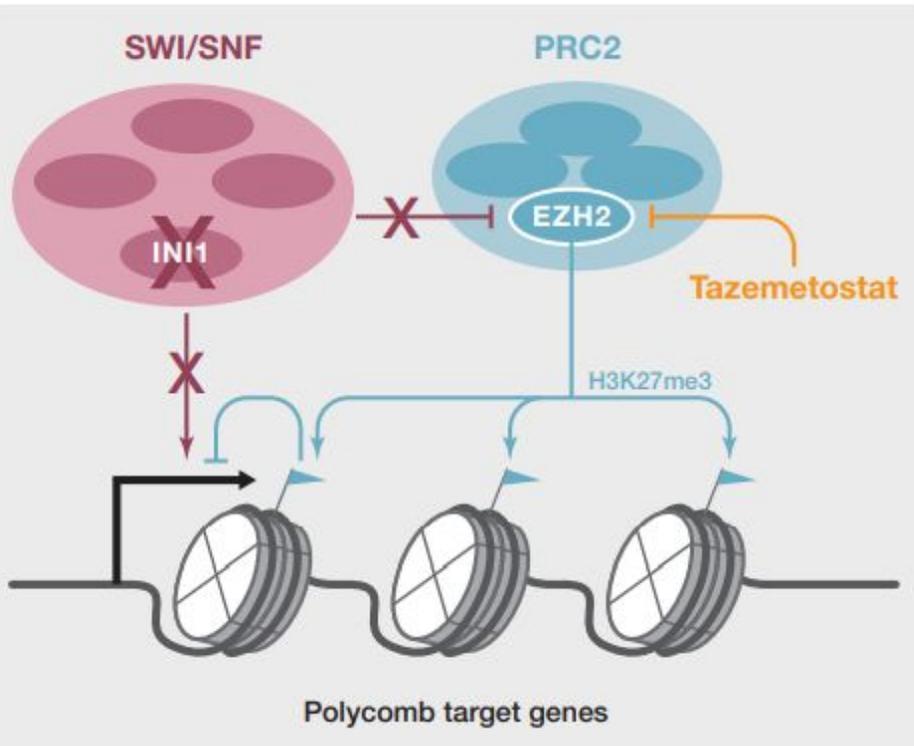


Science news

29.04.20

Подготовила
Князева Анастасия

Эпигенетическая терапия саркомы - первый подобный препарат, одобренный FDA



Prevalence

Epithelioid sarcoma
1.2%–1.5% of annual soft
tissue sarcoma diagnoses

Soft-tissue sarcoma
diagnoses are <1% of
annual cancer diagnoses

Approval stats

15%

Overall
response

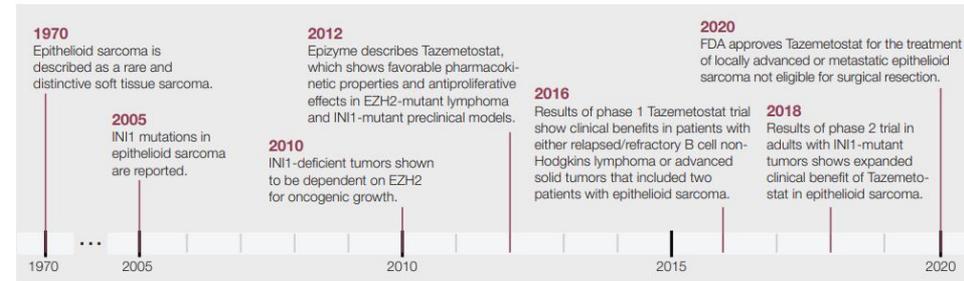
67%

Patients with
a response
of 6+ months

Duration of
response

