



# Management of feline chronic kidney disease

An RCVS Knowledge webinar produced for BSAVA Congress

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### Conflict of Interest Disclosure

I have no relevant financial interest, arrangement or affiliation with any company or organisation.













### Quality Improvement

What is it?













### Quality Improvement

It's about working together, gathering real information ...... and finding practical things that we can do to continuously improve as a whole.













### Learning Objectives

- To understand the advantages of creating clinical guidelines to improve the standards to which we work.
- To be able to design a clinical guideline and critically assess its value within your practice.











# Management of feline chronic kidney disease

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### Conflict of interest

I founded and am employed by Vet Professionals Ltd.

I provide consultancy services, lecturing, research etc. for a number of pharmaceutical companies and pet food companies including:

Boehringer Ingelheim; Ceva Animal Health; Dechra; Elanco; Eukanuba/Iams; Hill's Pet Nutrition; MSD Animal Health; Norbrook Laboratories; Royal Canin

I provide CPD for charities (Feline Friends, International Cat Care) and CPD organisations (e.g. BSAVA, BVNA, Central CPD, Webinar Vet, LVS, and others)













## Management of CKD



# What can we do to slow progression of CKD?

- Two major maladaptive processes on which we can have an impact:
  - Renal secondary hyperparathyroidism
    - Arises due to phosphate retention associated with CKD
  - Glomerular hyper filtration
    - RAAS activation results in glomerular hyper filtration and proteinuria
- And, we can address other factors thought to have an influence on progression
  - Systemic hypertension, dehydration etc.













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### Renal secondary hyperparathyroidism

- Renal excretion of phosphate depends on glomerular filtration
- $CKD \rightarrow \rightarrow \rightarrow$  phosphate retention
- Retained phosphate complexes with ionized calcium reducing the ionized calcium fraction
- PTH production and release is stimulated
- Early CKD: compensatory mechanisms help to maintain normophosphataemia
  - — ↑Parathyroid hormone (PTH) → phosphatiuria
- PTH levels are increased before changes in plasma Ca and P levels are detected
- PTH levels may be increased before azotaemia develops in some cats











### Renal secondary hyperparathyroidism

- As disease progresses, hyperphosphataemia develops
- $\rightarrow$   $\rightarrow$  development of renal secondary hyperparathyroidism (R2HPTH)
  - $-\downarrow$  calcitriol production by the kidneys is the other major trigger
- R2HPTH triggers Ca and P release from bone (renal osteodystrophy), soft tissue mineralisation, progression of renal disease
- PTH is considered to be a 'uraemic' toxin
- R2HPTH is detrimental to survival.















### How can we prevent/reverse this?

- Phosphate restriction is currently considered the key to successful management
- Aim for blood phosphate levels in the lower half of the reference range to remove the stimulus for PTH production













Stage	Description	Creatinine results	Target phosphate levels
1	Non-Azotaemic	< 140 μmol/l < 1.6 mg/dl SDMA > 14 μg/dL	0.8–1.5 mmol/l 2.5–4.5 mg/dl
2	Mild renal azotaemia	140-249 μmol/ 1.6–2.8 mg/dl	< 1.5 mmol/l < 4.5 mg/dl
3	Moderate renal azotaemia	250-439 μmol/l <i>2.9–5.0 mg/dl</i> SDMA ≥25 μg/dl	< 1.6 mmol/l < 5.0 mg/dl
4	Severe renal azotaemia	> 440 μmol/l > 5.0 mg/dl SDMA ≥45 μg/dl	< 1.9 mmol/l < 6.0 mg/dl

### NB many labs have inappropriate phosphate ranges for CKD patients

Sub-stage according to BP and proteinuria

www.iris-kidney.com















### Aims of phosphate restriction

- Your patient will feel better
- Your patient will live longer
- Their renal disease is less likely to progress
- **Current IRIS recommendations** 
  - Phosphate restriction recommended for all azotaemic CKD patients (i.e. Stages 2, 3 and 4)
  - Phosphate restriction recommended irrespective of patient's blood phosphate levels











### Where is the evidence for this?

- Many publications looking at survival statistics for cats with CKD\*
- Phosphate targets quoted are based on expert consensus (www.iris-kidney.com)

