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Name



Department of Animal Biology  
Faculty of Natural Sciences  
University of Tabriz

**M.Sc. thesis**

***Study of antioxidant effect of Lino–curcumin  
on spatial memory and oxidative stress parameters  
during Multiple sclerosis disease in male rats***

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**Advisor:**

**Dr. G. Dehghan**

**Researcher:**

**B. Barzegarzadeh**

**Feb 2017**

# Main Titles of Presentation

1

**Introduction**

2

**Materials & Methods**

3

**Results**

4

**Discussion**



# Main Titles of Presentation

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**Introduction**

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**Discussion**

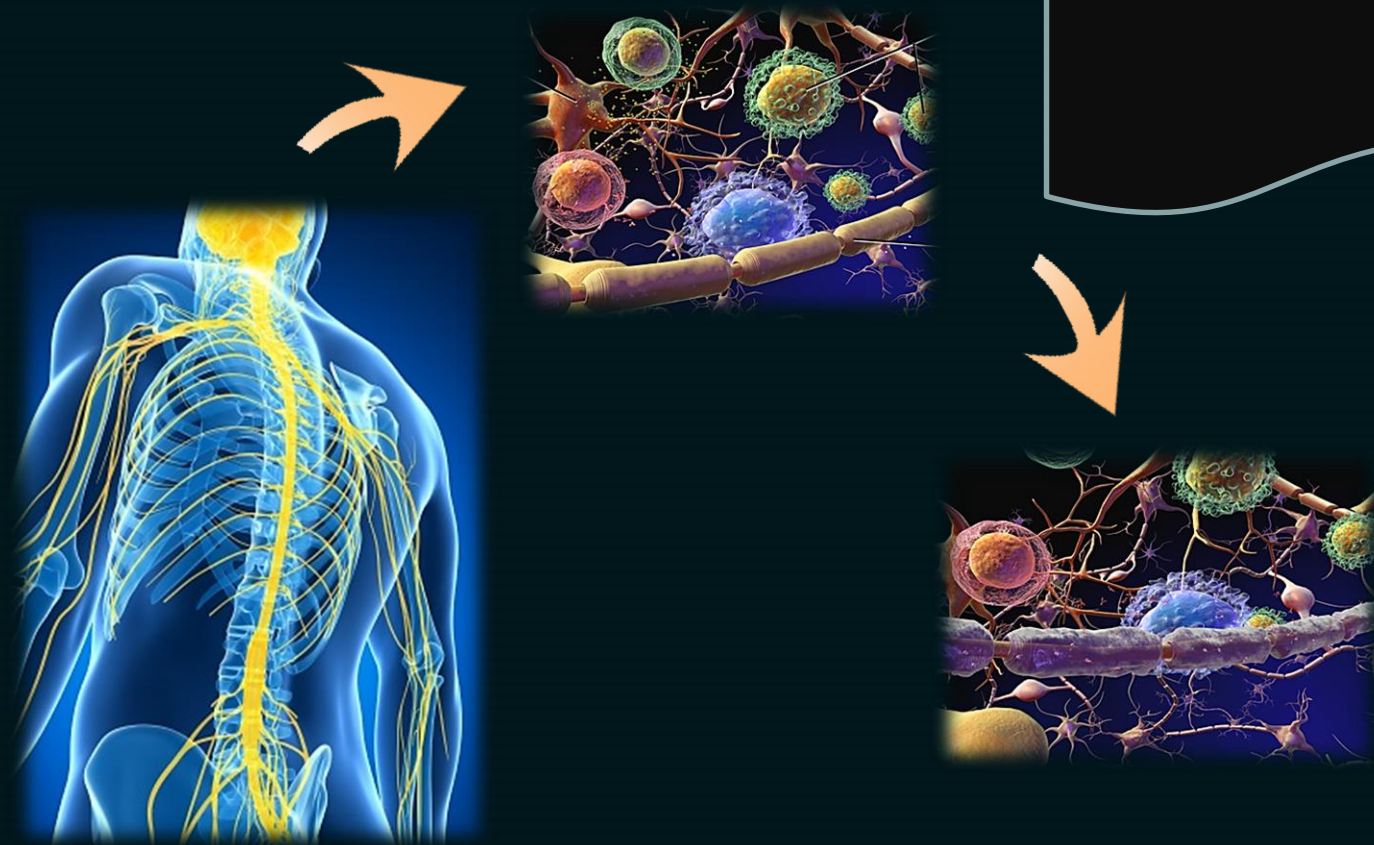


# INTRODUCTION



# Multiple Sclerosis Disease ( MS )

- Neurodegenerative
  - Autoimmune
  - Inflammatory
- 
- Demyelination
  - Neuroinflammation

