Supplementary file 7: GRADE evidence tables PTH replacement therapy

QoL

Certainty assessment						
Participants (studies)	**Risk of bias**	**Inconsistency**	**Indirectness**	**Imprecision**	**Publication bias**	**Overall certainty of evidence**
203 (3 RCTs)	serious[a]	serious[b]	not serious	serious[c]	none	⊕◯◯◯ Very low[a,b,c]
154 (6 non-randomised studies)	serious[d]	not serious	not serious	serious[c]	none	⊕◯◯◯ Very low[c,d]

Explanations

a. Some concerns on randomisation process.

b. QoL improvement with PTH[1-34], but not with PTH[1-84].

c. Small sample sizes.

d. Concerns on selection of patients.

Calcium and phosphate levels

Certainty assessment						
Participants (studies)	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Overall certainty of evidence
327 (4 RCTs)	serious[a]	not serious	not serious	serious[b]	none	⊕⊕◯◯ Low[a,b]
257 (9 non-randomised studies)	serious[c]	not serious	not serious	serious[b]	none	⊕◯◯◯ Very low[b,c]

Explanations

a. Some concerns on randomisation process, adhering to intervention and missing outcome data.

b. Small sample sizes.

c. Concerns on selection of patients.

CKD and renal calcifications

Certainty assessment						
Participants (studies)	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Overall certainty of evidence

Certainty assessment						
82 (1 RCT)	serious[a]	not applicable[b]	not serious	serious[c]	none	⊕◯◯◯ Very low[a,b,c]
253 (6 non-randomised studies)	serious[d]	serious[e]	not serious	serious[c]	none	⊕◯◯◯ Very low[c,d,e]

Explanations

a. Some concerns on randomisation process and possible confounders.

b. Can not reliably be judged with only one study.

c. Small sample sizes.

d. Some concerns on selection of patients and possible confounders.

e. Various effects on kidney function reported.

Cramps, tetany, seizures, and neuropsychological endpoints

Certainty assessment						
Participants (studies)	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Overall certainty of evidence
113 (4 non-randomised studies)	serious[a]	not serious	not serious	serious[b]	none	⊕◯◯◯ Very low[a,b]

Explanations

a. Concerns on selection of patients and no standardized assessment of outcome.

b. Small sample sizes.

CVD

Certainty assessment						
Participants (studies)	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Overall certainty of evidence

Certainty assessment						
127 (2 non-randomised studies)	serious[a]	not serious	not serious	serious[b]	none	⊕◯◯◯ Very low[a,b]

Explanations

a. Concerns on selection of patients, possible confounders and assessment of outcome.

b. Small sample sizes.

Bone markers, fracture, BMD

Certainty assessment						
Participants (studies)	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Overall certainty of evidence
148 (3 RCTs)	very serious[a]	not serious	not serious	serious[b]	none	⊕◯◯◯ Very low[a,b]
194 (7 non-randomised studies)	serious[c]	serious[d]	not serious	serious[b]	none	⊕◯◯◯ Very low[b,c,d]

Explanations

a. Concerns on selection and randomisation process, and possible confounders.

b. Small sample sizes.

c. Concerns on selection of patients, possible confounders.

d. Inconsistent results for BMD effects.

Pill burden

Certainty assessment						
Participants (studies)	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Overall certainty of evidence
265 (3 RCTs)	serious[a]	not serious	not serious	serious[b]	strong association	⊕⊕⊕◯ Moderate[a,b]
225 (8 non-randomised studies)	serious[c]	not serious	not serious	serious[b]	strong association	⊕◯◯◯ Very low[b,c]

Explanations

a. Concerns on randomisation process, adhering to intervention and missing outcome data.

b. Small sample sizes.

c. Concerns on selection of patients.

Increased susceptibility to infection

Certainty assessment						
Participants (studies)	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Overall certainty of evidence
71 (2 non-randomised studies)	serious[a]	not applicable[b]	not serious	serious[c]	none	⊕◯◯◯ Very low[a,b,c]

Explanations

a. Concerns on selection of patients and no standardized assessment of outcome.

b. Can not reliably be judged with only two studies.

c. Small sample sizes.

Cataract

Certainty assessment						
Participants (studies)	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Overall certainty of evidence

Certainty assessment						
14 (1 non-randomised study)	serious[a]	serious[b]	not serious	very serious[c]	none	⊕◯◯◯ Very low[a,b,c]

Explanations

a. Concerns on selection of patients, and measurement of outcome not standardized.

b. Can not reliably be judged with only one study.

c. Very small sample size.