ISFM Consensus Guidelines on the Diagnosis and Management of Feline Chronic Kidney Disease

Practical relevance: Chronic kidney disease (CKD) is one of the most commonly diagnosed diseases in older cats. In most cats, CKD is also a progressive disease and can be accompanied by a wide range of clinical and clinicopathological changes. These ISFM Consensus Guidelines have been developed by an independent panel of clinicians and academics to provide practical

advice on the diagnosis and management of this complex disease. Clinical challenges: Although CKD is a common clinical problem in cats, the manifestations of disease vary between individuals. Thus there is a need for careful and repeat evaluation of cats with CKD and adjustment of therapy according to individual needs. In addition to addressing problems arising from CKD and improving quality of life (QoL) for the patient, therapy may also target slowing the underlying progression of disease and hence prolonging life. While maintaining QoL is of paramount importance in our patients, this can be challenging when multiple therapies are indicated. In some cases it is necessary to prioritise therapy, given an understanding of what is likely to most benefit the individual patient. Evidence base: In preparing these Guidelines, the Panel has carefully reviewed the existing published literature, and has also graded the quality of evidence for different interventions to help to provide practical recommendations on the therapeutic options for feline CKD. This is a field of veterinary medicine that has

benefited from some excellent published clinical research and further research findings will undoubtedly

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\* Barber et al 1999; Elliott et al 2000; Ross et al 2006











modify the recommendations contained in these Guidelines in the future.



### How do we achieve phosphate restriction?

- (Maintain normal hydration)
- Phosphate restricted diet
  - Therapeutic renal diet
  - Home prepared diet
  - Senior diets
  - Avoid high phosphate treats (dairy, meat)
- Oral phosphate binders
  - In combination with renal diet or normal food













### Therapeutic renal diets

- THE single most effective treatment for cats with CKD: proven to prolong life by 2-3 times AND improve quality of life
  - Recommended for all cats in IRIS Stages 2, 3 and 4 CKD
  - Recommended for IRIS Stage 1 cats if proteinuric or hyperphosphataemic
- Many key characteristics including
  - Palatable, high calorie
  - High quality protein, restricted levels
  - Phosphate restricted
  - Non acidifying
  - Potassium & vitamins added, low sodium











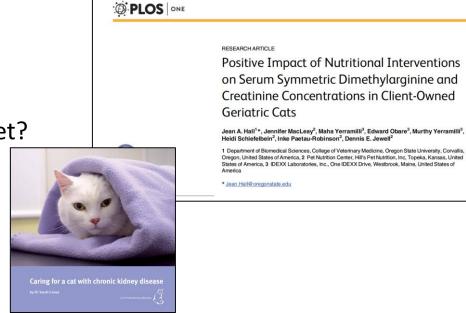






# What changes (if any) are indicated for CKD IRIS Stage 1 cats?

- Owner education
- **Monitoring**
- Diet change?
  - Standard therapeutic renal diet?
  - Early renal diet?
  - Senior diet?













### Therapeutic renal diets

#### Quality of evidence as an intervention

Increased longevity: GOOD

Improved QoL: GOOD

ISFM Consensus Guidelines on the Diagnosis and Management of Feline Chronic Kidney Disease

en individuals. Thus there is a need for careful and repeat evaluation of cats with CKD and

#### Panel recommendations

 The Panel strongly recommends the feeding of a commercial renal diet in all cats with azotaemic (stages 2-4) CKD. Where possible this diet should be fed exclusively...Feeding a wet rather than dry diet...also recommended.....

ISFM Consensus Guidelines on the Diagnosis and Management of Feline CKD, 2016















## What level of dietary compliance is realistic?

- Research studies relating to feline CKD (1999 2006)
  - -46 94%
- Owners of cats with CKD (2015, 2016)
  - **-** 51% 66%
- Pet owner surveys: compliance to therapeutic diets in general (2006, 2009)
  - -12-21%













### Successful transition to a renal diet

- Make it a long-term aim gradual transition (4-8 weeks+)
- Introduce early (IRIS Stage 2)
- Persevere, don't be daunted
- Emphasise benefits in longevity and QoL
- Give medications separately e.g. in palatable treat
- Don't introduce when the cat is in hospital or is clinically unwell
- Home prepared diets: possible with input from a vet nutritionist
- Wet preferred to dry
- Renal > home prepared > senior > standard commercial cat food
- Consider feeding tube support







