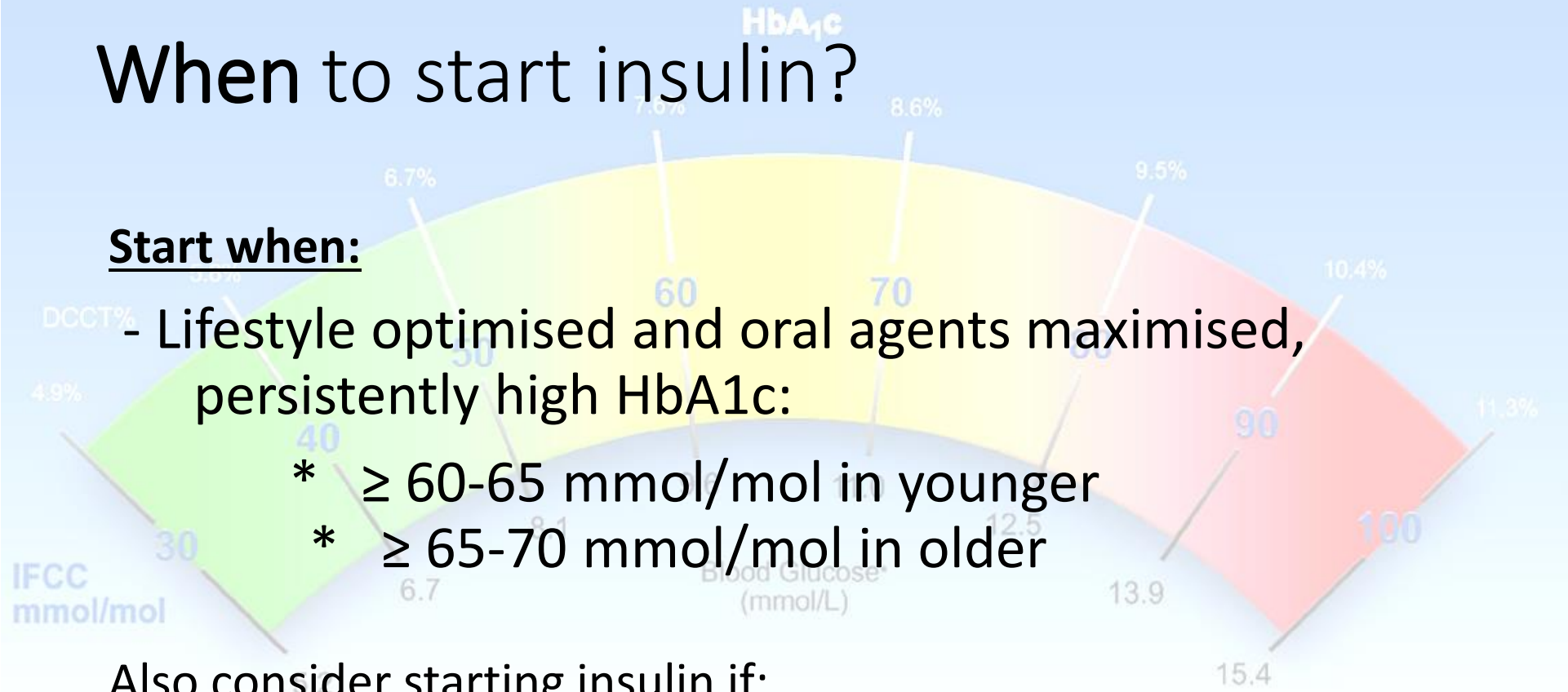
Start when:

- Lifestyle optimised and oral agents maximised, persistently high HbA_{1c}:

- * ≥ 60 -65 mmol/mol in younger
- * ≥ 65 -70 mmol/mol in older

Also consider starting insulin if:

- HbA_{1c} very high (>95mmol/mol) even if lifestyle/ oral treatment naïve
- Long duration of disease on near/maximum metformin (+/- sulphonylurea) with HbA_{1c} >75mmol/mol

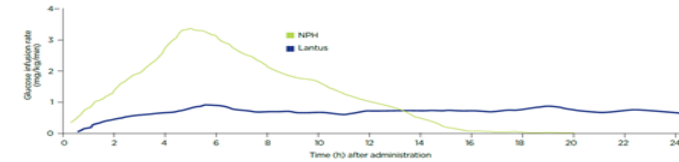


Insulins available on the Market

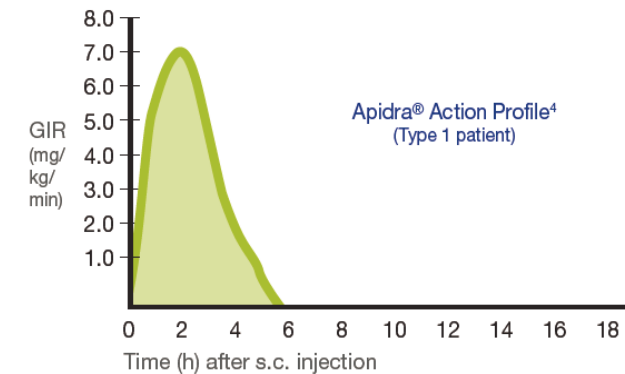


Lantus® (insulin glargine)

