

# Inflammation chronique dans les maladies cardiovasculaires liées à l'athérome : Place de l'alimentation

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Chirurgien des Hôpitaux

# L'athérome une maladie chronique non transmissible

- Maladie de rétention avec dépôt dans la paroi des artères
- Polygénique multifactorielle
- Inflammatoire et auto-immune
- Les lipoprotéines de basse densité (LDL), les chylomicrons et les macrophages au cœur de la rétention et de la constitution de la plaque

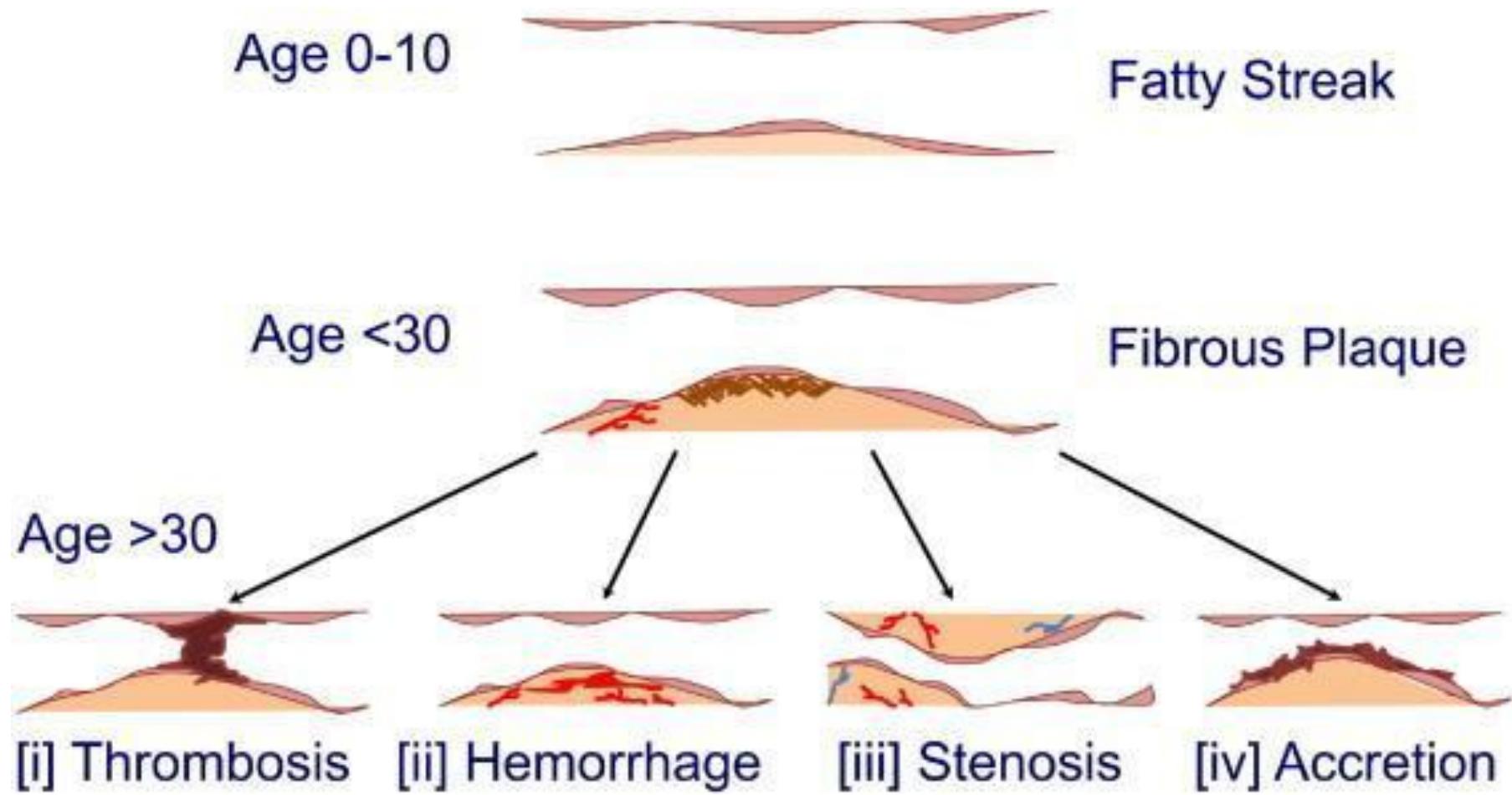


Figure 1 Evolution of human atherosclerosis. Fatty streaks represent one of the earliest visible lesions in atherogenesis, and have been observed early in life. These fatty streaks often evolve into fibrous plaques, coupled with intimal hyperplas...  
Brian W. Wong , Anna Meredith , David Lin , Bruce M. McManus

#### The Biological Role of Inflammation in Atherosclerosis

Canadian Journal of Cardiology Volume 28, Issue 6 2012 631 - 641

<http://dx.doi.org/10.1016/j.cjca.2012.06.023>

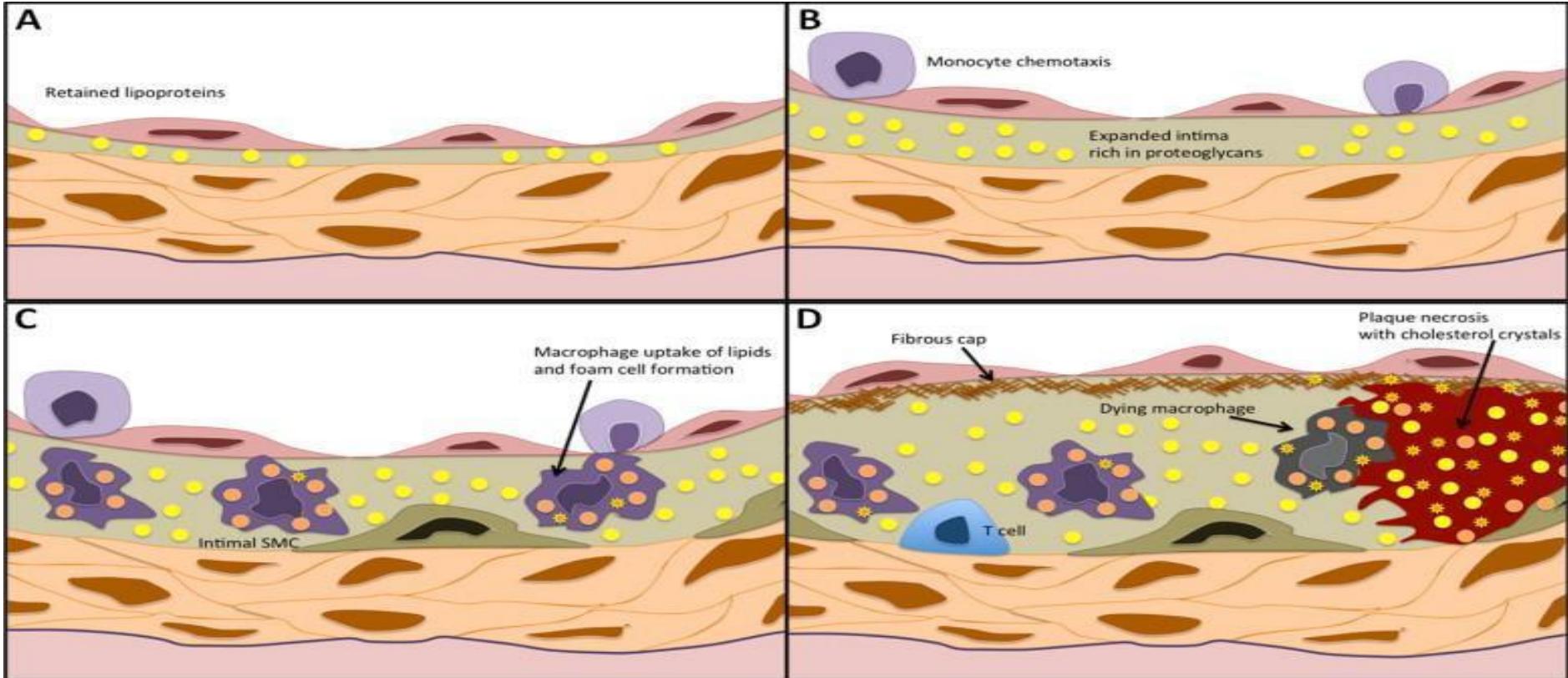


Figure 3 Progression of atherosclerosis and plaque formation. ( A ) The subendothelial retention of lipoproteins is an early initiating factor in atherogenesis. ( B ) This initiating process results in monocyte chemotaxis in response to trapped I...