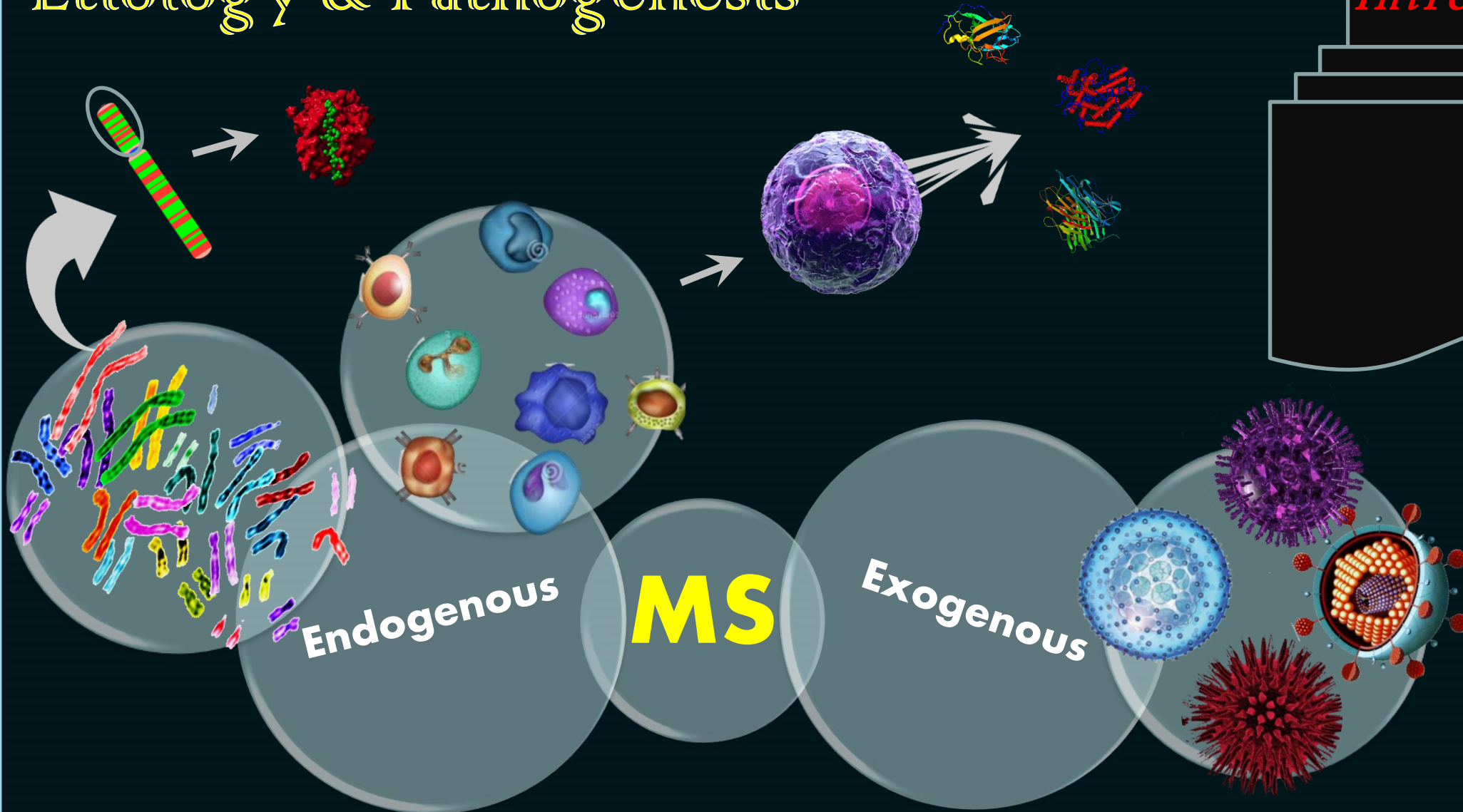


*Introduction*



# Etiology & Pathogenesis

*Introduction*

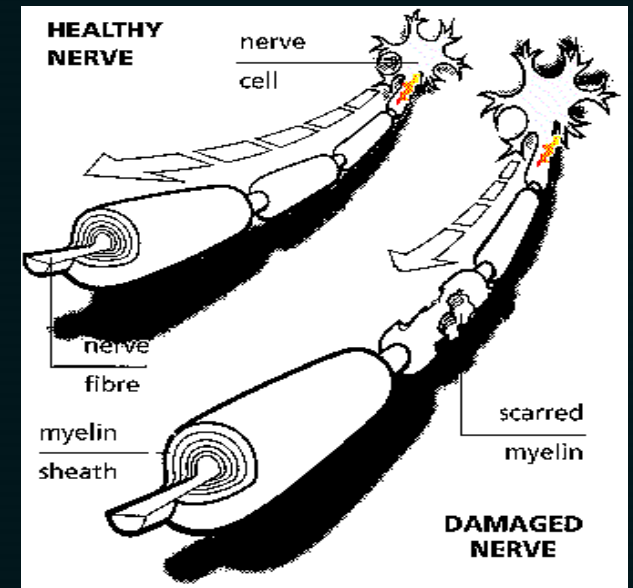
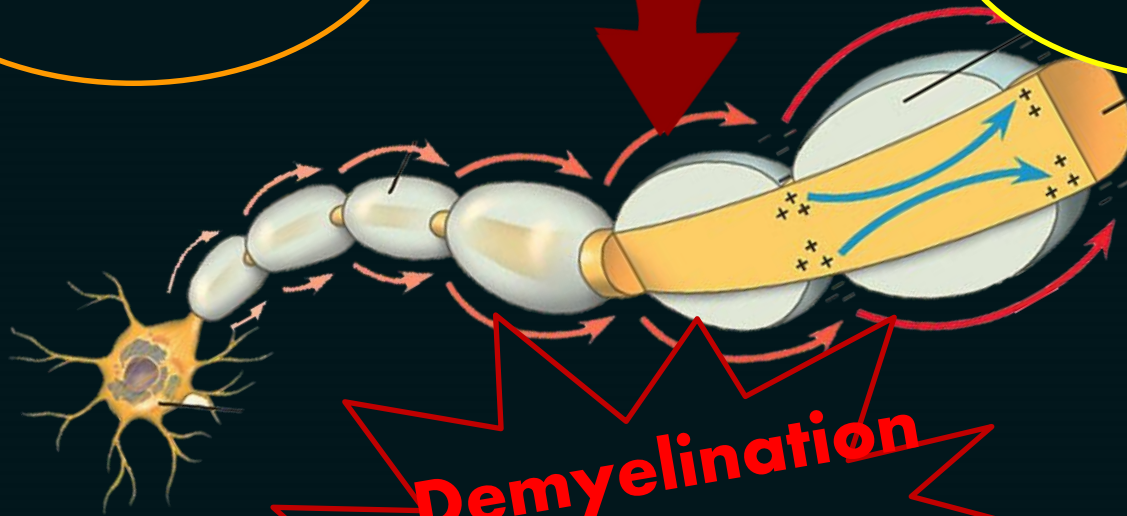


# Pathophysiology

*Introduction*

Inflammation

Oxidative Stress



# Symptoms

*Introduction*

## Central

- Depression
- Cognitive impairment
- Fatigue

## Visual

- Nystagmus
- Optic Neuritis
- Diplopia

## splanchnic

- Bladder dysfunction
- Bowel dysfunction

## Sensation

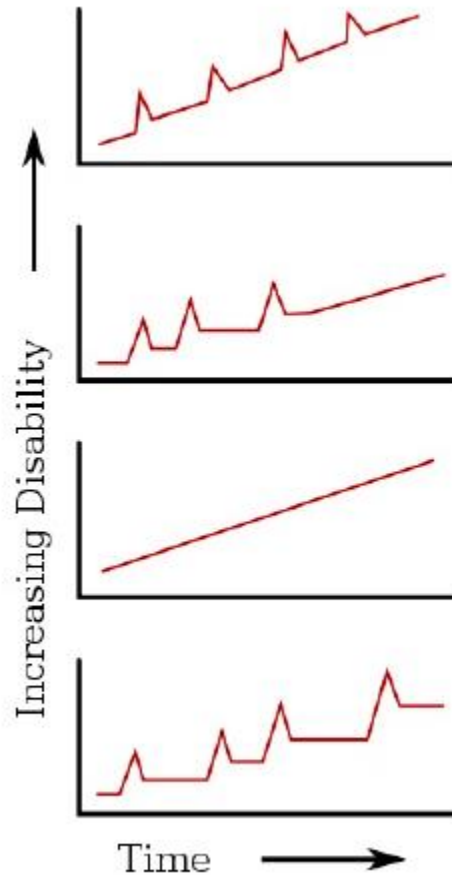
- Pain
- Numbness

## Musculoskeletal

- Weakness
- Ataxia

# Classification

*Introduction*



**PRMS** Progressive Relapsing MS  
Steady decline since onset with super-imposed attacks.

**SPMS** Secondary Progressive MS  
Initial RRMS that suddenly begins to decline without periods of remission and relapses.

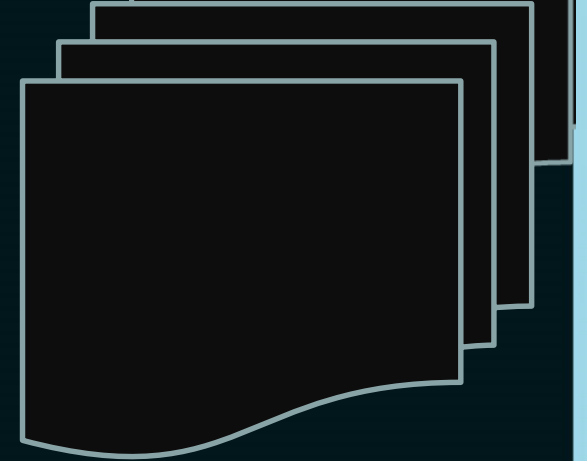
**PPMS** Primary Progressive MS  
Gradual progression of the disease from its onset with no relapses or remissions

**RRMS** Relapsing/ Remitting MS  
Unpredictable attacks which may or may not leave permanent deficits followed by periods of remission

# Necessity of MS Models

*Introduction*

- Common Reasons
- Biopsy Problems
- Complex Disease



# MS Models

- ✓ Myelin Mutants
- ✓ Viral & Autoimmune Models
- ✓ Chemically- induced Lesions

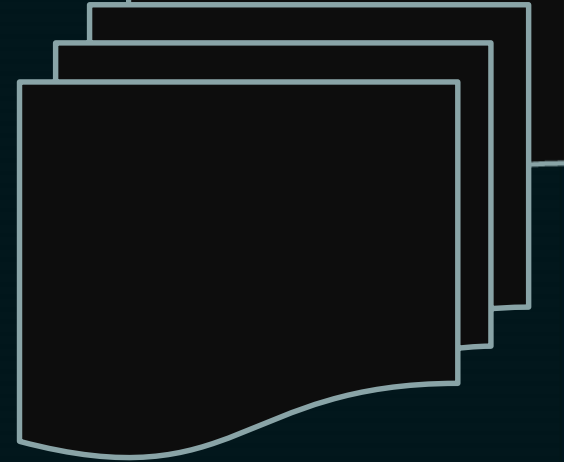


## Ethidium Bromide :

- Local Demyelination
- Oxidative Stress
- Neuroinflammation
- Astrocytes & Oligodendrocytes Damage



*Introduction*



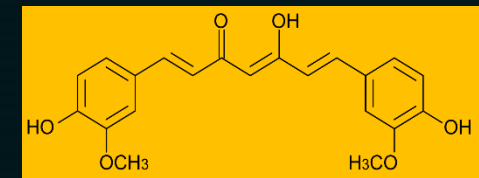
# Curcumin Origin

Class : Liliopsida  
Order : Zingiberales  
Family : Zingiberaceae  
Genus : Curcuma  
Species : C. longa (Turmeric)