Profile of Lantus vs NPH^{1,4}



Lepore M et al. Diabetes 2000;49:2142-2148



NOVO NORDISK DIABETES TREATMENT RANGE¹

	Brand	Presentations	Schematic Insulin Profile
Modern Insulins	NovoMix® 30 30% Rapid-acting & 70% Protamined insulin aspart (lys)	FlexPen® 3mL, PenFill® 3mL, Vial 10mL	Onset: 10 - 20 minute Peak: 1 - 4 hours Duration: 24 hours
	NovoRapid® Rapid-acting insulin aspart (lys)		Onset: 10 - 20 minute Peak: 1 - 3 hours Duration: 3 - 5 hours
	Levemir® Long-acting insulin detemir (lys)		Peak: 2 - 14 hours Duration: Up to 24 hours
Human Insulins	Protaphane® Long-acting human insulin (lys)		Onset: 30 minutes Peak: 1 - 8 hours Duration: 8 hours
	Protaphane® Long-acting human insulin (lys)		Onset: 1.5 hours Peak: 4 - 12 hours Duration: Up to 24 hours
	Permix® 30 & Mixard® 30 30% Short-acting & 70% Long-acting human insulin (lys)		Onset: 30 minutes Peak: 2 - 12 hours Duration: Up to 24 hours
	Permix® 40 40% Short-acting & 60% Long-acting human insulin (lys)		Onset: 30 minutes Peak: 2 - 8 hours Duration: Up to 24 hours
	Permix® 50 50% Short-acting & 50% Long-acting human insulin (lys)		Onset: 30 minutes Peak: 4 - 8 hours Duration: Up to 24 hours

NovoCare® Customer Care Centre 0800 733 737 | www.novonordisk.co.nz

Lilly Insulin Range¹

Brand Name ^{1a}	Type of Insulin (generic name) / Product Description	Presentation	Schematic Action Profile ^a
Humalog®	insulin lispro Rbe RAPID-ACTING	10mL vials and 3mL cartridges	Onset: 0-15 mins Peak: 1 hour Duration: 3.5-4.5 hours
Humulin® R	insulin neutral (Soluble) SHORT-ACTING	10mL vials and 3mL cartridges	Onset: 30 mins Peak: 2-4 hours Duration: 6-8 hours
Humulin® NPH	isophane (NPH) INTERMEDIATE ACTING	10mL vials and 3mL cartridges	Onset: 1 hour Peak: 4-10 hours Duration: 16-18 hours
Humalog® Mix25®	25% insulin lispro, 75% insulin lispro protamine suspension Rbe PREMIXED INSULIN LISPRO	3mL cartridges	Onset: 0-15 mins Peak: 1 hour Duration: 16-18 hours
Humalog® Mix50®	50% insulin lispro, 50% insulin lispro protamine suspension Rbe PREMIXED INSULIN LISPRO	3mL cartridges	Onset: 0-15 mins Peak: 2 hour Duration: 16-18 hours
Humulin® 30/70	30% insulin neutral, 70% isophane (NPH) PREMIXED INSULIN	10mL vials and 3mL cartridges	Onset: More rapid than NPH alone Peak: 2-12 hours Duration: 16-18 hours

HUMAPEN LUXURA can only be used with Lilly 3mL insulin cartridges. BEFORE PRESCRIBING PLEASE REVIEW THE PRODUCT DATA SHEET.

How to start?

More than one way to skin a cat!

1. Conventional approach:

- **Bedtime basal insulin** (10 units of Protaphane / Humulin NPH / Glargine(Lantus), titrate dose up with an aim to normalise (4-7mmol/L) fasting glucose
- Particularly appropriate in those waking up with raised fasting glucose

How to start?

More than one way to skin a cat!

2. Alternative approach:

- **Pre-dinner premixed insulin:**

e.g. 12 units of NovoMix30 or HumalogMix25

- Good for those with high post prandial glucoses, large dinners, unlikely to manage more than 2 injections a day

How to start?

