# Cinacalcet Benefits in Chronic Kidney Disease Patients in Babylon Province

Received: 11-September-2022

Revised: 13-November-2022

Accepted: 12-December-2022

# Ali Jasim Mhaimeed Alsultani<sup>1</sup>, Mudhfar Mohammed Meizel<sup>2</sup>, Fadhil Kadhim Jassim<sup>3</sup>

1 College of medicine, University of Babylon, Iraq

2 College Of Medicine, Alqadisiya University, Iraq

3 Manager of Merjan medical City Babylon Health Directorate, Babil, Iraq.

alialsultany81@yahoo.com

#### **Abstract**

**Introduction:** Cinacalcetis calcium sensetive receptor and professionally reductions parathyroid hormone (PTH). The aim of study is to assessed the effect of Cinacalcet on the level of PTH, serum calcium and phosphate in patients they have CKD and on maintenance hemodialysis.

**Method:**Cross sectional experimental study of 60 patients with chronic kidney disease, group A; 30 patients on maintenance hemodialysis not receive cinacalcet 60 mg daily at night for 6 months' group B; 30 patients also on maintenance hemodialysis receive cinacalcet 60 mg daily at night for 6 months'. Patients age gender were taken, also measured the serum calisuim (Ca<sup>++</sup>), phosphate and parathyroid hormone levels.

**Results:** Mean age of patients in group A  $(50.83 \pm 14)$  years, while group B  $(48.9 \pm 15)$  years with no significant difference. There is significant difference in mean of serum Ca<sup>++</sup>level between group A and group B. There is significant difference in mean of serum phosphatelevel between group A and group B. There is significant difference in mean of parathyriodhormone level between group A and group B.

**Conclusion:** The serum Ca<sup>++</sup> level decrease in patients received cinacalcet, serum phosphate and parathyriod hormone levelsdecrease in patients received cinacalcet.

Keywords: cinacalcet, chronic kidney disease, hemodialysis, Babylon province

# 1. INTRODUCTION

Chronic kidney disease (CKD) is the most important illness with life time risk <sup>(1)</sup>, It associated with electrolyte disturbance and cardiovascular disease and death <sup>(2)</sup>, also it is associate with high level of parathyroid hormone and increase risk of hypercalcemiaand cardiovascular events<sup>(3)</sup>. Patients with final stagerenalillness, due to 2<sup>nd</sup>hyperparathyroidism this lead to bone disease and blood vessels calcifications and atherosclerosis<sup>(4, 5)</sup>. Cinacalcet is act as "calcium sensetive receptor" and help to decrease parathyroid hormone (PTH) <sup>(6)</sup>. Cinacalcet can also managing phosphate level to be normal and decrease hypercalcemia<sup>(7, 8, 9)</sup>. Cinacalcet and vitamin D postponements the calcification of "valvular, aortic and coronary"<sup>(10, 11)</sup>. Nemurous studies show lowering in the cardiovascular disease after used Cinacalcettrial <sup>(9)</sup>. The aim of study is to assessed the effect of Cinacalceton the level of PTH, serum calcium and phosphate in patients they have CKD and on maintenance hemodialysis.

#### 2. METHOD

Cross sectional experimental study of 60 patients with chronic kidney disease, group A; 30 patients onmaintenance hemodialysis not receive cinacalcet 60 mg daily at night for 6 months' group B; 30 patients also on maintenance hemodialysis receive cinacalcet 60 mg daily at night for 6 months'. The study done in Marjan medical city/ dialysis department from August 2021 to March 2022. Patients age gender were taken, also measured the serum calisuim (Ca<sup>++</sup>), phosphate and parathyroid hormone levels. The symptoms also diagnosed in 2 groups; (bone pain, Bone fracture Anemia (Hb<10), no fatal cardiovascular events, fatal cardiovascular events, Pruritus, Calciphylaxis, Nausea and vomiting). Statistical analysis done by SPSS 22, frequency and percentage, mean, SD. T test used for assessment differences between mean of continues variables. P-value less or equal to 0.05 is consider significant.