Brian W. Wong , Anna Meredith , David Lin , Bruce M. McManus

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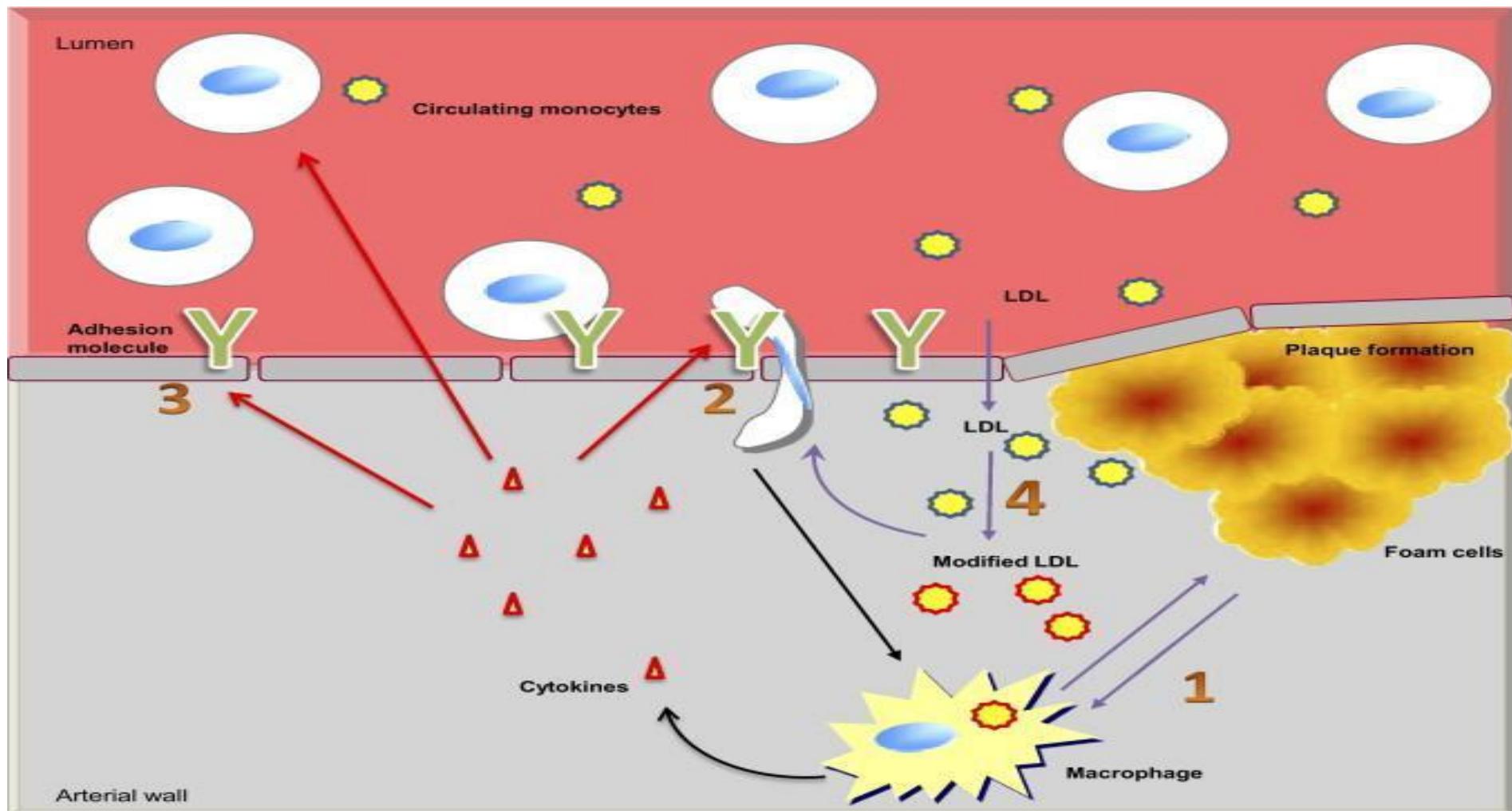


Figure 1 Inhibition of atherosclerosis by high-density lipoprotein (HDL). The diagram highlights the targets of the antiatherosclerotic actions of HDL. HDL inhibits atherosclerosis by promoting cholesterol efflux from foam cells (1), by regulatin...

Valérie Duchatelle , Ekaterini A. Kritikou , Jean-Claude Tardif

### Clinical Value of Drugs Targeting Inflammation for the Management of Coronary Artery Disease

Canadian Journal of Cardiology Volume 28, Issue 6 2012 678 - 686

<http://dx.doi.org/10.1016/j.cjca.2012.05.003>

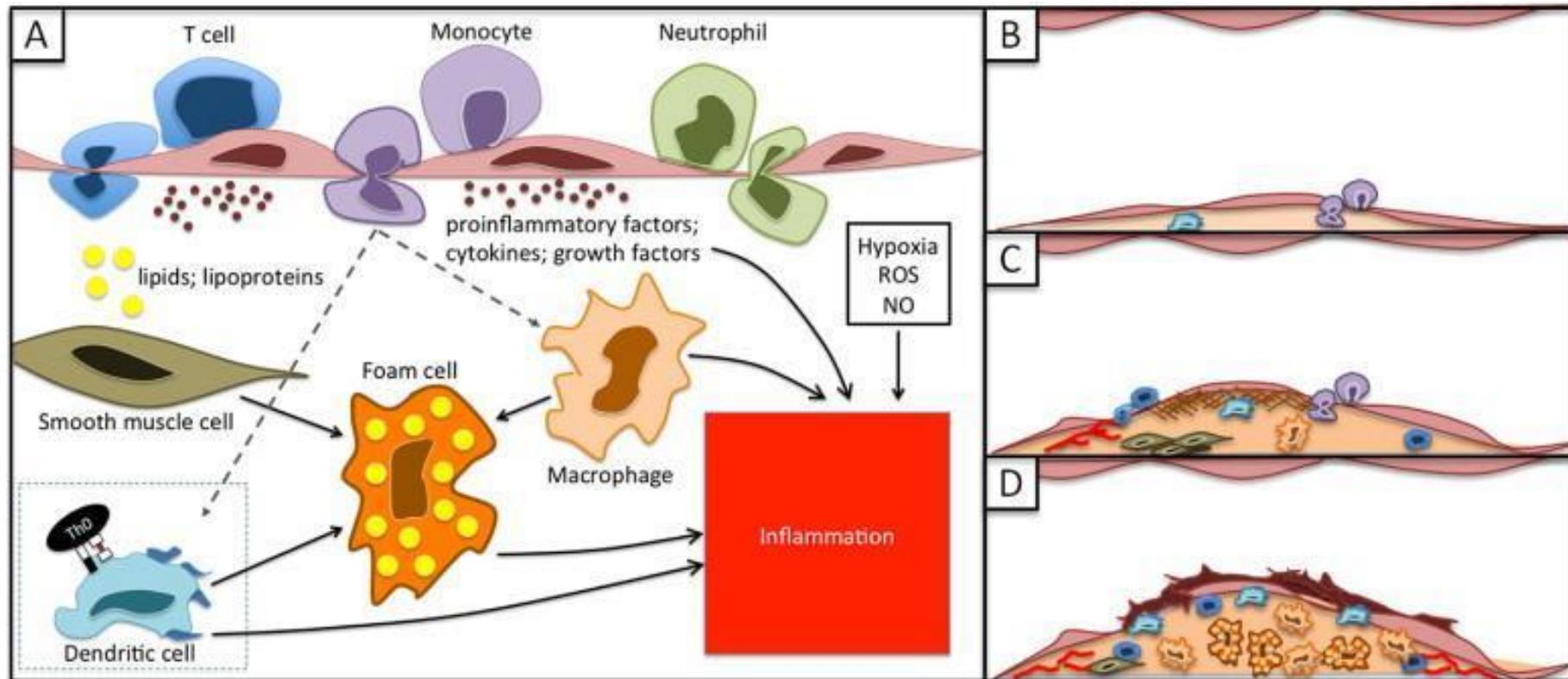


Figure 2 Inflammatory mediators in atherosclerosis. (A) Numerous inflammatory cell types play major roles in mediating the inflammatory response which is part of atherosclerosis, including T-cells, monocytes, and neutrophils. Early insudation a... Brian W. Wong<sup>1</sup>, Anna Meredith<sup>1</sup>, David Lin<sup>2</sup>, Bruce M. McManus<sup>1</sup>