Perennial plant

Used as home remedy  
for various diseases

*Introduction*

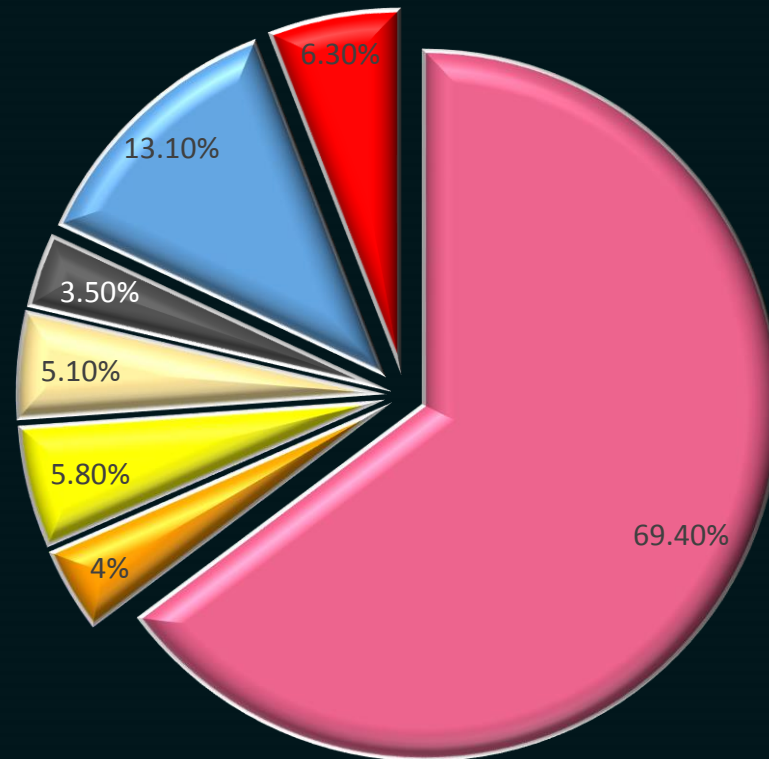


Curcumin is derived from  
*“Curcuma longa”*

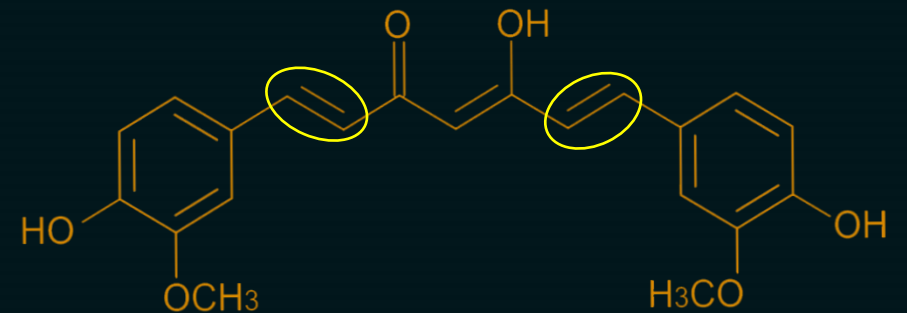
# Curcumin

## Introduction

- carbohydrates
- curcumin
- essential oil
- fat
- minerals
- moisture
- protein

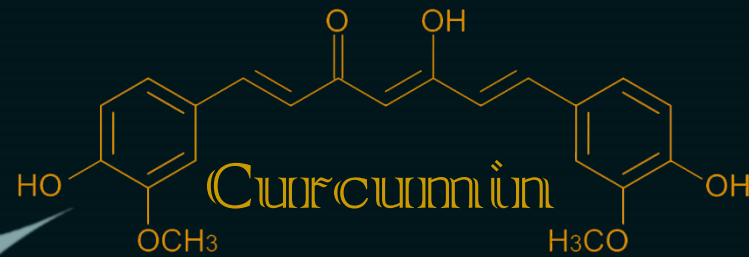


- Chemical composition of turmeric

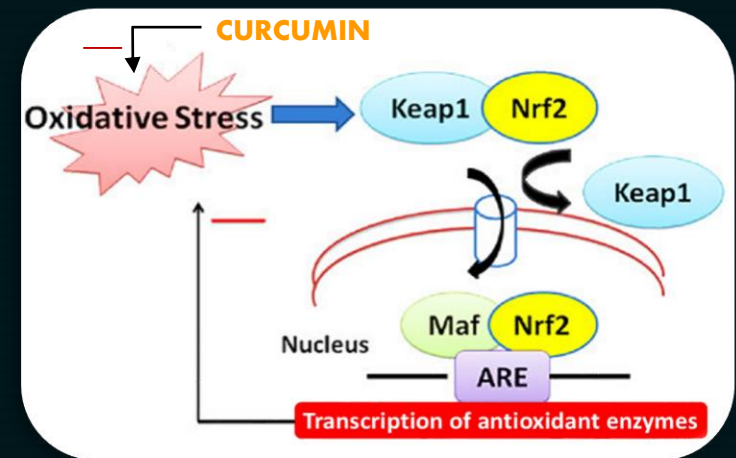
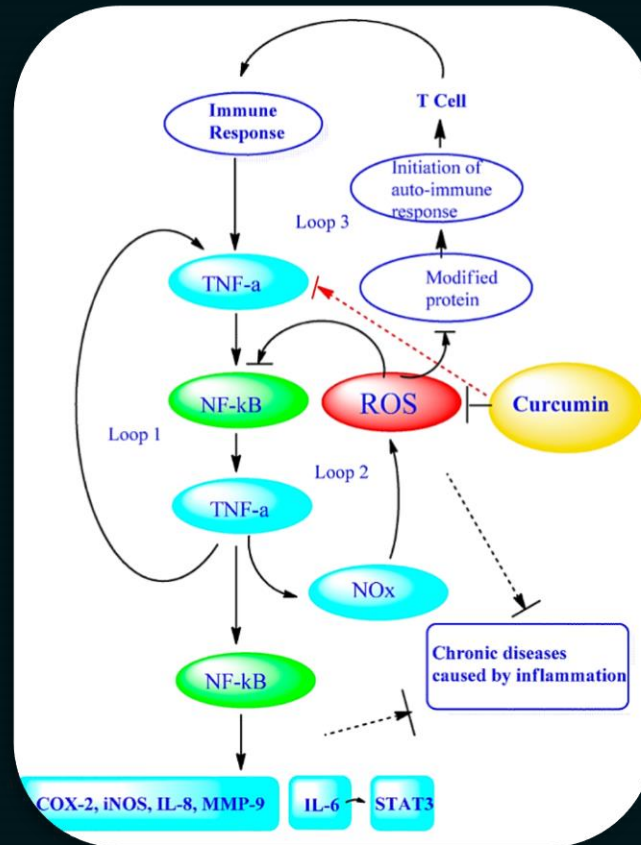




