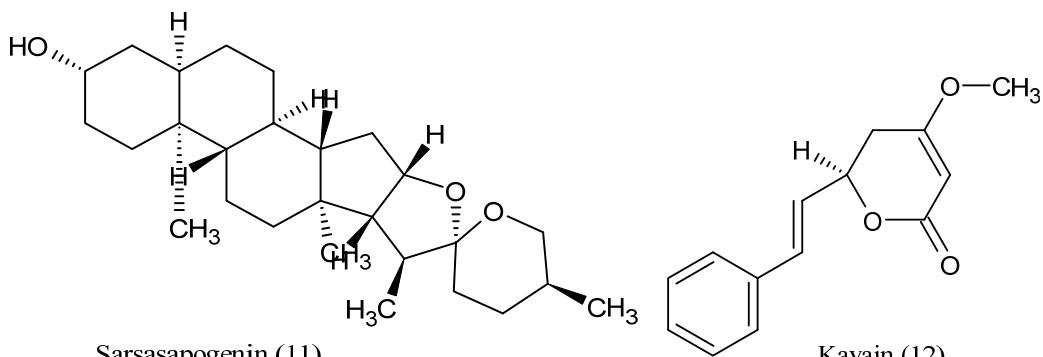


Cimiracemate A (8)

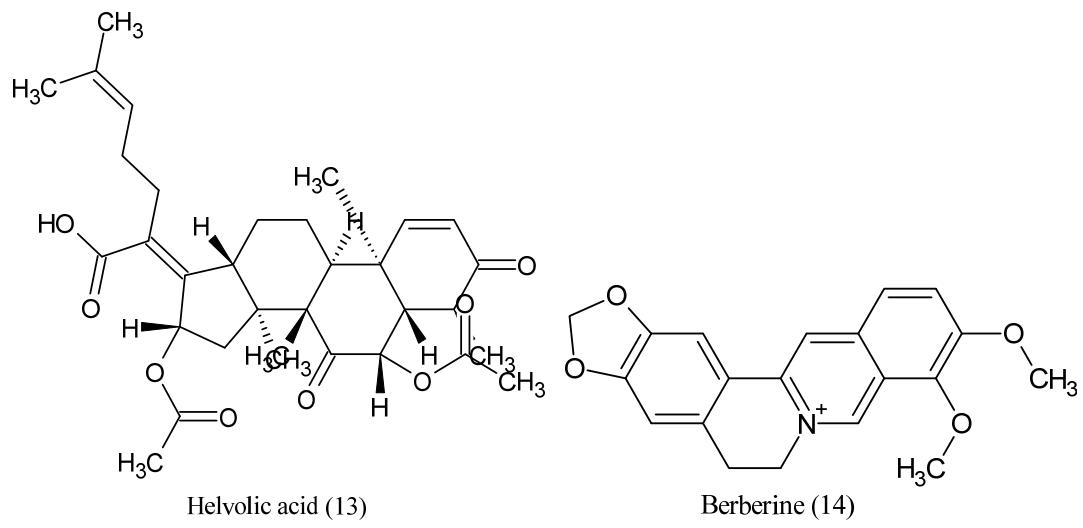
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