

Dr. István Gera

Periorodtal pocket as a risk factor
in the etiology of systemic
diseases

*Semmelweis University,
Department of
Periodontology, Budapest*

PERIODONTITIS

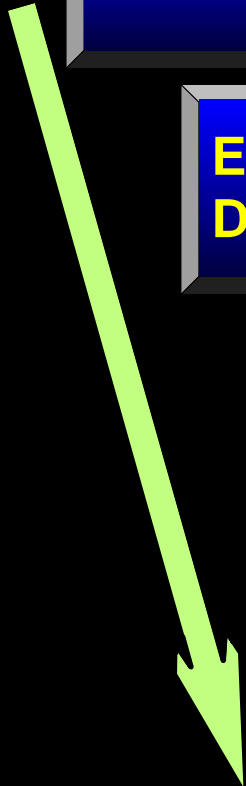
PATHOGENIC BACTERIA

ENDOTOXIN, TOXINS, MEMBRANE
DEGRADATION MATERIALS

INFLAMMATORY CASCADE

INFLAMMATORY MEDIATORS
PGE, TNF, IL

CONNECTIVE TISSUE AND BONE
DAMAGE



PERIODONTITIS AND SYSTEMIC DISEASES

PRETERM BIRTH

CORONARY HEART DISEASE

THROMBOEMBOLIC DISEASES

MYOCARDIAL INFARCT
STROKE

TYPE I DIABETES

TYPE II DIABETES

Pregnancy effects



**THE CONNECTION BETWEEN PREGNANCY
AND PERIODONTAL-GINGIVAL DISEASES HAS
LONG BEEN KNOWN**

**THE EFFECT OF PREGNANCY
*-PREGNANCY GINGIVITIS -***

**BUT THE CONNECTION BETWEEN THE
PREGNANT WOMENS' PERIODONTAL
CONDITION AND THE BIRTH WEIGHT OR
THE INCIDENCE OF PREMATURE DELIVERY
HAS JUST RECENTLY BEEN STUDIED**





Galloway as early as 1931 indicated that Gram negative bacteria from the periodontal pocket might have negative effects on pregnancy outcome

The periodontal state of mothers with preterm low birth weight (PLBW) and age matched controls

THE WORTH PERIODONTAL STATE	NORMAL BIRTH BETWEEN THE AGE 16-44	PRETERM LOW BIRTH WEIGHT AGE 16-44
HEALTHY PERIODONTIUM	27	0
MILD GINGIVITIS	14	1
SEVERE GINGIVITIS	30	12
SHALLOW POCKET	26	38
DEEP POCKET	3	49

The East London Study of Maternal Chronic Periodontal Disease and Preterm Low Birth Weight Infants: Study Design and Prevalence Data by *Davenport E.S. & Co. Annales of Periodontology 3: 213-221 1997*

The periodontal state of women with preterm low birth weight (PLBW) and age matched controls WHO CPITN INDEX

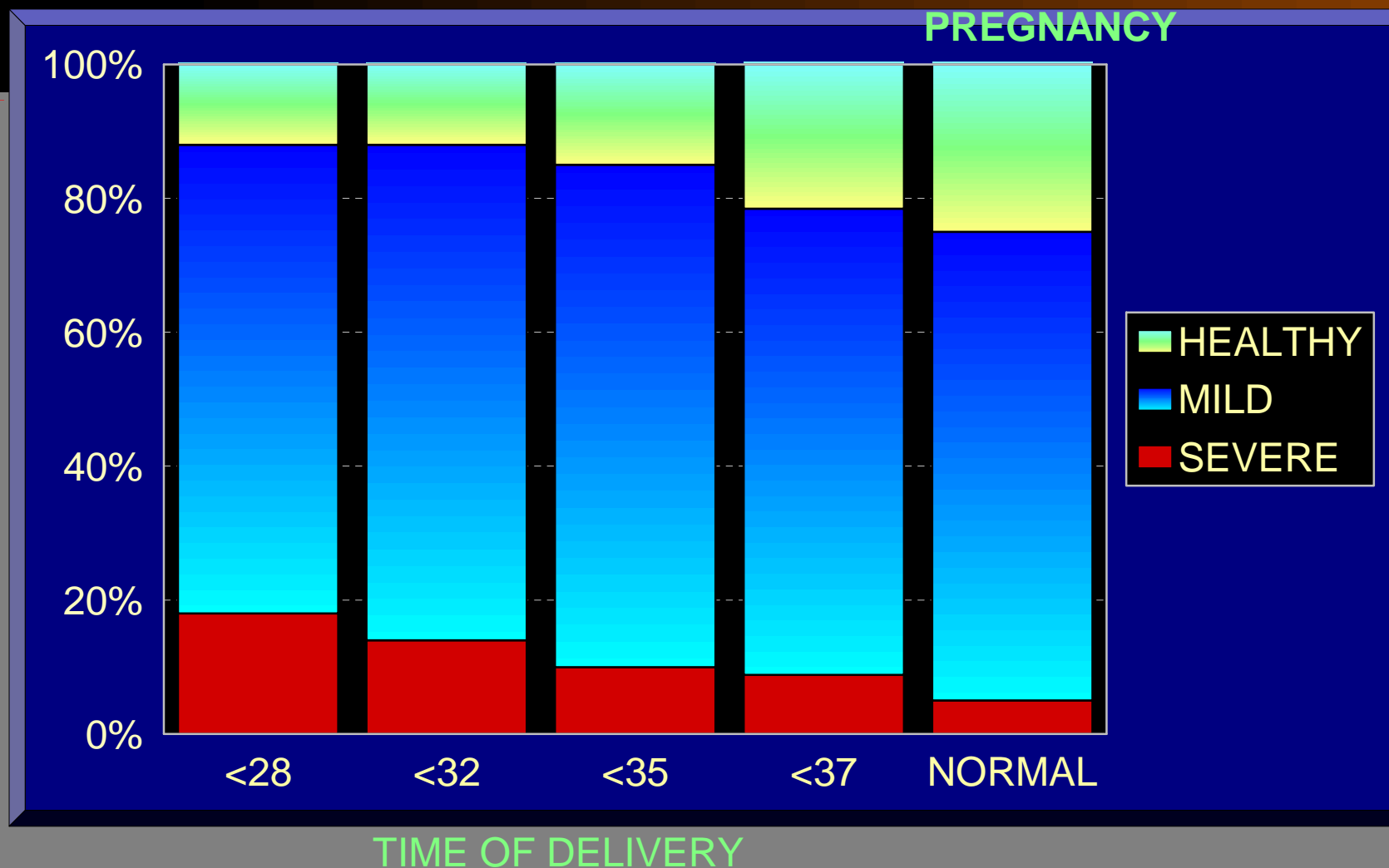
PLBW vs. NORMAL CONTROLS (THAILAND)

CPITN SCORES	PRTERM	CONTROL	P value
CPITN 0	0,4	1,1	0,001
CPITN 1	5,6	4,9	0,001
CPITN 2	4	3,2	0,01
CPITN 3-4	0,5	0,2	0,1
DMFT	4,8	3,4	0,17

Poor periodontal health of the pregnant woman as a risk factor for low birth weight Dasanayake A.P. Annals of Periodontology 3: 206-212 1997.

GESTATIONAL AGE AND PERIODONTITIS

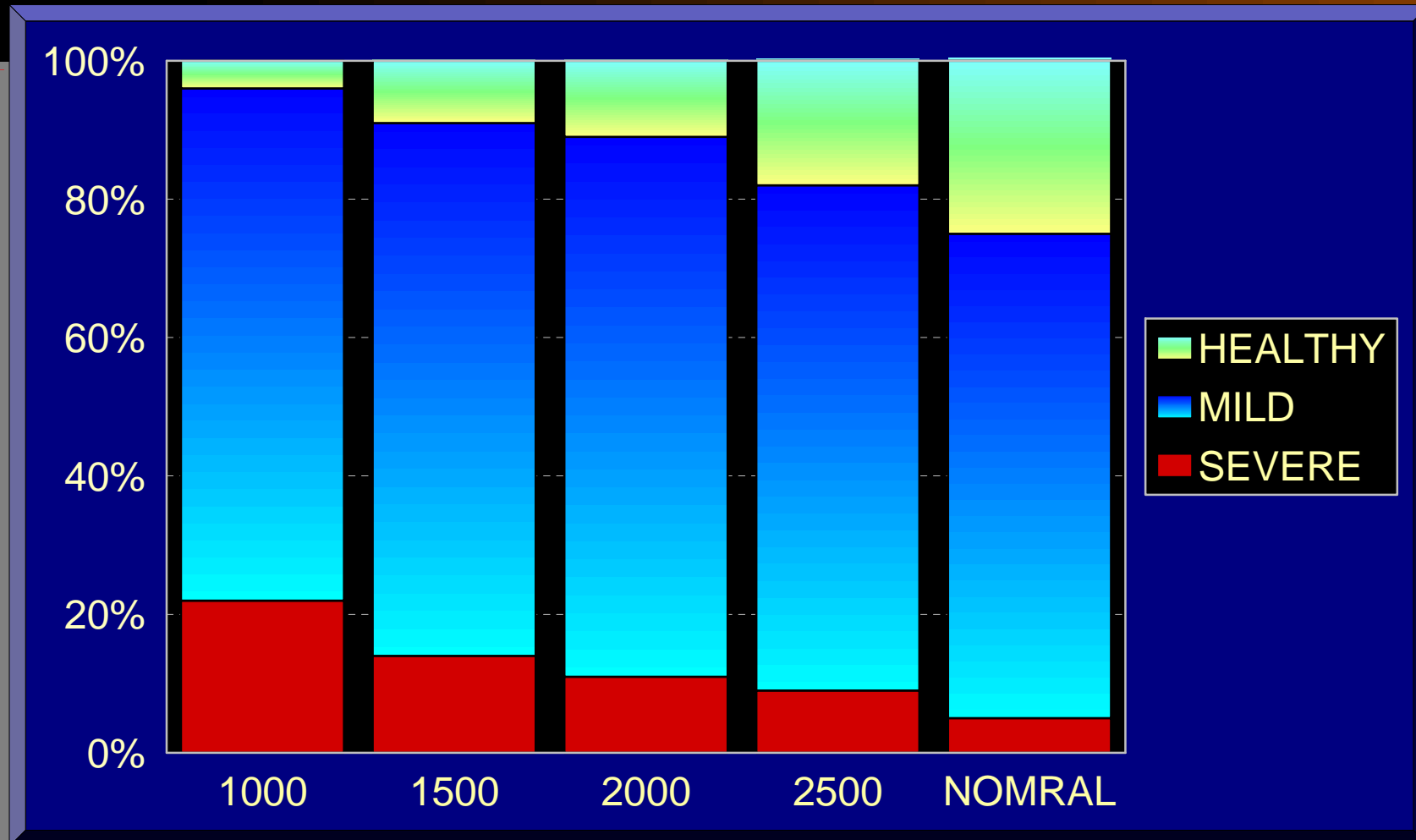
1300 EXAMINED
PREGNANCY



Offenbacher S. et al: Maternal periodontitis and Prematurity. Part I: Obstetric outcome of prematurity and growth restriction *Ann Periodontol* 2001;6:164-174

BIRTH WEIGHT AND PERIODONTITIS

1300 EXAMINED
PREGNANCY

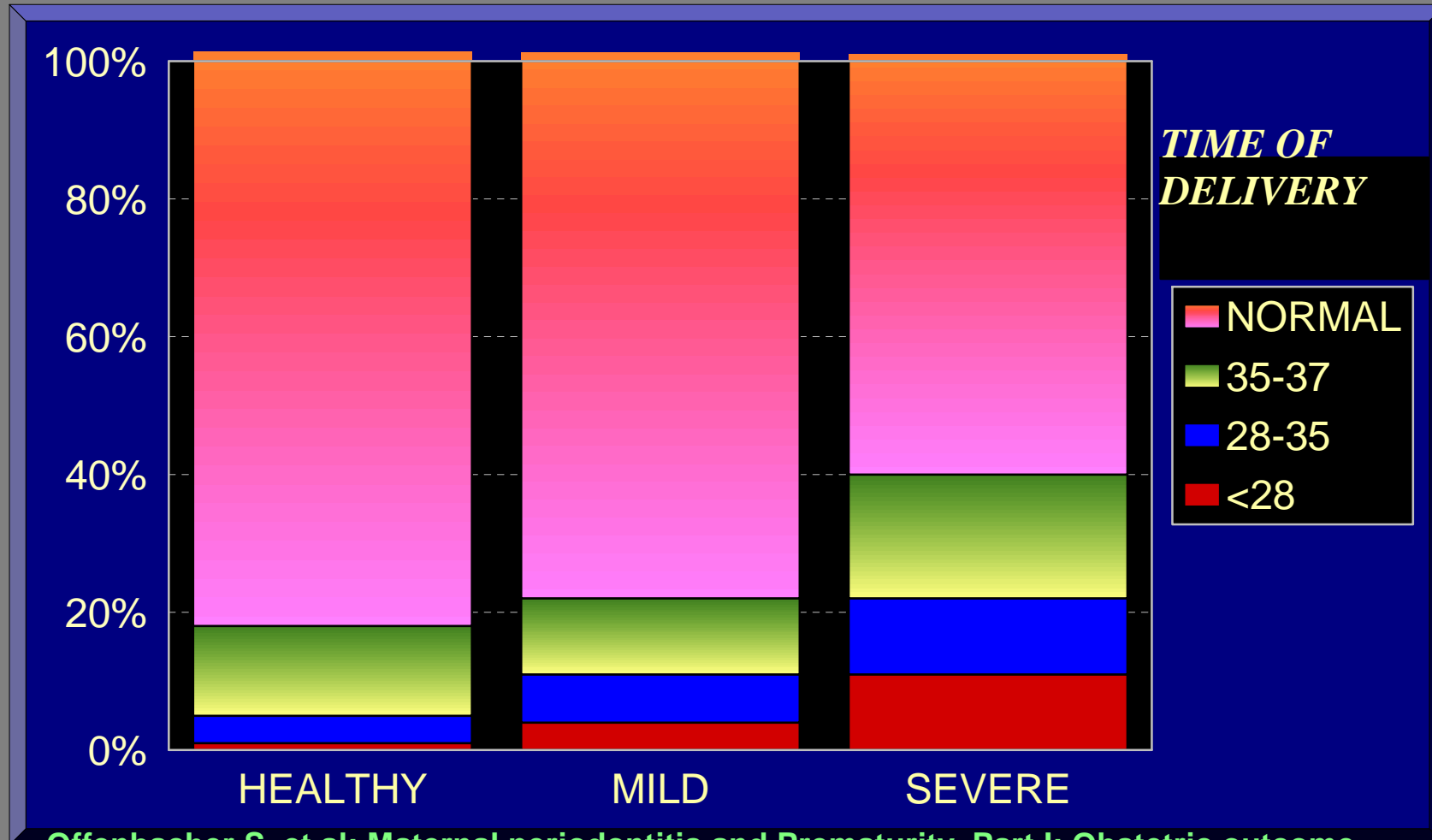


BIRTH WEIGHT (gramS)

Offenbacher S. et al: Maternal periodontitis and Prematurity. Part I: Obstetric outcome of prematurity and growth restriction *Ann Periodontol* 2001;6:164-174