# Introduction

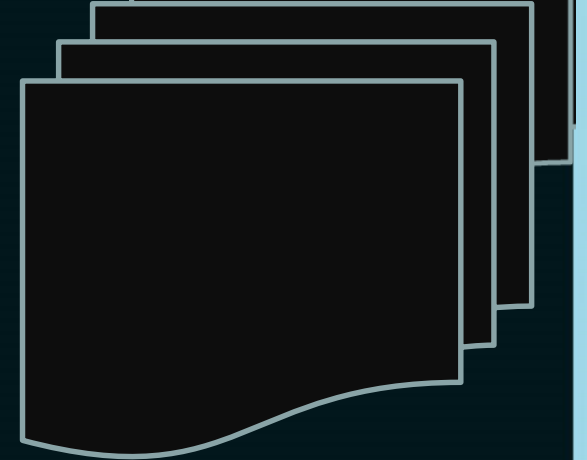
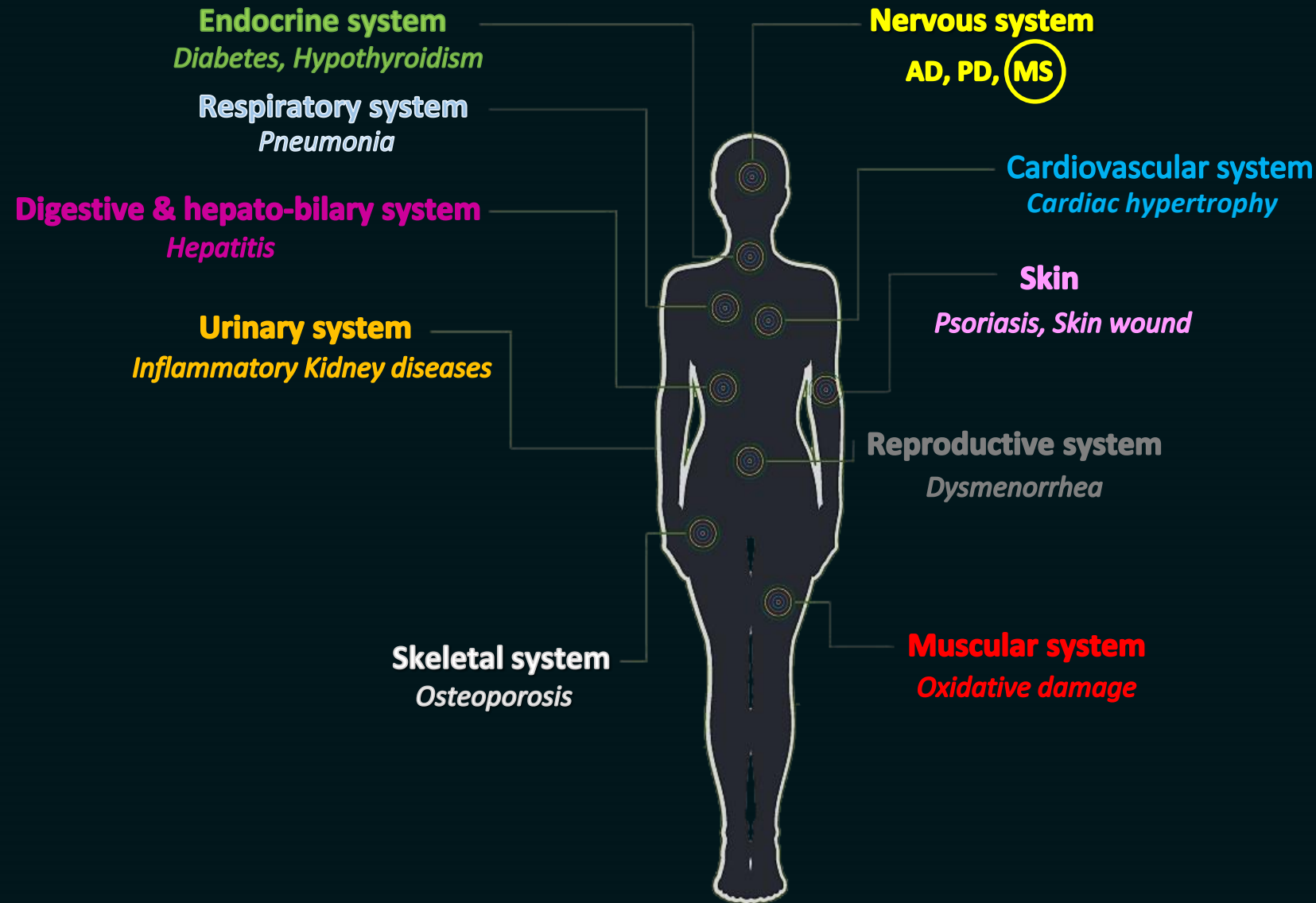


Antioxidant  
anti-inflammatory



# Curcumin

## Introduction



# Aim of This Study



*Introduction*

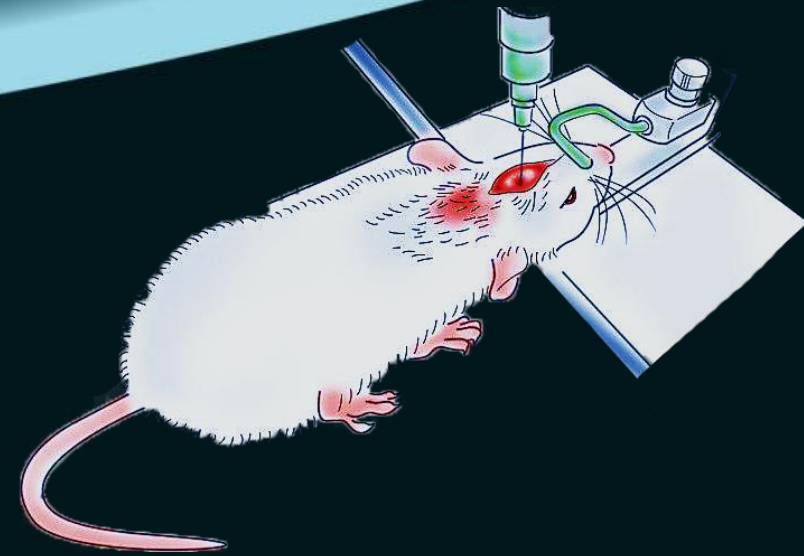


- ✓ Investigating memory improvement and antioxidant effect of curcumin and Lino-curcumin on EB-induced MS model.
- ✓ Compare Lino-curcumin efficacy with curcumin.

# MATERIALS

&

# METHODS



# Animals

- ❑ Rats were purchased from Pastor institute.
- ❑ **Situation:** temperature:  $22 \pm 2^\circ\text{C}$   
lights on 07:00–19:00 h  
ad libitum access to food and water
- ❑ **Subjects:** 56 Male Wistar rats  
weighing  $250 \pm 20$  gr  
divided in 7 groups (n=8)

*Materials  
&  
Methods*



# Groups

1. Control
2. Sham (DMSO/PBS 1% v/v)
3. EB (4  $\mu\text{g}/\text{rat}$ )
4. EB+ 5  $\mu\text{g}/\text{rat}$  Cur.
5. EB+ 10  $\mu\text{g}/\text{rat}$  Cur.
6. EB+ 5  $\mu\text{g}/\text{rat}$  Lino-cur.
7. EB+ 5  $\mu\text{g}/\text{rat}$  Lino-cur.



*Materials  
& Methods*

# Stereotaxic Operation

ICV co-ordination:

Coronal

AP: - 0.6 mm

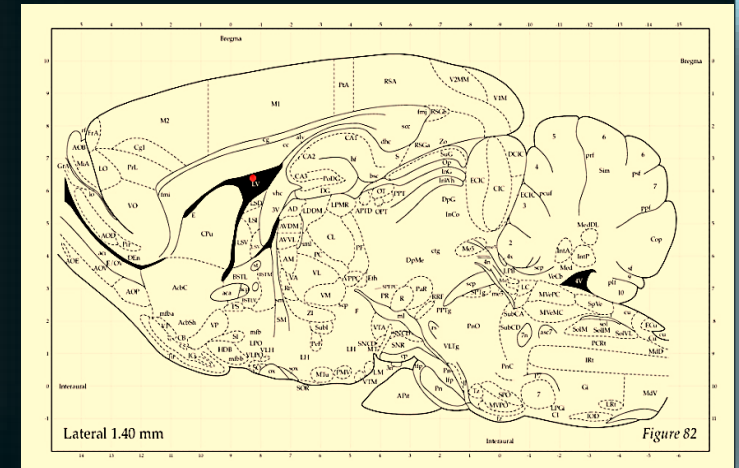
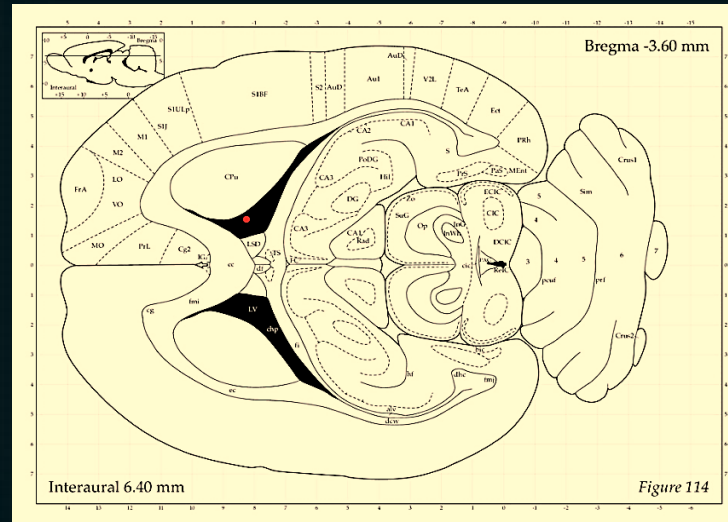
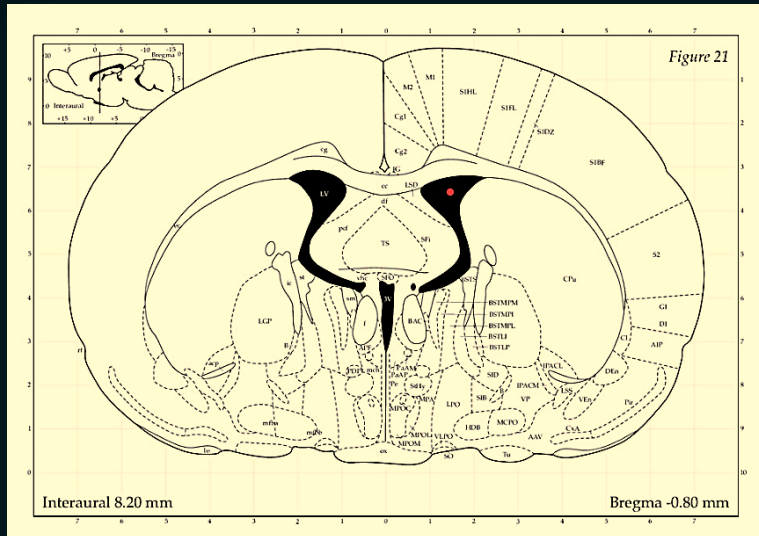
ML:  $\pm$  1.6 mm