### The Biological Role of Inflammation in Atherosclerosis

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# Athérome carotidien évolué avec hémorragie intraplaque



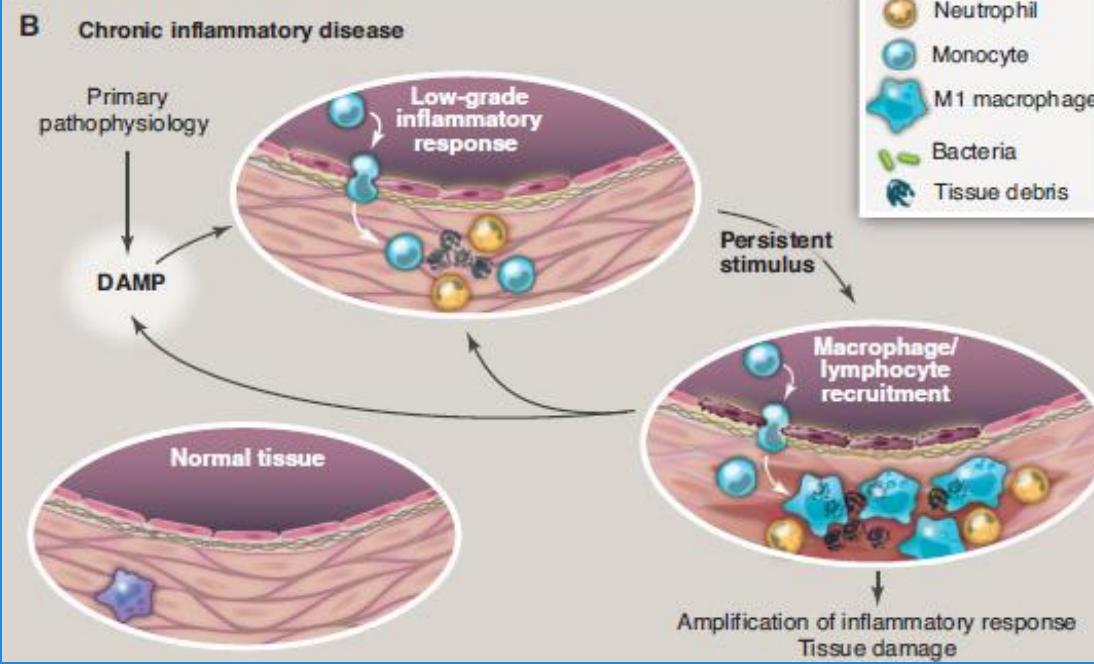
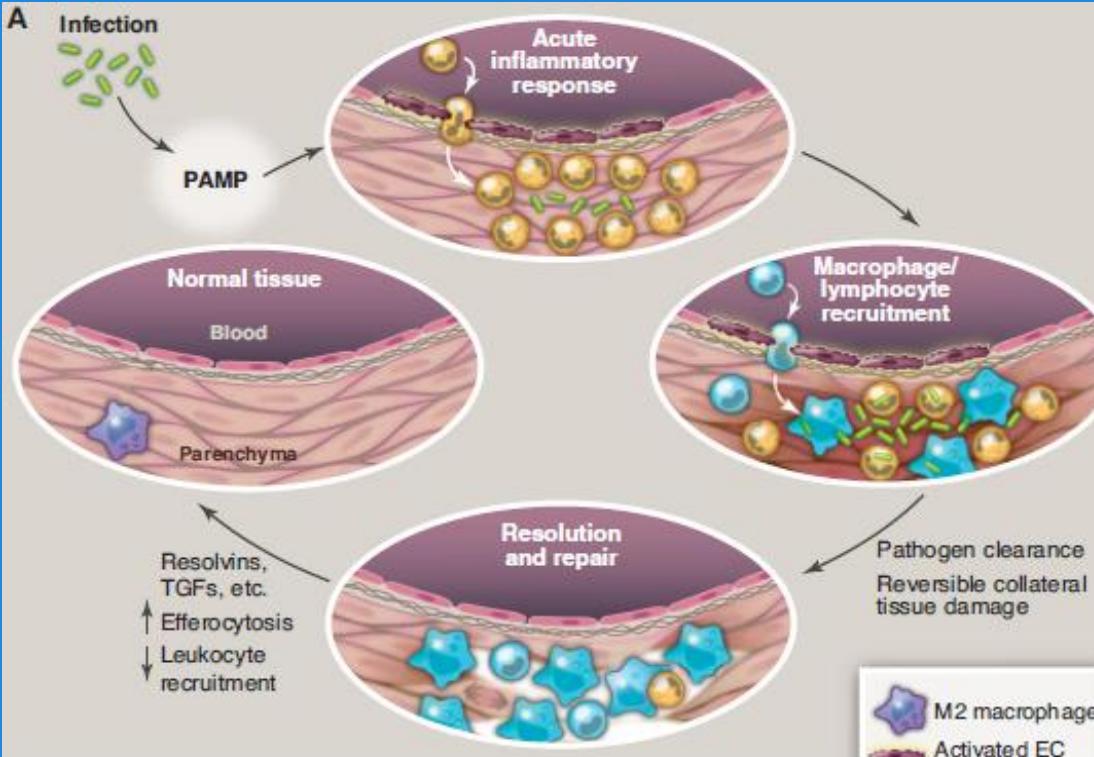
# Inflammation chronique et athérome

- Marqueurs de l'inflammation chronique  
VS  
CRP ultrasensible  
Fibrinogène
- Modèles cliniques d'inflammation chronique avec accélération de l'athérogénèse  
Diabète type 2  
Fumeur actif  
Polyarthrite rhumatoide

# Preuves diagnostiques de l'inflammation dans la plaque d'athérome

Table 1. Molecular imaging targets for the vulnerable plaque

Process	Target	Agent	Imaging modality	Phase (reference)
Inflammation	Macrophages	<sup>18</sup> F-FDG	PET or PET/CT	Clinical [39–43, 65–69]
		<sup>11</sup> C-PK11195	PET	Clinical [71–73]
		<sup>99m</sup> Tc-labeled anti-LOX-1	SPECT	Preclinical [46, 47]
		Nanoparticles	MRI	Preclinical [51, 53]
Apoptosis	P-selectin (EC and platelets)	<sup>99m</sup> Tc-fucoidan	SPECT/ MRI	Preclinical [59–61]
	VSMC, macrophages	<sup>99m</sup> Tc-anxA5	SPECT	Clinical [42, 83–85, 91]
Vascular remodeling	VSMC	Z2D3	SPECT	Preclinical [96–101]
	MMP	P947	MRI	Preclinical [23, 25, 54, 102]
	integrin $\alpha_v\beta_3$	Nanoparticles	MRI	Preclinical [26]
	ED-B	L19	SPECT	Preclinical [107]



