

**Table S1 Factors associated with osteoporosis in patients with primary biliary cholangitis**

Variable	Univariate		Multivariate	
	OR (95%CI)	P-value	OR (95%CI)	P-value
Female	1.840 (0.569–5.945)	0.308		
Age (years)	1.098 (1.046–1.153)	<0.001	1.090 (1.034–1.148)	0.001
BMI (kg/m <sup>2</sup> )	0.762 (0.653–0.890)	0.001	0.823 (0.690–0.982)	0.031
Liver cirrhosis	3.511 (0.991–12.436)	0.052		
Age at diagnosis (years)	1.063 (1.021–1.106)	0.003		
Duration of disease (years)	1.056 (0.993–1.123)	0.080		
AST (IU/L)	0.997 (0.967–1.029)	0.873		
ALT (IU/L)	0.953 (0.906–1.003)	0.064		
ALP (IU/L)	1.000 (0.998–1.003)	0.742		
GGT (IU/L)	0.998 (0.991–1.005)	0.517		
Total bilirubin (mg/dL)	1.105 (0.655–1.865)	0.708		
Albumin (g/dL)	0.277 (0.102–0.755)	0.012		
M2BPGi (C.O.I.)	1.125 (0.881–1.436)	0.347		
IGF-1 (ng/mL)	0.980 (0.967–0.993)	0.004		
BCAA (μmol/L)	0.994 (0.989–0.999)	0.018		
TRACP-5b (mU/dL)	1.002 (0.999–1.004)	0.218		
Total P1NP (ng/mL)	1.003 (0.983–1.024)	0.746		
SMI (kg/m <sup>2</sup> )	0.266 (0.144–0.491)	<0.001		
Handgrip strength (kg)	0.777 (0.693–0.870)	<0.001		
Lumbar spine BMD (g/cm <sup>2</sup> )	0.000 (0.000–0.005)	<0.001		
Femoral neck BMD (g/cm <sup>2</sup> )	0.000 (0.000–0.000)	<0.001		
Total hip BMD (g/cm <sup>2</sup> )	0.000 (0.000–0.000)	<0.001		
Sarcopenia	10.000 (3.778–26.470)	<0.001	4.126 (1.280–13.297)	0.018
Vertebral fracture	6.175 (2.251–16.939)	<0.001		

ALP, alkaline phosphatase; ALT, alanine aminotransferase; AST, aspartate aminotransferase; BCAA, branched-chain amino acids; BMD, bone mineral density; BMI, body mass index; CI, confidence interval; GGT, gamma-glutamyltransferase; IGF-1, insulin-like growth factor 1; M2BPGi, Mac-2 binding protein glycosylation isomer; OR, odds ratio; P1NP, procollagen type I N-terminal propeptide; skeletal muscle mass index; SMI, skeletal muscle mass index; TRACP-5b, tartrate resistant acid phosphatase-5b.

