

Welcome to the August Q&A!

Trusts represented today include: Cambridge, East and North Herts, Guy's and St Thomas', North Bristol, Cork, Glasgow, Bristol, Dundee, Devon and Exeter, Limerick, Lancashire, Northampton, Merseyside and the Wirral

Q+A monthly meeting new format



- First presentations in June/July met with success.... 😁
- 10-15 minute bite site presentation/update from an RPG member
 - Themes that regularly come up in the Whatsapp group (anticoagulation on HD)
 - Case presentations opportunity for portfolio entry
 - Other updates from UKKA
- June Q+A
 - Robert Brown to present the UKKA presentation on Roxadustat
- July Q+A
 - Natasha Moore presenting the ERA/UKKW overview
- Aug Q+A
 - Louise Condon/Robert Brown presenting the two sides of the denosumab story
- Sept Q+A
 - ?BK talk from Leeds consultant

Any feedback/requests gratefully received © New members to clinical sub-group welcome ©

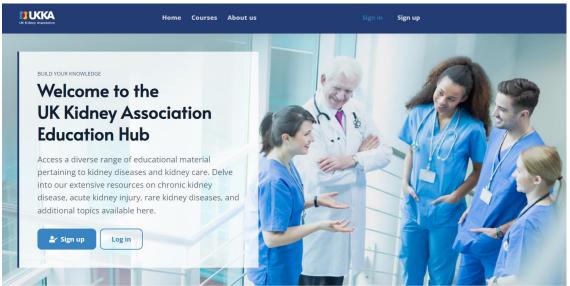
Updates



- RPG conference 2024 11th/12th October Registration currently open ©
 - Birmingham Arden Hotel, £250 for full programme
 - Programme circulated via whatsapp
 - Abstract submissions open until 1st September
- UKKA guideline living with frailty and CKD
 - See link in the chat shared by Clare great for portfolios ©
- Monthly teams chat for advanced/consultant portfolios
 - Email cathy.pogson@porthosp.nhs.uk if you would like to join
 - Members have been linked up into smaller groups to work together and then link back into the bigger group in time
 - Monthly meetings to suspend until over summer
- Reminder to check your RPG membership is up to date
 - Large number of lapsed memberships, currently being checked against the Q&A whatsapp group list
 - If you are not receiving the emails from UKKA, email the secretarial team (Caitlin Sewell)

Updates

- KRUK have opened applications for research grants
 - Pharmacy-led research and research into nephrotoxicity grants
 - Speak to Kathrine Parker, Dane Howard or Cathy Pogson for more information
- UKKA guidance on measles
- UKKA Education and Training Hub is now live
 - UKKA Learning Hub (ukkidney.org)
 - Resources on CKD, Glomerulonephritis and Transplantation
 - Other resources and links to follow



KQIP Knowledge Hubs

Our educational resources

eLearning Courses









Denosumab in CKD 4/5

Doable or disaster?

Louise Condon, Guys and St Thomas' NHS Foundation Trust Robert Brown, North Bristol NHS Trust

Overview



- Challenging area of care
- Most anti-fracture treatments contra-indicated in GFR<30ml/min
- FREEDOM trial (Denosumab) shows a significant reduction in vertebral, nonvertebral and hip fractures in post-menopausal osteoporosis
- No clear recommendations / guidance for the use of Denosumab in fracture prevention in CKD stage 4 + 5
- In CKD stage 4 + 5 denosumab can cause pronounced hypocalcaemia and increased
 PTH within 15 days
- Close monitoring required +/- calcium supplementation +/- vitamin D receptor agonists

Denosumab and hypocalcaemia - Case 1



• 83 yr female

admitted to hospital with severe hypocalcaemia. Preceded by one day history of some confusion with a UTI. Lab phoned adj Ca²⁺ 1.7

- Retired, lives with husband
- Hx:
 - ESKD secondary to Wegner's granulomatosis
 - DBD kidney transplant 1993
 - Anti-cardiolipin antibody positive on anticoag. Hx VTE
 - Osteoporosis on denosumab since approx. 2015
 - Parastomal hernia
 - Hartmann's 2017

