# Book-3.1

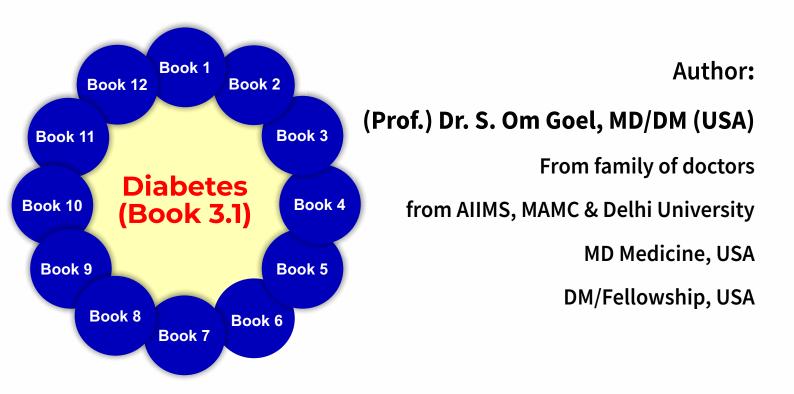


ADD 15 YEARS TO OUR LIFE CAN WE? OF COURSE, WE CAN!!

# Add 15 Years Diabetes - Book 3.1 SULFONYLUREAS (1<sup>st</sup> generation medicine)

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

# USA/India Edition 2021 | ENGLISH





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

These "Medical Books" can really help in Adding "15 Healthier Years"

# to Our Lifespan!

<u>"The recommended age is 13yrs to 80yrs</u>

<u>The sooner you start, more</u> <u>years you add to lifespan."</u>



# PREFACE

#### Think About It!

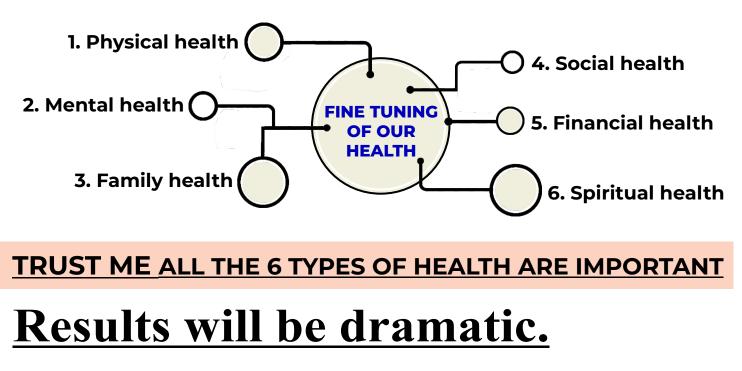
Miracles do not happen in real life.

At least in India (NOT very common in U.S.) we seek medical care only when we have so called "alarming symptoms" and MD/Physicians provide care with their focus on these "alarming symptoms". We call it "Crisis Medicine".

Personally speaking, in 2020/2021 and in the coming years, these "alarming symptoms" (called "Crisis Medicine") should be RARE before we get into the 75 to 85 years of age group.

### PROVIDED

We keep **fine tuning our health** (medical definition) and gain insight about fundamental medical knowledge, and we actively plan and pay attention to our:



# DISCLAIMER

In all the books that I am writing as a physician and a MD with a lifetime of experience (along with my team)-

I have mentioned the most common medical facts which each one of us need to know in our day to day life.

Ignorance is never a bliss and 2020 has shown us that.

I highly recommend that anybody entering middle school (8th grade i.e. a 13-year-old) should start reading these books and try to read as many as he or she can <u>so that they can get an</u> <u>insight into the most common medical facts.</u>

These books are written in basic English and several other languages as well.

If we get an insight into medical facts before we get into chain-smoking and excessive alcohol use or another drug abuse, then ultimately, we can live a healthy and a long life.

All my research and common sense says that starting at the age of 13 years, (that is when we enter our teenage years, **our personality**, **our habits**, **our likings and our disliking -- all are pretty much shaped by the time we get to 26 years old or older**.

#### Both India and USA are very dear to me,

In the former I was born; I have my parents. In the latter, I have my wife and my children.

One thing became obvious to me as I live in USA, that the population is taking advantage of the latest medical advances.

Our Indian community, all our friends and families, when we have medical issues, are really putting themselves at the mercies of government and private hospitals, and doctors (with "zero" trust).

In India, people have to spend their own money. Realizing this, I have provided all the medical information, which is available to us doctors, so that one can make wise choices and confidently take their health in their own hands.

But still I sincerely request that you should not take any medications without the supervision of your own family doctors.



The facts that I have provided in my books are available in every other medical book, but I them in very simple English or in your language **because how are you going to make right choices in relation to your health, if you do not know what the answer is.** 

Our horoscope/stars are not enough, and consulting a pandit, priest or maulana is never enough.