**Table S2** Factors associated with sarcopenia in patients with primary biliary cholangitis

Variable	Univariate		Multivariate	
	OR (95%CI)	P-value	OR (95%CI)	P-value
Female	2.000 (0.541–7.388)	0.298		
Age (years)	1.053 (1.009–1.099)	0.019		
BMI (kg/m <sup>2</sup> )	0.695 (0.576–0.839)	<0.001	0.708 (0.568–0.883)	0.002
Liver cirrhosis	4.857 (1.351–17.457)	0.015		
Age at diagnosis (years)	1.051 (1.009–1.095)	0.018		
Duration of disease (years)	0.998 (0.932–1.070)	0.963		
AST (IU/L)	1.015 (0.985–1.045)	0.327		
ALT (IU/L)	0.966 (0.918–1.016)	0.182		
ALP (IU/L)	1.001 (0.998–1.004)	0.542		
GGT (IU/L)	0.999 (0.992–1.006)	0.709		
Total bilirubin (mg/dL)	1.626 (0.868–3.049)	0.129		
Albumin (g/dL)	0.210 (0.072–0.611)	0.004		
M2BPGi (C.O.I.)	1.362 (1.013–1.831)	0.041		
IGF-1 (ng/mL)	0.985 (0.972–0.999)	0.033		
BCAA ( $\mu$ mol/L)	0.997 (0.993–1.002)	0.215		
TRACP-5b (mU/dL)	1.001 (0.998–1.004)	0.432		
Total P1NP (ng/mL)	0.999 (0.978–1.021)	0.935		
SMI (kg/m <sup>2</sup> )	0.024 (0.005–0.108)	<0.001		
Handgrip strength (kg)	0.617 (0.505–0.755)	<0.001		
Lumbar spine BMD (g/cm <sup>2</sup> )	0.004 (0.000–0.069)	<0.001		
Femoral neck BMD (g/cm <sup>2</sup> )	0.001 (0.000–0.036)	<0.001		
Total hip BMD (g/cm <sup>2</sup> )	0.000 (0.000–0.025)	<0.001		
Osteoporosis	8.875 (3.376–23.334)	<0.001	3.420 (1.057–11.067)	0.040
Vertebral fracture	9.518 (3.340–27.126)	<0.001	10.936 (2.678–44.663)	0.001

ALP, alkaline phosphatase; ALT, alanine aminotransferase; AST, aspartate aminotransferase; BCAA, branched-chain amino acids; BMD, bone mineral density; BMI, body mass index; CI, confidence interval; GGT, gamma-glutamyltransferase; IGF-1, insulin-like growth factor 1; M2BPGi, Mac-2 binding protein glycosylation isomer; OR, odds ratio; P1NP, procollagen type I N-terminal propeptide; skeletal muscle mass index; SMI, skeletal muscle mass index; TRACP-5b, tartrate resistant acid phosphatase-5b.

**Table S3 Comparison of clinical characteristics between female patients with and without osteoporosis**

Variable	All patients	Osteoporosis	Non-osteoporosis	P-value
Patients, n (%)	96	29 (30.2)	67 (69.8)	
Age (years)	67.0 (57.0–73.0)	71.0 (66.5–77.0)	63.0 (55.0–70.0)	<0.001
BMI (kg/m <sup>2</sup> )	22.3 (20.1–24.9)	20.3 (18.6–22.8)	22.7 (20.7–26.0)	0.002
Liver cirrhosis, n (%)	7 (7.3)	5 (17.2)	2 (3.0)	0.014
Age at diagnosis (years)	60.5 (50.0–68.0)	64.0 (54.0–72.0)	57.0 (49.0–66.0)	0.019
Duration of disease (years)	3.0 (1.0–8.0)	5.0 (2.0–13.0)	2.0 (1.0–6.0)	0.038
Menopause, n (%)	83 (86.5)	28 (96.6)	55 (82.1)	0.057
Duration of menopause (years)	15.0 (6.3–24.0)	23.0 (14.5–27.0)	13.0 (3.0–18.0)	<0.001
AST (IU/L)	25 (21–30)	26 (23–30)	24 (20–29)	0.565
ALT (IU/L)	20 (14–24)	18 (14–22)	20 (14–24)	0.525
ALP (IU/L)	282 (216–348)	271 (206–319)	290 (224–363)	0.202
GGT (IU/L)	38 (24–60)	31 (25–54)	41 (23–65)	0.287
Total bilirubin (mg/dL)	0.6 (0.5–0.8)	0.5 (0.4–0.7)	0.6 (0.5–0.8)	0.092
Albumin (g/dL)	4.1 (3.9–4.3)	4.0 (3.9–4.3)	4.2 (3.9–4.4)	0.037
M2BPGi (C.O.I)	0.87 (0.57–1.36)	1.13 (0.62–1.66)	0.82 (0.57–1.23)	0.055
IGF-1 (ng/mL)	89 (61–112)	61 (46–101)	95 (68–116)	0.002
BCAA (μmol/L)	412 (356–460)	379 (326–430)	433 (379–482)	0.013
TRACP-5b (mU/dL)	351 (240–475)	384 (281–498)	343 (231–477)	0.402
P1NP (ng/mL)	47 (34–58)	47 (33–59)	48 (34–59)	0.780
SMI (kg/m <sup>2</sup> )	6.01 (5.32–6.48)	5.30 (4.97–5.86)	6.11 (5.74–6.57)	<0.001
Handgrip strength (kg)	21.8 (17.2–23.8)	17.0 (14.7–20.3)	22.7 (19.3–25.5)	<0.001
Lumbar spine BMD (g/cm <sup>2</sup> )	1.03 (0.87–1.17)	0.80 (0.71–0.91)	1.09 (0.97–1.21)	<0.001
Femoral neck BMD (g/cm <sup>2</sup> )	0.74 (0.67–0.84)	0.63 (0.58–0.69)	0.77 (0.72–0.89)	<0.001
Total hip BMD (g/cm <sup>2</sup> )	0.79 (0.71–0.88)	0.67 (0.60–0.71)	0.84 (0.77–0.93)	<0.001
Sarcopenia, n (%)	24 (25.0)	16 (55.2)	8 (11.9)	<0.001
Vertebral fracture, n (%)	16 (16.7)	10 (34.5)	6 (9.0)	0.002

