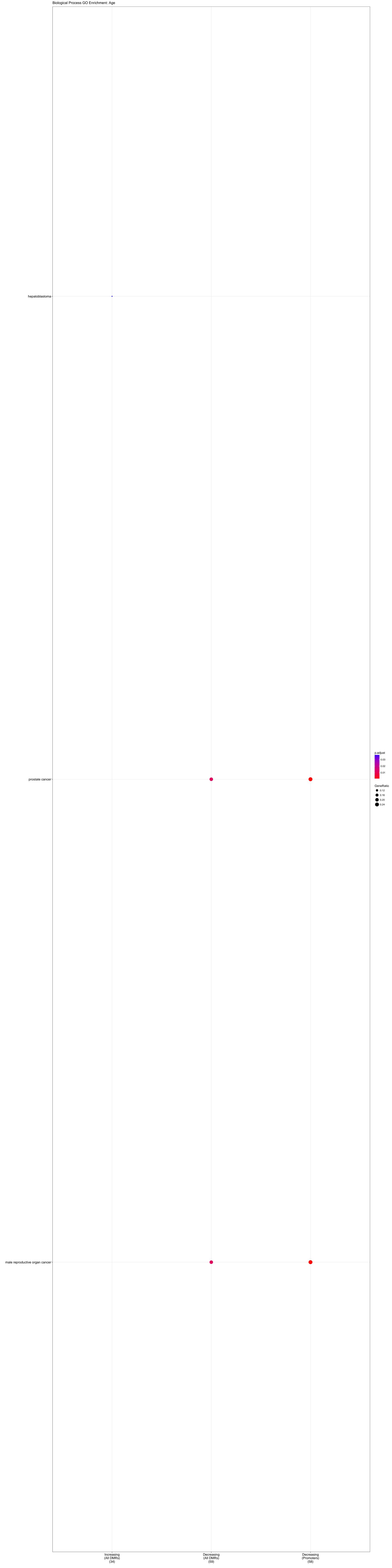
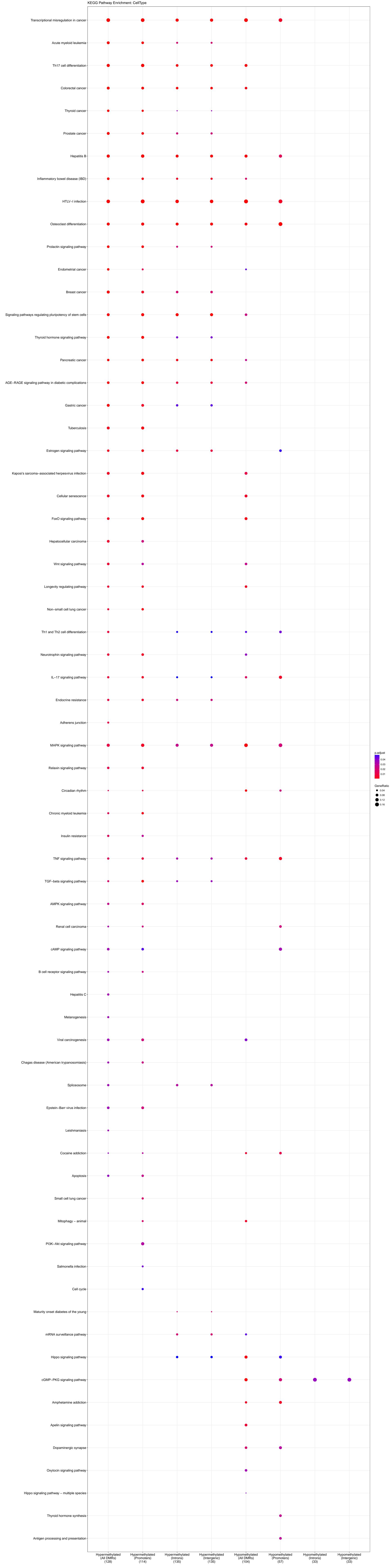
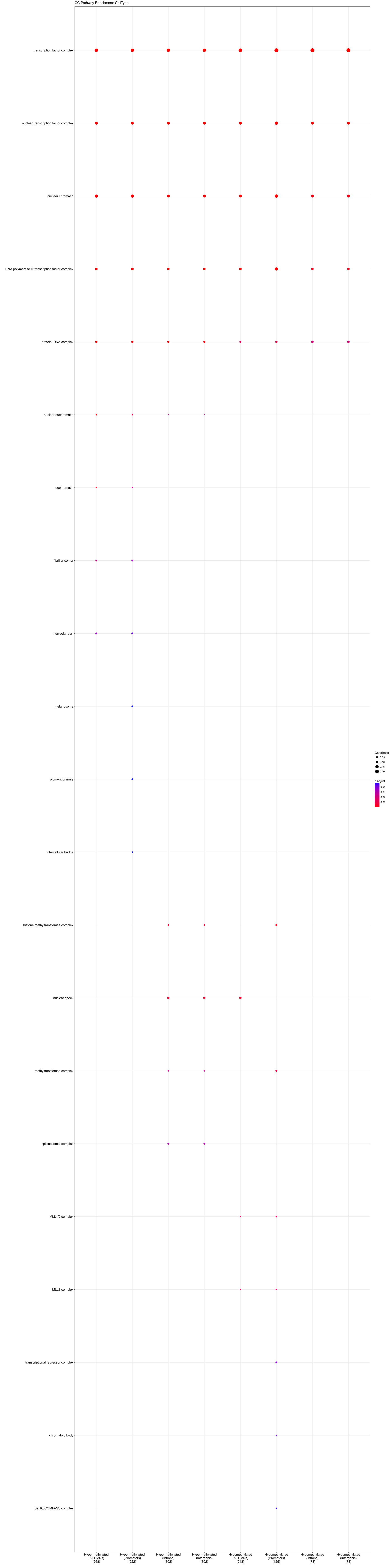


Biological Process GO Enrichment: Age myeloid leukocyte differentiation cellular response to extracellular stimulus regulation of myeloid leukocyte differentiation cellular response to external stimulus positive regulation of neuron death positive regulation of transcription from RNA polymerase II promoter in response to endoplasmic reticulum stress positive regulation of ossification cellular response to metal ionresponse to extracellular stimulus regulation of leukocyte differentiation positive regulation of neuron apoptotic process leukocyte differentiation cellular response to inorganic substance response to endoplasmic reticulum stress cellular response to nutrient levels myeloid cell differentiation glucose metabolic process intrinsic apoptotic signaling pathway in response to endoplasmic reticulum stress positive regulation of transcription from RNA polymerase II promoter in response to stress response to lipopolysaccharide liver development skeletal muscle cell differentiation carbohydrate homeostasis glucose homeostasis cellular response to starvationhepaticobiliary system development response to metal ion response to molecule of bacterial origin positive regulation of transcription from RNA polymerase II promoter involved in cellular response to chemical stimulus hexose metabolic process regulation of myeloid cell differentiation gluconeogenesis skeletal muscle tissue development hexose biosynthetic process monocyte differentiation pri-miRNA transcription from RNA polymerase II promoter mononuclear cell differentiation skeletal muscle organ development monosaccharide biosynthetic process response to starvation cellular response to glucose starvation monosaccharide metabolic process response to cAMP regulation of neuron death anion homeostasis regulation of ossification dicarboxylic acid metabolic process response to drug-ER-nucleus signaling pathway response to light stimulus circadian rhythm regulation of neuron apoptotic process response to nutrient levels negative regulation of myeloid leukocyte differentiation response to gravity regulation of transcription from RNA polymerase II promoter in response to stress response to radiation rhythmic process neuron death lipid homeostasis regulation of hemopoiesis fat cell differentiation regulation of DNA-templated transcription in response to stress regulation of fat cell differentiation oxaloacetate metabolic process transcription from RNA polymerase III promoter multi-multicellular organism process neuron apoptotic process circadian regulation of gene expression fatty acid homeostasis eyelid development in camera-type eyeendoplasmic reticulum unfolded