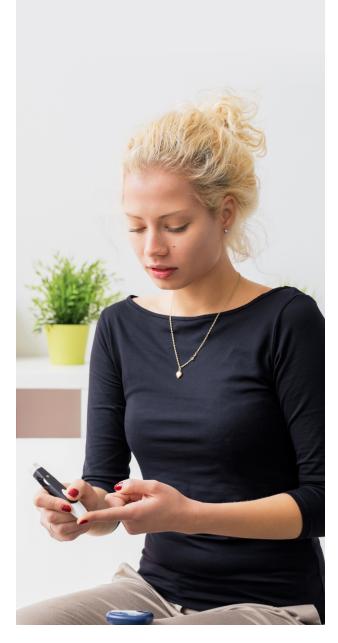
Blind faith is never an option.

Prayers help, yes! when we do not know the answer to a situation.

If our car runs out of petrol, then eventually it will not move; does not matter how much we may pray, it is never going to happen unless we fill the tank.

If you anticipate and choose wisely, you will not have a crisis. So, our disclaimer is that we give you the insights but please always consult your physicians before starting any prescription medications.





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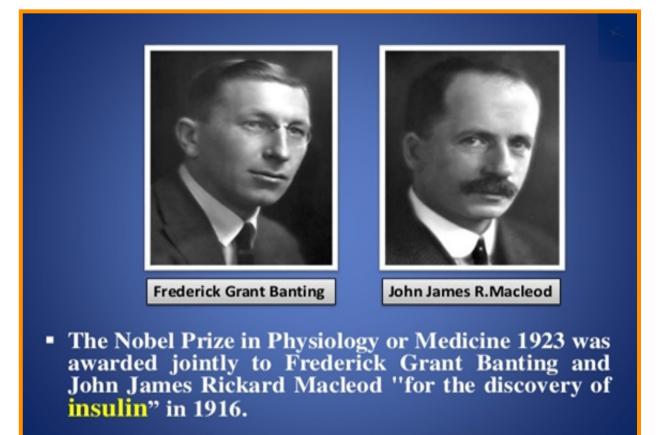
7 First Generation Sulfonylureas



**CAN WE? OF COURSE, WE CAN!!** 

# Referring Diabetes - Book 1 **HIGH BLOOD SUGAR** IN 2021: No reason for kidneys to fail!! Before 1921: Yes, You would have died. (year of Nobel Prize for Insulin)

# Introduction



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!

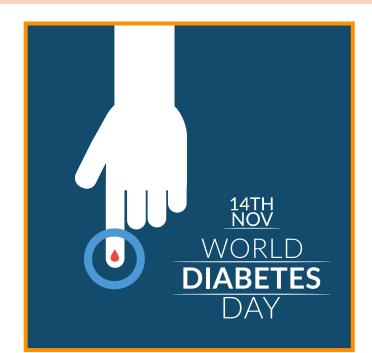
# Before 1921

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!!

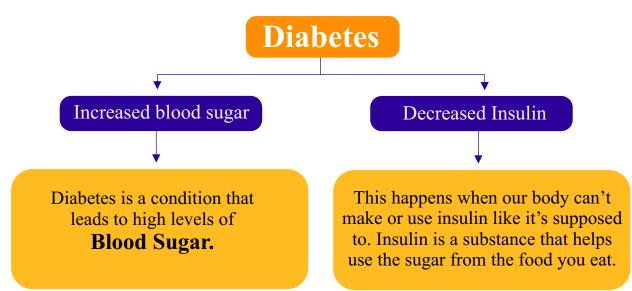


But yes, we have 12 to 13 different groups of medicines for high blood sugar.

<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

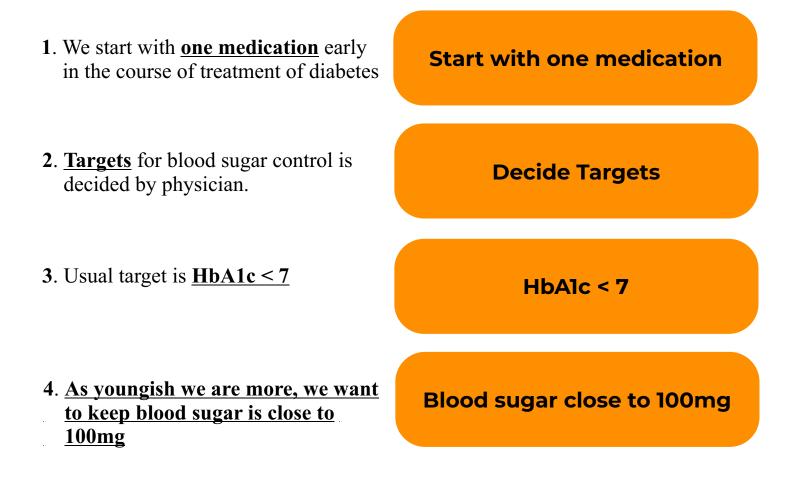


# The Golden Principle in Management of high blood sugar (Diabetes)

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!



