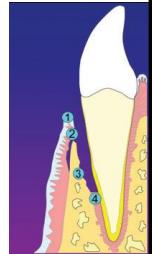


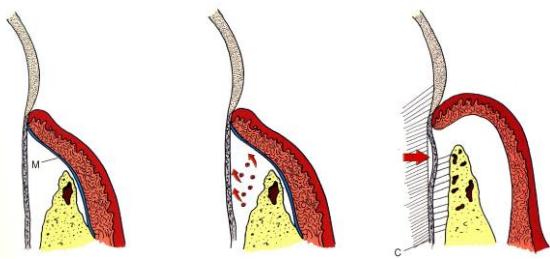
### Biologic Foundation

Compartmentalization (*Melcher AH, 1976*)

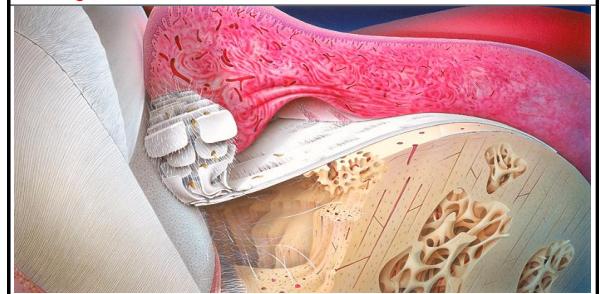
Cell type	Effect
Gingival epithelium	Long junctional epithelium
Gingival connective tissue	Connective tissue attachment Root resorption
PDL cells (mesenchymal cells)	Cementum & PDL fibers
Alveolar bone	Ankylosis



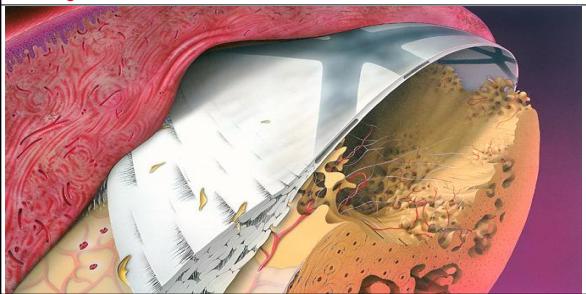
### Biologic Foundation



### Biologic Foundation

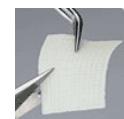


### Biologic Foundation



### Membranes

Non-resorbable membranes



Bio-resorbable membranes



### Membranes... Requirements

- Biocompatibility
- Cell exclusion
- Space maintenance
- Tissue integration
- Ease of use & handling
- Biological activity



### Membranes... Non-Resorbable

Expanded polytetrafluoroethylene (ePTFE)

Miscellaneous membranes  
Millipore membrane  
Rubber dam

### Membranes... Non-Resorbable

Preservation of keratinized gingiva

Thick overlying surgical flap



### Membranes... Non-Resorbable

Healing is allowed for 4-6 weeks after membrane placement (longer periods are better; 12-16 weeks)

No probing for 3-6 months after membrane removal  
Radiographic evidence of bone formation 6-12 months

PERFECT ORAL HYGIENE / TISSUE PERFORATION

Infection → Membrane removal

### Membranes... Non-Resorbable



### Membranes... Bio-Resorbable

Polyglycoside synthetic membranes  
(Polylactic acid, polylactate/polygalactide copolymers)

Collagen

Type I or Types I + III (Porcine or Bovine)

Calcium sulfate

## Membranes... Bio-Resorbable

They are easier to manage

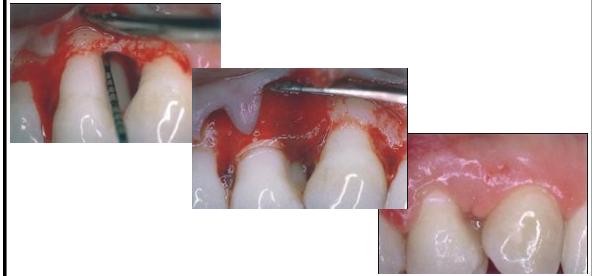
More tissue compatible than non-resorbable membranes

Timing for resorption can be regulated

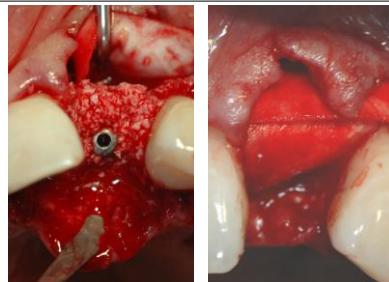
Second surgery for membrane removal is not required