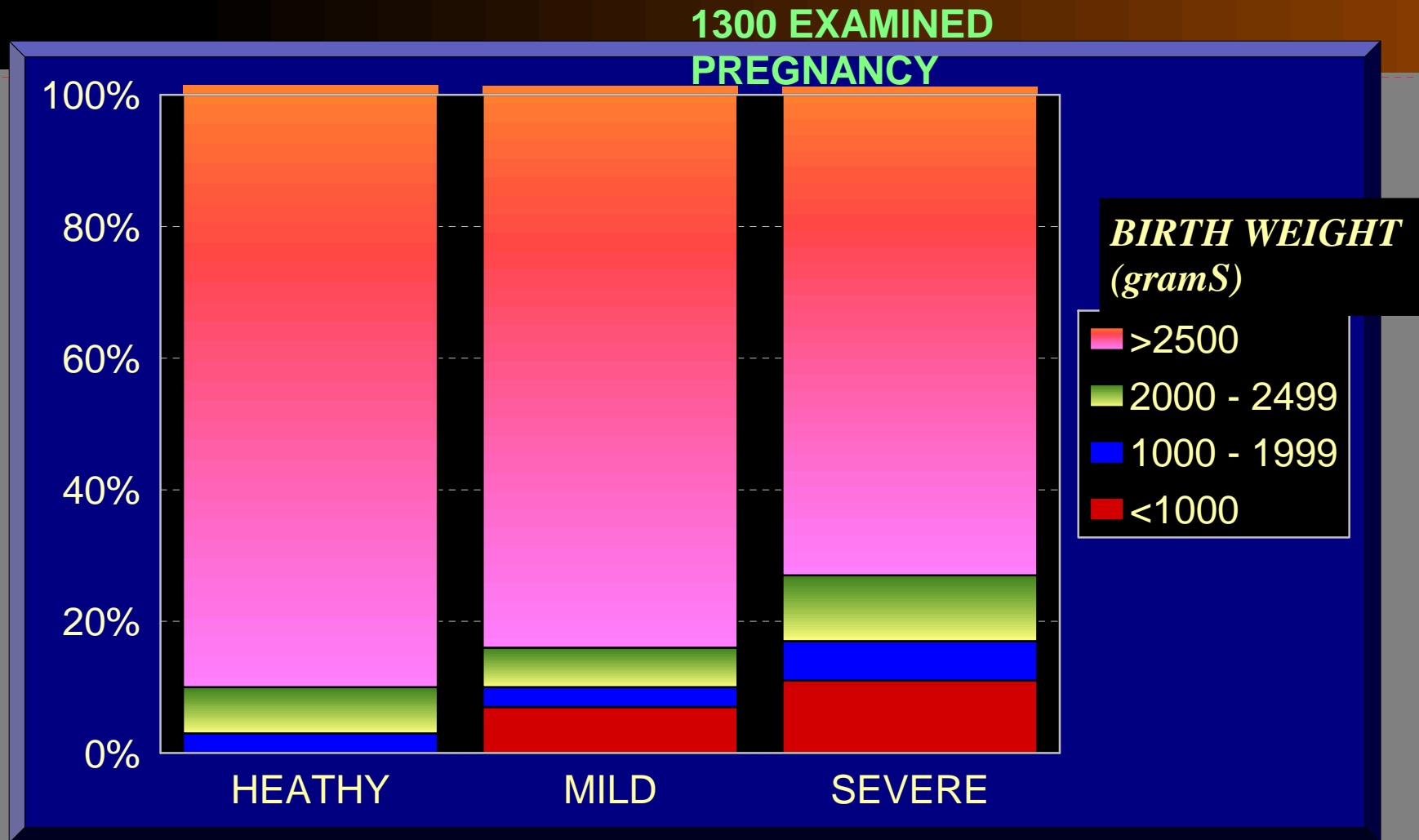
MATERNAL ANTEPARTUM AND PERIODONTAL STATUS

1300 EXAMINED PREGNANCY



Offenbacher S. et al: Maternal periodontitis and Prematurity. Part I: Obstetric outcome of prematurity and growth restriction *Ann Periodontol* 2001;6:164-174

MATERNAL ANTEPARTUM PERIODONTAL STATUS



Offenbacher S. et al: Maternal periodontitis and Prematurity. Part I: Obstetric outcome of prematurity and growth restriction *Ann Periodontol* 2001;6:164-174

The periodontopathogenic microorganisms in the gingival sulcus of mothers with preterm low birth weight (PLBW) and age matched controls

BIRTH WEIGHT	<i>B. forsythus</i>	<i>P. gingivalis</i>	AA.	<i>T. denticola</i>
NORMAL	0,10 \pm 0,11	0,0 \pm 0,0	0,0 \pm 0,0	0,30 \pm 0,16
PLBW	0,96 \pm 0,15	0,26 \pm 0,09	0,15 \pm 0,07	1,67 \pm 0,24

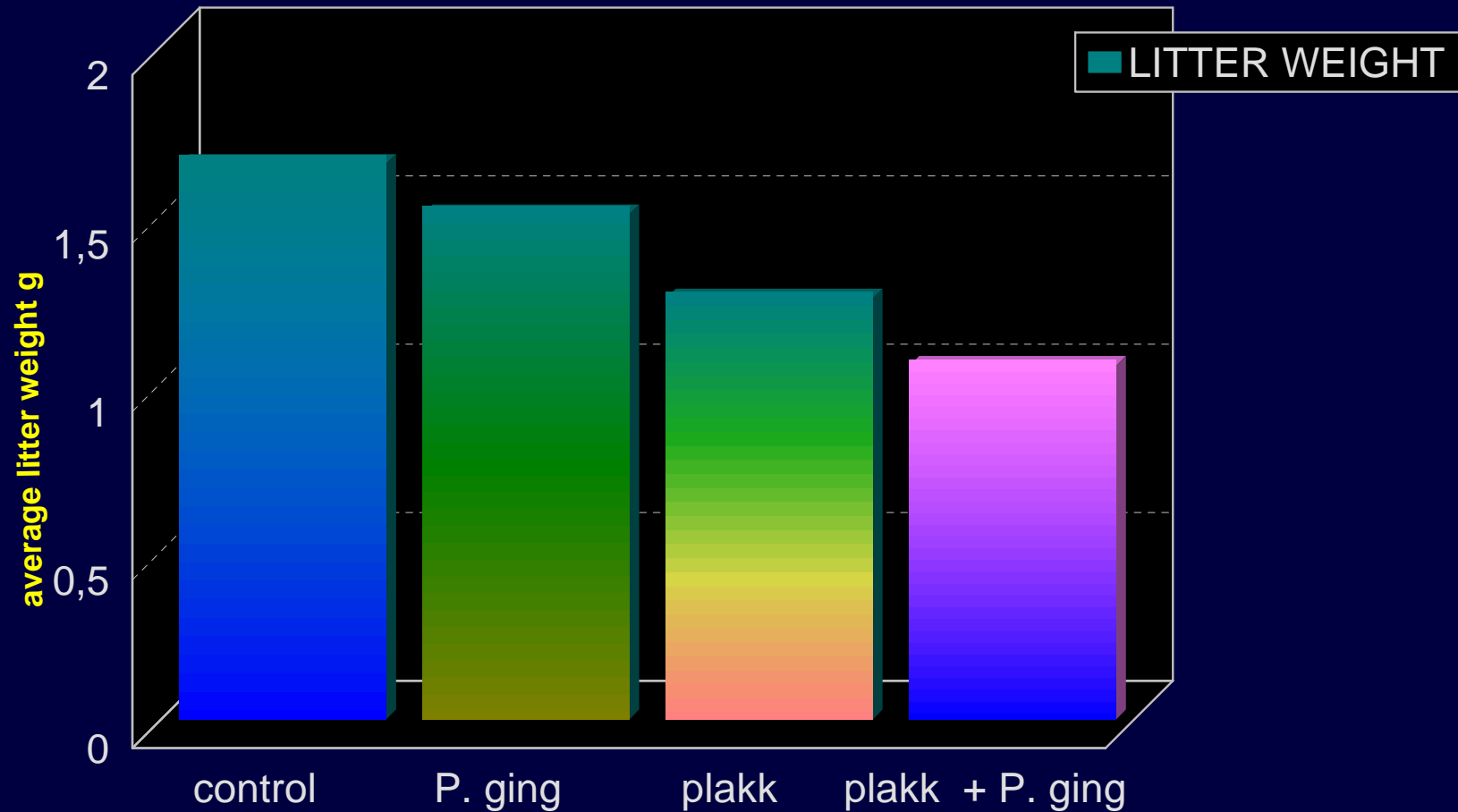
Potential pathogenic mechanisms of periodontitis associated pregnancy complications
 Offenbacher S. et al. *Annales of Periodontology* 3: 233-250 1997

PRETERM LOW BIRTH WEIGHT MOTHER'S AND THE CONTROL MOTHER'S GINGIVAL SULCUS PGE2 és IL-1B CONCENTRATION

BIRTH WEIGHT	SULCUS PGE2 ng/ml + SE	SULCUS IL-1B ng/ml + SE
NORMAL	62,6 ± 10,2	720 ± 105,2
PLBW	131,4 ± 21,8	1217 ± 281,3

Potential pathogenic mechanisms of periodontitis associated pregnancy complications
Offenbacher S. et al. *Annales of Periodontology* 3: 233-250 1997

THE EFFECT OF EXPERIMENTAL PERIODONTITIS ON THE AVERAGE LITTER WEIGHT OF GOLDEN HAMSTERS



Effects of E. Coli and P. gingivalis lipopolysaccharide on pregnancy outcome in golden hamster by Collins & al. *Infect Immun* 1994; 62: 4652-4655

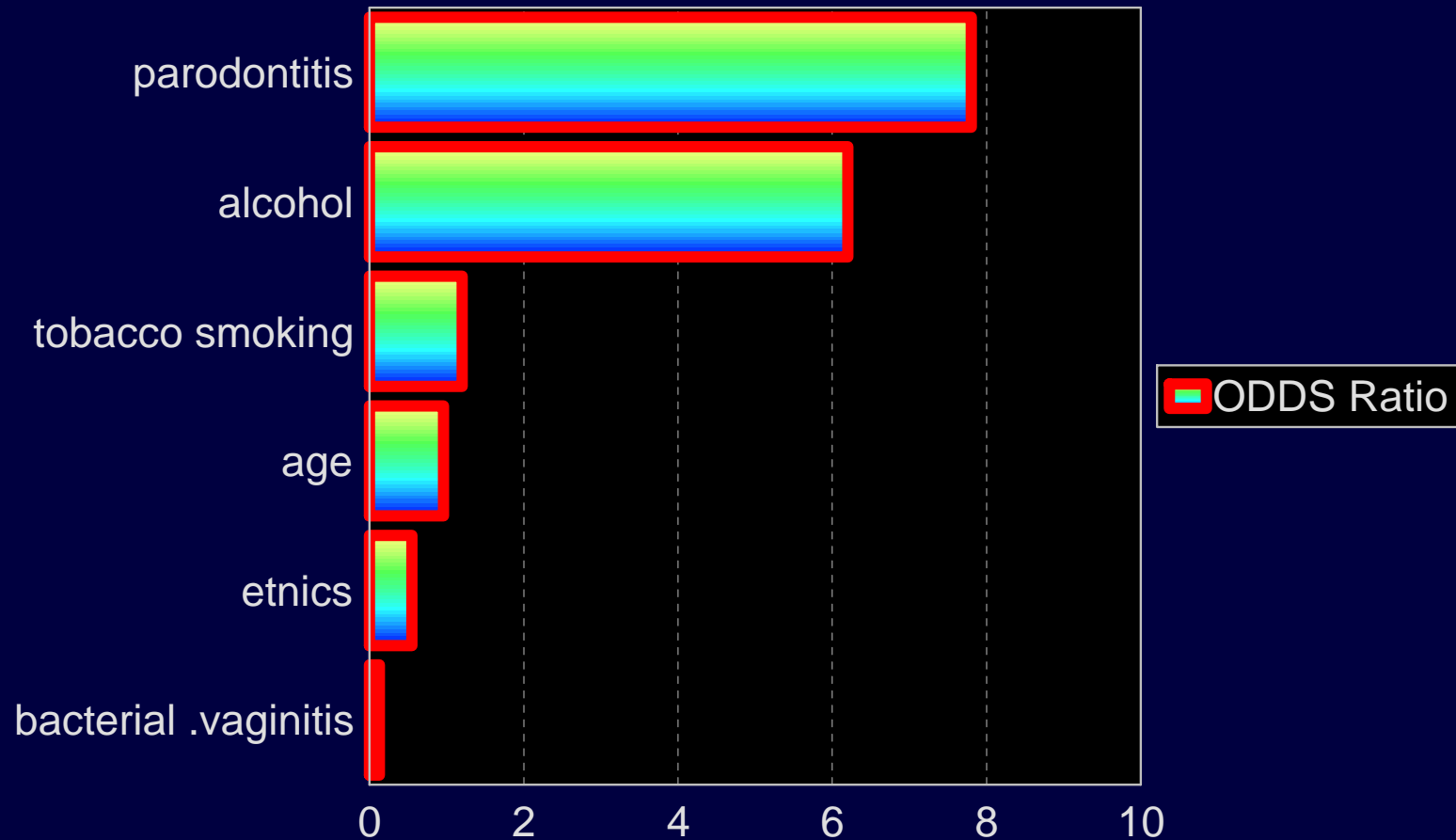
THE PERIODONTAL TREATMENT'S EFFECT ON THE INCIDENCE OF PLBW

*390 pregnant women - 195 treated before week 22
195 non treated*

	TREATED	NON TREATED
PLBW	1,8%	10,2%
NORMAL	98,2%	89,2%

Lopez NJ et al.: Periodontal therapy reduces the risk of preterm low birth weight *J Dent Res.* 2001;80:188 (Abstr-1223)

DIFFERENT RISK FACTORS IN THE ETIOLOGY OF PRETERM LOW BIRTH WEIGHT



Periodontal disease as a possible risk factor for preterm low birth weight

Offenbacher et al. J. Periodontol 1996; (suppl) 67: 1003-1113

conclusion

sulcus PGE2 level is significantly higher in PLBW mothers than in normal controls

There is a negative correlation between the sulcus PGE2 level and the birth weight of the newborn babies

Bacterioides forsythus, Porphyromonas gingivalis, Actinobacillus actinomycetemcomitans and Treponema denticola were much more frequently occurring in the gingival sulcus of PLBW mothers than in normal controls

During normal pregnancy the amniotic PGE concentration is steadily increasing till reaches the threshold that will initiate the labor and finally the delivery

The ascending urogenital infection will locally increase the amniotic PGE concentration that can lead to preterm labor

***F. nucleatum* can frequently be detected from amniotic fluid and that seldom originates from the vagina .**

Many times the vaginal bacterial culture is negative while the amniotic culture is positive for *F. nucleatum*

As F. nucleatum is the most common member of the periodontal pocket it can be assumed the those bacteria can hematologically spread and infect the amnion .

The blood samples taken from 60 new born babies' umbilical cord 23,3% was positive for anti *C.rectus*, *P.gingivalis*, *E.corrodens*, *A.a.* and *Veilonella parvula* IgM antibodies

As maternal IgM cannot pas the placental barrier, this antibody should originate from the fetus produced against the bacteria or bacterial antigens infecting the fetus

Pregnant women suffering with periodontitis might produce high antibody titer against *Porphyromonas gingivalis*, *T. forsythia*, *Prevotella intermedia* originating from periodontal pockets. Others produce low antibody against periodontopathogenic bacteria – in those women the bacterial infection can easily spread

In those cases antibodies against maternal periodontopathogenic bacteria can be detected from the umbilical cord of the fetus.

In those pregnant women who does not produce high anti periodontopathogenic antibodies the incidence of preterm low weight birth was 66.7%-

The putative pathomechanism of preterm low weight birth

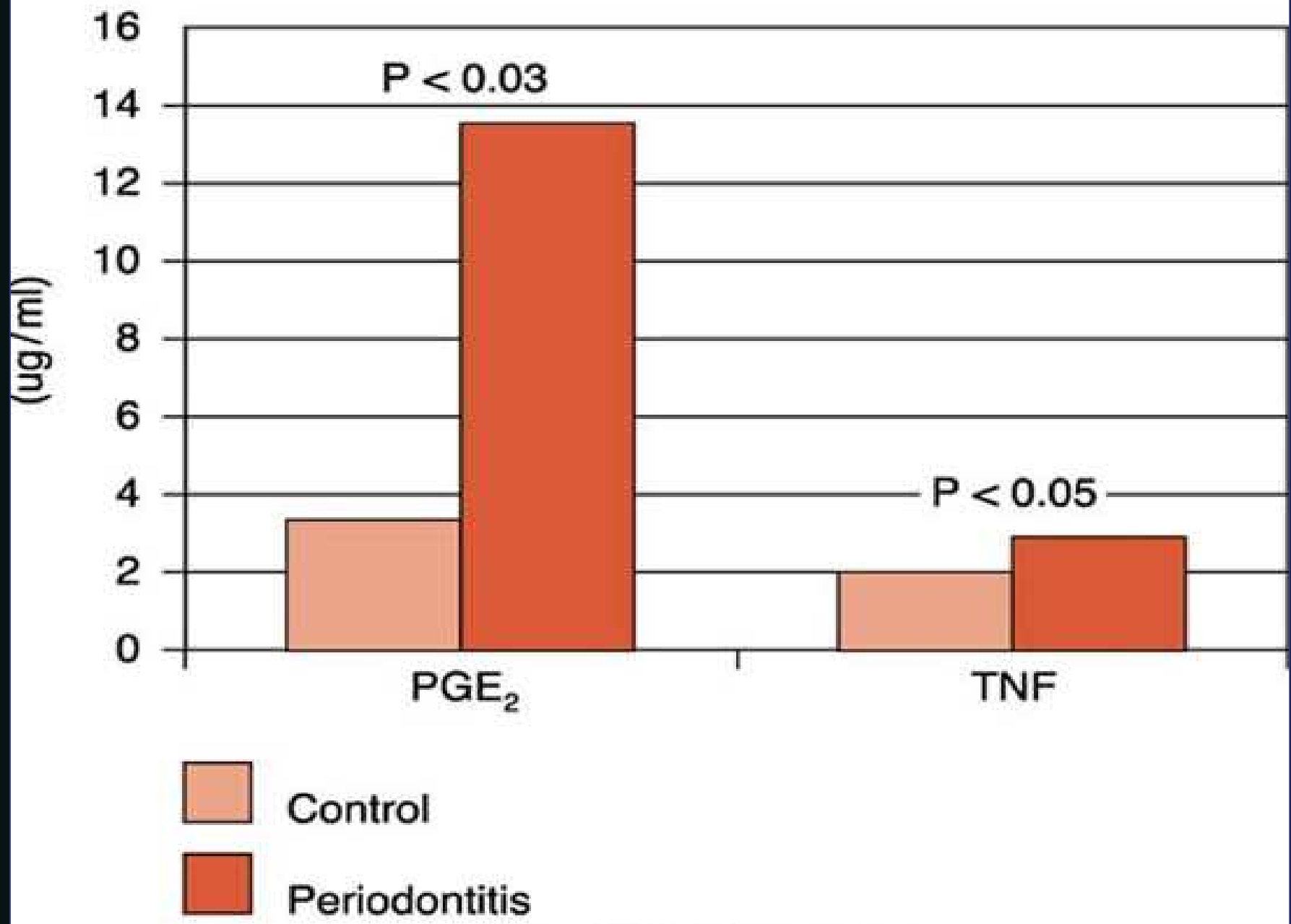
Inflammatory cytokines (PGE2, TNF α , IL-1 IL- 6) can enter into the systemic blood circulation and get into the womb and the placenta .