Denosumab and hypocalcaemia - Case 1



- Osteoporosis:
 - DEXA 2007 osteoporosis
 - DEXA 2009 osteopenia hip, osteoporosis R and L femoral neck
 - DEXA 2013 osteoporosis # T8, T11, T12
 - Intolerant to bisphosphonates
 - started denosumab approx. 2015

Denosumab and hypocalcaemia - Case 1

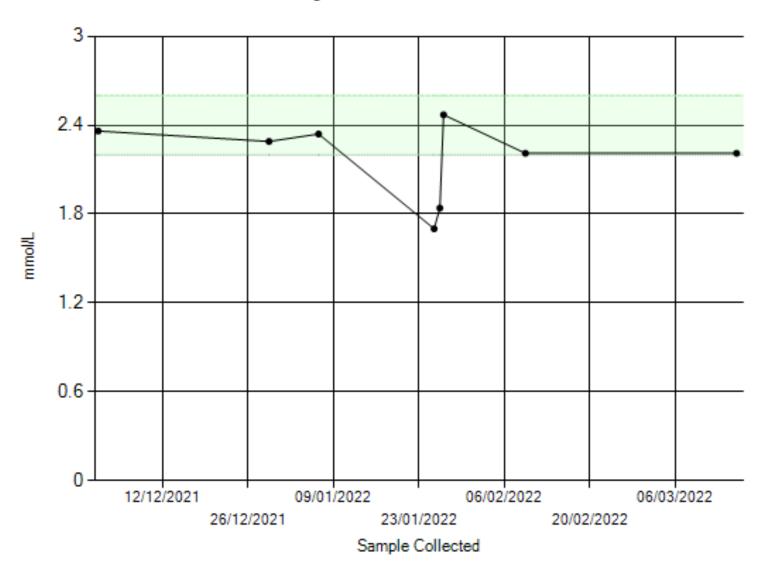


DHx

18.03.22 DARBEPOETIN ALFA 30micg Monthly SC 16.03.22 CO-CARELDOPA 125mg 4 x daily Oral 02.03.22 ENSURE PLUS ACBS 220ml 1 x daily Oral 27.01.22 ALLOPURINOL 200mg 1 x daily 27.01.22 CALCEOS 1.00 Tab 2 x daily Oral 17.01.22 DENOSUMAB 60mg Every 6 month SC 14.07.21 STEROID WARNING CARD 24.10.18 VANQUORAL 25mg 2 x daily Oral 21.03.17 GABAPENTIN 100mg 3 x daily Oral 21.03.17 BUPRENORPHINE 20micg 1 x week Topic 21.09.16 APIXABAN 2.5mg 2 x daily Oral 13.05.15 MYCOPHENOLATE MOFETIL 500mg 2 x daily Oral 24.05.94 PREDNISOLONE 5mg 1 x daily