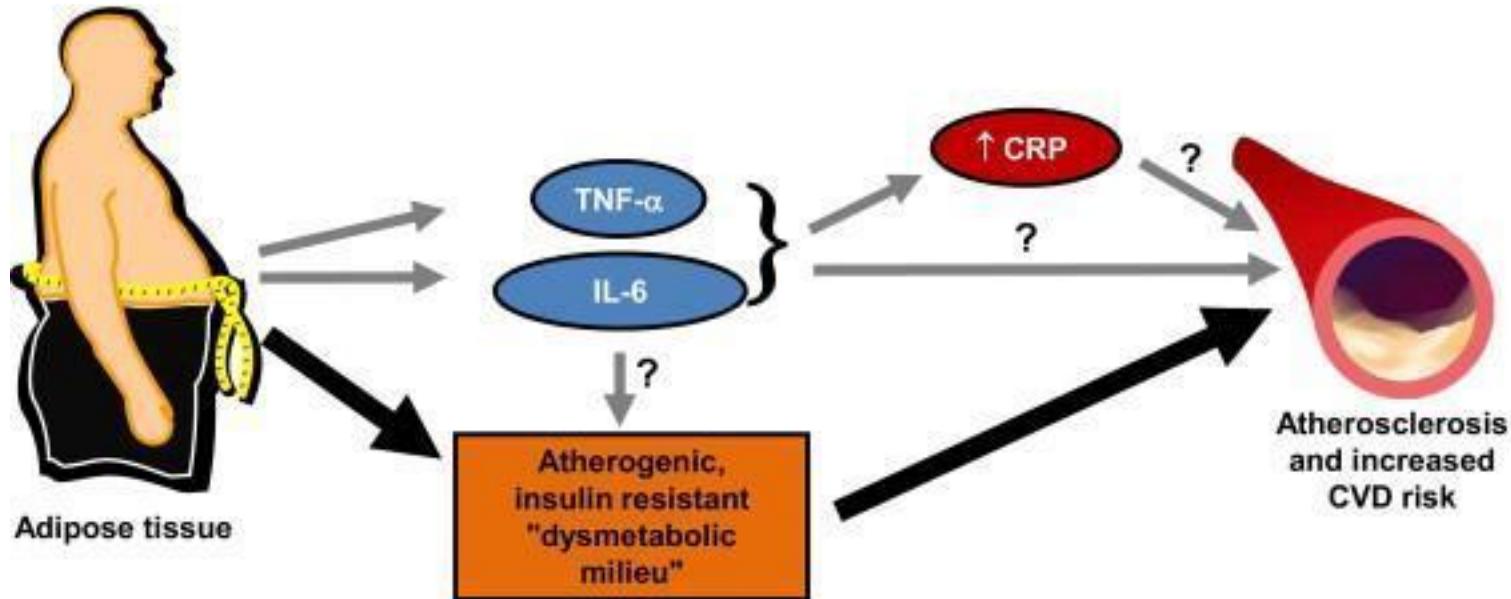


Figure 4 Working model by which an elevated C-reactive protein (CRP) concentration predicts the risk of an acute coronary syndrome. First, CRP per se may have a direct impact on processes involved in the etiology of coronary heart disease. Second...

Jean-Pierre Després

### **Abdominal Obesity and Cardiovascular Disease: Is Inflammation the Missing Link?**

Canadian Journal of Cardiology Volume 28, Issue 6 2012 642 - 652

<http://dx.doi.org/10.1016/j.cjca.2012.06.004>

GENETICS/ENVIRONMENT

?

LIPID ENZYMES

Therapy

Disease

LIPIDS

Resolve

Inflame

LIPID RECEPTORS

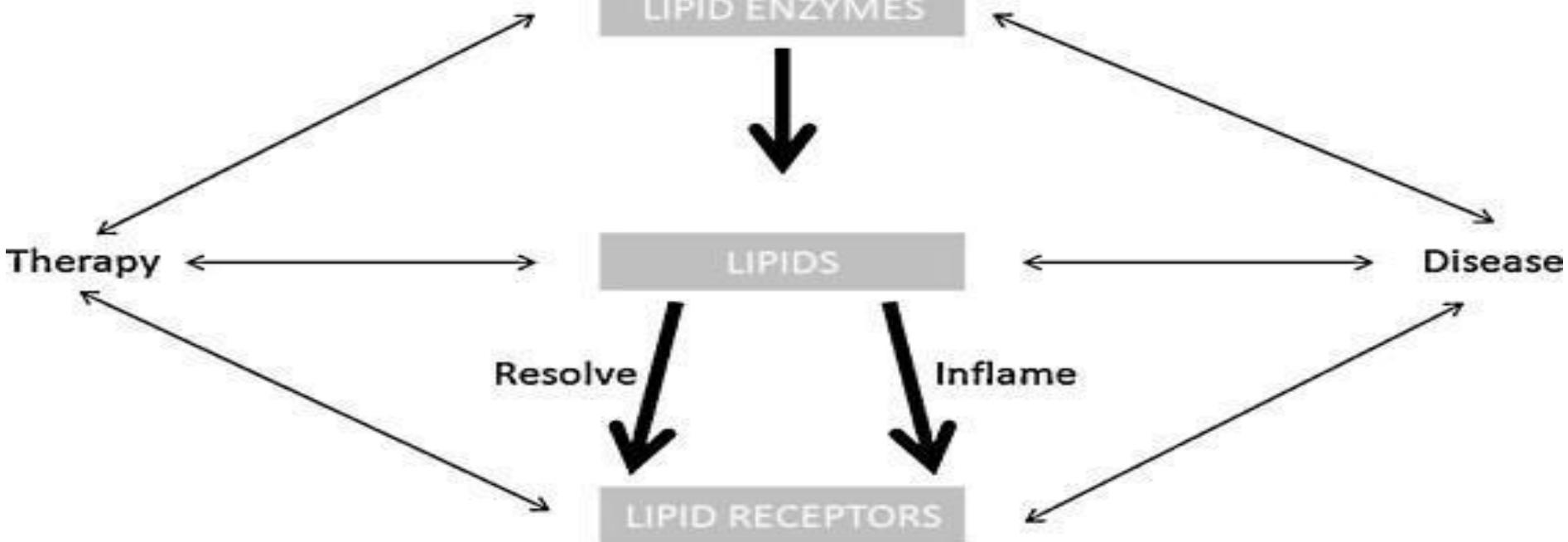


Fig. 1 The continuum of bioactive lipid pathways. Whether prostaglandins or resolvins bind lipid receptors in a therapeutic context, or HETE molecules interact with receptors to push cancer toward metastasis, there is evidence to support nearly ...

Stephanie C. Tucker , Kenneth V. Honn

### Emerging targets in lipid-based therapy

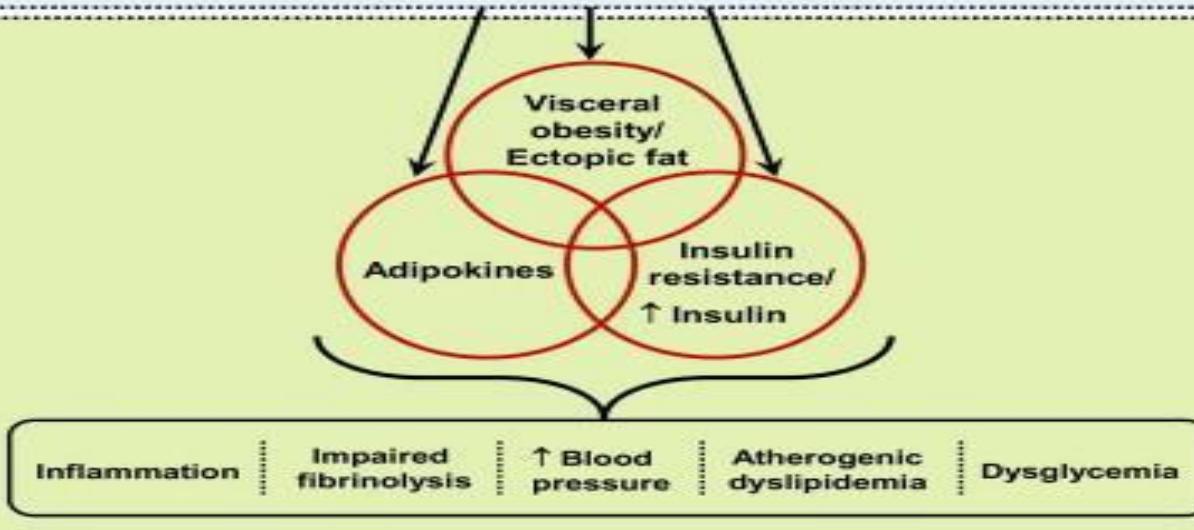
Biochemical Pharmacology Volume 85, Issue 5 2013 673 - 688