Cytokines will enhance the local PGE production in the placenta

If the local PGE2 level reaches a certain threshold the labor will start independent of the origin of the high PGE concentration

The LPS can also increase the local PGE production in the womb-

Amniotic Fluid Levels in Experimental Periodontitis



Bacterial
infection

Bacteria & products in amnion

Inflammatory response with
cytokine production in amnion

Increased amniotic
prostaglandin production

Preterm labor



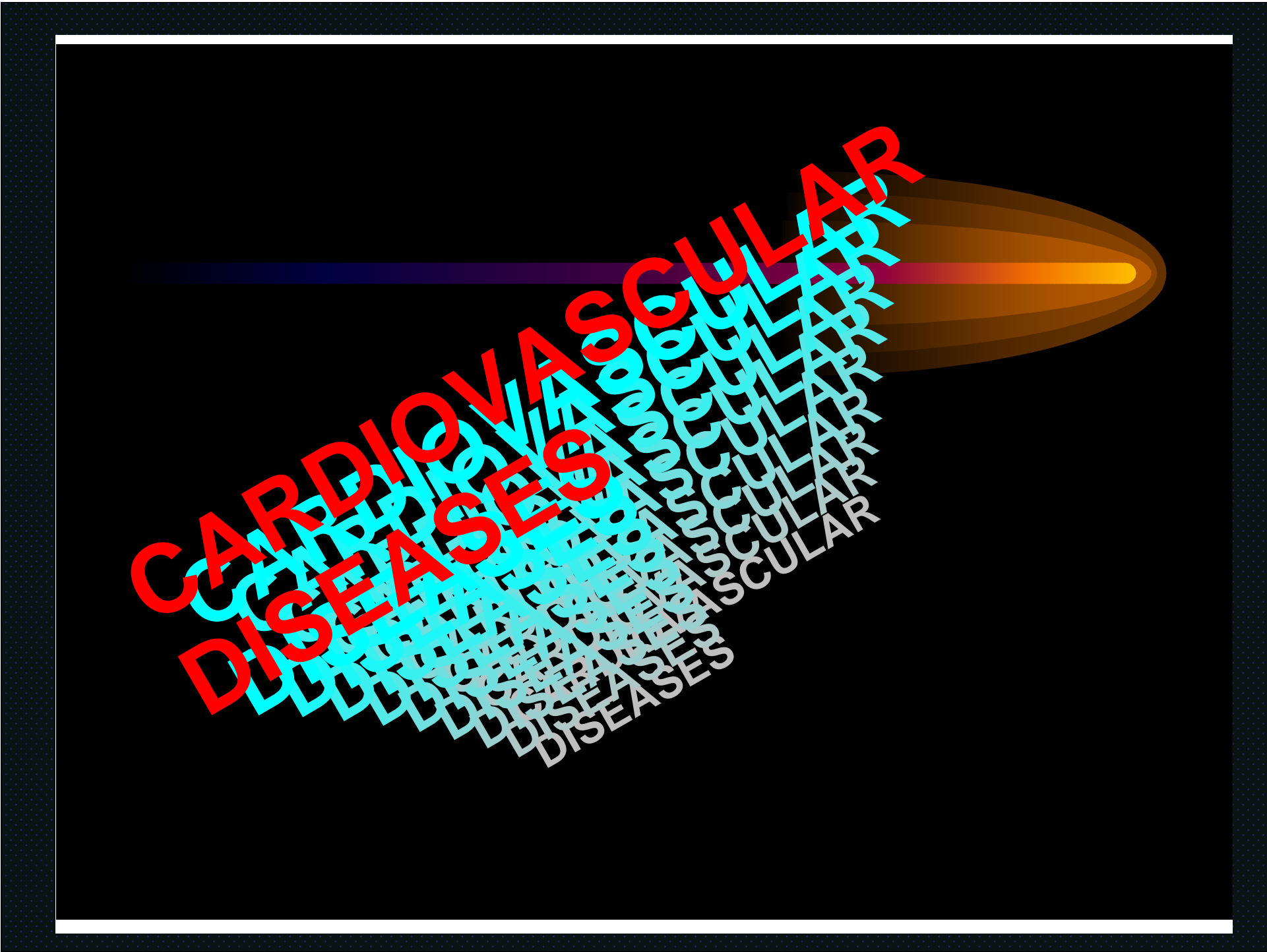
CONCLUSION



**THE POOR PERIODONTAL
CONDITION OF PREGNANT WOMEN
IS AN INDEPENDENT RISK FACTOR
FOR PRETERM LOW BIRTH
WEIGHT**

(PLBW - < 2500 g, < 37 week)

Poor periodontal health of the pregnant woman as a risk factor for low birth weight Dasanayake A.P. Annals of Periodontology 3: 206-212 1997.



**CARDIOVASCULAR
DISEASES**

The image is a graphic design on a black background. It features a word cloud of the words 'CARDIOVASCULAR' and 'DISEASES'. The top-most word, 'CARDIOVASCULAR', is in a large, bold, red font. Below it, 'DISEASES' is also in a large, bold, red font. The rest of the words are in a smaller, cyan or light blue font, and they are arranged in a way that they appear to be falling or trailing away from the top left towards the bottom right. The words are repeated multiple times in various orientations. In the background, there is a stylized light effect that looks like a beam of light or a lens flare, with a gradient from purple to orange and yellow, pointing towards the right. The entire graphic is framed by a thin white border, which is itself set within a larger black frame with a fine dotted pattern.

CLASSIC DENTAL FOCAL THEORY

DENTAL FOCUS
PERIODONTAL POCKET IS
NOT A FOCUS !!!!

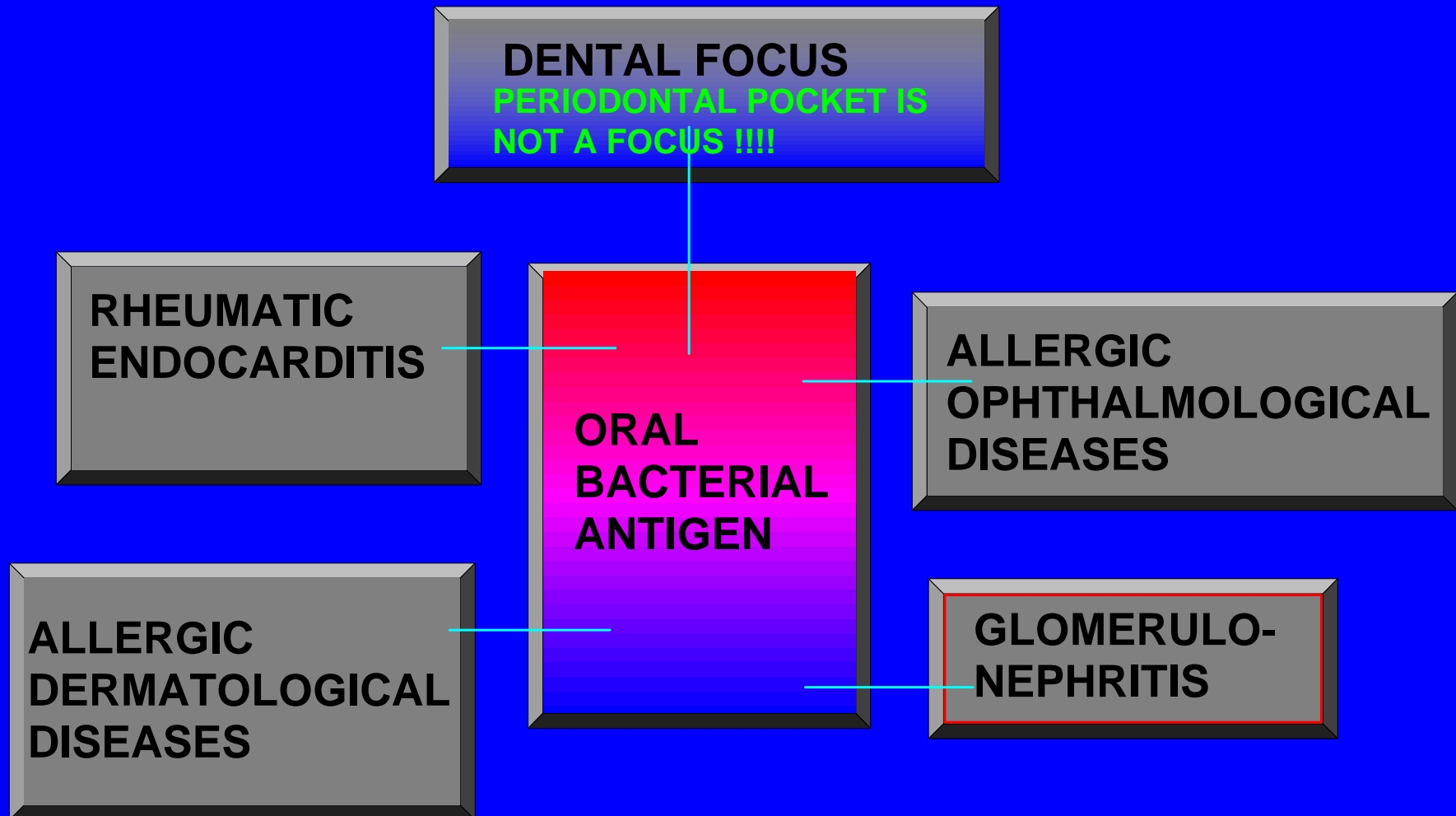
RHEUMATIC
ENDOCARDITIS

ALLERGIC
OPHTHALMOLOGICAL
DISEASES

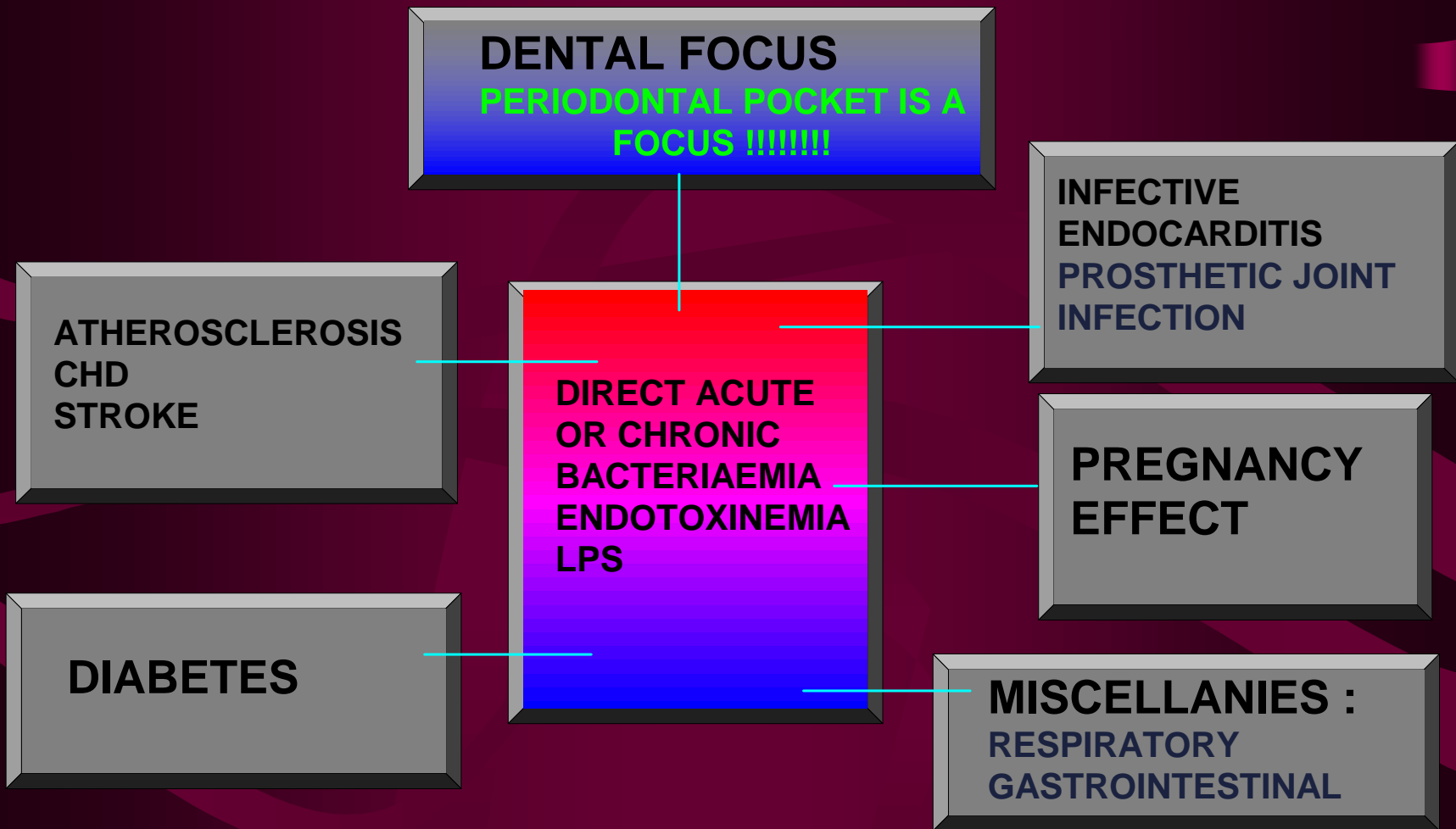
ORAL
BACTERIAL
ANTIGEN

ALLERGIC
DERMATOLOGICAL
DISEASES

GLOMERULO-
NEPHRITIS



PRESENT FOCAL THEORY



Position Paper : Periodontal disease as a potential risk factor for systemic diseases J Periodontol 1998; 69: 841-850.