#### 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> <u>by different mechanisms of action.</u>
- **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.
   <u>We can add a second medicine.</u>

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

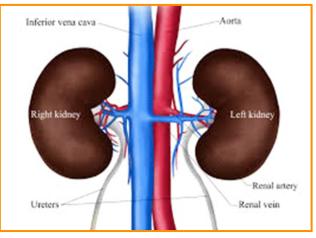


Chap3Fig2

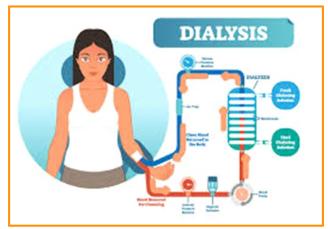


Chap3Fig3

• And HbA1c < 7



Chap3Fig4



Chap3Fig5



Chap3Fig7

## • To prevent kidney failure

• And no dialysis

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



Chap3Fig8



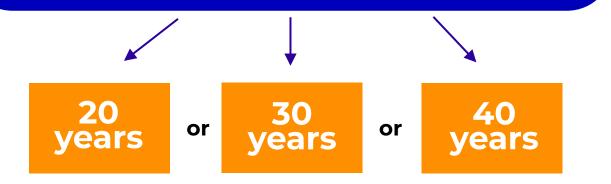
Chap3Fig9

• 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



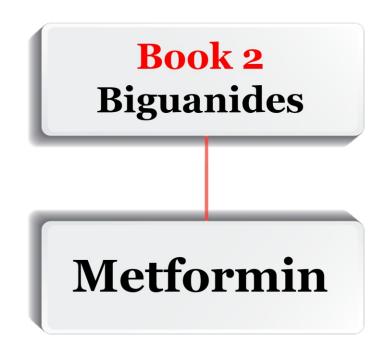
# Is it true? ✓Really Yes! it is True

# Today in 2020/2021 we have excellent medicines available

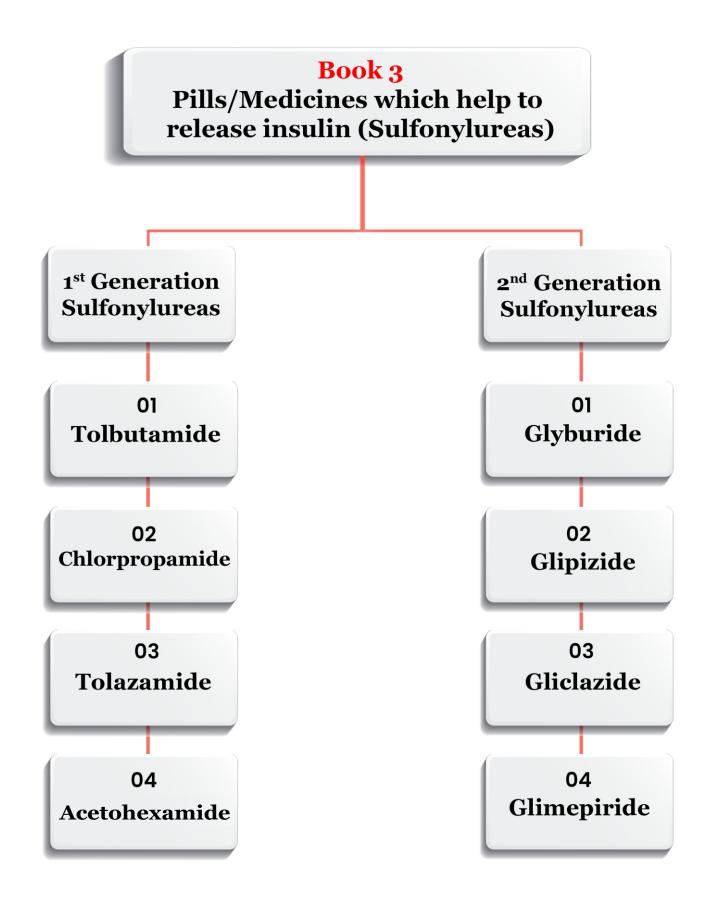
There are a number of medicines available in the market today for the treatment of Diabetes. The main groups which these medicines belong to are:

### **Book-2** Biguanides

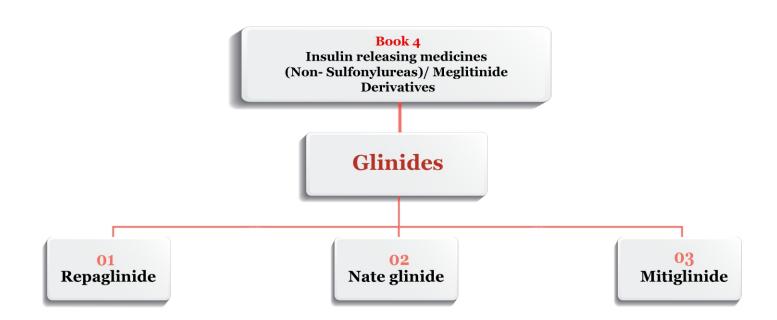
**Metformin:** The most famous and most used medicine and 1st medicine to start the treatment of diabetes.



