

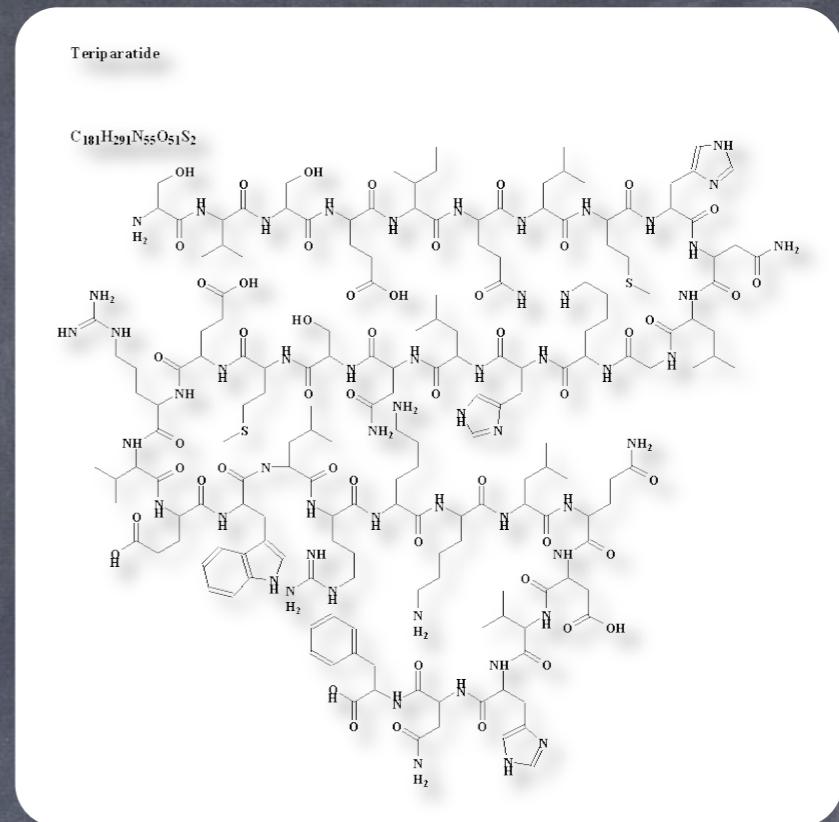


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TERIPARATIDE THERAPY FOR BISPHOSPHONATE-RELATED OSTEONECROSIS OF THE JAW: OUR EXPERIENCE ON THREE CASES

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Teriparatide is a synthetic polipeptidic hormone, which contains the amino-acidic fragment of the 1-34 parathyroid recombinant hormone (rhPTH 1-34).



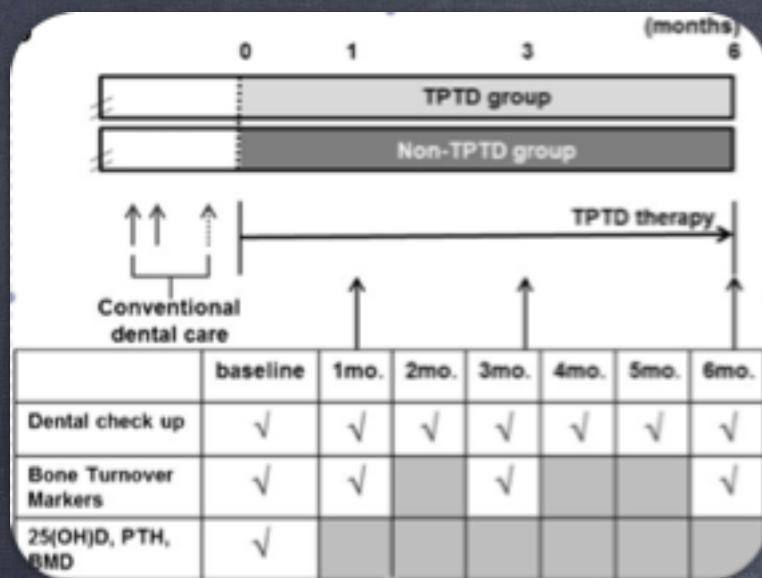
Pulsatile exposure to lower doses actually results in increased bone mineral density (BMD) and bone mass. The off-label use of Teriparatide in BRONJ patients affected from osteoporosis has recently been reported in literature.