Lack of rigidity

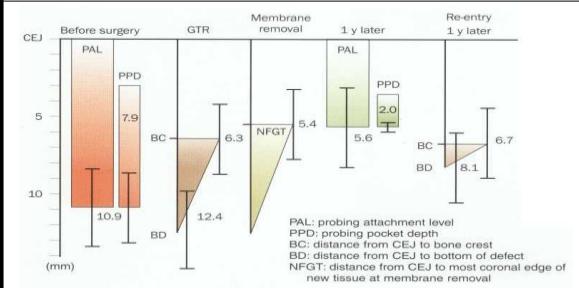
## Membranes... Bio-Resorbable



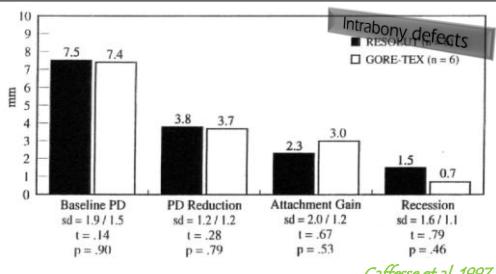
## Membranes... Bio-Resorbable



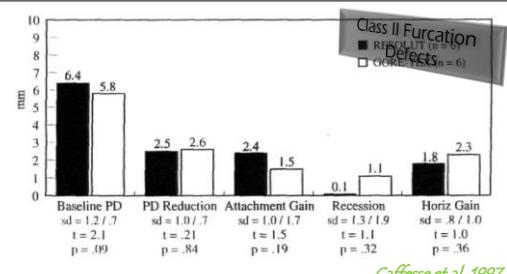
## Membranes... Results



## GTR... Results



## GTR... Results



Membranes... Non-Resorbable vs. Bio-Resorbable			
Membrane category	Advantages	Disadvantages	Commercial examples
Nonresorbable	<ul style="list-style-type: none"> <li>Numerous studies demonstrate their success</li> <li>May be titanium reinforced</li> <li>Hemostatic intact until removal</li> <li>Easy to place with titanium or resorbable tacks</li> <li>Greater bone fill if membrane not exposed</li> <li>Minimal tissue response if membrane not exposed</li> </ul>	<ul style="list-style-type: none"> <li>Require a second surgery for removal</li> <li>Increase patient morbidity</li> <li>If exposed, must be removed</li> <li>Can be technique sensitive</li> </ul>	<ul style="list-style-type: none"> <li>ePTFE membranes, e.g., Gore-Tex (Gore Medical, Flagstaff, Ariz.)</li> <li>Titanium-reinforced Gore-Tex</li> </ul>
Resorbable	<ul style="list-style-type: none"> <li>Numerous studies demonstrate their success</li> <li>Does not require surgical removal</li> <li>Decreased patient morbidity</li> <li>Improved soft-tissue healing</li> <li>Tissue ingrowth reduction to membrane exposure</li> <li>Cost effective; one surgery only</li> <li>Does not have to be removed if exposed</li> </ul>	<ul style="list-style-type: none"> <li>Uncertain duration of barrier membrane function</li> <li>Difficult to tack down</li> <li>Slightly less bone fill than nonresorbable membranes</li> <li>Inflammatory reaction from healing GBR</li> <li>Can be technique sensitive</li> </ul>	<ul style="list-style-type: none"> <li>Nescon (bovine collagen matrix; Citegenix Inc., Laval, Que.)</li> <li>Bio-Gide (porcine collagen matrix; Geistlich AG, Wolhusen, Switzerland)</li> <li>Ossis (cross-linked collagen barrier; Implant Innovations Inc., Palm Beach Gardens, Fla.)</li> </ul>