protein response cellular response to calcium ionresponse to organophosphorus positive regulation of fat cell differentiation positive regulation of leukocyte differentiation dicarboxylic acid biosynthetic process positive regulation of osteoblast differentiation cellular response to unfolded proteincellular response to arsenic-containing substance positive regulation of exosomal secretion regulation of osteoclast differentiation striated muscle tissue developmentregulation of exosomal secretion response to purine-containing compound response to glucocorticoidmorphogenesis of embryonic epithelium negative regulation of myotube differentiation hepatocyte proliferationepithelial cell proliferation involved in liver morphogenesis muscle tissue development regulation of monocyte differentiation response to corticosterone liver morphogenesis muscle organ development cellular response to topologically incorrect proteinkidney development response to steroid hormone response to muscle stretch exosomal secretion regulation of protein catabolic process response to corticosteroidextracellular exosome biogenesis extracellular vesicle biogenesis positive regulation of pri-miRNA transcription from RNA polymerase II promoter renal system development-PERK-mediated unfolded protein response positive regulation of hemopoiesis cellular response to oxidative stress positive regulation of epithelial cell proliferation aspartate family amino acid biosynthetic process retina layer formation positive regulation of interleukin-4 production negative regulation of translational initiation response to unfolded protein response to oxidative stress response to lithium ionglandular epithelial cell development osteoclast differentiation negative regulation of myeloid cell differentiation negative regulation of neuron death negative regulation of leukocyte differentiation in utero embryonic developmentregulation of transcription from RNA polymerase III promoter regulation of interleukin-4 production positive regulation of DNA-templated transcription, initiation urogenital system developmentprimary neural tube formation response to topologically incorrect proteincarbohydrate biosynthetic process response to mineralocorticoid response to arsenic-containing substance regulation of pri-miRNA transcription from RNA polymerase II promoter p.adjust granulocyte differentiation 0.04 negative regulation of exocytosis 0.03 neural tube formation 0.02 cellular response to cadmium ion 0.01 positive regulation of viral process response to mechanical stimulus GeneRatio eye development-0.08 gland development 0.12 regulation of DNA-templated transcription, initiation 0.16 DNA-templated transcription, initiation endocrine system development mammary gland epithelial cell proliferation