PUBBLICAZIONE	ETÀ/SEX	CONDIZIONE MEDICA E ALTRI FARMACI	BISFOSFONATO	SPONTANEA VS POSTESTRATTIVA	STADIO DELLA BRONJ	DURATA TERAPIA	DOCUMENTAZIONE RADIOGRAFICA
Harper e Fung (2007)	75/F	Osteoporosi simvastatina, rofecoxib, docusate sodium, zolpidem	Alendronato (70mg giornalieri) per 1 anno	Postestrattivo	Stadio II Mascellare	3 mesi	10 mesi OPT
Lau ed Adachi (2009)	56/F	Fibromialgia, osteoporosi indotta da glucocorticoidi, osteoartrite prednisone 18 mesi	Etidronato, calcitonina, clodronato, pamidronato, zoledronato per 3 aa	Postestrattivo	Stadio III Mascellare	Non riferito (10 mesi OPT)	10 mesi OPT
Narongroeknawn et al. (2010)	63/M	Osteoporosi esomeprazolo, glucosamina, chondroitina, atorvastatina, cardioaspirina, mirtazapina	Alendronato per 5 aa	Postimpianto	Stadio I Mascellare	4 mesi	4 mesi peripendale
Lee et al (2010)	78/F	Osteoporosi cardioaspirina	Alendronato 70 mg/set per 5 aa	Postestrattivo	Stadio II Mandibolare	4 sett.	6 mesi OPT
Tsai et al (2010)	72/F	Non dichiarata	Alendronato settimanale per 4 aa	Postimpianto	Stadio III Mandibolare	4 mesi	5 mesi OPT
Cheung et al (2010)	88/F	Osteoporosi prednisone per 20 aa	Alendronato settimanale per 10 aa	Postestrattivo	Stadio II/III Mandibolare	8 sett.	CT scan a 8 settimane
Kwon et al (2011)	88/M	Osteoporosi	Risendronato per 5 anni		Stadio II Mandibolare	1 mese	
Kwon et al (2011)	74/F	Osteoporosi methotrexate, prednisolone, mizoribine	Alendronato per 8 anni		Stadio III Mandibolare	3 mesi	
Kwon et al (2011)	80/F	Osteoporosi	Alendronato per 3 anni		Stadio III Mandibolare	2 mesi	
Kwon et al (2011)	76/F	Osteoporosi glicazide	Alendronato e ibandronato per 3 anni		Stadio II Mandibolare	1 mese	
Kwon et al (2011)	79/F	Osteoporosi amlodipina, metformina	Alendronato per 5 anni		Stadio II Mandibolare	2 mesi	
Kwon et al (2011)	68/M	Osteoporosi	Alendronato per 4 anni		Stadio II Mascellare	3 mesi	

