



76 year-old female presents
with muscle cramps

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12/6/12

HPI

- Worked up for outpatient hypercalcemia
- Calcium had been 10.3-11.1, PTH ~120
- No h/o osteoporosis, CKD, kidney stones
- Not taking calcium supplements
- Sestamibi scan: no evidence of parathyroid adenoma
- Diagnosed w/hyperparathyroidism secondary to a parathyroid adenoma

More history

Past Medical

HTN

Dyslipidemia

Asthma

Hyperparathyroidism

Family

Father- TB @ 36

Mother- CHF

Social

No tobacco

No EtOH

Medications

ASA 81mg daily

Losartan 50mg daily

Vitamin D 2000 IU

Referred for parathyroidectomy

- Immediate pre-op sestamibi scan: re-demonstrated no evidence of parathyroid adenoma.
 - Pre-op notes: probable location of the parathyroid adenoma was in the R inferior quadrant.
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Surgery

- Neck exploration of the R inferior region- R inferior parathyroidectomy
 - “Attention was directed to the L side where the pre-op studies indicated a probable adenoma. Dissection was carried out on the R inferior thyroid surrounding tissue in addition to the L side.”
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Post-op course

- POD # 0, extubated but suffered acute respiratory failure
 - Required tracheostomy placement for B vocal cord paralysis
 - G-tube placed
 - R inferior parathyroid pathology- abnormal parathyroid tissue consistent w/parathyroid adenoma
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Physical Exam

Vitals: 36.3, 127/71, 70, 16, 98% trach collar

Gen: no apparent distress

HEENT: hearing aids

Neck: tracheostomy

CV: RRR, no murmurs

Pulm: clear bilaterally

GI: g-tube, soft non-tender

MSK: normal ROM joints

Neuro: alert and oriented, normal reflexes

No Chvostek or Trousseau's Sign

Labs

141	98	16	96	9.9
4.1	34	0.6		
			6.4	7.8
			4.1	275
			1.7	30

7.0	3.7
0.3	57
31	29

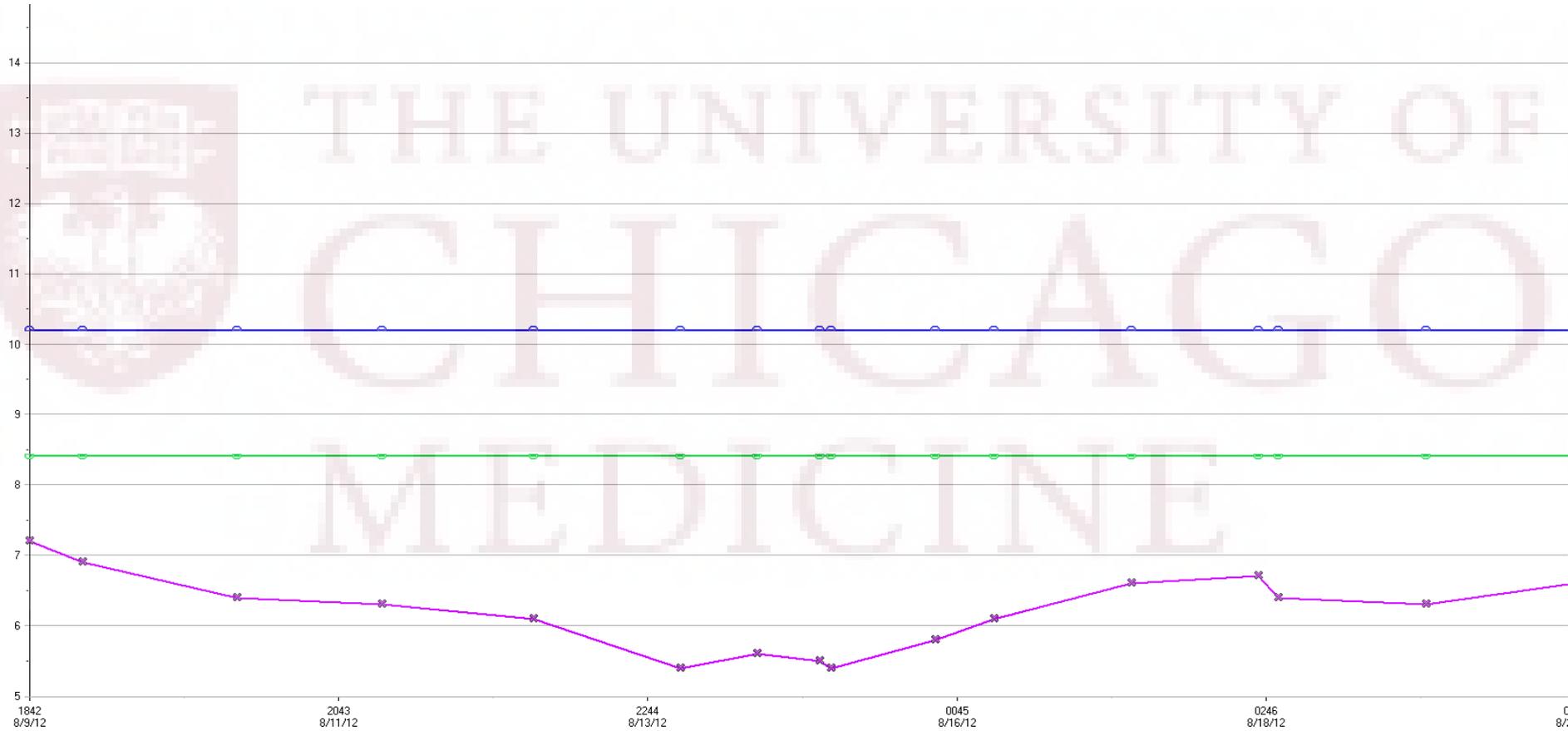
PTH 6 (15-75)

