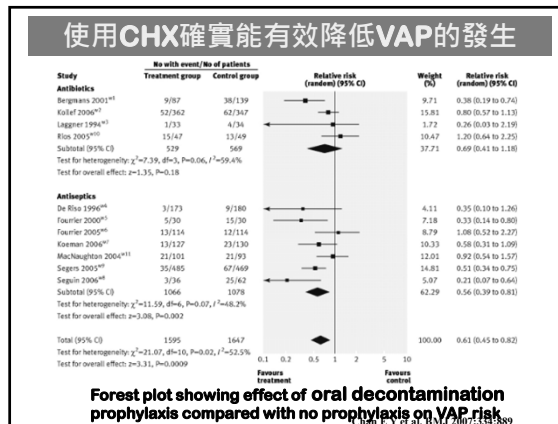


中國醫藥大學
China Medical University

口腔微生物與全身感染: 牙科照護策略的實證探討

盧敏吉 教授

中國醫藥大學微生物免疫學科
中國醫藥大學附設醫院感染科
luminch@mail.cmu.edu.tw



口腔的照護是重要的議題

牙齒

- 口腔的細菌是造成蛀牙與感染的要素!
- 定殖菌 vs. 致病菌
- 口腔局部感染! 全身感染!
- 哪些是重要的致病菌? 造成感染的證據?
- 不同口腔照護方式影響臨床床預後? 實證

口咽的定殖菌 (GNB)

Community Hospital Environment

N Engl J Med 1969; 281:1137

ADDRESSES AND ORIGINAL ARTICLES

ON CERTAIN SEPTICEMIAS DUE TO ANAEROBIC ORGANISMS *

By A. LEMIERRE, M.D.

PROFESSOR IN RADIOLOGY OF THE FACULTY OF MEDICINE, PARIS; ATTENDING TO THE CLAUDE BERNARD HOSPITAL.

Lemierre's syndrome

“在出現和重複發燒反應，之前有冷顫發生的喉嚨痛（特別是扁桃腺腫），伴隨發作肺梗塞和關節炎表現的綜合病徵.....”

急性口咽感染
合併頸內靜脈敗血症
血栓性靜脈炎
和轉移性感染。

Lemierre's syndrome 臨床表現

- 發熱· 毒性症狀; 通常是健康的年輕人
- 口咽感染
- 咽喉炎· 乳突炎· 牙科感染· 手術· 外傷
- 內頸靜脈血栓通常在口咽感染後4-8天發生
- 腫脹在下頷角
- 大約50%的患者沒有血栓形成
- 膿毒性栓塞來自內頸靜脈靜脈的血栓
- 肺· 化膿性關節炎· 內臟膿腫· 腦膜炎等
- 死亡率: Lemierre描述的系列研究為80%; 在最近的研究為4-12%

致病菌

Fusobacterium necrophorum most commonly recovered
Peptostreptococcus species
Bacteroides species
Haemophilus aphrophilus

Gram stain of *Fusobacterium necrophorum*

細菌性心內膜炎 MICROBIOLOGY OF NATIVE vs. PROSTHETIC VALVE

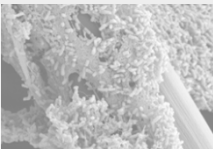
TABLE 1. MICROBIOLOGIC FEATURES OF NATIVE-VALVE AND PROSTHETIC-VALVE ENDOCARDITIS.

PATHOGEN	NATIVE-VALVE ENDOCARDITIS			PROSTHETIC-VALVE ENDOCARDITIS		
	2 MO-15 YR OF AGE	16-60 YR OF AGE	>60 YR OF AGE	EARLY (<60 DAYS AFTER PROCDURE)	INTERMEDIATE (60 DAYS-12 MO AFTER PROCDURE)	LATE (>12 MO AFTER PROCDURE)
	approximate percentage of cases					
<i>Streptococcus</i> species	15-20	40-50	45-65	30-45	1	7-10
<i>Staphylococcus aureus</i>	40-50	22-27	30-40	25-30	30-24	10-15
Cocci, negative, alpha-hemolytic	8-12	4-7	4-8	3-5	30-35	30-35
<i>Enterococcus</i> species	<1	3-6	5-8	14-17	5-10	10-15
Gram-negative bacilli	8-12	4-6	4-10	5	10-15	2-4
Fungi	8-12	1-3	1-3	1-2	5-10	10-15
Culture-negative and HACEK organisms*	2-6	0-15	3-10	5	3-7	2-7
Diphtheroids	<1	<1	<1	<1	5-7	2-5
Polymicrobial	3-5	<1	1-2	1-3	2-4	3-7