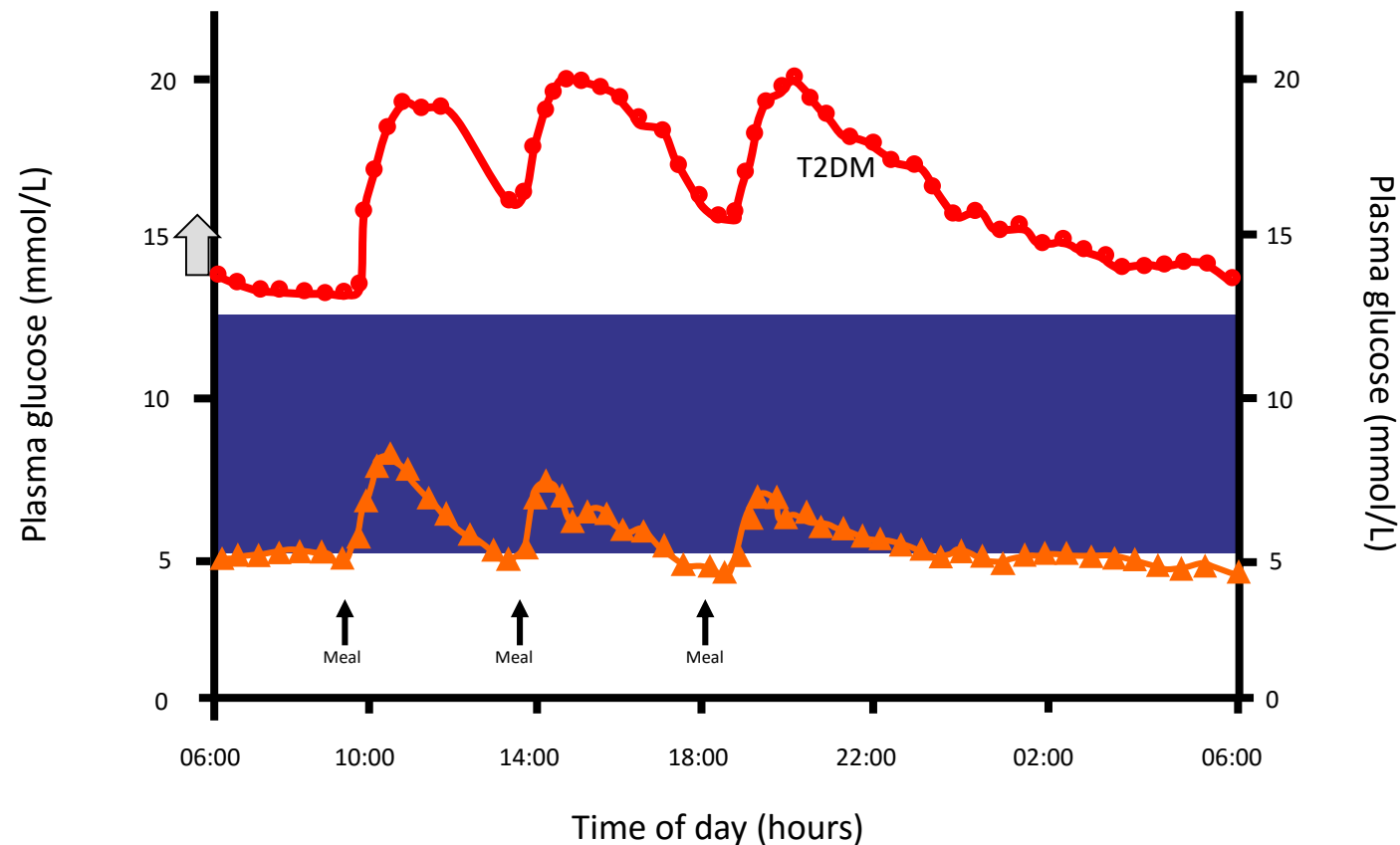
More than one way to skin a cat!

3. Other approaches:

- **BD premixed insulin:** consider in those with high post prandial glucoses, likely minimal β -cell function +/- very poor glycaemic control $>95\text{mmol/mol}$ (11%). *(No need to be on a sulphonylureas but continue Metformin)*
- **Prandial** (3x rapid acting insulin with meals) **or Basal bolus insulin** (Basal + prandial rapid acting insulin) – almost never used when initiating insulin in *T2DM* *(No need to be on Sulphonylureas but continue Metformin)*

Why start with basal insulin?

Comparison of 24-hour glucose levels in untreated vs treated patients with diabetes



Adapted from Hirsch I et al. *Clin Diabetes* 2005; 23: 78–86.

Dose adjustment – first fix fasting

Two dose adjustment schedules possible:

1. SLOW SCHEDULE (CAN BE PATIENT-LED)

Increase by 2 units of insulin every
3 days

continue until fasting BGL is ≤ 6.0 mmol/L

Increase dose only if FBG >4 mmol/L and accordingly decrease dose if FBG is <4 mmol/L.
Titration reviewed by HCPs at each contact.

Adapted from RACGP 2009/10 and Davies et al 2005.

Dose adjustment – first fix fasting

2. FAST SCHEDULE (PHYSICIAN-MANAGED)

Increase by 2–8 units of insulin depending on fasting BGL over previous 2–3 days

Mean fasting blood glucose (mmol/L)	Increase in insulin dose
<4	* See below
4–5.9	No change
6–6.9	2 units
7–7.9	4 units
8–10	6 units
>10	8 units