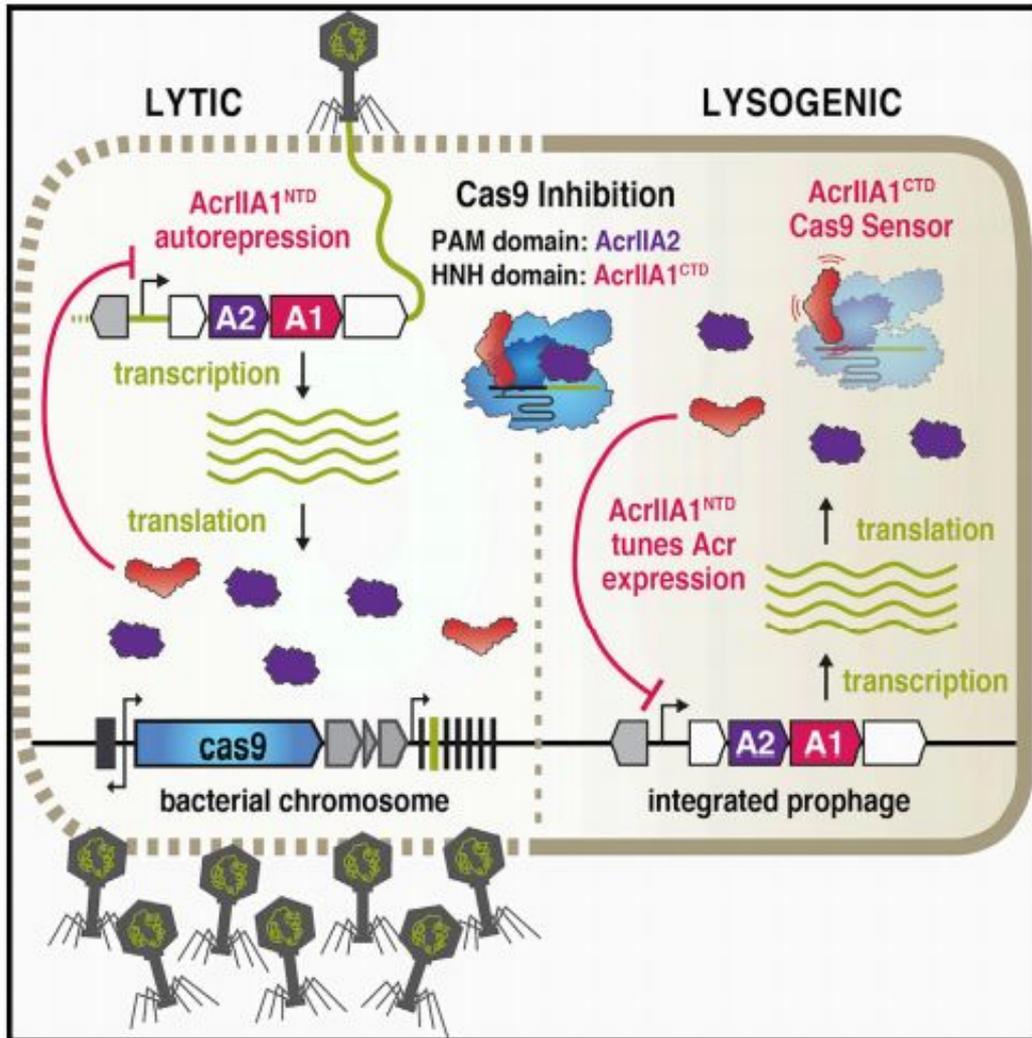
First approval

- In its class (KMT inhibitor)
- Epigenetic therapy for a solid tumor



Rothbart, Baylin/ Cell, 2020/

<http://dx.doi.org/10.1016/j.cell.2020.03.042>



Critical Anti-CRISPR Locus Repression by a Bifunctional Cas9 Inhibitor

- anti-CRISPR protein AcrIIA1 serves as an antiCRISPR and a vital autorepressor
- AcrIIA1 allows prophages to tune Acr expression to Cas9 levels
- AcrIIA1 homologs have been co-opted by host bacteria as “anti-anti-CRISPRs”

