

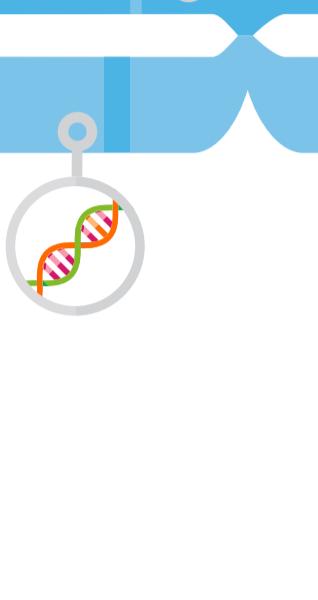
Predicting Cancer Immunotherapy Response with Tumor Mutational Burden (TMB)

Tumor Mutational Burden, or TMB, is a new, quantitative clinical marker that can help predict responses to certain cancer immunotherapies. Let's see how it works.

Cancer Immunotherapy

Cancer immunotherapies have the potential to treat cancer by harnessing the power of our **own immune systems**.

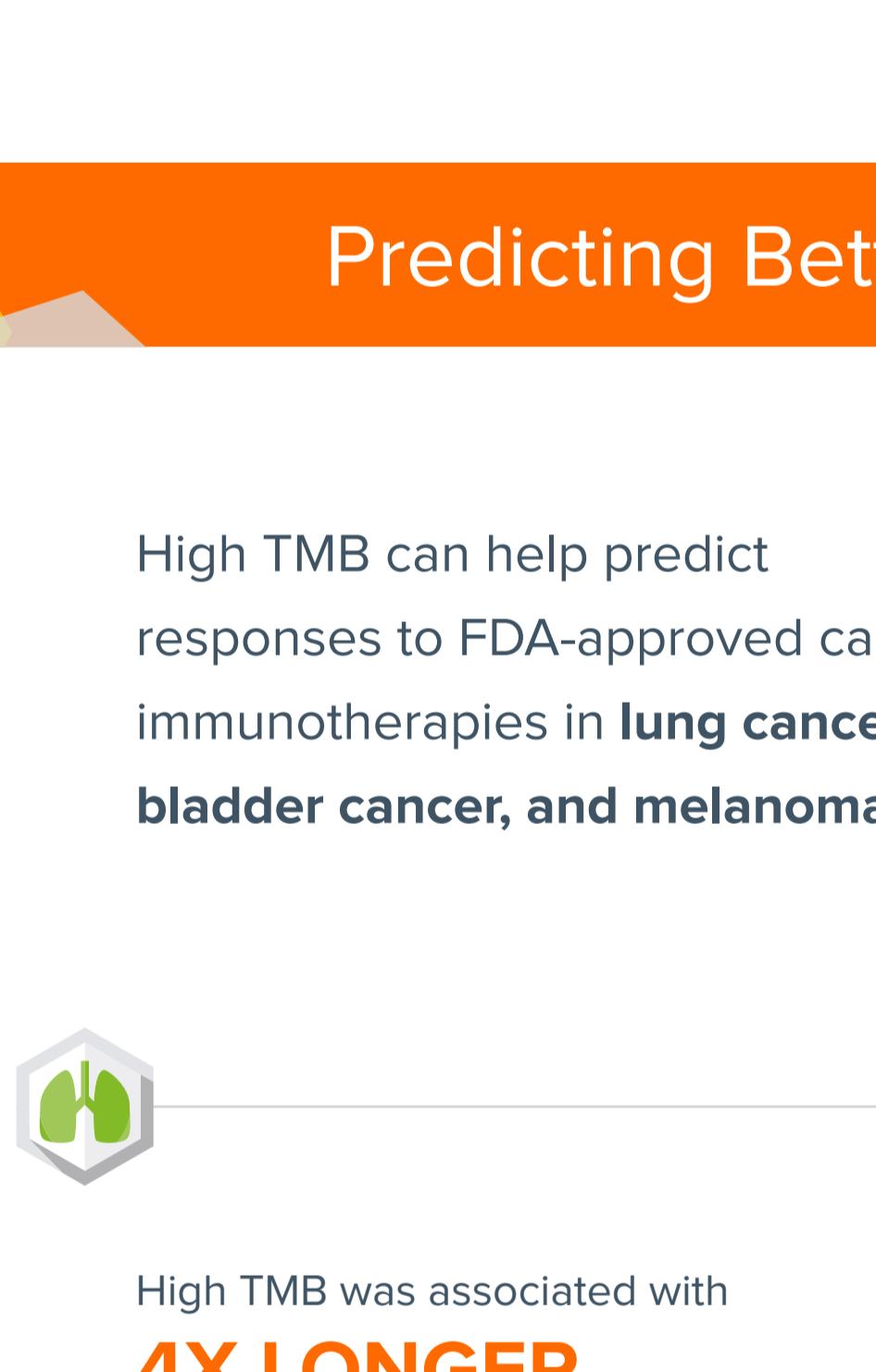
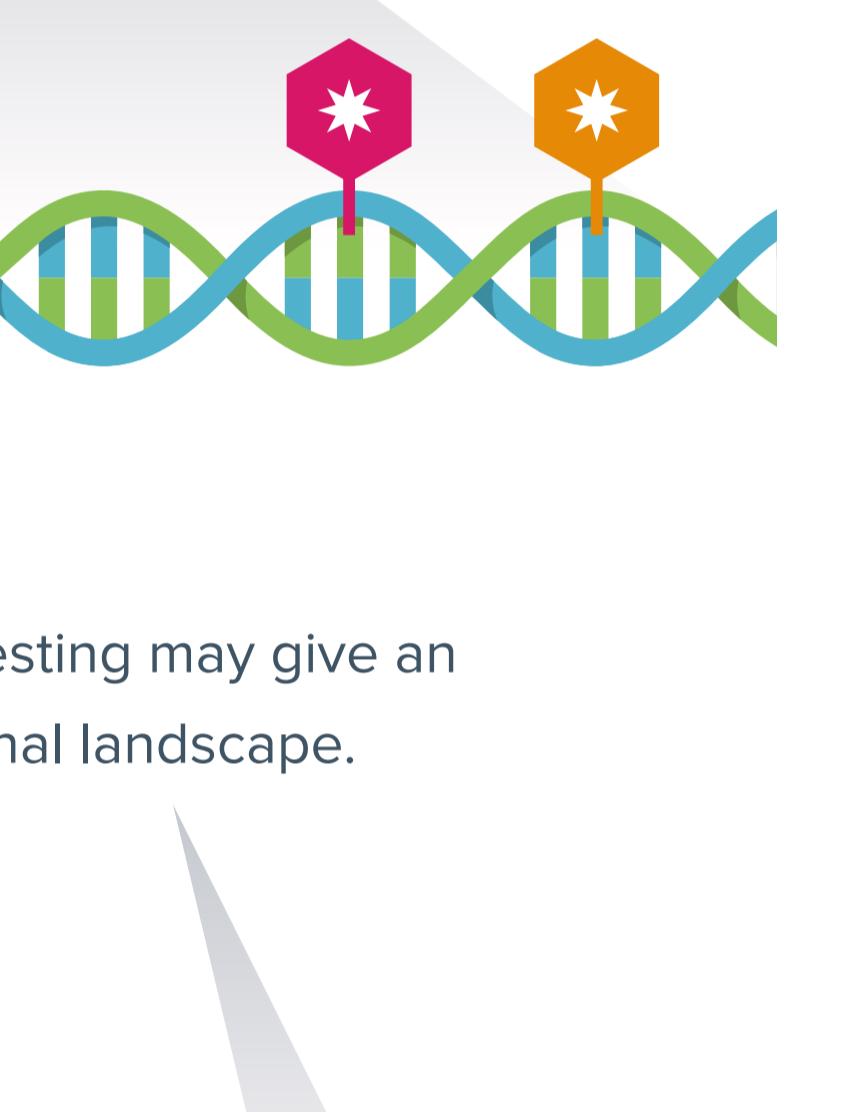
But right now, only about **20-40%** of people respond to this important new class of medicine.^{1,2}



