#### Edition 2024 | English



# **NEXT STEP LIVING LONGER BOOKS**

## Diabetes - Book 2

# **METFORMIN** (Biguanides group of medicine)

The most famous and well known medicine 1<sup>st</sup> medicine to start even before we have diabetes

Author:

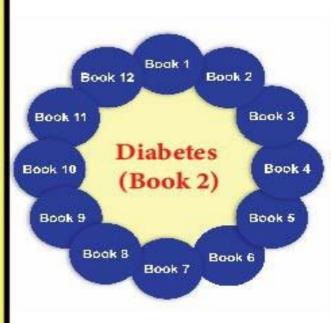
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From family of doctors

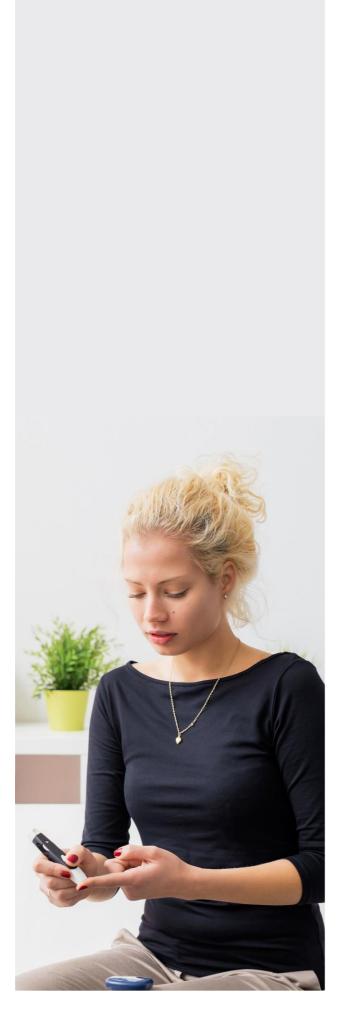
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Book 2



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IN 2021: No reason for kidneys to fail!! Before 1921: Yes, You would have died. (year of Nobel Prize for Insulin)

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#### Introduction



**Frederick Grant Banting** 



John James R.Macleod

 The Nobel Prize in Physiology or Medicine 1923 was awarded jointly to Frederick Grant Banting and John James Rickard Macleod "for the discovery of insulin" in 1916.

Chap1Fig1

Insulin was discovered in 1921 leading to a nobel prize. In 2021 we have 12 groups of medicines for medicine diabetes.

#### Think About It!

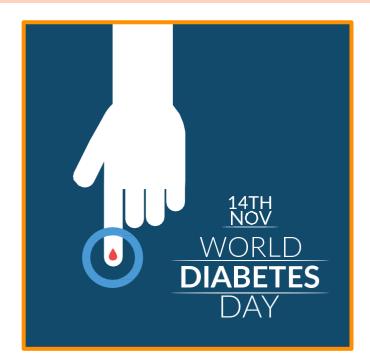
# **Before 1921**

Every child who was born with Type 1 diabetes or lack of insulin would eventually die.

The discovery of insulin or should we say the discovery of insulin as a medicine has changed the history of diabetes and its treatment.

## In 2021

#### We can now have a healthy life with functioning kidneys till very end!!

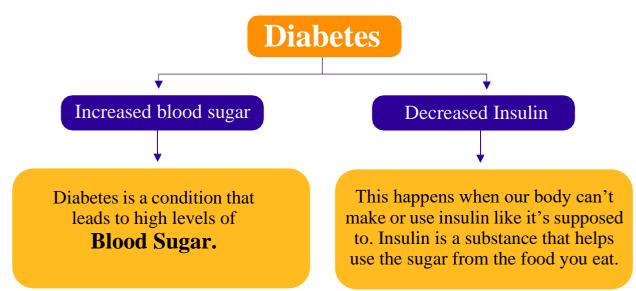


But yes, we have 12 to 13 different groups of medicines for high blood sugar.

<u>The golden principle</u> of the medicinal treatment of diabetes is that we start with one medication early in the course of treatment of diabetes.

We keep adding medicines as needed to control our blood sugar (along with diet & exercise)

Chap1Fig1



# The Golden Principle in Management of high blood sugar (Diabetes)

The golden principle of the medical treatment of diabetes is that Once we have high blood sugar (Diabetes) – exercise and diet management are needed life-long.

#### **Together diet, exercise and medications – all will help!!**

Risk of taking medications are minimal as compared to the benefits that we receive from these medicines!

1. We start with <u>one medication</u> early<br/>in the course of treatment of diabetesStart with one medication2. Targets for blood sugar control is<br/>decided by physician.Decide Targets3. Usual target is <u>HbA1c < 7</u>HbA1c < 7</td>4. As voungish we are more, we want<br/>to keep blood sugar is close to<br/>100mgBlood sugar close to 100mg

#### Continued

The golden principle of the medical treatment of diabetes is that Once we have high blood sugar (Diabetes) – exercise and diet management are needed life-long.

- 5. <u>Today in 2021</u> are several groups of excellent medicines are available:
- a) which can <u>bring down our blood sugar</u> by different mechanisms of action.
- **b**) As a golden rule, what we should start with <u>one medicine</u> if our blood sugar is not extremely high.
- c) And as time goes on. We can add a second medicine.

