Medical Oncology: the unmet needs and how we meet them

John Bridgewater and Juan Valle CCA-UK Inaugural Meeting 07 December 2018





Acknowledgement | 1301 UK patients have participated in NCRI Academic studies*

*ABC-01 to -08; BILCAP; PHOTOSTENT-2

CHOLANGIOCARCINOMA



Advanced disease | limited treatment options



- For our current treatment
 - Who are the "rapid nonresponders"?
 - Are there sub-groups who benefit most?
- Need for new agents
 - Acelarin
 - Immunotherapy
- Should we intensify?
 - FOLFIRINOX?
 - CisGem + nab-paclitaxel?
 - Chemosaturation?
- Efficacy vs quality of life

Need to accelerate progress





Second-line | no established treatment options



- Systematic review¹
 - n=895 patients
 - No conclusion
 regarding best
 approach
 - Patients are willing to participate in clinical trials
- Need to accelerate progress



Completed recruitment² | results ASCO 2019



¹ Lamarca et al 2014 Ann Oncol; ² ClinicalTrials.gov | NCT01926236

Incorporating radiation therapy

ABC-07



- Need to determine optimal sequencing
 - Need to explore novel approaches





Chemo and surgery | getting more patients to cure

Lessons from CRLM





- Neo-adjuvant
- Borderline resectable





Folprecht G et al. Ann Oncol 2005;16:1311-1319

Anatomical heterogeneity | lump or split?

ABC-02



ABC-02 + BT22



ADVANCED DISEASE



BILCAP



EARLY-STAGE DISEASE



Driver mutations | emerging and new







Driver mutations | targeted in practice

MOSCATO study | 1035 patients ≥1 prior systemic treatment, PS 0-1 Available tissue

| High-throughput molecular | analysis |
|---------------------------|----------|
| > BTC | n=43 |
| > Evaluable | n=34 |
| Druggable alteration | n=23 |
| > Treated | n=18 |

ORR 33%

DCR 88%

Median OS 17 mo vs. 5 mo

[HR 0.29; 95% CI, 0.11-0.76; p< 0.008).



Verlingue et al Eur J Cancer 2017;87:122e130



Translational unmet needs and how we meet them

- Optimising translational research
- Molecular understanding of CCA (prognostic and therapeutic)
- Accessibility of cancer centre care (research and expertise)
- International collaboration

- Co-ordinated basic science
- Translationally driven clinical studies
- Profile and education (and politics?)
- Harmonisation of regulation and funding

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A translational framework



Adapted from Wareham et al, 2009