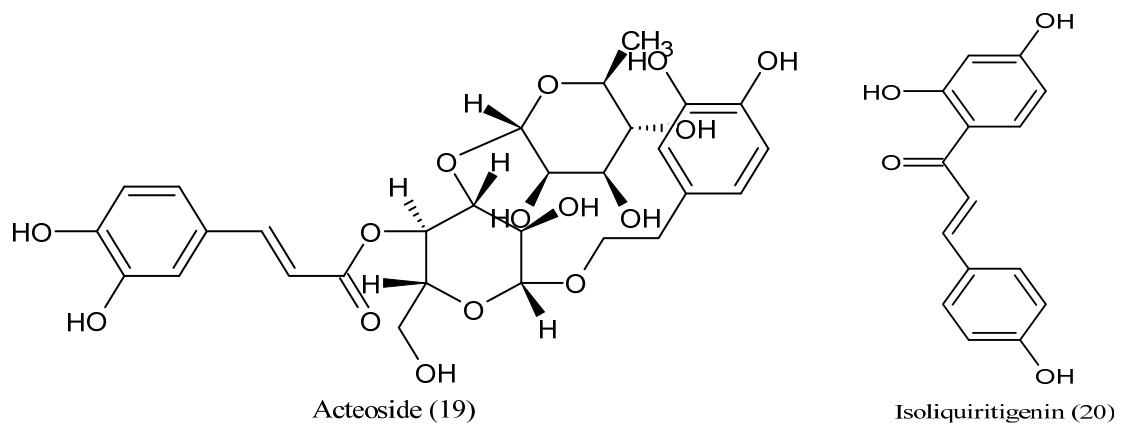
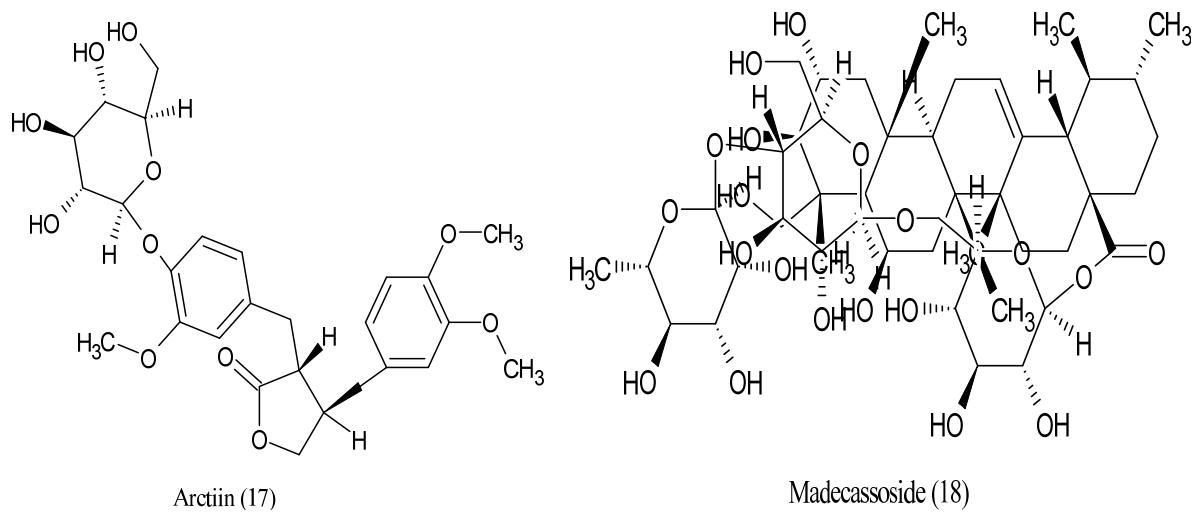
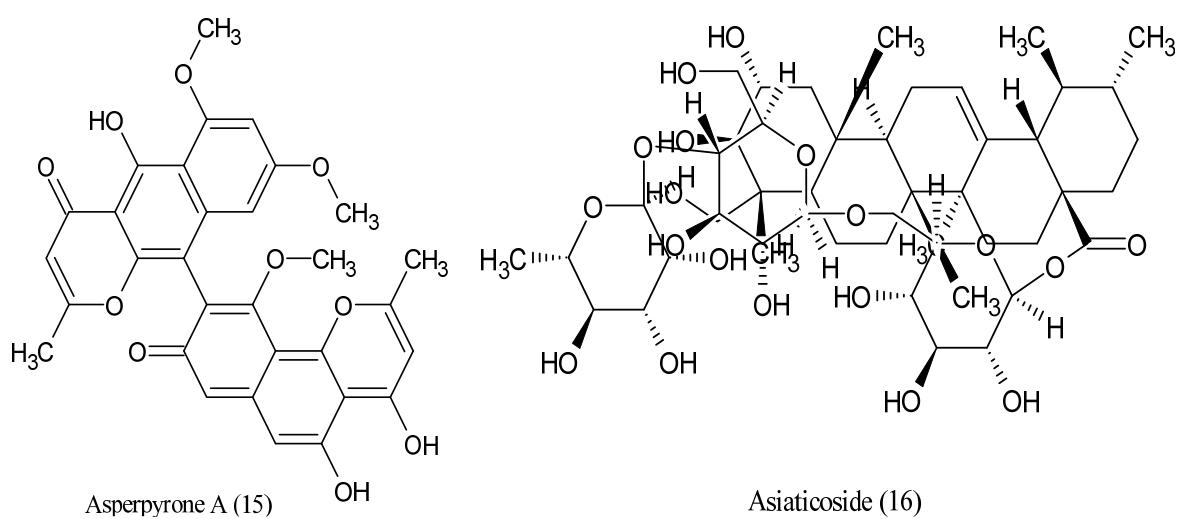