DV: 4.0 mm

Horizontal

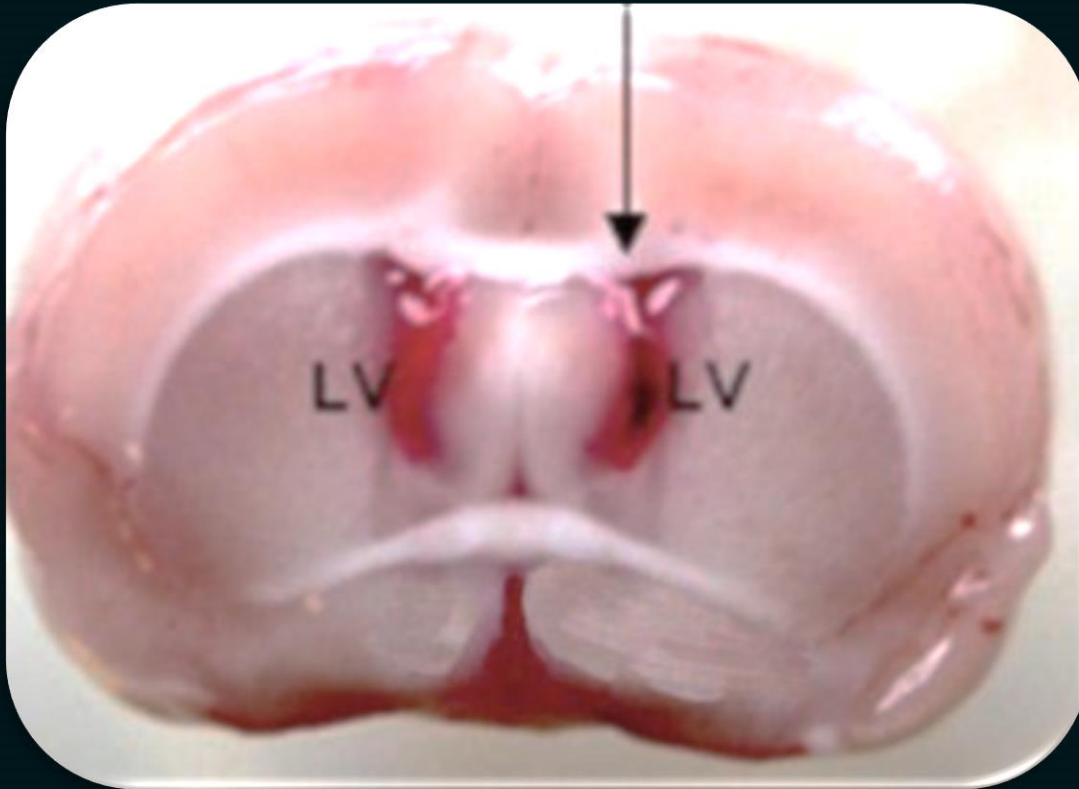
Materials  
& Methods

Sagittal



# Histological confirmation of Stereotaxic

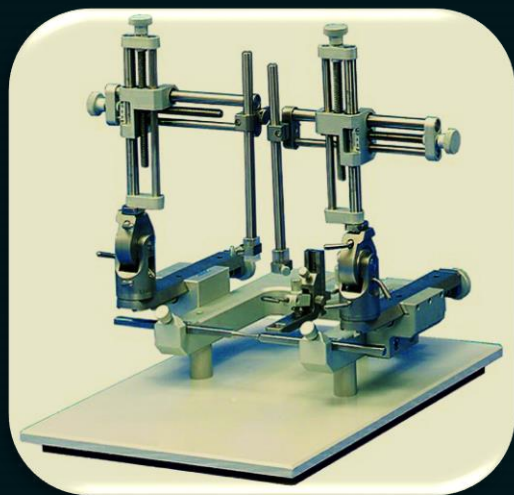
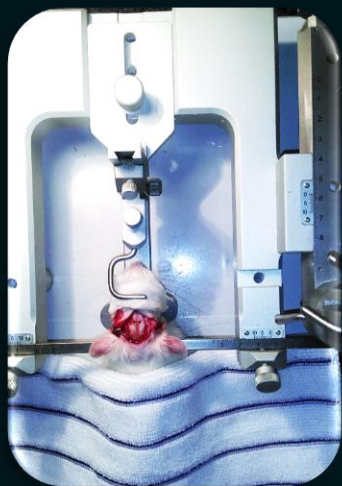
*Materials  
& Methods*