Start with one medicine, if blood sugar is not very high

Start with one medicine, Add a second medicine, if needed And add a third medicine, if necessary

d) Or we can even add a third medicine which works by a different mechanism.

Diet and exercise are an integral part of high blood sugar management!

# Diet and exercise are an integral part of high blood sugar management!

Purpose of medical management and medici nes for diabetes is:

• To manage our blood sugar

• As close to 100mg as possible



Chap3Fig1

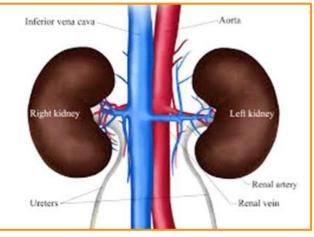


Chap3Fig2

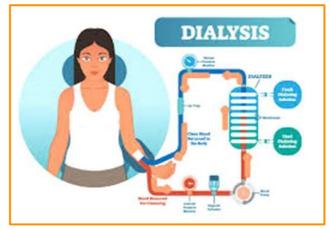


Chap3Fig3

• And HbA1c < 7



Chap3Fig4



Chap3Fig5



Chap3Fig7

• To prevent kidney failure

• And no dialysis

• No heart attack (Diabetes is literally one of the reasons)



Chap3Fig8



Chap3Fig9

• No blindness (Diabetes is one of the reasons)

• No amputation of leg (Diabetes is one of the reasons)

Yes, Diabetes management can really reduce complications

If we manage diabetes from day one or <u>even before</u> <u>we have actual diabetes</u> (called prediabetes), Complication are dramatically delayed by



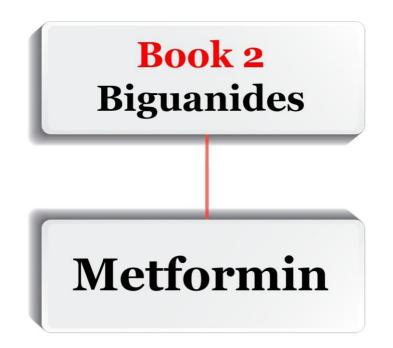
# Is it true? ✓Really Yes! it is True

#### **Today in 2020/2021 we have excellent medicines** available

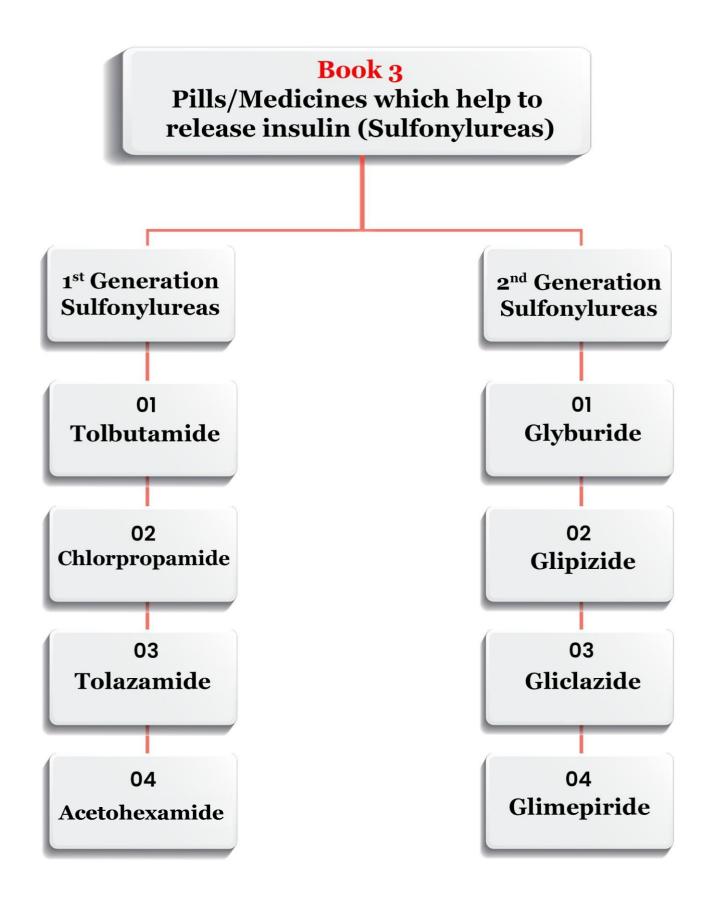
There are a number of medicines available in the market today for the treatment of Diabetes. The main groups which these medicines belong to are:

#### **Book-2** Biguanides

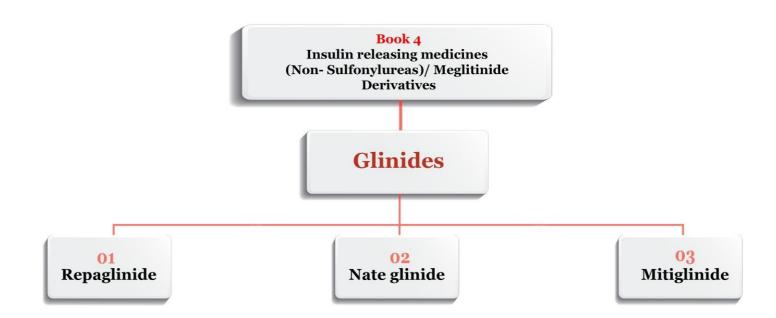
**Metformin:** The most famous and most used medicine and 1st medicine to start the treatment of diabetes.