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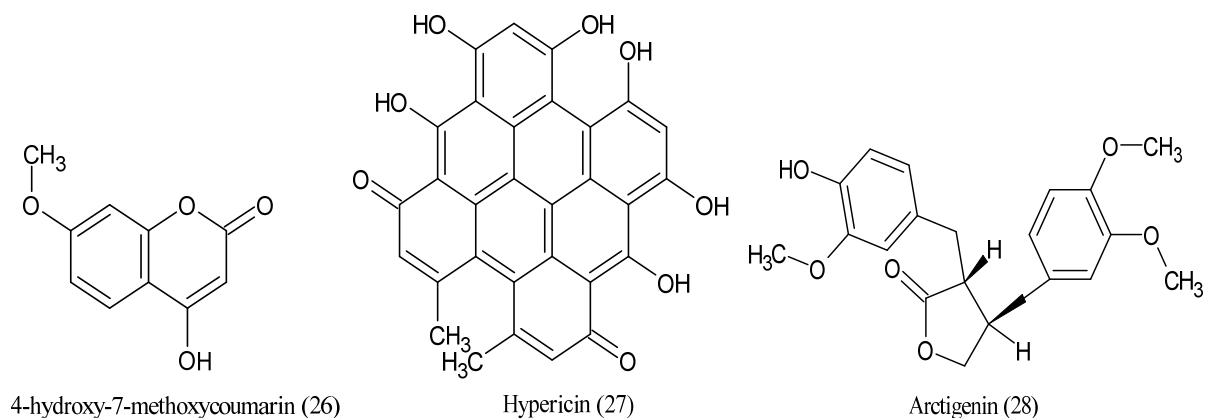
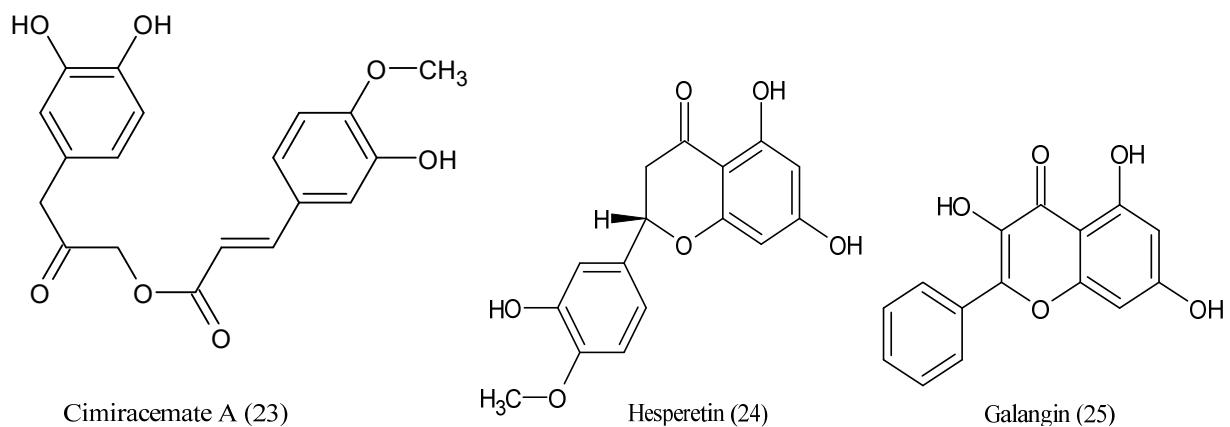
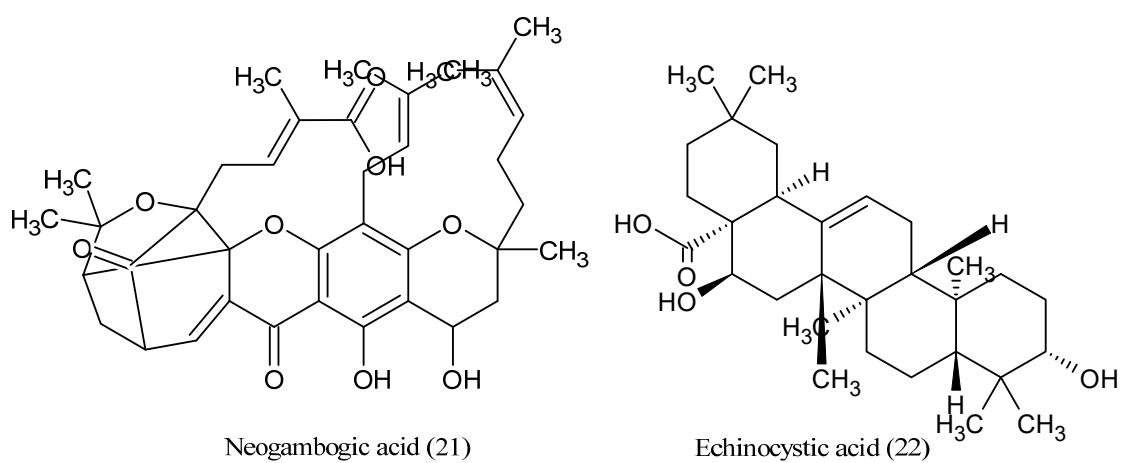