<http://dx.doi.org/10.1016/j.bcp.2012.11.028>

**A**

## **Genetic and environmental determinants**

- Energy-dense diet
- Genetic variation
- Age and sex
- Lack of physical activity
- Smoking
- Stress
- Neuroendocrine abnormalities
- Steroid hormones
- Susceptible endocannabinoid system
- Drugs

**B**

- Syndrome X (Reaven)
- Insulin resistance syndrome

⇒ **Based on key pathophysiological markers**

**C**

- Metabolic syndrome (NCEP-ATP III or IDF)

⇒

**Based on clinical tools:**

- |                 |                   |                  |
|-----------------|-------------------|------------------|
| • Waist girth   | • HDL cholesterol | • Blood pressure |
| • Triglycerides | • Glucose         |                  |

Figure 1 Simplified model showing the possible correlates ( A ) of insulin resistance often found among individuals with excess visceral/ectopic fat. ( B ) Illustrates the notion that the syndrome X/insulin resistance syndrome concept was based o...  
Jean-Pierre Després

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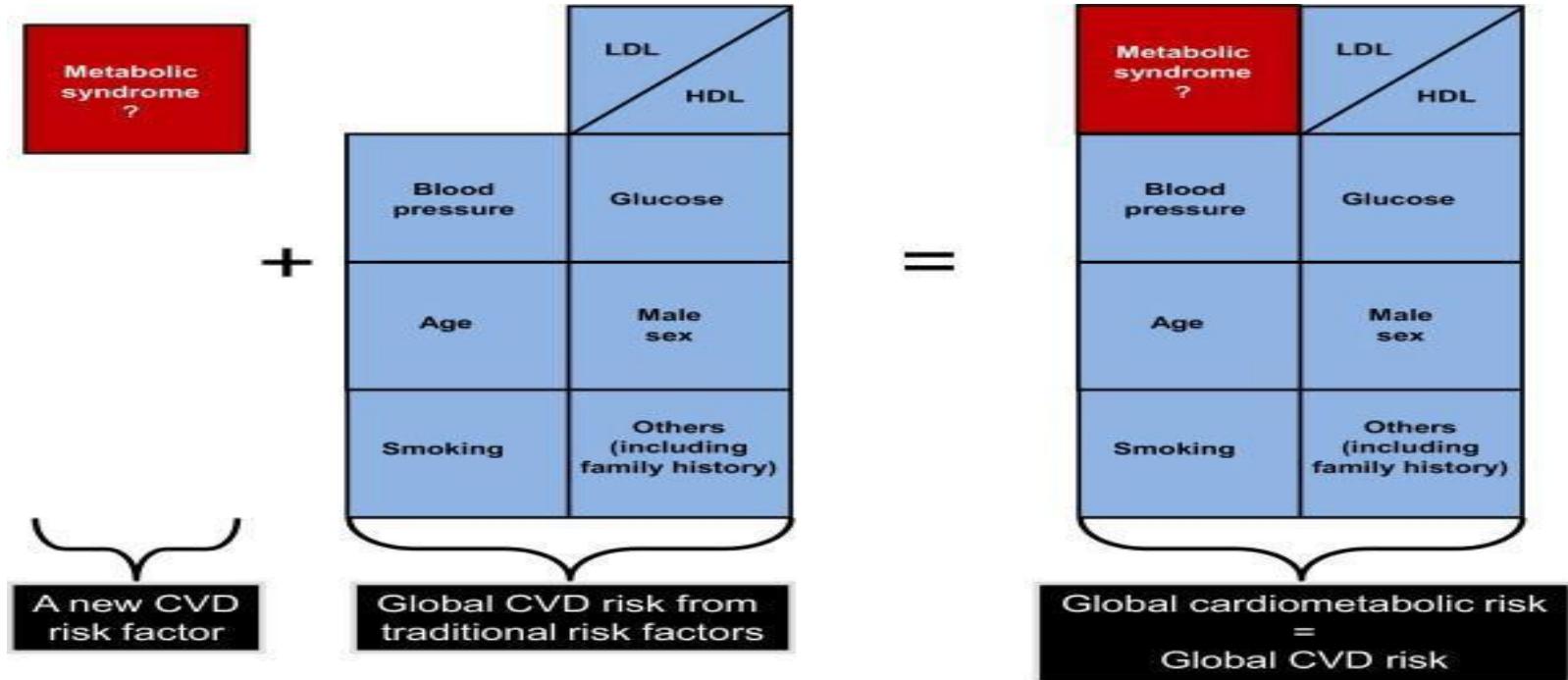


Figure 2 The ?building blocks? of global cardiometabolic risk. Cardiometabolic risk is the overall risk of cardiovascular disease (CVD) resulting from the presence of the metabolic syndrome but also of traditional risk factors such as lipids (low...

Jean-Pierre Després

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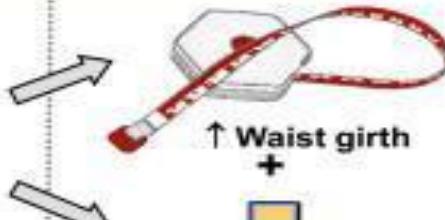
## FUNCTIONAL ADIPOSE TISSUE

### OBESITY PHENOTYPE

### SCREENING TOOL

#### Subcutaneous

#### Elevated waist girth



- Favourable genotype
- Healthy diet
- Physically active
- Insulin sensitive

Normal triglycerides

## DYSFUNCTIONAL ADIPOSE TISSUE

### OBESITY PHENOTYPE

### SCREENING TOOL

#### Visceral

#### Hypertriglyceridemic waist



- Unfavourable genotype
- Unhealthy diet
- Sedentary
- Insulin resistant

Elevated triglycerides

## HYPERTRIGLYCERIDEMIC WAIST: RELATIONSHIPS WITH METABOLIC ABNORMALITIES/CLINICAL OUTCOMES

- |  |   |
|--|---|
| <ul style="list-style-type: none"> <li>▪ Presence of atherogenic metabolic triad</li> <li>▪ ↑ Cholesterol/HDL cholesterol</li> <li>▪ Postprandial hyperlipidemia</li> <li>▪ Glucose intolerance</li> <li>▪ Hyperinsulinemia</li> <li>▪ ↑ Blood pressure</li> </ul> | <ul style="list-style-type: none"> <li>▪ ↑ Risk of cardiovascular disease</li> <li>▪ ↑ Risk of coronary artery disease</li> <li>▪ ↑ Annual progression rate of aortic calcification</li> <li>▪ ↑ Risk of type 2 diabetes</li> </ul> |
|--|---|

Jean-Pierre Després

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<http://dx.doi.org/10.1016/j.cjca.2012.06.004>

Figure 3 Use of hypertriglyceridemic waist as a screening tool to identify individuals likely to be characterized by the cluster of abnormalities of the metabolic syndrome. It is proposed that in presence of an elevated waist circumference, fasti...