epithelial cell proliferation mammary gland epithelium development regulation of DNA binding transcription factor activity DNA-templated transcriptional preinitiation complex assembly regulation of RNA polymerase II transcriptional preinitiation complex assembly mammary gland development reproductive structure development reproductive system development regulation of fibroblast proliferation fibroblast proliferation negative regulation of phosphorylation type B pancreatic cell development columnar/cuboidal epithelial cell development ovarian follicle development lymphocyte differentiation negative regulation of protein phosphorylation cell fate commitment type B pancreatic cell differentiation response to axon injury transcription initiation from RNA polymerase II promoter enteroendocrine cell differentiation regulation of transcription initiation from RNA polymerase II promoter positive regulation of cysteine-type endopeptidase activity involved in apoptotic process steroid hormone mediated signaling pathway protein-DNA complex subunit organization positive regulation of cysteine-type endopeptidase activity T cell activation RNA polymerase II transcriptional preinitiation complex assembly regulation of intracellular steroid hormone receptor signaling pathway response to hydrogen peroxide intracellular steroid hormone receptor signaling pathway negative regulation of DNA binding transcription factor activity regulation of protein complex assembly activation of cysteine-type endopeptidase activity involved in apoptotic process gonad development extrinsic apoptotic signaling pathway via death domain receptors placenta development ovulation cycle process development of primary sexual characteristics positive regulation of cell cycle arrest hormone-mediated signaling pathway homeostasis of number of cells positive regulation of endopeptidase activity ATF6-mediated unfolded protein response negative regulation of fibrinolysis vagina development embryonic organ development protein-DNA complex assembly positive regulation of RNA polymerase II transcriptional preinitiation complex assembly positive regulation of DNA binding transcription factor activity female gonad development endocrine pancreas development mammary gland morphogenesis cell cycle arrest regulation of circadian rhythm intrinsic apoptotic signaling pathway in response to oxidative stress mRNA cleavage involved in mRNA processing regulation of IRE1-mediated unfolded protein response glandular epithelial cell differentiation negative regulation of fat cell differentiation development of primary female sexual characteristics positive regulation of peptidase activity modification of morphology or physiology of other organism involved in symbiotic interaction cellular response