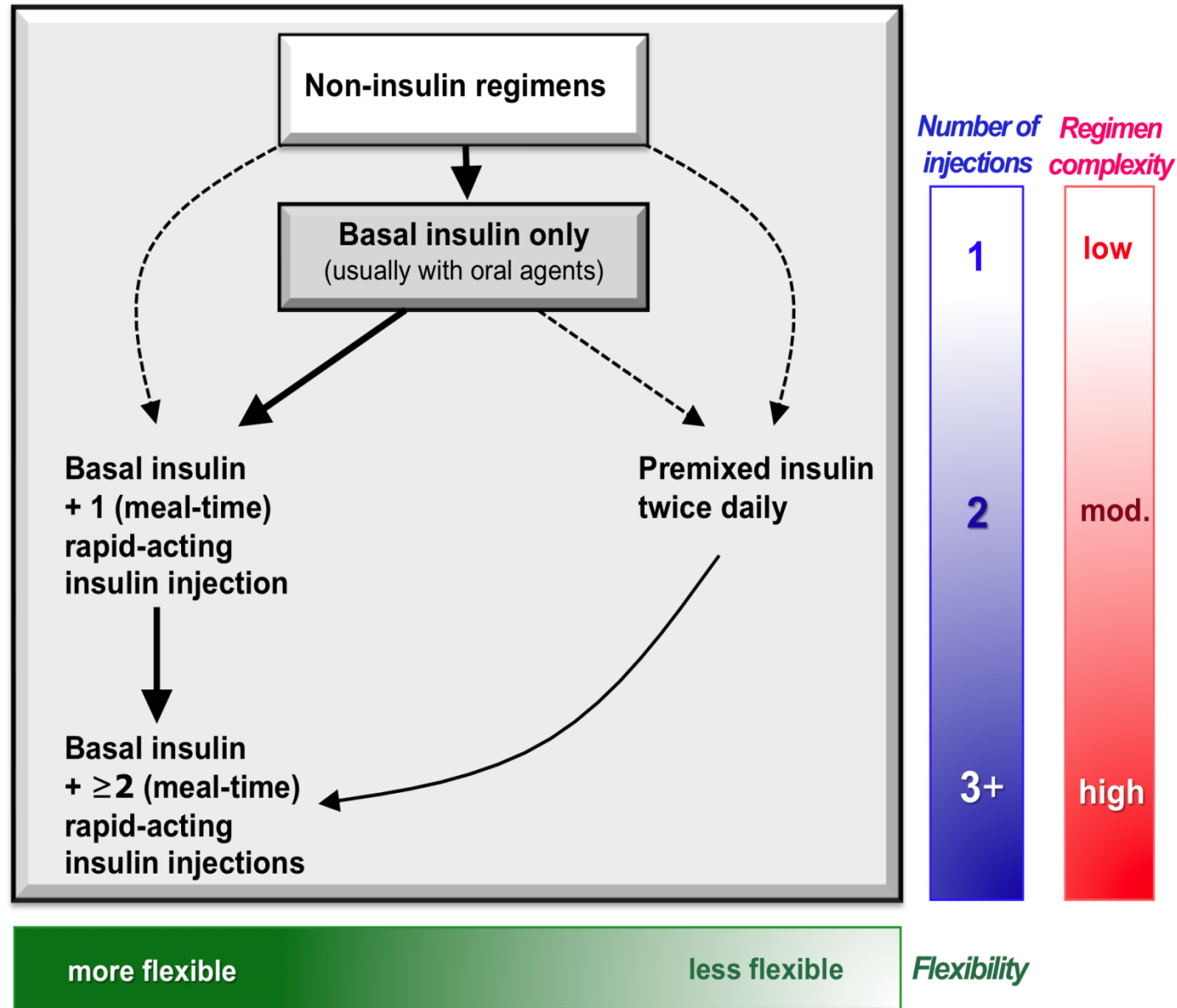
Starting dose 10 units, adjust dose twice weekly to reach the target FBG of <6mmol/L

Insulin dose may be decreased (small decreases of 2 to 4 units) if there is severe hypoglycaemia (requiring assistance) or if BGL <3.0 mmol/L in preceding week.

Do not increase insulin dose if fasting BGL <4 mmol/L at any time in preceding week.