DISTINCTIVE ROLE OF 6-MONTH TERIPARATIDE TREATMENT ON INTRACTABLE BISPHOSPHONATE-RELATED OSTEONECROSIS OF THE JAW

K.M.Kim, W.Park, S.Y.Oh, H.-J.Kim, W.Nam, S.-K.Lim, Y.Rhee, I.H.Cha

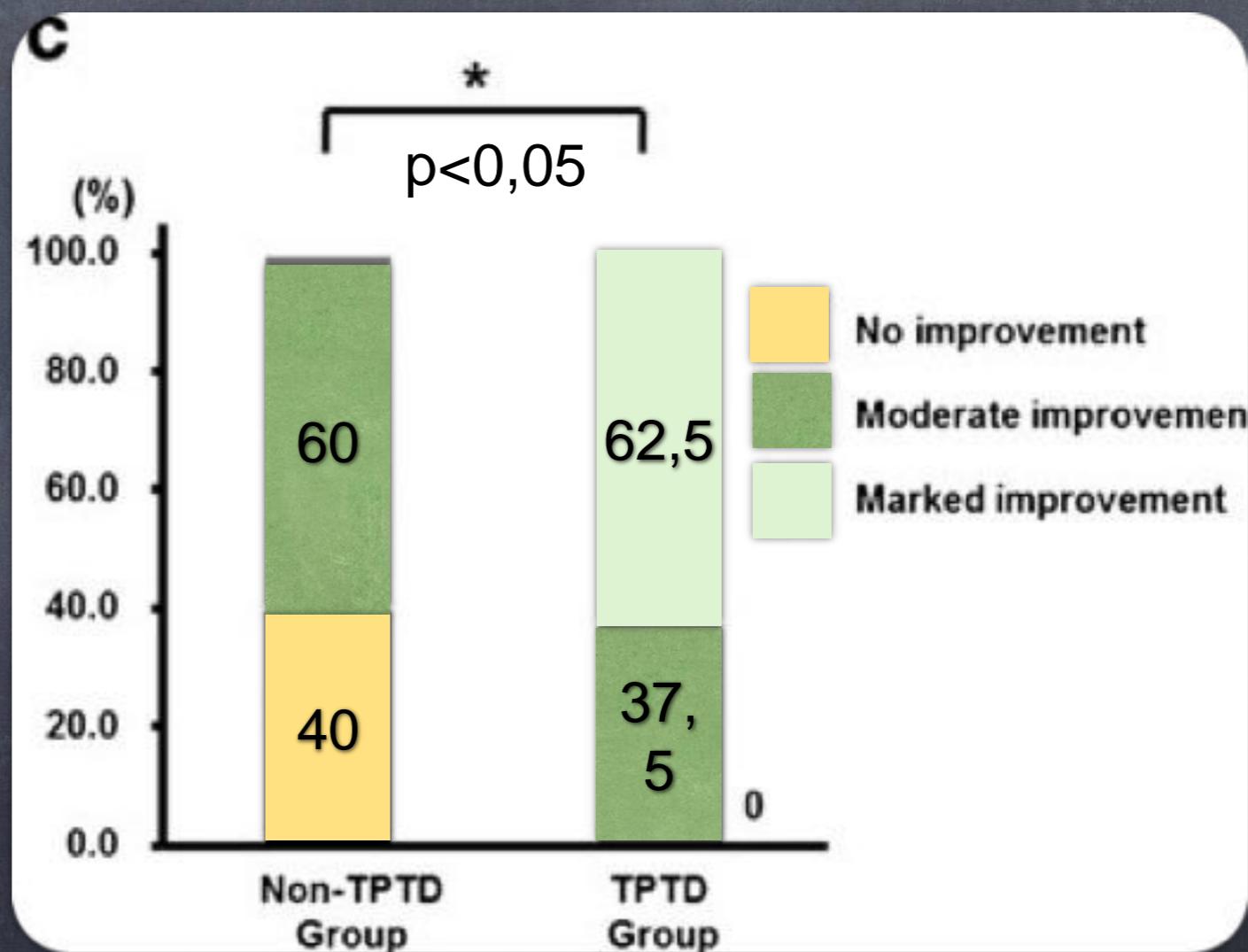
Osteoporos Int 2014



24 BRONJ

15
TPTD +
Ca-Vit D

9
TPTD-
Ca-Vit D



Patients with higher baseline serum 25(OH)D levels showed better clinical therapeutic outcomes with TPTD

METHODS

- Visit and anamnesis
- Clinical and radiological assessment of BRONJ
- Drug holiday
- Endocrinologist advice
 - 1. spine x-ray, phosphatase,
 - 2. complete blood count, 8. uric acid ,
 - 3. serum protein electrophoresis, 9. TSH,
 - 4. determinations of serum calcium, 10. PTH,
 - 5. creatinine, 11. 24-h urinary
 - 6. phosphorus, calcium