2022 December; 5 (2s): 488-493

# 3. RESULTS

60 patients with chronic kidney disease, group A; 30 patients on maintenance hemodialysis not received cinacalcet 60 mg daily at night for 6 months' group B; 30 patients also on maintenance hemodialysis received cinacalcet 60 mg daily at night for 6 months'. Mean age of patients in group A ( $50.83 \pm 14$ ) years, while group B ( $48.9 \pm 15$ ) years with no significant difference. There is significant difference in mean of serum Ca<sup>++</sup> level between group A and group B; patients received cinacalcethave decrease in the mean of serum phosphatelevel between group A and group B; patients received cinacalcet have decrease in the mean of serum phosphate level less than in patients not received cinacalcet. There is significant difference in mean of serum phosphate level less than in patients not received cinacalcet. There is significant difference in mean of parathyriodhormone level between group A and group B; patients received cinacalcet have decrease in the mean of parathyriodhormone level less than in patients not received cinacalcet. As show in table 1.

**Table 1:** mean difference of Age, serum Ca<sup>++</sup>, phosphate and parathyriod hormone levels according to groups of study.

Variables	hypothyroidism	N	Mean	SD	P-value
Age	Group A	30	50.83	14.03	0.6
	Group B	30	48.90	15.03	
Ca++	Group A	30	1.96	0.15	0.0001
	Group B	30	1.78	0.18	
Phosphate	Group A	30	5.93	1.93	0.0001
	Group B	30	4.05	1.83	
PTH	Group A	30	491.63	308.91	0.0001
	Group B	30	171.34	168.55	

# P-value $\leq 0.05$ (significant).

According to fig 1; gender distribution in 2 groups as the following; group A: (53.3%) are males and (46.7%) are females. In group B; (56.7%) are males and (43.3%) are females.



489

2022 December; 5 (2s): 488-493

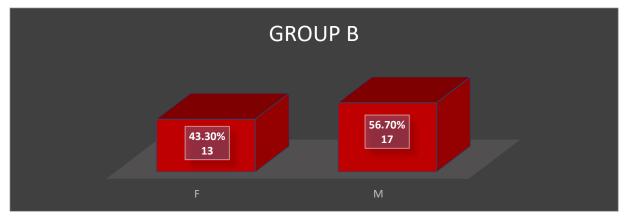


Figure 2: distribution accoprding gender in groups A and B.

Table 2 show; there is no significant difference mean of serum phosphate, serum Ca<sup>++</sup>, PTH level between males and females in patients they received cinacalcet (group B).

Variables	hypothyroidism	N	Mean	SD	P-value
Age	female	13	45.46	16.551	0.3
	male	17	51.53	13.675	
Ca	female	13	1.992	.1891	0.4
	male	17	1.935	.1869	
Phosphate	female	13	3.7408	1.80584	0.4
	male	17	4.2941	1.87919	
PTH	female	13	122.2231	83.50461	0.1
	male	17	208.9112	206.89774	

P-value  $\leq 0.05$  (significant).

According to fig 2; When compate with clinical manifistations of patients not received cinacalcet (group A); 12 patients have bone ache, 15 patients have anemia, 5 patients have non fatal cardiovascular disease, 17 patients have pruritis.

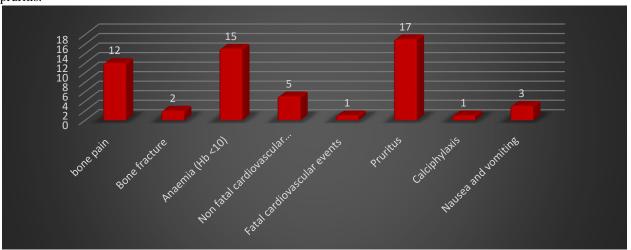


Figure 2: Clinical data for patients not received cinacalcet (group A).

The clinical manifistations of patients received cinacalcet (group B) are; 4 patients have anemia, 2 patients have non fatal cardiovascular disease, 10 patients have nausea and vomiting, 6 patients have pruritis and 5 patients have bone ache. As show in fig 3.

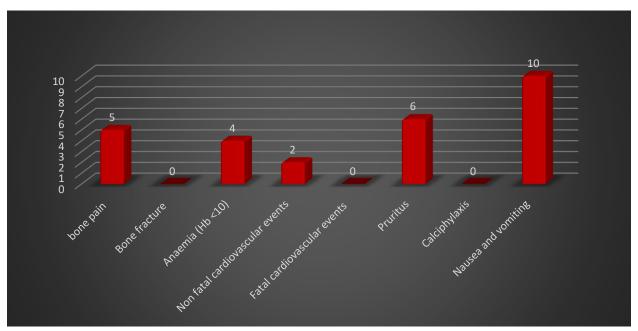


Figure 3: Clinical data for patients received cinacalcet (group B).