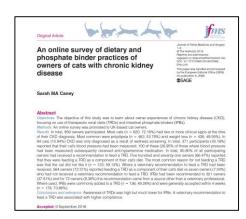






### Know the power of your recommendation!

'Where a veterinary recommendation to feed a therapeutic renal diet (TRD) had been received, 72% owners reported feeding a TRD as a component of their cat's diet versus 7% of owners who had not received a veterinary recommendation to feed a therapeutic renal diet'



Survey of 859 UK owners of cats with CKD, published online JFMS 2016











### What is the role of phosphate binders?

- P restriction recommended for IRIS Stage 2, 3 and 4 cats
- **Indications** 
  - Cats/owners where transition to a renal diet not possible
  - Where phosphate levels remain 'high' in spite of renal diet
- Oral phosphate binders
  - In combination with renal diet/normal cat food
  - Bind P present in the diet, bound P excreted in the faeces
  - Check P levels after 6-8 weeks and adjust dose, add second agent if needed
  - (avoid sucralfate: V+, constipation, lack of efficacy) Quimby & Lappin 2016 JAAHA 52:8-12



















### Owner top tips for acceptance of IPB

(859 owners of cats with CKD)

- Start with a low dose and then increase
- Mix thoroughly with food
- Use wet food if possible
- Use empty gelatin capsules to dose the cat
- Syringe down the phosphate binder
- Add to dry food in a bag and leave overnight
- Use strong tasting food to disguise the binder
- Warm the food
- Use a low dose several times a day
- Add to the food when the cat is not looking!

















# Use of phosphate binders

### Quality of evidence as an intervention

- Increased longevity: No data, likely to be GOOD
- Improved QoL: No data

#### Panel recommendations

 If a renal diet cannot be used, or is insufficient to control serum phosphate, phosphate binders should be used (given with food), the response monitored, and the dose adjusted accordingly ...adopting the IRIS target phosphate concentrations

ISFM Consensus Guidelines on the Diagnosis and Management of Feline CKD, 2016











### Inappetence, nausea and vomiting

- Consider possible causes:
  - 'Uraemia'
  - Hypergastrinaemia
  - Dehydration
  - Electrolyte disturbances e.g. hypokalaemia
  - Anaemia
  - Chronic pain

- Find and fix whatever you can
- General advice for the owner
- Symptomatic options
- Don't introduce new diet when 'unwell'
- More aggressive support if needed















Find and fix whatever you can

- Correct dehydration
- Correct electrolyte imbalances
- Support anaemia where present
- Treat nausea and vomiting
  - Maropitant (1 mg/kg q24h)
  - Mirtazapine (1-2 mg/cat q48h)
- ☐ Is analgesia required (e.g. osteoarthritis)
  - e.g. meloxicam, buprenorphine
- Other options?
  - e.g. B12 supplementation















### Managing inappetence, nausea & vomiting

#### Quality of evidence as an intervention

- Increased longevity: No data
- Improved QoL: Likely to be GOOD if cat symptomatic

#### Panel recommendations

 Vomiting should be actively managed...and nausea always be considered as a potential contributory cause in cats with inappetence...centrally acting antiemetics likely to be most valuable and mirtazapine may have additional benefits.....

ISFM Consensus Guidelines on the Diagnosis and Management of Feline CKD, 2016













### Appetite support: tips for owners

- Offer food little and often
  - Remove if not eaten within 1-2 hours
  - Buffet not recommended
- Choice of bowl type
  - Metal/glass/china
  - Wide, shallow
- **Bowl location** 
  - Quiet, away from other resources
- Room temperature or slightly warmer
- Try popular foods
  - Chicken, fish, prawns...