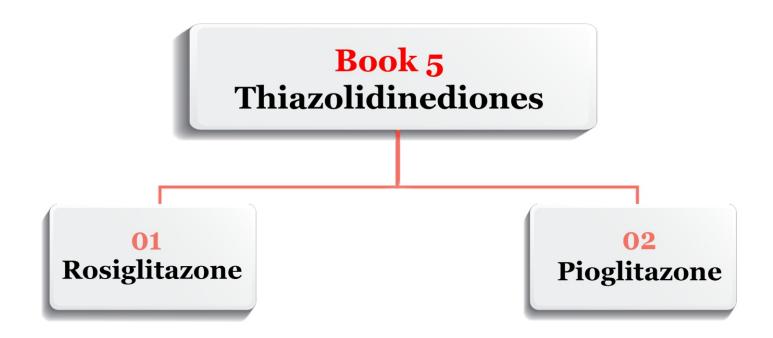
## **Book-3** Pills/Medicines which help to insulin (Sulfonylureas)



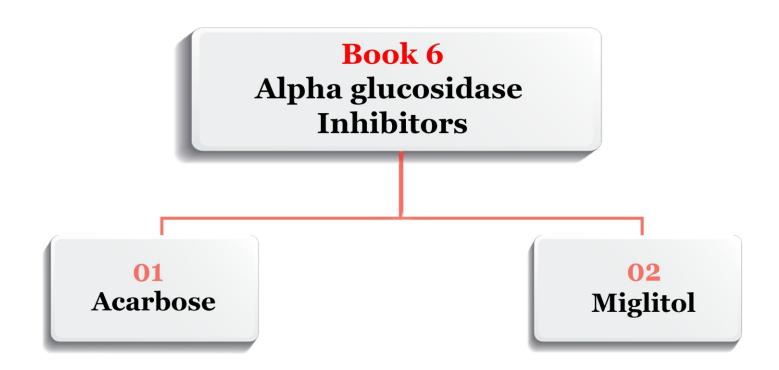
**Book-4** Insulin releasing medicines (Non- Sulfonylureas)



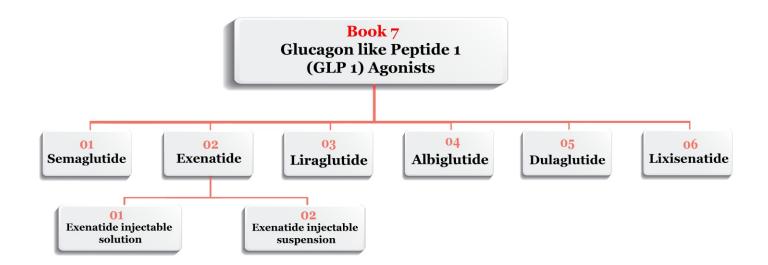
**Book-5** Thiazolidinediones



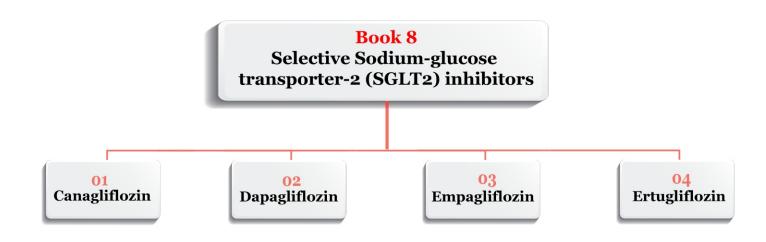
**Book-6** Alpha glucosidase Inhibitors



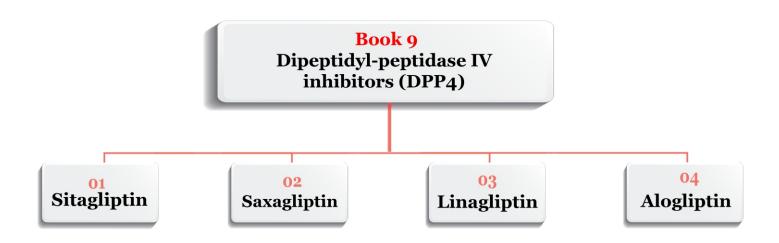
Book-7 Glucagon like Peptide 1 (GLP 1) Agnostics



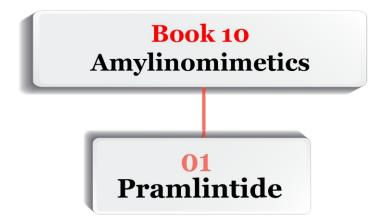
# **Book-8** Selective Sodium-glucose transporter-2 (SGLT2) inhibitors