 - 7. alkaline
- Monthly follow-up

	SEX/ AGE	SINTOMATOLOGY OSTEOPOROSIS	BF	DURATION DEL OF THE TREATMEN T WITH BF	CONCURRENT PATHOLOGIES	CONCURREN T PHARMACOL OGICAL THERAPIES	TERIPARATID E
Pz. 1	F/69	2007 right wrist fracture. NOT OSTEOPOROTIC PATIENT!!	Alendronate e Vit D3 (Fosavance)	11/2009 12/2010 1 YEAR	arterial hypertension, arthrosis, mild hyperglycemia, premature menopause	ASA, flosinopril, spironolactone and furosemide, allopurinol and pantoprazole	7 months



Extraction s 46 - 47



II stadium (AAOMS)
II stadium (SIPMO-SICMF)



Starts FORSTEO

Pain and swelling disappeared

Ends FORSTEO
(7th month)

APRIL 2

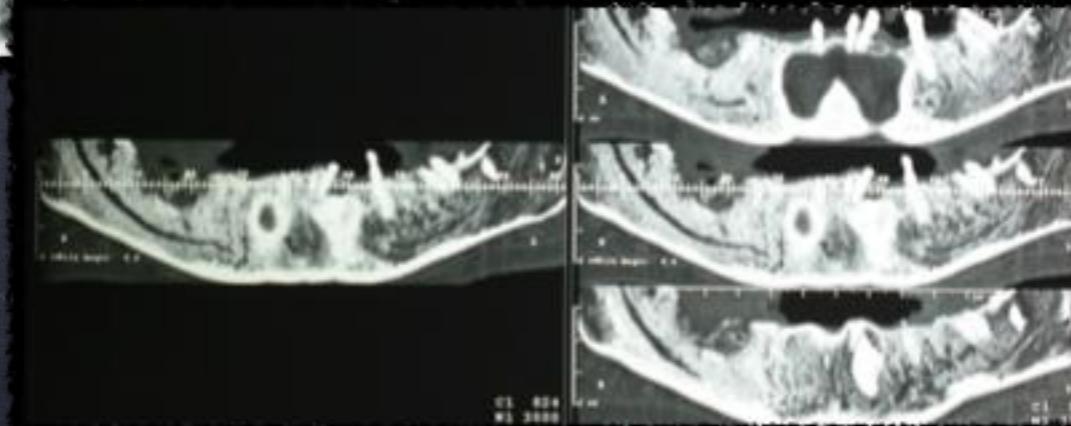
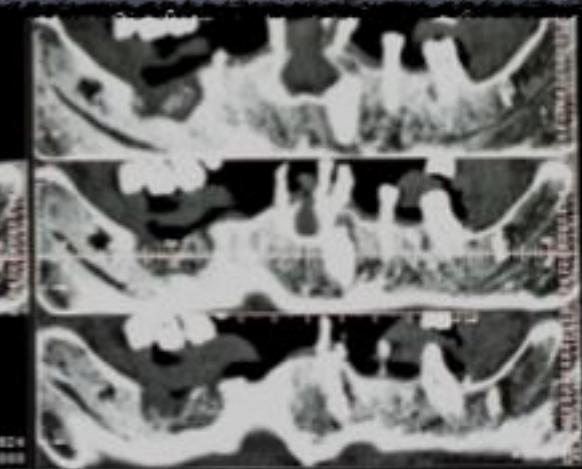
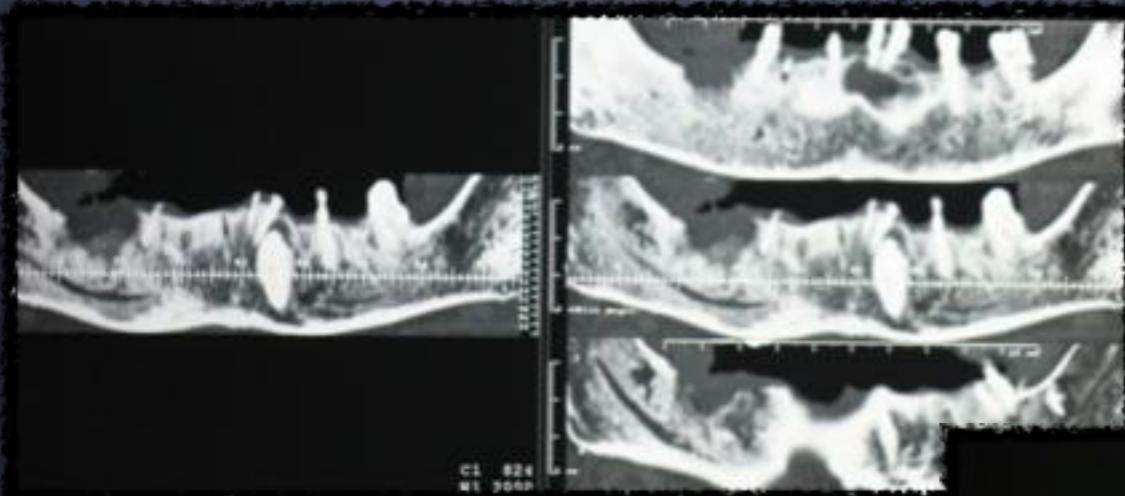
MAY