- **Mattila KJ, Valle MS, Neiminen MS, Valtonen VV, Hietantemi KL: Dental infections and coronary atherosclerosis. Atherosclerosis 103, 205, 1993.**
- **Mattila KJ, Nieminen MS, Valtonen VV. Association between dental health and acute myocardial infarction. Br Med J 1989; 298:779-782.**
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- **Syrjanen J, Peltola J, Valtonen V, Livanainen M, Kaste M, Huttunen JK: Dental infections in association with cerebral infarction in young and middle-aged men. J Intern Med 225, 179. 1989.**
- **Beck J, Garcia R, Heiss G, Vokonas P, Offenbacher S: Periodontal disease and cardiovascular disease .J Periodontol 1996; 67: 1123-1137.**
- **Grau AJ, Buggle F, Ziegler C és mts. Association between acute cerebrovascular ischemia and chronic and recurrent infections Stroke 1997; 28: 1724-1729.**
- **Paunio K, Impivaara O, Tiekso J, Maki J: Missing teeth and ischaemic heart disease in men aged 45- 64 years. Eur Heart J 14, 54. 1993**
- **Loesche WJ: Periodontal disease as a risk factor for heart disease. Compendium 25, 976. 1994.**

THE CORRELATION BETWEEN THE ATHEROSCLEROSIS (THE INCIDENCE OF CORONARY HEART DISEASES) AND PERIODONTAL CONDITION

RELATIVE ODDS RATIO

STUDY	CORRELATION	(ODDS RATIO)
Matilla -Finland	total dentition / heart attack	1,3
Matilla - Finland	total dentition / atherosclerosis	1,4
Matilla-Finland	total dentition / CHD incidence	1,2
DeStefano - USA	plaque, periodontium / lethal CHD	1,7
Beck - USA	periodontal bone level/new CHD	1,5
Beck - USA	periodontal bone level/ lethal CHD	1,9
Beck - USA	periodontal bone level/ stroke	2,7
Joshiyura - Japan	missing teeth / CHD	1,7


National Health and Nutrition Examination Survey

14 years follow-up study
10.000 participants

Conclusion:

subjects with severe periodontal condition had 25% more chance to develop ischemic heart disease than their age matched counterparts with normal periodontium

44 119 male subjects having no heart condition at baseline were followed up for 6 years



***THE RELETIVE ODDS RATIO WAS 1.4
HIGHER FOR THOSE WHO HAD 0-10
NATURAL TEETH AT BASELINE THAN
THOSE WHO HAD 25 HATURAL TEETH***

Normative Aging Study VA Hospitals Boston USA

1147 men, 18 years follow-up

207 developed ischemic heart disease

59 died of heart attack

40 men had stroke

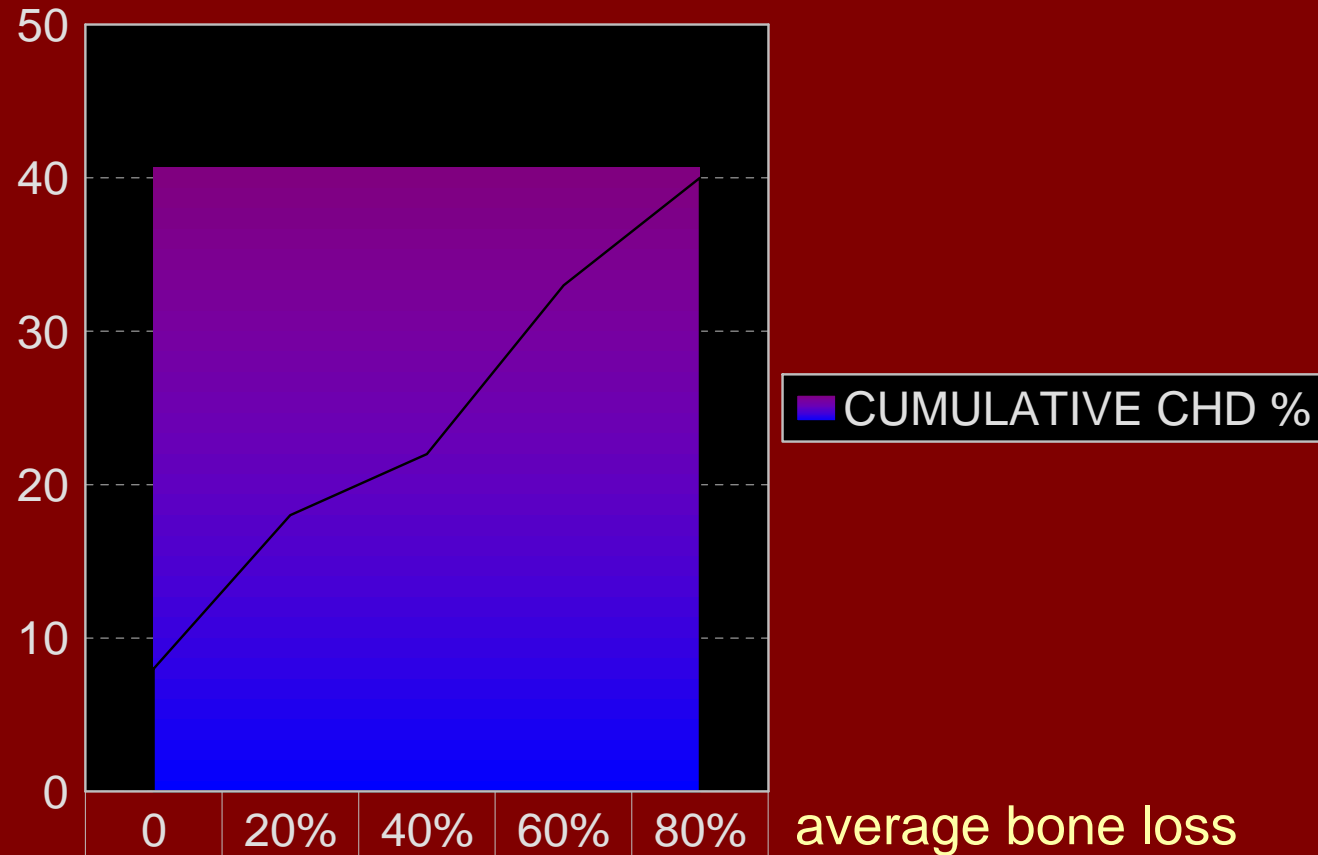
periodontitis as a relative risk factor

- coronary diseases ----- 1,5*
- lethal coronary diseases ----- 1.9*
- stroke ----- 2,8*

*Beck et al. Periodontal disease and cardiovascular disease J.
Periodontol 1996; 67(suppl) 1123-1137*

THE CORRELATION BETWEEN THE PERIODONTAL BONE LOSS AND THE CUMULATIVE PREVALENCE OF CHD

18 YEARS FOLLOW-UP

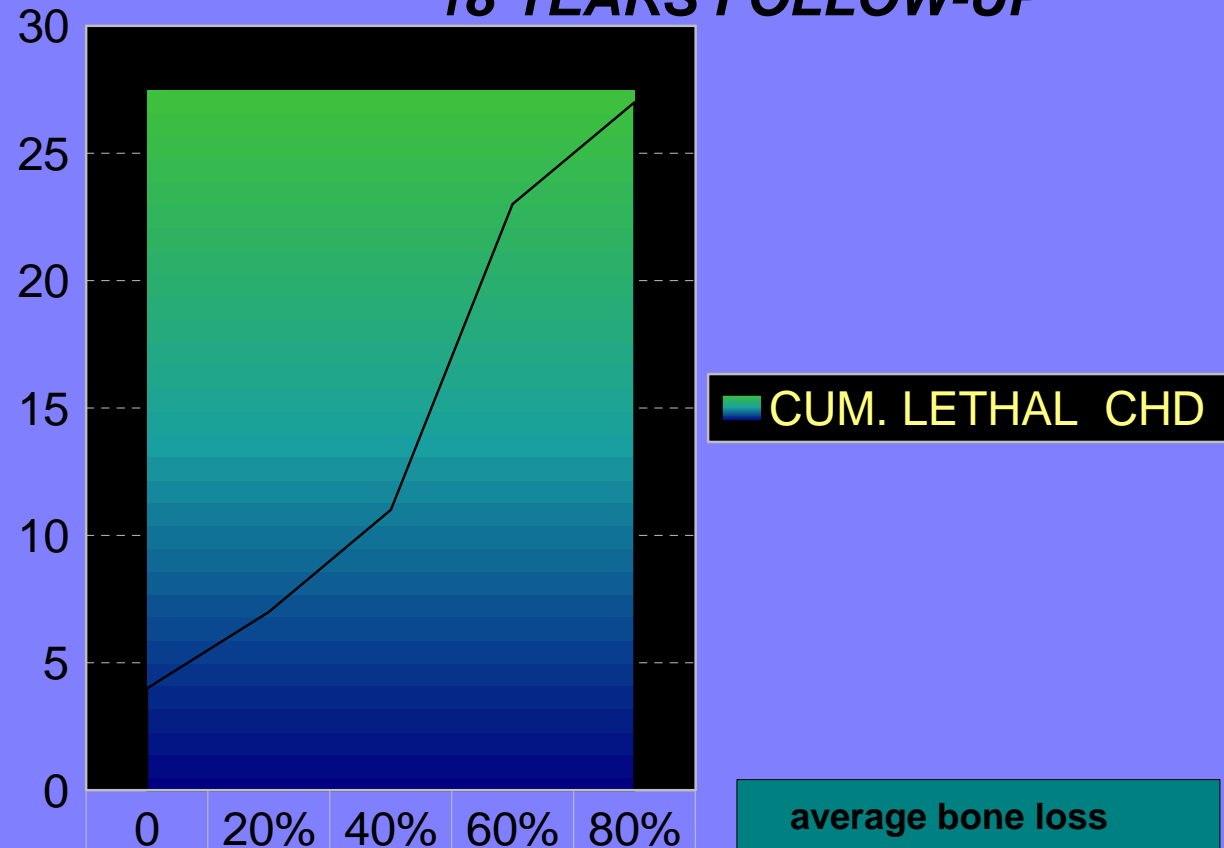


average bone loss	0	20%	40%	60%	80%
CUMULATIVE CHD %	8	18	22	33	40

Beck JD. et al.: Periodontal disease and cardiovascular disease *J. Periodontol* 1996;
67(suppl): 1123-1137

THE CORRELATION BETWEEN THE PERIODONTAL BONE LOSS AND THE CUMULATIVE PREVALENCE OF LETHAL CHD

18 YEARS FOLLOW-UP

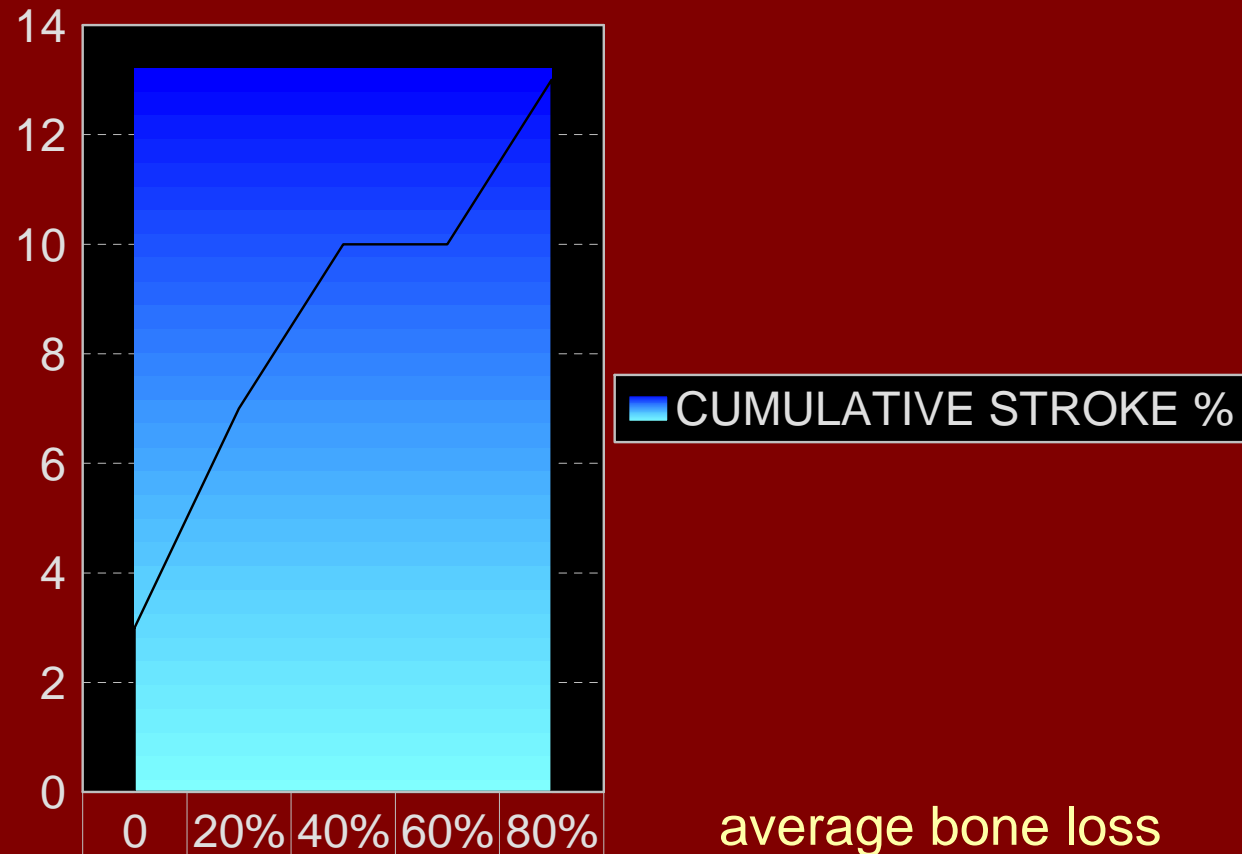


	0	20%	40%	60%	80%
CUM. LETHAL CHD	4	7	11	23	27

Beck JD. et al.: Periodontal disease and cardiovascular disease J. Periodontol 1996; 67(suppl): 1123-1137

