Supplementary Table 2. Other potential pharmacological options and therapeutic targets for NAFLD/NASH or CKD.

| | NAFLD/NASH | CKD |
|---|---|--|
| Targeting inflammation | | |
| NLRP3 inflammasome inhibitors | Diacerein : Phase-3-trial (NCT02242149): reduced liver stiffness in NAFLD with T2DM after two-year treatment [174]. | Canakinumab (IL-1β inhibition): CANTOS trial (NCT01327846): safe, reduced CVD event rates in moderate CKD [175]. |
| NF-κB inhibitors | Preclinical studies | Bindarit : Phase-2-trial (NCT01109212): investigate its effects in DN, but no results yet. |
| Targeting extracellular mediators of inflan | nmation | |
| Chemokine antagonists | Cenicriviroc: Phase-2b-CENTAUR (NCT02217475): well tolerated, improved fibrosis [176] Phase-3-AURORA (NCT03028740): terminated and undisclosed | PF-04634817: Phase-3-trial (NCT01712061): discontinued clinical development [177] BMS-813160: Phase-2a-trial (NCT01752985): terminated and undisclosed |
| Targeting a liver-derived amine oxidase | | |
| VAP-1 inhibitors | Preclinical studies | ASP8232 : Phase-2-ALBUM (NCT02358096): safe, reduced albuminuria in DN [178]. |
| Targeting common fibrogenic pathways | | |
| Galectin-3 antagonists | Belapectin (GR-MD-02): Phase-2b-NASH-CX (NCT02462967): reduced HVPG and development of varices in NASH cirrhosis patient without esophageal varices [179] Phase-2-NASH-FX (NCT02421094): no improvement in non-invasive biomarkers of liver inflammation or fibrosis [180] Phase-2/3-NAVIGATE (NCT04365868): ongoing | GCS-100: Phase-2b-trial (NCT02312050): completed but no result. |
| ROCK inhibitors | Preclinical studies | SAR407899: Phase-1-trial (NCT01485900):no |

| EGF inhibitors | Eulatinik, Phase 1/2 toial (MCT02272262), and an amiting | result | |
|--|--|--|--|
| EGF innibitors | Erlotinib: Phase-1/2-trial (NCT02273362): not recruiting | Preclinical studies | |
| Other current and emerging medications on NAFLD/NASH independently of renal outcomes | | | |
| | NAFLD/NASH | CKD | |
| Vitamin E (benefit for steatosis and inflammation) | Vitamin E : may be used for the treatment of some NASH patients, recommended by EASL, AASLD and KASL[1,2,181]. | Tocotrienol-Rich Vitamin E : Phase-2b-trial:improved serum creatinine and eGFR but not uACR[182]. | |
| Orlistat (approved to anti-obesity) | may be used for the improvement of steatosis and inflammation in obese people[183] | may induce acute kidney injury[184] | |
| Fibroblast growth factor 21 analogues (improve lipid and glucose metabolism) | LY2405319:Phase-1-trial (NCT01869959): improved lipid profile, body weight and fasting insulin levels in T2DM[185] PF-05231023:Phase-1b-trial (NCT01396187): improved lipid profile, body weight, no effect in glycemic control, in T2DM[186] Pegbelfermin:Phase-2-trial (NCT02413372):well-tolerated, reduced hepatic fat fraction in NASH[187] Phase-2-trial (NCT03486912/ NCT03486899):planning | Preclinical studies | |
| Acetyl-CoA carboxylase inhibitors (attenuate hepatic steatosis) | Firsocostat (GS-0976):Phase-2-trial (NCT02856555):decreased hepatic steatosis in NASH[188] | None | |
| β-selective THR agonists (benefit for steatosis and inflammation) | Resmetirom (MGL-3196): Phase-2-(NCT02912260): reduced hepatic fat in NASH[189]. Phase-3-MAESTRO-NASH (NCT03900429): ongoing | None | |
| Caspase inhibitors (anti-apoptotic) | Emricasan: Phase-2b-trial (NCT02686762): may worsen fibrosis and ballooning[190] | Preclinical studies | |

Abbreviations: NAFLD, non-alcoholic fatty liver disease; NASH, non-alcoholic steatohepatitis; CKD, chronic kidney disease; NLRP3, nucleotide-binding oligomerization domain-like receptor family pyrin domain containing 3; T2DM, type 2 diabetic mellitus; CVD, cardiovascular disease; DN, diabetic nephropathy; VAP-1, vascular adhesion protein-1; HVPG, hepatic venous pressure gradient; ROCK, rho-associated kinases; EGF, epidermal growth factor; EASL, European Association for the Study of the Liver; AASLD, American Association for the Study of Liver Diseases; KASL, Korean Association for the Study of the Liver; eGFR, estimated glomerular filtration gate; uACR, urinary albumin-to-creatinine ratio; KDOQI, Kidney Disease Outcomes Quality Initiative; β-selective THR, β-selective thyroid hormone receptor.