Practice points

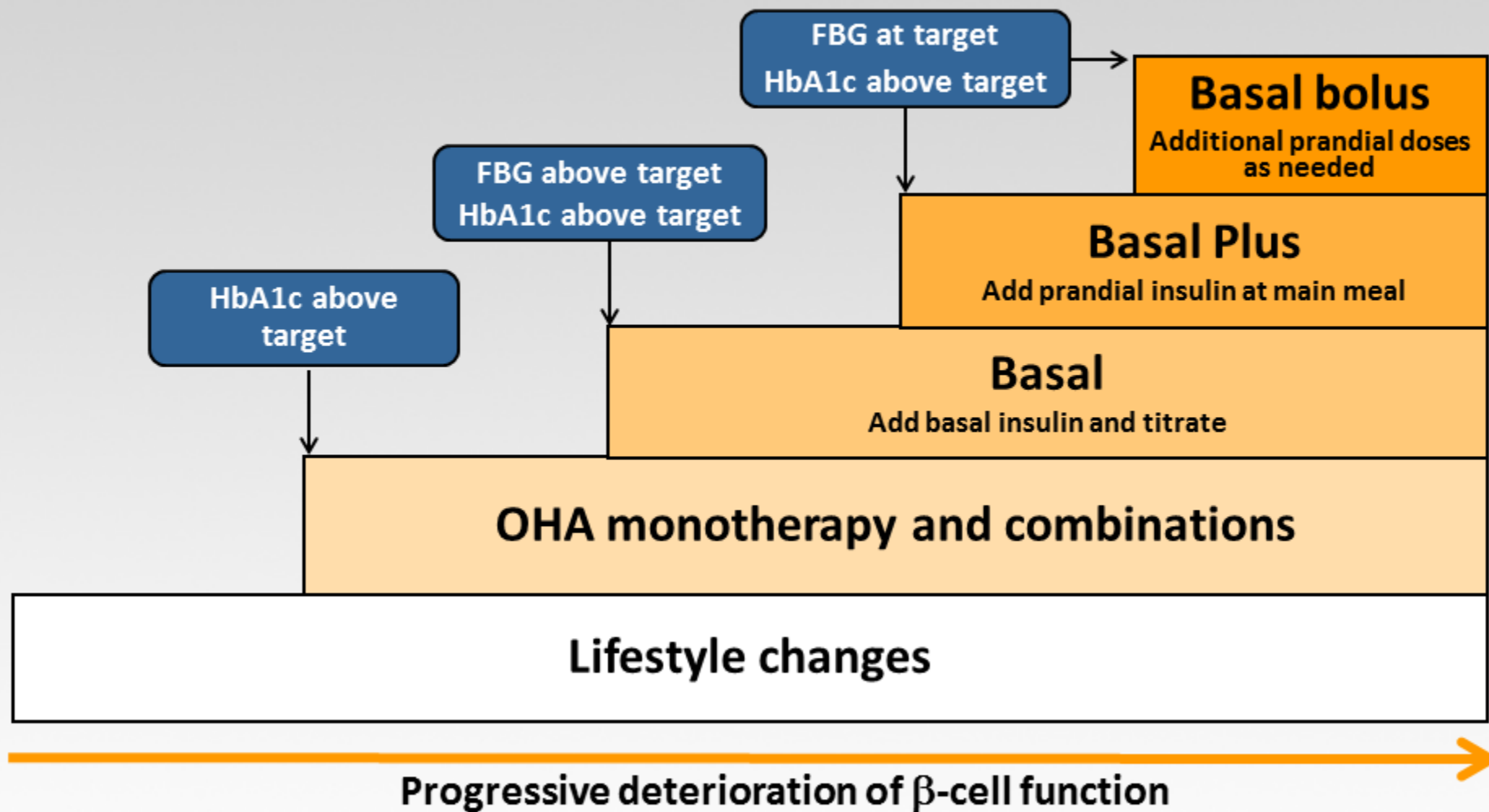
- Don't delay insulin initiation
- Keep it simple for you and patient – 10 units basal insulin (almost always at bedtime)
- Titrate dose up - Fix the fasting first, then look for hypers at other times



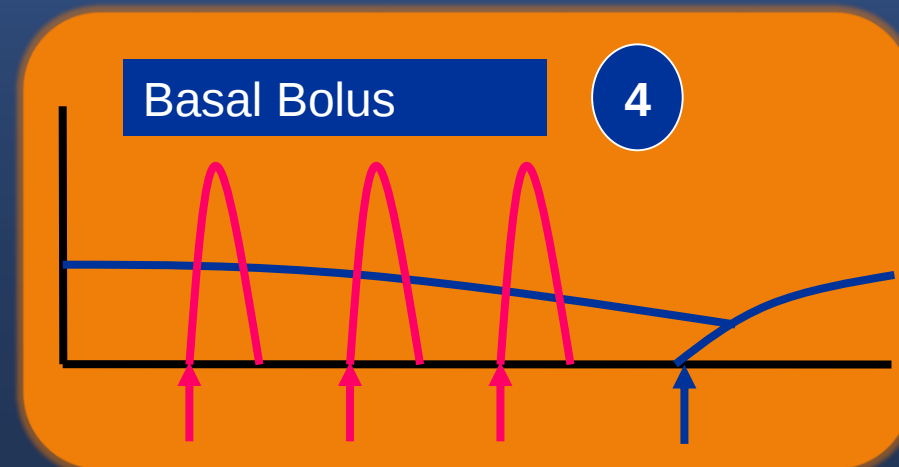
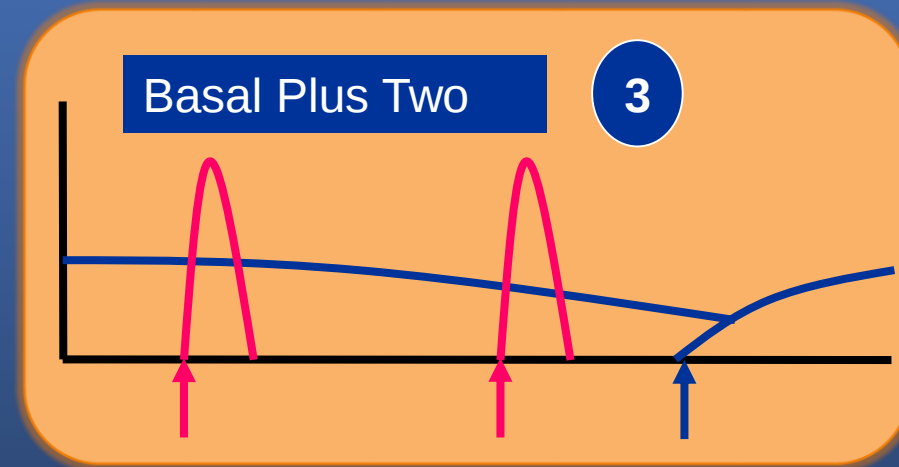
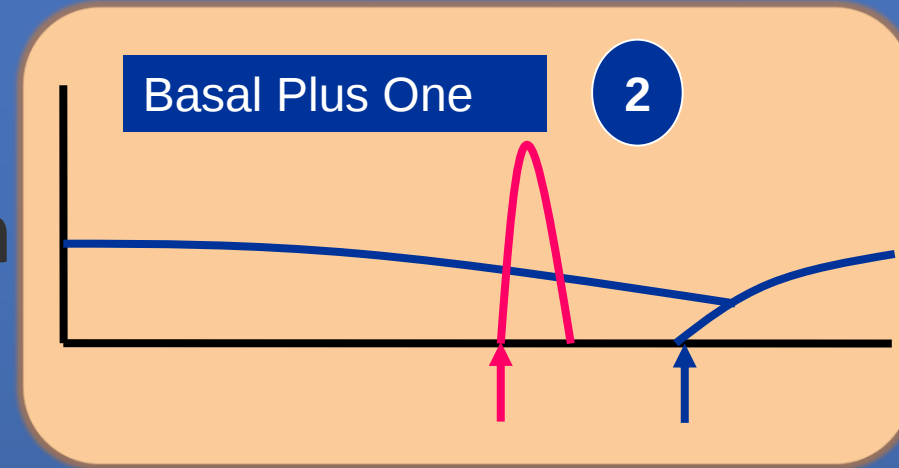
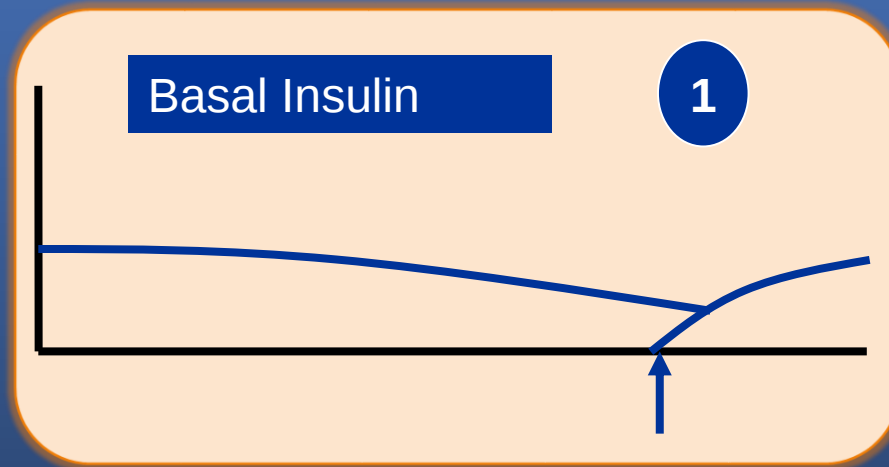
Sequential Insulin Strategies in T2DM