Values are shown as median (interquartile range) or number (percentage). Statistical analysis was performed using the chi-squared test or the Mann-Whitney U test, as appropriate. ALP, alkaline phosphatase; ALT, alanine aminotransferase; AST, aspartate aminotransferase; BCAA, branched-chain amino acids; BMD, bone mineral density; BMI, body mass index; GGT, gamma-glutamyltransferase; IGF-1, insulin-like growth factor 1; M2BPGi, Mac-2 binding protein glycosylation isomer; P1NP, procollagen type I N-terminal propeptide; SMI, skeletal muscle mass index; TRACP-5b, tartrate-resistant acid phosphatase 5b.

**Table S4 Comparison of clinical characteristics between female patients with and without sarcopenia**

Variable	Sarcopenia	Non-sarcopenia	P-value
Patients, n (%)	24 (25.0)	72 (75.0)	
Age (years)	73.0 (65.0–76.8)	65.5 (56.0–71.0)	0.008
BMI (kg/m <sup>2</sup> )	19.6 (18.0–21.5)	22.8 (20.7–25.9)	<0.001
Liver cirrhosis, n (%)	4 (16.7)	3 (4.2)	0.041
Age at diagnosis (years)	66.5 (51.5–73.0)	57.0 (50.0–66.0)	0.022
Duration of disease (years)	3.5 (1.0–8.8)	2.0 (1.0–8.0)	0.698
Menopause, n (%)	22 (91.7)	61 (84.7)	0.389
Duration of menopause (years)	22.5 (13.3–28.8)	13.5 (5.0–20.0)	0.006
AST (IU/L)	25 (22–32)	26 (20–30)	0.946
ALT (IU/L)	18 (14–21)	20 (14–24)	0.358
ALP (IU/L)	284 (217–337)	281 (216–352)	0.800
GGT (IU/L)	34 (21–60)	39 (24–63)	0.449
Total bilirubin (mg/dL)	0.5 (0.5–0.7)	0.6 (0.5–0.8)	0.144
Albumin (g/dL)	4.0 (3.8–4.2)	4.2 (3.9–4.4)	0.077
M2BPGi (C.O.I.)	1.27 (0.70–1.79)	0.83 (0.55–1.22)	0.008
IGF-1 (ng/mL)	77 (48–104)	94 (66–114)	0.067
BCAA (μmol/L)	381 (326–493)	421 (377–458)	0.331
TRACP-5b (mU/dL)	390 (215–510)	344 (240–469)	0.682
P1NP (ng/mL)	38 (31–61)	48 (36–58)	0.268
SMI (kg/m <sup>2</sup> )	5.08 (4.84–5.92)	6.27 (5.88–6.57)	<0.001
Handgrip strength (kg)	16.6 (14.1–17.2)	22.6 (20.0–25.2)	<0.001
Lumbar spine BMD (g/cm <sup>2</sup> )	0.88 (0.64–0.74)	1.08 (0.92–1.21)	<0.001
Femoral neck BMD (g/cm <sup>2</sup> )	0.68 (0.64–0.74)	0.77 (0.70–0.89)	<0.001
Total hip BMD (g/cm <sup>2</sup> )	0.72 (0.66–0.78)	0.82 (0.73–0.92)	0.001
Osteoporosis, n (%)	16 (66.7)	13 (18.1)	<0.001
Vertebral fracture, n (%)	10 (41.7)	6 (8.3)	<0.001