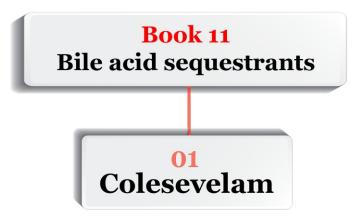
**Book-9** Dipeptidyl-peptidase IV inhibitors (DPP4)



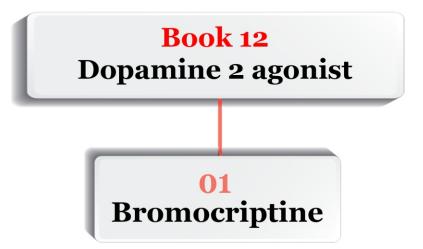
**Book-10** Amylinomimetic group



Book-11 Bile acid sequestrants



Book-12 Dopamine 2 agonist



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

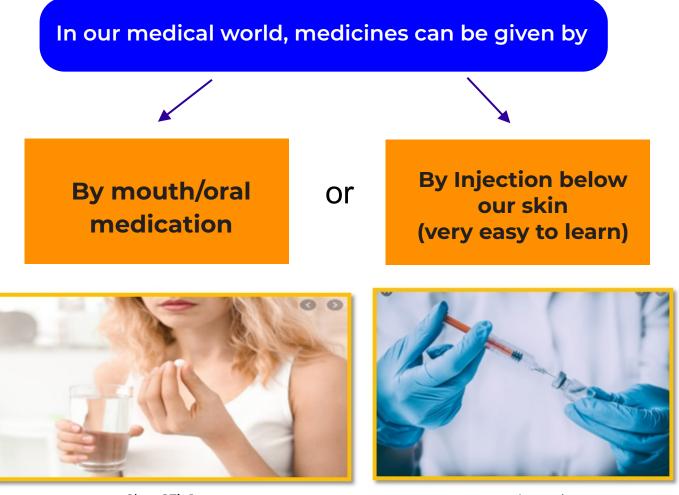
#### Think About It!

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.



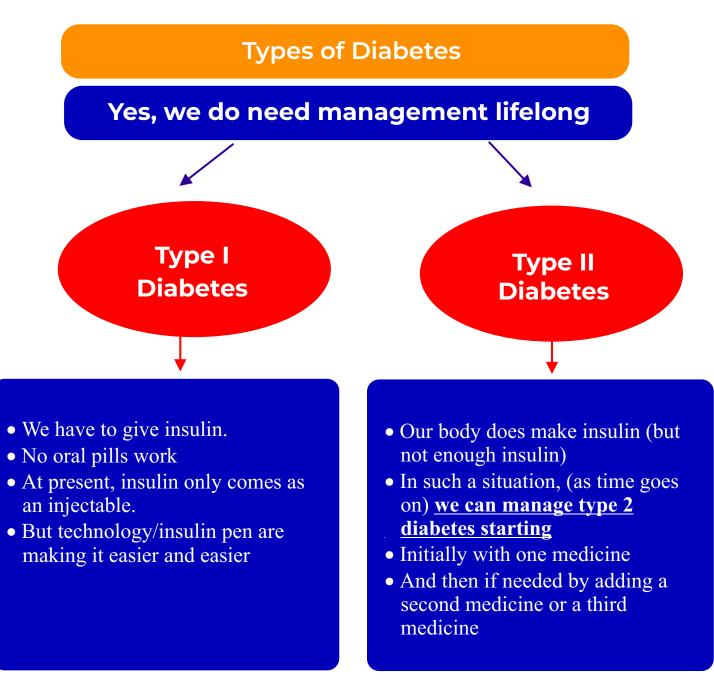
Chap6Fig1

Chap6Fig2

# Type 1 vs Type 2 Diabetes

#### Think About It!

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.



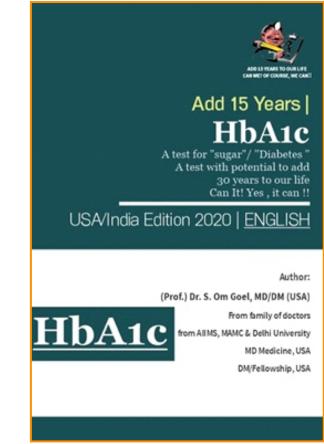
# How do we decide less dose or more dose of medicines

#### 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

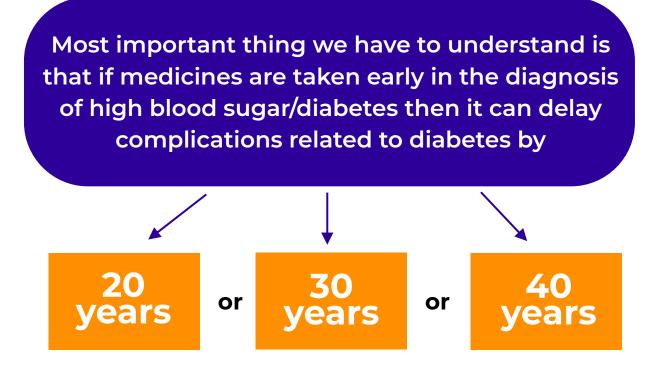
# Early diagnosis and early treatment: an absolute necessity in 2021

#### Think About It!

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

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.