THE CORRELATION BETWEEN THE PERIODONTAL BONE LOSS AND THE CUMULATIVE PREVALENCE OF STROKE

18 YEARS FOLLOW-UP

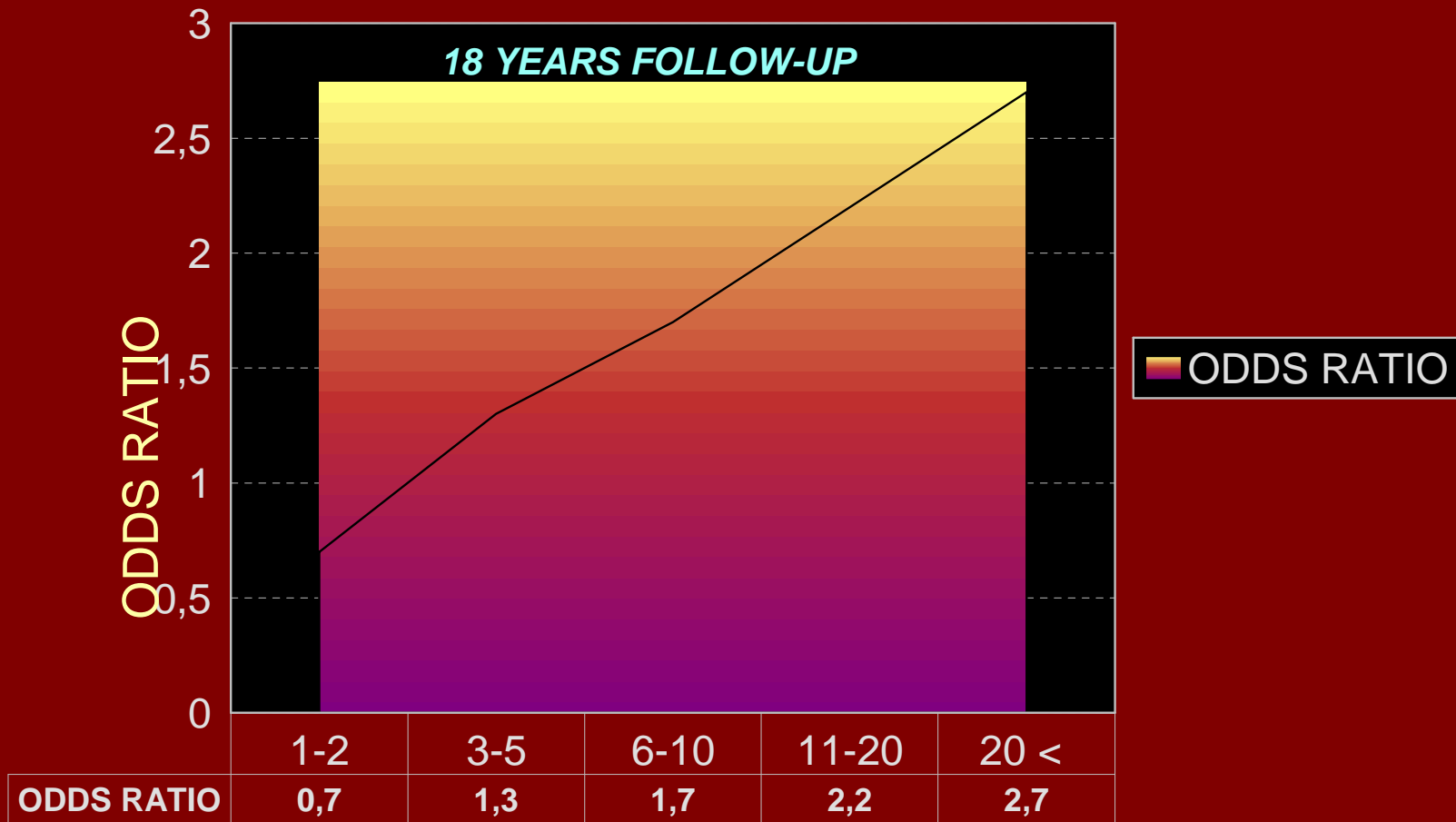


CUMULATIVE STROKE %	0	20%	40%	60%	80%
	3	7	10	10	13

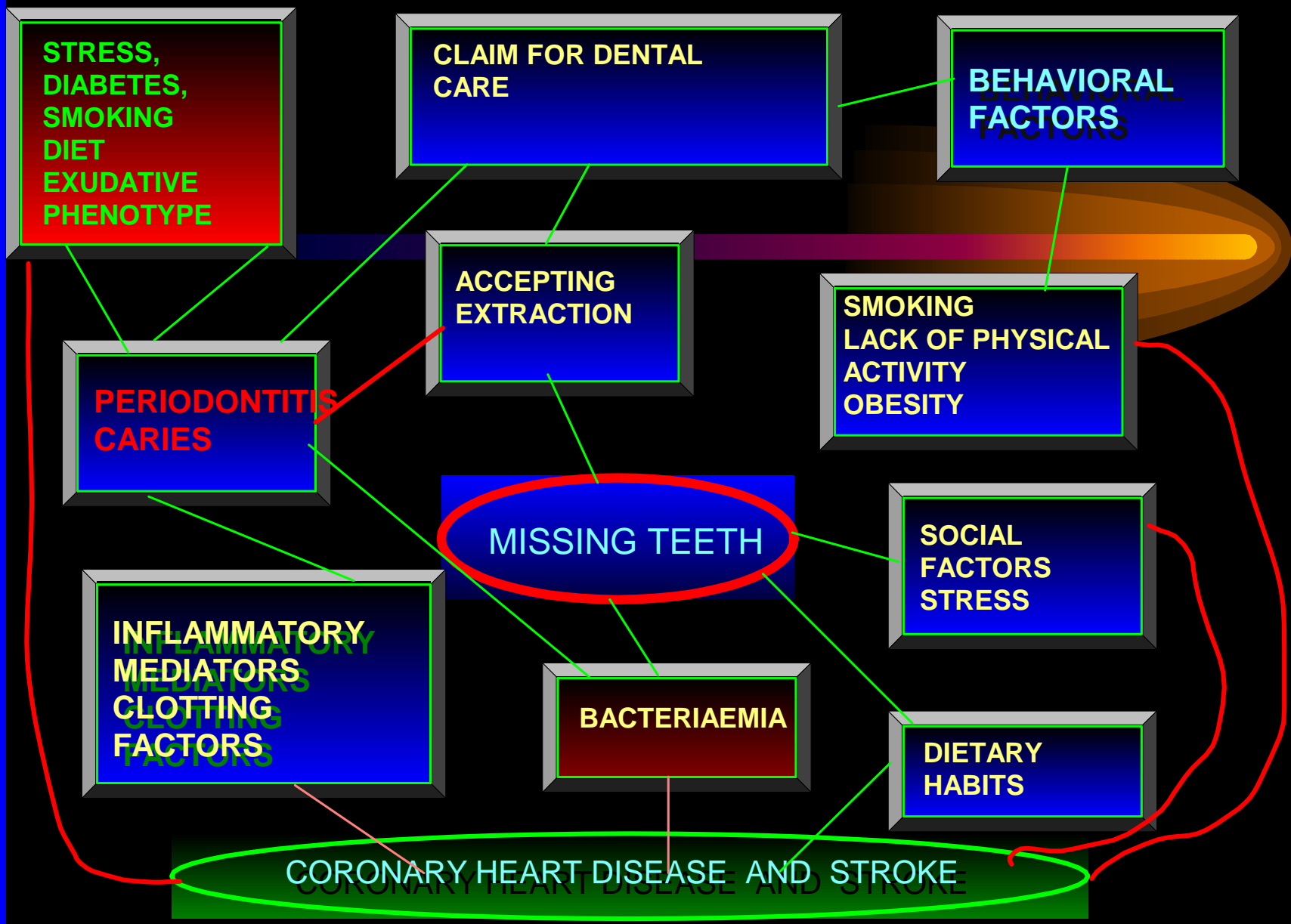
average bone loss

Beck JD. et al.: Periodontal disease and cardiovascular disease *J. Periodontol* 1996; 67(suppl): 1123-1137

THE CORRELATION BETWEEN THE NO OF TEETH WITH MORE THAN 20% ALVEOLAR BONE LOSS AND THE CUMULATIVE PREVALENCE OF CHD



Beck JD. et al.: Periodontal disease and cardiovascular disease J. Periodontol 1996; 67(suppl): 1123-1137



Joshipura et al.: Possible explanations for the tooth loss and cardiovascular disease relationship Ann. Period. 1998; 3.: 175-183, 1998

THE CORRELATION BETWEEN THE ATHEROSCLEROSIS (THE INCIDENCE OF CONGESTIVE HEART DISEASES) AND PERIODONTAL CONDITION

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Joshiyura - Japan	missing teeth / CHD	1,7

PUBLICATION	EXPOSITION	VASCULAR EFFECT	ASSOCIATION
DeStefano et al 1993/ USA	Russell PI	CHD/ lethal CHD	Sign. +
Hujoel et al 2000.2001 /USA	Gingivitis, Russell PI	CHD/ lethal CHD	negatív
Wu et al 2000/ USA	Gingivitis,missing tooth , deep pocket	Ischemic stroke	Gingivitis – Pocket sign. +
Hujoel et al 2002/USA	Russell PI gingivitis	Halálos CHD, kórházban kezelt CHD	negatív
Howel et al 2001/USA	periodontitis survey	Halálos CHD, kórházban kezelt CHD	negatív
Joshi-pura et al 1996/ USA	periodontitis survey	Halálos CHD, vagy nem halálos infarctus	Gingivitis negatív Parodontitis +
Joshi-pura et al 2003 USA	Numer of teeth periodontitis survey	Ischemias stroke	Sign. +
Beck 1996/USA	Periodontal bone level	New CHD, lethal CHD, stroke	Sign. +

PUBLICATION	EXPOSITION	VASCULAR EFFECT	ASSOCIATION
Mendez et al 1998/ USA	Total bone loss	Periferial atherosclerosis	Sign. +
Morrison et al 1999/Canada	Gingivitis, Periodontitis	lethal CHD, stroke	Sign. +
Mattila et al 1995/ Finland	TDI	New MI or lethal CHD	Sign. +
Touminen et al 2003/ Finland	CPITN, PPD, caries	Lethal CHD,	negative
Janson et al 2001/ Sweden	10%< bone loss	Lethal CHD,	Sign. +
Ajwani et al 2003/Finland	CPITN 4mm< pocket	Lethal CHD,	Sign. +

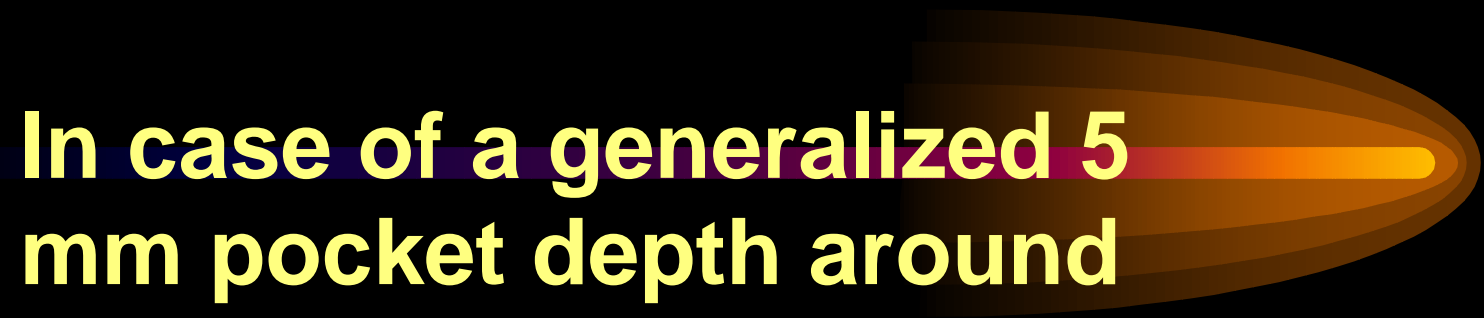
subgingival bacterial biofilm

gram negative bacterial deposits
endotoxine - LPS
high concentration of
inflammatory mediators
TNF-alpha, IL-1beta, PGE₂



***BACTERIAEMIA OCCURRED
IN 55% OF INDIVIDUALS WITH
SEVERE PERIODONTITIS
AFTER PARAFFIN CHEWING***

Murray M & Moonsnick F.: Incidence of bacteriemia in patients with dental plaque J. Lab Clin Med 1941; 26: 801-802.



In case of a generalized 5 mm pocket depth around teeth the subgingival bacterial flora communicates through an approx. 70-80 cm² open wound surface with the systemic circulation

BACTERIAEMIA



**EVEN WITH RELATIVELY
SOUND PERIODONTIUM A
TRANSIENT BACTERIAEMIA
CAN OCCUR DURING
CHEWING, TOOTHBRUSHING,
FLOSSING**

Silver JG et al.: Experimental transient bacteraemias in human subjects with varying degrees of plaque accumulation and gingival inflammation. J Clin Periodontol 1977;4: 92-99.

BACTERIAEMIA



IN SEVERE DESTRUCTIVE
PERIODONTITIS THE
BACTERIAL INVASION HAS
BEEN PROVEN IN THE
DENTAL LITERATURE

Allenspach-Petrzilka GE, Guggenheim B. Bacterial invasion of the periodontium: an important factor in the pathogenesis of periodontitis? J Clin Periodontol 1983;10:609-617.

infective endocarditis

IE

THE MOST COMMON CAUSE OF IE IS

***Streptococcus sanguis*, that is very common in dental plaque even in healthy periodontal condition**

Many times there were isolated from patients with IE

Gram - negative periodontopathogenic strains:

- *Actinobacillus actinomycetemcomitans*,**
- *Eikenella corrodens*,**
- *Fusobacterium nucleatum***
- *Capnocytophaga* strains**

Geraci JE, Wilson JR. Symposium on infective endocarditis III.

Endocarditis due to Gram-negative bacteria. Report of 56 cases. Mayo

Clinic Proc. 1982;57:145-148.

- **Mattila KJ, Valle MS, Neiminen MS, Valtonen VV, Hietantemi KL: Dental infections and coronary atherosclerosis. Atherosclerosis 103, 205, 1993.**
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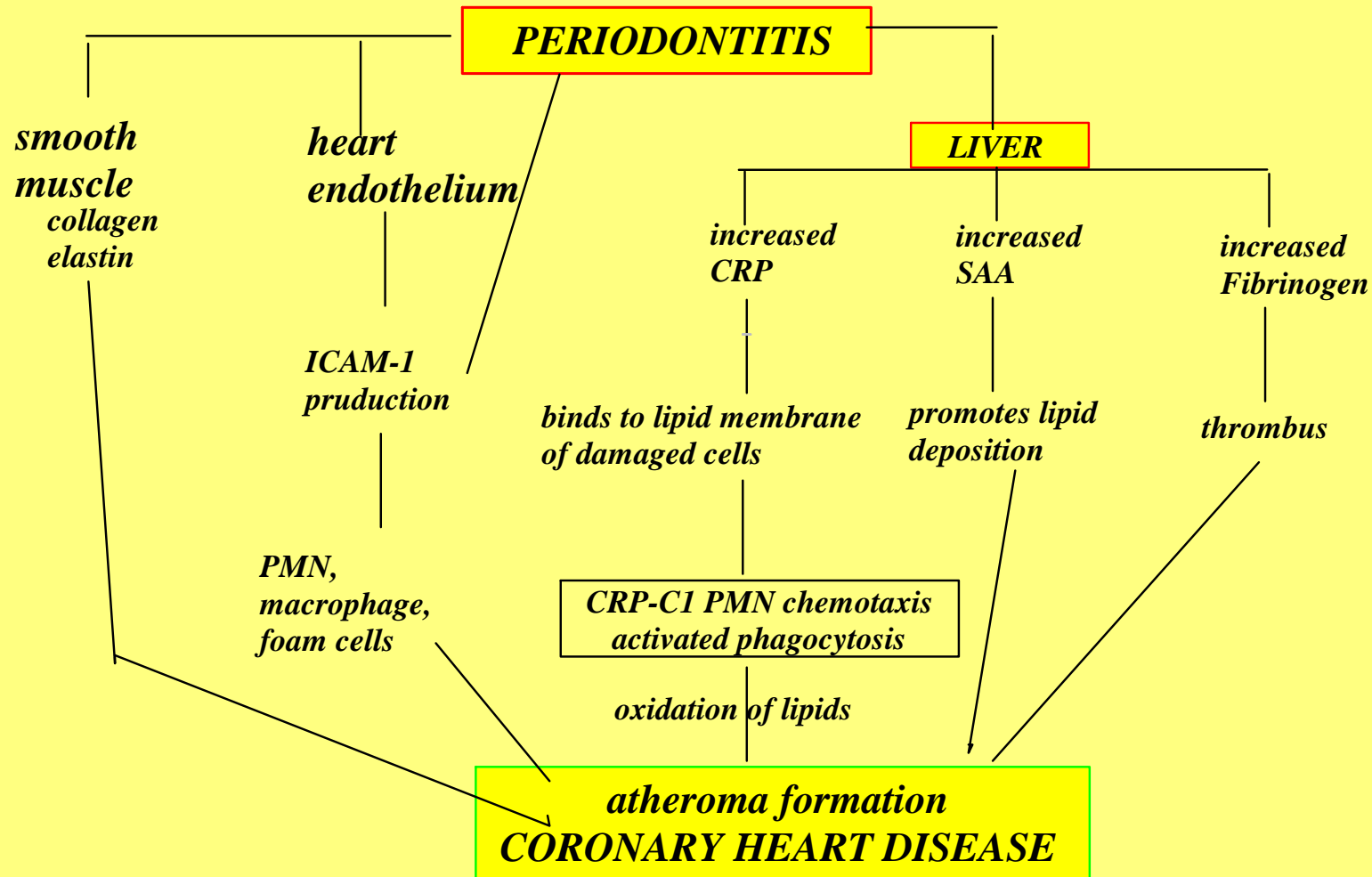
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- Beck J, Garcia R, Heiss G, Vokonas P, Offenbacher S: Periodontal disease and cardiovascular disease .J Periodontol 1996; 67: 1123-1137.
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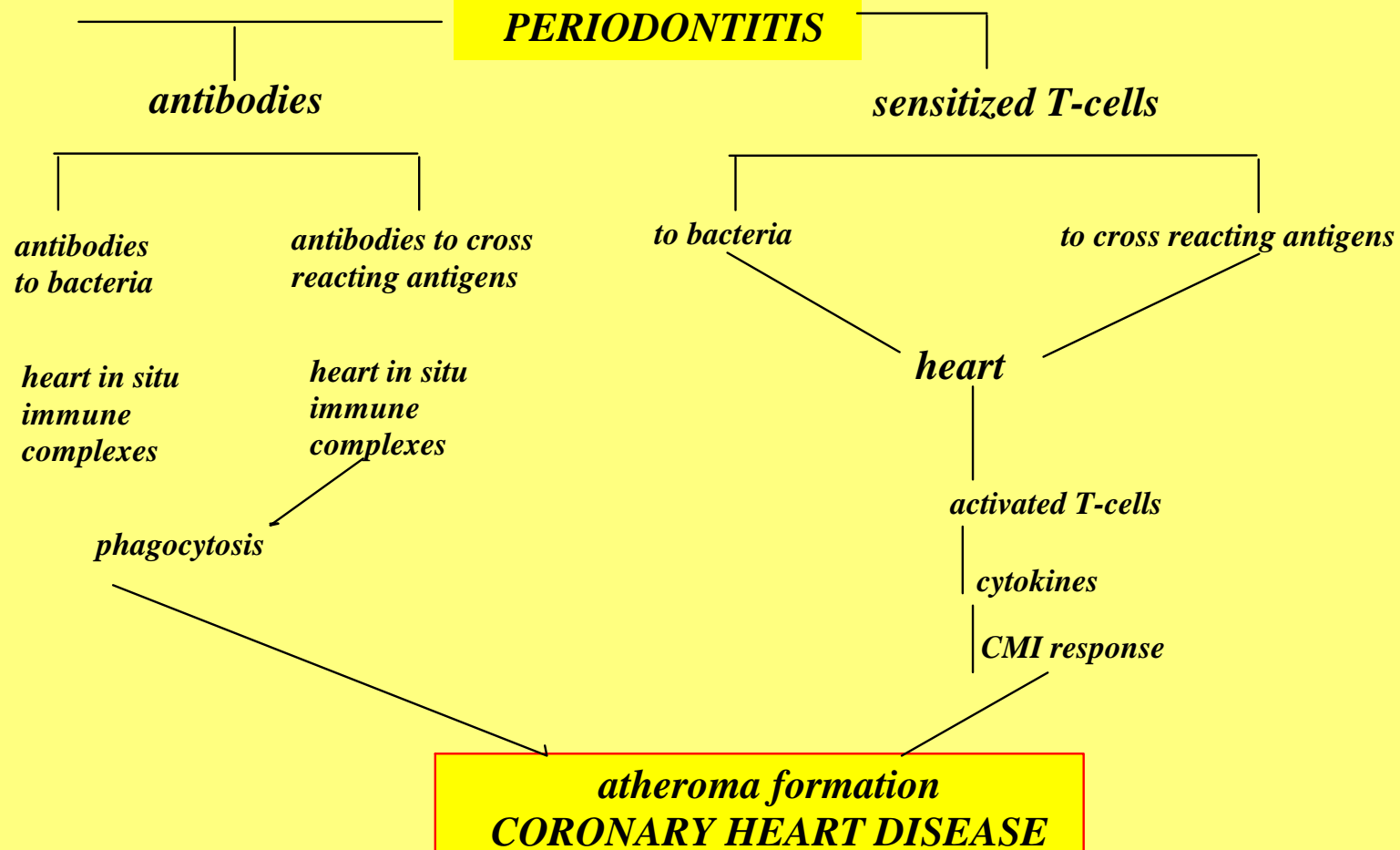
The image features a blue pen nib pointing towards a word cloud. The word cloud contains the words 'CARDIOVASCULAR' and 'diseases' in various sizes and orientations. The words are arranged in a way that they appear to be written by the pen nib. The word 'CARDIOVASCULAR' is written in red, while 'diseases' is written in cyan. The background is black with a blue border.

