Most environmentalcauses of cancer are mutagens: mutagenic compounds, $X$-rays, uv


Cancer tends to arise in actively dividing cells

- Epithelialcells (lining of intestine, lungs etc.) = carcinoma
- Blood and lymphatic cells = lukemia, meyloma, lymphoma
- Connective tissue (bones, tendons muscle) = sarcoma

Cancer is agenetic disease of somatic cells

The underlying cause is mutations that release cells from the normal constraints that exist in well organized tissues allowing uncontrolled growth

Which are the keygenes that are mutated?

## Incidence of stomach cancer as a function of age



Figure by MIT OCW.

Major complications in understanding the genetic basis of cancer

- Multiple mutations are necessary to produce a tumor cell
- Different types of tumor have different genes mutated
- Early initiating events occur rarely in complex tissues and are therefore extremely difficult to detect
- The key initiating event oftenleads to an increase in mutation rate thus tumor cells often bear many fortuitous mutations

Important aspects of the disease we won't discuss

$$
\begin{gathered}
\text { relate to the spread of cancer cells } \\
\text { and the formation of large tumors } \\
\text { (metastasis and angiogenesis) }
\end{gathered}
$$

$3 T 3$ cells in culture

# Transformed 3T3 cells 

Images removed due to copyright reasons.

Isolation of the Ras
oncogene from fuman tumor cells


Focus of transformed 3T3 cells growing among untransformed cells


Extract genomic DNA $\downarrow$


Plate phage on
E. coli $\downarrow$

Figure by MIT OCW.


Synergistic effect of oncogenic forms of myc and ras