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

# Diabetes - Book 3.1 SULFONYLUREAS (1<sup>st</sup> generation medicine) IN 2021: No reason for kidneys to fail!!

**Before 1921**: 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?

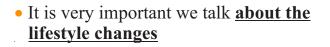
1. They <b>work by directly stimulating the</b> <b>release of insulin</b> from beta cells of Langerhans which lie in the pancreas	Release Insulin directly in Pancreases.
<ol> <li>You can take these medicines with food or before food. They lead to the lowering of blood glucose level indirectly by stimulating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

exercising

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 Blindness

**Because Diabetes is so common** 



Chap3Fig2



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



Chap3Fig3

# Diabetes #1 cause of Loss of Limbs

**Because Diabetes is so common** 



Chap3Fig4



#### **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

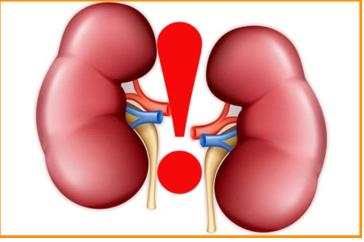
• If patients consume <u>too much alcohol then</u> <u>it leads to poor nutrition status</u> and affects the liver.

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 <u>kidney failure actually need less insulin</u> (as insulin stays longer in the blood.)



Chap4Fig3

#### • Some medication can cause drug

**interaction** 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

### **Chapter 5**

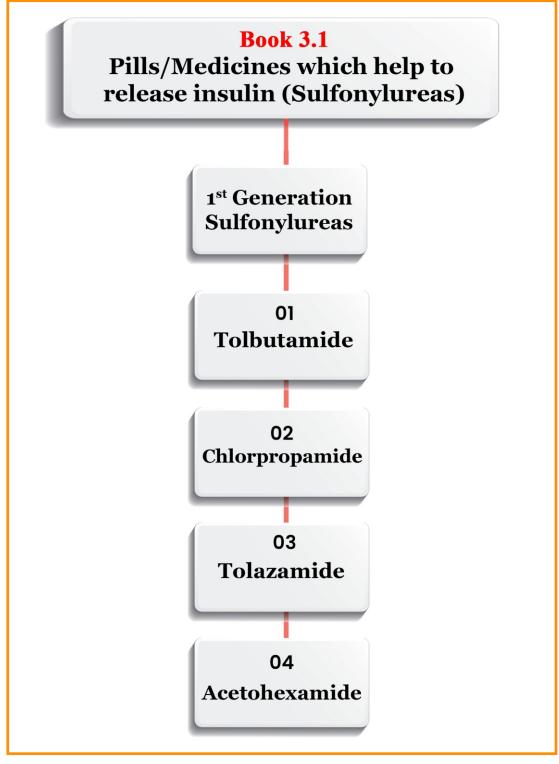
Introduction to First and Second Generation Sulfonylureas

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



### **Chapter 6**

### **First Generation Sulfonylureas**



Chap6Fig1

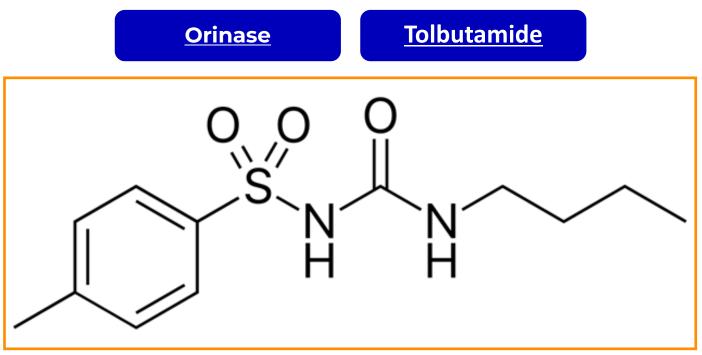
### <u>Orinase</u> Generic Name: Tolbutamide

- 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.