Trejo et al, 2000

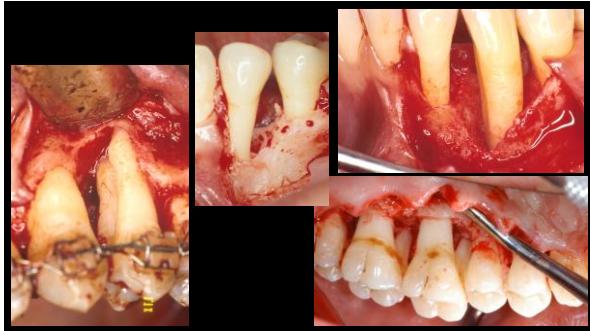
GTR... Membranes +/- Bone Grafts			
	PD Reduction	CAL Gain	REC Change
6 Months			
GTR with DFDBA (Test) (N = 16)	3.64 (1.23) P = 0.41	2.72 (0.84) P = 0.32	-0.85 (0.98) P = 0.92
GTR alone (Control) (N = 14)	4.00 (1.12)	3.11 (1.19)	-0.88 (0.89)
12 Months			
GTR with DFDBA (N = 15)	3.37 (1.16) P = 0.06	2.29 (0.61) *P = 0.008	-1.08 (1.07) P = 0.54
GTR alone (N = 14)	4.12 (0.84)	3.27 (1.10)	-0.85 (0.91)

Trejo et al, 2000

GTR... Membranes +/- Bone Grafts				
	Resorption	Bone Fill	Defect Resolution	Percent Defect Fill
GTR with DFDBA (N = 15)	-1.10 (1.22) P = 0.223	3.72 (2.03) P = 0.262	4.73 (1.18) P = 0.455	65.5
GTR alone (N = 13)	-0.61 (0.77)	4.85 (3.14)	5.35 (2.91)	58.4

Trejo et al, 2000

## New Approaches to Periodontal Regeneration



New Approaches to Periodontal Regeneration			
Enamel Matrix Derivative (EMD)	Growth factors	Platelet-Rich Plasma (PRP)	Bone Morphogenetic Proteins (BMPs)
			Gene Therapy
			Tissue Engineering

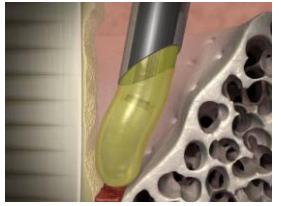
... Enamel Matrix Derivative

- Induction of cementogenesis
- Hertwig's Epithelial Root Sheath
- Porcine origin
- Amelogenins + Ameloblastin + Enamelin + PGA
- Precipitates to the root surface



... Enamel Matrix Derivative

- Root conditioning with PrefGel®
- Application of Emdogain®



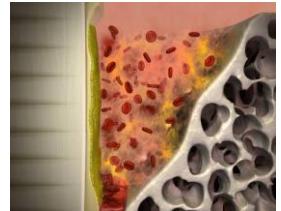
... Enamel Matrix Derivative

- Precipitation of amelogenins on root surface (matrix formation)



... Enamel Matrix Derivative

- Clot formation
- Granulation tissue



... Enamel Matrix Derivative

- Migration & adhesion of mesenchymal stem cells (MSCs)
- Proliferation of MSCs



... Enamel Matrix Derivative

- Cytokine production
- Proliferation & differentiation of MSCs

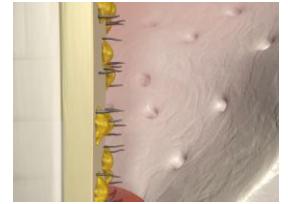


**... Enamel Matrix Derivative**

Differentiation into cementoblasts  
Deposition of cementum

**... Enamel Matrix Derivative**

Insertion of periodontal ligament fibers into newly-formed cementum

**... Enamel Matrix Derivative**

Filling of defect with newly-formed periodontal tissues

**... Enamel Matrix Derivative**

Parallel formation of alveolar bone in the defect

**... Enamel Matrix Derivative**

Periodontal regeneration with a new functional attachment

**... Enamel Matrix Derivative**

Regeneration of cementum  
Bone formation  
PDL formation  
No recession  
No junctional epithelium

### ... Enamel Matrix Derivative

Table 5. Baseline defect characteristics expressed in mm (mean  $\pm$  SD)

Treatment	PPD	GR	CAL	CEJ-BBD	CEJ-crest	Intrabony depth
EMD	8.2 $\pm$ 1.1	1.7 $\pm$ 1.3	9.9 $\pm$ 1.4	11.0 $\pm$ 1.9	7.1 $\pm$ 1.2	3.9 $\pm$ 1.5
GTR	8.3 $\pm$ 1.3	1.6 $\pm$ 1.4	9.9 $\pm$ 1.7	10.8 $\pm$ 1.8	7.0 $\pm$ 1.3	3.8 $\pm$ 1.7
EMD+GTR	8.4 $\pm$ 1.0	1.4 $\pm$ 0.8	9.8 $\pm$ 1.2	10.9 $\pm$ 1.8	7.2 $\pm$ 1.3	3.7 $\pm$ 1.5
OFD	8.2 $\pm$ 1.1	1.5 $\pm$ 0.7	9.7 $\pm$ 0.8	10.7 $\pm$ 1.9	6.9 $\pm$ 1.8	3.8 $\pm$ 1.2