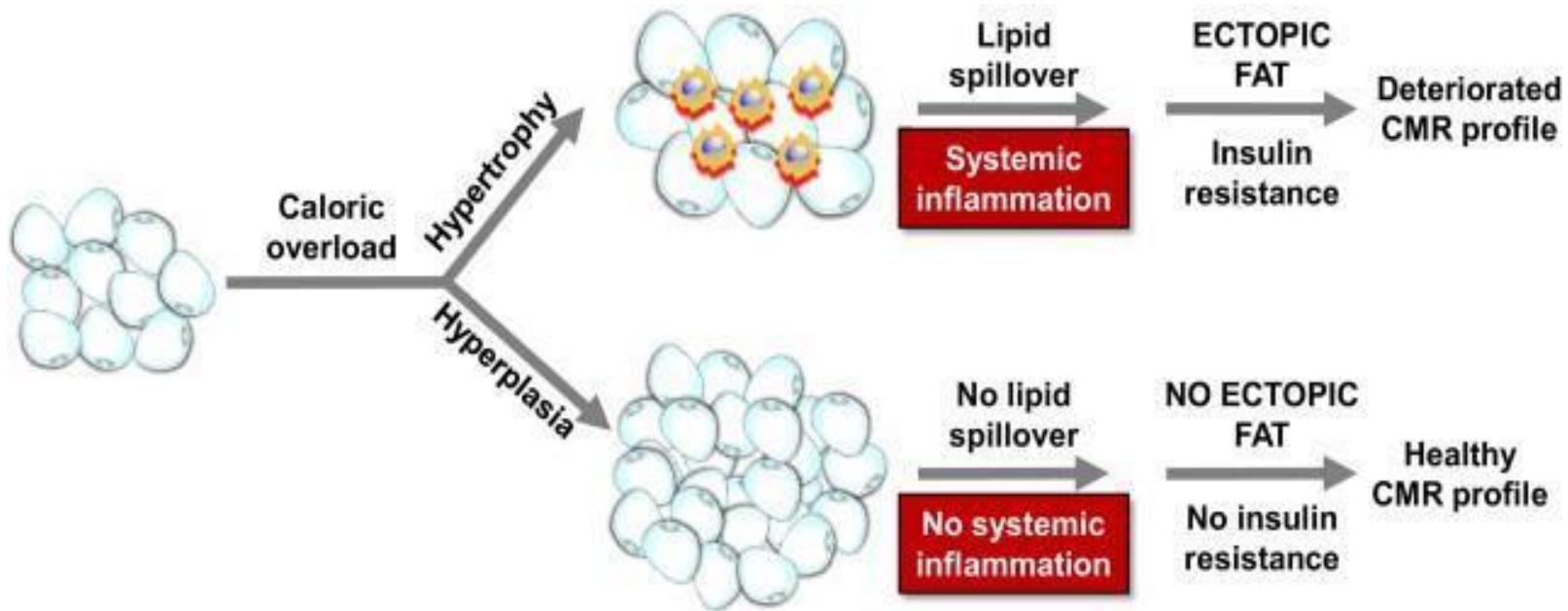


Figure 5 Simplified model to illustrate the concept that when facing a chronic energy surplus, inability of subcutaneous adipose tissue to expand through hyperplasia may lead to hypertrophic and ?inflamed? adipose tissue. Resulting systemic infla...

Jean-Pierre Després

#### **Abdominal Obesity and Cardiovascular Disease: Is Inflammation the Missing Link?**

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# Quels sont les facteurs non alimentaires pro-inflammatoires?

1. Tabac fumé actif ou passif ou fumée de cheminée
2. Sédentarité
3. Pollution environnementale par les particules fines: PM2.5 (*particulate matter less than 2.5 µm in diameter*)
4. Gingivite, Foyers infectieux chroniques (dentaires, osseux ...)
5. Faible exposition solaire et faible Vit D circulante

# Obésité et insuline

- Le gras stocké est plus souvent un excès calorique qu'une ingestion de graisses
- Le signal de stockage est amplifié par l'insuline
- Les aliments insulino-sécréteurs sont pro-inflammatoires
- Le rôle de la génétique est secondaire comme le démontre l'explosion du D2 et de l'obésité dans le monde industriel

# Proinflammatory diet

1. High temperature cooking
2. Charred, charcoaled food on barbecue for instance
3. High insulinic foods (high GI and high GL)
4. Low raw/cooked ratio
5. High intake of refined and/or industrial foods
6. High W6/W3 ratio
7. High intake of W6 PUFA (Sunflower, corn, soy oils)

# Does Red Meat Cause Inflammation? It just depends on you...

- Grassfed beef or lamb
- Raw meat (or tartare)
- Gently cooked (Gigot de sept heures)
- Rich in W3 PUFA (preserved)
- Rich in lipoic acid
- Grainfed beef or lamb
- High temperature cooking
- Richer in W6 PUFA
- Low content in lipoic acid

# Résultats

- Tartare de veau Rosée des Pyrénées (viande d'estive)
- Viande de bœuf aux farines de céréales cuite à haute température



# Alimentation anti-inflammatoire

1. Aliments entiers et crus
2. Cuissons basse température ou vapeur
3. Aliments peu insulinosécréteurs (IG bas & CG basse)
4. Alimentation riche en micronutriments
5. Omega 3 longue chaîne
6. Rapport oméga 6/oméga 3 bas
7. Epices, herbes, aromates et autres micronutriments (Curcuma, Ail, acide lipoïque...)

# Epices et herbes



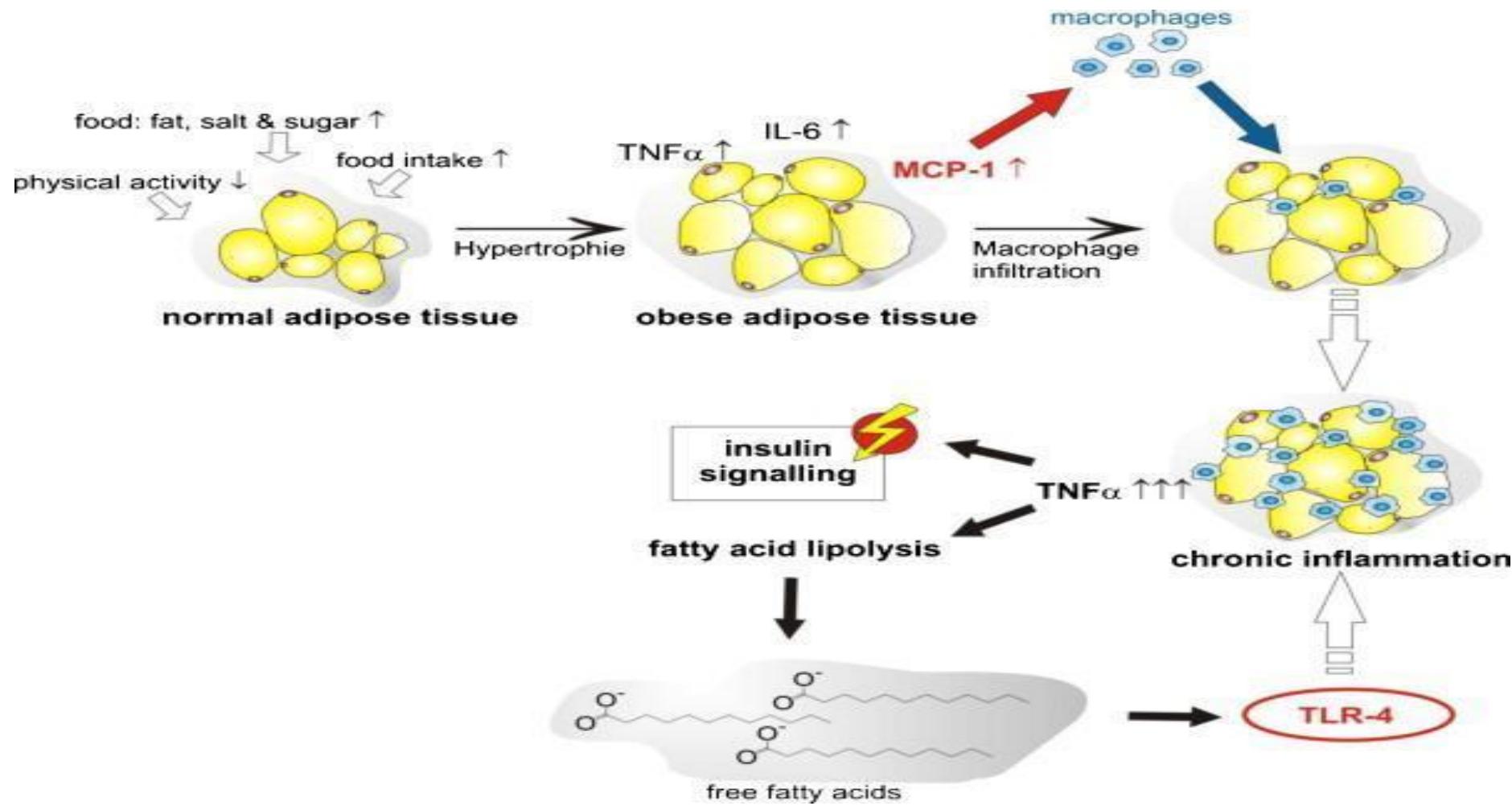
# Boissons anti-inflammatoires

- Eaux
- Jus de légumes
- Thé vert
- Kéfir de fruit ou de lait cru (non pasteurisé)
- Vins
- Bières (Non filtrées et non pasteurisées)