Values are shown as median (interquartile range) or number (percentage). Statistical analysis was performed using the chi-squared test or the Mann-Whitney U test, as appropriate. ALP, alkaline phosphatase; ALT, alanine aminotransferase; AST, aspartate aminotransferase; BCAA, branched-chain amino acids; BMD, bone mineral density; BMI, body mass index; GGT, gamma-glutamyltransferase; IGF-1, insulin-like growth factor 1; M2BPGi, Mac-2 binding protein glycosylation isomer; P1NP, procollagen type I N-terminal propeptide; SMI, skeletal muscle mass index; TRACP-5b, tartrate-resistant acid phosphatase 5b.

**Table S5** Factors associated with osteoporosis in female patients with primary biliary cholangitis

Variable	Univariate		Multivariate	
	OR (95%CI)	P-value	OR (95%CI)	P-value
Age (years)	1.088 (1.034–1.145)	0.001	1.074 (1.018–1.132)	0.008
BMI (kg/m <sup>2</sup> )	0.790 (0.676–0.924)	0.003		
Liver cirrhosis	6.771 (1.230–37.262)	0.028		
Age at diagnosis (years)	1.054 (1.010–1.099)	0.015		
Duration of disease (years)	1.055 (0.989–1.125)	0.107		
Menopause	6.109 (0.756–49.398)	0.090		
Duration of menopause (years)	1.092 (1.039–1.148)	0.001		
AST (IU/L)	1.002 (0.964–1.041)	0.925		
ALT (IU/L)	0.959 (0.907–1.014)	0.145		
ALP (IU/L)	1.000 (0.997–1.003)	0.975		
GGT (IU/L)	0.996 (0.988–1.005)	0.424		
Total bilirubin (mg/dL)	1.332 (0.677–2.621)	0.406		
Albumin (g/dL)	0.247 (0.075–0.818)	0.022		
M2BPGi (C.O.I.)	1.197 (0.881–1.627)	0.250		
IGF-1 (ng/mL)	0.978 (0.963–0.993)	0.005		
BCAA (μmol/L)	0.996 (0.991–1.001)	0.140		
TRACP-5b (mU/dL)	1.001 (0.999–1.004)	0.375		
Total P1NP (ng/mL)	1.000 (0.979–1.021)	0.969		
SMI (kg/m <sup>2</sup> )	0.277 (0.137–0.560)	<0.001		
Handgrip strength (kg)	0.740 (0.643–0.852)	<0.001		
Lumbar spine BMD (g/cm <sup>2</sup> )	0.000 (0.000–0.004)	<0.001		
Femoral neck BMD (g/cm <sup>2</sup> )	0.000 (0.000–0.000)	<0.001		
Total hip BMD (g/cm <sup>2</sup> )	0.000 (0.000–0.000)	<0.001		
Sarcopenia	9.077 (3.209–25.672)	<0.001	6.510 (2.100–20.180)	0.001
Vertebral fracture	5.351 (1.719–16.657)	0.004		

ALP, alkaline phosphatase; ALT, alanine aminotransferase; AST, aspartate aminotransferase; BCAA, branched-chain amino acids; BMD, bone mineral density; BMI, body mass index; CI, confidence interval; GGT, gamma-glutamyltransferase; IGF-1, insulin-like growth factor 1; M2BPGi, Mac-2 binding protein glycosylation isomer; OR, odds ratio; P1NP, procollagen type I N-terminal propeptide; skeletal muscle mass index; SMI, skeletal muscle mass index; TRACP-5b, tartrate resistant acid phosphatase-5b.