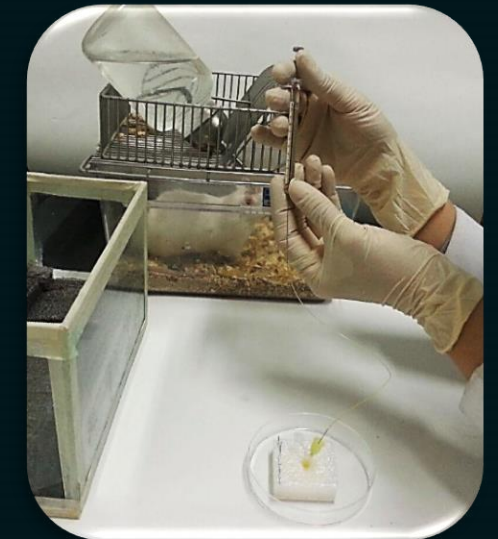
# Stereotaxic Operation

*Materials  
& Methods*

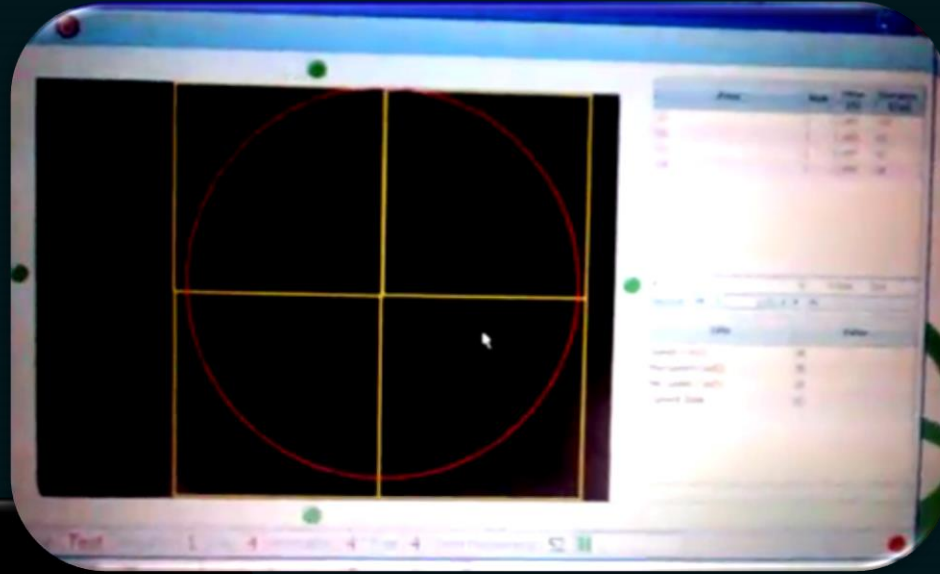


# MS model induction & Injections

*Materials  
& Methods*



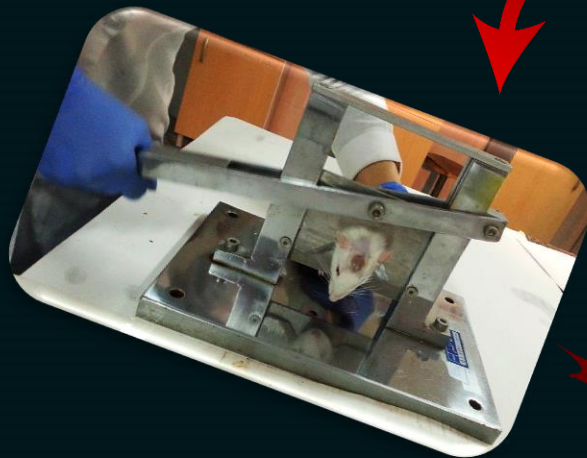
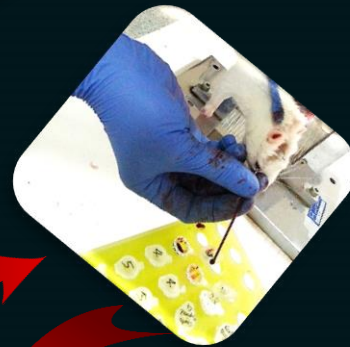
# Morris Water Maze test



*Materials  
& Methods*



# dissection & Sampling



*Materials  
& Methods*

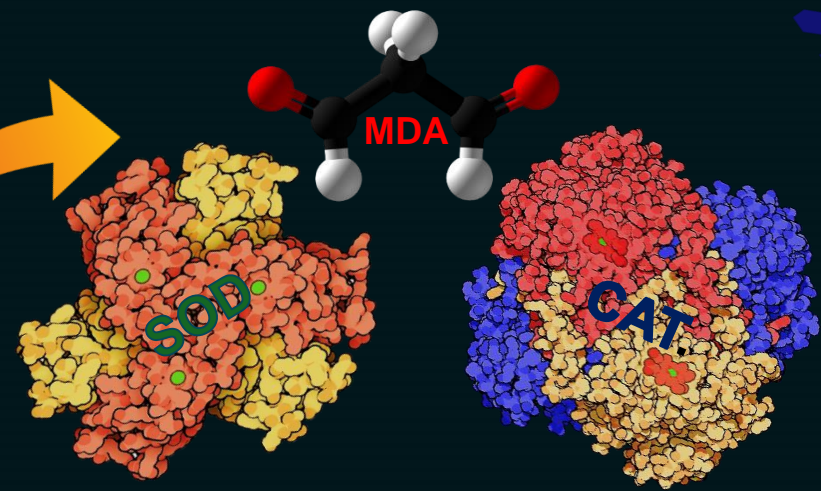
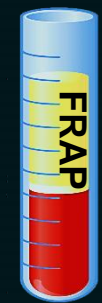
# Homogenization

*Materials  
& Methods*

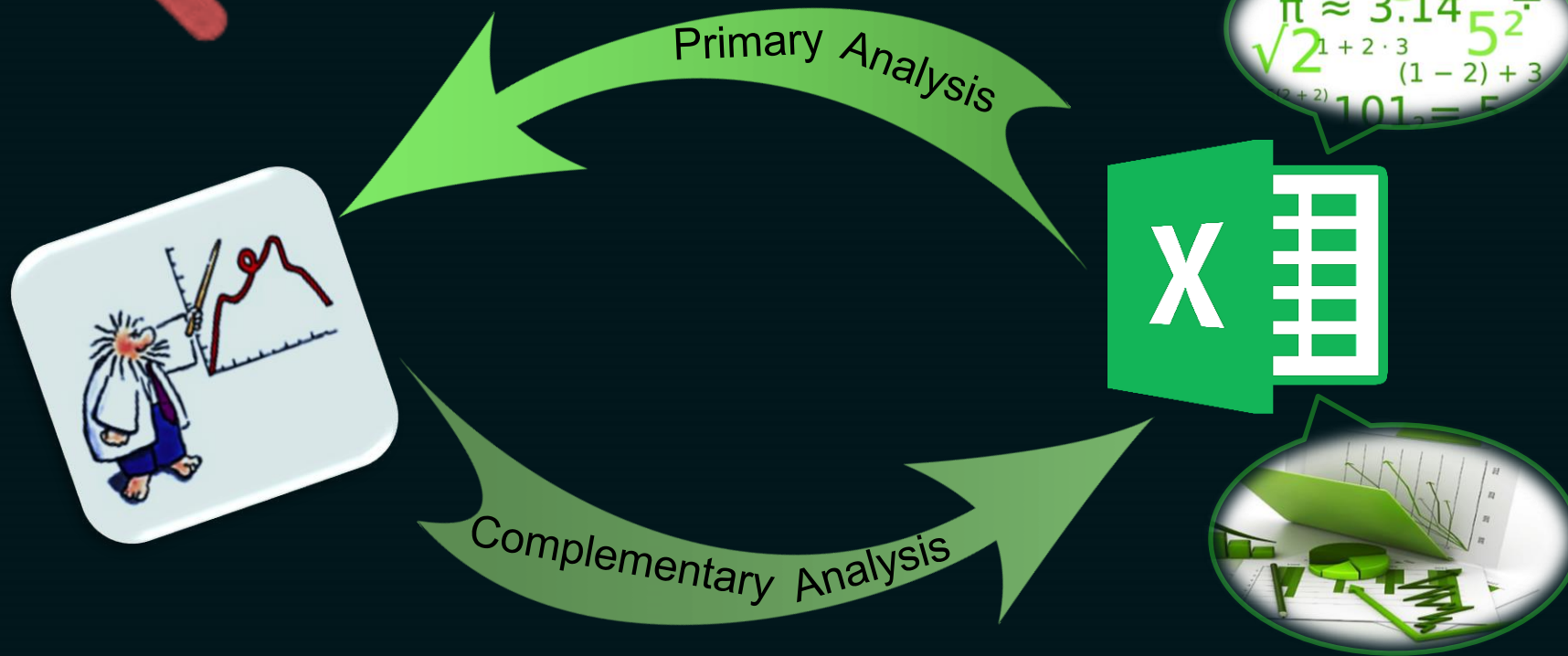


# Enzymes Assay

*Materials  
& Methods*



# Statistical Analysis



*Materials  
&  
Methods*

- ▼ The data were expressed as means  $\pm$  SEM.
- ▼ One-way analysis of variance (ANOVA), followed by Tukey post-test
- ▼ Differences were considered significant if  $P < 0.05$