# Jus de légumes

- Préparation du jus de légumes
- Résultat





Alois Jungbauer , Svjetlana Medjakovic

#### **Anti-inflammatory properties of culinary herbs and spices that ameliorate the effects of metabolic syndrome**

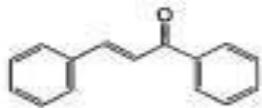
Maturitas Volume 71, Issue 3 2012 227 - 239

<http://dx.doi.org/10.1016/j.maturitas.2011.12.009>

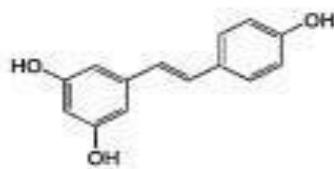
Fig. 1 Schematic overview of how metabolic syndrome develops from adipose tissue [7]

# POLYPHENOLS

## Chalcones



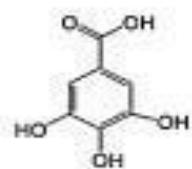
## Stilbenes



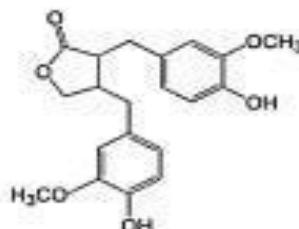
## Flavonoids



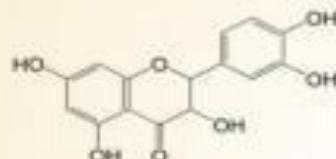
## Phenolic acids



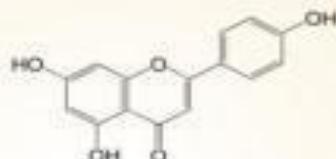
## Lignans



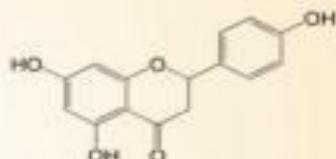
### Flavanonols



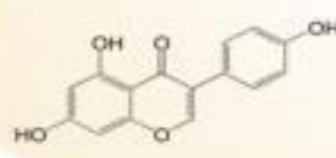
### Flavones



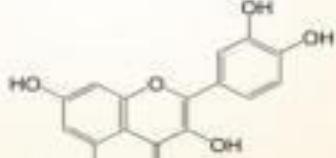
### Flavanones



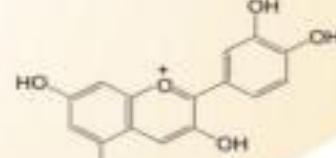
### Isoflavonoids



### Flavonols



### Anthocyanidins



Alois Jungbauer , Svetlana Medjakovic

**Anti-inflammatory properties of culinary herbs and spices that ameliorate the effects of metabolic syndrome**

Maturitas Volume 71, Issue 3 2012 227 - 239

<http://dx.doi.org/10.1016/j.maturitas.2011.12.009>

Fig. 2 Overview of the different classes of polyphenols, though some phenolic acids are not strictly polyphenols. Examples for the different classes are given, being matairesinol as lignan, gallic acid as phenolic acid, *trans*...