The ability to predict who is most likely to respond to cancer immunotherapies could save **significant cost and precious time**.

A New, Quantitative Clinical Marker

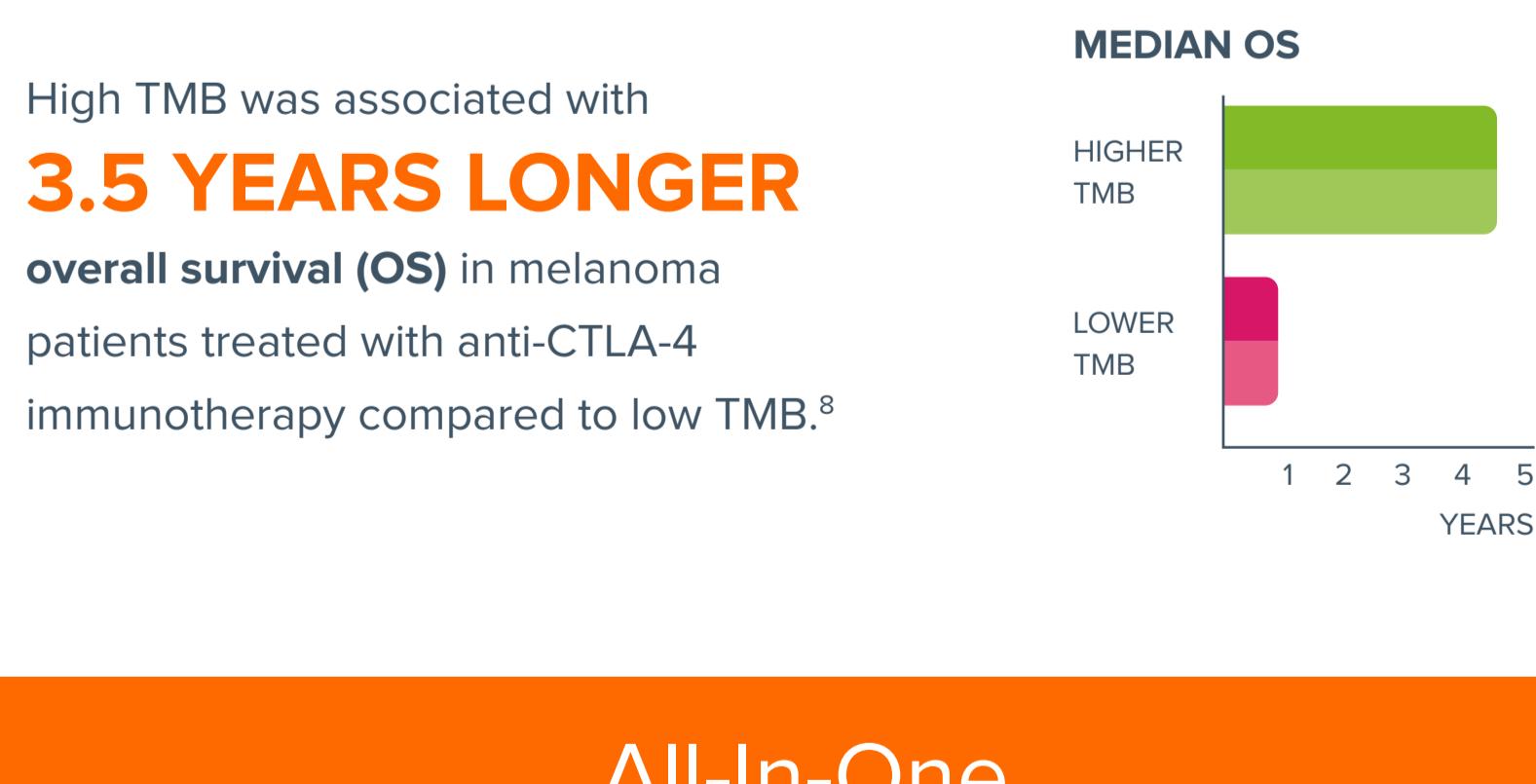
Tumor Mutational Burden (TMB) is the **TOTAL NUMBER OF MUTATIONS** per coding area of a tumor genome.



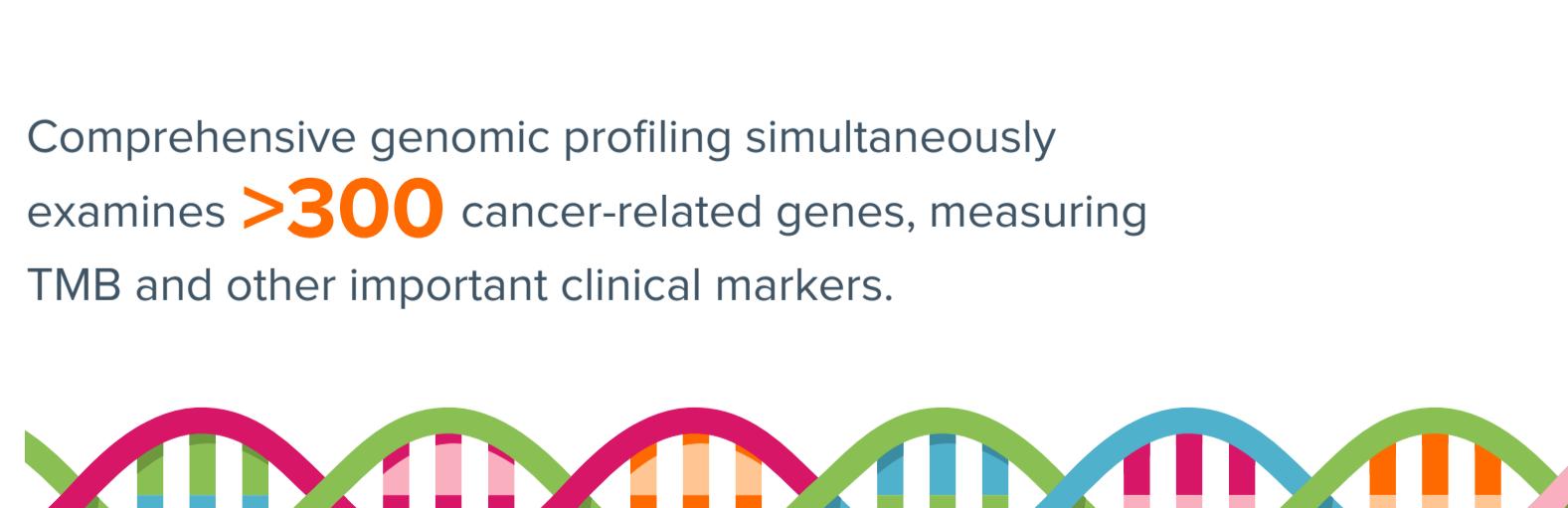
Higher TMB levels are correlated with **HIGHER LEVELS OF NEOANTIGENS** which help our immune system to recognize tumors.^{3,4}

