Attachment 4

Additional Information on Biospecimens

November 16, 2012

Analyte	Preferred Matrix	Biomarker	
Tobacco Biomarkers			
Nicotine and nicotine metabolites Tobacco user: cotinine and trans-3'- hydroxycotinine Non-tobacco user : cotinine	Serum	Exposure	
Nicotine and nicotine metabolites Tobacco user: cotinine, trans-3'-hydroxycotinine, cotinine N-oxide, nicotine N-oxide, nornicotine, norcotinine; analogues: anabasine, anatabine Non-tobacco user: cotinine and trans-3'- hydroxycotinine	Urine	Exposure	
Tobacco specific nitrosamines (TSNAs): NNAL, NNN, NNA, NAT, NAB, NNK	Urine	Exposure	
Polycyclic aromatic hydrocarbons (pyrene, naphthalene, phenanthrene, fluorene)	Urine	Combustion	
Cadmium, cobalt, uranium, lead, strontium, beryllium, manganese, and thallium	Urine, prescreened container	Toxicity	
Speciated arsenic (As III, As V, dimethylarsinic acid (DMA), and monomethylarsonic acid (MMA))	Urine, prescreened container	Toxicity	
Creatinine	Urine	For correction	
4-ABP hemoglobin	Red blood cells, EDTA	Toxicity	
VOCs metabolites	Urine	Combustion	
Volatile nitrosamines	Urine	Combustion	
Aromatic amines	Urine	Combustion	
Cyanide	Urine	Combustion	
Other Biomarkers			
C-reactive protein	Serum, red top or SST	Inflammation; Cardiovascular risk	
Fibrinogen	Plasma, citrate	Cardiovascular risk	
Interleukin 6	Plasma	Inflammation	
sICAM (soluble intercellular adhesion molecule)	Plasma, EDTA or Serum, SST	Cardiovascular risk	

PATH Biospecimen Analytes by Specimen and Biomarker

F2-isoprostane / 8-epi-prostaglandin F2a	Urine	Oxidative stress
MicroRNA profile	Plasma	Epigenetic effects
Metabolomic profile	Plasma, EDTA	Metabolic effects
Proteomic profile	Plasma	Toxicity, risk, stress
DNA (genotyping, sequencing)	Buffy coat	Role of genetics
RNA (gene expression)	PAXgene	Epigenetic effects
Metabolomic profile	Urine	Metabolic effects
microRNA, total RNA	Buccal cells	Epigenetic effects
Epigenetic marks	Buccal cells	Epigenetic effects