CARDIOVASCULAR
diseases

INFLAMMATORY MEDIATORS AND ITS RELATION TO CORONARY HEART DISEASES



IMMUNE RESPONSE TO LINK TO PERIODONTAL DISEASE AND CORONARY HEART DISEASE



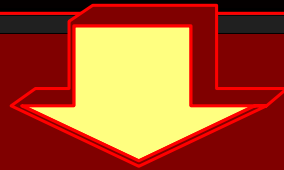
GILA RIVER INDIAN RESERVATION STUDY

**1440 AMERICAN NON-SMOKING MALE INDIANS
WERE FOLLOWED-UP FOR 15 YEARS
STRONG POSITIVE CORRELATION BETWEEN
PERIODONTAL STATE AND INCIDENCE OF
CARDIOVASCULAR DISEASES**

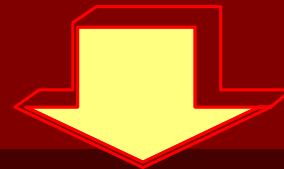
**THE INCIDENCE OF CORONARY HEART
DISEASE WAS 168%- HIGHER AMONG THOSE
WHO HAD SEVERE PERIODONTAL DISEASE
AT BASELINE**

**Genco RJ: . Periodontal disease and risk for myocardial infarction and
cardiovascular disease *Cardiovasc Rev Rep* 1998: March 34-40**

Experimental periodontitis in monkeys significantly increased serum (LPS), CRP and IL-8 concentration



elevated serum cholesterol, triglycerid and lipoprotein level



increased risk for atherosclerosis

ATHEROSCLEROSIS

LIPID ACCUMULATION

ATHEROMA RUPTURE, THROMBUS

FOCAL ENDOTHEL DAMAGE

INFLAMMATORY CELLULAR PROLIFERATION

PARTIAL OR TOTAL OBLITERATION

ATHEROMA

CYTOKINE PRODUCTION

CONTINUOUS GROWTH

ADHERENT, FIBROTIC ATHEROMA

PDGF

MACROPHAGES
T LYMPHOCYTE
THROMBOCYTE

SMOOTH MUSCLE PROLIFERATION

RISK FACTORS FOR CARDIOVASCULAR DISEASES

- **ELEVATED LDL CHOLESTEROL > 160mg/dL**
- **LOWER HDL CHOLESTEROL < 35mg/dL**
- **SMOKING**
- **OBESITY > 30% OVERWEIGHT**
- **MALE GENDER**
- **DIABETES MELLITUS**
- **ELEVATED HOMOCYSTEIN LEVEL**

Smoking is a serious risk factors in the etiology of cardiovascular and cardiorespiratoric diseases

It is also an important risk factor in the development of periodontitis



consequently the significantly higher incidence of cardiovascular diseases among heavy smokers can be attributed to a complex effect

Haber, J. et al: Evidence for cigarette smoking as a major risk factor for periodontitis. J. Periodontol 64, 16. 1993.

Coultchin J et al: Association of smoking with periodontal treatment needs. J Periodontol 61, 364. 1990.

**MORE THAN 100 YEARS AGO SIR
WILLIAM OSLER HYPOTHESIZED
THE INFECTIOUS ORIGIN OF
CARDIOVASCULAR DISEASES**

THAT TIME THERE WAS NO SCIENTIFIC
EVIDENCE TO SUPPORT THIS HYPOTHESIS

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ??!!

THE FIRST EVIDENCE WAS :

***CYTOMEGALOVIRUS INFECTION
IN IMMUNOLOGICALLY COMPROMISED
INDIVIDUALS LED TO SEVERE
CARDIOVASCULAR ATHEROSCLEROSIS***

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ???!!

- POSITIVE CORRELATION BETWEEN PEPTIC ULCER AND THE INCIDENCE OF CORONARY HEART DISEASES
- ***HELICOBACTER PYLORI*** INFECTION IN STOMACH ULCERS CAN BE RESPONSIBLE FOR THE INCIDENCE OF CORONARY HEART DISEASE

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ??!!

***CHLAMYDIA PNEUMONIAE INFECTION
CORRELATED WITH THE INCREASED INCIDENCE
OF CORONARY HEART DISEASE***

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ??!!

**BY CAMPYLOBACTER PNEUMONIAE INFECTION
ATHEROMA WAS EXPERIMENTALLY DEVELOPED
IN RABBITS**

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ???!!

POSSIBLE MECHANISMS

● BACTERIAL THROMBUS HYPOTHESIS

STREPTOCOCCUS SANGUIS

PORPHYROMONAS GINGIVALIS

**PLATELET-AGGREGATION-ASSOCIATED-PROTEIN
(PAAP)**

A SURFACE COLLAGEN-LIKE PROTEIN THAT
FACILITATES PLATELET AGGREGATION AND
FINALLY THROMBUS FORMATION

Herzberg MC et al The platelet as an inflammatory cell in periodontal disease: interactions with Porphyromonas gingivalis. *Molecular Pathogenesis of Periodontal disease* 1994: 247-255

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ???!!

POSSIBLE MECHANISMS

- BACTERIAL THROMBUS HYPOTHESIS

STREPTOCOCCUS SANGUIS GIVEN
INTRAVENOUSLY CAUSED HEART ATTACK AND
FINALLY DEATH IN RABBITS

Herzberg MC et al Effects of oral flora on platelets: possible consequences in cardiovascular disease J Periodontol 1996;67: 1138-1142

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ???!!

POSSIBLE MECHANISMS

● BACTERIAL THROMBUS HYPOTHESIS

50 carotis atheroma examined

CMV

H. PYLORI

CLAMIDIA PNEUMONIAE

P. GINGIVALIS

PREVOTELLA INTERMEDIA

A. ACTINOMYCETEMCOMITANS

were detected by DNAPOLYMERAZ CHAIN REACTION

**72% CONTAINED AT LEAST ONE PERIODONTOGENIC
PLAQUE BACTERIAL STRAIN**

Haraszthy VI et al: Identification of pathogens in atheromatous plaques J. Dent Res. 77: 1998

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ???!!

POSSIBLE MECHANISMS

- INFLAMMATORY MEDIATORS
C REACTIVE PROTEINS - (CRP)

NORMAL CONTROL :	1.14 ug/ml
PERIODONTITIS + CORONARY H DISEASE:	8.70 ug/ml

EFFECTIVE PERIODONTAL TREATMENT DECREASED SERUM
CRP LEVEL BY 65%

Genco et al : Overview of risk factors for periodontal disease and implications for diabetes and cardiovascular disease Compendium of Continuing Education in Dentistry 2000:

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ???!!

POSSIBLE MECHANISMS

● INFLAMMATORY MEDIATORS

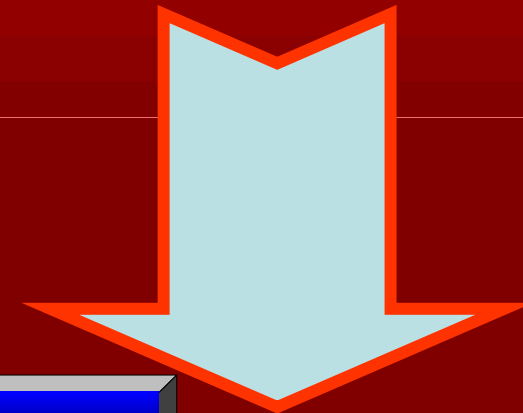
C-REACTIVE PROTEINS - (CRP)

PERIODONTITIS

BACTERIAEMIA

INCREASED TNF- α AND IL-6 PRODUCTION

INCREASED CRP PRODUCTION IN THE LIVER



Genco et al : Overview of risk factors for periodontal disease and implications for diabetes and cardiovascular disease *Compendium of Continuing Education in Dentistry 2000*:

ARE CARDIOVASCULAR DISEASE INFECTIOUS DISEASES ???!!

POSSIBLE MECHANISMS

● INFLAMMATORY MEDIATORS

C-REACTIVE PROTEINS - (CRP)

CRP DEPOSITED IN DAMAGED ENDOTHELIA

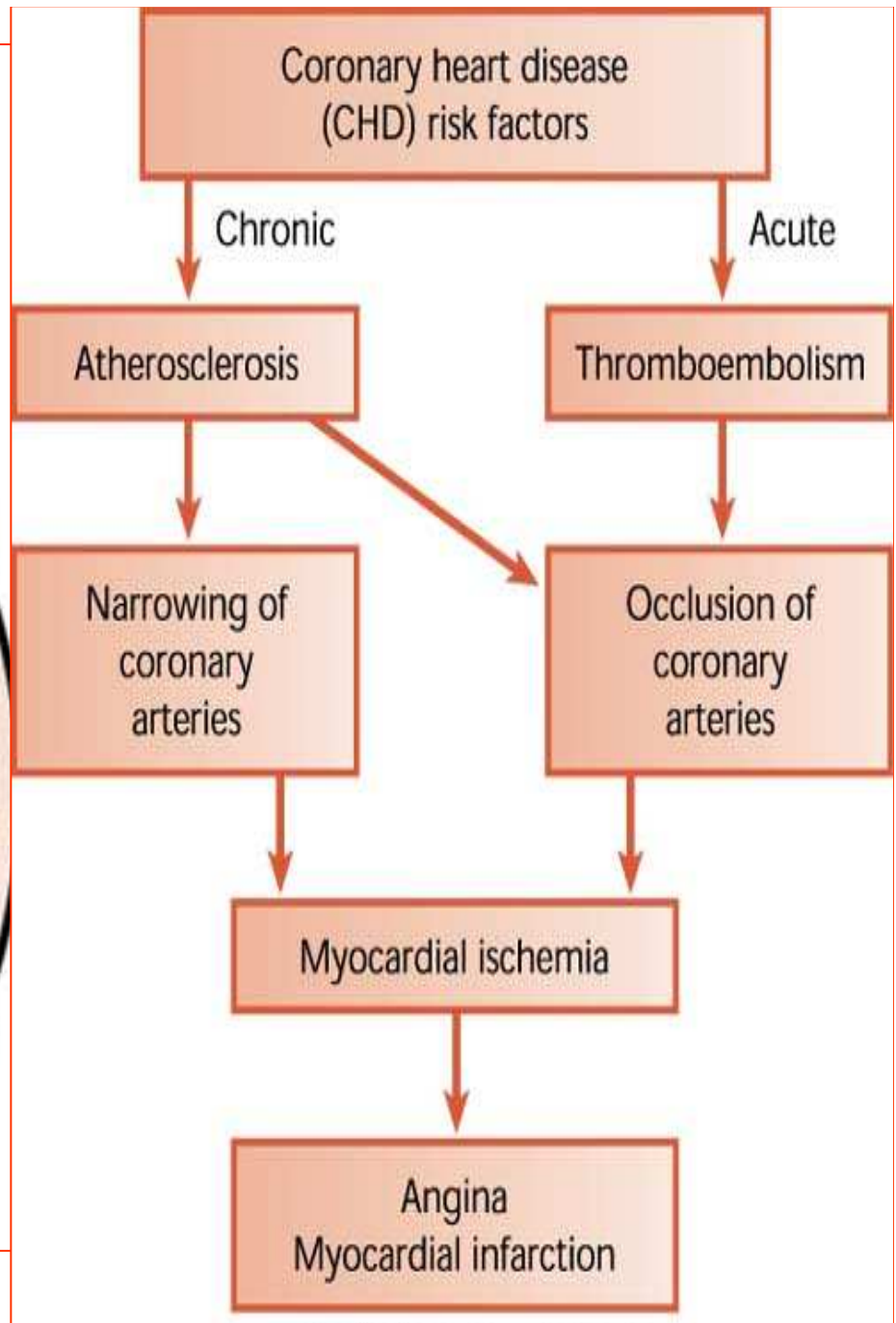
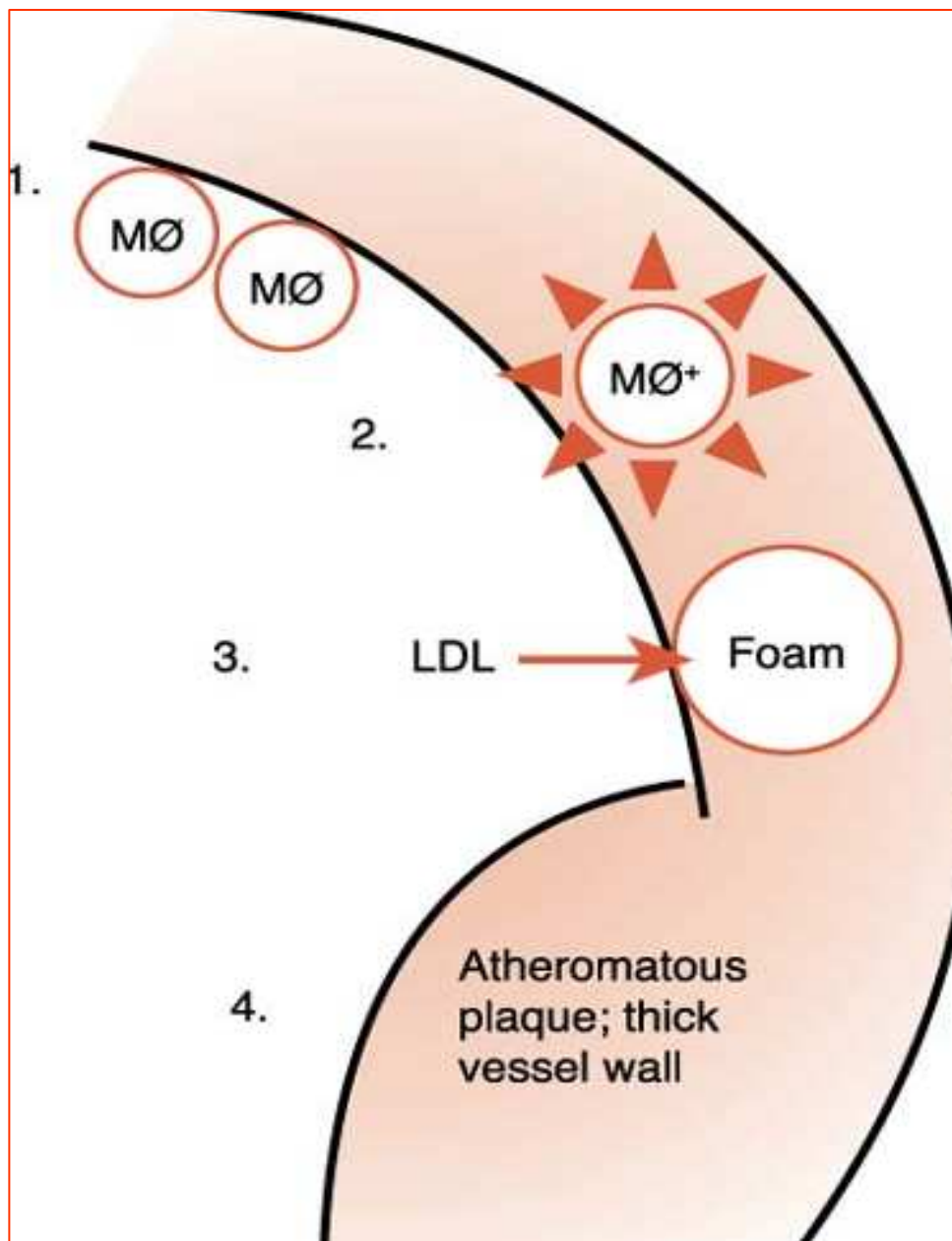
COMPLEMENT FIXATION