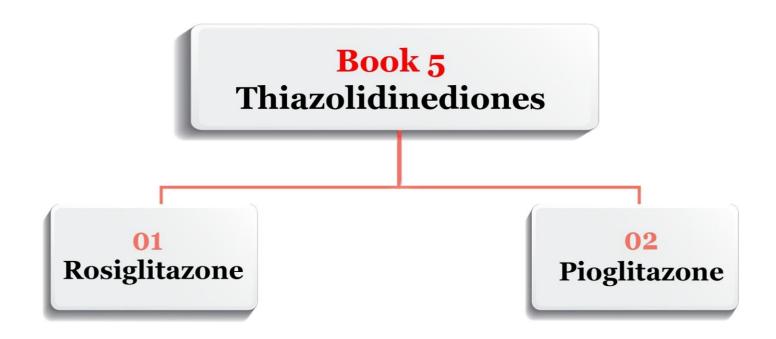
#### **Book-3** Pills/Medicines which help to insulin (Sulfonylureas)



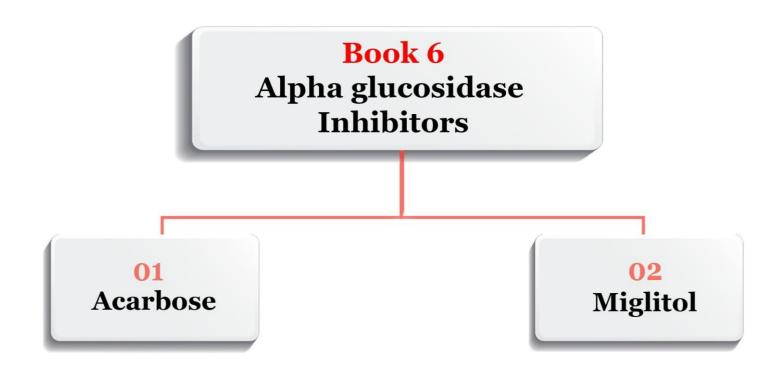
#### Book-4 Insulin releasing medicines (Non- Sulfonylureas)



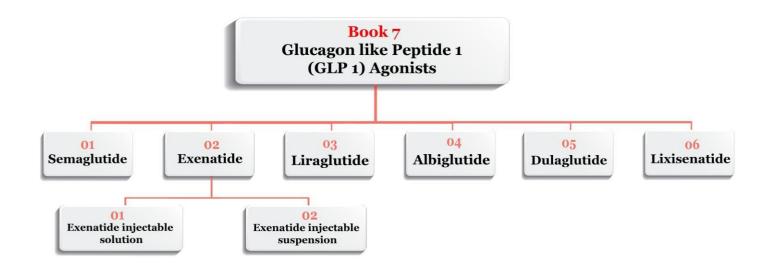
**Book-5** Thiazolidinediones



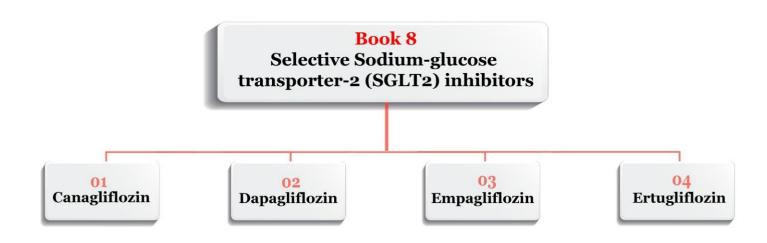
**Book-6** Alpha glucosidase Inhibitors



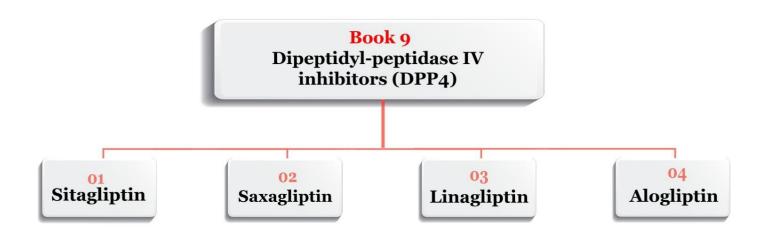
**Book-7** Glucagon like Peptide 1 (GLP 1) Agnostics



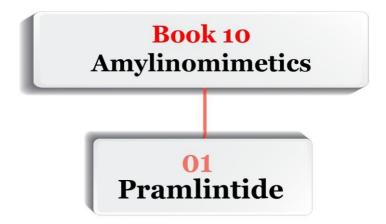
# **Book-8** Selective Sodium-glucose transporter-2 (SGLT2) inhibitors