JUNE

1

OCTOBER

Control
(4th month)



Starts FORSTEO

Surgery

APRIL

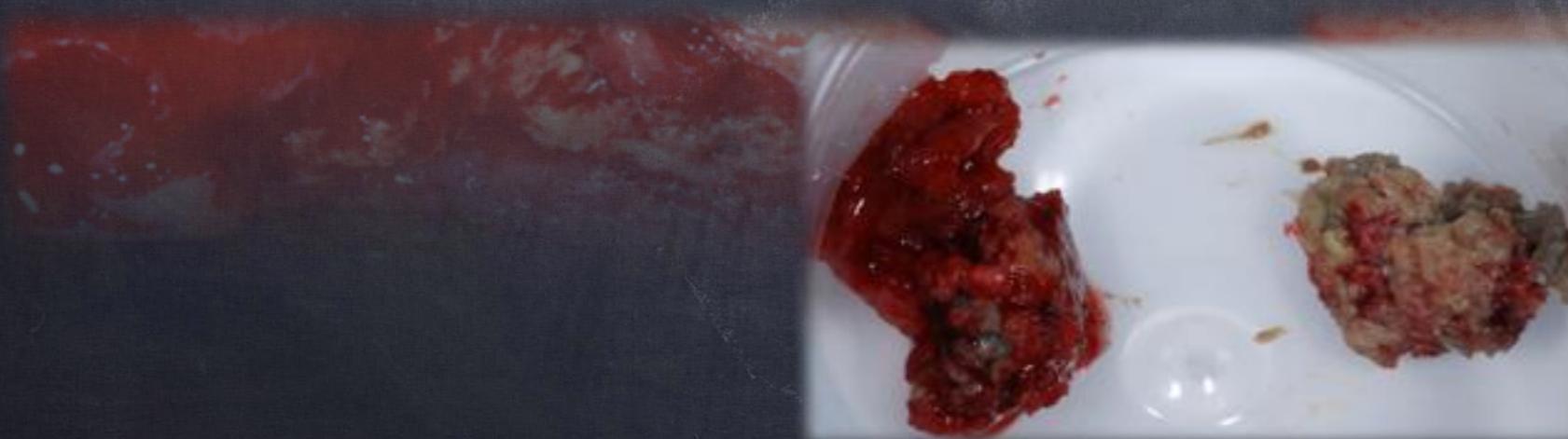
2

0

1

1

NOVEMBER



Starts FORSTEO
Ends FORSTEO
(7th month) Surgery

Control

Control

2 APRIL 0

1 OCTOBER 1

NOVEMBER

APRIL

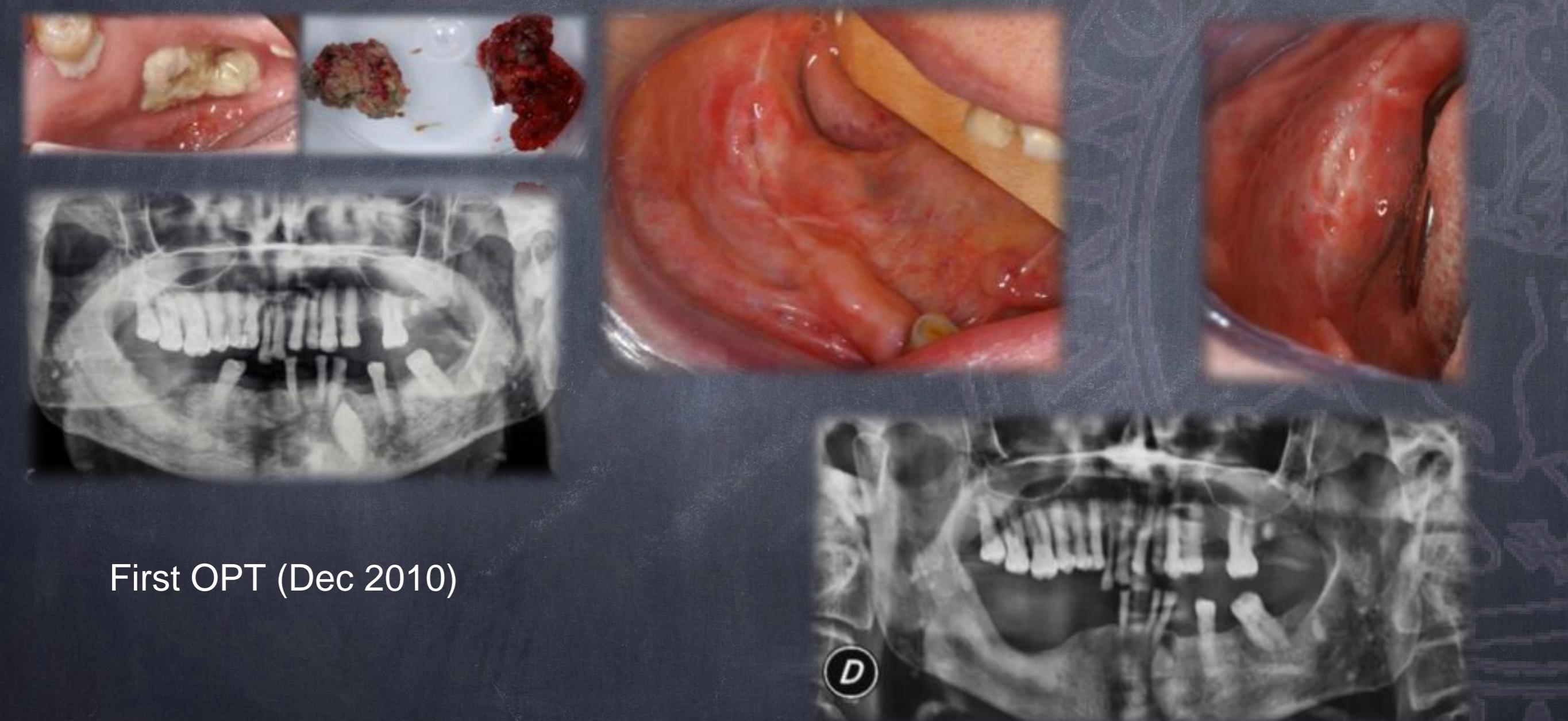
2

0

1

2

OCTOBER



OPT 12 month after surgery (10/12)

	SEX/ AGE	SINTOMATOLOGY	BF	DURATION DEL OF THE TREATMEN T WITH BF	CONCURRENT PATOLOGIES	CONCURRENT PHARMACOLO GICAL THERAPIES	TERIPARATID E
Pz. 2	F/80	Several bone fractures (femur, humerus, ribs) Multiple vertebral collapses	Alendronate (Adronat)	2004 9/2011 7 ANNI	Epilepsy, psoriasis and osteoporosis	Fenobarbital, lorazepam and loratadine, Prednisone	9 months



Extractions 36-37



II stadium (AAOMS)
II stadium (SIPMO-SICMF)

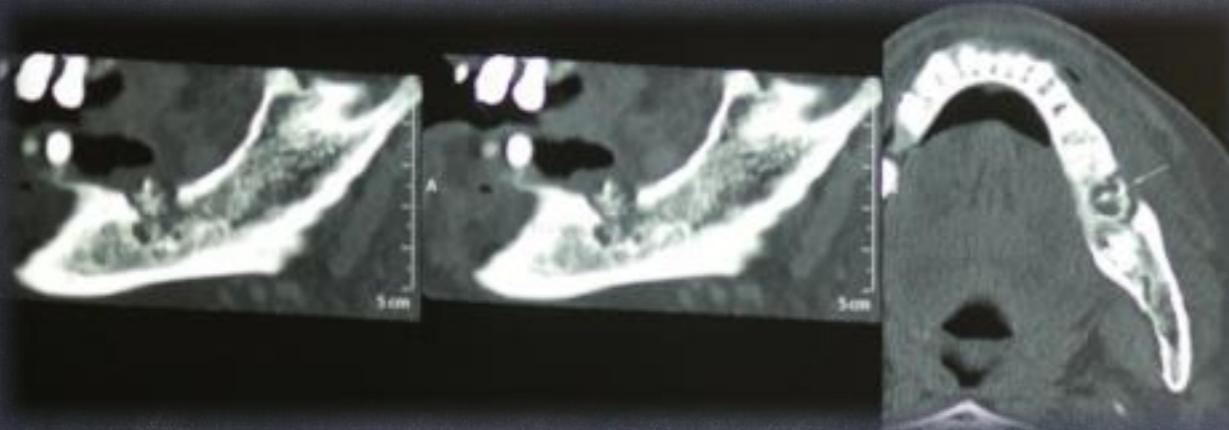
Starts FORSTEO

Pain and swelling disappeared

2 OCTOBER 1 NOVEMBER 1 DECEMBER

FEBRUARY

0 1 2



Control
(4th month)