# 4. DISCUSSION

Cinacalcetdecrease the cardiovascular disease (11).In current study there is significant difference in mean of serum Ca<sup>++</sup> level between group A and group B; patients received cinacalcet have decrease in the mean of serum Ca<sup>++</sup> level less than in patients not received cinacalcet. This is similar to other study that state the srum calcium values increased beforecinacalcet start, although they persisted within the normal before cinacalcet management ongoing. But after cinacalcet start, calcium fallen shortly to slightly minor values (11). Marcocci et al. also show that decrease in serum calcium to the usual range "[≤10.3 mg/dl (2.6 mmol/liter)]" at the finalestage was completed in 9 patients (53%)<sup>12</sup>. Also Yasuhiro et al. agreed with current results and stated that serum Ca<sup>++</sup> concentration at the standard was  $12 \pm 1$  mg/dL and decreased to  $10.1 \pm 1.6$  mg/dL at the end of the titration phase with cinacalcetdosage of up to 75 mg/3 times in a day<sup>(13)</sup>. In addition there is significant difference in mean of parathyriod hormone level between group A and group B; patients received cinacalcet have decrease in the mean of parathyriod hormone level less than in patients not received cinacalcet. This is agreed with Evans et al. (11) and Marcocci (12) and Peacock et al. (14) they stated that the decrease in PTH concentration. Also there is significant difference in mean of serum phosphate level between group A and group B; patients received cinacalcet have decrease in the mean of serum phosphate level less than in patients not received cinacalcet. This ia agreed with Evans et al. (11) that stated there is dropped of serum phosphate after treatment cinacalcet, other study show serum phosphate alterations in dialysis patients have cinacalcet, (41%) had decreased, (4%) unchanged and (25%) increased, serum P at Month 12 (15). In the current study the clinical manifistations of patients received cinacalcet (group B) are; 4 patients have anemia, 2 patients have non fatal cardiovascular disease, 10 patients have nausea and vomiting, 6 patients have pruritis and 5 patients have bone ache. Nausea, vomiting, and paresthesias were reflected to be knowledgeable management linked side effectes in 9 (53%), 4 (24%), and 3 (18%) patients, correspondingly. Paresthesiashappened when concentrations of serum Ca<sup>++</sup> were within or morethan the normal range (12, 16, 17).

#### , , ,

#### 5. CONCLUSION

The serum Ca<sup>++</sup> level decrease in patients received cinacalcet, in addition to that serum phosphate and parathyriod hormone levels decrease in patients received cinacalcet in patients with chronic kidney disease they are on maintenance hemodialysis.

#### 6. ACKNOWLEDGEMENTS

To Dr. Qais Ajam who contributed towards the study by making substantial contributions to conception, design, acquisition of data, or analysis and interpretation of data, or who was involved in drafting the manuscript or revising it critically for important intellectual content.

No conflict of interest. No funding.

