PROSTATE CANCER Prostatic epithelial cell Normal prostate epithelium Genetic alterations Proliferative inflammatory atrophy Hypermethylation — ►O-AR Oncogenes: DNA Tumor suppressors: CDKN1B, NKX3.1, PTEN GSTP1 0 ►O——►Genomic damage Carcinogens DNA Cell cycle PI3K-Akt signaling pathway Apoptosis Prostatic intraepithelial neoplasia K04399 DNA K09348 Apoptosis inhibition K17454... ►O——► G1/S progression K02158 K07201 Localized prostate K06625 cancer p53 signaling pathway Cell cycle progression Cell cycle K06624 Impaired G1 and G2 arrest Reduced apoptosis Genomic instability PIP₃ DNA Cytokine-cytokine receptor interaction K04374... MAPK signaling pathway K02620... Cell proliferation DNA Testosterone Steroid hormone K12344 biosynthesis — 🖚 Cell survival DNA Tumor growth Dihydrotestosterone Metastatic prostate cancer K01351 K08557 → Cell proliferation Survival (Therapeutic decreased testosterone and dihydrotestosterone) K08557 K08557 Other target DNA Other ligands events Androgen-independent cancer 05215 1/26/16 (c) Kanehisa Laboratories