FEBRUARY

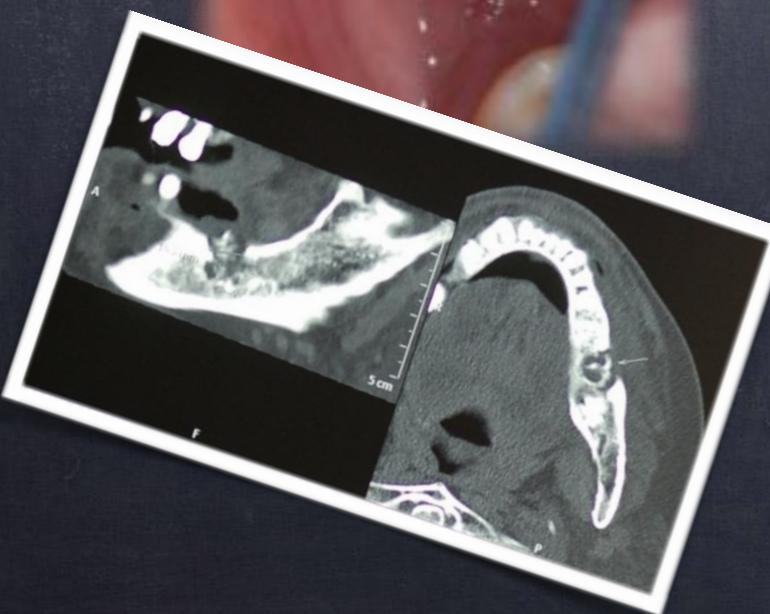
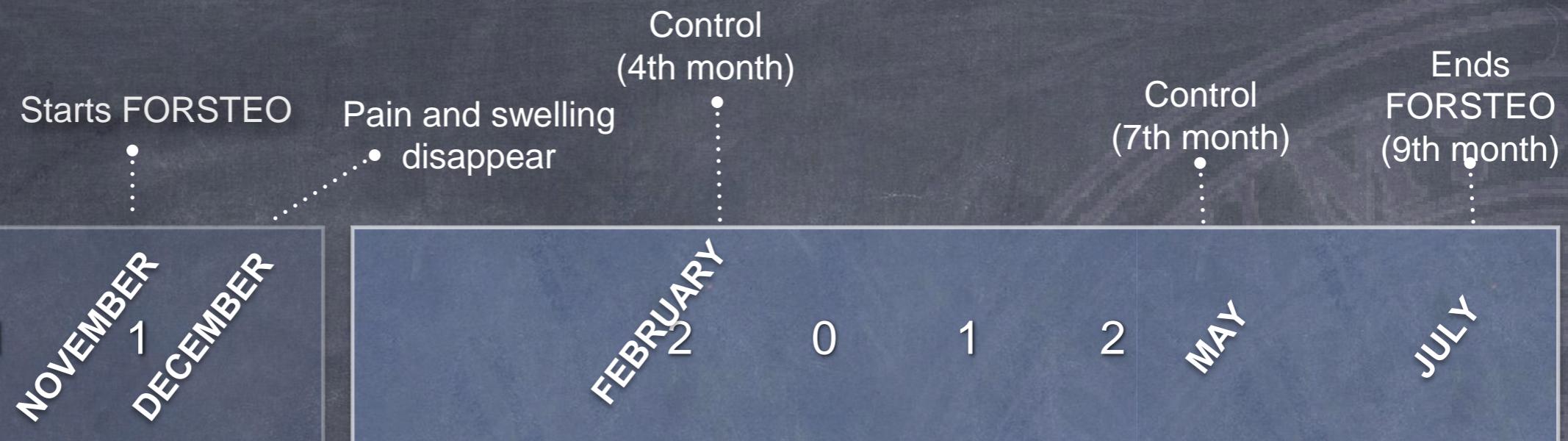
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2