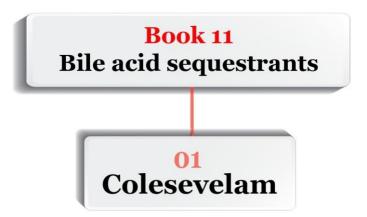
**Book-9** Dipeptidyl-peptidase IV inhibitors (DPP4)



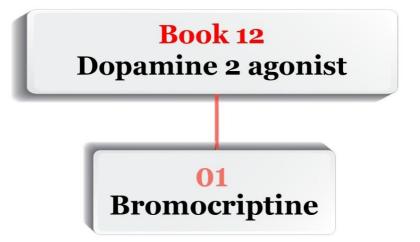
**Book-10** Amylinomimetic group



**Book-11** Bile acid sequestrants



Book-12 Dopamine 2 agonist



# There are dramatic advances in treatment of diabetes today in 2021

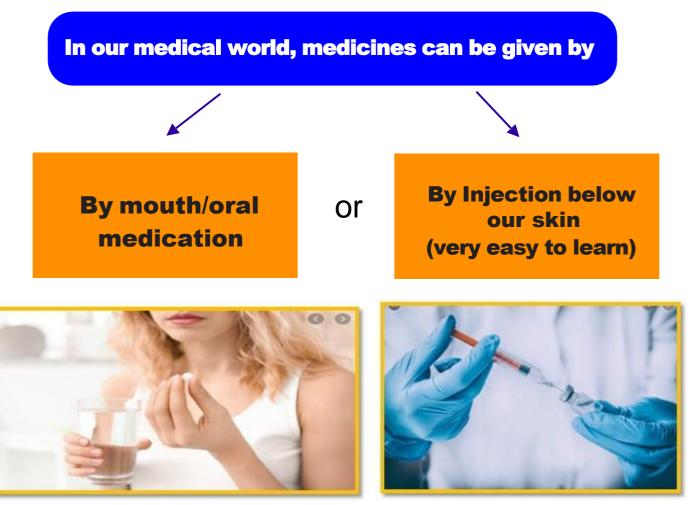
#### Think About It!

We are **NOT** perfect; we don't need to be. But we **CANNOT** quit trying either.

Every high sugar management prescription always includes:

- 1.Medicine(s),
- 2. Diet, and
- 3. Exercise

Without these, it is impossible to achieve our targets.



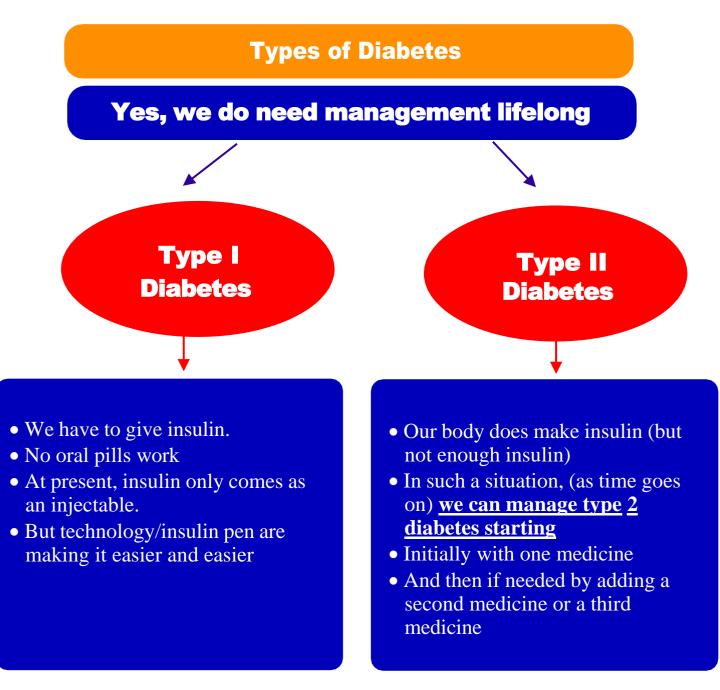
Chap6Fig1

Chap6Fig2

#### **Type 1 vs Type 2 Diabetes**

#### <u>Think About It!</u>

We were always scared of needles and injection needs some degree of training so that it does not lead to any infection on the injected place. We have to follow the sterile techniques that do not lead to any infection of the injection sites.



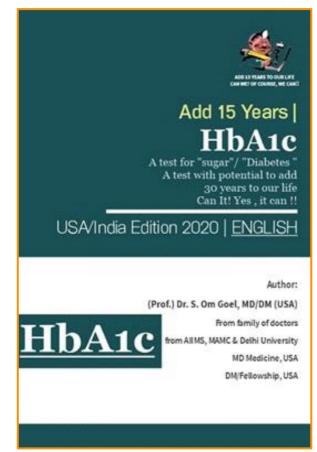
How do we decide less dose or more dose of medicines

#### Think About It!

Our blood sugar changes all day based on

- Our meals,
- Level of activity etc.
- HbA1c remains stable over three months.