**Table S6** Factors associated with sarcopenia in female patients with primary biliary cholangitis

Variable	Univariate		Multivariate	
	OR (95%CI)	P-value	OR (95%CI)	P-value
Age (years)	1.055 (1.007–1.106)	0.025		
BMI (kg/m <sup>2</sup> )	0.682 (0.555–0.839)	<0.001	0.706 (0.562–0.888)	0.003
Liver cirrhosis	4.600 (0.950–22.278)	0.058		
Age at diagnosis (years)	1.049 (1.004–1.097)	0.032		
Duration of disease (years)	1.005 (0.937–1.077)	0.899		
Menopause	1.984 (0.407–9.664)	0.397		
Duration of menopause (years)	1.072 (1.021–1.126)	0.005		
AST (IU/L)	1.003 (0.963–1.045)	0.887		
ALT (IU/L)	0.960 (0.904–1.020)	0.183		
ALP (IU/L)	1.001 (0.997–1.004)	0.771		
GGT (IU/L)	0.998 (0.990–1.007)	0.713		
Total bilirubin (mg/dL)	1.464 (0.711–3.014)	0.301		
Albumin (g/dL)	0.289 (0.089–0.941)	0.039		
M2BPGi (C.O.I.)	1.245 (0.909–1.705)	0.171		
IGF-1 (ng/mL)	0.987 (0.973–1.001)	0.074		
BCAA (μmol/L)	0.999 (0.995–1.004)	0.745		
TRACP-5b (mU/dL)	1.001 (0.998–1.003)	0.614		
Total P1NP (ng/mL)	0.990 (0.966–1.014)	0.411		
SMI (kg/m <sup>2</sup> )	0.007 (0.001–0.059)	<0.001		
Handgrip strength (kg)	0.527 (0.391–0.711)	<0.001		
Lumbar spine BMD (g/cm <sup>2</sup> )	0.009 (0.001–0.147)	0.001		
Femoral neck BMD (g/cm <sup>2</sup> )	0.002 (0.000–0.158)	0.006		
Total hip BMD (g/cm <sup>2</sup> )	0.001 (0.000–0.084)	0.002		
Osteoporosis	9.077 (3.209–25.672)	<0.001	4.012 (1.186–13.576)	0.026
Vertebral fracture	7.857 (2.451–25.187)	0.001	8.898 (1.828–43.305)	0.007

ALP, alkaline phosphatase; ALT, alanine aminotransferase; AST, aspartate aminotransferase; BCAA, branched-chain amino acids; BMD, bone mineral density; BMI, body mass index; CI, confidence interval; GGT, gamma-glutamyltransferase; IGF-1, insulin-like growth factor 1; M2BPGi, Mac-2 binding protein glycosylation isomer; OR, odds ratio; P1NP, procollagen type I N-terminal propeptide; skeletal muscle mass index; SMI, skeletal muscle mass index; TRACP-5b, tartrate resistant acid phosphatase-5b.