#### REFERENCES

- [1] Grams, M. E., Chow, E. K. H., Segev, D. L. & Coresh, J. Lifetime Incidence of CKD Stages 3–5 in the United States. *Am J Kidney Dis*, 2013 **62**, 245–252. <a href="https://pubmed.ncbi.nlm.nih.gov/23566637/">https://pubmed.ncbi.nlm.nih.gov/23566637/</a>
- [2] van der Velde, M. *et al.* Association of estimated glomerular filtration rate and albuminuria with mortality and end-stage renal disease: a collaborative meta-analysis of kidney disease cohorts. *Kidney Int*, 2011, **79**, 1331–1340.https://pubmed.ncbi.nlm.nih.gov/21289598/
- [3] Block, G. A. *et al.* Mineral Metabolism, Mortality, and Morbidity in Maintenance Hemodialysis. *J Am SocNephrol*, 2004, **15**, 2208–2218.https://pubmed.ncbi.nlm.nih.gov/15284307/
- [4] Martín, M., Valls, J., Betriu, A., Fernández, E. &Valdivielso, J. M. Association of serum phosphorus with subclinical atherosclerosis in chronic kidney disease. *Sex makes a difference*. *Atherosclerosis*, 2015, **241**, 264–270. <a href="https://pubmed.ncbi.nlm.nih.gov/25748053/">https://pubmed.ncbi.nlm.nih.gov/25748053/</a>
- [5] Yamamoto, D. *et al.* Predictors of abdominal aortic calcification progression in patients with chronic kidney disease without hemodialysis. *Atherosclerosis*, 2016, 253, 15–21. <a href="https://pubmed.ncbi.nlm.nih.gov/27573734/">https://pubmed.ncbi.nlm.nih.gov/27573734/</a>
- [6] Block, G. A. *et al.* Cinacalcet for Secondary Hyperparathyroidism in Patients Receiving Hemodialysis. *N Engl J Med*, 2004, **350**, 1516–1525.https://www.nejm.org/doi/full/10.1056/NEJMoa031633
- [7] Ureña, P. *et al.* Cinacalcet and achievement of the NKF/K-DOQI recommended target values for bone and mineral metabolism in real-world clinical practice—the ECHO observational study. *Nephrol Dial Transplant*, 2009, **24**, 2852–2859. <a href="https://pubmed.ncbi.nlm.nih.gov/19369690/">https://pubmed.ncbi.nlm.nih.gov/19369690/</a>
- [8] Li, D. et al. The efficacy of cinacalcet combined with conventional therapy on bone and mineral metabolism in dialysis patients with secondary hyperparathyroidism: A meta-analysis. Endocrine, 2013, 43, 68– 77.https://pubmed.ncbi.nlm.nih.gov/22669774/
- [9] Chertow, G. et al. Effect of Cinacalcet on Cardiovascular Disease in Patients Undergoing Dialysis. N Engl J Med, 2012, 367, 2482–2494.https://www.nejm.org/doi/full/10.1056/nejmoa1205624
- [10] Raggi, P. *et al.* The ADVANCE study: a randomized study to evaluate the effects of cinacalcet plus low-dose vitamin D on vascular calcification in patients on hemodialysis. *Nephrol Dial Transplant*, 2011, 26, 1327–1339. <a href="https://pubmed.ncbi.nlm.nih.gov/21148030/">https://pubmed.ncbi.nlm.nih.gov/21148030/</a>
- [11] Evans, M., Methven, S., Gasparini, A. *et al.* Cinacalcet use and the risk of cardiovascular events, fractures and mortality in chronic kidney disease patients with secondary hyperparathyroidism. *Sci Rep*, 2018, **8**, 2103.https://www.nature.com/articles/s41598-018-20552-5
- [12] Marcocci C, Chanson P, Shoback D, et al. Cinacalcet reduces serum calcium concentrations in patients with intractable primary hyperparathyroidism. *J ClinEndocrinolMetab*. 2009;94(8):2766-2772. <a href="https://pubmed.ncbi.nlm.nih.gov/19470620/">https://pubmed.ncbi.nlm.nih.gov/19470620/</a>

2022 December; 5 (2s): 488-493

- [13] Takeuchi Y, Takahashi S, Miura D, et al. Cinacalcet hydrochloride relieves hypercalcemia in Japanese patients with parathyroid cancer and intractable primary hyperparathyroidism. *J Bone Miner Metab*. 2017;35(6):616-622.https://pubmed.ncbi.nlm.nih.gov/27873072/
- [14] Peacock M, Bilezikian JP, Klassen PS, Guo MD, Turner SA, Shoback D. Cinacalcet hydrochloride maintains long-term normocalcemia in patients with primary hyperparathyroidism. J ClinEndocrinolMetab, 2005, 90:135–141.https://pubmed.ncbi.nlm.nih.gov/15522938/
- [15]Zitt E, Fouque D, Jacobson SH, et al. Serum phosphorus reduction in dialysis patients treated with cinacalcet for secondary hyperparathyroidism results mainly from parathyroid hormone reduction. *Clin Kidney J*. 2013;6(3):287-294.https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3665312/
- [16] Cetani F, Saponaro F, Banti C, et al. Cinacalcet efficacy in patients with moderately severe primary hyperparathyroidism according to the European Medicine Agency prescription labeling. *J Endocrinol Invest*. 2012;35(7):655-660.https://pubmed.ncbi.nlm.nih.gov/21971564/
- [17] Marcocci C, Cetani F. Update on the use of cinacalcet in the management of primary hyperparathyroidism. *J Endocrinol Invest*. 2012;35(1):90-95. <a href="https://pubmed.ncbi.nlm.nih.gov/22104762/">https://pubmed.ncbi.nlm.nih.gov/22104762/</a>