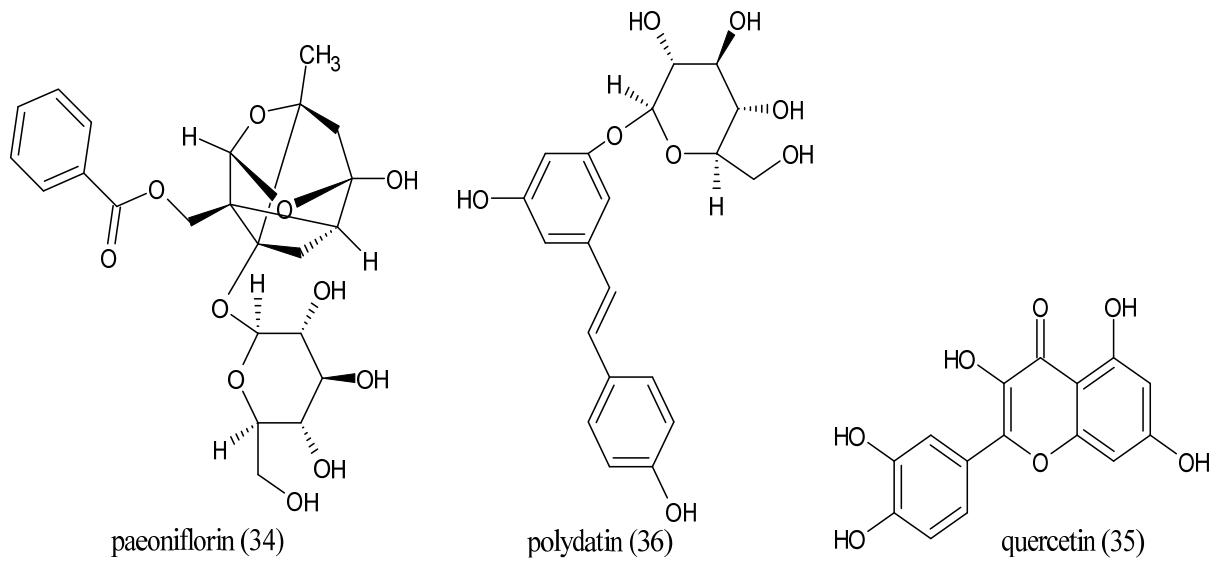
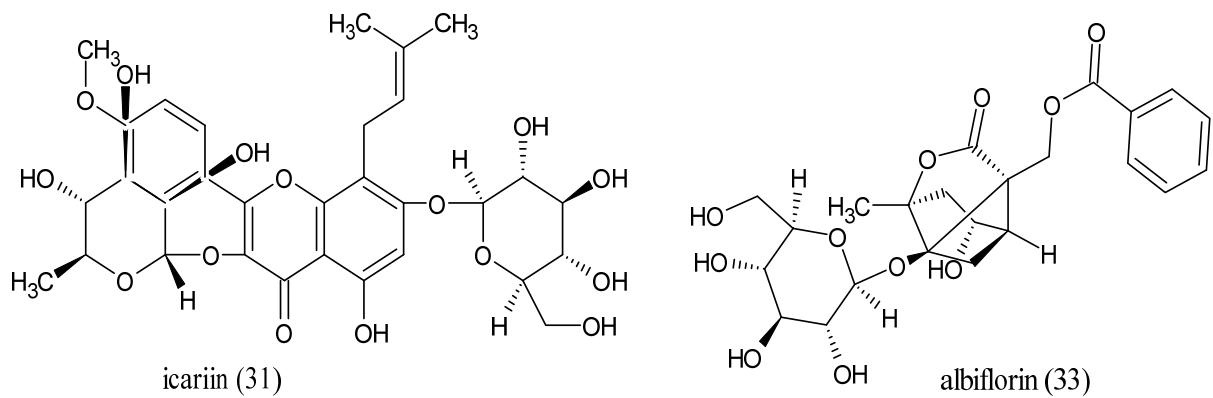
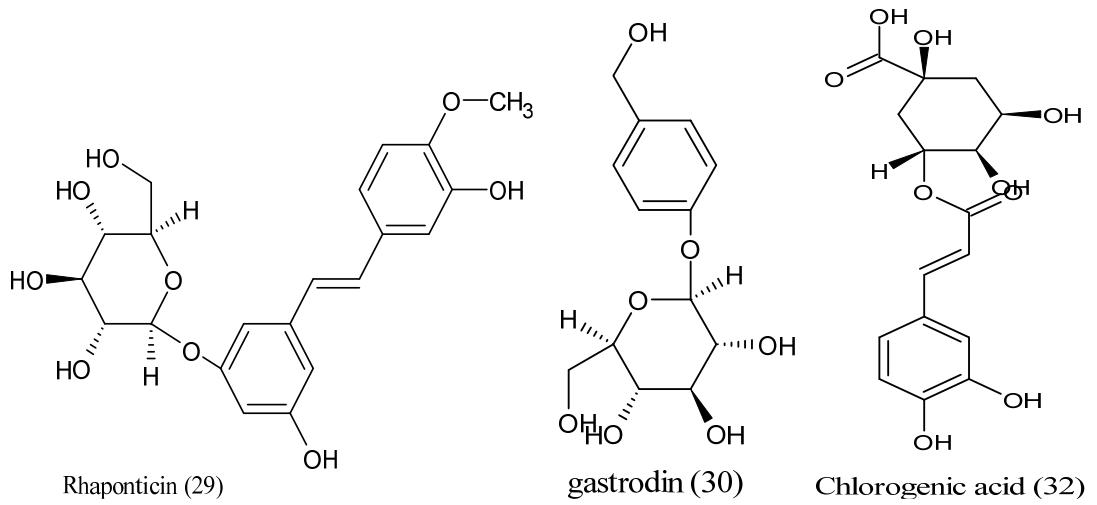
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