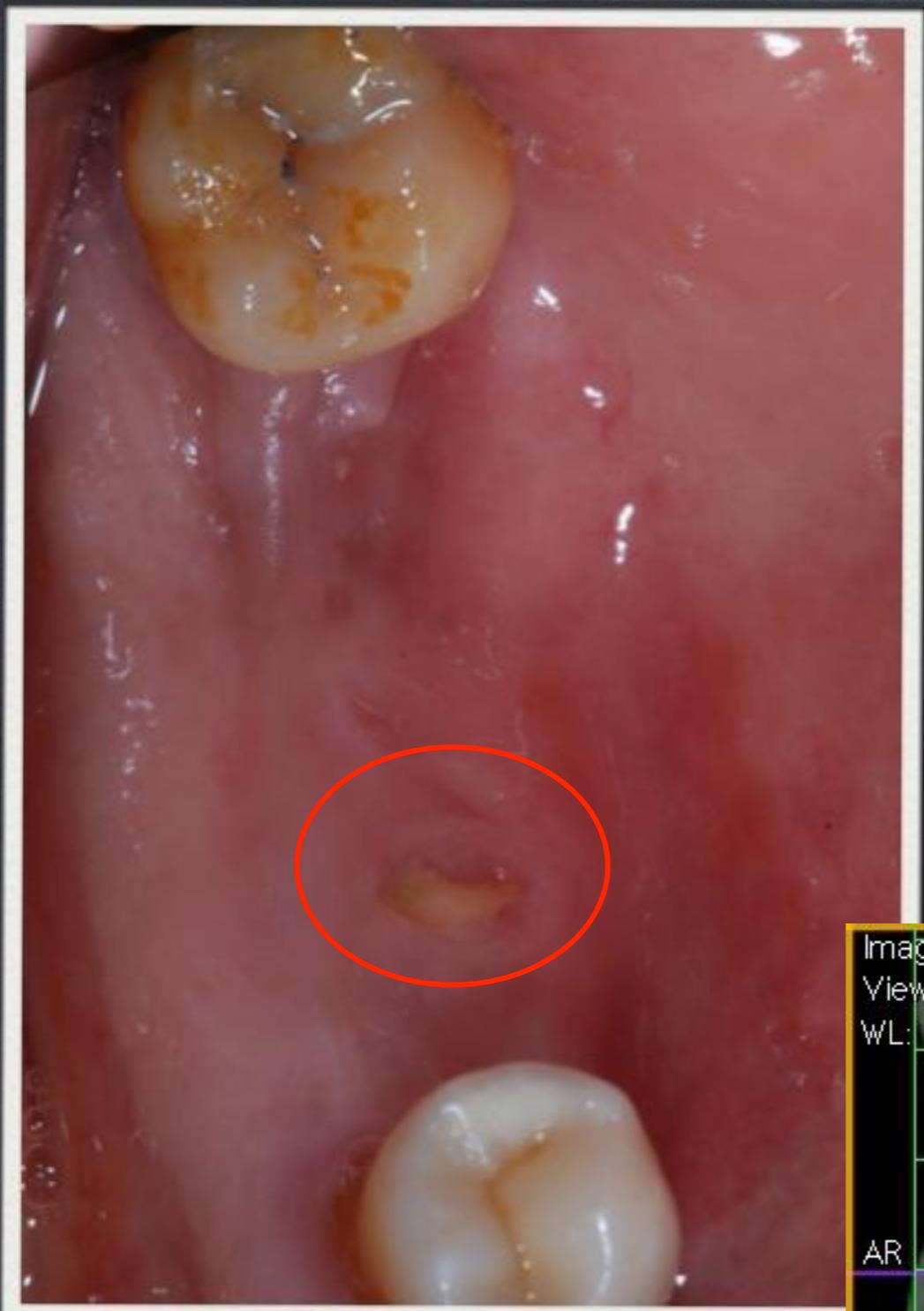






SEPTEMBER 2012





OCTOBER 2012

Amoxicillin 1 gr X 3 X 6



SEX/A SINTOMATOLOGY
GE OSTEOPOROSIS

BF

DURATION
DEL OF THE
TREATMENT
WITH BF

CONCURRENT
PATOLOGIES

CONCURRENT
PHARMACOLO
GICAL
THERAPIES

TERIPARATID
E

Pz. 3

F/90

multiple vertebral
collapses

Ibadronato
(Bonviva)

2005
10/2012
7 YEARS

Hypertensive
cardiomyopathy,
mitral fibrosis,
dispnea,
esophagitis and
iron deficiency
anemia

Oxybutin, eosina,
homeoplasmine,
haloperidol,
promazine and
quetiapine

3 months

SEPTEMBER

2 0 1 2

III stadium (AAOMS)
III stadium (SIPMO-SICMF)



Start FORSTEO

Control
(3rd month)