Ionized calcium 3.28 (4.6-5.4)

25-OH vitamin D 40

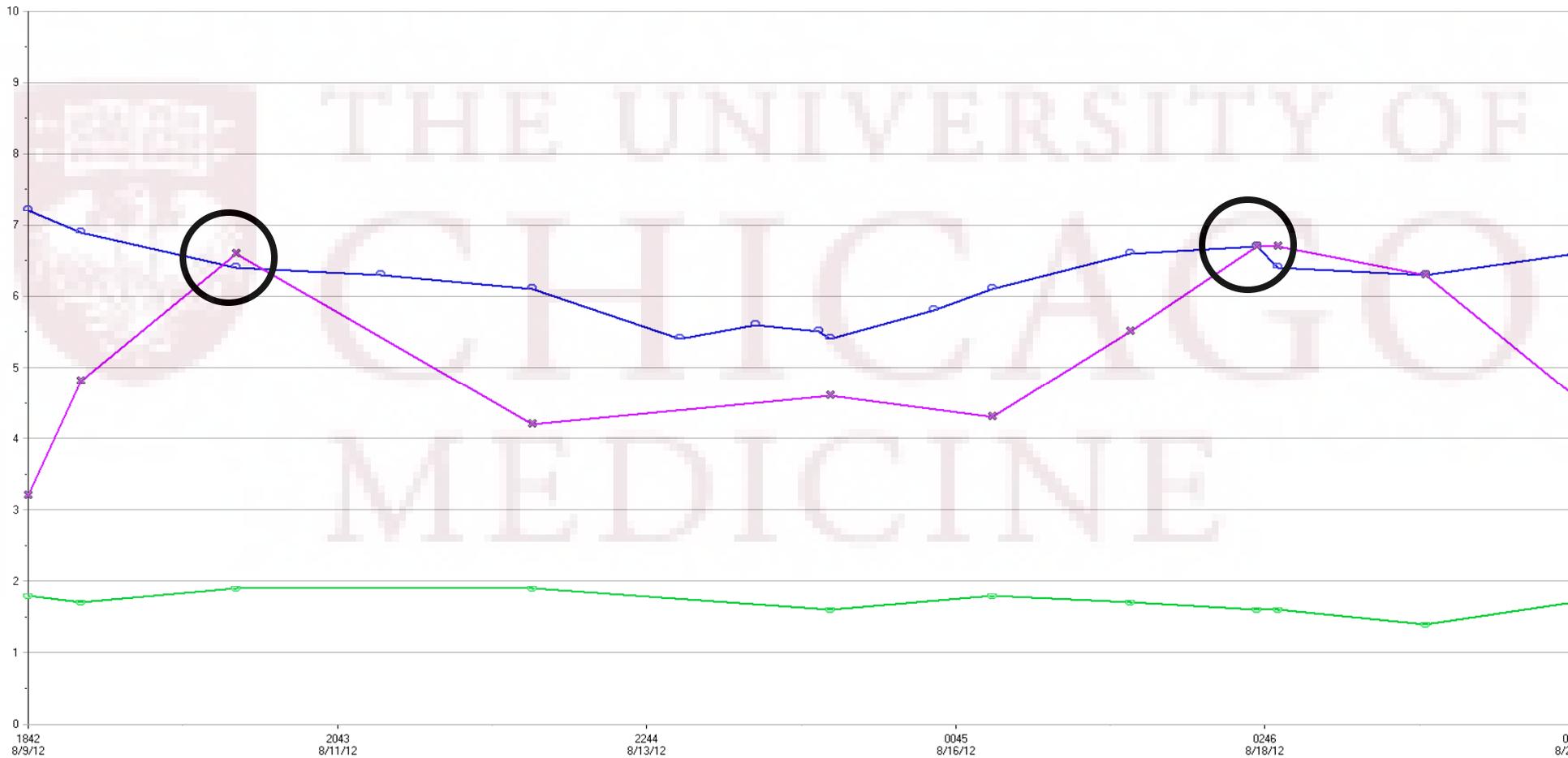
1,25-OH vitamin D 27

1st admission: Calcium trend



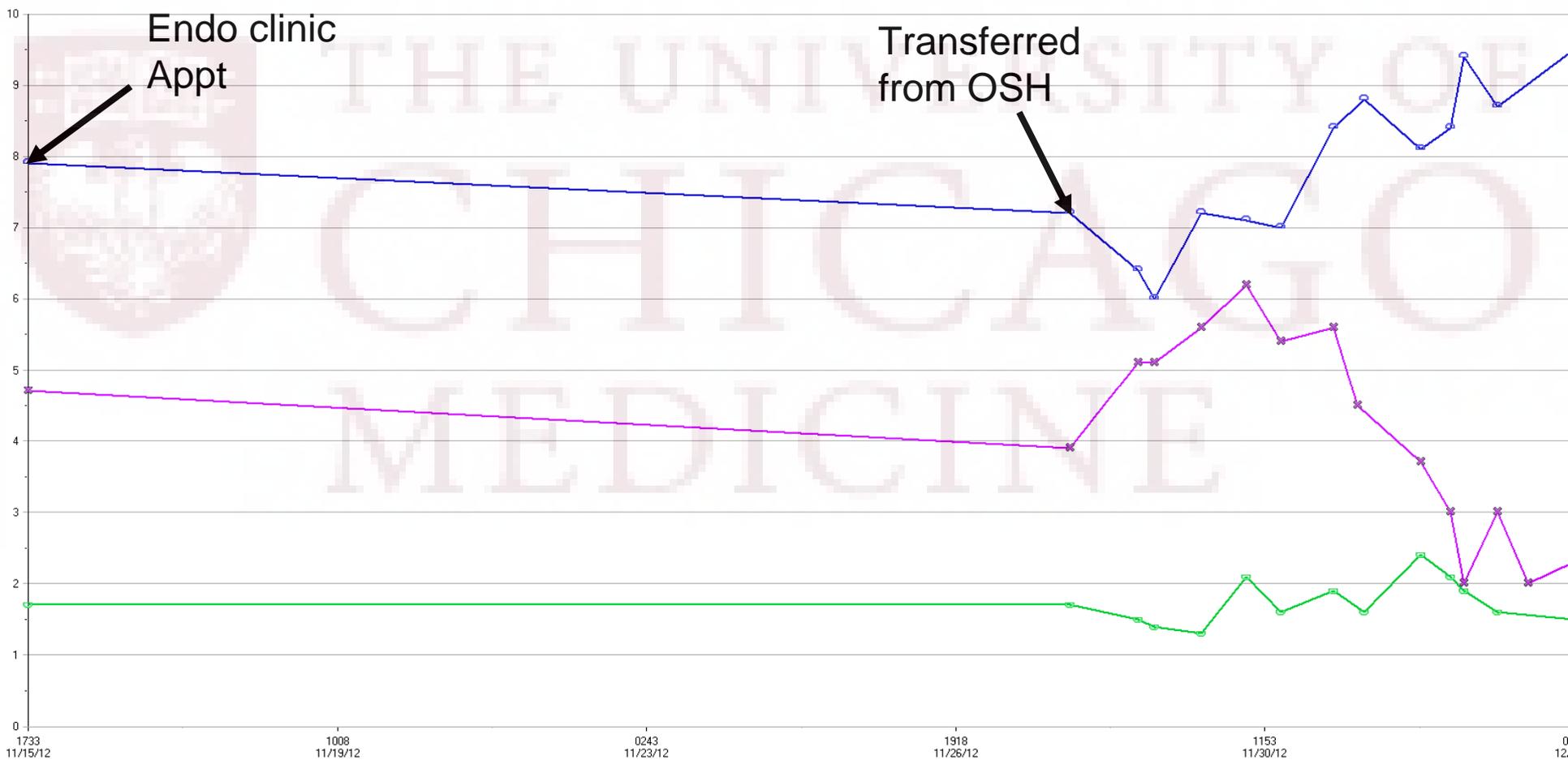
Calcium/Phosphate/Magnesium

Electrolyte trend in MICU



From MICU, was only discharged home on Calcium Acetate, no calcitriol

Calcium/Phosphate/Magnesium



Hypocalcemia

- Gen: altered mental status
 - HEENT: premature cataracts