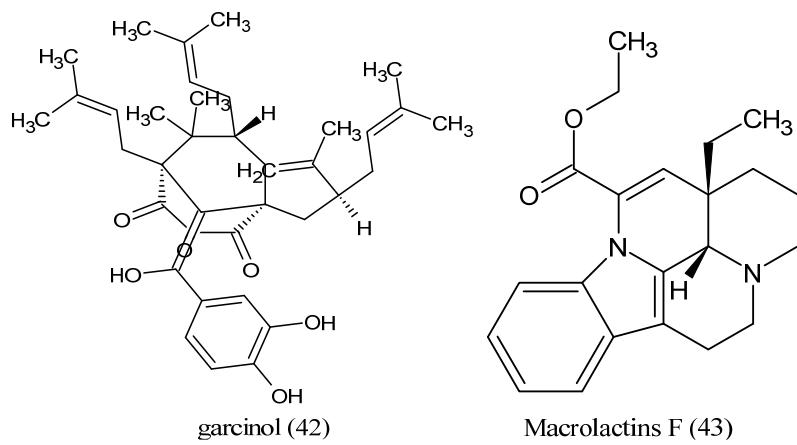
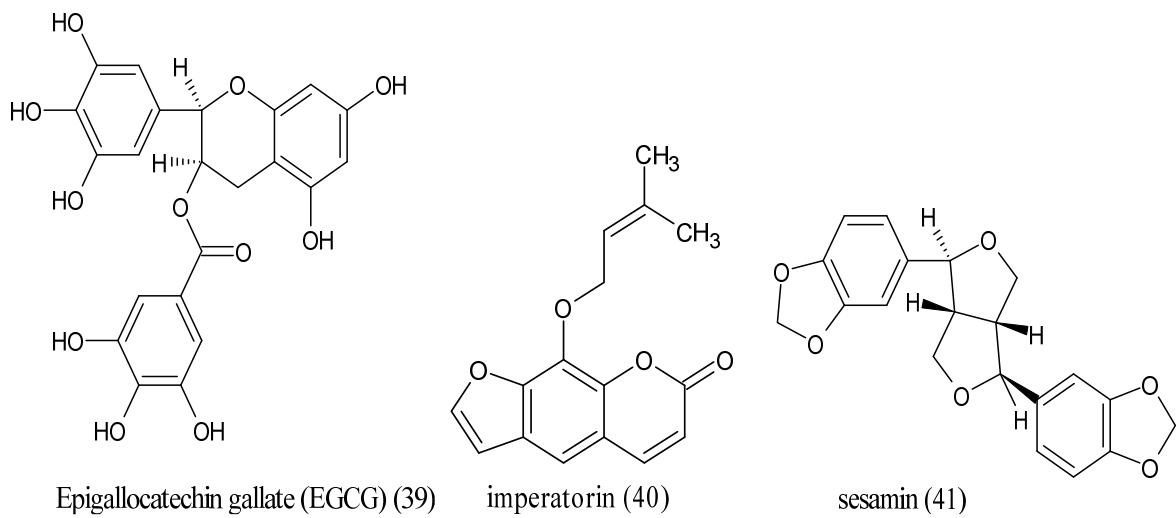
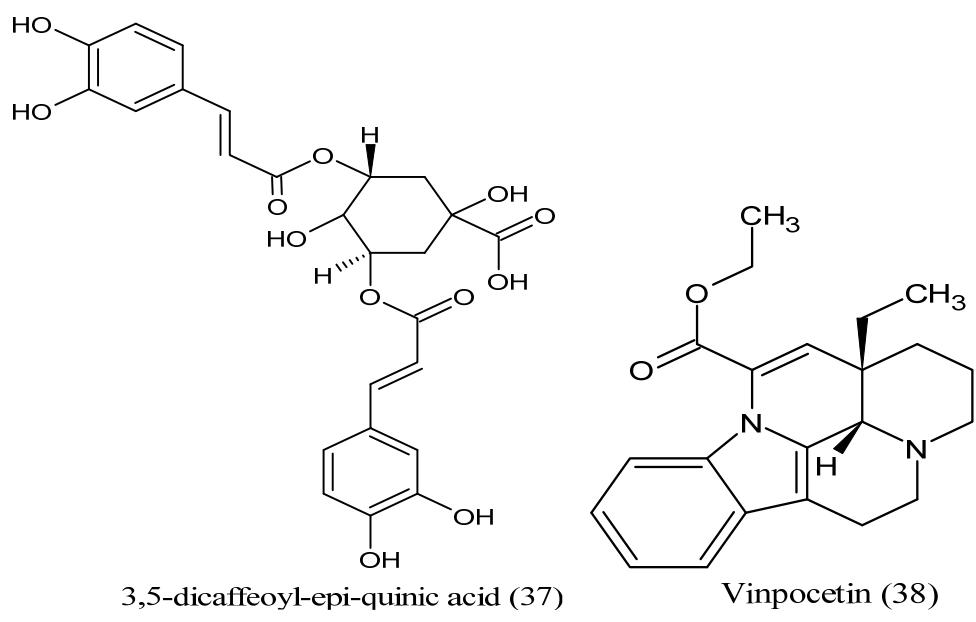
Kavain (12)