*Patients whose blood cultures were rendered negative by prior antibiotic treatment are excluded. HACEK denotes *Haemophilus parainfluenzae*, *H. aphrophilus*, and *H. parrophilus*; *Acinetobacter anitruminus*; *Cardiobacterium hominis*; *Eikenella corrodens*; and *Klebsiella* sp.

由人工髖關節prosthetic hips分離出產生生物膜(Biofilm)的微生物

Coagulase-negative Staphylococci
Hemolytic streptococci
P. mirabilis
Bacterioides species
S. aureus
Viridans streptococci
E. coli
P. aeruginosa



血行來源 (20-40%)

- S. aureus* 菌血症:
佔有PJI的34%
- 牙齒牙齦感染或局部處理:
viridans streptococci & anaerobes
- GU 或 GI 手術或感染:
GNB, enterococci & anaerobes

Mandell, Bennet and Dolin, Principles and Practice of Infectious Diseases, 6th ed. Elsevier, 2005

細菌性心內膜炎 Bacterial Endocarditis Predisposing Factors

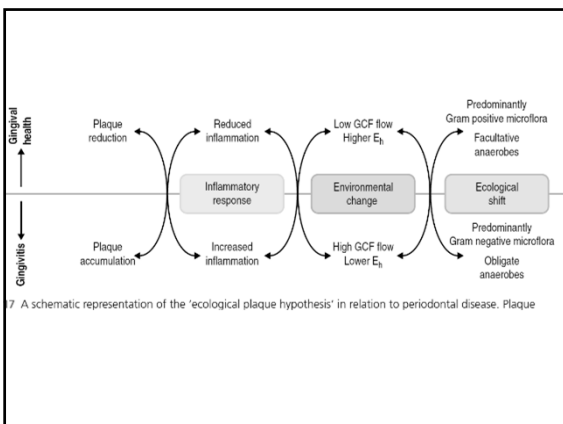
- 牙科操作
- 牙齒疾病(齲齒caries, 膿腫abscess)
- 心臟外感染(肺, 尿道, 皮膚, 骨, 膿腫)
- 侵入性措施(尿道, 胃腸道, 輸液)
- 心臟手術
- 使用注射毒品
- 沒有明顯的因素

人工關節感染 Prosthetic Joint Infection (PJI) 危險因子

- 皮膚潰瘍、壞死
- 類風濕關節炎
- 之前的臀部、膝蓋手術
- 反覆性泌尿道感染
- 慢性腎功能不全
- 需要化療的腫瘤
- 糖尿病
- 口服皮質類固醇
- 拔牙

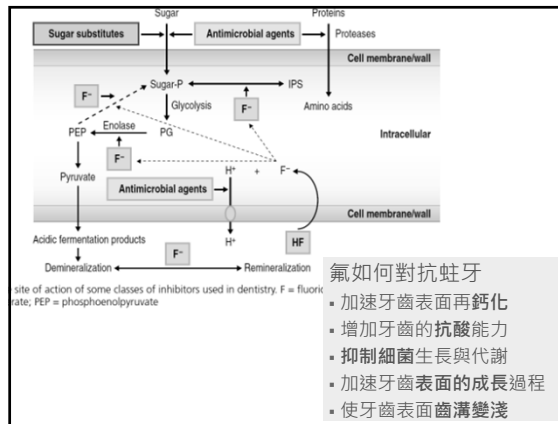
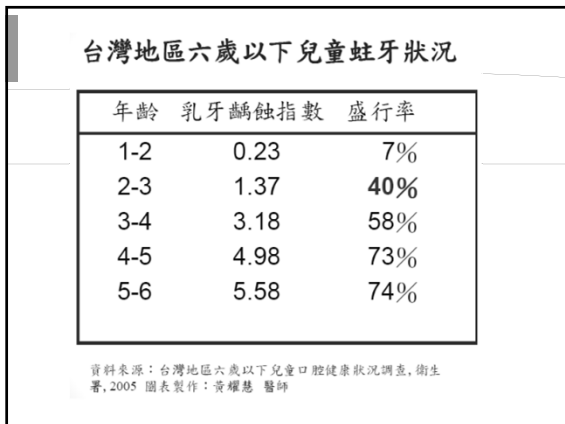
Table 6.6 Some of the predominant bacteria that have been commonly cultured from sites with chronic periodontitis in adults