 - CV: prolonged QT, CHF
 - Pulm: bronchospasm
 - Neuro: basal ganglia calcifications, paresthesias, seizure
 - MSK: muscle twitching, cramps
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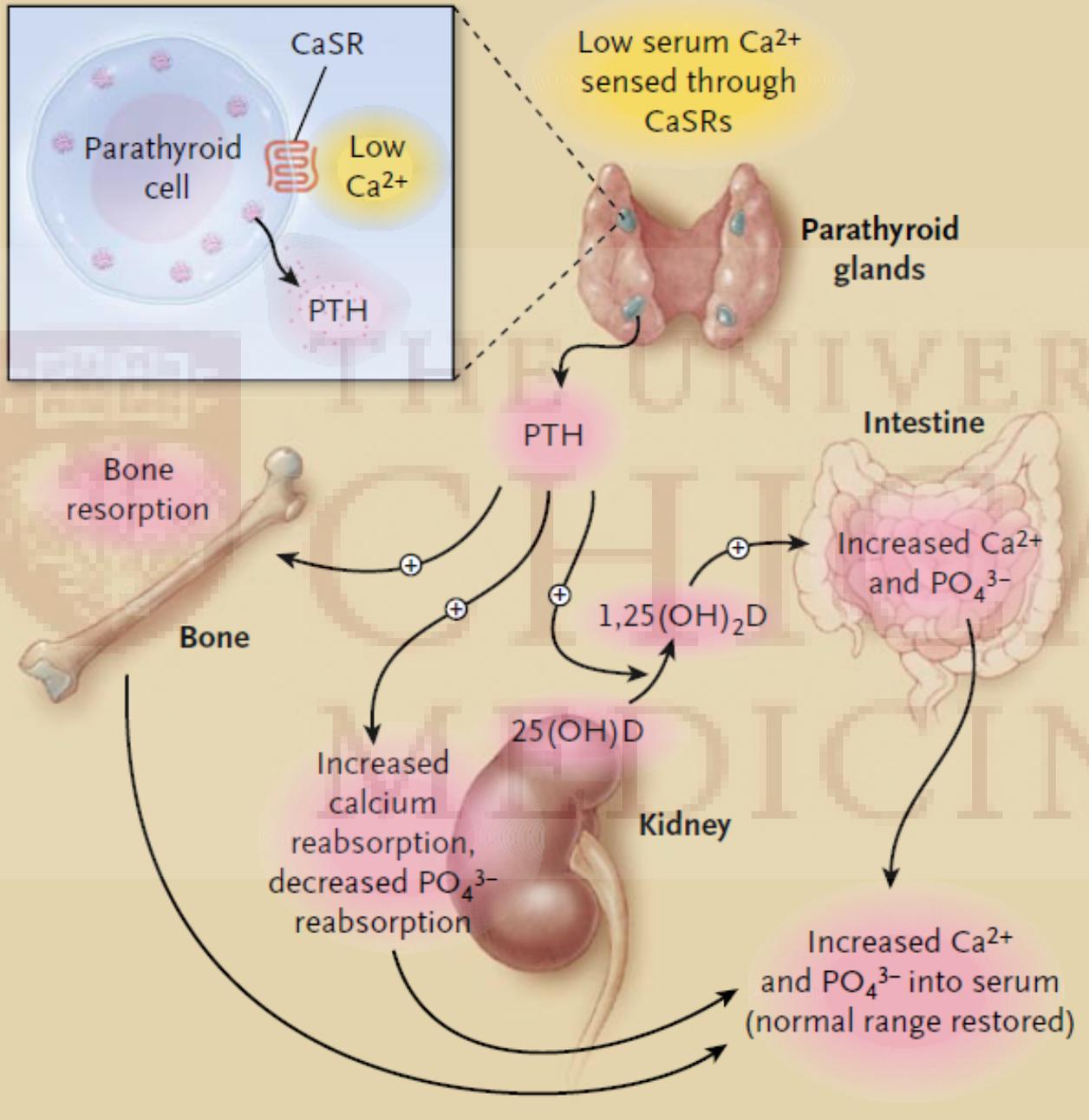
Post-surgical hypoparathyroidism

- Transient hypoparathyroidism occurs in 10% patients who undergo total thyroidectomy.
 - Permanent hypoparathyroidism occurs in <5% of patients who undergo total thyroidectomy.
 - Risk of hypoparathyroidism after initial surgery for primary hyperparathyroidism is <1% but increases to up to 30% after successive surgeries.
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Clinical Questions

- Goals of therapy in hypoparathyroidism?
- Using PTH for refractory cases?
- Effect of PTH on bone in hypoparathyroidism?
- Hypoparathyroidism in patients G-tube?