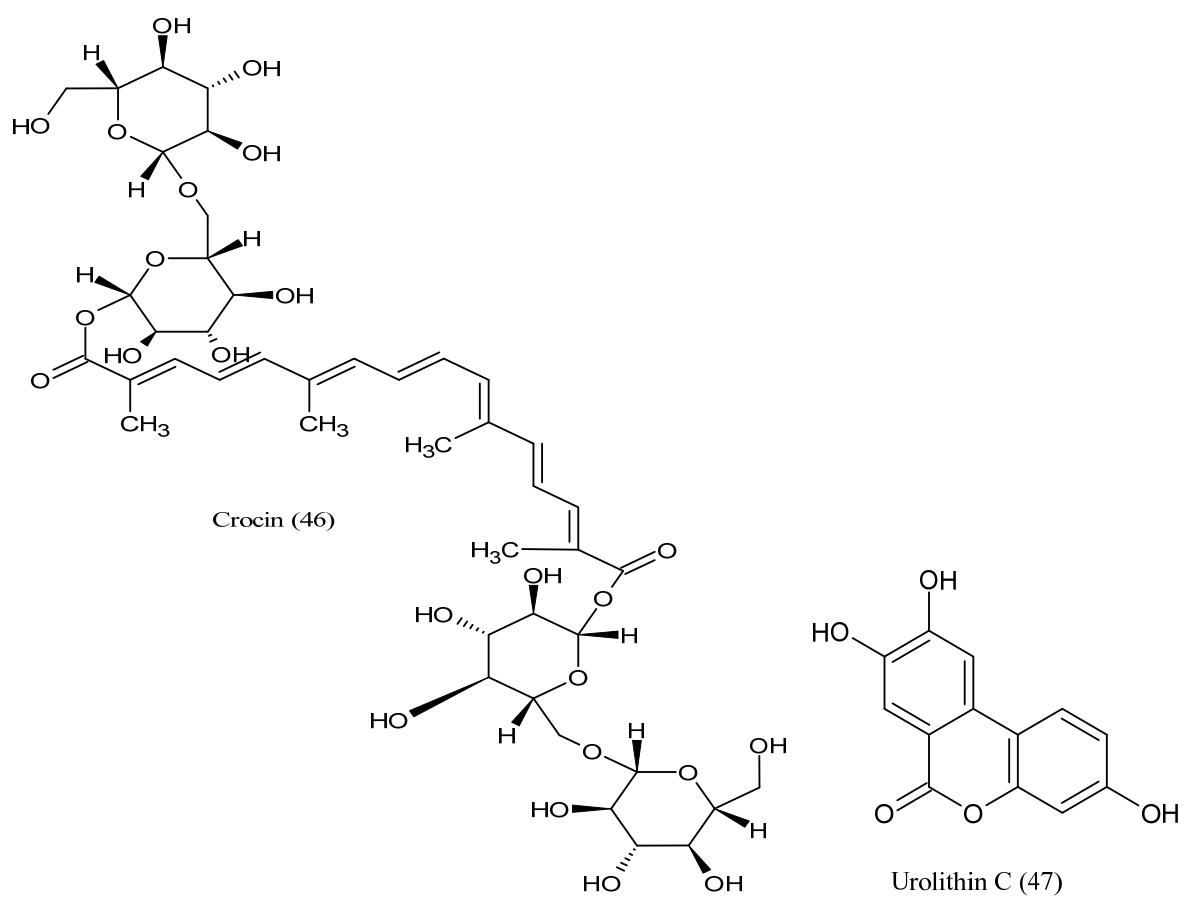
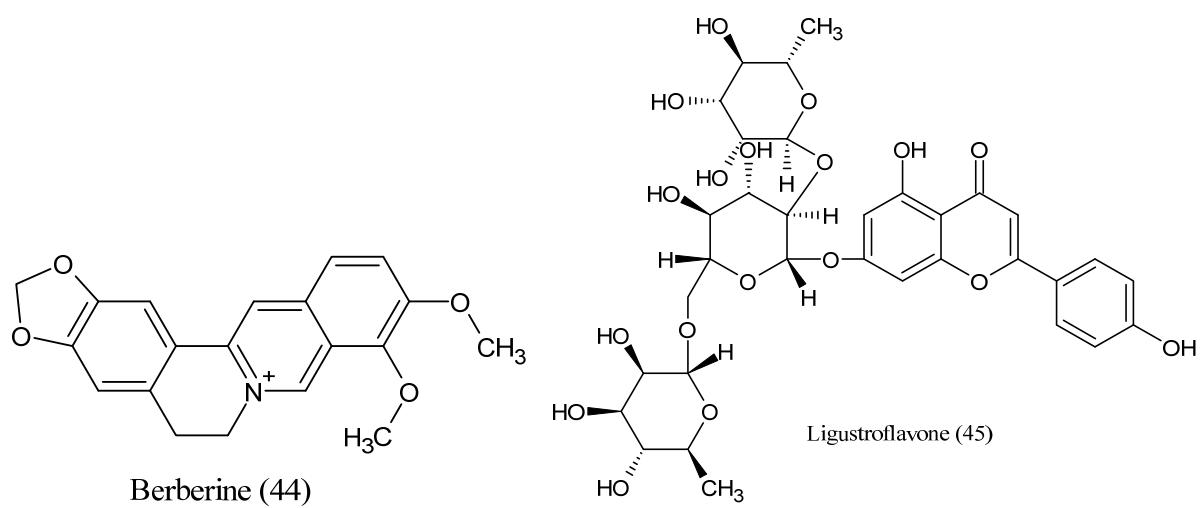