Osuna et al., 2020, Cell Host & Microbe, 2020

<https://doi.org/10.1016/j.chom.2020.04.002>

Микробиом ЖКТ важен для лечения рака

| Deleterious microbiota | | Favorable microbiota | |
|------------------------|---|-------------------------------|--|
| Diversity | Low | | High |
| |  <p>Antibiotics-related dysbiosis: Probably associated with decreased diversity</p> | |  |
| Cancer | | Mechanism | Mechanism |
| NSCLC RCC | <div style="border: 1px solid red; padding: 5px;"> - <i>Parabacteroides distasonis</i> - <i>Clostridiales bacterium</i> VE202-14 </div> | Unknown | <div style="border: 1px solid green; padding: 5px;"> - <i>Akkermanasia muciniphila</i> - <i>Alistipes indistinctus</i> - <i>Ruminococcus</i> spp </div> <ul style="list-style-type: none"> • Tumor: ↑CD4⁺ CCR9⁺, ↓T_{reg} • Peripheral Blood ↑INFγ production of CD4⁺ and CD8⁺ |
| Melanoma | <div style="border: 1px solid red; padding: 5px;"> - <i>Roseburia intestinalis</i> - <i>Bacteroidales</i> </div> | ↓ intratumor CD8 ⁺ | <div style="border: 1px solid green; padding: 5px;"> - <i>Collinesella aerofaciens</i> - <i>Bifidobacterium longum</i> - <i>Faecalibacterium</i> </div> <ul style="list-style-type: none"> • Tumor ↑CD8⁺ • Peripheral Blood ↑INFγ, ↑CD8⁺, ↓T_{reg} |

Deleterious microbiota

Favorable microbiota

Fecal microbiota transplantation

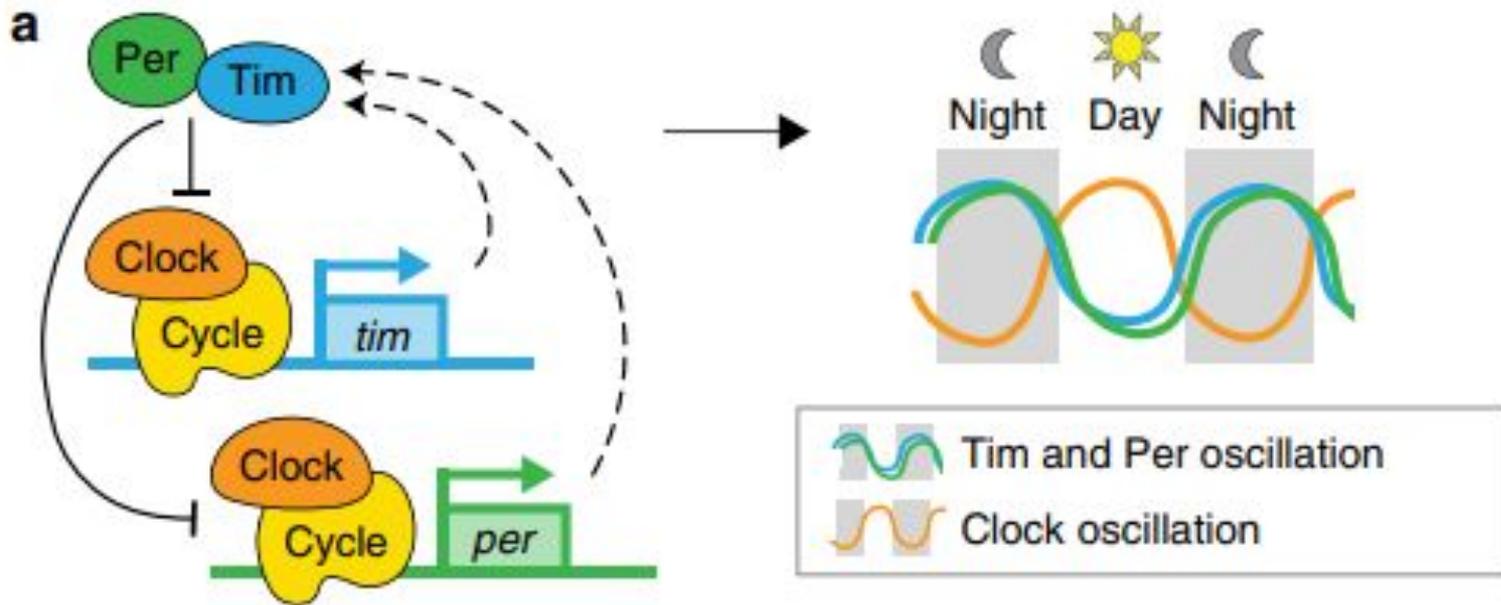
Dietary intervention

Probiotics: *Akkermanasia muciniphila*

Prebiotics

Co-medication

“Отключение” двух компонентов циркадных часов увеличило продолжительность жизни самцов дрозофил



Ulgherait et al., NatCom, 2020

<https://doi.org/10.1038/s41467-020-15617-x>