Gram positive	Gram negative
<i>Eubacterium brachy</i>	<i>Tannerella forsythia</i>
<i>Eubacterium nodatum</i>	<i>Fusobacterium nucleatum</i>
<i>Mogibacterium timidum</i>	<i>Porphyromonas gingivalis</i>
<i>Parvimonas micra</i>	<i>Prevotella intermedia</i>
<i>Peptostreptococcus stomatis</i>	<i>Prevotella loeschii</i>
<i>Parvimonas micra</i>	<i>Dialister pneumosintes</i>
	<i>Campylobacter rectus</i>
	<i>Treponema spp.</i>



牙周病

- 「牙周」是指「牙齒周圍的組織」，也就是支持牙齒穩固於齒槽骨的組織，包括：牙齦(肉)、牙周韌帶(使牙根附著於牙槽骨的纖維)、齒槽骨、以及牙骨質(覆蓋於牙根表面的鈣化組織，能使牙周韌帶附著其上)，這四種組織有了毛病便稱為「牙周病」。
- 臺灣地區18歲以上的成年人，牙周病的盛行率達99.1%；定期看牙醫的人卻不到20%。



Clinical Infectious Diseases
MAJOR ARTICLE

Oral Streptococcal Endocarditis, Oral Hygiene Habits, and Recent Dental Procedures: A Case-Control Study

Xavier Duval,¹ Sarah Millet,² Catherine Chirouze,³ Christine Selton-Suty,⁴ Vanessa Moly,⁵ Pierre Tettevin,⁶ Christophe Strady,⁷ Edouard Evrard,⁸ Nelly Agrillon,⁹ Daniel Thomas,¹⁰ Bruno Heen,¹¹ and François Alla,^{12,13} for the Ii-dents Association pour l'Etude et la Prévention de l'Endocardite Infectieuse (AEPFI) Study Group

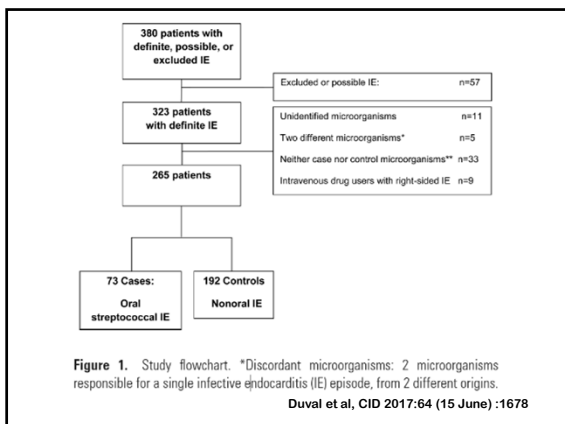
¹Service CIC 1425, AP-HP Hôpital Universitaire Bicêtre, Inserm UMR 1127 IAME, Université Paris Diderot, UFR de Médecine-Bicêtre, and UMR 1149-Inserm, CR, Université Paris Diderot, Faculté de Médecine Bicêtre, Paris; ²UMR 8249 Laboratoire Choro-environnement Université de Bourgogne Franche Comté, Service de maladies infectieuses, CHU Bicêtre; ³Centre Hospitalier Régional Universitaire, and ⁴Service Epidémiologie-Centre Hospitalier Régional Universitaire Nancy; ⁵Maladies Infectieuses et Immunologie Médicale, Centre Hospitalier Universitaire, Sorbonne; ⁶Cabinet Clinique, Clinique Saint André-Groupes Couvables, Reims, France; ⁷CIC 1431, Service de Stomatologie, Chirurgie Maxillo-faciale et Odontologie Hospitalière, CHU Bicêtre; ⁸Tourcoing; ⁹CIC 1433 Epidémiologie Clinique, Centre Hospitalier Régional Universitaire Nancy; ¹⁰AP-HP Hôpital Pitié-Salpêtrière, Département de Cardiologie, Paris; ¹¹Université des Antilles et de la Guyane, Faculté de Médecine Necker-Bichat, EA 4020, Centre Hospitalier Universitaire de Pointe-à-Pitre, Inserm CIC 1425, Service de Maladies Infectieuses et Tropicales, Stomatologie, Médecine Interne, Pointe-à-Pitre and ¹²Université de Lorraine, Université Paris Descartes, Apejac, EA4002 Inserm, CIC-1433, Nancy, France