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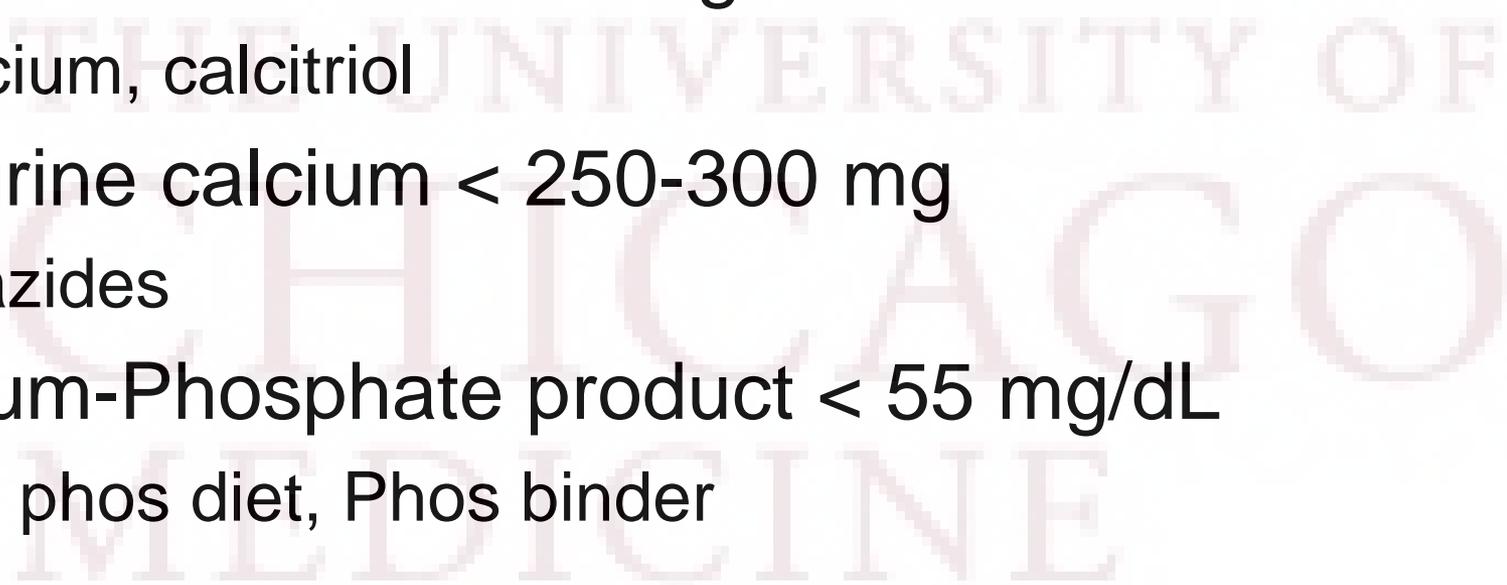
PTH effect on:

	Ca	Phos
Kidney	↑	↓
Bone	↑	↑
Gut	↑	↑

Treatment	Preparations available/dosing	Notes
Parenteral calcium supplements		
10% calcium gluconate	10-mL ampules (94 mg elemental calcium); 1-2 ampules intravenously diluted in 100-200 mL of 5% dextrose or normal saline, infused over 1-2 hours	Duration of effect, 2-3 hours
Calcium gluconate drip	10-mL ampules (94 mg elemental calcium); 10 ampules diluted in 1 L 5% dextrose solution. Initially infused at 50 mL/h (or 1-3 mg/kg per h) and adjusted to maintain a corrected serum calcium ≥ 8.0 mg/dL	...
Oral calcium supplements		
Calcium salts	Starting dose of elemental calcium, 2 g orally 3 times daily	...
Calcium carbonate	Calcium content, 400 mg/g; liquid form, 500 mg elemental calcium/5 mL	Gastrointestinal adverse effects, including constipation, are common
Calcium citrate	Calcium content 211 mg/g	Used in patients with achlorhydria or gastrointestinal intolerance of calcium carbonate
Vitamin D supplements		
Calcitriol	0.25 mcg, 0.5 mcg; liquid, 1 mcg/mL; starting dosage, 0.5 mcg orally 3 times daily	Maximum effect, 10 hours; duration of action, 2-3 days
Ergocalciferol	50 000 IU; liquid, 8000 IU/mL; 25 000-100 000 IU orally weekly	Onset of action, 10-14 days; duration of action, 14-75 days
Cholecalciferol	400, 800, 1000, or 2000 IU; 25 000-100 000 IU orally daily	...