#### How do we decide less dose or more dose or combination of medicines?



Chap8Fig1

#### I. <u>We actually use HbA1c as our guideline for</u> the treatment of diabetes.

II. We set up a <u>target for HbA1c</u> in diabetic managementand we want that the <u>HbA1c</u> <u>always remains 7 or less than 7.</u>

#### HbA1c < 7

III. <u>If we met our target</u>, we can lower the dose.

Start with one medicine, if blood sugar is not very high

**IV. <u>If our HbA1c is high</u>**, then we can give the increased dose or add a 2nd pill or add even a 3rd pill. Start with one medicine, Add a second medicine, if needed And add a third medicine, if necessary

# Early diagnosis and early treatment: an absolute necessity in 2021

#### Think About It!

Every mom knows how her baby will start walking around in 1 year of age.

or

Every physician knows how to diagnose diabetes (called Prediabetes) year before you have any "symptoms".

Actually, if we can diagnose Diabetes, even before it happens to us (called as Prediabetes), we can really delay complications as mentioned and it will be a very wise thing to do.

Most important thing we have to understand is that if medicines are taken early in the diagnosis of high blood sugar/diabetes then it can delay complications related to diabetes by

or



ADD 15 YEARS TO OUR LIFE CAN WE? OF COURSE, WE CAN!!

# Diabetes - Book 3.2 SULFONYLUREAS

# (2<sup>nd</sup> generation medicine)

IN 2021: No reason for kidneys to fail!! Before 1921: Yes, You would have died. (year of Nobel Prize for Insulin)

#### **Introduction to Sulfonylureas**



Chap1Fig1

#### Think About It!

<u>Sulfonylureas Insulin secretagogues</u> are a group of medicines which every doctoris very familiar with.

These medicines have been used for years and years.

When we were young back in 1970s-80s, we were using the 'first generation of sulfonylureas.'

As years went by, drug companies/pharmaceutical companies manufactured better acting medicines belonging to this group which we call 'second generation of sulfonylureas.'

#### **Sulfonylureas**

#### How do they work?

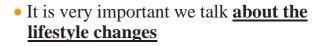
1. They work by directly stimulating the **Release Insulin directly** release of insulin from beta cells of in Pancreases. Langerhans which lie in the pancreas 2. You can take these medicines with food or Lower the blood before food. They lead to the lowering of glucose level. **blood glucose level** indirectly by stimulating the release of the insulin 3. They are only useful for type 2 diabetes **Only useful for** where our beta cells in pancreas make some **Type 2 Diabetes.** degree of insulin 4. These medicines do lead to weight gain Lead to weight gain. which is the effect of the insulin released **Excellent medicines to** 5. And they are very good medicines in lowering our blood sugar lower blood sugar.

If we miss our meal or we take too much of the medicine, then it can lead to actually lower blood sugar than the normal and we have to be really careful about very low blood sugar.

#### How to treat type 2 Diabetes?

#### Treatment of the type 2 diabetes always includes

So, we must educate as much as we can, the patient giving him as much insight into diabetes (high blood sugar).





Chap2Fig1



Chap2Fig2

• We talk about <u>the importance of diet</u>



Chap2Fig3



• Weight loss always helps

• We talk about <u>the importance of</u> <u>exercising</u>

Chap2Fig4

#### **Complications of suffering from Diabetes**

#### **Complications of diabetes**

What are the complications of diabetes?

Diabetes #1 cause of <u>Kidney Damage</u>

**Because Diabetes is so common** 



Chap3Fig1

Diabetes #1 cause of <u>Blindness</u>

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**Because Diabetes is so common** 



Chap3Fig2

Diabetes #1 cause of <u>Nerve Problem</u>

**Because Diabetes is so common** 



Chap3Fig3

Diabetes #1 cause of Loss of Limbs

**Because Diabetes is so common** 



Chap3Fig4

Diabetes #1 cause of Sexual Dysfunction

#### **Because Diabetes is so common**



Chap3Fig5

#### Diabetes #1 cause of Increased Risk of Heart Attack

**Because Diabetes is so common** 



Chap3Fig6

#### How to minimize the risk of Low Blood Sugar?

<u>Sulfonylureas</u> work by releasing insulin. Insulin lowers blood sugar but if our blood sugar is already low, it can make our sugar levels very, very low which is not safe for us.

It is important to know how to minimize risk of low blood sugar

• Exercise which increases the utilization of the blood sugar. If we take the medicine with heavy exercise, it can further lower the blood sugar



Chap4Fig1

• <u>We miss our meal.</u> We still take the medicine, but we do miss our meal. (We take our food, it raises the blood sugar and the medicines bring it down.)



Chap4Fig2

• Another thing is when <u>our medication</u> <u>starts</u>, we start with smaller doses and we carefully raise the medicine as needed.

These are long acting medicines which keep working for a long time



Chap4Fig3

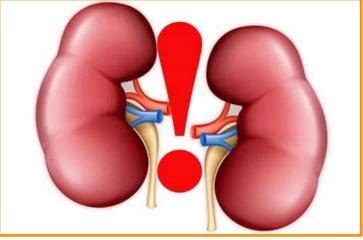
• If patients consume <u>too much alcohol then</u> <u>it leads to poor nutrition status</u> and affects the liver.

This affects the storage of glucose in our liver and one becomes very sensitive to low blood sugar



Chap4Fig4

• We have to understand that people with kidney failure actually need less insulin (as insulin stays longer in the blood.)