SEPTEMBER

OCTOBER

2

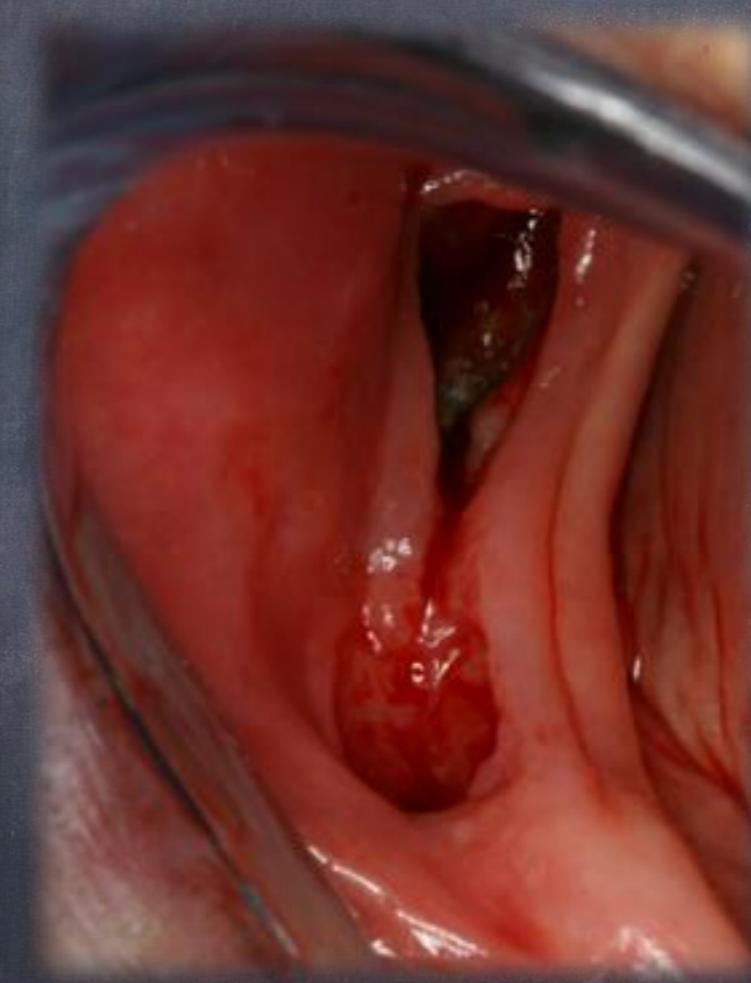
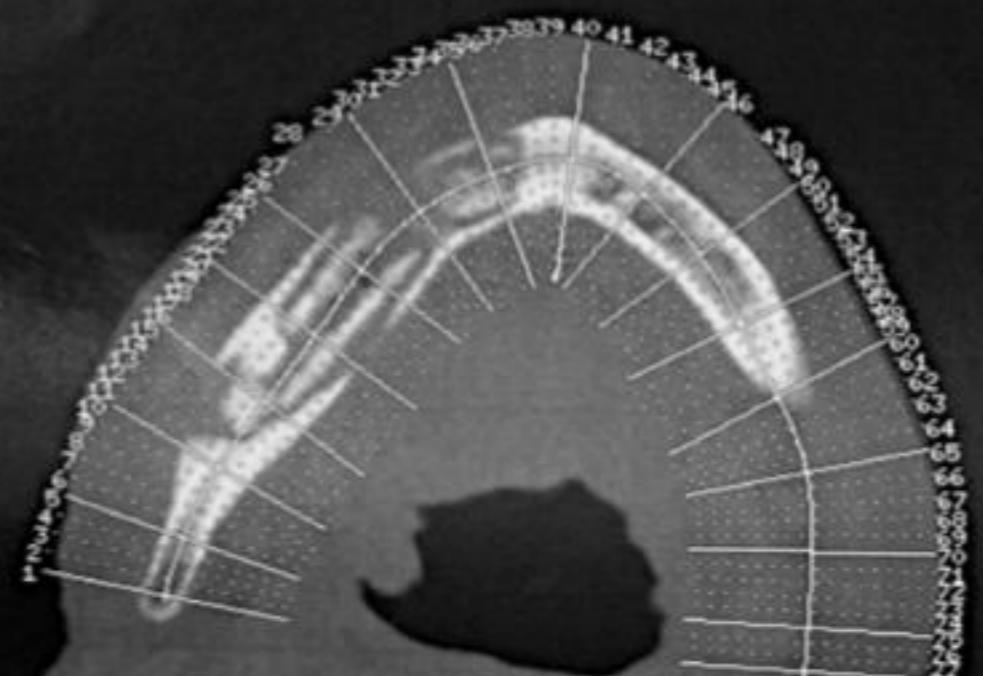
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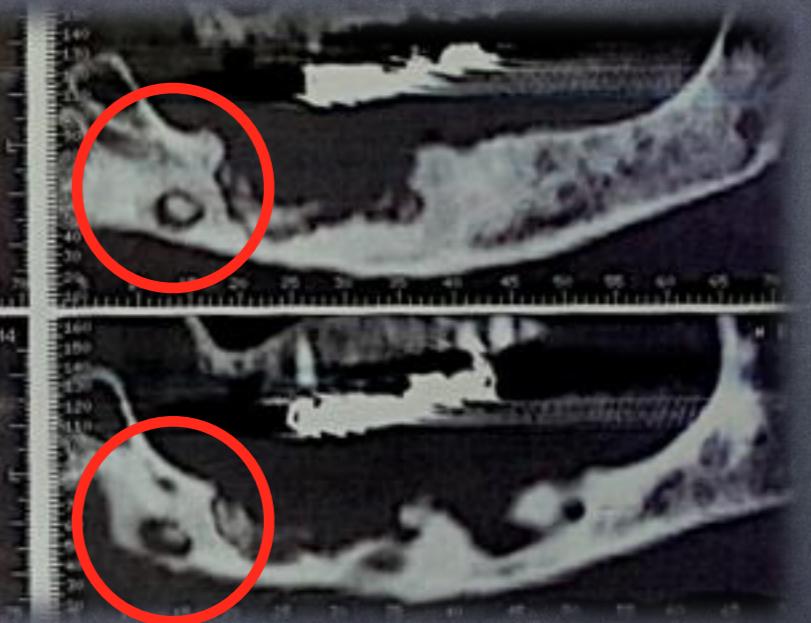
1

2

DECEMBER

: 51
OV 15.3 cm
ME





The cutaneous fistula did not resolved

CONCLUSIONS

- Reduction of pain and swelling in the early two weeks from the beginning of the therapy
- Reprise of bone remodeling, evidenced by the release of bone sequestra and by a better definition of the cortical contours in TC and OPT
- Although the application of teriparatide as an adjunctive modality is an opinion-based approach rather than evidence-based one, we might validate this short-term teriparatide therapy for the BRONJ patients as an adjunctive modality.

**FOR THIS PURPOSE FURTHER INVESTIGATIONS AND
CONTROLLED STUDIES SHOULD BE PERFORMED.**

Grazie per l'attenzione...



Bone homeostasis involves 3 key biological pathways:

- the estrogen-endocrine pathway that preserves BMD
- the canonical Wnt/ β -catenin signaling pathway, a major signaling pathway that facilitates bone formation
- the receptor activator of NF- $\kappa\beta$ ligand/receptor activator of NF- $\kappa\beta$ /osteoprotegerin (RANKL/RANK/OPG) pathway that determines the balance between bone formation and resorption