Treatment goals

- Serum calcium 8.0-8.5 mg/dL
 - Calcium, calcitriol
- 24h urine calcium < 250-300 mg
 - Thiazides
- Calcium-Phosphate product < 55 mg/dL
 - Low phos diet, Phos binder



PTH replacement

- PTH 1-34 BID
- PTH 1-84
- Subcutaneous infusion PTH 1-34

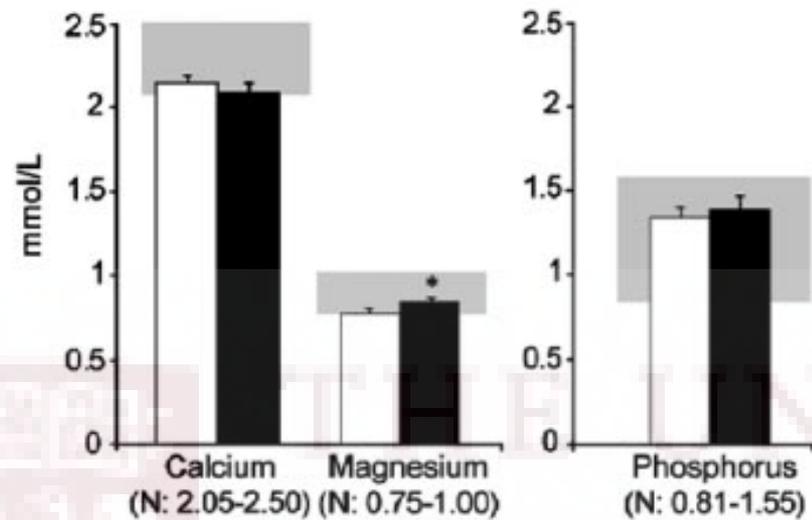
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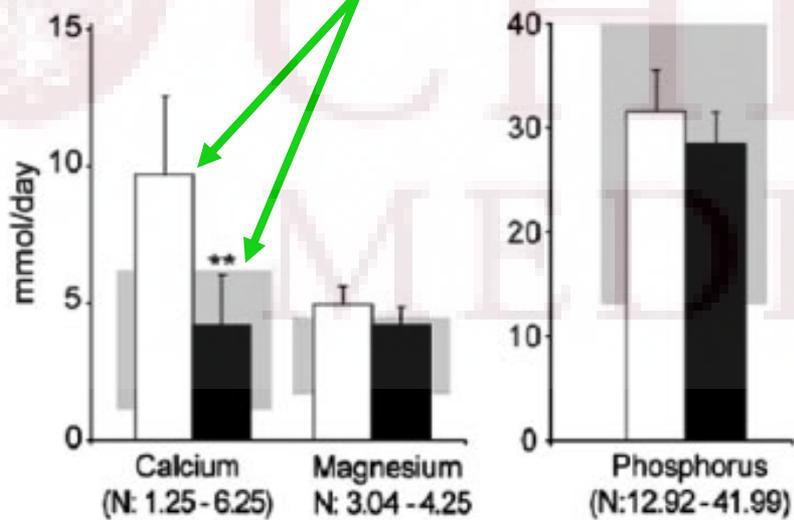
PTH 1-34 BID vs SC infusion