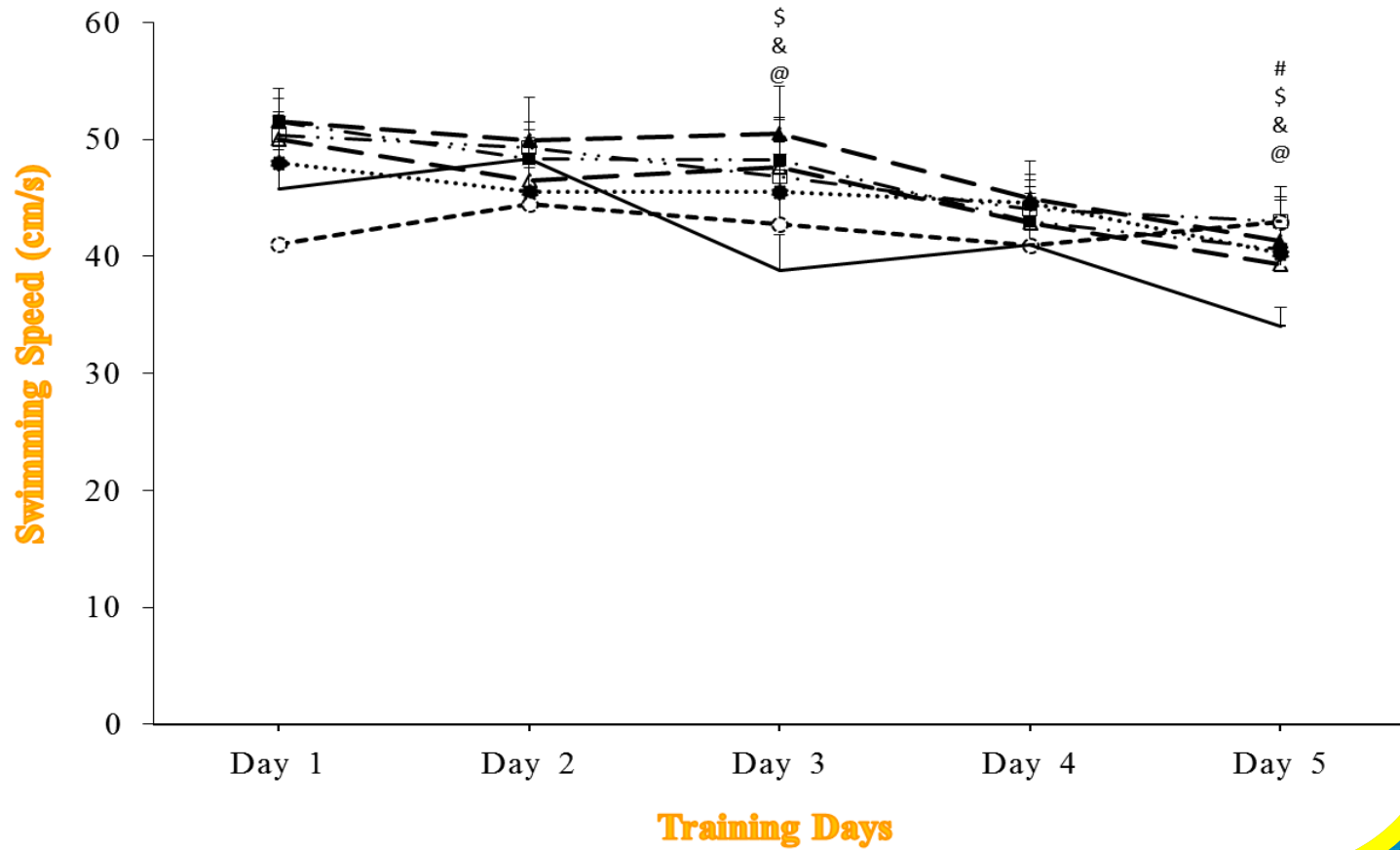
# RESULTS





# Spatial Memory

## MWM (Swimming Speed)

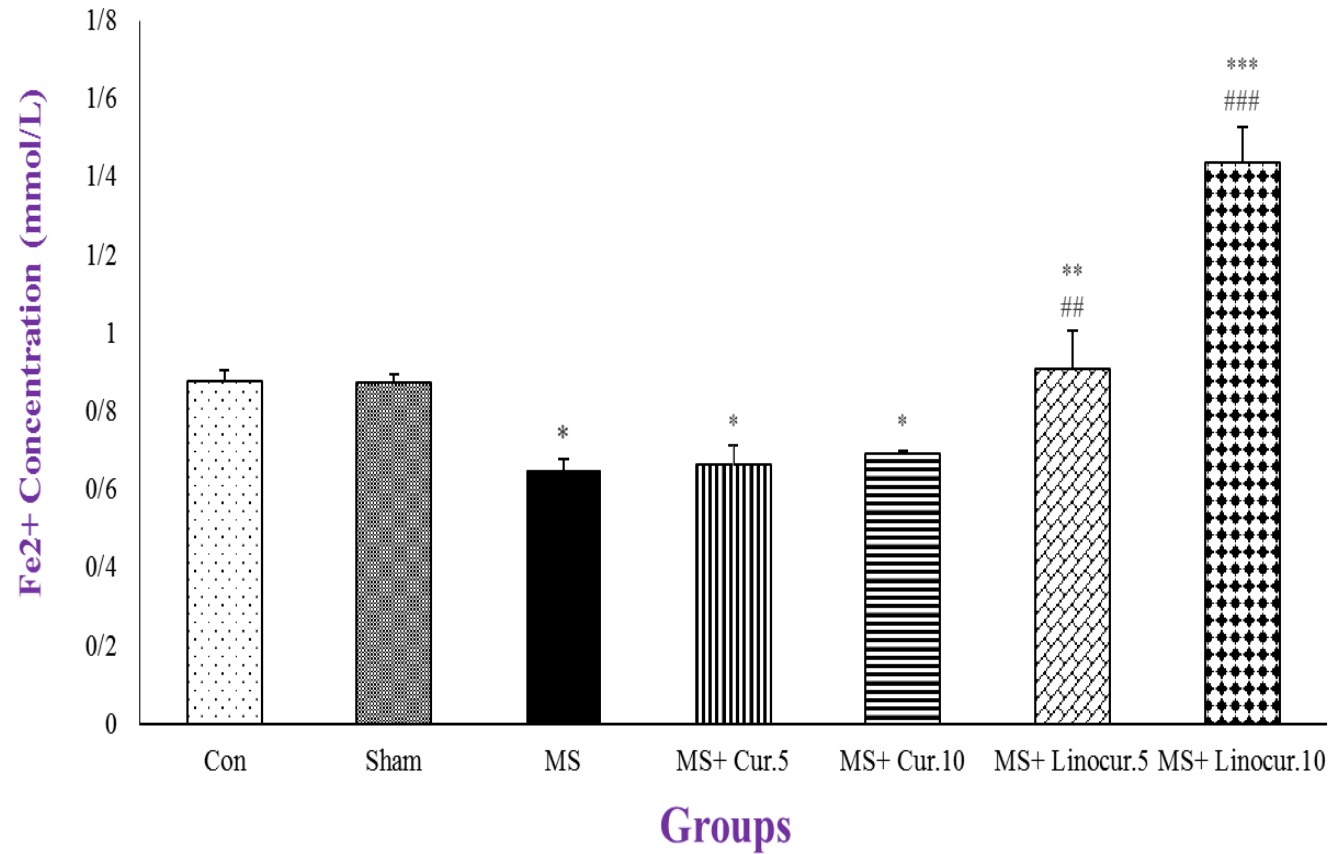


*Results*

- Con.
- Sham
- EB
- ▲ EB+Cur.5
- △ EB+Cur.10
- EB+Lino-cur.5
- EB+Lino-cur.10

# Oxidative Stress

## Ferric Reducing Ability of Plasma (FRAP)



*Results*

- Con
- Sham
- MS
- ▨ MS+ Cur.5
- ▨ MS+ Cur.10
- ▨ MS+ Linocur.5
- ▨ MS+ Linocur.10

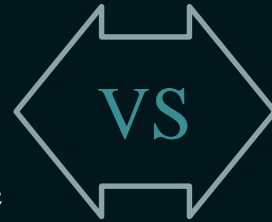
# DISCUSSION



# Spatial memory (current study)

**EB**  
injection

- Swimming speed
- ↑ Time Latency
- ↑ Travelled Distance



**Control**

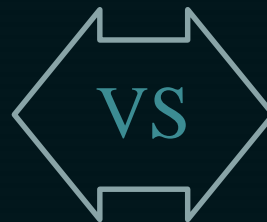
*Discussion*

*Spatial memory  
impairment*

*Spatial memory  
improvement*

**Cur.**  
&  
**Lino-cur.**  
treats

- Swimming speed
- ↓ Time Latency
- ↓ Travelled Distance



**EB**

*Lino-cur > Cur.*

# Spatial memory (Previous Studies)

**EB**  
injection

- Swimming speed
- ↑ Time Latency
- ↑ Travelled Distance



Ziehn. et al, 2010  
Kim. et al, 2012  
Ghaffary. et al, 2013

**Cur.**  
&  
**Lino-cur.**  
treats

- Swimming speed
- ↓ Time Latency
- ↓ Travelled Distance



Sethi. et al , 2009  
Kuhad. et al, 2007  
Belviranlı. et al, 2013  
Tang. et al, 2009  
Xu. et al, 2009

*Discussion*

# Oxidative Stress Parameters (current study)



# Oxidative Stress Parameters (Previous studies)

Cur.  
&  
Lino-cur.  
treats

↑ FRAP  
↓ MDA  
↓ SOD  
↓ CAT.