PHAGOCYTE ACTIVATION

NITRIC OXID LIBERATION - ATHEROMA



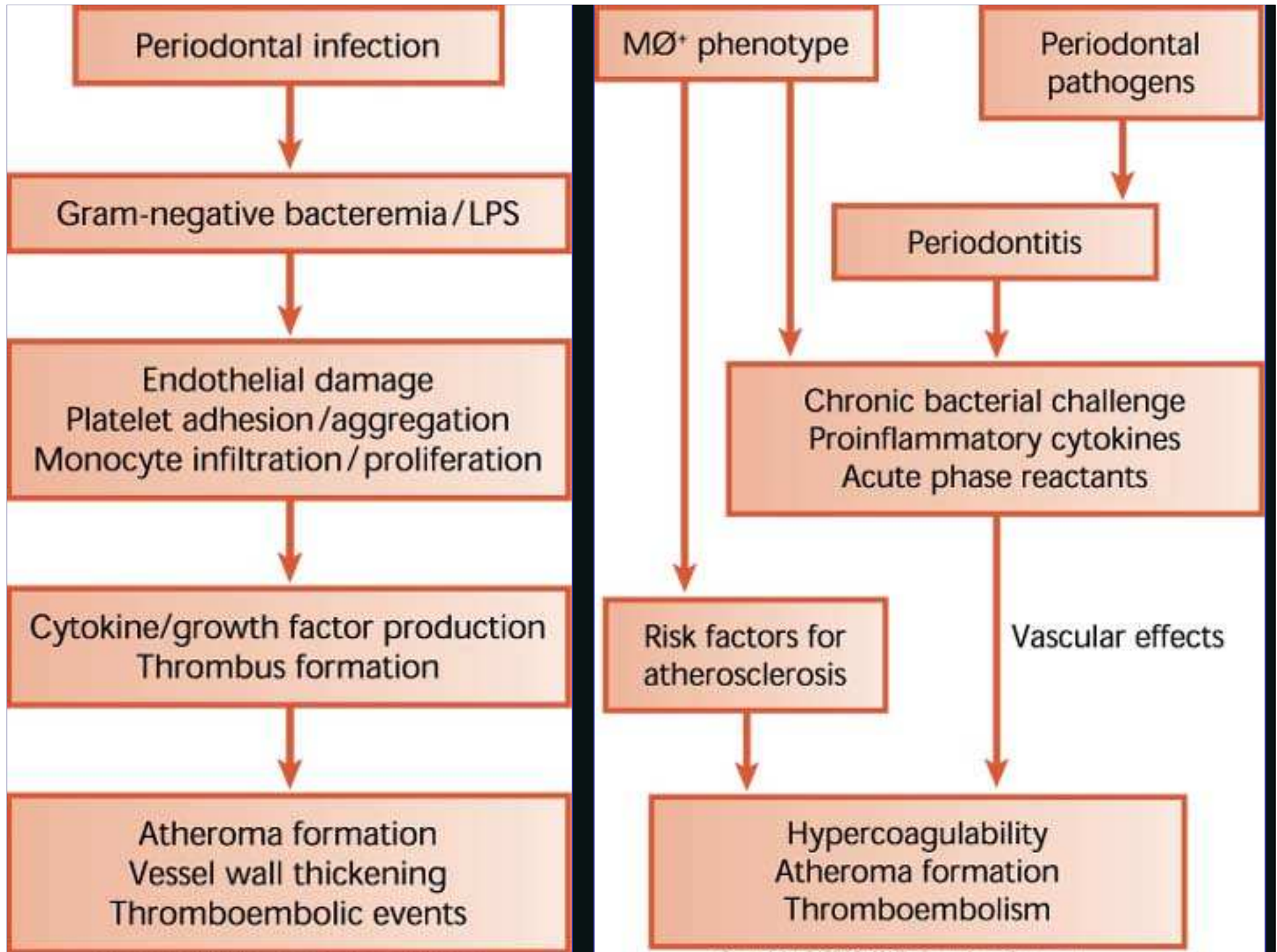
Genco et al : Overview of risk factors for periodontal disease and implications for diabetes and cardiovascular disease Compendium of Continuing Education in Dentistry 2000:



- Plasma fibrinogen
- Plasma lipoproteins (LDL/VLDL)
- White blood cell count



Blood viscosity



CAUSE OF BACTERIAL ENDOCARDITIS

BASED ON THE STATISTICS OF 450 PUBLICATIONS

95%PROBABILITY	%
DENTAL TREATMENT	7,5% (5,5-9,4)
DENTAL AND PERIODONTAL INFECTIONS	7,6% (5,2-9,6)
MEDICAL TREATMENT	14,5% (12,5-17,8)
EXTRAORAL INFECTIONS	15,5% (13,5-16,5)
INTRAVENOUS DRUG	4,5% (2,3-6,8)
UNKNOWN	52,5% (48,0-54,1)

Drangshot MT: A New Causal Model of Dental Diseases Associated With Endocarditis J. Periodontol, Annales of Periodontology 3:1. 185-196

SYSTEMIC RISK FACTORS FOR IE

alcoholism
intravenous drugs
chronic haemodialysis
immunosuppression
diabetes

**THE INDICATION OF
ANTIBIOTIC
PROPHYLAXIS BEFORE
SCALING**

**GENERALLY 15% OF ALL
KNOWN INFECTIVE
ENDOCARDITIS CASES ARE
PRECEDED BY INVASIVE
DENTAL TREATMENT ONE
MONTHS PRIOR TO THE
INCIDENCE OF IE**

**BESIDES TOOTH EXTRACTION
THE HIGHEST RISK IS
IMPOSED BY SUBGINGIVAL
SCALING AND CURETTAGE
AND INTRALIGAMENTAL
ANAESTHESIA**

**THE DESTRUCTIVE
PERIODONTITIS IS A
SERIOUS RISK FACTOR
FOR CERTAIN HIGHLY
SUSCEPTIBLE
INDIVIDUALS
(HIGH RISK GROUP)**

It is very important to anticipate the invasion of the plaque bacteria and to prevent them entering into the systemic circulation.

If it should occur the attachment of circulating bacteria onto endothelium or endocardium must be blocked by antibiotics

It is even more important to maintain excellent oral hygiene in high risk individuals than just to provide antibiotic prophylaxis during invasive therapy.

**Periodontitis is caused by plaque
bacteria**

**In dental plaque more than 500
bacterial strains were detected by
culturing or DNA PCR**

**One gram dental plaque contains
 2×10^{11} bacteria**

GibbonsRJ, van Houte J. Bacterial adherence and formation of dental plaques. Beachey EH. Ed. Bacterail Adherence. London: Chapman and Hall Ltd. 1980: 62-104.

CARDIOLOGICAL CONDITIONS REQUIRING ANTIBIOTIC PROPHYLAXIS BEFORE INVASIVE DENTAL TREATMENT:

HIGH RISK GROUPS

- ARTIFICIAL VALVE (mechanical or biological)
- PAST IE IN MEDICAL HISTORY
- COMPLEX CARDIAC DEVELOPMENTAL DISORDERS
- SYSTEMIC PULMONARY SHUNT
- CARDIAC VALVE CORRECTING OPEN HEART SURGERY

Dajani et al. Prevention of bacterial endocarditis Recommendations by the American Heart Association Circulation 1997;96:358-366

CARDIOLOGICAL CONDITIONS REQUIRING ANTIBIOTIC PROPHYLAXIS BEFORE INVASIVE DENTAL TREATMENT:

MEDIUM RISK GROUP

- RHEUMATOID FEVER
- KAWASAKI DISEASE
- AUTÓIMMUNE COLLAGEN DISEASES (SLE)
- HEART VALVE FUNCTIONAL DISORDERS
- ANY PATIENT AFTER OPEN HEART SURGERY

Dajani et al. Prevention of bacterial endocarditis Recommendations by the American Heart Association Circulation 1997;96:358-366

CARDIOLOGICAL CONDITIONS REQUIRING ANTIBIOTIC PROPHYLAXIS BEFORE INVASIVE DENTAL TREATMENT:

MEDIUM RISK GROUP

- HYPERTROPHIC CARDIOMYOPATHY
- MITRAL VALVE PROLAPS WITH REGURGITATION
- SEVERAL DEVELOPMENTAL DISORDER INFLUENCING THE FUNCTION OF THE VALVES.

Dajani et al. Prevention of bacterial endocarditis Recommendations by the American Heart Association Circulation 1997;96:358-366

CARDIOLOGICAL CONDITIONS REQUIRING NO ANTIBIOTIC PROPHYLAXIS BEFORE INVASIVE DENTAL TREATMENT:

- ATRIAL SEPTUM DEFECT
- SURGICALLY CORRECTED SEPTUM DEFECT
- CORONARY BYPASS OPERATIONS
- MITRAL VALVE PROLAPSE WITHOUT REGURGITATION
- PHYSIOLOGICAL HEART MURMUR
- PAST RHEUMATIC FEVER WITHOUT VALVE DAMAGE AND HEART MURMUR
- PACEMAKER

Dajani et al. Prevention of bacterial endocarditis Recommendations by the American Heart Association Circulation 1997;96:358-366

DENTAL PROCEDURES LEADING TO BACTERIAEMIA

HIGH RISK PROCEDURES

- extraction
- periodontal surgery
- subgingival curettage
- intraligamental anaesthesia
- subgingival scaling
- dentoalveolar surgery
- periodontal probing
- certain endodontal procedures
- supragingival scaling provoking gingival bleeding
- any procedure causing gingival bleeding

Dajani et al. Prevention of bacterial endocarditis Recommendations by the American Heart Association Circulation 1997;96:358-366

DENTAL PROCEDURES LEADING TO BACTERIAEMIA

HIGH RISK PROCEDURES

- intraligamental anaesthesia

Intraligamental anaesthesia causes more frequently bacteriaemia than extraction

Roberts et al. Odontogenic bacteriaemia ind intraligamental analgesia Br Dent J 1992;173:195-200

DENTAL PROCEDURES LEADING TO BACTERIAEMIA

HIGH RISK PROCEDURES

- periodontal pocket diagnostics with probe

IN PATIENTS WITH SEVERE PERIODONTITIS
BACTERIAEMIA OCCURED IN 43% OF ALL
CASES AFTER POCKET PROBING

Daly et al. Bacteraemia caused by periodontal probing Aust Dent J 1997;42:77-80

DENTAL PROCEDURES LEADING TO BACTERIAEMIA

HIGH RISK PROCEDURES

- periodontal surgery
- subgingival curettage
- subgingival scaling
- periodontal probing

In most of the cases bacteriaemia is transient lasting no longer than 15 min but in less severe cases no longer than 2-3 min. The shorter the transient bacteriaemia the less the chance to attach bacteria to the damaged endocardium

Dajani et al. Prevention of bacterial endocarditis Recommendations by the American Heart Association Circulation 1997;96:358-366

Mealey BL. Periodontal implications: medically compromised patients Ann Periodontol 1996;1:256-321

DENTAL PROCEDURES CAUSING NO BACTERIAEMIA

low risk procedures

- restorative work without retraction cord
- local anaesthesia
- rubber dam
- suture removal
- orthodontic treatment
- impression without retraction cord
- endodontic therapy without periapical involvement
- supragingival scaling without gingival bleeding
- intraoral radiographs

Dajani et al. Prevention of bacterial endocarditis Recommendations by the American Heart Association Circulation 1997;96:358-366

**THROUGH THE LARGE POCKET WALL
SURFACE A GREAT AMOUNT OF BACTERIA
AND BACTERIAL PRODUCTS CAN ENTER THE
BLOOD STREAM AND THIS MIGHT HAVE
VERY SERIOUS EFFECTS ON THE CONDITION
OF CERTAIN REMOTE ORGANS**

**EVEN IN CASES OF SEEMINGLY GOOD ORAL
HYGIENE A GREAT AMOUNT OF
SUPRAGINGIVAL PLAQUE COVERS THE
TEETH IN THE GINGIVAL REGIONS
CONSEQUENTLY EVERY TIME WE MIGHT BE
EXPOSED TO SYSTEMIC BACTERIAL
CHALLENGES**

**A TRANSIENT BACTERIAEMIA CAN BE
PROVOKED BY TOOTHBRUSHING, FLOSSING
ETC. EVEN IN CASE OF CLINICALLY
HEALTHY GINGIVA**

**THE SEVERITY OF BACTERIAEMIA DEPENDS
ON THE POCKET DEPTH AND GINGIVAL
INFLAMMATION**

Silver JG, Martin AW, McBride BC. Experimental transient bacteraemias in human subjects with varying degrees of plaque accumulation and gingival inflammation. J Clin Periodontol 1977;4: 92-99.

**amoxicillin SHOULD BE GIVEN ONE
HOUR PRIOR TO INVASIVE DENTAL
PROCEDURES**

**IN CASE OF PENICILLIN ALLERGY
600mg Clindamycin (Dalacin C) IS TO
BE GIVEN**

**This protocol is primarily against *alpha
haemolytic streptococcal infections***

*Dajani AS, Taubert KA, Wilson W et al. Prevention of bacterial endocarditis.
Recommendation by the American Heart Association JAMA 1997; 277: 1794-1801.*

The most common cause of IE is *Streptococcus sanguis*, this is the most common and numerous member of the supragingival plaque

This bacteria can easily attach to the sterile thrombotic plaques of the endothelium

Bayliss R, Clarke C, Oakley CM et al. The microbiology and pathogenesis of infective endocarditis Br Heart J 1983; 50: 513-519.

Herzberg MC, Meyer MW. Effects of oral flora on platelets: possible consequences in cardiovascular disease. J Periodontol 1996; 67: 1138-1142.

The second most common cause of IE is *Staphylococcus aureus*, that is the primary cause of the antibiotic resistant nosocomial IE

Bayliss R, Clarke C, Oakley CM et al The microbiology and pathogenesis of infective endocarditis Br Heart J 1983; 50: 513-519.

**Many times Gram negative anaerobic bacteria
can be cultivated from the haemocultures of
patients with IE
and also the so called periodonto- pathogenic
microorganisms**

***Actinobacillus actinomycetemcomitans,
Eikenella corrodens,
Fusobacterium nucleatum
Capnocytophaga subsp .***

Geraci JE, Wilson JR. Symposium on infective endocarditis III. Endocarditis due to Gram-negative bacteria. Report of 56 cases. Mayo Clinic Proc. 1982;57:145-148.