Table 6. Changes of clinical parameters at 1 and 5 years compared to baseline (mean  $\pm$  SD)

Parameter	OFD	EMD	GTR	EMD+GTR
$\Delta$ PPD (mm) 1 year	3.3 $\pm$ 1.1	4.6 $\pm$ 1.2	4.4 $\pm$ 1.4	4.4 $\pm$ 0.8
$\Delta$ PPD (mm) 5 years	2.7 $\pm$ 1.2	4.3 $\pm$ 1.7	3.9 $\pm$ 1.6	4.0 $\pm$ 1.0
$\Delta$ GR (mm) 1 year	1.7 $\pm$ 0.5	1.3 $\pm$ 0.6	1.3 $\pm$ 1.0	1.5 $\pm$ 0.7
$\Delta$ GR (mm) 5 years	1.7 $\pm$ 0.5	1.3 $\pm$ 0.7	1.2 $\pm$ 1.0	1.5 $\pm$ 0.7
$\Delta$ CAL (mm) 1 year	1.6 $\pm$ 1.0	3.4 $\pm$ 1.1	3.2 $\pm$ 0.8	3.0 $\pm$ 1.0
$\Delta$ CAL (mm) 5 years	1.3 $\pm$ 1.2	2.9 $\pm$ 1.6	2.7 $\pm$ 0.9	2.6 $\pm$ 0.7

Sculean et al, 2006

### ... Growth Factors

	Platelet-derived growth factor	Fibroblast growth factor-2	Bone morphogenetic proteins	Enamel matrix derivative	Transforming growth factor-beta	Insulin-like growth factor-1, 2
Periodontal ligament cells						
Cell proliferation	++	+++	++	++	-	+
Chemotaxis	++	+++	+	++	0	++
Collagen synthesis	+	-	+	+	+	+
Protein synthesis	+	+	+	+	+	+
Matrix gene expression	++	++/-	?	+	+	+

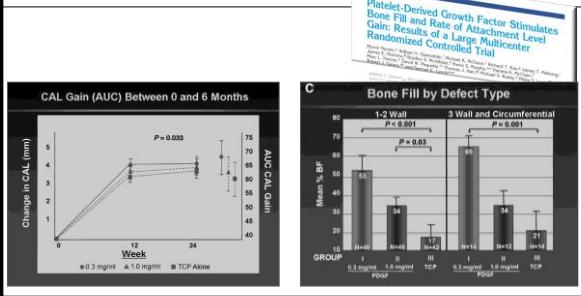
### ... Growth Factors

	Platelet-derived growth factor	Fibroblast growth factor-2	Bone morphogenetic proteins	Enamel matrix derivative	Transforming growth factor-beta	Insulin-like growth factor-1, 2
Cementoblasts						
Cell proliferation	+++	?	-	++	++	++
Chemotaxis	++	?	?	?	?	?
Collagen synthesis	+	?	++	++	+	+
Protein synthesis	+	?	++	++	+	+
Matrix gene expression	+/-	?	++	++/-	+/-	+/-

### ... Growth Factors

	Platelet-derived growth factor	Fibroblast growth factor-2	Bone morphogenetic proteins	Enamel matrix derivative	Transforming growth factor-beta	Insulin-like growth factor-1, 2
Osteoblasts						
Cell proliferation	++	+++	0	++	+++	++
Chemotaxis	+++	+++	+	++	+++	+
Collagen synthesis	0	++	0	+	++	+
Protein synthesis	0	+	ND	+	+/-	0
Matrix gene expression	+/-	++/-	++	++/-	++	++
Alkaline phosphatase synthesis	0	-	++	++	+/-	0