The Basal/Basal Plus Strategy for T2DM

Stepwise intensification of treatment for continuity of control



Basal Plus Strategy: Stepwise Intensification



Case 1 - Mr x

Persistent
hyperglycaemia (HbA1c
89) on Lantus 50u nocte,
Apidra 16u pre-dinner,
Metformin 1.5g BD,
Pioglitazone 45mg daily

What next??

[illegible]

Case 1 - Mr x

- Added Apidra 3 units pre-breakfast and pre-lunch
- Persistent high CG's (~10-15) at pre-lunch/dinner - Titrated from 4 units to 6 units,
- BS low bef bed so reduced the dinnertime Apidra
- BS too low fasting reduced the lantus to 46 units
- Off Pioglitazone, continued on Metformin, latest HbA1c 67 mmol/mol (CGs 3.5 – 15 mmol/L)

[illegible]

Case 2 - Mr T

- 46 yr old man with 17 year hx of T2DM, on combination of metformin (1.5g BD), pioglitazone (30mg daily) and Lantus 30u nocte (formerly on other insulins)
- Poor HbA1c, mostly >78 mmol/mol in preceding 2 years

Blood tests Diary

[illegible]

Case 2 - Mr T

What was done:

- Introduced rapid acting insulin (Apidra) - 4 units at pre-dinner
- Lantus reduced to 26 units then 22 units nocte due to hypos, pre-dinner Apidra titrated gradually up to 8 units
- Pioglitazone increased to 45mg daily

Thu																	
Fri																	
Sat																	
Sun																	

WEEK BEGINNING: (DATE)

	Type of Insulin	Insulin Injections				Monitoring Blood Glucose						Remarks Activity, stress, diet changes, time of hypos (noting blood glucose and treatment).	
		Units given				Breakfast		Lunch		Dinner			Before Supper or Bed
		Breakfast	Lunch	Dinner	Before Bed	Before	After	Before	After	Before	After		
Mon	Lantus				30	9.8		7.8		8.3		15.3	
Tues	Lantus				30	10.2		6.9		8.9		18.9	
Wed	Apidra		4		30	7.9		7.7		7.8		16.3	Hypo 4 am
Thu	Apidra		6		26	6.2		7.5		6.8		15.5	Hypo 3 am
Fri	Lantus		6		28	15.2		6.8		7.1		13.4	
Sat	Lantus		8		22	7.1		6.9		7.3		8.9	
Sun	Lantus		8		22	6.5							