Duval et al, CID 2017;64 (15 June) :1678

Background. We aimed to compare oral hygiene habits, orodental status, and dental procedures in patients with infective endocarditis (IE) against background (BG) patients. The study was a case-control study comparing oral hygiene habits, orodental status, and dental procedures in patients with infective endocarditis (IE) against background (BG) patients. The study was a case-control study comparing oral hygiene habits, orodental status, and dental procedures in patients with infective endocarditis (IE) against background (BG) patients.

於2008年5月至2013年1月期間，在6家法國三級醫院進行了一項評估員盲法病例對照研究。口腔衛生習慣是使用自填式問卷記錄。口腔牙齒的狀態由訓練有素的牙科醫生以盲法分析微生物。使用標準化的臨床檢查和牙科全景成像分析。牙科程序的歷史是通過患者和牙醫訪談獲得。病例和對照有確定的IE，在研究過程中一個專家驗證的保密清單，導致的微生物被分類為口腔鏈球菌或非口腔病原體。streptococci or nonoral pathogens, respectively. Participants were enrolled between May 2008 and January 2013.

Duval et al, CID 2017;64 (15 June) :1678



Characteristic	Whole Population		Cases: Oral Streptococcal IE		Controls: Nonoral IE		P Value
	N = 265	n = 73 (27.5%)	n = 192 (72.5%)				
Age, y, mean (SD)	60.8 (16.7)	55.5 (17.6)	62.8 (15.9)		.002		
Age ≥65 y	123 (46.4)	21 (28.8)	102 (53.1)		.001		
Male sex	212 (80.0)	63 (72.6)	159 (82.8)		.003		
At least 1 comorbidity	94 (35.5)	16 (20.5)	79 (41.1)		.002		
Diabetes mellitus	48 (18.1)	9 (12.3)	39 (20.3)		.136		
Cancer	41 (15.5)	6 (8.2)	35 (18.2)		.044		
Dialysis	5 (1.9)	0	5 (2.6)		.323		
Intravenous drug use	6 (2.3)	0	6 (3.1)		.019		
Active smoking	51 (19.8)	15 (21.7)	36 (19.0)		.631		
Cardiac history							
Underlying valve disease (HD)					.014		
Prosthetic valve	67 (25.3)	18 (24.7)	49 (25.5)				
Previously known native HD	76 (28.1)	30 (41.1)	46 (24.0)				
No previously known HD	122 (46.0)	25 (34.2)	97 (50.5)				
Pacemaker and/or implantable cardioverter defibrillator	29 (10.9)	2 (2.7)	27 (14.1)		.008		
History of IE	21 (7.9)	7 (9.6)	14 (7.3)		.536		

Data are presented as No. (%) unless otherwise indicated. Bold indicates statistically significant values. Abbreviations: HD, heart disease; IE, infective endocarditis; SD, standard deviation.

Duval et al, CID 2017;64 (15 June) :1678

Cardiac conditions associated with the highest risk of adverse outcome from endocarditis for which prophylaxis with dental procedures is reasonable.

- Prosthetic cardiac valve or prosthetic material used for cardiac valve repair
- Previous infective endocarditis
- Congenital heart disease (CHD)^{*}
 - Unrepaired cyanotic CHD, including palliative shunts and conduits
 - Completely repaired congenital heart defect with prosthetic material or device, whether placed by surgery or by catheter intervention, during the first six months after the procedure[†]
 - Repaired CHD with residual defects at the site or adjacent to the site of a prosthetic patch or prosthetic device (which inhibit endothelialization)
- Cardiac transplantation recipients who develop cardiac valvulopathy

^{*} Except for the conditions listed above, antibiotic prophylaxis is no longer recommended for any other form of CHD.
[†] Prophylaxis is reasonable because endothelialization of prosthetic material occurs within six months after the procedure.