to steroid hormone stimulus positive regulation of apoptotic signaling pathway regulation of histone phosphorylation cellular response to lithium ion activation of cysteine-type endopeptidase activity involved in apoptotic signaling pathway columnar/cuboidal epithelial cell differentiation ovulation cyclepositive regulation of myeloid leukocyte differentiation digestive tract morphogenesis regulation of lipid metabolic process protein folding in endoplasmic reticulumregulation of fibrinolysis sex differentiation positive regulation of muscle cell differentiation regulation of striated muscle cell differentiation alternative mRNA splicing, via spliceosome regulation of osteoblast differentiation regulation of cell cycle arrest ossification · positive regulation of cell cycle positive regulation of nuclease activity glucocorticoid receptor signaling pathway response to gamma radiation positive regulation of fibroblast proliferation female sex differentiation B cell differentiation release of cytochrome c from mitochondria interleukin-12-mediated signaling pathway response to cadmium ion cellular response to interleukin-12 positive regulation of intrinsic apoptotic signaling pathway epithelial cell development gland morphogenesis response to calcium ionregulation of muscle cell differentiation intracellular receptor signaling pathway response to interleukin-12 osteoblast development corticosteroid receptor signaling pathway mammary gland epithelial cell differentiation positive regulation of transforming growth factor beta production intrinsic apoptotic signaling pathway positive regulation of proteolysis female genitalia development mammary gland alveolus development mammary gland lobule developmentpositive regulation of cysteine-type endopeptidase activity involved in apoptotic signaling pathway osteoblast differentiation regulation of cysteine-type endopeptidase activity involved in apoptotic process regulation of myotube differentiation positive regulation of epithelial cell differentiation entrainment of circadian clock by photoperiod regulation of histone modification regulation of mitochondrial membrane potential modification of morphology or physiology of other organism modulation by virus of host morphology or physiology response to acid chemical prostate glandular acinus development modification by symbiont of host morphology or physiology branching involved in prostate gland morphogenesis response to amino acid interaction with host muscle cell differentiation signal transduction involved in regulation of gene expression muscle cell cellular homeostasis positive regulation of transcription initiation from RNA polymerase II promoter Increasing (All DMRs) (56) Decreasing (All DMRs) (85) Decreasing (Promoters) (87)