**Table S7 Characteristics of patients with and without sarcopenia/osteoporosis**

Variable	Osteoporosis (-)	Osteoporosis (+)	Osteoporosis (-)	Osteoporosis(+)	P-value
	Sarcopenia (-)	Sarcopenia(-)	Sarcopenia (+)	Sarcopenia (+)	
Patients, n (%)	75 (64.1)	15 (12.8)	9 (7.7)	18 (15.4)	
Female, n (%)	59 (78.7)	13 (86.7)	8 (88.9)	16 (88.9)	0.646
Age (years)	63.0 (53.0–70.0)	73.0 (68.0–77.0)	73.0 (56.0–76.0)	73.5 (65.8–81.5)	<0.001
BMI (kg/m <sup>2</sup> )	22.9 (21.2–26.0)	22.6 (20.2–24.2)	21.2 (18.9–24.2)	19.3 (18.1–21.4)	<0.001
Liver cirrhosis, n (%)	3 (4.0)	2 (13.3)	2 (22.2)	4 (22.2)	0.045
Age at diagnosis (years)	57.0 (48.0–66.0)	66.0 (57.0–71.0)	72.0 (53.0–74.0)	66.5 (52.5–74.3)	0.003
Duration of disease (years)	2.0 (1.0–6.0)	5.0 (2.0–15.0)	2.0 (1.0–3.5)	4.0 (1.0–10.5)	0.223
AST (IU/L)	24 (20–33)	27 (23–30)	24 (21–31)	26 (23–33)	0.964
ALT (IU/L)	20 (15–27)	18 (14–22)	18 (13–26)	18 (14–22)	0.478
ALP (IU/L)	290 (229–363)	271 (200–314)	300 (240–379)	272 (215–327)	0.645
GGT (IU/L)	43 (29–74)	28 (24–52)	41 (18–66)	33 (24–71)	0.391
Total bilirubin (mg/dL)	0.6 (0.5–0.8)	0.6 (0.4–0.8)	0.6 (0.5–1.0)	0.5 (0.4–0.6)	0.150
Albumin (g/dL)	4.2 (3.9–4.3)	4.0 (3.9–4.3)	4.0 (3.4–4.5)	3.9 (3.7–4.1)	0.037
M2BPGi (C.O.I.)	0.77 (0.57–1.12)	0.96 (0.52–1.34)	0.87 (0.67–3.59)	1.47 (0.99–1.71)	0.009
IGF-1 (ng/mL)	95 (73–117)	61 (51–101)	88 (57–114)	61 (45–96)	0.010
BCAA (μmol/L)	442 (396–500)	380 (325–436)	407 (321–486)	366 (310–443)	0.039
TRACP-5b (mU/dL)	328 (237–429)	384 (309–444)	440 (210–500)	390 (227–484)	0.499
P1NP (ng/mL)	44 (35–58)	48 (35–68)	52 (30–77)	40 (33–63)	0.837
SMI (kg/m <sup>2</sup> )	6.48 (6.03–7.12)	5.90 (5.75–6.35)	5.27 (4.91–5.68)	5.00 (4.43–5.24)	<0.001
Handgrip strength (kg)	23.9 (21.9–28.1)	21.9 (16.8–22.5)	17.0 (14.8–17.3)	16.6 (12.8–17.5)	<0.001
Lumbar spine BMD (g/cm <sup>2</sup> )	1.11 (1.01–1.23)	0.81 (0.70–0.87)	0.96 (0.85–1.05)	0.84 (0.74–0.94)	<0.001
Femoral neck BMD (g/cm <sup>2</sup> )	0.82 (0.74–0.90)	0.60 (0.57–0.67)	0.75 (0.70–0.83)	0.65 (0.58–0.71)	<0.001
Total hip BMD (g/cm <sup>2</sup> )	0.85 (0.79–0.93)	0.69 (0.58–0.71)	0.79 (0.77–0.88)	0.67 (0.62–0.72)	<0.001
Vertebral fracture, n (%)	5 (6.7)	3 (20.0)	3 (33.3)	10 (55.6)	<0.001

Values are shown as median (interquartile range) or number (percentage). Statistical analysis was performed using the chi-squared test or the Kruskal-Wallis test, as appropriate. ALP, alkaline phosphatase; ALT, alanine aminotransferase; AST, aspartate aminotransferase; BCAA, branched-chain amino acids; BMD, bone mineral density; BMI, body mass index; GGT, gamma-glutamyltransferase; IGF-1, insulin-like growth factor 1; M2BPGi, Mac-2 binding protein glycosylation isomer; P1NP, procollagen type I N-terminal propeptide; SMI, skeletal muscle mass index; TRACP-5b, tartrate-resistant acid phosphatase 5b.