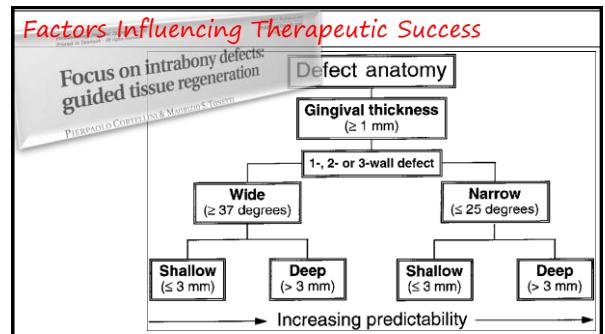
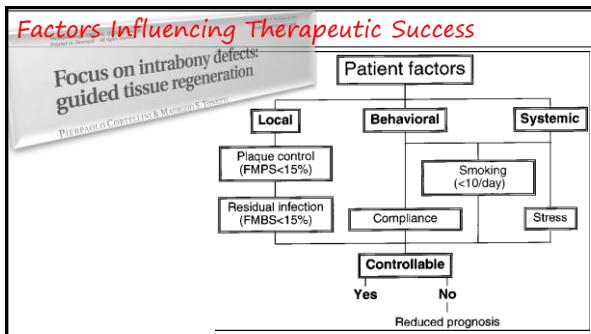
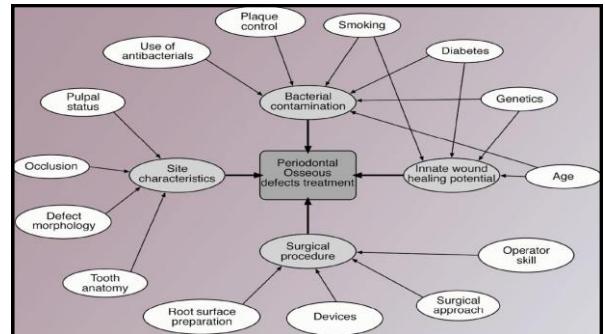
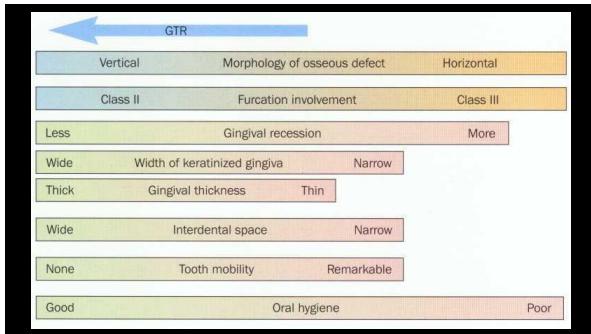
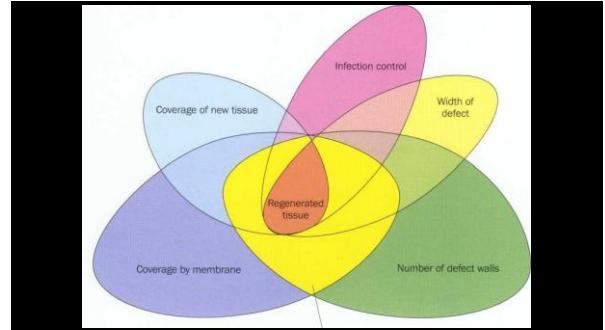
### ... Growth Factors



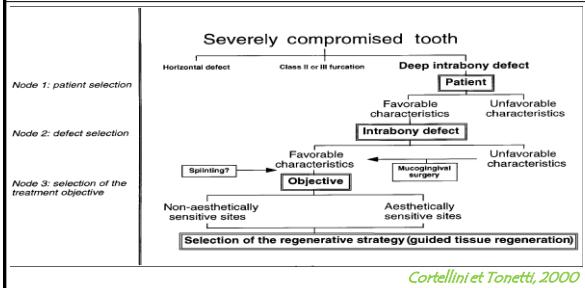
### ... Growth Factors



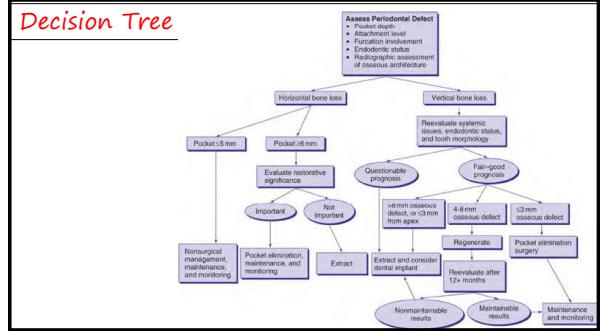
## Factors Affecting GTR



### Selection... Patient, Defect, Objective



### Decision Tree



Thank You...