**if serial treatments are
indicated the procedures
should be made in 9-14 days
intervals to minimize the
possibility of the development
of antibiotic *resistance***

*Dajani AS, Taubert KA, Wilson W et al Prevention of bacterial endocarditis.
Recommendation by the American Heart Association JAMA 1997; 277: 1794-1801.*

before major cardiac surgery or joint prosthesis surgeries the patients need dental prophylaxis and comprehensive dental and periodontal treatment

following surgery patients also need regular periodontal maintenance care and excellent individual oral hygiene

It is wise to use chlorhexidine rinse before dental treatments to decrease the total bacterial count in the oral cavity nevertheless no scientific evidence support this statement

Barco CT. Prevention of infective endocarditis: a review of the medical and dental literature J Periodontol 1991;62:510-563

**IF A NON PREDICTED GINGIVAL
BLEEDING OCCURS DURING
DENTAL/ PERIODONTAL
PROCEDURES THE ANTIBIOTIC
PROPHYLAXIS CAN BE EFFECTIVE
WITHIN TWO HOURS
AFTER FOUR HOURS THE
ADMINISTERED ANTIBIOTIC CANNOT
EFFECTIVELY PROTECT THE
SUBJECT**

*Dajani AS, Taubert KA, Wilson W et al. Prevention of bacterial endocarditis.
Recommendation by the American Heart Association JAMA 1997; 277: 1794-1801.*

**THE ANTIBIOTIC
PROPHYLAXIS
DECREASES THE RISK TO
DEVELOP IE BUT IT
TOTALLY CANNOT BE
ANTICIPATED**

*Dajani AS, Taubert KA, Wilson W és mts. Prevention of bacterial endocarditis.
Recommendation by the American Heart Association JAMA 1997; 277: 1794-1801.*

**CARDIOVASCULARIS
BETEGSÉGEK**

**DIABETES
MELLITUS**

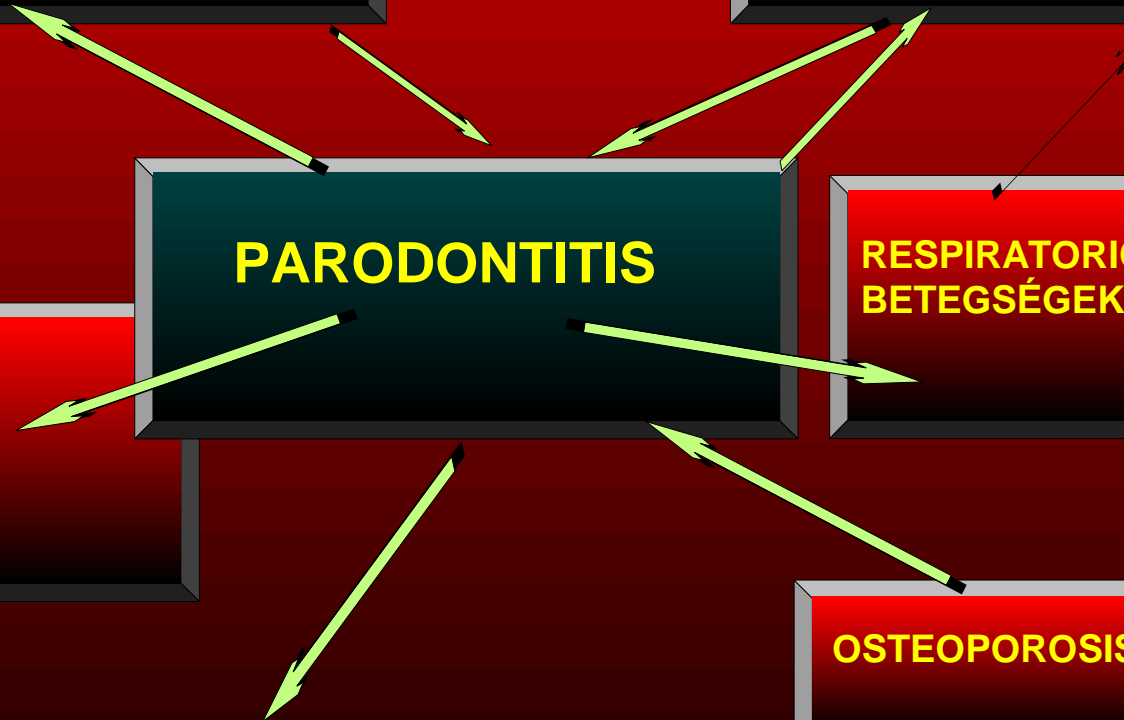
PARODONTITIS

**RESPIRATORICUS
BETEGSÉGEK**

**GASTRO-
INTESTINALIS
BETEGSÉGEK**

OSTEOPOROSIS

KORASZÜLÉSEK



THE DEMOGRAPHIC DATA OF NEW PATIENTS ADMITTED TO THE DEPARTMENT OF PERIODONTOLOGY IN 6 MONTHS

GENDER	MEAN AGE	NUMBER OF CASES
MALE	43,2	406
FEMALE	45,7	758
TOTAL	44,8	1164

ARTIFICIAL VALVE

	MALE	FEMALE
10-20	0	0
21-40	0	0
41-60	1	2
60 <	0	1
MEAN AGE	57	53
TOTAL	1	3

ENDOCARDITIS

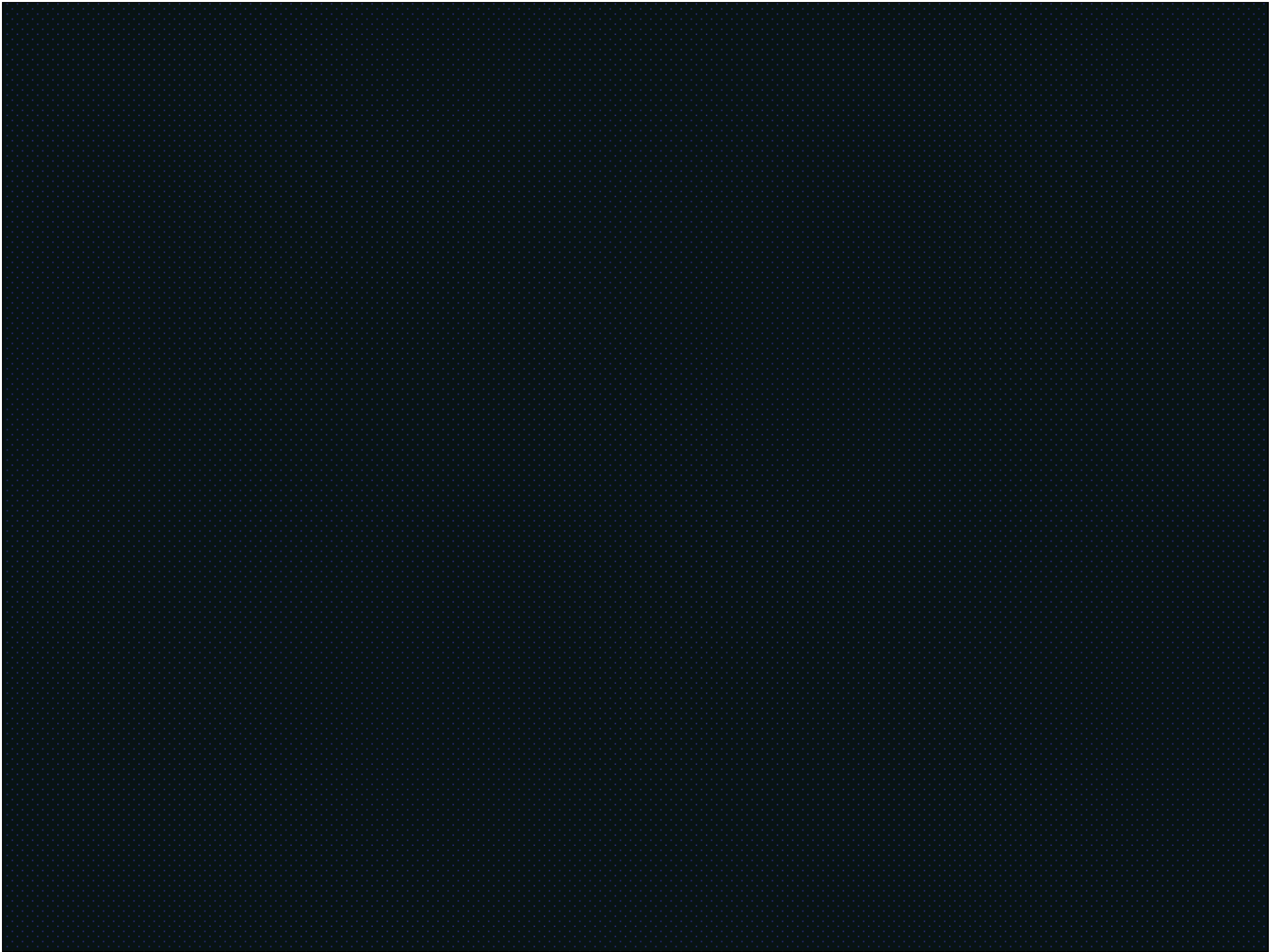
	MALE	FEMALE
10-20	0	0
21-40	0	0
41-60	3	8
60 <	2	0
MEAN AGE	58	48,5
TOTAL	5	8

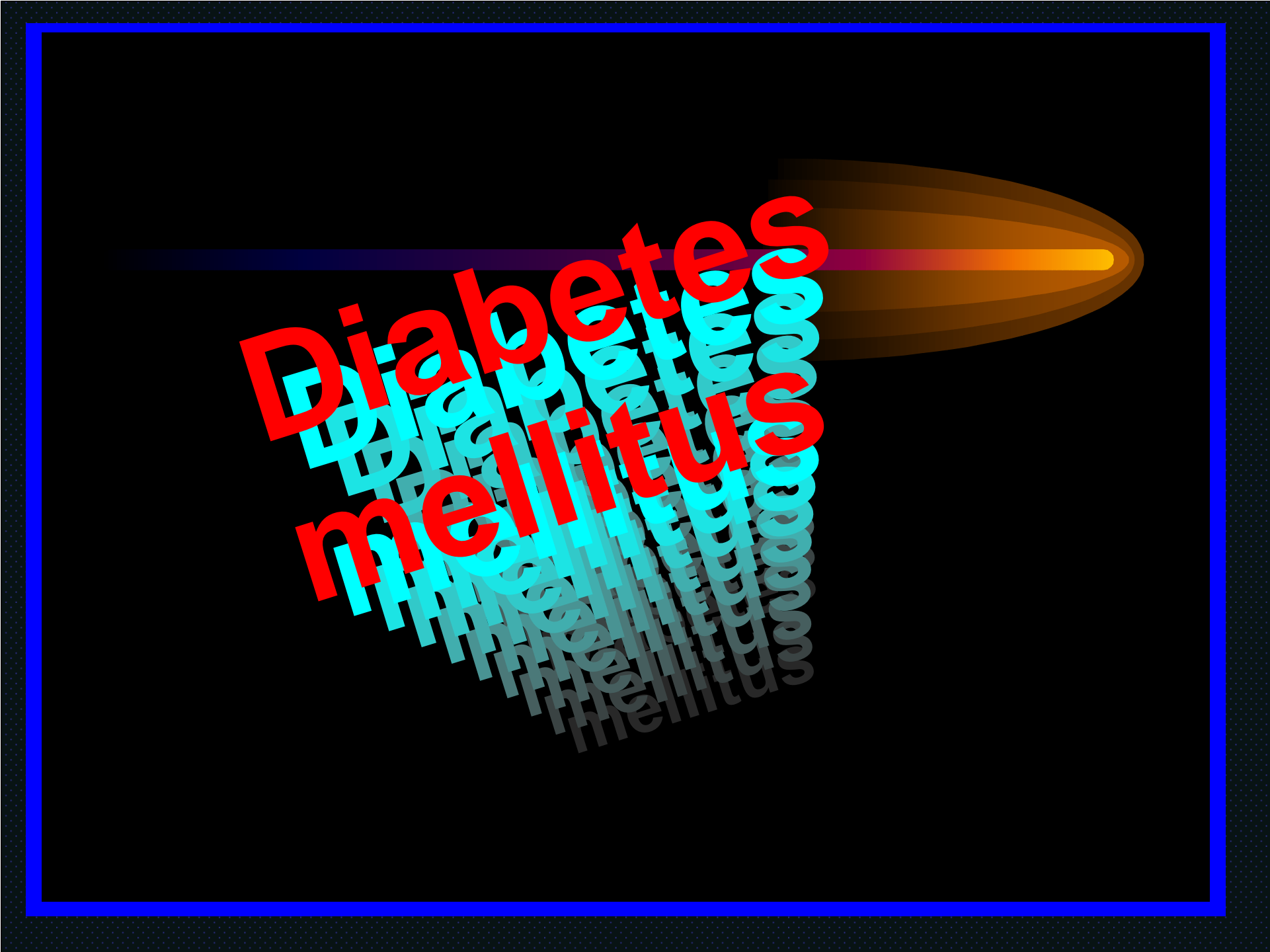
OPEN HEART SURGERY

	MALE	FEMALE
10-20	0	0
21-40	0	0
41-60	2	3
60 <	1	2
MEAN AGE	63	53
TOTAL	3	5

PACEMAKER

	MA,E	FEMALE
10-20	0	0
21-40	0	0
41-60	0	0
60 <	0	5
MEAN AGE	0	71
TOTAL	0	5





Diabetes
mellitus

Diabetes, especially that of the uncontrolled IDDM is a severe risk factor in the pathogenesis of destructive periodontitis

Also the chronic destructive periodontitis has a major negative effect on the metabolic disorders in diabetes mellitus

Löe H. Periodontal disease. The sixth complication of diabetes mellitus. Diabetes care 1993; 16: 329-334.

Hugoson A, Thorstensson H, Falk H, Kuylensterna J: Periodontal conditions in insulin-dependent diabetics. J Clin Periodontol 16, 215. 1989.

Miller LS, Manwell MA, Newbold D et al. The relationship between reduction in periodontal inflammation and diabetes control: a report of 9 cases. J Periodontol 1992; 63: 343-849.

RHEUMATOID ARTHRITIS ÉS PARODONTITIS

SÚLYOS PARODONTITISBEN
SZENVEDŐK KÖRÉBEN 4X
NAGYOBB A VALÓSZÍNŰSÉGE
ANNAK, HOGY **RA** SZEREPELJEN
AZ ANAMNÉZISÜKBEN

Mercado és mts.: Is there a relationship between rheumatoid arthritis and periodontal disease ? *J Clin Periodontol 2000;27: 267-272*

PERIODONTITIS AND SYSTEMIC DISEASES

PRETERM BIRTH

CORONARY HEART DISEASE

THROMBOEMBOLIC DISEASES

MYOCARDIAL INFARCT
STROKE

TYPE I DIABETES

TYPE II DIABETES

CAUSATIVE FACTORS :

EXCESSIVE PRO-INFLAMMATORY
CYTOKINE PRODUCTION IN
PERIODONTAL FOCUS

PROSTAGLANDIN E₂

TUMOR NECROSIS FACTOR (TNF) alfa

INTERLEUKIN (IL) - 6

DIRECT BACTERIAEMIA

RHEUMATOID ARTHRITIS AND PERIODONTITIS

PATIENTS WITH RHEUMATOID ARTHRITIS MOSTLY HAVE SEVERE PERIODONTITIS AS WELL

PROBABLE NO CAUSATIVE CORRELATION BETWEEN THE TWO IN BOTH DISEASES THE GENETIC PRO-INFLAMMATORY PHENOTYPE CAN BE ONE OF THE RISK FACTORS

Mercado és mts: Relationship between Rheumatoid Arthritis and Periodontitis *J Periodontol* 2001; 72: 779-787

Periodontitis and type II diabetes mellitus

**SEVERE PERIODONTAL INFLAMMATION
INCREASES TISSUE INSULIN RESISTANCE**

***IN PERIODONTITIS THE GRAM-NEGATIVE
INFECTION AND CHRONIC ENDOTOXEMIA
ELEVATE TISSUE INSULIN RESISTANCE AND
WORSENS PATIENTS' METABOLIC CONTROLS***

Grossi és mts. Response to periodontal therapy in diabetics and smokers *J Periodontol* 1996; 67: 1094-1102

Periodontitis and type II diabetes mellitus metabolic control

successful comprehensive periodontal therapy and regular periodontal maintenance in patients with type II DM potentiated the effect of anti-hyperglycemic therapy

Stewart et al. The effect of periodontal treatment on glycemic control in patients with type 2 diabetes mellitus *J Clin Periodontol* 2001; 28: 306-310

Iwamoto Y et al: The effect of antimicrobial periodontal treatment on circulating tumor necrosis factor alpha and glycated hemoglobin level in patients with Type 2 diabetes *J. Periodontol* 2001;72: 774-778.

Periodontitis and chronic respiratory diseases

bacteria being responsible for periodontitis might be directly or indirectly risk factors for upper respiratory diseases

Scannapio FA, et al.: Relationship between periodontal disease and bacterial pneumonia. J. Periodontol 1996; 67(Suppl): 1114-1122

Scannapio FA, Role of oral bacteria in respiratory infections J. Periodontol 1999; 70: 793-802

Periodontitis and chronic respiratory diseases

The data base of the III. National Health and Nutrition Examination Survey in the USA indicated that the presence of severe periodontitis increased the incidence of upper respiratory diseases by 50%

Scannapio FA, et al.: Potential associations between chronic respiratory disease and periodontal disease: Analysis of National Health and Nutrition Examination Survey III. *J. Periodontol* 2001; 72: 50 -56.

Periodontal Treatment: Effects on Glycemic Control⁴⁹