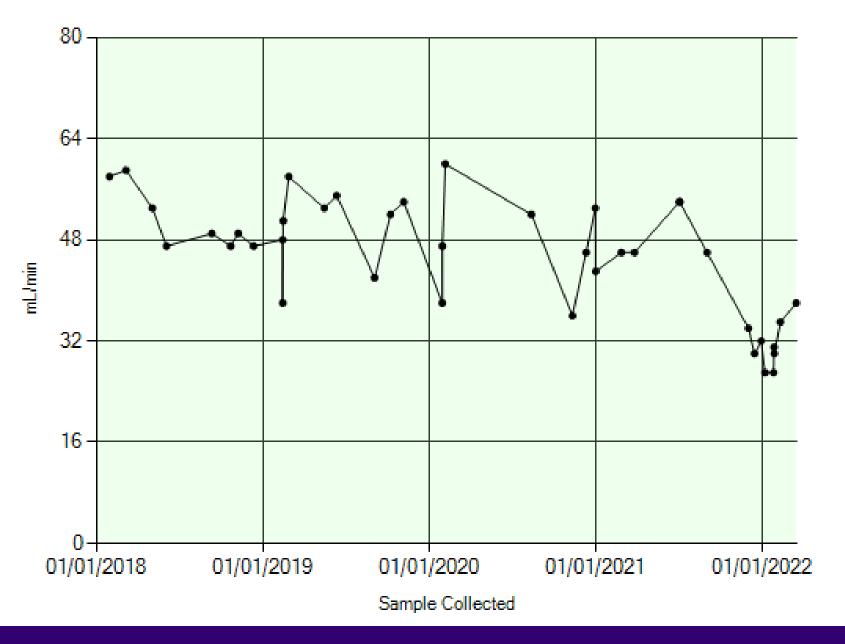
Adjusted Calcium





eGFR/1.73m2 (CKD-EPI)





Denosumab and hypocalcaemia – Case 2



74 yr female

admitted to hospital with severe hypocalcaemia. HD bloods lab phoned adj Ca²⁺ 1.7

1 of many admission from April to July 2021 with hypocalcaemia

- Hx:
 - ESKD secondary to diabetic nephropathy
 - HD commenced Dec 2018
 - Type 1 diabetes with recurrent diabetic ketoacidosis
 - AKI 2001 recurring acute HD secondary to anti-TB med (Px TB contact)
 - Hypothyroidism
 - Coeliac disease 2016
 - C. diff. Oct 2018
 - Osteoporosis on DEXA scan, following a fractured pubic rami Sep 2019

Denosumab and hypocalcaemia – Case 2



- Osteoporosis:
 - DEXA Jan 2020 osteoporosis
 - fractured pubic rami in Sep 2019
 - Denosumab started March 2021

Denosumab and hypocalcaemia – Case 2

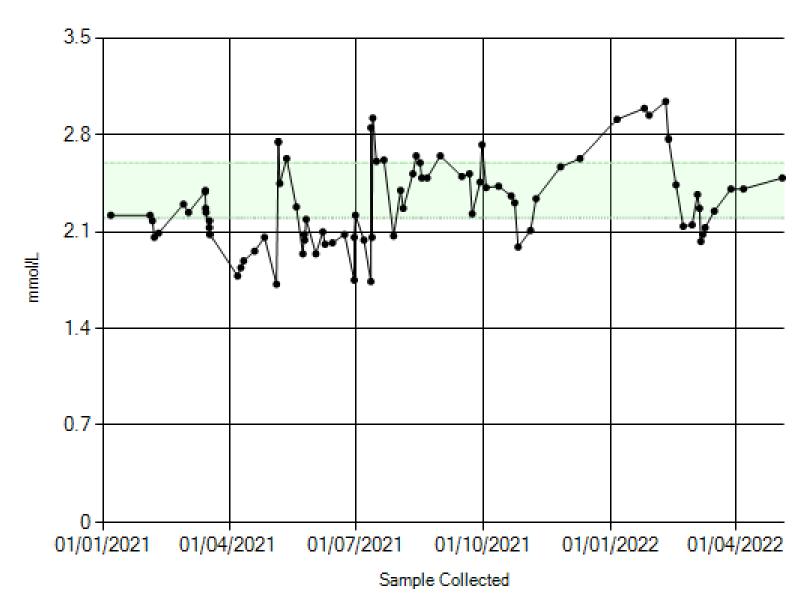


DHx

17.03.22 DARBEPOETIN ALFA 30micg 1 x week IV **ALFACALCIDOL 500nang 1 x daily Oral** 11.02.22 DOXAZOSIN 4mg 1 x daily Oral **12.07.21 ↑**A ALFACALCIDOL **1.25** micg **1** x daily Oral STOP **21.01.22** 01.04.21 Denosumab 60 mg s/c 6 monthly STOP 12.05.22 16.03.20 FOLIC ACID 5mg 1 x daily Oral 06.02.20 FELODIPINE 10mg 1 x daily Oral 10.07.19 EZETIMIBE 10mg 1 x daily Oral 08.04.19 GLANDOSANE LEMON 1 Dose PRN other Oral 14.03.19 ABASAGLAR 100iu/ml 3ml 2 x daily SC 07.12.18 VENOFER (PER-PROTOCOL) PER PROTOCOL IV 21.06.18 LEVOTHYROXINE SODIUM 225micg 1 x daily Oral 10.05.18 NOVORAPID 100iu/ml VAR unit 3 x daily SC