- Nursing tactics
  - Sit with the cat, groom, hand feed
- Elderly cats
  - Raise the food bowl
  - Mash the food
- Others
  - ? Catnip
  - Avoid stressful events with feeding
  - Avoid items toxic to cats e.g. onion

















Attention to detail, an individualised approach and dedicated carer → → → optimal outcome







### Summary: key points

- Slowing progression of disease
  - Phosphate restriction
    - Key target for all azotaemic patients
    - Achieved through using renal diet and/or phosphate binders
    - Use the IRIS target ranges for phosphate levels
  - Consider RAAS suppression
    - Prioritise for proteinuric patients
- Symptomatic support improves quality of life













- Signalment: 12y FN DSH
- Referring VS history
  - Healthy weight around 4.5 kg, some dental disease in the past
  - Feb and Oct 2016: senior health checks: all OK, SBP 138 mmHg, USG 1.045
  - Jan and July 2017: senior health check: USG 1.035, bloods OK, weight 4.6 kg
    - Dental July 2018
  - Aug 2018: USG 1.035, creatinine û, weight 4.5 kg, therapeutic renal diet prescribed













- Presenting history Jan 2019
  - Sula's appetite has reduced and she is now rapidly losing weight
    - Oct 2018: weight 4.2 kg
    - Dec 2018: weight 3.85 kg
    - Jan 2019: weight 3.7 kg (compared to her healthy weight, Sula has lost 18% of her bodyweight)
  - Sula prefers wet food to dry but even her appetite for wet food is poor
  - Waxing and waning loose faeces reported
  - Renal parameters stable, IRIS Stage 2-3 CKD, non proteinuric and non hypertensive, phosphate 1.6, mild hypercalcaemia (2.96 mmol/l)











- What's a sensible approach?
  - Detailed assessment (physical exam, lab work, renal imaging) to make sure we find as many reasons as possible for the poor appetite
  - Reassess calcium, ionized calcium
  - 'Find and fix' as many things as possible
    - Complications from CKD
      - Dehydration, electrolyte disturbances (K+), anaemia, pyelonephritis...
    - Other causes of poor appetite?
      - Chronic pain (dental disease, osteoarthritis?), other illnesses?











- What if appetite remains poor in spite of this....?
  - Symptomatic options?
    - Appetite stimulants e.g. mirtazapine: 1-2 mg per cat every 48 hours?
    - Anti-emetic e.g. maropitant: 1 mg every 24 hours?
    - Nursing care and TLC from owners
    - Nutritional assessment: calorie dense foods
    - Tube feeding?













- What happened?
  - Sula's owners keen to proceed with investigations
  - EPOC calcium analyser out of slides...decided to await these before completing investigations (ionized calcium test)
  - Mirtazapine 1 mg PO EOD prescribed with advice on nursing care and support of appetite, phosphate binder supplied for use with non TRD food
  - Progress report (phone, 1 week later): improvements seen, owners happy with initial progress











- Reviewed Sula 3 weeks after first appointment
  - Physical exam: kidneys a little small, 0.45 kg bodyweight gained in 3 weeks!
    - Blood results: SDMA 21, Creatinine 231, Phosphate 1.7, Potassium 3.4
    - B12, folate, SpecFPL: all normal
    - Ionised calcium normal, T4 24, USG 1.028, UPC 0.1
    - Imaging: Left kidney very small, some mineralisation, both kidneys had reduced corticomedullary definition. Some mild diffuse thickening of small intestine

















- What's our assessment?
  - Poor appetite consistent with IRIS Stage 2-3 CKD
  - No further complications at this point
  - Cause of loose faeces not determined: ? IBD, dietary, neoplasia ?
  - Excellent response to symptomatic support of CKD
- Initial plan
  - Continue with supportive approach, monitor progress













- Treatment plan
  - Potassium supplement (Kaminox)
  - Phosphate binder (Renate)
  - Continued appetite support (mirtazapine)
  - Good progress but not keen on tablets
  - Transdermal mirtazapine discussed
    - No licensed preparation available in the UK but has been studied in the USA (healthy cats and cats with CKD)



















#### Transdermal mirtazapine

- Assessment of compounded transdermal mirtazapine as an appetite stimulant in cats with chronic kidney disease. Quimby et al 2019, JFMS online
  - Study 1, 9 client owned cats with IRIS Stage 2 or 3 CKD
    - 3.75 mg TD mirtazapine or placebo TD EOD for 3 weeks; 4 day washout then crossed over to alternate treatment.
  - Study 2, 10 client owned cats with IRIS Stage 2 or 3 CKD
    - 1.88 mg TD mirtazapine or placebo TD EOD for 3 weeks then crossed over
  - Statistically significant increase in appetite, food consumption and weight with both. No significant difference in activity/vocalization between groups although some cats had increased vocalization