Measuring TMB

TMB can be measured by sequencing the genome of a tumor by **comprehensive genomic profiling**.⁵



Traditional "hotspot" genomic testing may give an **incomplete view** of the mutational landscape.

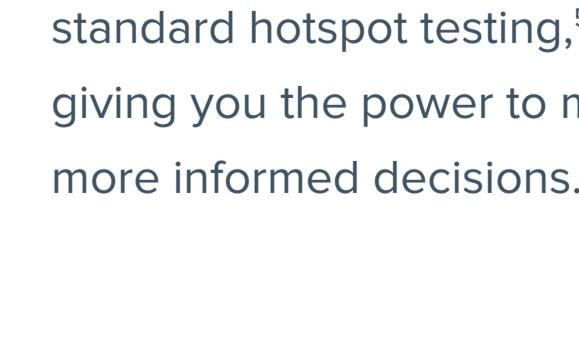


Predicting Better Responses

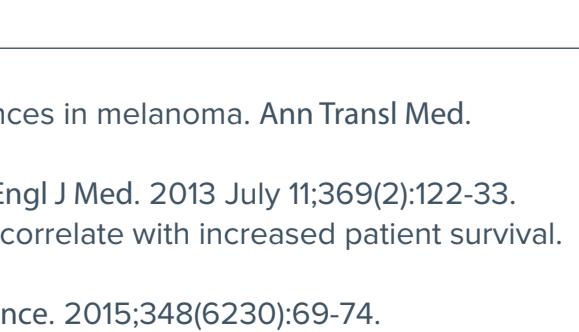
High TMB can help predict responses to FDA-approved cancer immunotherapies in **lung cancer, bladder cancer, and melanoma**.



High TMB was associated with **4X LONGER progression-free survival (PFS)** with anti-PD-1 immunotherapy in lung cancer compared to low TMB.⁶



In bladder cancer, high TMB predicted **BETTER RESPONSES** to anti-PD-L1 immunotherapy.⁷



All-In-One

Comprehensive genomic profiling simultaneously examines **>300** cancer-related genes, measuring TMB and other important clinical markers.

FOUNDATIONONE® provides insights for **CLINICAL TRIALS**, **TARGETED THERAPIES**, and **CANCER IMMUNOTHERAPIES**.

The comprehensive FoundationOne® test is designed to deliver

3X MORE actionable insights than standard hotspot testing,⁵ giving you the power to make more informed decisions.

1. Márquez-Rodas et al. Immune checkpoint inhibitors: therapeutic advances in melanoma. Ann Transl Med. 2015 October;3(18):267.

2. Wolchok et al. Nivolumab plus ipilimumab in advanced melanoma. N Engl J Med. 2013 July 11;369(2):122-33.

3. Brown et al. Neo-antigens predicted by tumor genome meta-analysis correlate with increased patient survival. Genome Res. 2014;24:743-750.

4. Schumacher & Schreiber. Neoantigens in cancer immunotherapy. Science. 2015;348(6230):69-74.

5. Frampton et al. Development and validation of a clinical cancer genomic profiling test based on massively parallel DNA sequencing. Nature Biotech. 2013;31:1023-1031.

6. Rizvi et al. Mutational landscape determines sensitivity to PD-1 blockade in non-small cell lung cancer. Science. 2015;348(6230):124-128.

7. Rosenberg et al. Atezolizumab in patients with locally advanced and metastatic urothelial carcinoma who have progressed following treatment with platinum-based chemotherapy: a single-arm, multicentre, phase 2 trial. Lancet. 2016;387(10031):1909-1920.

8. Snyder et al. Genetic basis for clinical response to CTLA-4 blockade in melanoma. NEJM. 2014;371:2189-2199.