

King Saud University College of Medicine 1<sup>st</sup> Year, 2<sup>nd</sup> block

# Diseases Modifying Anti-Rheumatic Drugs



# Musculoskeletal Block

### **Objectives**

### Define DMARDs

Describe the classification of this group of drugs

Describe the general advantages & criteria of this group of drugs List some examples of drugs related to DMARDS.

Describe the mechanism of action , specific clinical uses , adverse effects & contraindications of individual drugs.

Describe the general clinical uses

Used when the disease is progressing & causing deformities

Used when the inflammatory disease is not responding to NSAIDs Low doses are commonly used early in the course of the disease

#### Have no analgesic

**effects** (not directly) but they reduce pain because they are anti-inflammatory (when inflammation is reduced the pain is reduced too)

General Features & Conditions to use Antirheumatic

**General Clinical** 

Uses

Slow onset their effects take from 6 weeks up to 6 months to be evident

Can not repair the existing damage( not treating ), but prevent further deformity

Treatment of rheumatic disorders Combination therapies are both safe & efficacious

#### Types of Anti-rheumatic drugs

# Non Biologic disease modifiers(Old drugs)

#### **Biologic disease modifiers**

Genetically engineered drugs that are used to modify imbalances of the immune system in autoimmune diseases.(they make these drugs in the laboratories by genetic modification )

What are they ? Types Or blocking certain messenger agents block, or modify the proteins known as cytokines, activity of selected cells in that send signals between the immune system( **Directly** those cells.(block affecting the cell communication between the cells) **T-cell modulating** Anti-IL-6 receptor Abatacept Tocilizumab antibody drug **TNF-**blocking **B-cell cytotoxic** Rituximab Infliximab agents agent

#### FIRST :NON BIOLOGIC DISEASE MODIFIERS(OLD DRUGS)

Name of the drug	Mechanism of action	Pharmacokinetics and features	Adverse Effects
Hydroxychloroquin e Note: when we give this drug we do vision examinations to avoid side effects .	<ul> <li>Trapping free radicals(anti oxidant).</li> <li>Suppression of T lymphocyte cells.</li> <li>Stabilization of lysosomal enzyme activity.</li> </ul>	<ul> <li>■ Rapidly &amp; completely absorbed following oral administration.</li> <li>■ Penetrates into C.N.S. &amp; traverse the placenta ( that's why this drug isn't given to pregnant women)</li> <li>■ Metabolized → liver</li> </ul>	<ul> <li>Pruritus</li> <li>GIT upset (Nausea &amp; vomiting)</li> <li>Discoloration of nail beds &amp; mucous membranes</li> <li>Headaches</li> <li>Blurred vision (Corneal deposits)</li> <li>Irreversible retinal damage ( the most dangerous side affect)</li> </ul>
Methotrexate (anti cancer drug , inhibit the growth of cancer cells )	<ul> <li>Inhibition of T- Cells (cell- mediated immune reactions)</li> <li>Inhibition of polymorpho- nuclear chemotaxis</li> </ul>	<ul> <li>Immunosuppressant drug</li> <li>Used mainly as chemotherapy for cancer treatment</li> <li>The doses needed as anti- rheumatic are much lower than those needed in cancer chemotherapy</li> <li>Given once a week</li> </ul>	<ul> <li>Nausea</li> <li>Liver cirrhosis (ONLY with chronic administration of the drug) &amp; Hepatotoxicity</li> <li>Acute pneumonia –like syndrome (result from Cytopenia)</li> <li>Mucosal ulceration.(GIT)</li> <li>bone marrow depression         <ul> <li>Cytopenia.(reduction in the number of BC)</li> <li>(Mucosal ulceration and Cytopenia are the most dangerous side effects)</li> </ul> </li> </ul>

#### SECONDS: BIOLOGIC DISEASE MODIFIERS(NEW DRUGS)



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#### SECONDS: BIOLOGIC DISEASE MODIFIERS (NEW DRUGS)



# COMPARISON BETWEEN NSAIDS & DMARDS

DMARDs	NSAIDs
Slow onset of action used in chronic cases when deformity is exciting	Rapid onset of action used in acute cases to relief inflammation & pain
Arrest progression of the disease	No effect
Prevent formation of new deformity	Can not stop formation of new deformity

## SUMMARY

Define DMARDs	Describe the general clinical uses	Describe the classification of this group of drugs	Describe the general advantages & criteria of this group of drugs
Disease –Modifying Antirheumatic drugs Used when the disease is progressing & causing deformities	Treatment of rheumatoid arthritis	Non Biologic disease modifiers(Old drugs)	<ul> <li>used in chronic cases when deformity is exciting</li> <li>Arrest progression</li> <li>Prevent formation of new</li> </ul>
prevent further deformity and have no analgesic effect	Combination therapies	Biologic disease modifiers	deformity - Have no analgesic effects - Slow onset

#### Know some examples of drugs related to DMARDS.

OLD drugs: Hydroxychloroquine Methotrexate

New drugs: Tocilizumab

## Describe the mechanism of action , specific clinical uses , adverse effects &contraindications of individual drugs.

- Hydroxychloroquine acts mainly through suppression of the activity of lysosomal enzymes and trapping free radicals
- Its main adverse effects is irreversible retinal damage & hepatic toxicity.

- Methotrexate acts mainly through suppression of phagocytic cells & T cells- Its adverse effects are bone marrow depression & mucosal ulceration

- Infliximab is a chimeric TNF- $\alpha$  blocking agent
- Given with methotrexate to reduce antichimeric effect



1- Which drug is used with special precautions to guard against the development of irreversible retinopathy:

- a) Methotrexate
- B) Infliximab
- C) Hydroxychloroquine
- d) rituximab

2-inhibition of polymorphonuclear chemotaxin is the mechanism of action in which DMARDs

- a) methotrexate
- b) Hydroxychloroquine
- c) Infliximab.
- d) Tocilizumab

**3- Drug Its main adverse effects are upper respiratory tract infections & reactivation of latent TB:** 

- a) Infliximab
- b) Tocilizumab
- c) Methotrexate
- d) Hydroxychloroquine

4- which of the following Biologic disease modifiers that works Directly affecting the cell :

- a) rituximab
- b) Infliximab
- c) Abatacept
- d) Both a & c

5- which of the following DMARDs is used with high doses as anti cancer drug ?a) Hydroxychloroquine

- b) Methotrexate
- c) Tocilizumab
- d) Infliximab
- 6- Blood tests will be used monthly for

increase in cholesterol, liver enzymes & decrease in WBCs :

- a) Hydroxychloroquine
- b) Methotrexate
- c) Tocilizumab
- d) Infliximab

#### 1-C 'S-Y ' 3-Y't-D'2-B'6-C

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Nada Dammas Nada Bin Dawood Ahmad AlDakhil Faris Almoammarie