#### MIRTAZAPINE 40 mg/ml

Transdermal Gel

Mirataz™ (mirtazapine transdermal ointment)

- Current treatment recommendations
  - 2 mg/cat starting dose (same as oral dose)
  - EOD treatment for cats with CKD
  - Use licensed product where available
  - Reduce dose if side-effects seen



Transdermal mirtazapine

















- Follow-up
  - Remained on oral mirtazapine (2 mg eod)
  - Compliance to TRD greatly improved
  - May 2019: 4.4 kg
  - Owners great at recording and sharing data on her appetite and weight measurements at home...

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#### Outcome

- Last update Jan 2020 and Sula continues to remain stable and doing well
  - SDMA 27, creatinine 228, phosphate 1.5, potassium 4.5, USG 1.028
  - Continued soft stool have resulted in some dietary 'tweaks' - RC sensitivity + RC renal combination currently

















- Main lesson from Sula...
  - Supportive treatments can make a massive difference in some cases













#### Useful resources

www.rcvsknowledge.org/QIResources

Bit.ly/WebinarQI

ISFM Consensus Guidelines on the Diagnosis and Management of Feline Chronic Kidney Disease

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Catherine Langston

















#### Over to Paul....



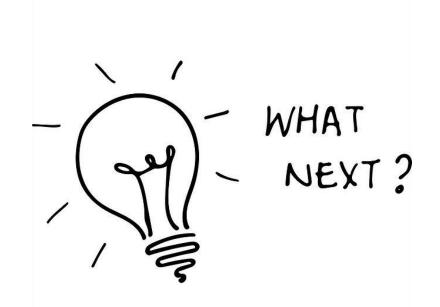
























#### Good CPD

• It is not just what you hear...... it is what you **do** as a result of what you hear.





























#### Aims of phosphate restriction

- Your patient will feel better
- Your patient will live longer
- Their renal disease is less likely to progress
- **Current IRIS recommendations** 
  - Phosphate restriction recommended for all azotaemic CKD patients (i.e. Stages 2, 3 and
  - Phosphate restriction recommended irrespective of patient's blood phosphate levels

















I have learned that phosphate restriction is very important in the management of renal disease.

















## What will I do differently when I go back into practice?















I am going to recommend renal diets for all my renal patients and explain the benefits to my clients.















### But this CPD is important - and expensive... Is there anything else I can do?









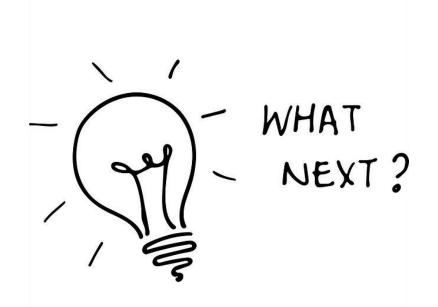






The CPD Pyramid

















### Why bother with a guideline?













#### Why do we feed a renal diet?

- I don't know
- It is kinder to the kidneys
- It slows down the disease
- It has less protein
- It isn't that tasty
- It is tastier than other foods
- I only know that you can't feed it to sick cats if they are not eating.













### Clinical Guidelines - Why bother?

- It's a complex progressive disease.
  - Your patient will feel better
  - Your patient will live longer
  - Their renal disease is less likely to progress
- Source of reference.
- Guide to best practice.
- Increase in clinical standards.
- Should be reviewed and adjusted with changing opinion.