Chap4Fig3

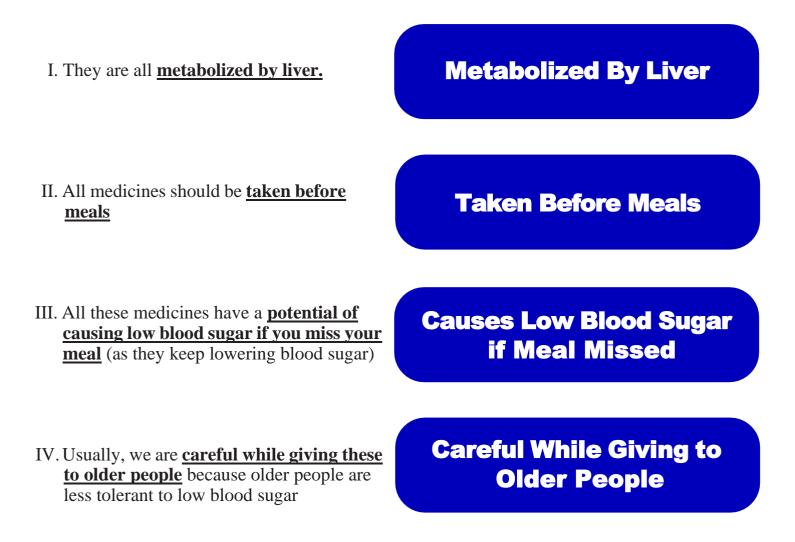
#### • <u>Some medication can cause drug</u> <u>interaction</u> which decreases the effect of the medication. There is a risk of high blood sugar. If we take higher dose of medication once admitted to the hospital, then we must monitor our blood sugar levels and balance future doses



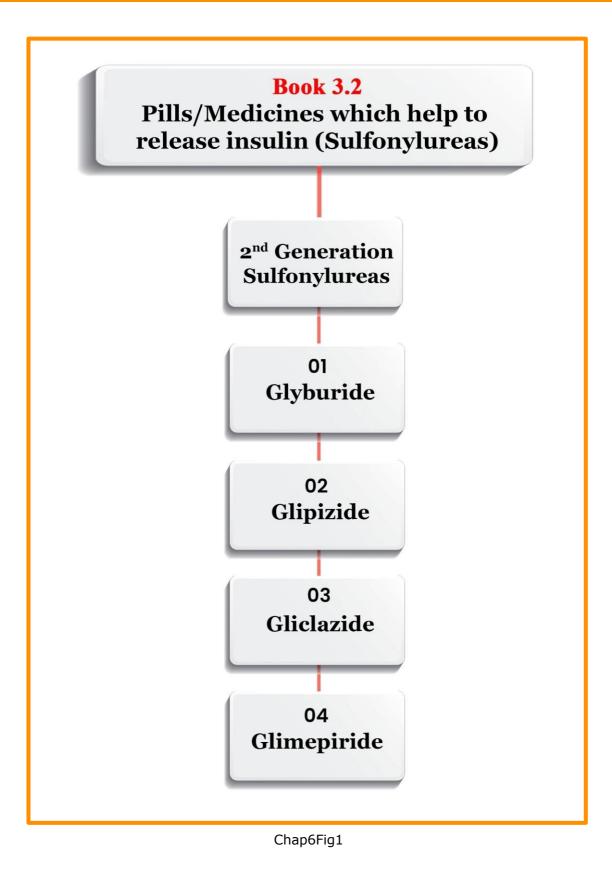
Chap4Fig4

Introduction to First and Second Generation Sulfonylureas

All the second-generation sulfonylureas and first-generation sulfonylureas, are very potent (they lower blood sugar)

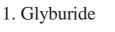


#### **Second Generation Sulfonylureas**

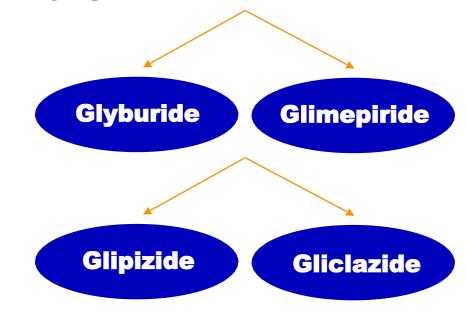


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#### There are 4 medicines in this group:



- 2. Glipizide
- 3. Gliclazide
- 4. Glimepiride



The best sulfonylureas to be used are usually glipizide and glimepiride. (Gliclazide is also good sulfonylureas to use, but it is not available in US.)