- Winer et al. JCEM 2012.
 - Randomized crossover trial lasting 6 months.
 - N = 8 patients, ~on calcitriol 0.6 mcg/d, calcium supplementation 3100 mg/d, cholecalciferol 1000 IU/d and magnesium 578 mg/d
 - Results: 50% reduction urine calcium, better maintenance of magnesium and normalized bone turnover markers with pump
 - Conclusion: pump delivery of PTH 1,34 provides most physiologic replacement therapy
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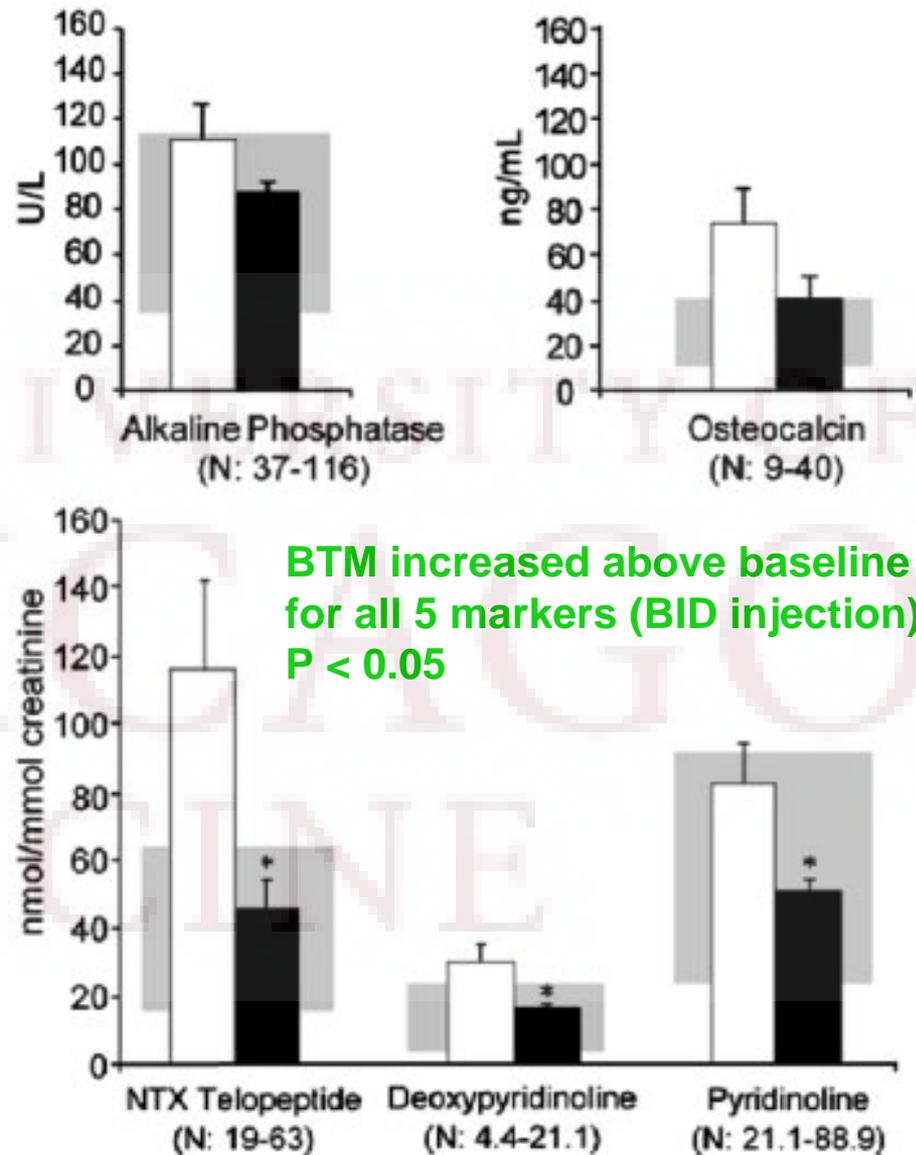
A Serum