# How to write a guideline

- Pick a topic
- Gather a team and do some research
  - -Our nurses are invaluable. Speak to the reception team.
  - −Get ownership with a bottom up approach.
- Write the guideline keep it simple. One that works for you
- Supply references
- Release it and ask for feedback
- Launch it with a review date
- Undertake a process audit





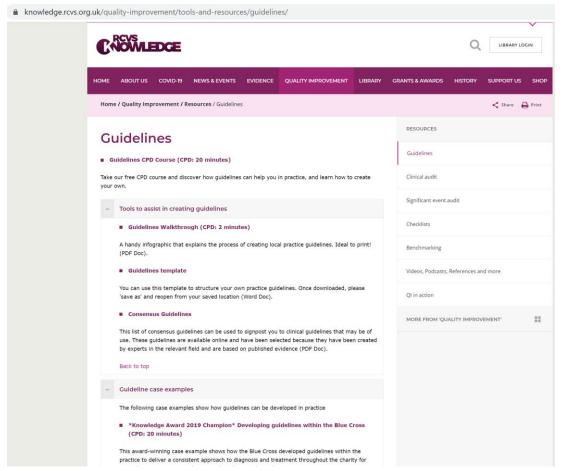














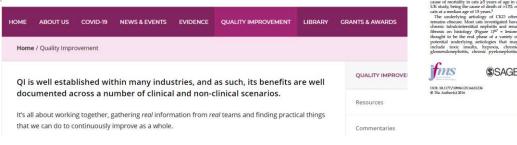














Journal of Feline Medicine and Surgery (2016) 18, 219-239













#### How to write a guideline on....

- We have the key members of our team.
- We have done our research.
- Now let's create a guideline to fit your practice!















#### Does anyone know who this man is?















#### John Biggs

"Learning is constructed by what activities the students carry out; learning is about what they do, not about what we teachers do."













#### Things I have learned...

- It's important and it takes time. It needs to be protected and designated.
- Think about your team and clients values. Financial, better medicine, animal welfare.
- Buy in makes it work.
- Educate your team and give them examples.
- Undertake a process audit and ask for feedback.











#### Things to consider

- Keep it simple.
- Follow up with a phone call
- Create reminders
- Make diet an important question in your history taking
- Encourage your clients, give them options, know their values
- Educate them and explain the value of the diet.
- Ask open questions "tell me about Fluffy's diet" and listen
- Audit your process reflect and adjust. Consider benchmarking.















#### Owner top tips for acceptance of IPB

(859 owners of cats with CKD)

- Start with a low dose and then increase
- Mix thoroughly with food
- Use wet food if possible
- Use empty gelatin capsules to dose the cat
- Syringe down the phosphate binder
- Add to dry food in a bag and leave overnight
- Use strong tasting food to disguise the binder
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#### Appetite support: tips for owners

- Offer food little and often
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  - Mash the food
- Others
  - ? Catnip
  - Avoid stressful events with feeding
  - Avoid items toxic to cats e.g. onion















#### How can QI improve what we do

A real life example













#### Guideline on heart murmurs

If a murmur is noted in an older cat, always recommend an echo.

Give a handout explaining SRR, the disease process and how an echo works











#### Process audit - heart murmurs in cats:

- 11 year old cat presented for a health check. On cardiac auscultation a grade 2/6 heart murmur was heard for the first time.
- Discussed heart murmur and recommended monitor SRR at home. Handout given. Owner to call in with SRR values. Owner is to consider an echo.











#### Process audit - heart murmurs in cats:

- 10 year old cat with murmur No discussion or recommendation in the notes handout given
- 12 year old cat in for health check murmur detected dental needed. No handout given
- 9 year old cat with breathing issue Grade 1 murmur no handout or recommendation
- 19 year old cat Heart murmur and gallop rhythm. No handout given. No discussion in clinical notes - why. Was owner not interested or was vet not interested?
- 2nd opinion 14 year old cat. Arrhythmia with dropped beat. Breathing issues. Was PTS
- Primary care vets: 4 months previously mms pink and moist, crt<2s, no skin tent, chest/heart- heart murmur 2/6, abdo comf on palpation. Sore leg. Treated with pain relief













- Templates and walkthroughs to get you started
- Real case studies to download and read
- Insightful articles and opinion pieces
- QI podcasts you can listen to on the go

www.rcvsknowledge.org/QIResources

#### In Summary

- Gather a team and do some reading.
- Come up with a plan and give it a go.
- Reassess in a few months and tweak it.

Good CPD is not defined by what you hear, but by what you do as a result of what you hear.















### Further questions?

- www.rcvsknowledge.org/QIResources
- sarah@vetprofessionals.com
- Vet Professionals Ltd; www.vetprofessionals.com













# Time to reflect



- What is your take home message for this session?
- Is there anything you are going to do differently as a result of this session? Why?