Ms Blood sugar Diary

Case 3: Ms A

52 yr old woman with T2DM for 23 yrs, chronic poor control, HbA1c >80 for over a year

Metformin 1g TDS + on basal insulin since 5 years ago – Protaphane at bedtime, dose gradually increased to 48 units

Long history of patchy compliance, occasional CG testing, mostly fasting 8 - 15, pre-dinner 10 - 20 mmol/L

Date	Insulin	am	Lunch	Bed	Bef Bkfast	Bef Lunch	Bef Dinner	Bef Bed
Mon	Protaphane			48	8.9	12.4	8.2	14.5
Tues					8.2	13.1	7.9	18.2
Wed					10.2			
Thurs								

Case 3: Ms A

Management?

- Lifestyle intervention?
- Add/ adjust oral agents?
- **Optimise/ intensify insulin**
 - - Increase protaphane dose?
 - Add prandial insulin?
 - Switch to twice daily premixed insulin?

Case 3 Mrs A

- Simplify Metformin to 1.5g BD
- Switched insulin to premix (NovoMix30) – 20 units pre-breakfast and 20 units pre-dinner
- Got her to test CG's more intensively; pre-breakfast and pre-dinner, with a follow up phone call the following week
- Increase morning Novomix to 24 units continue 20 units at dinner

[illegible]

Steroids

Case 4 Mrs P

- 67 yrs old lady on metformin 1 gm bd + Gliclazide 80 mg bd + lantus 26 units nocte
- Fairly well controlled Hba1c of 62 mmols
- Recently started on Prednisone 15 mg for asthma since then BS gone crazy!!!

WEEK BEGINNING: (DATE)

	Type of Insulin	Insulin Injections				Monitoring Blood Glucose								Remarks Activity, illness, diet changes, time of hypoglycaemia (noting blood glucose and treatment).
		Units given				Breakfast		Lunch		Dinner		Before Supper or Bed	Over night	
		Breakfast	Lunch	Dinner	Before Bed	Before	After	Before	After	Before	After			
Mon	Lantus				26	7.2		6.9		7.8		9.8		
Tues	Lantus				26	6.9		18.9		22		13.5		Started Prednisone 15 mg
Wed	Lantus				26	7.1		19.6		23		15.5		Started Prednisone 15 mg
Thu	Lantus													
Fri	Lantus													
Sat	Lantus													
Sun	Lantus													

WEEK BEGINNING: (DATE)

	Type of Insulin	Insulin Injections				Monitoring Blood Glucose								Remarks Activity, illness, diet changes, time of hypoglycaemia (noting blood glucose and treatment).
		Units given				Breakfast		Lunch		Dinner		Before Supper or Bed	Over night	
		Breakfast	Lunch	Dinner	Before Bed	Before	After	Before	After	Before	After			

Steroids

Case 4 Mrs P

- Stop Gliclazide
- Start on 4 units of Novorapid then titrate up
- Add 3 units of Novorapid at lunch titrate

WEEK BEGINNING: _____ (DATE)														
	Type of Insulin	Insulin Injections				Monitoring Blood Glucose								Remarks Activity, illness, diet changes, time of hypot (noting blood glucose and treatment).
		Breakfast	Lunch	Dinner	Before Bed	Breakfast		Lunch		Dinner		Before Supper or Bed	Over night	
		Before	After	Before	After	Before	After	Before	After	Before	After	Before	After	
Sat	<i>Novorapid</i>													
Sun	<i>Novorapid</i>													
Mon	<i>Novorapid</i>	4			<i>20</i>	6.1		13.1		15.6		9.8		<i>Hypo 3 am</i>
Tues	<i>Novorapid</i>	6	3		<i>20</i>	5.1		10.3		12.7		9.2		
Wed	<i>Novorapid</i>	8	4		<i>20</i>	6.1		8.1		9		8.7		
Thu														
Fri														
Sat														
Sun														

2.30
3.00
3.30
4.00

Case 5 MR BD

- Continue metformin 500 mg OD
- Stop Gliclazide
- Start 4 units of Apidra with Breakfast and dinner
- Titrate Insulin dose until BS stabilises
- Ended up on Apidra 6 units amm, 4 units at lunch and 6 units at dinner
- Lantus reduced to 40 units nocte .

[illegible]

Insulin doses in T2D

- Requirements depend on insulin (body) resistance
- Duration of DM will affect remaining beta cell function
- Correct dose of insulin is when you achieve target blood sugars.

Insulin doses in T2D

When might changes need to be made?