B Urine



C Markers of Bone Turnover



BTM increased above baseline for all 5 markers (BID injection) P < 0.05

□ Injection ■ Pump ■ Normal range

* P<0.05 Data are mean +SEM

** P<0.01 Data are mean +SEM

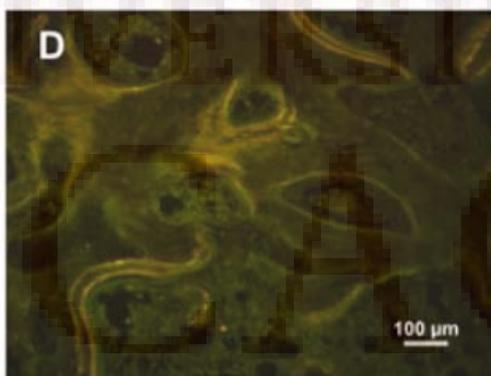
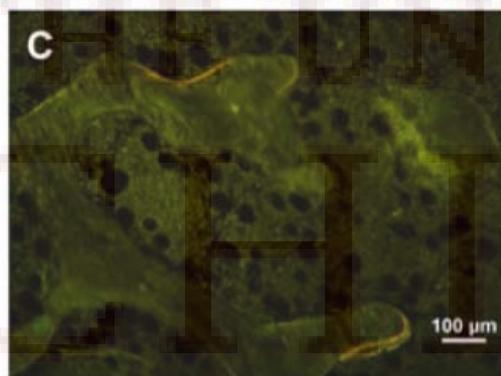
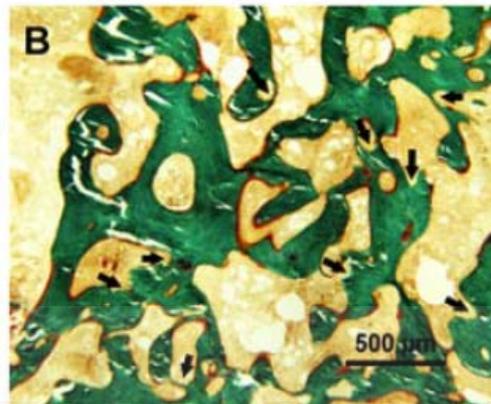
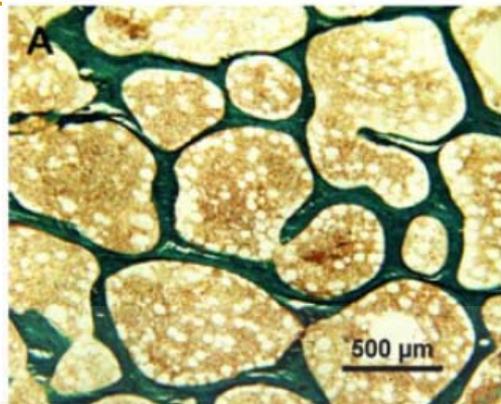
Additional points to consider

- Mean TDD PTH 1-34 was 65% less during pump than BID SC delivery (13 vs 37 mcg/d)
 - 7 of 8 patients preferred pump to SC due to convenience and less calcium-related symptoms
 - Blackbox warning about not using for >2 years due to osteosarcome occurrence in rats
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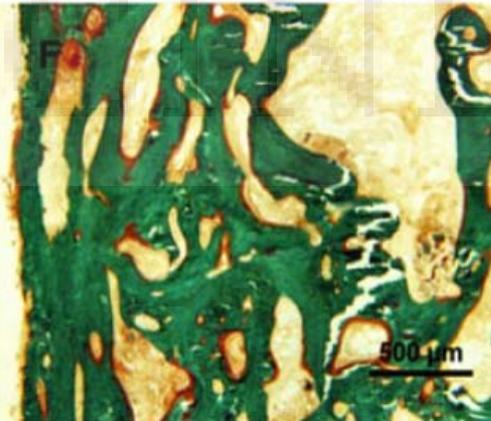
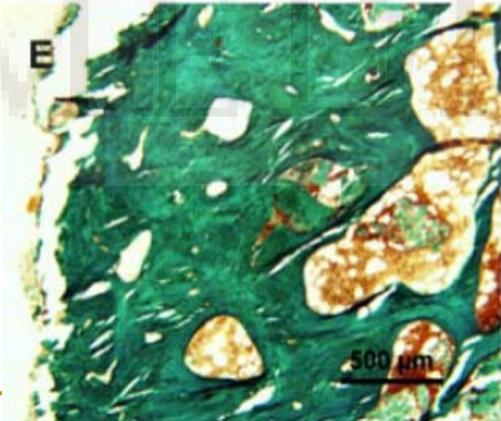
Chronic PTH on bones

- N = 5 hypoparathyroid subjects
 - Human PTH 1-34 BID-TID for 18 mos
 - Bone turnover markers, DXA, iliac crest biopsy
 - Long-term studies on skeletal changes as well as effects of withdrawal of therapy are needed
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Cancellous Bone



Cortical Bone



Baseline

1 year of hPTH 1-34

Tube Feeds & Hypoparathyroidism

- No studies on treating hypoparathyroidism in patients with G-tube feeding



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Take Home Points

- Targets of hypoparathyroidism therapy
- PTH replacement through injection or pump is being studied
- Chronic PTH 1-34 Replacement Might induce Bone Structure

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References

- Winer KK et al. Synthetic Human Parathyroid Hormone 1-34 Replacement Therapy: A Randomized Crossover Trial Comparing Pump versus Injections in the Treatment of Chronic Hypoparathyroidism. JCEM 2012;97(2).
- Shoback D. Hypoparathyroidism. NEJM 2008;359(4):391-403.
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- Gafni RI et al. Daily PTH 1-34 Replacement Therapy for Hypoparathyroidism Induces Marked Changes in Bone Turnover and Structure. JBMR 2012;27(8):1811-1820.
- Cusano NE et al. Mini-review: new therapeutic options in hypoparathyroidism. Endo 2012;41:410-414.