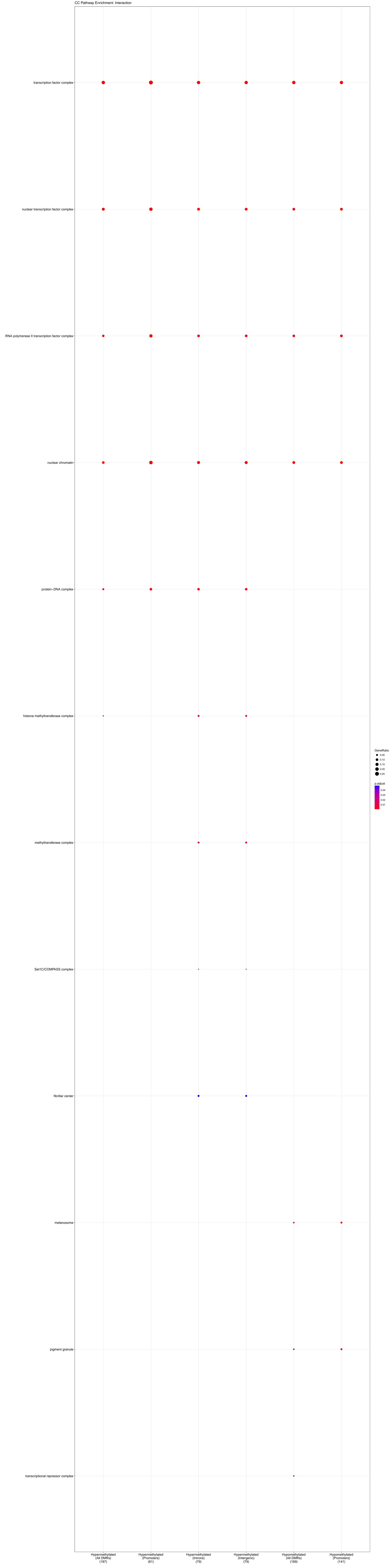


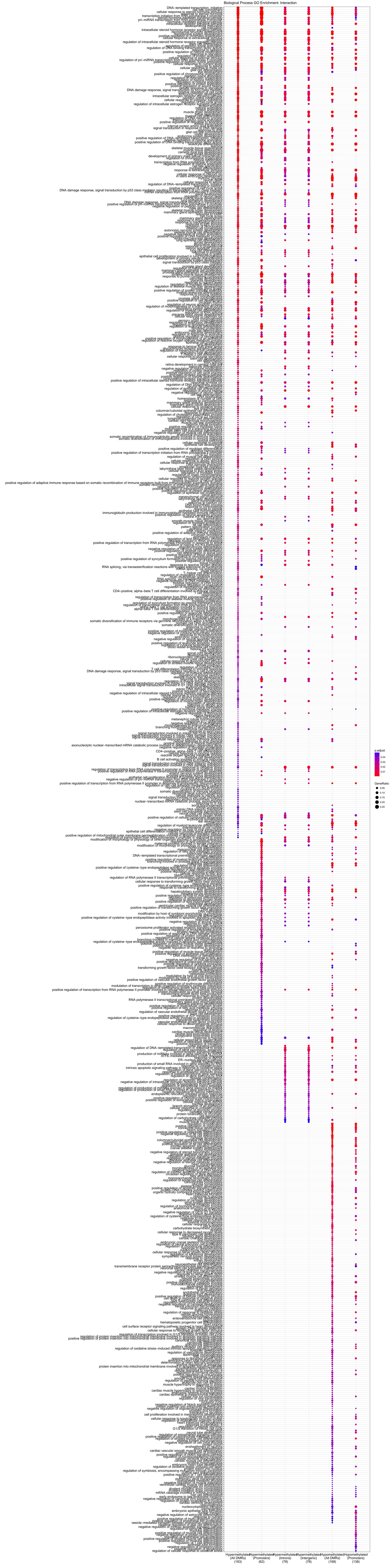




| musculoskeletal system cancer- | Biological Process GO Enrichm | nent: CellType | | | | | |
|---|-------------------------------|--------------------------------|---------------------------|--|------------------------------|---------------------------------------|----------------------------------|
| hepatitis B - connective tissue cancer - | | | | | | | |
| gastric adenocarcinoma - obesity - | | | | | | | |
| overnutrition - cell type benign neoplasm - | | | | | | | |
| nutrition disease - female reproductive organ cancer- | | | | | | | |
| adenoma - non-small cell lung carcinoma - breast adenocarcinoma - | | | | | | | |
| kidney cancer- autosomal dominant disease- | | | | | | | |
| reproductive organ benign neoplasm - progressive multifocal leukoencephalopathy - | | | | | | | |
| mammary Paget's disease - | | | | | | | |
| stomach carcinoma - | | | | | | | |
| malignant ovarian surface epithelial-stromal neoplasm ovary epithelial cancer- | | | | | | | |
| ovarian carcinoma - urinary system cancer - | | | | | | | |
| physical disorder - lymphoblastic leukemia - breast carcinoma - | | | | | | | |
| retinoblastoma - | | | | | | | |
| urinary system disease - | | | | | | | |
| placenta cancer- choriocarcinoma- | | | | | | | |
| nephroblastoma - stomach cancer - | | | | | | | |
| prostate cancer- uterine benign neoplasm- | | | | | | | |
| uterine fibroid- kidney disease- | | | | | | | |
| hepatitis - ovarian cancer - renal cell carcinoma - | | | | | | | |
| male reproductive organ cancer- cervix carcinoma- | | | | | | | |
| female reproductive organ benign neoplasm - cervical cancer- | | | | | | | |
| hair disease - muscular atrophy - | | | | | | | |
| peripheral nervous system neoplasm - sensory system cancer - | | | | | | | |
| ocular cancer- alopecia- | | | | | | | |
| leiomyoma - alveolar rhabdomyosarcoma - clear cell sarcoma - | | | | | | | |
| gallbladder disease - | | | | | | | |
| renal carcinoma - prostate carcinoma - | | | | | | | |
| hypospadias - | | | | | | | |
| familial hyperlipidemia - thoracic disease - | | | | | | | |
| breast disease - | | | | | | | |
| lipomatous cancer- bone giant cell tumor- | | | | | | | |
| neurofibroma - bone cancer - muscle cancer - | | | | | | | GeneRatio ● 0.05 |
| chondrosarcoma - | | | | | | | ● 0.10 ● 0.15 p.adjust |
| neuroblastoma - atherosclerosis - | | | | | | | 0.03 0.02 0.01 |
| arteriosclerotic cardiovascular disease - congenital nervous system abnormality - | | | | | | | |
| colon carcinoma - cholelithiasis - | | | | | | | |
| lipid metabolism disorder - arteriosclerosis - | | | | | | | |
| esophageal cancer- organ system benign neoplasm- | | | | | | | |