Abdel-salam. et al, 2012  
Spanevello. et al, 2009

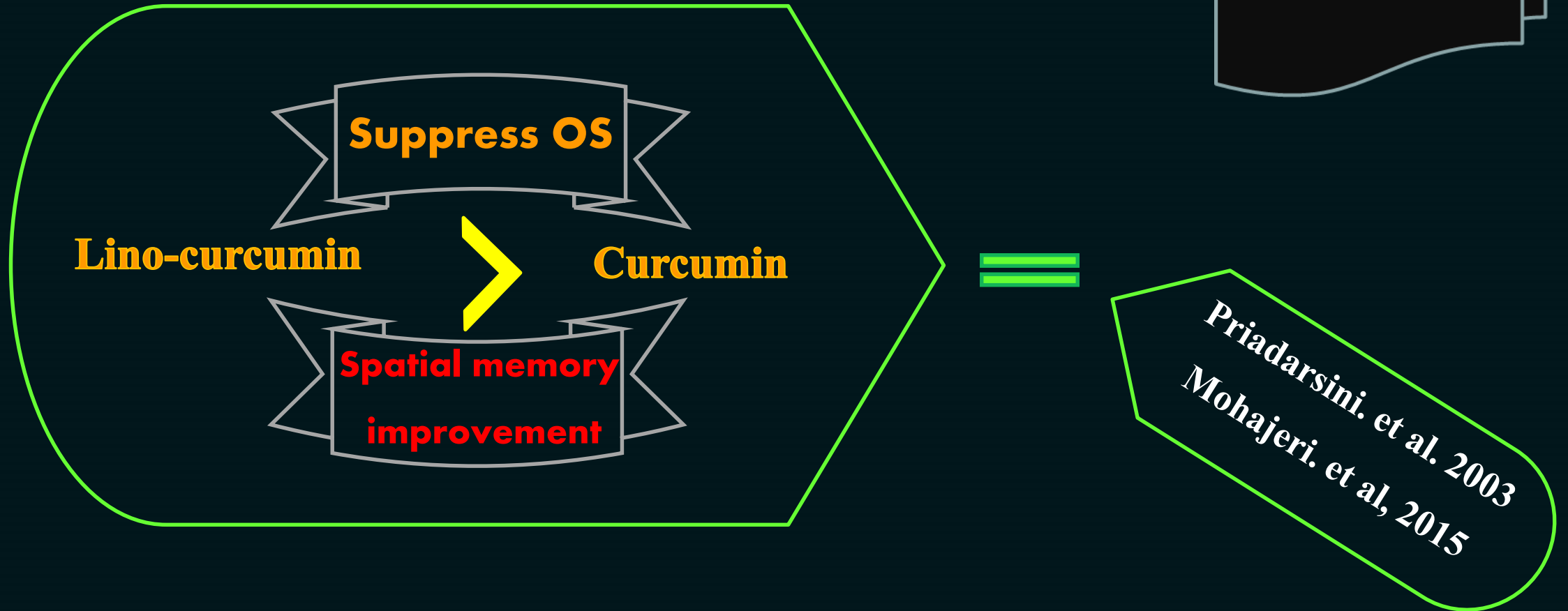


Sethi. et al , 2009  
Kuhad. et al, 2007  
Belviranlı. et al, 2013  
Mohajeri. Et al, 2015

*Discussion*

# Spatial memory & Oxidative stress

*Discussion*





# Conclusion

## *EB Model of MS*

- ⊗ Oxidative stress
- ⊗ Spatial memory impairment

## *Lino-cur. & Cur. treatment*

- ⊗ Oxidative parameters decreased
- ⊗ Spatial memory improved

*Discussion*

**Using Lino-curcumin was more effective on normalizing oxidative stress parameters and improvement of spatial memory in compare with curcumin during MS model in male rats.**

# Suggestions



*Discussion*



- **Investigating the long term of this treatment**
- **Test this treatment on the other models of MS, e.g. EAE**
- **Analyzing the results of this method on spinal cord tissue**
- **Analyzing gene expression of MBP during MS + Lino-cur.**
- **Investigating the brain histological study during MS + Lino-cur.**



PARTICIPATED



CONGRESSES

بِسْمِ اللَّهِ الرَّحْمَنِ الرَّحِيمِ



دانشگاه علوم پزشکی کاشان  
Kashan University  
of Medical Sciences



*CERTIFICATE OF ATTENDANCE*

*Poster Presentation*

This is to certify that  
**Behnaz Barzegarzadeh**  
attended the

**1<sup>st</sup> International and 22<sup>nd</sup> Iranian Congress of Physiology and Pharmacology**  
held in Kashan University of Medical Sciences, 7-11 September, 2015

*Dr. Mohammad Hossein Aarabi*

*The President*

*Dr. Mahmoud Salami*

*The Secretary General*

اولین کنگره بین‌المللی  
فیزیولوژی و فارماکولوژی  
و بیست و دومین کنگره  
فیزیولوژی و فارماکولوژی ایران  
۱۶ - ۲۰ شهریور ماه ۱۳۹۴  
دانشگاه علوم پزشکی کاشان  
  
1<sup>st</sup> International and  
22<sup>nd</sup> Iranian Congress of  
Physiology and Pharmacology  
7-11 Sept 2015  
Kashan University  
of Medical Sciences

بناام خدا

گواهی می شود آقای /خانم بهناز برزگزاده، حمیرا حاتمی، غلامرضا دهقان

در نوزدهمین کنگره ملی و هفتمین کنگره بین المللی زیست شناسی ایران

که در تاریخ ۹-۱۱ شهریور ۱۳۹۵ در دانشگاه تبریز برگزار گردید، با ارائه سخنرانی با عنوان زیر شرکت نمودند:

اثر ریزتزریق کورکومین بر بهبود حافظه فضایی طی بیماری مالتیپل اسکلروزیس در رت های نر

This is to certify that Behnaz Barzegarzadeh, Homeira Hatami, Golam-Reza Dehgan  
Have attended the 19<sup>th</sup> National and 7<sup>th</sup> International Biology Congress,  
University of Tabriz, Iran and presented the following paper:  
The effect of curcumin microinjection on spatial memory improvement during multiple sclerosis in male rats

Dr M. Sharifi  
Head of Iranian Biology Society



نوزدهمین و هفتمین

کنگره ملی و بین المللی  
زیست شناسی ایران  
19th National & 7th International  
Congress of Biology, Iran

Dr R. Hajiboland  
Secretary of Congress



شماره:

تاریخ:



دانشگاه تبریز



انجمن زیست شناسی ایران



نوزدهمین و هفتمین  
کنگره ملی و بین المللی  
زیست شناسی ایران  
19th National & 7th International  
Congress of Biology, Iran





IUMS



CMRC



INSS

5<sup>th</sup>

Basic and Clinical  
**NEUROSCIENCE**  
Congress 2016

December 7-9, 2016

Razi Hall, Tehran, Iran

This is to certify that

**Behnaz Barzegarzadeh**

*Had a Poster Presentation Entitled:*

**“Investigating the Microinjection Effect of Curcumin-Linoleic Acid on Spatial Memory in Multiple Sclerosis Model of Male Rats”**

*Participated at the 5<sup>th</sup> Basic and Clinical Neuroscience Congress*

*7 - 9 Dec. 2016 Tehran, Iran*

*Prof. Mohammad Taghi Joghataie*  
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*Basic Scientific Secretary*

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*Clinical Scientific Secretary*



Submitted

Article



## Submission Confirmation



Thank you for your submission

**Submitted to** Archives Of Physiology And Biochemistry

**Manuscript ID** NAPB-2017-0016

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**Authors** barzegarzadeh, behnaz  
Hatami, homira

**Date Submitted** 31-Jan-2017



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Dr. F. Sheikhzadeh

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Miss. S.N. Fotuhi

Mr. Y. Alipour

Dr. M.A. Doctorzadeh



Thanks  
for  
Your  
Consideration

Any   
Question 