JADA, Vol. 139 <http://jada.ada.org> January 2008

Dental procedures for which endocarditis prophylaxis is reasonable for patients in Box 3.

All dental procedures that involve manipulation of gingival tissue or the periapical region of teeth or perforation of the oral mucosa.^{*}

^{*} The following procedures and events do not need prophylaxis: routine anesthetic injections through noninfected tissue, taking dental radiographs, placement of removable prosthodontic or orthodontic appliances, adjustment of orthodontic appliances, placement of orthodontic brackets, shedding of primary teeth, and bleeding from trauma to the lips or oral mucosa.

JADA, Vol. 139 <http://jada.ada.org> January 2008

Primary reasons for revision of the infective endocarditis prophylaxis guidelines.

- Infective endocarditis (IE) is much more likely to result from frequent exposure to random bacteremias associated with daily activities than from bacteremia caused by a dental, gastrointestinal (GI) tract or genitourinary (GU) tract procedure
- Prophylaxis may prevent an exceedingly small number of cases of IE, if any, in people who undergo a dental, GI tract or GU tract procedure
- The risk of antibiotic-associated adverse events exceeds the benefit if any, from antibiotic prophylaxis
- Maintenance of optimal oral health and hygiene may reduce the incidence of bacteremia from daily activities and is more important than prophylactic antibiotics for a dental procedure to reduce the risk of IE

JADA, Vol. 139 <http://jada.ada.org> January 2008

CHLORHEXIDINE FORMULATIONS Mouthrinses

Chlorhexidine (CHX) 洗必泰, 氯己定 口腔沖洗液的形式為0.2%和0.12%。當以相應的相似劑量使用時, 0.2%和0.12%的沖洗效果相同。沖洗時間為30或60秒, 取決於消毒劑對口腔表面的吸附速率(50%的CHX在15秒內與受體結合), 但情況因個體而異。在72小時非刷牙期後, 使用0.2%CHX的沖洗時間為15、30和60秒, 其斑塊抑制作用並沒有明顯差異。理想的治療方案是每天早晚兩次, 作用將持續12小時。

Hoffman et al. *Clin Oral Invest* 2001, 5: 89-95.
 Van de Weijden GA, Timmerman MF, Novrny GA, Rosema N, Verkerk A. Three different rinsing times and inhibition of plaque accumulation with chlorhexidine. *J Clin Periodontol* 2005, 32(1):89-92.

Chlorhexidine: The Gold Standard Antiplaque Agent

CHLORHEXIDINE FORMULATIONS Gel 凝膠

氯己定凝膠製劑的可用濃度為1%、0.2%、0.12%, 以托盤和牙刷交付。每天施用一次的氯己定凝膠具有治療效果, 例如減少口腔惡臭, 並可減少氯己定染色。

牙膏: 0.12%的CHX與百萬分之一的氟化物具有類似於CHX漱口水的抗斑效果。然而, 將CHX加入凝膠和牙膏中存在困難。1%CHX用作液態並每天沖洗一次, 持續一分鐘導致牙菌斑和牙齦評分顯著降低, ……

Dolles, Gjermo. The effects of a chlorhexidine toothpaste on the development of plaque, gingivitis and tooth staining. *J Clin Periodontol*, January 1993, 20:59-62.

Chlorhexidine: The Gold Standard Antiplaque Agent

預防口腔相關感染: 口腔照護

The effect of two different dental gels and a mouthwash on plaque and gingival scores: a six-week clinical study

M R Pai, L D Acharya, N Udupa
 Karnataka, India

Table 1 The mean baseline plaque and gingival scores and standard deviations for the four study

Group No.	Formulation	No. of subjects	Baseline mean plaque score	Baseline mean gingival score
I	Control (Placebo Gel)	12	1.66 ± 0.39	1.26 ± 0.24
II	Chlorhexidine mouthwash	12	1.62 ± 0.28	1.24 ± 0.19
III	Chlorhexidine - gel	12	1.63 ± 0.30	1.21 ± 0.22
IV	Neem extract-gel	12	1.55 ± 0.25	1.28 ± 0.31

Table 2 The mean 3-week plaque and gingival scores and standard deviations for the four study