Adjusted Calcium





Denosumab in CKD 4



Pre-initiation: CrCl 15-30 ml/min

- Assess for contra-indications: recurrent UTIs, LRTI or soft tissue infection, on concurrent immunosuppressants; consider risedronate (lower dose 35 mg every 2 weeks)
- Measure serum corrected calcium (optimize intake), phosphate, PTH (<100ng/L), 25(OH)vit D
- If 25(OH) < 50 nmol/L; replete with colecalciferol D3) to concentrations > 50 nmol/L
- Aim for PTH <100ng/L (consider addition of active vitamin D (alfacalcidol), optimization of calcium and phosphate)
- If PTH < 40 ng/L: consider reducing or stopping alfacalcidol

Post dose monitoring

- Repeat serum corrected calcium within 2 weeks, if low, increase calcium supplementation or if taking active vitamin D (one-alfa calcidol), increase dose
- Further follow up in 6 weeks for repeat calcium/PTH
- ?Role of MDT / GP follow up?

Denosumab in CKD 5



Pre-initiation: CrCl <15ml/min or dialysis dependent

- Assess risk: recurrent UTIs, LRTI or soft tissue infection, on concurrent immunosuppressants
- Measure serum corrected calcium (optimize intake), phosphate, PTH, 25(OH)vit D
- If 25(OH) < 50 nmol/L; replete with colecalciferol D3) to concentrations > 50 nmol/L
- Aim for PTH <300ng/L (consider addition of active vitamin D (alfacalcidol), cinacalcet, optimization
 of calcium and phosphate)
- If PTH < 100 ng/L: consider reducing or stopping alfacalcidol

Post dose monitoring

- **CKD 5:** repeat serum corrected calcium within 2 weeks, if low, increase calcium supplementation or if taking active vitamin D (one-alfa calcidol), increase dose. Further follow up in 6 weeks for repeat calcium/PTH.
- **CKD 5D:** repeat corrected calcium **weekly**. If low, increase calcium supplementation (oral +/-dialysate) and/or active vitamin D (alfacalcidol)
- ?Role of MDT / Dialysis unit

Communication is key



- Dear Dr X,
- I hope you are well.
- I'm one of the rheumatology pharmacists at Guy's and I'm emailing to just flag that your patient X had her denosumab dose on 16th August. We conducted the first post-dose calcium check yesterday on dialysis and her corrected calcium has dropped to 2.06mmol/l just one week after the dose.
- Ordinarily, the calcium tends to dip up to 2 weeks post dose then recover but I wanted to flag to you in case you recommend any change to her supplementation or dialysate?
- We will continue to monitor weekly for the next 3 weeks and update you if anything else happens.
- Thank you for your help in advance,
- Kind Regards
- Separate comms to dialysis unit nurses, bloods ordered in advance

References



Cummings SR, Martin JS, Mc Clung MR et al Denosumab for prevention of fractures in post menopausal women with osteoporosis. N Engl J Med 2009; 361: 756-765

Jamal SA, Ljunggren O, Stehman-Breen C et al Effects of Denosumab on fracture and bone mineral density by level of kidney function. J Bone Miner Res 2011; 26: 1829-1835.

Sprague SM, Bellorin-Font E, Jorgetti V, Carvalho AB, Malluche HH, Ferreira A, et al. .: Diagnostic accuracy of bone turnover markers and bone histology in patients with CKD treated by dialysis. Am J Kidney Dis 2016; 67: 559–566.

Miller PD, Roux C, Boonen S, et al. Safety and efficacy of risedronate in patients with agerelated reduced renal function as estimated by the Cockcroft and Gault method: a pooled analysis of nine clinical trials. J Bone Miner Res. 2005;20:2105-2115.