#### These medicines (second generation) are very effective.

#### **Glyburide**

- I. All diabetes medicines should be taken before meals as a golden rule.
- II. We start on a low dose and increase the dose if needed

III. We should usually take medication at the same time every day.

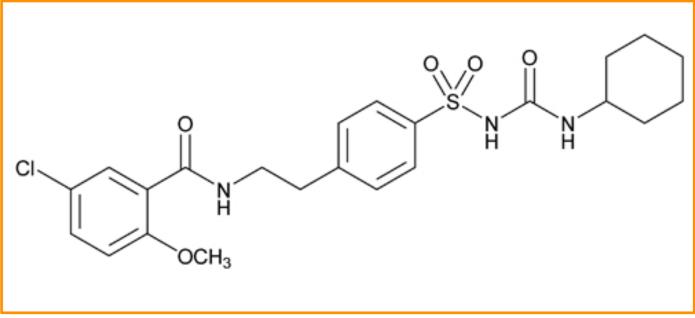
#### **Glyburide FDA Approval History**

FDA Approved: Yes (Approved In September, 2002) Brand name: Diabeta (India), Glycron (India), Glynase (USA), Micronase (USA) Generic name: Glyburide Dosage form: Tablets Treatment for: <u>Diabetes - Type 2</u>



#### <u>Glybuide</u>

Chap6Fig2



Chap6Fig3

#### What we need to know about Glyburide:

Usual Dose: 2.5mg to 10mg
Usual dose: 2.5 mg
We can give it once a day or we can give it in divided doses. As a physician, I will prefer divided doses i.e. 2.5 mg in morning with breakfast and 2.5mg with lunch, and 5 mg with dinner

**Once a day or in divided doses** 

• There is a long-acting formulation of glipizide, also **glipizide XL which obviously we give once a day only** 

#### **Glipizide XL is given once a day**

• Maximum Dose: 20mg

#### **Maximum Dose: 20mg**

### Trade Name- Dose- Cost in USA- Cost in India

# GLYBURIDE

Please always combine any blood sugar medicine with diet and exercise.

#### All medicines continue life-long

Generic Name & Trade Name	About Glyburide	Cost of Glyburide for one month (in USA)	Cost of Glyburide for one month (in India)
Generic Name: Glyburide	<ul> <li>Usual dose: 2.5-5mg once a day</li> <li>Can be increased to</li> </ul>	Dose = Once a day (5mg) 1-month =	<mark>Dose</mark> = Once a day (5mg) <b>1-month</b> =
Trade Name:	20 mg once a day	100 tablets	100 tablets
Diabeta (India)	As a Golden Rule – before starting any	<u>Cost</u> = \$20	Cost = Rs 80
Glycron (India)	medicine, liver functions and kidney		
Glynase (USA)	functions should be screened.		
Micronase (USA)			
		*All costs are meant for a rough estimate of one-month expense.	*All costs are meant for a rough estimate of one-month expense.



Chap6Fig4

#### <u>Glipizide</u>

- I. All diabetes medicines should be taken before meals as a golden rule.
- II. We start on a low dose and increase the dose if needed
- III. We should usually take medication at the same time every day.

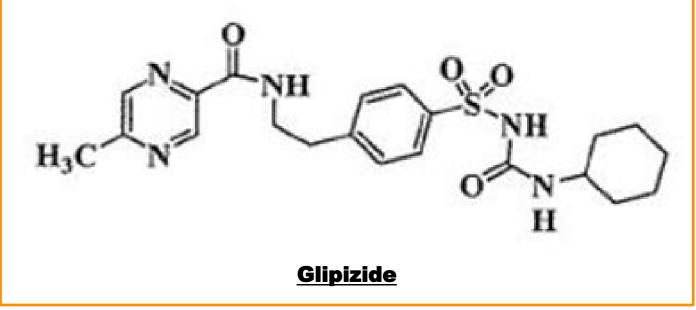
#### **Glipizide FDA Approval History**

FDA Approved: Yes (Approved In October, 2002) Brand name: Diabeta (India), Glycron (India), Glynase (USA), Micronase (USA) Generic name: Glipizide Dosage form: Tablets Treatment for: <u>Diabetes - Type 2</u>



Chap6Fig5





Chap6Fig6

#### What we need to know about Glipizide:

• Usual Dose: 10mg

#### Usual dose: 10 mg

• We can give it once a day or we can give it in divided doses if the patient is consuming more than 20 mg.

Once a day or in divided doses if more than 20 mg

• Maximum Dose: **20mg** 

#### **Maximum Dose: 20mg**

### Trade Name- Dose- Cost in USA- Cost in India

# GLIPIZIDE

Please always combine any blood sugar medicine with diet and exercise.

#### All medicines continue life-long

Generic Name & Trade Name	About Glipizide	Cost of Glipizide for one month (in USA)	Cost of Glipizide for one month (in India)
Generic Name: Glipizide Trade Name: Glucotrol (USA) Glucotrol XL (USA) Diacon (India) Dibizide (India)	<ul> <li>Usual dose: 5 mg twice day</li> <li>Can be increased to 20 mg</li> <li>As a Golden Rule – before starting any medicine, liver functions and kidney functions should be screened.</li> </ul>	Dose = Once a day (10 mg) 1-month = 60 tablets Cost = \$228	Dose = Once a day (10 mg) 1-month = 60 tablets Cost = Rs 85
		*All costs are meant for a rough estimate of one-month expense.	*All costs are meant for a rough estimate of one-month expense.



Chap6Fig7

#### **Gliclazide**

- I. All diabetes medicines should be taken before meals as a golden rule.
- II. We start on a low dose and increase the dose if needed
- III. We should usually take medication at the same time every day.