### **Orinase FDA Approval History**

FDA Approved: Yes (First approved May, 1957) Brand name: Orinase Generic name: Tolbutamide Dosage form: Tablets Treatment for: <u>Diabetes - Type 2</u>







#### What we need to know about Tolbutamide (Orinase):

• It is a first-generation **sugar lowering potassium channel blocker**, sulfonylurea oral hypoglycemic medication

Eirct	generation	CUGOR	lowering	modicino
FIISL	generation	Sugar	lowening	medicine

• This drug may be used in the **management of type 2 diabetes** if diet alone is not effective

### Manages Type 2 diabetes

• It stimulates the **secretion of insulin by the pancreas** and lowers blood sugar levels while helping the body use Insulin efficiently

Stimulates secretion of insulin to lower blood sugar

• It is used along with **diet and exercise** 

### Used with diet and exercise

• Available as **500 mg pill**, Daily Dose: **1gm-2gm once a day** 

Usual dose- 1-2gm a day

• Maximum dose: Usually more than **2** gm is not needed

### Maximum Dose: 2gm a day

• Tolbutamide comes as a tablet to **take by mouth** 

### Taken by mouth

• It is usually taken once a day in the morning

### Taken once a day

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

## ORINASE (TOLBUTAMIDE)

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

### All medicines continue life-long

Generic Name & Trade Name	About Orinase	Cost of Orinase for one month (in USA)	Cost of Orinase for one month (in India)
Generic Name: Tolbutamide <u>Trade Name:</u>	<ul> <li>Usual dose: 1000mg daily</li> <li>Can be increased 2000mg</li> <li>As a Golden Rule –</li> </ul>	Dose = 2 times daily (500mg) 1-month = 92 tablets	Dose = 2 times daily (500mg) 1-month = 92 tablets
Orinase	As a Golden Rule – before starting any medicine, liver functions and kidney functions should be screened.	<u>Cost</u> = \$85	<u>Cost</u> = Rs 65
		*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

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### <u>Diabinese</u>

### Generic Name: Chloropropamide

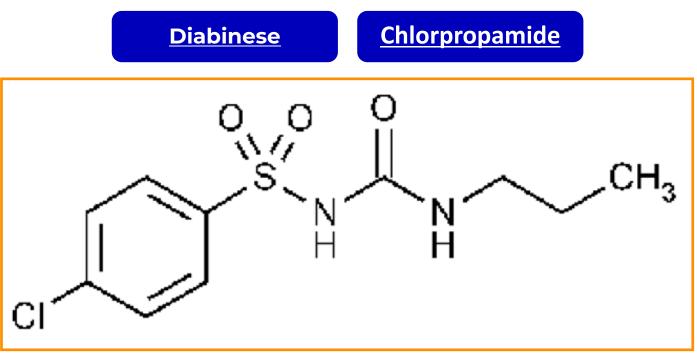
- I. All diabetes medicines should be taken before meals as a golden rule.
- II. We start on a low dose and increase the dose if needed
- III. We should usually take medication at the same time every day.

### **Diabinese FDA Approval History**

FDA Approved: Yes (First approved October, 1958) Brand name: Diabinese Generic name: Chlorpropamide Dosage form: Tablets Treatment for: <u>Diabetes - Type 2</u>



Chap6Fig5



#### What we need to know about Chlorpropamide (Diabinese):

• It is a first-generation **sugar lowering potassium channel blocker**, sulfonylurea oral hypoglycemic medication

First	generation	sugar	lowering	medicine
FIISC	generation	Sugai	lowering	medicine

• This drug may be used in the **management of type 2 diabetes** if diet alone is not effective

### Manages Type 2 diabetes

• It stimulates the **secretion of insulin by the pancreas** and lowers blood sugar levels while helping the body use Insulin efficiently

Stimulates secretion of insulin to lower blood sugar

• It is used along with **diet and exercise** 

### Used with diet and exercise

• Available as **100 mg pill**, Daily Dose: **250gm once a day** 

#### Usual dose- 250gm a day

• Maximum dose: Usually more than **250 gm is not needed** 

### Maximum Dose: 250gm a day

• Tolbutamide comes as a tablet to **take by mouth** 

### Taken by mouth

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

# DIABNESE

### (CHLORPROPAMIDE)

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

### All medicines continue life-long

Generic Name & Trade Name	About Diabnese	Cost of Diabnese for one month (in USA)	Cost of Diabnese for one month (in India)
Generic Name: Chlorpropamide <u>Trade Name:</u> Diabnese	<ul> <li>Usual dose: 250mg once daily</li> <li>Can be increased 250mg</li> <li>As a Golden Rule – before starting any medicine, liver functions and kidney functions should be screened.</li> </ul>	Dose = 250mg once a day 1-month = 75 tablets Cost = \$69	Dose = 250mg once a day 1-month = 75 tablets Cost = Rs 25
		*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

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### <u>Tolinase</u>

### Generic Name: Tolazamide

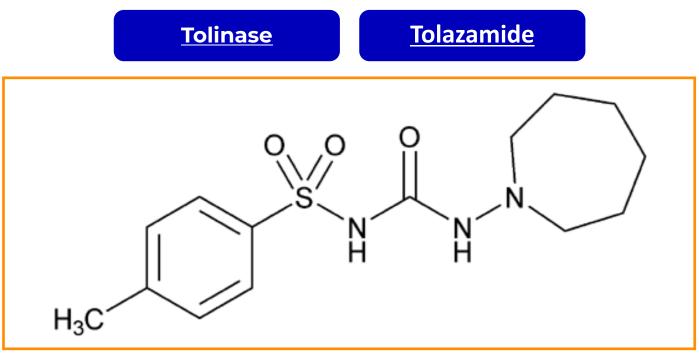
- I. All diabetes medicines should be taken before meals as a golden rule.
- II. We start on a low dose and increase the dose if needed
- III. We should usually take medication at the same time every day.