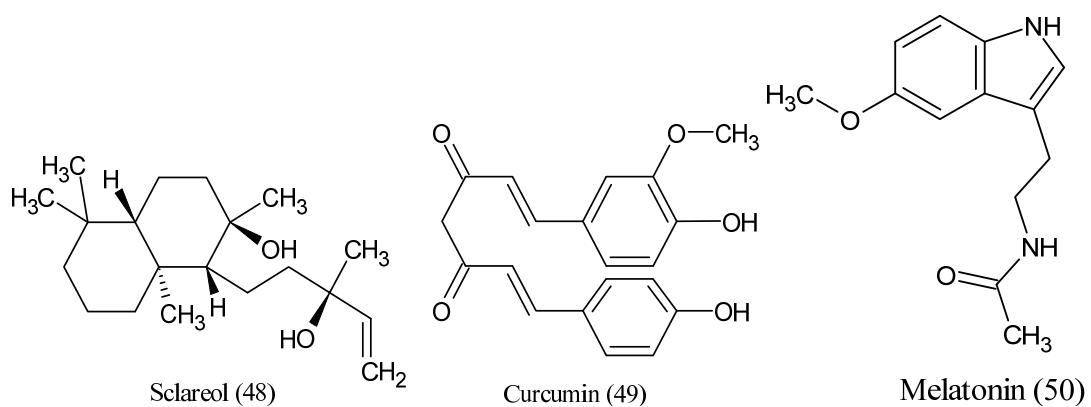


Figure S1: Structures of selected natural products with osteoprotective properties

Abbreviations

AGE/RAGE (advanced glycated end-products/receptor)

Akt (protein kinase-B)

ALP (alkaline phosphatase)

AMPK (AMP-dependent kinase)

AP-1 =(activator protein 1)

Apc (adenomatosis polyposis coli)

Apo (apolipoprotein)

ARE (antioxidant response element)