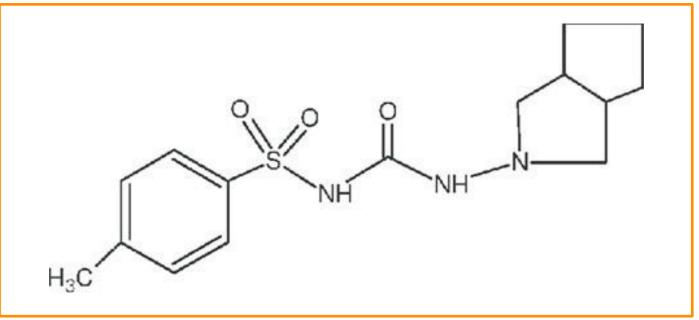
#### **Gliclazide FDA Approval History**

FDA Approved: Yes (Approved In 1972) Brand name: Diamicron (USA) Bilxona (USA) Vamju (India) Zicron (India) Generic name: Gliclazide Dosage form: Tablets Treatment for: <u>Diabetes - Type 2</u>



#### <u>Gliclazide</u>

Chap7Fig8



Chap7Fig9

#### What we need to know about Gliclazide:

• Usual Dose: 40mg

#### Usual dose: 40mg

• We can give it **once a day or twice a day** 

#### Given once or twice a day

• Maximum Dose: **320mg** 

**Maximum Dose: 320mg** 

### Trade Name- Dose- Cost in USA- Cost in India

# GLICLAZIDE

Please always combine any blood sugar medicine with diet and exercise

#### All medicines continue life-long

Generic Name & Trade Name	About Gliclazide	Cost of Gliclazide for one month (in USA)	Cost of Gliclazide for one month (in India)
Generic Name: Gliclazide <u>Trade Name:</u> Diamicron (USA) Bilxona (USA) Vamju (India) Zicron (India)	<ul> <li>Usual dose: 40 mg twice a day</li> <li>Can be increased to 320 mg once a day</li> <li>As a Golden Rule - before starting any medicine, liver functions and kidney functions should be screened.</li> </ul>	Not available in USA	Dose= 40 mg 1month = 30 tablets Cost= Rs 210
		*All costs are meant for a rough estimate of one-month expense.	*All costs are meant for a rough estimate of one-month expense



Chap6Fig10

#### **Glimepiride**

- I. All diabetes medicines should be taken before meals as a golden rule.
- II. We start on a low dose and increase the dose if needed
- III. We should usually take medication at the same time every day.

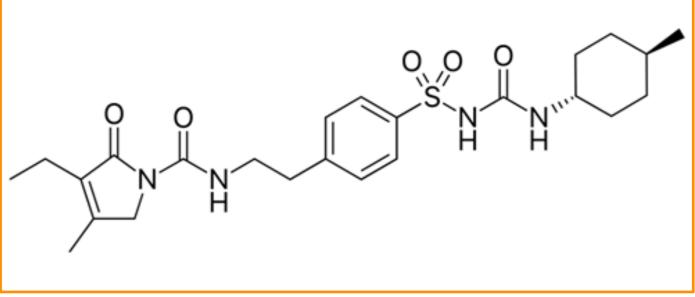
#### **Glimepiride FDA Approval History**

FDA Approved: Yes (Approved In Febuary, 1999) Brand name: Amaryl(USA), Dexasone(USA) Decdan (India), Dexona (India) Generic name: Glimepiride Dosage form: Tablets Treatment for: <u>Diabetes - Type 2</u>



#### <u>Glimepiride</u>

Chap6Fig11



Chap6Fig12

#### What we need to know about Glimepiride:

• Usual Dose: 2-4 mg

#### Usual dose: 2 mg

• We can give it once a day or we can give it in divided doses if the patient is consuming till 8 mg

Once a day or in divided doses if more than 8 mg

• Maximum Dose: **8 mg** 

#### Maximum Dose: 8 mg

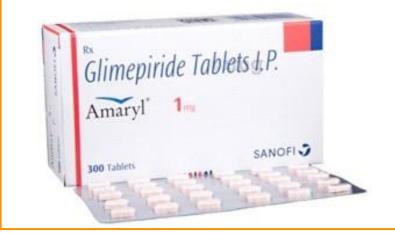
### Trade Name- Dose- Cost in USA- Cost in India

# GLIMEPIRIDE

Please always combine any blood sugar medicine with diet and exercise.

#### All medicines continue life-long

Generic Name & Trade Name	About Glimepiride	Cost of Glimepiride for one month (in USA)	Cost of Glimepiride for one month (in India)
Generic Name: Glimepiride <u>Trade Name:</u> Amaryl(USA) Dexasone(USA) Decdan (India) Dexona (India)	<ul> <li>Usual dose: 1 mg twice a day</li> <li>Can be increased to 8 mg</li> <li>As a Golden Rule – before starting any medicine, liver functions and kidney functions should be screened.</li> </ul>	Dose = Once a day (2mg) <u>1-month</u> = 60 tablets <u>Cost</u> = \$66	Dose = Once a day (2mg) <u>1-month</u> = 60 tablets <u>Cost</u> = Rs 660
		*All costs are meant for a rough estimate of one-month expense.	*All costs are meant for a rough estimate of one-month expense.



Chap6Fig13