### **Tolinase FDA Approval History**

FDA Approved: Yes (First approved July, 1966) Brand name: Tolinase Generic name: Tolazamide Dosage form: Tablets Treatment for: <u>Diabetes - Type 2</u>



Chap6Fig8



Chap6Fig9

#### What we need to know about Tolazamide (Tolinase):

• It is a first-generation **sugar lowering potassium channel blocker**, sulfonylurea oral hypoglycemic medication

First	generation	sugar	lowering	medicine
FIISC	generation	Sugai	lowering	medicine

• This drug may be used in the **management of type 2 diabetes** if diet alone is not effective

### Manages Type 2 diabetes

• It stimulates the **secretion of insulin by the pancreas** and lowers blood sugar levels while helping the body use Insulin efficiently

Stimulates secretion of insulin to lower blood sugar

• It is used along with **diet and exercise** 

### Used with diet and exercise

• Available as 250 mg pill, Daily Dose: 100gm once a day

#### Usual dose- 100gm a day

• Maximum dose: Usually more than **250gm is not needed** 

### Maximum Dose: 250gm a day

• Tolbutamide comes as a tablet to **take by mouth** 

### Taken by mouth

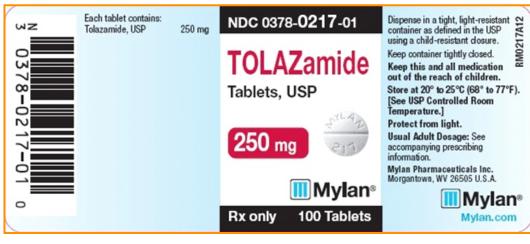
### Trade Name- Dose- Cost in USA- Cost in India

# TOLINASE (TOLAZAMIDE)

Please always combine any blood sugar medicine with diet and exercise

### All medicines continue life-long

Generic Name & Trade Name	About Tolinase	Cost of Tolinase for one month (in USA)	Cost of Tolinase for one month (in India)
<b>Generic Name:</b> Tolazamide <b>Trade Name:</b> Tolinase	<ul> <li>Usual dose: 100 mg once a day</li> <li>Can be increased to 250 mg once a day</li> <li>As a Golden Rule – before starting any medicine, liver functions and kidney functions should be screened.</li> </ul>	Dose = 100 mg 1 month = 15 tablets Cost = \$9 *All costs are meant for a rough estimate of one-month expense.	Dose = 100 mg 1 month = 15 tablets Cost = Rs 2175 *All costs are meant for a rough estimate of one-month expense



Chap6Fig10

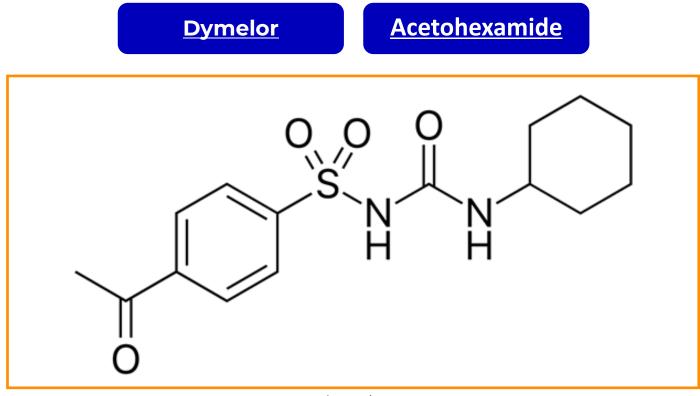
### <u>Dymelor</u>

### **Generic Name: Acetohexamide**

- 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.

### **Dymelor FDA Approval History**

FDA Approved: No longer in use Brand name: Dymelor Generic name: Acetohexamide Dosage form: Tablets Treatment for: <u>Diabetes - Type 2</u>



Chap6Fig11

#### What we need to know about Acetohexamide (Dymelor):

• It is a first-generation **sugar lowering potassium channel blocker**, sulfonylurea oral hypoglycemic medication

-			• • • • •	
First	generation	sudar	lowering	medicine
	3011010101			

• This drug may be used in the **management of type 2 diabetes** if diet alone is not effective

### Manages Type 2 diabetes

• It stimulates the **secretion of insulin by the pancreas** and lowers blood sugar levels while helping the body use Insulin efficiently

Stimulates secretion of insulin to lower blood sugar

• It is used along with **diet and exercise** 

### Used with diet and exercise

• Not used anymore as better medicines are available in the market

### Not used anymore

• Acetohexamide comes as a tablet to **take by mouth** 

### Taken by mouth

# This medicine is no longer used as better medicines are available in the market.