| head and neck squamous cell carcinoma- head and neck carcinoma- osteoporosis- | | | | | | | |
| hyperthyroidism - | | | | | | | |
| inherited metabolic disorder- bone resorption disease- | | | | | | | |
| biliary tract cancer- | | | | | | | |
| skin cancer- head and neck cancer- | | | | | | | |
| neuroectodermal tumor- osteosarcoma- | | | | | | | |
| giant cell tumor- lipid storage disease- Parkinson's disease- | | | | | | | |
| Parkinson's disease - osteoarthritis - rhabdomyosarcoma - | | | | | | | |
| mouth disease - | | | | | | | |
| germ cell and embryonal cancer- embryonal rhabdomyosarcoma- | | | | | | | |
| colorectal cancer- | | | | | | | |
| endocrine pancreas disease - skeletal muscle cancer - | | | | | | | |
| pleomorphic adenoma - gestational diabetes - hyperglycemia - | | | | | | | |
| cervical adenocarcinoma - | | | | | | | |
| bile duct cancer- | | | | | | | |
| bile duct disease - salivary gland adenoid cystic carcinoma - | | | | | | | |
| cholestasis - multiple myeloma - | | | | | | | |
| endocrine system disease - biliary tract disease - | | | | | | | |
| germ cell cancer- | | | | | | | |
| familial combined hyperlipidemia - osteonecrosis - cervical squamous cell carcinoma - | | | | | | | |
| cervical squamous cell carcinoma- thyroid carcinoma- embryoma- | | | | | | | |
| myeloma - ovarian disease - | | | | | | | |
| bone marrow cancer- esophageal carcinoma | | | | | | | |
| brain cancer- embryonal cancer- | | | | | | | |
| premature ovarian failure - ischemic bone disease - | | | | | | | |
| seminoma - peripheral nerve sheath neoplasm - sarcoma - | | | | | | | |
| sarcoma- telangiectasis- bipolar disorder- | | | | | | | |
| pilocytic astrocytoma | | Hypermethylated (Promoters) | Hypermethylated (Introns) | Hypermethylated (Intergenic) (182) | Hypomethylated (All DMRs) | Hypomethylated (Promoters) (84) | |
| | אואום וורץ) (169) | (Promoters) (143) | (Introns) (182) | (182) | (All DMRs) (151) | (84) | |







| Biol hereditary breast ovarian cancer | logical Process GO Enrichme | ent: Interaction | | | | | |
|---|--|--|--------------------------------------|---|---------------------------------------|---------------------------------------|-----------|
| musculoskeletal system cancer-seminoma | | | | | | | |
| congenital nervous system abnormality | | | | | | | |
| gastric adenocarcinoma | | | • | | | | |
| stomach cancer physical disorder | | | | | | | |
| autosomal dominant disease - gallbladder disease - | | | | | | | |
| malignant peripheral nerve sheath tumor- breast carcinoma | | | | | | | |
| organ system benign neoplasm | | | | | | | |
| stomach carcinoma uterine benign neoplasm | | | | | | | |
| uterine fibroid hepatitis B | | | | | | | |
| thoracic disease | | | | | | | |
| female reproductive organ benign neoplasm connective tissue cancer | | | | | | | |