Q&A - UKRPG Conference 11th/12th October 2024



- 40th Anniversary of RPG Conference!
- Conference registration currently open
- £250 for both days (including dinner/celebrations and accommodation on Friday evening)
- Abstract submission opens until 1st September
- All posters/abstracts encouraged including if presented at UKKW
- Looking for more volunteers to join the conference committee
- Looking for abstract markers and facilitators for the conference
- Contact <u>linda.ross@gstt.nhs.uk</u> / poojamehta.gudka@nhs.net to get involved
- A number of bursaries may be available further details to follow, however please do start exploring local funding options ASAP

Supply Problems



- Sando K/slow Na/sando phos
 - Supplies appear resolved
- Taurolock U/Alteplase
 - Taurolock U back?
 - Alteplase 2mg still OOS options include ULM alteplase (cost pressure) or urokinase
- Mycophenolic acid 360mg
 - Generic issues ongoing, Sandoz are limiting supplies to instalments for homecare companies
- Ibandronate/Pamidronate
 - Unlicensed pamidronate being explored
 - Some centres looking into using zolendronic acid/denosumab depending in indication

Supply Problems

- Ganciclovir
 - Vials in stock, some supplies of pre made bags



- 5% HAS
 - Restrict supply, IVIG panels alerted to likely increase use due to the impact on plasma exchange
- Daliport 0.5 mg strength
 - Round up/down where target range allows. Switching to Advagraf/Envarsus an option
- Budesonide (kinpeygo)– no supply issue but issues getting accounts set up, now available through Phoenix.
- Basiliximab transplant centres on monthly allocation, unlicensed supplies available but very £££

Q&A Themes - Guidelines



- Treatment of renal artery thrombosis catheter-directed thrombosis
- UKKA version of the anaemia guideline
- IV heparin for ACS in HD patients

Q&A Themes - Renal

UKRPG

- Dartimumab for cresc IgA
- Tolvaptan on homecare
- Baricitinib for monogenic interferonopathies
- Oxycodone/fentanyl in ESRD
- Apixaban for AF in CKD5 pre-dialysis/monitoring
- Sparsentan through the managed access scheme
- Rituximab and high dose methylprednisolone for vasculitis
- Sodium thiosulfate IV to non-dialysis patient
- Molnupiravir in CrCl < 30 ml/min
- Switching between DOACs, dosing interval extended as per NICE
- Inadine dressings CI in renal impairment
- GLP 1RA in CKD patients
- Cyclophosphamide cumulative dose
- Tolvaptan frequency of LFT monitoring

Q&A Themes - Dialysis



- Ciprofloxacin dosing for peritonitis
- Xylocaine 10 % spray in needle phobic dialysis patients
- Ivermectin is it dialysed?
- IV heparin to treat ACS in HD patients
- Kitelock 4 % as line lock
- Lokelma for high potassium in dialysis patients
- Cyclophosphamide dosing on dialysis/filtration
- Etelcalcetide supply for home haemo patients
- Fondaparinux in dialysis patients
- DDAVP prior to tunnelled line insertion
- Vitamin a and e supplementation in dialysis patients with deficiency
- Rivaroxaban 2.5 mg BD and aspirin in dialysis patients on as per COMPASS trial, now on HD

Cefazolin dosing for endocarditis in HD (?4/4/6g)

Q&A Themes - Transplant

UKRPG

- ATG peripherally
- High dose MMF for acute rejection
- Polarspeed for transplant immunosupporession homecare supplier
- Switching from tacrolimus to sirolimus
- Roxadustat in transplant patients
- Campath for T cell mediated rejection in renal transplant
- IVIG in BK

June Q&A - Close



Thank you for attending!

Next Q&A: Wednesday 18th September

Keep in touch throughout the month on WhatsApp Q&A Group

- >100 participants throughout UK and Ireland
- Useful real time forum for clinical Q's
- Please consider joining!



RPG Oct Q&A - Topic: Open Session 28