Group No.	Formulation	No. of subjects	Mean plaque score	Mean gingival score
I	Control (Placebo Gel)	12	1.43 ± 0.23	1.13 ± 0.26
II	Chlorhexidine mouthwash	12	1.28 ± 0.20*	1.05 ± 0.21*
III	Chlorhexidine - gel	12	1.11 ± 0.19 ^{ab}	0.82 ± 0.16 ^{ab}
IV	Neem extract-gel	12	1.06 ± 0.13 ^{ab}	0.80 ± 0.24 ^{ab}

Table 3 The mean 6-week plaque and gingival scores and standard deviations for the four study groups

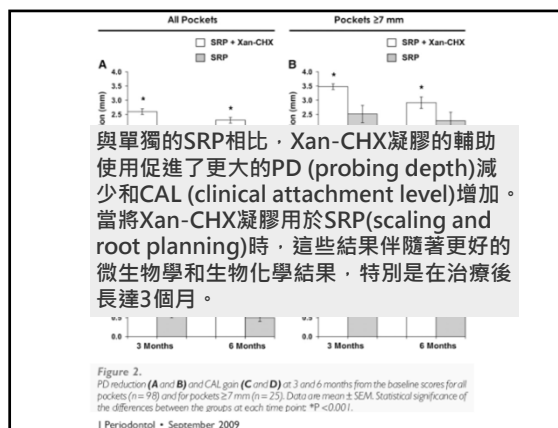
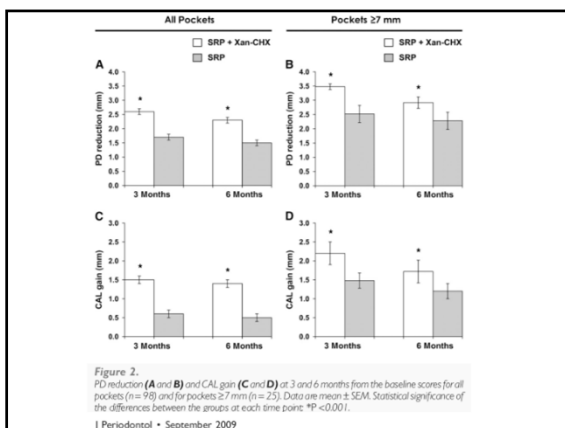
Group No.	Formulation	No. of subjects	Mean plaque score	Mean gingival score
I	Control (Placebo gel)	12	1.31 ± 0.20	1.150 ± 0.26
II	Chlorhexidine mouthwash	12	0.98 ± 0.20*	0.92 ± 0.21*
III	Chlorhexidine gel	12	0.62 ± 0.29 ^{ab}	0.52 ± 0.25 ^{ab}
IV	Neem extract gel	12	0.63 ± 0.24 ^{ab}	0.60 ± 0.28 ^{ab}

International Dental Journal (2004) 54, 219-223

Periodontol • September 2009

Clinical, Microbiologic, and Biochemical Effects of Subgingival Administration of a Xanthan-Based Chlorhexidine Gel in the Treatment of Periodontitis: A Randomized Multicenter Trial

Michele Paolantonio,* Simonetta D'Ercole,[†] Andrea Piloni,[‡] Domenico D'Archivio,* Luca Lisanti,* Filippo Graziani,[§] Beatrice Femminella,* Gilberto Sammartino,^{||} Letizia Perillo,[¶] Stefano Tete,[¶] Giorgio Perfetti,[¶] Giuseppe Spoto,** Raffaele Piccolomini,^{||} and Giuseppe Perinetti^{||††}



- ### CLINICAL APPLICATIONS OF CHLORHEXIDINE
- 輔助口腔衛生和專業預防
 - 牙齦下灌洗
 - 治療義齒性口炎，過敏症與口腔惡臭。
 - 全口腔消毒，具有除牙結石和根部規劃
 - 用於頷間固定的患者和齦齒高風險的患者
 - 在牙周手術或牙根部計劃中進行口腔外科手術：與刷牙和每日牙線相比，每日使用CHX漱口水結合刷牙可減少鄰間牙菌斑
 - 維護牙齒即時功能中的方式：斑塊和出血指數之間的相關性顯示，每日植入物自我護理使用0.2% CHX凝膠，在6個月時有良好結果
 - 易患口腔念珠菌病的高危險患者
 - 限制口腔細菌的菌血症和操作性污染，並作為抗生素的輔助手段。

Thank you for your attention!