| neurofibroma - | | | | | | | |
| cell type benign neoplasm | | | | | | | |
| peripheral nerve sheath neoplasm | | | | | | | |
| hereditary Wilms' tumor | | | | | | | |
| leiomyoma - | | | | | | | |
| infertility | | | | | | | |
| megacolon - germ cell and embryonal cancer - | | | | | | | |
| kidney cancer | | | | | | | |
| pleomorphic adenoma | | | | | | | |
| neuromuscular junction disease urinary system cancer | | | | | | | |
| lung adenocarcinoma biliary tract cancer | | | | | | | |
| cervical adenocarcinoma bile duct cancer | | | | | | | |
| bile duct carcinoma bilateral breast cancer | | | | | | | |
| esophageal cancer | | | | | | | |
| hypospadias | | | | | | | |
| dementia | | | | | | | |
| gastrointestinal system benign neoplasm- retinoblastoma | | | | | | | |
| retinal cell cancer | | | | | | | |
| non-small cell lung carcinoma | | | | | | | |
| malignant ovarian surface epithelial-stromal neoplasm ovary epithelial cancer | | | | | | | |
| ovarian carcinoma | | | | | | | p.adjust |
| alopecia | | | | | | | GeneRatio |
| osteonecrosis - bile duct disease - | | | | | | | |
| renal cell carcinoma | | | | | | | |
| lymphoblastic leukemia | | | | | | | |
| biliary tract disease | | | | | | | |
| autoimmune disease of the nervous system- | | | | | | | |
| peripheral nervous system neoplasm | | | | | | | |
| urinary system disease | | | | | | | |
| prostate cancer-male reproductive organ cancer- | | | | | | | |
| female reproductive organ cancer ovarian cancer | | | | | | | |
| esophageal carcinoma | | | | | | | |
| nephritis - | | | | | | | |
| clear cell sarcoma | | | | | | | |
| lipomatous cancer | | | | | | | |
| muscular atrophy - embryoma - | | | | | | | |
| cryptorchidism - osteoarthritis - | | | | | | | |
| follicular thyroid carcinoma cerebrovascular disease | | | | | | | |
| breast lobular carcinoma | | | | | | | |
| invasive lobular carcinoma | | | | | | | |
| hypotrichosis - myopathy - | | | | | | | |
| muscle tissue disease | | | | | | | |
| intrinsic cardiomyopathy | | | | | | | |
| muscular disease | | | | | | | |
| germ cell cancer- obesity- | | | | | | | |
| esophagus adenocarcinoma | | | | | | | |
| overnutrition - | | | | | | | |
| cardiomyopathy | | | | | | | |
| familial combined hyperlipidemia - glomerulonephritis - | | | | | | | |
| pancreatic carcinoma | | | | | | | |
| familial hyperlipidemia | | | | | | | |
| cholestasis - | | | | | | | |
| autonomic nervous system neoplasm | | | | | | | |
| progressive multifocal leukoencephalopathy | Hypermethylated (All DMRs) (107) | Hypermethylated (Promoters) (44) | Hypermethylated (Introns) (52) | Hypermethylated (Intergenic) (52) | Hypomethylated (All DMRs) (118) | Hypomethylated (Promoters) (94) | |
| | (107) | (44) | (52) | (32) | (118) | (94) | |