

# Mechanisms of melanoma-targeting antibody therapy in mice Benonisson, H.

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#### Stellingen behorende bij het proefschrift

### 'Mechanisms of melanoma-targeting therapy in mice

- Inflammatory macrophages expressing Fcγ receptor I are important for antibodymediated killing of tumor cells in mice (this thesis).
- The TA99 antibody combined with TLR ligands and IL2 induces tumor-specific CD8 T cell responses that are required for tumor control (this thesis).
- 3. The CD3-targeting bispecific antibody CD3xTRP1 does not induce protective immune memory against subsequent tumor challenge (this thesis).
- An mCMV-based vaccine induces an endogenous polyclonal antibody response against tumor surface antigens mediating Fcγ receptor-dependent killing of tumor cells (this thesis).
- A role of Fcγ receptors always need to be considered as a mechanism of antibodymediated treatments against cancer, contributing in a positive or negative way (Labrijn et al., Sci rep, 7: p2476, 2017).
- 6. Mice lacking the Fc-associated common γ chain are not suitable to study Fcγ receptors, as multiple other innate immune receptors, like MINCLE, are hampered as well (Yamasaki et al., Nat Immunol, 9: p1179, 2008).
- Extrapolation of mouse studies on Fcγ receptors to understand the human system is possible although a careful evaluation is always required due to species differences (Bruhns et al., Blood, 119: p5640, 2012).
- 8. CD3-targeting antibodies perform better when they do not bind Fcγ receptors (Labrijn, Sci rep, 7: p2476, 2017).
- 9. Unexpected results lead to scientific progress.
- 10. It is easier to design studies afterwards.

Hreinn Benonisson Leiden. 19 november 2019