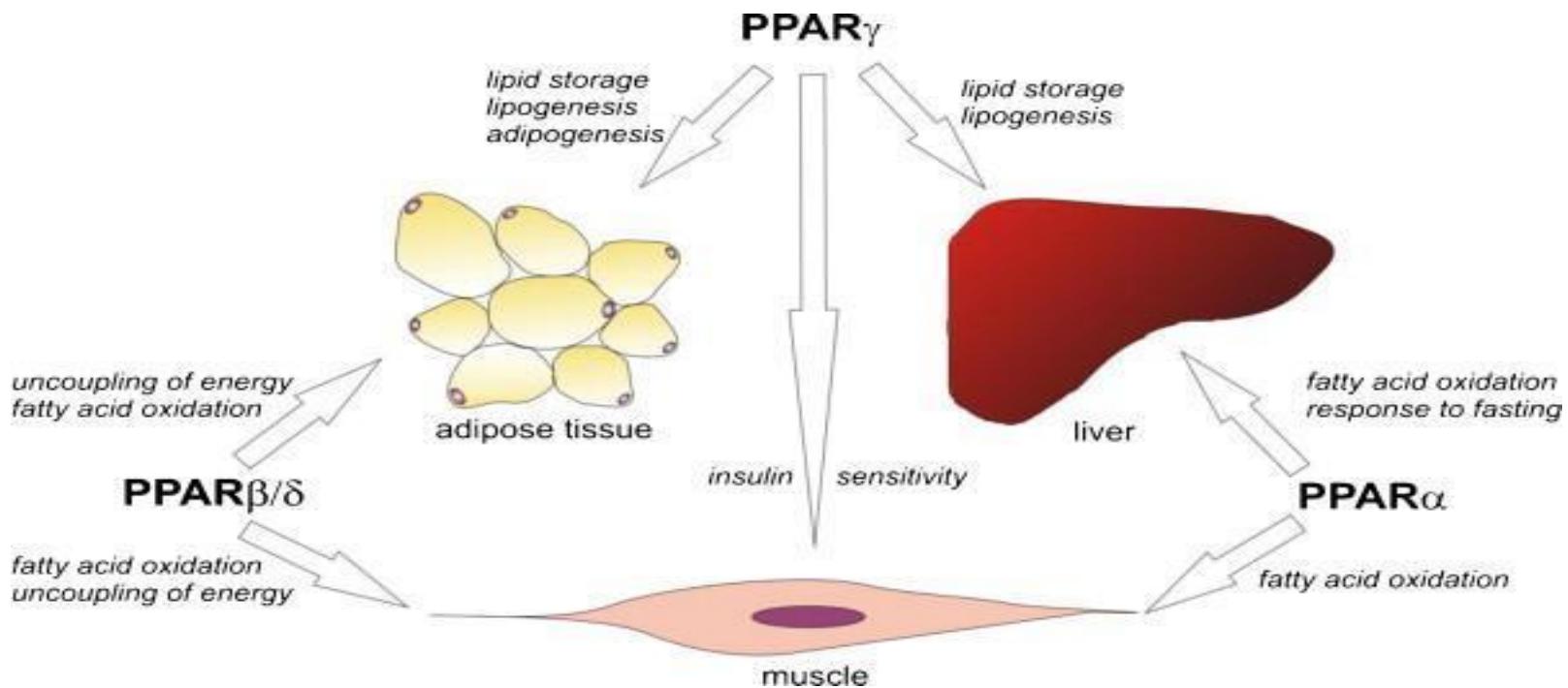


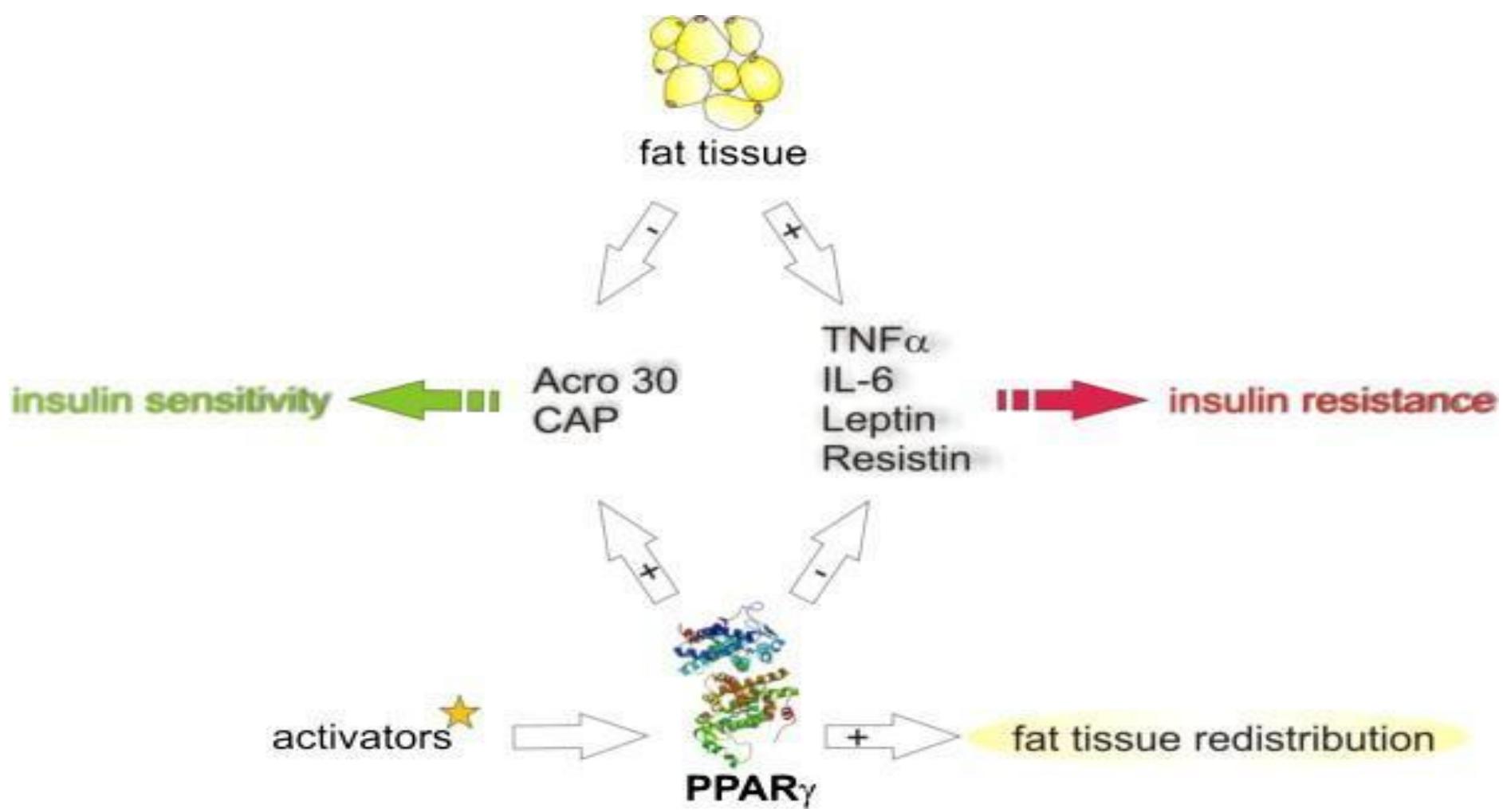
Fig. 3 Overview of how PPARs are involved in fat storage, lipogenesis, and adipogenesis.

Alois Jungbauer , Svjetlana Medjakovic

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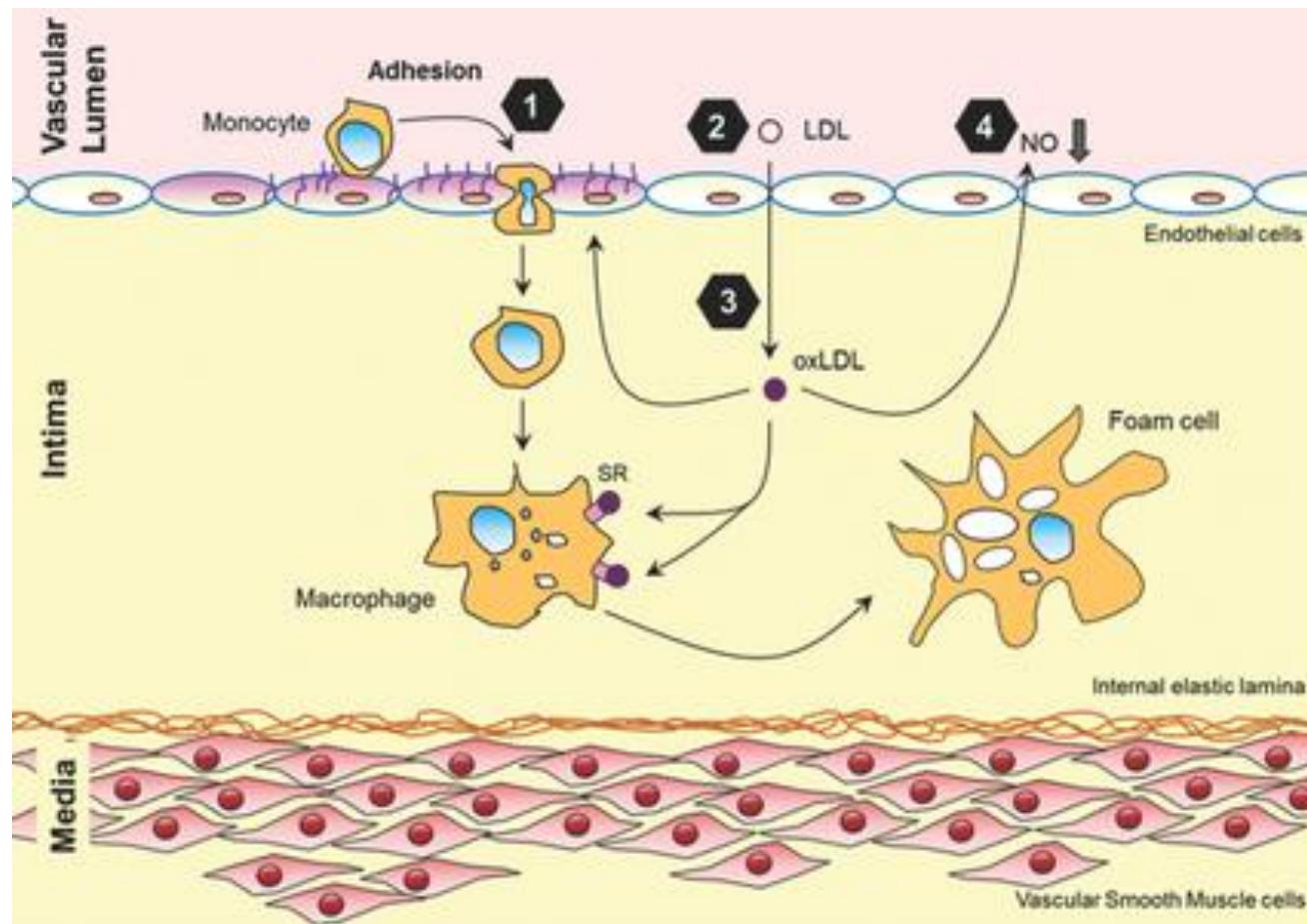
#### Anti-inflammatory properties of culinary herbs and spices that ameliorate the effects of metabolic syndrome

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Fig. 4 Effects induced by PPAR $\gamma$  that affect fat tissue remodelling or insulin resistance or sensitivity.

## Dietary oats and modulation of atherogenic pathways



### 1. Anti-inflammatory effects

Oat polyphenols reduce the amount of adhesion molecules on the endothelial surface, resulting in less monocyte recruitment to the vascular wall.

### 2. Reduction of plasma cholesterol

The cholesterol-lowering effect of oats reduce the chance of LDL being trapped in the vascular wall.

### 3. Antioxidative capacity

Oat antioxidants prevent oxidation of LDL and scavenge other reactive oxygen species.

### 4. Maintained endothelial function

Oats may preserve nitric oxide production and thereby counteract endothelial dysfunction.

# Conclusion

- L'inflammation est une adaptation de survie aux agressions en particulier des micro-organismes
- Malgré la disparition du péril infectieux nous restons très enclin à l'inflammation et tout particulièrement chronique
- En effet le mode de vie sédentaire et l'alimentation industrielle sont pro-inflammatoires
- En prévention primaire ou secondaire un régime anti-inflammatoire est utile dans les maladies chroniques
- Sous réserve d'une bonne compréhension les régimes crétois, japonais, paléo sont anti-inflammatoires

# Mon blog préféré:

<http://dysnutrition.blogspot.fr/>

Merci de votre attention