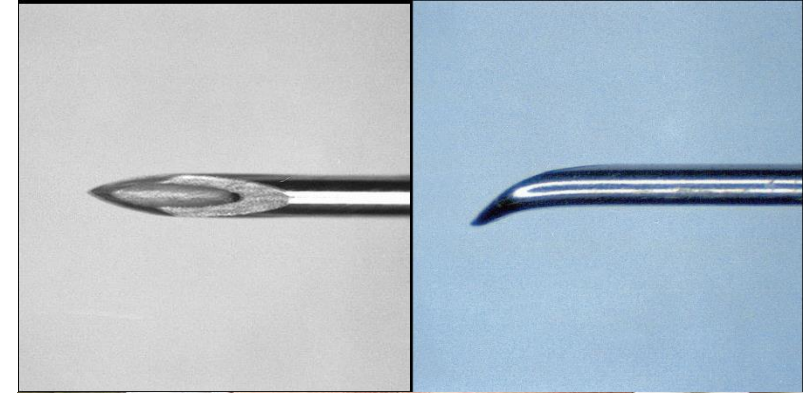
- Hypos
- BGLs consistently > 15 mmol/L
- Illness
- Steroids/medications causing hyperglycaemia
- HbA1c not on target

Pen needles

- Needles-100 needles for 3 months

After 2nd Use

- ☆ First use: Lubricant removed
- ☆ Needle hooking second time
- ☆ After 6 uses~
 - ☆ fishing anyone?



Hypoglycaemia

Need to explain what Hypoglycaemia is!!!!!!

Pt can freak out!!!

- *Was insulin taken at appropriate time?*
- *Are they missing doses and then overcompensating later?*
- *Are they missing meals?*
- *Are they changing quantity or quality of food in order to lose weight-without changing the dose?*
- *Are hypos occurring on particular days of the week, i.e at week ends??*
- *Was Alcohol a factor??*
- *Was exercise a factor?/ delayed hypo due to exercise during the day??*
- *Pre and post menstrual changes in BS levels??*
- *Have injection sites been checked for signs of lipohypertrophy??*
- *Rotating sites frequently??*
- *Mixing insulin correctly??*
- *Hot weather/Hot bath ??*
- *Do they need to review their diet??*
- *Is the Insulin regimens right for the patient??*



Hypoglycaemia

**Caused by the
body's initial reaction**
(trying to
increase BGL)

Irritability

Anxiety

Hunger/ feeling sick

Paleness

Shakiness

Heart pounding

Sweating

Numbness – fingers, lips, toes



From the brain

Inability to concentrate

Weakness, dizziness

Headache

Confusion

Blurred or double vision

Odd behaviour

Slurred speech

Lack of coordination/ unsteady

Lapses in consciousness

Convulsions



HYPOGLYCAEMIA - Treatment

STEP 1 – 15-20g of fast acting carbohydrate.

Regular fizzy drinks, jellybeans (6-8), glucose tablets, 3 teaspoons of sugar. [**NOT** chocolate, cakes or biscuits!]

STEP 2 – Retest blood sugar if back above 4mmol then move to step 3, if not repeat step 1.

STEP 3 – Meal if due or 15-20g of slow acting carbohydrate. Piece of fruit, slice of bread, 2 plain biscuits

SICK DAYS – KEY MESSAGES for T2DM



Key Messages:

- Tell someone, anyone (family, friend, work colleague) that you are sick
- TEST FREQUENTLY (3-4x daily or more)
- Continue your diabetes medications, do not reduce them without advice. (However, do not take Metformin if vomiting and or diarrhoea).
- Have plenty to drink . Replace meals with drinks (May need sweet drinks / soft foods if necessary).
- Contact GP or diabetes team for further advice
- Go to hospital if feeling drowsy & confused, experiencing breathing difficulties, stomach pain, vomiting, bsl's >15mmol consistently or unable to keep bsl's >4.0

Drury, P. & Gatling, W. (2005). *Diabetes: Your questions answered*. Edinburgh, New York: Churchill Livingstone.

Travel

Preparation & precautions:



- Vaccinations
- Customs letter detailing all medication
- Customs (200mls maximum rule for liquids)
- Hypo treatment
- All diabetes supplies i.e. Meds, insulin, pens, needles and testing equip **MUST** be in hand luggage.
- First aid kit
- Sick day action plan
- Consider Insulin storage while away
- Possible changes in lifestyle practices while away.

Everyone starting insulin **MUST** be made aware of their responsibilities regarding driving.

Regular testing and hypo management / avoidance is of the utmost importance.

Any Type 2 patients on insulin who wish to hold a passenger license, heavy vehicle license or driving instructor license are likely to require :

1. **Six-monthly medical certificate from a GP documenting:**
 - adherence to treatment
 - proof of regular self-testing of blood glucose with satisfactory levels
 - the absence of hypoglycaemic episodes or unawareness
 - the absence of significant diabetic complications
2. **A regular pattern of shifts with adequate meal breaks**
3. **A satisfactory annual specialist review**

Reference: New Zealand Transport Agency. (2009). *Medical aspects of fitness to drive.*

SHARPS DISPOSAL



- Lancets and pen needles are sharps that need safe disposal.
- Advise patients to speak with their pharmacist about safe disposal options.
- Diabetes Auckland or Comprehensive Care provide sharp containers for \$5.00 (1 L) and \$8.70 (1.5L) for sharps disposal.
- Auckland now can dispose of sharps in any Chemists

Take home messages

- Start insulin in a timely fashion, employ techniques for a smooth initiation
- Early intensive CG testing with up-titration of dose
- Expect insulin regimen intensification with time – initially dose escalation of single daily injection, later more regular injections
- Individualise glycaemic targets and insulin regimen

Thank you

Questions