ATP6v0d2 (ATPase H⁺ transporting V0 subunit d2)

BAX (Bcl-2-associated X protein)

BDs (bone diseases)

BMD (bone mass density)

BMMs (primary murine-derived bone marrow macrophages)

BMP-2 (bone morphogenetic protein-2)

BSAP (bone-specific alkaline phosphatase)

BSP (bone sialoprotein)

BV/TV (bone volume/total volume)

BVD (bone volume density)

CaMKII (Ca2+/calmodulin (CaM)-dependent protein kinases)

CaSR (Calcium sensing receptors)

CAT (catalase)

Cbfa1(Core-binding factor alpha 1)

c-Fms (Colony-stimulating factor-1 receptor)

c-Src (proto-oncogene tyrosine-protein kinase)

CTR (calcitonin receptor)

CTSK (cathepsin K)

CTX (carbo-terminal telopeptide of type I collagen)

cyt c (cytochrome C)

DEX (dexamethasone)

DMP-1 (dentin matrix acidic phosphoprotein 1)

EGCG (Epigallocatechin gallate)

ERK (extracellular signal-regulated kinase)

FasL (Fas ligand)

FOXP3+ (forkhead box P3+)

GCS (γ -glutamylcysteine synthetase)

GSK3 β (glycogen synthase kinase-3 β)

GSR (glutathione S- reductase)

hAOBs (human alveolar osteoblaststs)

HDL (high-density lipoproteins)

HIF-1 (hypoxia-inducible factor 1)

HMGB1 (High mobility group box 1)

hPDLCs (human periodontal ligament cells)

i.p (intraperitoneal injection)

IFN- γ (interferon gamma)
IGF (insulin-like growth factor)
IL-1 (interleukin 1)
IL-17 (interleukin 17)
IL-1b/6 (interleukin 1 beta)
IL-6 (interleukin 6)
I κ B α (kappa light polypeptide gene enhancer in B-cells inhibitor)
JNK (c-Jun N-terminal kinase)
Keap-1 (Kelch-like ECH-associated protein 1)
LDL (low-density lipoproteins)
MAPKs (mitogen-activated protein kinase)
MCPs (MC3T3-E1 preosteoblasts)
M-CSF (monocyte/macrophage colony stimulating factor)
MMCs (mouse macrophage cells)
MMP-9 (matrix metalloproteinase-9)
N.Oc (OC number per bone surface)
NFATc1 (nuclear factor of activated T Cells 1)
NF-kb (nuclear factor kappa B)
NOX-1/2 (NADPH oxidase 1/2)
NP (number of porosity)
NPs (Natural products)
NQO1 (NAD(P)H:quinone reductase)
NR1D (nuclear receptor subfamily 1, group D, members)
NR1D1 (nuclear receptor subfamily-1 group D member-1)
Nrf2 (nuclear factor erythroid 2-related factor 2)
OA (osteoarthritis)
Oc.S/BS (osteoclast surface per bone surface)

OCG (osteoclastogenesis)
OCN (osteocalcin)
OP (osteoporosis)
OPG (osteoprotegerin)
OSCAR (osteoclast-associated receptor)
OVX (ovariectomized)
p.o (per oral injection)
P1NP (N-terminal propeptide of type I collagen)
p38 MAPK (mitogen-activated protein kinase)
PGC-1 α (peroxisome proliferator-activated receptor gamma coactivator 1-alpha)
PGE2 (prostaglandin E2)
PI3K (phosphoinositide 3-kinase)
PP (percentage porosity)
PTH (parathyroid hormone)
PUVA (Psoralen plus ultra violet radiation)
RA (Rheumatoid arthritis)
RANKL (receptor activation of NF- κ B ligand)
ROS (reactive oxygen species)
SASP (senescence-associated secretory phenotype)
Sc (subcutaneous injection)
SERMs (selective estrogen receptor modulators)
SIM (structure model index)
sirt-1 (sirtuin-1)
SMAD (SMA (small worm phenotype); MAD (mothers against decapentaplegic))
SOST (sclerostin)
TAG (triacylglycerol)
TC (total cholesterol)

Th17 (T helper 17)

TN (trabecular number)

TNF- α (tumor necrosis factor-alpha)

TRAC (T Cell Receptor Alpha Constant)

TRAF6 (tumor necrosis factor (TNF) receptor-associated factor-6)

TRAF6TNF (receptor-associated factor-6)

TRAP (tartrate-resistant acid phosphatase)

TS (trabecular separation/space)

TT (trabecular thickness)

UVA (ultra violet radiation)

VDR (vitamin D receptor)

VEGF-A (Vascular endothelial growth factor alpha)

γ -EV (γ -Glutamyl valine)